

Experiment Number: 00058 - 03
Test Type: CHRONIC
Route: GAVAGE
Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA
Black Cohosh
CAS Number: 84776-26-1

Date Report Requested: 11/13/2020
Time Report Requested: 09:33:45
First Dose M/F: 07/03/12 / 07/02/12
Lab: BAT

Rats Final 2

NTP Study Number: C00058B
Lock Date: 10/02/2018
Cage Range: ALL
Date Range: ALL
Reasons For Removal: ALL
Removal Date Range: ALL
Treatment Groups: Include ALL
Study Gender: Both
TDMSE Version: 3.0.2.3_002
PWG Approval Date: NONE

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Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

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First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 1

TRT#: 1

SEX: Male

DAY ON TEST: 731

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402740

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Bone | * Bone Marrow | * Brain | * Epididymis |
| * Esophagus | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Pancreas | * Parathyroid Gland | * Pituitary Gland | * Preputial Gland |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|-------------------|---------------------------|------------------------------|------------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Mild |
| * Adrenal Medulla | | Hyperplasia | Focal, Marked |
| | | Pheochromocytoma Benign | |
| * Eye | | Cataract | Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Lipoma | |
| | | Nephropathy | Chronicprogr, Moderate |
| | | Oncocytoma Benign | |
| * Liver | | Clear Cell Focus | |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Nose | Nasolacrim Dct | Inflammation | Chronic, Mild |
| * Prostate | | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| Testis | Bilateral, Germinal Epith | Degeneration | Marked |
| | | Granuloma Sperm | Minimal |
| | | Polyarteritis Nodosa | Mild |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | | Ectopic Thymus | Minimal |

* PROTOCOL REQUIRED TISSUE

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First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 2

TRT#: 1

SEX: Male

DAY ON TEST: 726

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402741

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Bone Marrow
- * Brain
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lung
- * Lymph Node, Mandibular
- * Mammary Gland
- * Parathyroid Gland
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- Testis
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel Mineral Mild
 - Note: The aorta and cardiac and renal arteries are mineralized.
- * Epididymis Mesothelioma Malignant
- * Heart Mineral Minimal
- * Kidney Nephropathy Chronicprogr, Moderate
 - [Nephropathy TGLs = 1 - 8]
- * Liver Bile Duct Hyperplasia Mild
 - Necrosis Moderate
 - [Necrosis TGLs = 3,4,5 - 18+19+20]
- Lymph Node Mediastinal Infiltration Cellular Histiocyte, Minimal
 - [Infiltration Cellular TGLs = 2 - 17]
- * Lymph Node, Mesenteric Infiltration Cellular Histiocyte, Minimal
 - [Infiltration Cellular TGLs = 6 - 9]
- * Nose Foreign Body Inflammation Suppurative, Minimal
 - Hyperplasia Mild
- * Pancreas Acinus Atrophy Mild
- * Spleen White Pulp Extramedullary Hematopoiesis Mild
 - Pigment Minimal
- * Stomach, Glandular Mineral Minimal
- * Thymus Atrophy Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

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P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

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Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 2

TRT#: 1

SEX: Male

DAY ON TEST: 726

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402741

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

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Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

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Date Report Requested: 11/13/2020

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First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 3

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402742

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|--------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Pancreas | * Parathyroid Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

MISSING

- * Mammary Gland

OBSERVATIONS

- | | | | |
|---------------------------|----------------------------------------|-------------------------|----------------------|
| * Adrenal Medulla | Bilateral | Pheochromocytoma Benign | |
| * Heart | | Cardiomyopathy | Minimal |
| * Intestine Large, Rectum | | Polyarteritis Nodosa | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 6 - 8] | | |
| * Liver | | Angiectasis | Moderate |
| | | Clear Cell Focus | |
| | Bile Duct | Dilation | Moderate |
| | | Eosinophilic Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| | [Angiectasis TGLs = 7 - 18] | | |
| | [Dilation TGLs = 3 - 17] | | |
| * Lung | Alveolar Epith | Hyperplasia | Mild |
| | | Infiltration Cellular | Histiocyte, Minimal |
| | Perivascular | Infiltration Cellular | Lymphoid, Mild |
| | [Infiltration Cellular TGLs = 4 - 7] | | |
| * Nose | | Foreign Body | |
| | | Inflammation | Acute, Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 5 - 11] | | |

* PROTOCOL REQUIRED TISSUE

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Test Type: CHRONIC

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Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

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Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 3

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402742

ORGAN AND ACCOUNTABLE SITE STATUS

* Skin		Lipoma	
	[Lipoma TGLs = 1 - 19]		
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
Testis	Interstit Cell	Adenoma	
	Bilateral, Germinal Epith	Degeneration	Minimal
* Thymus		Atrophy	Mild

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Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 4

TRT#: 1

SEX: Male

DAY ON TEST: 674

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402743

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Preputial Gland |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | Testis | * Thyroid Gland | * Trachea |

OBSERVATIONS

- | | | | |
|---------------------|------------------------------|------------------------------|-------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| | | Vacuolization Cytoplasmic | Mild |
| * Adrenal Medulla | | Pheochromocytoma Benign | |
| * Bone Marrow | | Hypercellularity | Moderate |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Prostate | | Inflammation | Chronic Active, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Moderate |
| * Urinary Bladder | | Inflammation | Chronic, Minimal |
| Zymbal's Gland | | Carcinoma | |
| | [Carcinoma TGLs = 2 - 17] | | |

PRIMARY CAUSE OF DEATH - Zymbal's Gland Carcinoma

* PROTOCOL REQUIRED TISSUE

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Species/Strain: RATS/HSD

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Black Cohosh

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First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 5

TRT#: 1

SEX: Male

DAY ON TEST: 654

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402744

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Parathyroid Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Trachea | * Urinary Bladder |

MISSING

- * Thymus

OBSERVATIONS

- | | | | |
|-------------------------------------------------------|----------------|-------------------------------|-----------------------------------|
| Blood Vessel | | Mineral | Minimal |
| Note: The aorta and cardiac arteries are mineralized. | | | |
| * Bone Marrow | | Hypercellularity | Moderate |
| * Heart | | Cardiomyopathy | Minimal |
| * Intestine Large, Colon | | Parasite Metazoan | |
| * Kidney | Renal Tubule | Accumulation, Hyaline Droplet | Minimal |
| | Renal Tubule | Cyst | |
| | | Nephropathy | Chronicprogr, Marked |
| | | | [Cyst TGLs = 2,4 - 8+17] |
| | | | [Nephropathy TGLs = 3,5 - 8+17] |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Mammary Gland | | Fibroadenoma | |
| | | | [Fibroadenoma TGLs = 1 - 18] |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Stomach, Glandular | | Mineral | Moderate |
| Testis | Germinal Epith | Degeneration | Minimal |
| * Thyroid Gland | C Cell | Adenoma | |
| | C Cell | Hyperplasia | Mild |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

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Species/Strain: RATS/HSD

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First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 6

TRT#: 1

SEX: Male

DAY ON TEST: 704

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402745

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Preputial Gland
- * Stomach, Glandular
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lung
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Trachea

OBSERVATIONS

- * Adrenal Medulla [Pheochromocytoma Benign TGLs = 2 - 11] Pheochromocytoma Benign
- * Bone Marrow Hypercellularity Moderate
- * Eye Retina Degeneration Mild
- Cornea Inflammation Chronic, Mild
- * Heart Cardiomyopathy Minimal
- * Kidney Nephropathy Chronicprogr, Moderate
- * Liver Clear Cell Focus
- * Pituitary Gland Pars Distalis Adenoma
- [Adenoma TGLs = 3 - 11]
- * Prostate Inflammation Chronic Active, Minimal
- * Skin Fibroma
- [Fibroma TGLs = 1 - 17]
- * Spleen Extramedullary Hematopoiesis Moderate
- Testis Interstit Cell Hyperplasia Mild
- * Thymus Atrophy Marked

PRIMARY CAUSE OF DEATH - Skin Fibroma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

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First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 7

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402746

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Epididymis | * Esophagus |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Pancreas | * Parathyroid Gland | * Preputial Gland | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-------------------------|
| * Eye | Retina | Degeneration | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| * Lung | [Infiltration Cellular TGLs = 1 - 6+7] | Infiltration Cellular | Histiocyte, Mild |
| * Nose | Nasolacrim Dct | Inflammation | Chronic Active, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Prostate | | Inflammation | Chronic Active, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Hyperplasia | Marked |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

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First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 8

TRT#: 1

SEX: Male

DAY ON TEST: 598

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402747

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Medulla | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Parathyroid Gland |
| * Pituitary Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|-------------------|------|------------------------------|-------------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Nose | | Inflammation | Chronic Active, Minimal |
| * Pancreas | | Polyarteritis Nodosa | Mild |
| * Preputial Gland | Duct | Hyperplasia | Squamous, Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | | Polyarteritis Nodosa | Mild |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

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Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 9

TRT#: 1

SEX: Male

DAY ON TEST: 680

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402748

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Bone
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Bone Marrow
- * Eye
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
Blood Vessel
Note: The aorta, cardiac arteries, and pulmonary vein are mineralized.
 - * Heart
 - * Intestine Large, Cecum
 - * Kidney
[Nephropathy TGLs = 1 - 8]
 - * Liver
 - * Lung
 - * Pancreas
 - * Pituitary Gland
 - * Spleen
 - * Stomach, Glandular
 - Testis
 - * Thymus
- | | | |
|--|----------------------|----------------------|
| | Degeneration | Cystic, Minimal |
| | Mineral | Minimal |
| | Cardiomyopathy | Mild |
| | Erosion | Mild |
| | Inflammation | Chronic, Minimal |
| | Mineral | Mild |
| | Polyarteritis Nodosa | Mild |
| | Nephropathy | Chronicprogr, Marked |
| | Bile Duct | Hyperplasia |
| | Alveolar Epith | Hyperplasia |
| | Acinus | Adenoma |
| | Acinus | Hyperplasia |
| | Pars Distalis | Hyperplasia |
| | | Pigment |
| | | Mineral |
| | | Degeneration |
| | | Polyarteritis Nodosa |
| | | Atrophy |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 10

TRT#: 1

SEX: Male

DAY ON TEST: 562

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402749

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Prostate
- * Stomach, Forestomach
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Seminal Vesicle
- * Trachea
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta, and cardiac, mammary, and salivary arteries are mineralized.
- * Bone Marrow
- * Epididymis
- * Heart
- * Kidney
[Nephropathy TGLs = 2 - 8]
- * Liver
- * Lung
- * Pituitary Gland
- * Spleen
- * Stomach, Glandular
- Testis
[Degeneration TGLs = 1 - 14]
- * Thymus
- Pheochromocytoma Benign
- Mineral
- Hemorrhage
- Hypercellularity
- Mesothelioma Malignant
- Cardiomyopathy
- Mineral
- Nephropathy
- Polyarteritis Nodosa
- Mineral
- Hyperplasia
- Atrophy
- Pigment
- Mineral
- Degeneration
- Polyarteritis Nodosa
- Atrophy
- Mild
- Moderate
- Mild
- Mild
- Mild
- Mild
- Mild
- Moderate
- Chronicprogr, Marked
- Minimal
- Minimal
- Minimal
- Moderate
- Mild
- Mild
- Moderate
- Minimal
- Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 11

TRT#: 1

SEX: Male

DAY ON TEST: 718

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402750

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Bone
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Forestomach
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Epididymis
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Eye
- * Heart
- * Intestine Small, Duodenum
- * Kidney
- * Liver
- * Lung
- Lymph Node
- * Mammary Gland
- * Nose
- * Parathyroid Gland
- * Spleen
- Testis
- Cornea
- Renal Tubule
- Bile Duct
- Interstitial
- Mediastinal
- Nasolacrim Dct
- Bilateral, Germinal Epith
- Hyperplasia
- Hypercellularity
- Inflammation
- Cardiomyopathy
- Polyarteritis Nodosa
- Cyst
- Nephropathy
- Clear Cell Focus
- Hyperplasia
- Necrosis
- Fibrosis
- Infiltration Cellular
- Inflammation
- Thrombus
- Hyperplasia
- Inflammation
- Hyperplasia
- Extramedullary Hematopoiesis
- Degeneration
- Focal, Moderate
- Moderate
- Acute, Moderate
- Minimal
- Mild
- Chronicprogr, Marked
- Minimal
- Minimal
- Minimal
- Histiocyte, Moderate
- Acute, Moderate
- Minimal
- Chronic, Mild
- Diffuse, Moderate
- Moderate
- Minimal

[Cyst TGLs = 2 - 17]
[Nephropathy TGLs = 1 - 8]

[Thrombus TGLs = 5 - 19]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 11

TRT#: 1

SEX: Male

DAY ON TEST: 718

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402750

ORGAN AND ACCOUNTABLE SITE STATUS

Polyarteritis Nodosa

Mild

Atrophy

Marked

* Thymus

* Thyroid Gland

C Cell

Adenoma

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 12

TRT#: 1

SEX: Male

DAY ON TEST: 573

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402751

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Urinary Bladder
- Blood Vessel
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- Testis
- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Thyroid Gland
- * Bone Marrow
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Heart
- * Kidney [Nephropathy TGLs = 1 - 17+18]
- * Lung
- * Nose
- * Pituitary Gland
- * Spleen
- * Thymus
- Hyperplasia
- Cardiomyopathy
- Nephropathy
- Infiltration Cellular
- Foreign Body
- Inflammation
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Focal, Mild
- Minimal
- Chronicprogr, Marked
- Histiocyte, Minimal
- Chronic, Minimal
- Minimal
- Mild
- Minimal
- Moderate

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 13

TRT#: 1

SEX: Male

DAY ON TEST: 711

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402752

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Esophagus | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|-------------------------------------------------------|---------------|--------------------|----------------------|
| Blood Vessel | | Mineral | Minimal |
| Note: The aorta and cardiac arteries are mineralized. | | | |
| * Bone Marrow | | Hemorrhage | Moderate |
| | | Hypercellularity | Moderate |
| * Epididymis | Bilateral | Atrophy | Mild |
| [Atrophy TGLs = 1,2 - 14] | | | |
| * Eye | Retina | Degeneration | Minimal |
| | Retina | Fibrosis | Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| [Nephropathy TGLs = 5,6 - 8] | | | |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Cyst | |
| | Bile Duct | Dilation | Marked |
| | | Eosinophilic Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| | | Mixed Cell Focus | |
| [Dilation TGLs = 9 - 18] | | | |
| [Mixed Cell Focus TGLs = 8 - 17] | | | |
| * Lung | | Inflammation | Acute, Minimal |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 13

TRT#: 1

SEX: Male

DAY ON TEST: 711

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402752

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Mild
* Stomach, Glandular		Mineral	Mild
Testis	Bilateral, Germinal Epith	Degeneration	Marked
		Polyarteritis Nodosa	Minimal
	[Degeneration TGLs = 3,4 - 14]		
* Thymus		Atrophy	Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 14

TRT#: 1

SEX: Male

DAY ON TEST: 643

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402753

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Medulla | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Parathyroid Gland | * Pituitary Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|------------------|---------------------------------------------|------------------------------|--------------------|
| * Adrenal Cortex | Bilateral | Hyperplasia | Focal, Mild |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Infarct | Minimal |
| | | Nephropathy | Chronicprogr, Mild |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Skin | | Cyst Epithelial Inclusion | |
| | [Cyst Epithelial Inclusion TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Moderate |
| | | Polyarteritis Nodosa | Minimal |
| * Thymus | | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH - UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 15

TRT#: 1

SEX: Male

DAY ON TEST: 646

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1401754

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lung
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Preputial Gland
- * Seminal Vesicle
- * Stomach, Forestomach
- * Testis
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel Mineral Minimal
Note: A renal artery is mineralized.
- * Bone Marrow Hypercellularity Marked
- * Kidney Nephropathy Chronicprogr, Marked
[Nephropathy TGLs = 3,4 - 8]
- * Liver Necrosis Mild
- * Pituitary Gland Pars Distalis Adenoma
- [Adenoma TGLs = 5 - 11]
- * Prostate Inflammation Chronic Active, Moderate
- * Salivary Glands Fibrosarcoma Metastatic (Skin)
- * Skin Fibrosarcoma
- [Fibrosarcoma TGLs = 1 - 17]
- * Spleen White Pulp Atrophy Marked
Extramedullary Hematopoiesis Marked
[Extramedullary Hematopoiesis TGLs = 2 - 8]
- * Stomach, Glandular Mineral Minimal
- * Thymus Atrophy Marked

PRIMARY CAUSE OF DEATH - Skin Fibrosarcoma

CONTRIBUTORY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 16

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402755

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Cortex | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Parathyroid Gland |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thymus | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------------|------------------------------|----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 4 - 20] | | |
| * Liver | | Clear Cell Focus | |
| * Nose | Nasolacrim Dct | Inflammation | Chronic, Minimal |
| | | Inflammation | Acute, Minimal |
| * Pancreas | Acinus | Adenoma | |
| | Acinus | Atrophy | Mild |
| | Acinus | Hyperplasia | Marked |
| | [Adenoma TGLs = 2 - 18] | | |
| | [Hyperplasia TGLs = 1,3 - 17 + 19] | | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| Testis | | Polyarteritis Nodosa | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 17

TRT#: 1

SEX: Male

DAY ON TEST: 624

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402756

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|--------------------------|---------------------------|-----------------------------|
| * Adrenal Cortex | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver | * Lung |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Parathyroid Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thymus | * Trachea | * Urinary Bladder | |

MISSING

- * Intestine Small, Ileum

OBSERVATIONS

- | | | | |
|-------------------|----------------------------------------------------------------------|----------------------|----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Pancreas | Acinus | Atrophy | Minimal |
| | | Polyarteritis Nodosa | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| | Note: Hyperplasia of pars distalis cells but within pars intermedia. | | |
| * Spleen | | Pigment | Minimal |
| Testis | | Polyarteritis Nodosa | Minimal |
| * Thyroid Gland | C Cell | Hyperplasia | Moderate |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 19

TRT#: 1

SEX: Male

DAY ON TEST: 711

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402758

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Stomach, Glandular
- * Bone
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Seminal Vesicle
- * Thyroid Gland
- * Brain
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Preputial Gland
- * Skin
- * Trachea
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder

MISSING

- * Mammary Gland

OBSERVATIONS

- * Adrenal Medulla
[Pheochromocytoma Benign TGLs = 3 - 11] Pheochromocytoma Benign
- * Bone Marrow Hypercellularity Marked
- * Epididymis Atrophy Mild
[Atrophy TGLs = 8,9 - 14]
- * Eye Cornea Inflammation Acute, Moderate
- * Heart Atrium Cardiomyopathy Minimal
[Thrombus TGLs = 4 - 10] Thrombus
- * Kidney Nephropathy Chronicprogr, Marked
[Nephropathy TGLs = 5,6 - 8]
- * Liver Bile Duct Dilation Minimal
Bile Duct Hyperplasia Minimal
Necrosis Minimal
- * Lung Infiltration Cellular Histiocyte, Minimal
Inflammation Granulomatous, Minimal
[Infiltration Cellular TGLs = 10 - 6]
- * Nose Inflammation Suppurative, Minimal
- * Pancreas Acinus Carcinoma
- * Parathyroid Gland Hyperplasia Diffuse, Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 19

TRT#: 1

SEX: Male

DAY ON TEST: 711

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402758

ORGAN AND ACCOUNTABLE SITE STATUS

* Pituitary Gland	Pars Distalis	Hyperplasia	Minimal
* Spleen	White Pulp	Atrophy	Moderate
		Extramedullary Hematopoiesis	Mild
Testis	Germinal Epith	Degeneration	Marked
		Polyarteritis Nodosa	Moderate
[Degeneration TGLs = 1,2 - 14]			
* Thymus		Atrophy	Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 20

TRT#: 1

SEX: Male

DAY ON TEST: 538

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402759

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pancreas
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Testis
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

MISSING

- * Parathyroid Gland

OBSERVATIONS

- Blood Vessel Mineral Mild
- Note: The aorta and cardiac, prostatic, and testicular arteries are mineralized.
- * Bone Marrow Hypercellularity Mild
- * Heart Cardiomyopathy Minimal
- * Intestine Large, Cecum Inflammation Acute, Minimal
- * Kidney Hemorrhage Marked
- [Hemorrhage TGLs = 2 - 17]
- [Nephropathy TGLs = 1 - 8]
- * Liver Nephropathy Chronicprogr, Marked
- * Lung Necrosis Mild
- Infiltration Cellular Histiocyte, Minimal
- Inflammation Granulomatous, Minimal
- Mineral Minimal
- * Pituitary Gland Pars Distalis Hyperplasia Minimal
- * Spleen White Pulp Atrophy Mild
- Extramedullary Hematopoiesis Moderate
- * Stomach, Glandular Mineral Moderate
- * Thymus Atrophy Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 21

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402760

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone Marrow
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Stomach, Glandular
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
 - * Adrenal Medulla
 - * Heart
 - * Kidney
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 2 - 6]
 - * Pancreas
 - * Pituitary Gland
 - * Spleen
 - Testis
 - [Degeneration TGLs = 1 - 14]
 - * Thymus
- | | | | |
|--|----------------|------------------------------|------------------------|
| | | | |
| | Bilateral | Degeneration | Cystic, Minimal |
| | | Hyperplasia | Focal, Marked |
| | | Cardiomyopathy | Minimal |
| | | Nephropathy | Chronicprogr, Moderate |
| | | Clear Cell Focus | |
| | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Minimal |
| | Acinus | Hyperplasia | Minimal |
| | Pars Distalis | Hyperplasia | Minimal |
| | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| | Germinal Epith | Degeneration | Moderate |
| | | Polyarteritis Nodosa | Minimal |
| | | Atrophy | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 22

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402761

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- Testis
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Parathyroid Gland
- * Seminal Vesicle
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Lung
- * Preputial Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Prostate
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Medulla
[Pheochromocytoma Benign TGLs = 5 - 11]
- * Bone Marrow
- * Heart
- * Islets, Pancreatic
[Adenoma TGLs = 8 - 22]
- * Kidney
- * Liver
- * Mammary Gland
[Fibroadenoma TGLs = 1 - 17]
- * Nose
- * Pancreas
- * Pituitary Gland
- * Skin
[Fibroma TGLs = 3 - 19; 7 - 21]
[Lipoma TGLs = 2 - 18; 4 - 20]
- * Spleen
[Extramedullary Hematopoiesis TGLs = 6 - 8]
- * Thymus
- * Thyroid Gland
- Bile Duct
- Acinus
- Pars Distalis
- C Cell
- Pheochromocytoma Benign
- Hypercellularity
- Cardiomyopathy
- Adenoma
- Nephropathy
- Eosinophilic Focus
- Hyperplasia
- Fibroadenoma
- Inflammation
- Atrophy
- Adenoma
- Fibroma
- Lipoma
- Extramedullary Hematopoiesis
- Atrophy
- Adenoma
- Marked
- Minimal
- Chronicprogr, Mild
- Mild
- Chronic Active, Minimal
- Mild
- Multiple
- Multiple
- Marked
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 22

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402761

ORGAN AND ACCOUNTABLE SITE STATUS

[Adenoma TGLs = 10 - 11]

C Cell

Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 23

TRT#: 1

SEX: Male

DAY ON TEST: 577

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402762

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Prostate
- * Stomach, Glandular
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lung
- * Pancreas
- * Salivary Glands
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla [Pheochromocytoma Benign TGLs = 3 - 11] Pheochromocytoma Benign
- * Bone Marrow Hypercellularity Moderate
- * Heart Cardiomyopathy Minimal
- * Kidney Nephropathy Chronicprogr, Moderate
- * Liver Necrosis Mild
- * Mammary Gland Galactocele Mild
- [Galactocele TGLs = 4 - 18]
- * Pituitary Gland Pars Distalis Hyperplasia Minimal
- * Skin Fibroma Fibroma
- [Fibroma TGLs = 2 - 17]
- * Spleen White Pulp Atrophy Mild
- Testis Bilateral, Germinal Epith Extramedullary Hematopoiesis Moderate
- [Degeneration TGLs = 1 - 14] Degeneration Moderate
- * Thymus Atrophy Marked
- * Urinary Bladder Inflammation Chronic, Minimal

PRIMARY CAUSE OF DEATH - Skin Fibroma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 24

TRT#: 1

SEX: Male

DAY ON TEST: 672

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402763

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea
- * Brain
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Epididymis
- * Intestine Large, Rectum
- * Liver
- * Nose
- * Prostate
- * Stomach, Forestomach

OBSERVATIONS

- * Adrenal Cortex
Blood Vessel
Note: The aorta and cardiac arteries and pulmonary vein are mineralized.
- * Bone Marrow
- * Eye
Cornea
- * Heart
- * Intestine Large, Cecum
- * Islets, Pancreatic
- * Kidney
[Nephropathy TGLs = 1,2 - 8]
- * Lung
- * Parathyroid Gland
- * Spleen
White Pulp
- * Stomach, Glandular
Testis
Bilateral, Germinal Epith
- Hyperplasia
- Mineral
- Hemorrhage
- Hypercellularity
- Inflammation
- Cardiomyopathy
- Inflammation
- Mineral
- Ulcer
- Hyperplasia
- Nephropathy
- Infiltration Cellular
- Inflammation
- Mineral
- Hyperplasia
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Mineral
- Degeneration
- Focal, Minimal
- Mild
- Mild
- Mild
- Acute, Mild
- Moderate
- Chronic Active, Moderate
- Moderate
- Moderate
- Moderate
- Chronicprogr, Marked
- Histiocyte, Minimal
- Granulomatous, Minimal
- Minimal
- Diffuse, Moderate
- Marked
- Minimal
- Minimal
- Mild
- Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 24

TRT#: 1

SEX: Male

DAY ON TEST: 672

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402763

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Marked

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 26

TRT#: 1

SEX: Male

DAY ON TEST: 731

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402765

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Skin
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Stomach, Glandular
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Heart
- * Kidney
- * Liver
- Bile Duct
- * Lung
- * Pituitary Gland
- * Prostate
- * Spleen
- Testis
- * Thymus
- * Thyroid Gland
- Pars Distalis
- Germinal Epith
- C Cell
- Pheochromocytoma Benign
- Hypercellularity
- Cardiomyopathy
- Nephropathy
- Clear Cell Focus
- Hyperplasia
- Necrosis
- Infiltration Cellular
- Adenoma
- Inflammation
- Extramedullary Hematopoiesis
- Pigment
- Degeneration
- Atrophy
- Adenoma
- Mild
- Minimal
- Chronicprogr, Moderate
- Minimal
- Minimal
- Histiocyte, Minimal
- Chronic Active, Mild
- Moderate
- Minimal
- Minimal
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 27

TRT#: 1

SEX: Male

DAY ON TEST: 720

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402766

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Bone
- * Harderian Gland
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Brain
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Trachea
- * Epididymis
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta and cardiac, pulmonary, and renal arteries are mineralized.
- * Bone Marrow
- * Eye
* Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Kidney
[Nephropathy TGLs = 2 - 8]
- * Liver
* Lung
[Infiltration Cellular TGLs = 3 - 6]
[Mineral TGLs = 4,5 - 6+7]
- * Parathyroid Gland
- * Pituitary Gland
[Hyperplasia TGLs = 6 - 11]
- Cornea
- Bile Duct
- Pars Distalis
- Hyperplasia
- Mineral
- Hemorrhage
- Hypercellularity
- Inflammation
- Cardiomyopathy
- Mineral
- Erosion
- Mineral
- Ulcer
- Inflammation
- Hyperplasia
- Nephropathy
- Hyperplasia
- Infiltration Cellular
- Mineral
- Hyperplasia
- Hyperplasia
- Focal, Minimal
- Mild
- Moderate
- Mild
- Chronic Active, Marked
- Moderate
- Minimal
- Mild
- Mild
- Mild
- Chronic, Minimal
- Oncocytic, Marked
- Chronicprogr, Marked
- Minimal
- Histiocyte, Minimal
- Mild
- Diffuse, Marked
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 27

TRT#: 1

SEX: Male

DAY ON TEST: 720

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402766

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen	White Pulp	Atrophy	Moderate
		Pigment	Mild
* Stomach, Glandular		Mineral	Marked
Testis	Bilateral, Germinal Epith	Degeneration	Moderate
		Polyarteritis Nodosa	Minimal
	[Degeneration TGLs = 1 - 14]		
* Thymus		Atrophy	Marked
* Thyroid Gland	C Cell	Hyperplasia	Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 28

TRT#: 1

SEX: Male

DAY ON TEST: 730

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402767

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Lymph Node, Mesenteric | * Parathyroid Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|--------------------------|-----------------------------------------|------------------------------|------------------------|
| * Heart | | Cardiomyopathy | Mild |
| * Islets, Pancreatic | | Adenoma | |
| | [Adenoma TGLs = 4 - 17] | | |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Cyst | |
| | Bile Duct | Hyperplasia | Mild |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Lymph Node, Mandibular | | Infiltration Cellular | Plasma Cell, Mild |
| | [Infiltration Cellular TGLs = 1 - 12] | | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 2 - 18] | | |
| * Nose | Respirat Epith | Hyperplasia | Minimal |
| * Pancreas | Acinus | Hyperplasia | Marked |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Skin | | Fibroma | |
| | [Fibroma TGLs = 3 - 19] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 29

TRT#: 1

SEX: Male

DAY ON TEST: 731

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402768

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Bone Marrow | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Trachea |
| * Urinary Bladder | | | |

OBSERVATIONS

- | | | | |
|-------------------|---------------|------------------------------|------------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Adrenal Medulla | Bilateral | Hyperplasia | Focal, Mild |
| * Brain | | Glioma Malignant | |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Pancreas | Acinus | Atrophy | Minimal |
| | Acinus | Hyperplasia | Moderate |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Adenoma | |
| | C Cell | Hyperplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 30

TRT#: 1

SEX: Male

DAY ON TEST: 616

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402769

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Liver
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Pancreas
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel Mineral Moderate
- Note: The aorta and cardiac, mesenteric, pancreatic, pulmonary and renal arteries are mineralized.
- * Bone Marrow Hemorrhage Marked
- Hypercellularity Mild
- * Heart Cardiomyopathy Minimal
- * Kidney Nephropathy Chronicprogr, Marked
- * Lung Mineral Minimal
- * Nose Inflammation Suppurative, Minimal
- * Parathyroid Gland Hyperplasia Diffuse, Marked
- * Spleen White Pulp Atrophy Marked
- Pigment Mild
- * Stomach, Glandular Mineral Mild
- Testis Polyarteritis Nodosa Mild
- * Thymus Atrophy Marked
- Polyarteritis Nodosa Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 31

TRT#: 1

SEX: Male

DAY ON TEST: 618

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402770

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Seminal Vesicle
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Skin
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Parathyroid Gland
- * Stomach, Forestomach
- * Epididymis
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Preputial Gland
- Testis

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- [Nephropathy TGLs = 4 - 8]
- * Lung
- [Inflammation TGLs = 3 - 6+7]
- * Lymph Node, Mandibular
- [Infiltration Cellular TGLs = 2 - 12]
- * Nose
- * Pancreas
- [Adenoma TGLs = 5 - 9]
- * Pituitary Gland
- * Prostate
- * Salivary Glands
- * Spleen
- * Stomach, Glandular
- * Thymus
- * Thyroid Gland
- Pelvis
- Alveolus
- Nasolacrim Dct
- Acinus
- Pars Distalis
- C Cell
- Hypertrophy
- Hypercellularity
- Calculus Micro Observation Only
- Inflammation
- Nephropathy
- Foreign Body
- Inflammation
- Infiltration Cellular
- Inflammation
- Adenoma
- Hyperplasia
- Inflammation
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Mineral
- Atrophy
- Hyperplasia
- Focal, Minimal
- Marked
- Chronic Active, Mild
- Chronicprogr, Marked
- Granulomatous, Moderate
- Plasma Cell, Marked
- Chronic, Minimal
- Minimal
- Chronic Active, Moderate
- Mild
- Mild
- Minimal
- Mild
- Mild
- Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 31

TRT#: 1

SEX: Male

DAY ON TEST: 618

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402770

ORGAN AND ACCOUNTABLE SITE STATUS

* Urinary Bladder

Inflammation

Chronic, Mild

Zymbal's Gland

Adenoma

[Adenoma TGLs = 1 - 17]

PRIMARY CAUSE OF DEATH

- Zymbal's Gland Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 32

TRT#: 1

SEX: Male

DAY ON TEST: 693

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402771

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Parathyroid Gland
- * Preputial Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea

OBSERVATIONS

- Blood Vessel Intima Hyperplasia Mild
Note: Thymic artery has endocardial (intimal) hyperplasia (B06).
- * Bone Marrow Hypercellularity Minimal
- * Kidney Nephropathy Chronicprogr, Moderate
[Nephropathy TGLs = 3 - 8]
- * Liver Bile Duct Cyst
Bile Duct Hyperplasia Minimal
- * Lung Infiltration Cellular Inflammation Histiocyte, Mild
Acute, Minimal
[Infiltration Cellular TGLs = 2 - 6+7]
- Lymph Node Lumbar Hyperplasia Lymphoid, Moderate
[Hyperplasia TGLs = 4 - 17]
- * Nose Foreign Body
Inflammation Suppurative, Marked
Inflammation Chronic, Minimal
- * Pancreas Acinus Hyperplasia Minimal
Polyarteritis Nodosa Mild
- * Pituitary Gland Pars Distalis Adenoma
- [Adenoma TGLs = 5 - 11]
- * Prostate Inflammation Chronic Active, Moderate
- * Seminal Vesicle Inflammation Chronic Active, Mild
- * Spleen White Pulp Atrophy Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 32

TRT#: 1

SEX: Male

DAY ON TEST: 693

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402771

ORGAN AND ACCOUNTABLE SITE STATUS

Extramedullary Hematopoiesis

Minimal

Pigment

Mild

Testis

Bilateral, Germinal Epith

Degeneration

Minimal

Polyarteritis Nodosa

Mild

* Thymus

Atrophy

Marked

Polyarteritis Nodosa

Mild

Note: Thymus was so small none could be processed to slides, hence diagnosis is based upon gross finding only.

[Atrophy TGLs = 1 - 6]

* Urinary Bladder

Perivascular

Infiltration Cellular

Lymphocyte, Mild

PRIMARY CAUSE OF DEATH

- Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA
 Black Cohosh
 CAS Number: 84776-26-1

Date Report Requested: 11/13/2020
 Time Report Requested: 09:33:45
 First Dose M/F: 07/03/12 / 07/02/12
 Lab: BAT

ANIMAL ID: 33

TRT#: 1

SEX: Male

DAY ON TEST: 692

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402772

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Cortex	* Adrenal Medulla	* Bone	* Bone Marrow
* Epididymis	* Esophagus	* Eye	* Harderian Gland
* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum	* Intestine Small, Duodenum
* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic	* Liver
* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Mammary Gland	* Nose
* Parathyroid Gland	* Preputial Gland	* Prostate	* Salivary Glands
* Seminal Vesicle	* Skin	* Stomach, Forestomach	* Stomach, Glandular
* Trachea	* Urinary Bladder		

OBSERVATIONS

* Brain		Granular Cell Tumor Benign	
* Heart		Cardiomyopathy	Minimal
* Kidney		Nephropathy	Chronicprogr, Marked
	[Nephropathy TGLs = 2 - 8+17]		
* Lung		Infiltration Cellular	Histiocyte, Minimal
* Pancreas	Acinus	Adenoma	
	Acinus	Hyperplasia	Minimal
* Pituitary Gland	Pars Distalis	Adenoma	
	[Adenoma TGLs = 1 - 11]		
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
Testis		Polyarteritis Nodosa	Minimal
* Thymus		Atrophy	Marked
* Thyroid Gland	C Cell	Adenoma	

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

CONTRIBUTORY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 34

TRT#: 1

SEX: Male

DAY ON TEST: 718

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402773

ORGAN AND ACCOUNTABLE SITE STATUS

* Seminal Vesicle		Inflammation	Chronic Active, Mild
* Spleen		Pigment	Mild
* Stomach, Forestomach	Epithelium	Hyperplasia	Mild
		Inflammation	Chronic, Mild
	[Hyperplasia TGLs = 7 - 9]		
Testis	Bilateral, Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Minimal
	[Degeneration TGLs = 1 - 14]		
* Thymus		Atrophy	Moderate
		Ectopic Parathyroid Gland	Moderate
* Thyroid Gland	C Cell	Hyperplasia	Moderate

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 35

TRT#: 1

SEX: Male

DAY ON TEST: 731

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402774

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Salivary Glands
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Pituitary Gland
- * Seminal Vesicle
- Testis
- * Bone Marrow
- * Eye
- * Intestine Small, Duodenum
- * Lung
- * Nose
- * Preputial Gland
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla [Pheochromocytoma Benign TGLs = 1 - 11] Pheochromocytoma Benign
- * Heart Cardiomyopathy Minimal
- * Intestine Large, Colon Lymphoid Tiss Hyperplasia Moderate
- * Kidney Nephropathy Chronicprogr, Mild
- * Liver Clear Cell Focus
- * Spleen Bile Duct Hyperplasia Minimal
- * Thymus Extramedullary Hematopoiesis Mild
- * Thyroid Gland Bilateral, C Cell Pigment Minimal
- Atrophy Minimal
- Adenoma Multiple

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 36

TRT#: 1

SEX: Male

DAY ON TEST: 731

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402775

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Pancreas | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach |
| Testis | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|----------------------|--------------------------------|------------------------------|------------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| * Nose | | Inflammation | Chronic, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| * Skin | | Inflammation | Granulomatous, Marked |
| | [Inflammation TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Stomach, Glandular | | Inflammation | Chronic, Minimal |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Adenoma | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 37

TRT#: 1

SEX: Male

DAY ON TEST: 621

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402776

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lung
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Pancreas
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

MISSING

- * Thymus

OBSERVATIONS

- Blood Vessel
 - Note: The aorta and pulmonary vein are mineralized.
- * Bone Marrow
 - Mineral
 - Mild
- * Eye
 - Cornea
 - Hemorrhage
 - Mild
 - Inflammation
 - Hypercellularity
 - Marked
 - Cardiomyopathy
 - Acute, Moderate
 - Mineral
 - Minimal
- * Heart
 - Cornea
 - Cardiomyopathy
 - Acute, Moderate
- * Kidney
 - Renal Tubule
 - Cyst
 - Nephropathy
 - Chronicprogr, Marked
 - [Cyst TGLs = 2 - 8]
 - [Nephropathy TGLs = 1 - 8]
- * Liver
 - Bile Duct
 - Clear Cell Focus
 - Hyperplasia
 - Minimal
- * Nose
 - Foreign Body
 - Inflammation
 - Suppurative, Minimal
- * Parathyroid Gland
 - Hyperplasia
 - Diffuse, Marked
- * Pituitary Gland
 - Pars Distalis
 - Hyperplasia
 - Moderate
- * Spleen
 - White Pulp
 - Atrophy
 - Moderate
 - Extramedullary Hematopoiesis
 - Moderate
 - Inflammation
 - Chronic, Minimal
- * Stomach, Forestomach
 - Inflammation
 - Chronic, Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 37

TRT#: 1

SEX: Male

DAY ON TEST: 621

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402776

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Glandular
Testis

Germinal Epith

Mineral
Degeneration
Polyarteritis Nodosa

Moderate
Mild
Minimal

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 38

TRT#: 1

SEX: Male

DAY ON TEST: 481

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402777

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|---------------------------|---------------|-----------------------|------------------------|
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| [Adenoma TGLs = 1 - 11] | | | |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 39

TRT#: 1

SEX: Male

DAY ON TEST: 678

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402778

ORGAN AND ACCOUNTABLE SITE STATUS

Leukemia Granulocytic

Degeneration

Minimal

Atrophy

Moderate

Adenoma

Hyperplasia

Moderate

Inflammation

Chronic, Minimal

Testis

Germinal Epith

* Thymus

* Thyroid Gland

C Cell

C Cell

[Adenoma TGLs = 2 - 11]

* Urinary Bladder

PRIMARY CAUSE OF DEATH - Kidney Pelvis Inflammation

CONTRIBUTORY CAUSE OF DEATH - Brain Hemorrhage

Animal Note: Primary cause of death was likely kidney infection with death due to sepsis and complications, including meningitis and brain hemorrhage.

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 40

TRT#: 1

SEX: Male

DAY ON TEST: 155

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402779

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thymus | * Thyroid Gland | * Trachea |
| * Urinary Bladder | | | |

OBSERVATIONS

- | | | | |
|---------------|---------------------------|------------------------------|--------------------|
| * Bone Marrow | | Hypercellularity | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Skin | | Sarcoma | |
| | [Sarcoma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |

PRIMARY CAUSE OF DEATH - Skin Sarcoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 42

TRT#: 1

SEX: Male

DAY ON TEST: 354

DOSE: 0 mg/kg male

DISP: Natural Death

HISTO: 1402781

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver | * Lung |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Parathyroid Gland | * Pituitary Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|----------|--------------|------------------------------|--------------------|
| * Kidney | Pelvis | Dilation | Mild |
| | Renal Tubule | Dilation | Mild |
| | | Nephropathy | Chronicprogr, Mild |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH

- Kidney Pelvis Dilation

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 43

TRT#: 1

SEX: Male

DAY ON TEST: 599

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402782

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Seminal Vesicle
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lung
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Prostate
- * Stomach, Forestomach

OBSERVATIONS

- * Adrenal Cortex
 - Blood Vessel
 - Note: The aorta and cardiac arteries are mineralized.
 - * Bone Marrow
 - * Heart
 - * Kidney
 - [Nephropathy TGLs = 1 - 8]
 - * Liver
 - Bile Duct
 - * Nose
 - * Pancreas
 - * Parathyroid Gland
 - * Pituitary Gland
 - * Spleen
 - * Stomach, Glandular
 - Testis
 - Bilateral, Germinal Epith
 - * Thymus
- | | | |
|--|------------------------------|----------------------|
| | Hyperplasia | Focal, Minimal |
| | Mineral | Minimal |
| | Hypercellularity | Minimal |
| | Cardiomyopathy | Mild |
| | Nephropathy | Chronicprogr, Marked |
| | Eosinophilic Focus | |
| | Hyperplasia | Minimal |
| | Mixed Cell Focus | |
| | Inflammation | Chronic, Minimal |
| | Hyperplasia | Marked |
| | Hyperplasia | Diffuse, Marked |
| | Hyperplasia | Mild |
| | Extramedullary Hematopoiesis | Mild |
| | Pigment | Minimal |
| | Mineral | Mild |
| | Degeneration | Mild |
| | Polyarteritis Nodosa | Mild |
| | Atrophy | Marked |
| | Ectopic Parathyroid Gland | Moderate |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 44

TRT#: 1

SEX: Male

DAY ON TEST: 643

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402783

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Prostate
- * Stomach, Forestomach
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
 - Degeneration
 - Necrosis
 - Blood Vessel
 - Mineral
 - Note: The aorta and cardiac and renal arteries are mineralized.
- * Bone Marrow
 - Hemorrhage
 - Hypercellularity
- * Heart
 - Cardiomyopathy
- * Kidney
 - Nephropathy

[Nephropathy TGLs = 1 - 8]
- * Liver
 - Eosinophilic Focus
- * Lung
 - Infiltration Cellular
- * Parathyroid Gland
 - Hyperplasia
- * Spleen
 - White Pulp
 - Atrophy
 - Extramedullary Hematopoiesis
- * Stomach, Glandular
 - Mineral
- Testis
 - Polyarteritis Nodosa
- * Thymus
 - Atrophy

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 45

TRT#: 1

SEX: Male

DAY ON TEST: 623

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402784

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Preputial Gland
- * Urinary Bladder
- Blood Vessel
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Stomach, Glandular
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Mammary Gland
- * Thyroid Gland
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Parathyroid Gland
- * Trachea

MISSING

- * Thymus

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Epididymis
- * Heart
- * Kidney
- [Nephropathy TGLs = 3 - 8]
- * Liver
- Note: The histiocytic sarcoma in the liver (only) contains numerous multinucleate cells.
- [Histiocytic Sarcoma TGLs = 1,5 - 12+17+18]
- * Lung
- Lymph Node
- [Histiocytic Sarcoma TGLs = 4 - 20]
- [Histiocytic Sarcoma TGLs = 2 - 19]
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Nose
- * Pancreas
- * Pituitary Gland
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Renal Tubule
- Accumulation, Hyaline Droplet
- Histiocytic Sarcoma
- Nephropathy
- Histiocytic Sarcoma
- Mediastinal
- Renal
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Histiocytic Sarcoma
- Nasolacrim Dct
- Pars Distalis
- Metastatic (Liver)
- Marked
- Chronicprogr, Moderate
- Suppurative, Moderate
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 45

TRT#: 1

SEX: Male

DAY ON TEST: 623

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402784

ORGAN AND ACCOUNTABLE SITE STATUS

* Prostate		Histiocytic Sarcoma	
* Salivary Glands		Histiocytic Sarcoma	
* Seminal Vesicle		Histiocytic Sarcoma	
* Skin		Histiocytic Sarcoma	
	[Histiocytic Sarcoma TGLs = 6 - 21]		
* Spleen	White Pulp	Atrophy	Mild
		Extramedullary Hematopoiesis	Moderate
* Stomach, Forestomach		Inflammation	Granulomatous, Minimal
Testis	Bilateral, Germinal Epith	Degeneration	Marked
		Histiocytic Sarcoma	

PRIMARY CAUSE OF DEATH - Liver Histiocytic Sarcoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 47

TRT#: 1

SEX: Male

DAY ON TEST: 731

DOSE: 0 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402786

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Epididymis | * Esophagus | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular |
| * Trachea | * Urinary Bladder | | |

MISSING

- * Thymus

OBSERVATIONS

- | | | | |
|-------------------|-----------------------------------|------------------------------|----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Eye | Cornea | Inflammation | Chronic, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| * Skin | | Keratoacanthoma | |
| | [Keratoacanthoma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Moderate |
| | | Mineral | Mild |
| | [Degeneration TGLs = 3 - 18] | | |
| | [Mineral TGLs = 2 - 18] | | |
| * Thyroid Gland | C Cell | Adenoma | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 50

TRT#: 1

SEX: Male

DAY ON TEST: 676

DOSE: 0 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402789

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| Blood Vessel | * Bone | * Brain | * Epididymis |
| * Esophagus | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Mammary Gland | * Nose |
| * Pancreas | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|--------------------------|--------------------------------|-----------------------|----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hemorrhage | Marked |
| * Eye | Cornea | Inflammation | Acute, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| | Atrium | Thrombus | |
| | [Thrombus TGLs = 4 - 10] | | |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 2- 8] | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| * Lymph Node, Mesenteric | | Atrophy | Moderate |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Moderate |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Pigment | Mild |
| * Stomach, Forestomach | Epithelium | Hyperplasia | Basal Cell, Moderate |
| | [Hyperplasia TGLs = 3 - 9] | | |
| * Stomach, Glandular | | Infiltration Cellular | Lymphocyte, Minimal |
| Testis | Germinal Epith | Degeneration | Mild |
| | [Degeneration TGLs = 1 - 14] | | |
| * Thymus | | Atrophy | Marked |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 51

TRT#: 3

SEX: Male

DAY ON TEST: 488

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402790

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Liver
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Thyroid Gland
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel Mineral Mild
Note: Aorta and cardiac and pulmonary arteries are mineralized.
- * Bone Marrow Hemorrhage Minimal
- * Heart Cardiomyopathy Minimal
- * Kidney Mineral Mild
Nephropathy Chronicprogr, Marked
- [Nephropathy TGLs = 1 - 8]
- * Lung Infiltration Cellular Histiocyte, Mild
Inflammation Granulomatous, Minimal
Mineral Minimal
- * Nose Nasolacrim Dct Inflammation Chronic, Minimal
Inflammation Suppurative, Marked
- * Pancreas Acinus Hyperplasia Marked
- * Parathyroid Gland Hyperplasia Diffuse, Mild
- * Spleen White Pulp Atrophy Marked
Pigment Mild
- * Stomach, Glandular Mineral Moderate
Testis Polyarteritis Nodosa Minimal
- * Thymus Atrophy Marked
- * Trachea Mineral Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 52

TRT#: 3

SEX: Male

DAY ON TEST: 637

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402791

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Forestomach	Epithelium	Hyperplasia Inflammation Ulcer	Moderate Chronic Active, Mild Mild
* Stomach, Glandular		Mineral	Mild
Testis	Bilateral, Germinal Epith	Degeneration	Moderate
	[Degeneration TGLs = 1 - 14]		
* Thymus		Atrophy	Marked
Tooth	Dentine	Degeneration	Moderate
	Pulp	Inflammation	Chronic, Mild

Note: Osteoblasts and osteoid lining the pulp cavities of both incisors and starting in one molar.

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 53

TRT#: 3

SEX: Male

DAY ON TEST: 666

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402792

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Prostate
- * Stomach, Glandular
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lung
- * Pancreas
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex [Hyperplasia TGLs = 1 - 11] Hyperplasia Focal, Marked
- * Bone Note: Block 01 with femur is missing; bone and marrow evaluated based upon nasal section.
- * Bone Marrow Hypercellularity Mild
- * Eye Cornea Inflammation Chronic Active, Mild
- * Heart Perivascular Cardiomyopathy Mild
- * Kidney Infiltration Cellular Lymphocyte, Mild
- * Liver Nephropathy Chronicprogr, Moderate
- * Liver Polyarteritis Nodosa Mild
- * Liver Basophilic Focus
- * Nose Nasolacrim Dct Inflammation Chronic, Mild
- * Parathyroid Gland Hyperplasia Diffuse, Mild
- * Skin Hemangioma
- * Spleen Extramedullary Hematopoiesis Mild
- * Spleen Pigment Minimal
- * Testis Germinal Epith Degeneration Minimal
- * Testis Polyarteritis Nodosa Mild
- * Thymus Atrophy Minimal

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 54

TRT#: 3

SEX: Male

DAY ON TEST: 724

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402793

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|------------------------|------------------------------|------------------------------|----------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 2 - 8] | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 1 - 11] | | |
| * Spleen | White Pulp | Atrophy | Mild |
| | | Extramedullary Hematopoiesis | Minimal |
| | | Pigment | Mild |
| * Stomach, Forestomach | Epithelium | Hyperplasia | Minimal |
| | | Inflammation | Chronic Active, Mild |
| Testis | | Polyarteritis Nodosa | Minimal |
| * Thymus | | Atrophy | Marked |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 55

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402794

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Epididymis | * Esophagus | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|------------------------|---------------|-----------------------------------------------------|-------------------------|
| * Adrenal Medulla | Bilateral | Hyperplasia
Pheochromocytoma Benign | Focal, Moderate |
| * Bone Marrow | | Hypercellularity | Minimal |
| * Eye | Cornea | Inflammation | Chronic, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | Bile Duct | Clear Cell Focus
Hyperplasia
Mixed Cell Focus | Minimal |
| * Nose | | Inflammation | Chronic Active, Minimal |
| * Pancreas | Acinus | Atrophy | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis
Pigment | Mild
Minimal |
| * Stomach, Forestomach | | Infiltration Cellular | Lipocyte, Minimal |
| * Stomach, Glandular | | Infiltration Cellular | Lipocyte, Mild |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 56

TRT#: 3

SEX: Male

DAY ON TEST: 644

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402795

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Seminal Vesicle
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Stomach, Forestomach
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Prostate
- * Thyroid Gland
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta and cardiac, mesenteric, prostatic, and pulmonary arteries are mineralized.
- * Bone
* Bone Marrow
- * Heart
- * Intestine Large, Cecum
- * Kidney
[Cyst TGLs = 3 - 8]
[Nephropathy TGLs = 2 - 8]
- * Lung
- * Nose
- * Parathyroid Gland
- * Preputial Gland
- Hyperplasia
- Mineral
- Cyst
- Hemorrhage
- Hypercellularity
- Myelofibrosis
- Cardiomyopathy
- Mineral
- Inflammation
- Mineral
- Polyarteritis Nodosa
- Ulcer
- Cyst
- Nephropathy
- Mineral
- Hyperplasia
- Inflammation
- Hyperplasia
- Papilloma
- Focal, Mild
- Moderate
- Minimal
- Minimal
- Minimal
- Mild
- Minimal
- Chronic Active, Minimal
- Minimal
- Mild
- Mild
- Chronicprogr, Marked
- Mild
- Minimal
- Suppurative, Minimal
- Diffuse, Mild
- Squamous

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 56

TRT#: 3

SEX: Male

DAY ON TEST: 644

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402795

ORGAN AND ACCOUNTABLE SITE STATUS

* Skin		Keratoacanthoma	
	[Keratoacanthoma TGLs = 1 - 17]		
* Spleen	White Pulp	Atrophy	Mild
		Extramedullary Hematopoiesis	Mild
		Pigment	Mild
* Stomach, Glandular		Mineral	Marked
Testis	Bilateral, Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Mild
* Thymus		Atrophy	Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 57

TRT#: 3

SEX: Male

DAY ON TEST: 680

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402796

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Trachea
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
 - [Pheochromocytoma Benign TGLs = 3 - 11]
 - Blood Vessel
 - Note: The aorta and gastric arteries are mineralized.
 - * Bone Marrow
 - * Heart
 - * Kidney
 - [Nephropathy TGLs = 1 - 8]
 - * Liver
 - [Cholangioma TGLs = 2 - 17]
 - * Lung
 - Alveolar Epith
 - * Parathyroid Gland
 - * Pituitary Gland
 - * Spleen
 - Pars Distalis
 - White Pulp
 - * Stomach, Glandular
 - Testis
 - Bilateral, Germinal Epith
 - * Thymus
- Pheochromocytoma Benign
- Mineral
- Hemorrhage
- Hypercellularity
- Cardiomyopathy
- Nephropathy
- Cholangioma
- Hyperplasia
- Hyperplasia
- Hyperplasia
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Mineral
- Degeneration
- Polyarteritis Nodosa
- Atrophy
- Ectopic Parathyroid Gland
- Minimal
- Moderate
- Mild
- Mild
- Chronicprogr, Marked
- Minimal
- Diffuse, Mild
- Mild
- Moderate
- Minimal
- Mild
- Mild
- Marked
- Mild
- Marked
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 57

TRT#: 3

SEX: Male

DAY ON TEST: 680

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402796

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 58

TRT#: 3

SEX: Male

DAY ON TEST: 596

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402797

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | Testis | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|---------------------|------------------------------|------------------------------|----------------------|
| * Bone Marrow | | Hemorrhage | Moderate |
| | | Hypercellularity | Mild |
| * Heart | | Cardiomyopathy | Minimal |
| | Atrium | Thrombus | |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 2 - 8] | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | Bronchiole | Fibrosis | Mild |
| | | Hemorrhage | Marked |
| | | Infiltration Cellular | Histiocyte, Moderate |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Moderate |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Extramedullary Hematopoiesis | Minimal |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Marked |
| * Thyroid Gland | C Cell | Adenoma | |

PRIMARY CAUSE OF DEATH - Heart Atrium Thrombus

CONTRIBUTORY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 59

TRT#: 3

SEX: Male

DAY ON TEST: 617

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402798

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Small, Duodenum
- * Parathyroid Gland
- * Stomach, Forestomach
- Blood Vessel
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- Peripheral Nerve
- Testis
- * Bone
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Preputial Gland
- * Trachea
- * Epididymis
- * Intestine Large, Rectum
- * Mammary Gland
- * Salivary Glands

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Small, Jejunum
- * Kidney
- * Liver
- [Leukemia Mononuclear TGLs = 3 - 12+18]
- * Lung
- Lymph Node
- [Leukemia Mononuclear TGLs = 5 - 20]
- [Leukemia Mononuclear TGLs = 4 - 19]
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- [Leukemia Mononuclear TGLs = 7 - 9]
- Nerve Trigeminal
- Leukemia Mononuclear
- Leukemia Mononuclear
- Hemorrhage
- Leukemia Mononuclear
- Metaplasia
- Leukemia Mononuclear
- Leukemia Mononuclear
- Leukemia Mononuclear
- Nephropathy
- Hyperplasia
- Leukemia Mononuclear
- Necrosis
- Leukemia Mononuclear
- Leukemia Mononuclear
- Leukemia Mononuclear
- Leukemia Mononuclear
- Minimal
- Minimal
- Chronicprogr, Moderate
- Minimal
- Mild

Note: Mononuclear cell leukemia surround trigeminal nerve in nasal section, but could not be coded in TDMSE.

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 59

TRT#: 3

SEX: Male

DAY ON TEST: 617

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402798

ORGAN AND ACCOUNTABLE SITE STATUS

* Nose		Leukemia Mononuclear	
* Pancreas	Acinus	Hyperplasia	Minimal
* Pituitary Gland	Pars Distalis	Hyperplasia	Minimal
* Prostate		Leukemia Mononuclear	
* Seminal Vesicle		Leukemia Mononuclear	
* Skin		Leukemia Mononuclear	
Spinal Cord	Axon	Degeneration	Mild
		Hemorrhage	Moderate
* Spleen		Hemorrhage	Moderate
		Leukemia Mononuclear	
	[Leukemia Mononuclear TGLs = 1 - 17]		
* Stomach, Glandular		Leukemia Mononuclear	
* Thymus		Leukemia Mononuclear	
	[Leukemia Mononuclear TGLs = 6 - 6]		
* Thyroid Gland	C Cell	Adenoma	
	C Cell	Hyperplasia	Mild
* Urinary Bladder		Leukemia Mononuclear	

PRIMARY CAUSE OF DEATH - Spleen Leukemia Mononuclear

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 60

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402799

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Eye
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea
- * Epididymis
- * Intestine Small, Duodenum
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Esophagus
- * Islets, Pancreatic
- * Prostate
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Cortex
 - Hyperplasia
 - Thrombus
- * Adrenal Medulla
 - Pheochromocytoma Benign
- * Bone Marrow
 - Hypercellularity
- * Heart
 - Cardiomyopathy
- * Intestine Large, Cecum
 - Polyarteritis Nodosa
- * Intestine Large, Colon
 - Polyarteritis Nodosa
- * Intestine Large, Rectum
 - Parasite Metazoan
- * Intestine Small, Ileum
 - Polyarteritis Nodosa
- * Intestine Small, Jejunum
 - Peyers Patch
 - Hyperplasia
- * Kidney
 - Nephropathy

[Nephropathy TGLs = 2 - 8]
- * Liver
 - Clear Cell Focus
- * Lung
 - Infiltration Cellular
 - Inflammation

[Infiltration Cellular TGLs = 3,4 - 6+7]
- * Lymph Node, Mandibular
 - Infiltration Cellular

[Infiltration Cellular TGLs = 1 - 12]
- * Lymph Node, Mesenteric
 - Hyperplasia
- * Nose
 - Nasolacrimal Duct
 - Inflammation
- * Pancreas
 - Adenoma
 - Hyperplasia
 - Polyarteritis Nodosa
- * Parathyroid Gland
 - Hyperplasia

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 60

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402799

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
* Stomach, Forestomach		Polyarteritis Nodosa	Mild
Testis	Bilateral, Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Moderate
* Thymus		Atrophy	Minimal

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 61

TRT#: 3

SEX: Male

DAY ON TEST: 630

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402800

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Liver
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Pancreas
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Thymus
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel Mineral Mild
Note: The aorta and mediastinal arteries are mineralized.
- * Bone Marrow Hemorrhage Moderate
Hypercellularity Mild
- * Eye Cornea Inflammation Acute, Minimal
- * Heart Cardiomyopathy Moderate
- * Intestine Large, Cecum Erosion Minimal
Mineral Minimal
- * Kidney Nephropathy Chronicprogr, Marked
[Nephropathy TGLs = 3 - 8]
- * Lung Inflammation Acute, Minimal
Mineral Minimal
- * Nose Inflammation Chronic Active, Moderate
Nasopharyn Dct Inflammation Chronic Active, Moderate
Nasopharyn Dct Squamous Metaplasia Moderate
- * Parathyroid Gland Hyperplasia Diffuse, Marked
- * Spleen White Pulp Atrophy Moderate
Pigment Mild
- * Stomach, Glandular Mineral Mild
Testis Bilateral, Germinal Epith Degeneration Marked
[Degeneration TGLs = 1 - 14]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 61

TRT#: 3

SEX: Male

DAY ON TEST: 630

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402800

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 62

TRT#: 3

SEX: Male

DAY ON TEST: 669

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402801

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Stomach, Glandular

MISSING

- * Thymus

AUTO PRECLUDES DIAG.

- * Intestine Large, Cecum

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Heart
- * Kidney
- Renal Tubule
- Pelvis
- [Cyst TGLs = 2,3 - 8+17]
- [Nephropathy TGLs = 1 - 8]
- * Liver
- * Lung
- Alveolus
- Mesentery
- [Polyarteritis Nodosa TGLs = 4-17]
- * Parathyroid Gland
- Necrosis
- Hemorrhage
- Hypercellularity
- Cardiomyopathy
- Calculus Micro Observation Only
- Cyst
- Inflammation
- Nephropathy
- Basophilic Focus
- Fibrosis
- Infiltration Cellular
- Inflammation
- Polyarteritis Nodosa
- Hyperplasia
- Minimal
- Moderate
- Moderate
- Mild
- Chronic Active, Moderate
- Chronicprogr, Marked
- Minimal
- Histiocyte, Moderate
- Acute, Mild
- Moderate
- Diffuse, Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 62

TRT#: 3

SEX: Male

DAY ON TEST: 669

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402801

ORGAN AND ACCOUNTABLE SITE STATUS

* Prostate		Inflammation	Chronic Active, Minimal
* Seminal Vesicle		Inflammation	Chronic Active, Mild
* Spleen	White Pulp	Atrophy	Marked
		Extramedullary Hematopoiesis	Minimal
		Pigment	Minimal
Testis	Bilateral, Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 63

TRT#: 3

SEX: Male

DAY ON TEST: 611

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402802

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Prostate
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Trachea
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta and cardiac, gastric, pulmonary, renal, salivary, seminal vesicular, and dermal arteries are mineralized.
- * Bone Marrow
- * Heart
- * Intestine Large, Cecum
- * Kidney
[Nephropathy TGLs = 1 - 8]
- * Lung
- * Parathyroid Gland
- * Spleen
- * Stomach, Forestomach
[Cyst TGLs = 2 - 17]
- * Stomach, Glandular
- Testis
- Hyperplasia
- Mineral
- Hemorrhage
- Cardiomyopathy
- Mineral
- Erosion
- Mineral
- Polyarteritis Nodosa
- Nephropathy
- Mineral
- Hyperplasia
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Cyst
- Inflammation
- Mineral
- Degeneration
- Polyarteritis Nodosa
- Focal, Mild
- Marked
- Moderate
- Moderate
- Minimal
- Mild
- Moderate
- Minimal
- Chronicprogr, Marked
- Mild
- Diffuse, Moderate
- Moderate
- Minimal
- Minimal
- Squamous
- Acute, Mild
- Mild
- Mild
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 63

TRT#: 3

SEX: Male

DAY ON TEST: 611

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402802

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Marked

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 64

TRT#: 3

SEX: Male

DAY ON TEST: 713

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402803

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Cortex	* Adrenal Medulla	* Bone	* Bone Marrow
* Brain	* Epididymis	* Esophagus	* Harderian Gland
* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Small, Duodenum	* Intestine Small, Ileum
* Intestine Small, Jejunum	* Islets, Pancreatic	* Lung	* Lymph Node, Mandibular
* Lymph Node, Mesenteric	* Mammary Gland	* Nose	* Parathyroid Gland
* Preputial Gland	* Seminal Vesicle	* Stomach, Forestomach	* Stomach, Glandular
* Trachea	* Urinary Bladder		

OBSERVATIONS

Blood Vessel		Aneurysm	Moderate
[Aneurysm TGLs = 2 - 18]			
* Eye	Retina	Dysplasia	Mild
	Retina	Hyperplasia	Mild
* Heart		Cardiomyopathy	Mild
* Intestine Large, Rectum		Polyarteritis Nodosa	Mild
* Kidney		Nephropathy	Chronicprogr, Moderate
* Liver		Clear Cell Focus	
* Pancreas		Polyarteritis Nodosa	Mild
* Pituitary Gland	Pars Distalis	Hyperplasia	Mild
* Prostate		Inflammation	Chronic Active, Mild
* Salivary Glands		Atrophy	Mild
* Skin		Cyst Epithelial Inclusion	
		Inflammation	Chronic Active, Marked
		Ulcer	Marked
[Cyst Epithelial Inclusion TGLs = 1 - 17]			
[Ulcer TGLs = 3 - 19]			
* Spleen		Extramedullary Hematopoiesis	Mild
Testis		Polyarteritis Nodosa	Moderate
* Thymus		Atrophy	Minimal
* Thyroid Gland	C Cell	Adenoma	Multiple
	C Cell	Hyperplasia	Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 64

TRT#: 3

SEX: Male

DAY ON TEST: 713

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402803

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Skin Ulcer

CONTRIBUTORY CAUSE OF DEATH

- Blood Vessel Aneurysm

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 65

TRT#: 3

SEX: Male

DAY ON TEST: 711

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402804

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen		Extramedullary Hematopoiesis	Moderate
		Pigment	Minimal
* Stomach, Forestomach	Epithelium	Hyperplasia	Basal Cell, Mild
Note: Recut has lesion in forestomach. [Hyperplasia TGLs = 2 - 17]			
* Stomach, Glandular		Mineral	Marked
Testis	Bilateral, Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Mild
* Thymus		Atrophy	Moderate
Tooth	Dentine	Degeneration	Mild
* Urinary Bladder		Inflammation	Chronic Active, Mild
		Polyarteritis Nodosa	Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

CONTRIBUTORY CAUSE OF DEATH - Kidney Hemorrhage

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 66

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402805

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Pancreas |
| * Parathyroid Gland | * Preputial Gland | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

MISSING

- * Mammary Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|------------------------|
| * Bone Marrow | | Hypercellularity | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Nose | | Inflammation | Suppurative, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Prostate | | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 67

TRT#: 3

SEX: Male

DAY ON TEST: 553

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402806

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Urinary Bladder
- * Adrenal Medulla
- * Esophagus
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Skin
- * Bone
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Thyroid Gland
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Prostate
- * Trachea

OBSERVATIONS

Blood Vessel	Aorta	Dilation	Moderate
		Mineral	Moderate
Note: The aorta and cardiac, pancreatic, renal, and splenic arteries are mineralized.			
	[Dilation TGLs = 2 - 17]		
* Bone Marrow		Hemorrhage	Minimal
		Myelofibrosis	Minimal
* Eye	Cornea	Inflammation	Acute, Mild
* Heart		Cardiomyopathy	Moderate
		Mineral	Moderate
* Intestine Large, Colon		Mineral	Moderate
* Kidney		Nephropathy	Chronicprogr, Marked
	[Nephropathy TGLs = 4 - 8]		
* Lung		Mineral	Marked
* Mammary Gland		Mineral	Mild
* Parathyroid Gland		Hyperplasia	Diffuse, Marked
* Pituitary Gland	Pars Distalis	Adenoma	
	[Adenoma TGLs = 1 - 11]		
* Seminal Vesicle		Mineral	Mild
* Spleen	White Pulp	Atrophy	Marked
		Pigment	Mild
* Stomach, Forestomach		Mineral	Moderate
* Stomach, Glandular		Mineral	Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 67

TRT#: 3

SEX: Male

DAY ON TEST: 553

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402806

ORGAN AND ACCOUNTABLE SITE STATUS

Testis	Bilateral, Germinal Epith	Degeneration	Moderate
[Degeneration TGLs = 3 - 14]			
* Thymus		Atrophy	Marked
Tooth		Metaplasia	Osseous, Minimal

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 68

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402807

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Bone Marrow |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | Testis | * Thymus | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|------------------------------|---------------|------------------------------|----------------------|
| * Intestine Large, Colon | Lymphoid Tiss | Hyperplasia | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| [Nephropathy TGLs = 1 - 8] | | | |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Pancreas | | Polyarteritis Nodosa | Minimal |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 69

TRT#: 3

SEX: Male

DAY ON TEST: 618

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402808

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Preputial Gland
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Prostate
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Trachea
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta and cardiac arteries are mineralized.
 - * Bone Marrow
 - * Heart
 - * Intestine Large, Cecum
 - * Kidney
[Nephropathy TGLs = 1 - 8]
 - * Lung
 - * Parathyroid Gland
 - * Skin
[Keratoacanthoma TGLs = 2 - 17]
 - * Spleen
White Pulp
 - * Stomach, Glandular
Testis
Bilateral, Germinal Epith
 - * Thymus
- | | | |
|--|----------------------|----------------------|
| | Hyperplasia | Focal, Mild |
| | Mineral | Mild |
| | Hemorrhage | Minimal |
| | Cardiomyopathy | Moderate |
| | Mineral | Minimal |
| | Erosion | Mild |
| | Inflammation | Chronic, Mild |
| | Mineral | Mild |
| | Polyarteritis Nodosa | Minimal |
| | Nephropathy | Chronicprogr, Marked |
| | Mineral | Mild |
| | Hyperplasia | Diffuse, Moderate |
| | Keratoacanthoma | |
| | Atrophy | Moderate |
| | Pigment | Mild |
| | Mineral | Mild |
| | Degeneration | Minimal |
| | Polyarteritis Nodosa | Mild |
| | Atrophy | Marked |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 69

TRT#: 3

SEX: Male

DAY ON TEST: 618

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402808

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 70

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402809

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Parathyroid Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- * Bone Marrow
 - * Eye
 - * Heart
 - * Intestine Large, Colon
 - * Kidney
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 2 - 7]
 - Mesentery
 - [Pigment TGLs = 1 - 17]
 - * Pancreas
 - * Pituitary Gland
 - * Spleen
 - Testis
 - * Thymus
 - * Thyroid Gland
- | | | | |
|--|----------------|------------------------------|-------------------------|
| | | Hypercellularity | Mild |
| | Cornea | Inflammation | Chronic Active, Minimal |
| | | Cardiomyopathy | Mild |
| | | Parasite Metazoan | |
| | | Nephropathy | Chronicprogr, Moderate |
| | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| | | Infiltration Cellular | Histiocyte, Minimal |
| | | Pigment | Mild |
| | Acinus | Atrophy | Minimal |
| | Pars Distalis | Hyperplasia | Moderate |
| | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| | Germinal Epith | Degeneration | Minimal |
| | Interstit Cell | Hyperplasia | Mild |
| | | Atrophy | Mild |
| | C Cell | Adenoma | |
| | C Cell | Hyperplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 71

TRT#: 3

SEX: Male

DAY ON TEST: 274

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402810

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|---------------|--|------------------------------|---------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Extramedullary Hematopoiesis | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Skin | | Sarcoma | |

Note: Sarcoma may be arising in an area of abscessation; associated inflammation is marked.

[Sarcoma TGLs = 1 - 17]

- | | | | |
|----------|------------|------------------------------|----------|
| * Spleen | White Pulp | Atrophy | Mild |
| | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH - Skin Sarcoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 72

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402811

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Trachea
- * Bone
- * Harderian Gland
- * Islets, Pancreatic
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Intestine Small, Duodenum
- * Lung
- * Nose
- * Salivary Glands
- * Stomach, Glandular
- * Epididymis
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Eye
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Kidney
- [Nephropathy TGLs = 1 - 8+17]
- * Liver
- * Pancreas
- * Pituitary Gland
- * Spleen
- Testis
- * Thymus
- Cornea
- Pars Distalis
- Germinal Epith
- Hyperplasia
- Hypercellularity
- Inflammation
- Cardiomyopathy
- Polyarteritis Nodosa
- Polyarteritis Nodosa
- Polyarteritis Nodosa
- Nephropathy
- Eosinophilic Focus
- Polyarteritis Nodosa
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Degeneration
- Polyarteritis Nodosa
- Atrophy
- Focal, Minimal
- Minimal
- Chronic Active, Moderate
- Mild
- Moderate
- Minimal
- Mild
- Chronicprogr, Marked
- Mild
- Minimal
- Minimal
- Minimal
- Minimal
- Mild
- Minimal
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 73

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402812

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Salivary Glands
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Mammary Gland
- * Preputial Gland
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Heart
- * Islets, Pancreatic
- * Kidney
- * Liver
- [Inflammation TGLs = 2,3 - 12]
- * Lung
- * Pituitary Gland
- * Prostate
- * Skin
- [Keratoacanthoma TGLs = 1 - 17]
- * Spleen
- Testis
- * Thymus
- * Thyroid Gland
- Perivascular
- Pars Distalis
- C Cell
- Hyperplasia
- Hypercellularity
- Cardiomyopathy
- Adenoma
- Nephropathy
- Clear Cell Focus
- Inflammation
- Inflammation
- Hyperplasia
- Inflammation
- Keratoacanthoma
- Extramedullary Hematopoiesis
- Pigment
- Polyarteritis Nodosa
- Atrophy
- Hyperplasia
- Focal, Minimal
- Minimal
- Mild
- Mixed Cell
- Chronicprogr, Marked
- Granulomatous, Marked
- Chronic, Mild
- Mild
- Chronic Active, Minimal
- Mild
- Minimal
- Mild
- Moderate
- Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 74

TRT#: 3

SEX: Male

DAY ON TEST: 560

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402813

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Pituitary Gland
- * Seminal Vesicle
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Preputial Gland
- * Skin
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Prostate
- * Stomach, Forestomach
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
 - Necrosis Minimal
 - Thrombus
- Blood Vessel
 - Aorta
 - Dilation Moderate
 - Mineral Mild
- Note: The aorta is mineralized.
[Dilation TGLs = 3 - 10]
- * Bone Marrow
 - Hemorrhage Mild
 - Hypercellularity Mild
- * Heart
 - Cardiomyopathy Mild
- * Intestine Large, Cecum
 - Erosion Minimal
- * Kidney
 - Renal Tubule
 - Cyst
 - Nephropathy Chronicprogr, Marked
- [Cyst TGLs = 2 - 17]
[Nephropathy TGLs = 1 - 8+17]
- * Lung
 - Inflammation Granulomatous, Minimal
 - Mineral Minimal
 - Squamous Metaplasia Mild
- * Spleen
 - White Pulp
 - Atrophy Moderate
 - Pigment Minimal
- * Stomach, Glandular
 - Mineral Moderate
- Testis
 - Bilateral, Germinal Epith
 - Degeneration Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 74

TRT#: 3

SEX: Male

DAY ON TEST: 560

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402813

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

* Thyroid Gland

C Cell

Polyarteritis Nodosa

Atrophy

Adenoma

Minimal

Marked

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 75

TRT#: 3

SEX: Male

DAY ON TEST: 475

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402814

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Bone Marrow |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|----------|----------------|------------------------------|----------------------|
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Liver | | Clear Cell Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Nose | | Inflammation | Suppurative, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Mild |
| | | Polyarteritis Nodosa | Mild |
| * Thymus | | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 76

TRT#: 3

SEX: Male

DAY ON TEST: 637

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402815

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|------------------------------------------|---------------------------|------------------------------|----------------------|
| * Bone Marrow | | Hemorrhage | Mild |
| | | Hypercellularity | Mild |
| * Epididymis | | Atrophy | Mild |
| [Atrophy TGLs = 3 - 14] | | | |
| * Heart | | Cardiomyopathy | Moderate |
| | | Mineral | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| [Nephropathy TGLs = 1 - 8] | | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| [Infiltration Cellular TGLs = 5 - 6+7] | | | |
| Lymph Node | Mediastinal | Hemorrhage | Mild |
| [Hemorrhage TGLs = 4 - 17] | | | |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Marked |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Stomach, Glandular | | Mineral | Mild |
| Testis | Bilateral, Germinal Epith | Degeneration | Minimal |
| [Degeneration TGLs = 2 - 14] | | | |
| * Thymus | | Atrophy | Marked |
| * Thyroid Gland | C Cell | Adenoma | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 76

TRT#: 3

SEX: Male

DAY ON TEST: 637

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402815

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 77

TRT#: 3

SEX: Male

DAY ON TEST: 433

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402816

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | |
|-----------------------------------|------------------------------|------------------------|
| * Epididymis | Granuloma Sperm | Marked |
| [Granuloma Sperm TGLs = 1 - 17] | | |
| * Heart | Cardiomyopathy | Minimal |
| * Kidney | Nephropathy | Chronicprogr, Moderate |
| * Lung | Infiltration Cellular | Histiocyte, Minimal |
| * Salivary Glands | Inflammation | Chronic Active, Mild |
| * Spleen | Extramedullary Hematopoiesis | Mild |
| | Pigment | Minimal |
| * Thymus | Atrophy | Minimal |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 78

TRT#: 3

SEX: Male

DAY ON TEST: 575

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402817

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Trachea
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Cortex
 - Blood Vessel
 - Note: The aorta and cardiac, cecal, dermal, pancreatic, pulmonary, renal, tongue, seminal vesicular, and urinary arteries are mineralized.
- * Bone Marrow
- * Heart
- * Intestine Large, Cecum
- * Kidney
 - [Nephropathy TGLs = 1 - 8]
- * Liver
 - Bile Duct
- * Lung
- * Parathyroid Gland
- * Pituitary Gland
 - Pars Distalis
- * Spleen
 - White Pulp
- * Stomach, Forestomach
- * Stomach, Glandular
- Testis
 - Bilateral, Germinal Epith
- * Thymus
- Tongue

- Hyperplasia
- Mineral
- Hemorrhage
- Cardiomyopathy
- Mineral
- Polyarteritis Nodosa
- Nephropathy
- Hyperplasia
- Mineral
- Hyperplasia
- Hyperplasia
- Atrophy
- Pigment
- Mineral
- Mineral
- Degeneration
- Atrophy
- Edema
- Inflammation
- Focal, Mild
- Mild
- Mild
- Moderate
- Mild
- Mild
- Mild
- Chronicprogr, Marked
- Minimal
- Mild
- Diffuse, Moderate
- Mild
- Marked
- Mild
- Mild
- Moderate
- Minimal
- Marked
- Moderate
- Chronic Active, Mild

[Edema TGLs = 2 - 17]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 78

TRT#: 3

SEX: Male

DAY ON TEST: 575

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402817

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 79

TRT#: 3

SEX: Male

DAY ON TEST: 575

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402818

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Forestomach

Epithelium

Hyperplasia

Minimal

* Thymus

Leukemia Mononuclear

PRIMARY CAUSE OF DEATH

- Spleen Leukemia Mononuclear

CONTRIBUTORY CAUSE OF DEATH

- Brain Hemorrhage

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 80

TRT#: 3

SEX: Male

DAY ON TEST: 680

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402819

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Preputial Gland
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel Mineral Mild
Note: The aorta and cardiac, gastric, and renal arteries are mineralized.
- * Bone Marrow Hypercellularity Mild
- * Heart Cardiomyopathy Mild
- * Intestine Large, Cecum Polyarteritis Nodosa Minimal
- * Intestine Small, Jejunum Carcinoma
[Carcinoma TGLs = 2 - 17]
- * Kidney Nephropathy Chronicprogr, Marked
[Nephropathy TGLs = 1 - 8]
- * Liver Bile Duct Cyst
- * Lung Mineral Minimal
- * Pancreas Acinus Adenoma
- Acinus Hyperplasia Moderate
- * Parathyroid Gland Hyperplasia Diffuse, Marked
- * Pituitary Gland Pars Distalis Hyperplasia Mild
- * Prostate Inflammation Chronic Active, Minimal
- * Spleen Extramedullary Hematopoiesis Moderate
- * Stomach, Glandular Mineral Mild
- Testis Bilateral, Germinal Epith Degeneration Moderate
- Polyarteritis Nodosa Minimal
- * Thymus Atrophy Moderate

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 81

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402820

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Small, Duodenum
- * Liver
- * Mammary Gland
- * Preputial Gland
- * Stomach, Forestomach
- * Trachea
- * Bone
- * Esophagus
- * Intestine Small, Ileum
- * Lung
- * Nose
- * Salivary Glands
- * Stomach, Glandular
- * Urinary Bladder
- * Bone Marrow
- * Harderian Gland
- * Intestine Small, Jejunum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- * Thymus
- * Brain
- * Intestine Large, Cecum
- * Islets, Pancreatic
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Skin
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla
- * Eye
- * Heart
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Kidney
- [Nephropathy TGLs = 2 - 8]
- Lymph Node
- Note: TGL-1 = Large C-cell carcinoma, possibly arising in ectopic thyroid tissue, since thyroid glands in routine section are normal.
- [Carcinoma TGLs = 1 - 17]
- [Hemorrhage TGLs = 3 - 18]
- * Pancreas
- * Prostate
- * Spleen
- Testis
- Cornea
- Lymphoid Tiss
- Bronchial
- Renal
- Renal
- Hyperplasia
- Pheochromocytoma Benign
- Inflammation
- Cardiomyopathy
- Polyarteritis Nodosa
- Hyperplasia
- Parasite Metazoan
- Nephropathy
- Oncocytoma Benign
- Carcinoma
- Hemorrhage
- Pigment
- Focal, Mild
- Chronic, Mild
- Minimal
- Mild
- Mild
- Chronicprogr, Marked
- Metastatic (Thyroid Gland)
- Mild
- Mild
- Polyarteritis Nodosa
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Hyperplasia
- Mild
- Minimal
- Mild
- Mild
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 81

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402820

ORGAN AND ACCOUNTABLE SITE STATUS

Polyarteritis Nodosa

Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 82

TRT#: 3

SEX: Male

DAY ON TEST: 706

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402821

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lung
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Urinary Bladder
- Blood Vessel
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- Testis
- * Bone
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Skin
- * Thyroid Gland
- * Bone Marrow
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Mammary Gland
- * Preputial Gland
- * Stomach, Forestomach
- * Trachea

OBSERVATIONS

- * Adrenal Medulla [Pheochromocytoma Benign TGLs = 1 - 11] Pheochromocytoma Benign
- * Eye Cornea Inflammation Chronic Active, Mild
- * Heart Cardiomyopathy Mild
- * Kidney Nephropathy Chronicprogr, Marked
- * Nose Inflammation Chronic Active, Minimal
- * Prostate Olfactory Epi Metaplasia Respiratory, Minimal
- * Spleen Hyperplasia Minimal
- * Thymus Extramedullary Hematopoiesis Mild
- Pigment Minimal
- Atrophy Mild

PRIMARY CAUSE OF DEATH - Adrenal Medulla Pheochromocytoma Benign

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 83

TRT#: 3

SEX: Male

DAY ON TEST: 687

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402822

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Epididymis
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- Testis
- * Bone Marrow
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Trachea

MISSING

- * Thymus

OBSERVATIONS

- * Adrenal Medulla
- * Eye
- * Heart
- * Intestine Large, Rectum
- * Kidney
- * Liver
- * Lung
- * Pancreas
- * Pituitary Gland
- * Spleen
- * Thyroid Gland
- Bilateral
- Cornea
- Acinus
- Pars Distalis
- C Cell
- Hyperplasia
- Inflammation
- Cardiomyopathy
- Parasite Metazoan
- Nephropathy
- Clear Cell Focus
- Eosinophilic Focus
- Inflammation
- Hyperplasia
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Hyperplasia
- Focal, Mild
- Chronic Active, Mild
- Minimal
- Chronicprogr, Moderate
- Granulomatous, Minimal
- Minimal
- Minimal
- Minimal
- Mild
- Mild

PRIMARY CAUSE OF DEATH

- UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 84

TRT#: 3

SEX: Male

DAY ON TEST: 654

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402823

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Pituitary Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|---------------------|-----------------------------------------------|------------------------------|------------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| | [Nephropathy TGLs = 5,6 - 8] | | |
| * Liver | | Extramedullary Hematopoiesis | Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 4 - 6] | | |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Skin | | Cyst Epithelial Inclusion | |
| | [Cyst Epithelial Inclusion TGLs = 3 - 18] | Liposarcoma | |
| | [Liposarcoma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Marked |
| | [Extramedullary Hematopoiesis TGLs = 2 - 8] | | |
| Testis | Bilateral, Germinal Epith | Degeneration | Moderate |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Hyperplasia | Minimal |

PRIMARY CAUSE OF DEATH - Skin Liposarcoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 85

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402824

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| Blood Vessel | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Parathyroid Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|-------------------|----------------------------------------|------------------------------|------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| | | Hyperplasia | Focal, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Eosinophilic Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 2 - 7] | | |
| * Pancreas | Acinus | Atrophy | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Skin | | Lipoma | |
| | [Lipoma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 86

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402825

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Bone Marrow
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Testis
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- * Intestine Large, Cecum Infiltration Cellular Lymphoid, Mild
- * Intestine Large, Colon Infiltration Cellular Lymphoid, Mild
- * Kidney Nephropathy Chronicprogr, Mild
- [Nephropathy TGLs = 2 - 8]
- * Liver Clear Cell Focus
- Bile Duct Cyst
- Eosinophilic Focus
- [Cyst TGLs = 1 - 17]
- * Lung Infiltration Cellular Histiocyte, Minimal
- * Spleen Extramedullary Hematopoiesis Mild
- Pigment Minimal
- * Thymus Atrophy Mild
- * Thyroid Gland C Cell Hyperplasia Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 87

TRT#: 3

SEX: Male

DAY ON TEST: 653

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402826

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 88

TRT#: 3

SEX: Male

DAY ON TEST: 729

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402827

ORGAN AND ACCOUNTABLE SITE STATUS

CONTRIBUTORY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 89

TRT#: 3

SEX: Male

DAY ON TEST: 708

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402828

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thymus | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|---------------------|---------------------------------------------|------------------------------|----------------------|
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Nose | Nasolacrim Dct | Inflammation | Chronic, Minimal |
| * Pancreas | Acinus | Hyperplasia | Marked |
| | [Hyperplasia TGLs = 4 - 20] | | |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Moderate |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Skin | | Keratoacanthoma | Multiple |
| | [Keratoacanthoma TGLs = 1,2,3 - 17+18+19] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 90

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402829

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- Blood Vessel
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- Testis
- * Urinary Bladder
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- * Thymus
- * Brain
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Stomach, Forestomach
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Glandular
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Kidney
- [Nephropathy TGLs = 1 - 8]
- * Liver
- [Eosinophilic Focus TGLs = 7 = 22]
- * Lung
- * Pancreas
- [Adenoma TGLs = 3,4 - 18+19]
- [Hyperplasia TGLs = 5,6 - 20+21]
- * Pituitary Gland
- * Skin
- [Adenoma TGLs = 2 - 17]
- * Spleen
- Degeneration
- Hyperplasia
- Hemorrhage
- Inflammation
- Hyperplasia
- Nephropathy
- Clear Cell Focus
- Eosinophilic Focus
- Inflammation
- Adenoma
- Hyperplasia
- Adenoma
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Cystic, Minimal
- Focal, Minimal
- Minimal
- Chronic, Minimal
- Moderate
- Chronicprogr, Marked
- Granulomatous, Minimal
- Multiple
- Marked
- Mild
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 91

TRT#: 3

SEX: Male

DAY ON TEST: 676

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402830

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Seminal Vesicle
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lung
- * Parathyroid Gland
- * Stomach, Glandular
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla
 - [Pheochromocytoma Benign TGLs = 4 - 11]
 - Blood Vessel
 - Note: Renal arteries are mineralized.
 - * Bone Marrow
 - * Heart
 - * Kidney
 - [Nephropathy TGLs = 5 - 8]
 - * Liver
 - * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 17]
 - * Pancreas
 - * Pituitary Gland
 - [Adenoma TGLs = 6 - 11]
 - * Prostate
 - * Skin
 - [Fibroma TGLs = 2 - 18]
 - * Spleen
 - * Stomach, Forestomach
 - Testis
 - [Degeneration TGLs = 3 - 14]
- Pheochromocytoma Benign
- Mineral
- Hypercellularity
- Cardiomyopathy
- Nephropathy
- Hyperplasia
- Fibroadenoma
- Adenoma
- Adenoma
- Inflammation
- Fibroma
- Extramedullary Hematopoiesis
- Hyperplasia
- Degeneration
- Polyarteritis Nodosa
- Minimal
- Moderate
- Chronicprogr, Marked
- Minimal
- Chronic Active, Mild
- Mild
- Minimal
- Moderate
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 91

TRT#: 3

SEX: Male

DAY ON TEST: 676

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402830

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Mild

* Urinary Bladder

Transit Epithe

Hyperplasia

Mild

Inflammation

Chronic, Mild

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 92

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402831

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|--------------------------|----------------|------------------------------|----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Mild |
| * Heart | | Cardiomyopathy | Minimal |
| * Intestine Large, Colon | Lymphoid Tiss | Hyperplasia | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Liver | | Eosinophilic Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | Interstitial | Fibrosis | Minimal |
| | | Infiltration Cellular | Histiocyte, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Minimal |
| | | Pigment | Mild |
| Testis | Germinal Epith | Degeneration | Minimal |
| | | Polyarteritis Nodosa | Mild |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 93

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402832

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel
Note: The aorta and cardiac and gastric arteries are mineralized. Mineral Mild
- * Bone Marrow Hemorrhage Moderate
- * Eye Cornea Inflammation Moderate
- * Heart Cardiomyopathy Chronic Active, Mild
- * Kidney Nephropathy Chronicprogr, Marked
- * Liver
[Nephropathy TGLs = 4 - 8]
Clear Cell Focus
Eosinophilic Focus
Fatty Change Focal, Mild
Hepatocellular Adenoma
Hyperplasia Minimal
- * Lung Inflammation Acute, Minimal
- * Mammary Gland Adenocarcinoma
- * Nose Nasolacrim Dct Inflammation Chronic, Minimal
- * Pancreas Acinus Hyperplasia Moderate
- * Parathyroid Gland Adenoma
- * Spleen White Pulp Atrophy Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 93

TRT#: 3

SEX: Male

DAY ON TEST: 730

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402832

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Glandular		Extramedullary Hematopoiesis	Mild
Testis	Bilateral, Germinal Epith	Mineral	Marked
		Degeneration	Mild
		Polyarteritis Nodosa	Mild
[Degeneration TGLs = 3 - 14]			
* Thymus		Atrophy	Marked

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 94

TRT#: 3

SEX: Male

DAY ON TEST: 559

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402833

ORGAN AND ACCOUNTABLE SITE STATUS

* Thyroid Gland

C Cell

Hyperplasia

Minimal

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 95

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402834

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|----------------------|----------------------------------------------|------------------------------|------------------------|
| * Adrenal Medulla | Bilateral | Pheochromocytoma Benign | |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | Renal Tubule | Cyst | |
| | | Nephropathy | Chronicprogr, Marked |
| | [Cyst TGLs = 2 - 18] | | |
| | [Nephropathy TGLs = 1 - 8+17] | | |
| * Liver | | Hepatodiaphragmatic Nodule | |
| | Bile Duct | Hyperplasia | Mild |
| | [Hepatodiaphragmatic Nodule TGLs = 3 - 12] | | |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Minimal |
| * Stomach, Glandular | | Mineral | Minimal |
| Testis | Bilateral, Germinal Epith | Degeneration | Moderate |
| | | Polyarteritis Nodosa | Moderate |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | | Polyarteritis Nodosa | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 96

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402835

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Harderian Gland
- * Intestine Large, Rectum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea
- * Brain
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Parathyroid Gland
- * Salivary Glands
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Cortex
 - * Adrenal Medulla
 - * Bone Marrow
 - * Eye
 - * Islets, Pancreatic
 - * Kidney
 - [Nephropathy TGLs = 3 - 8]
 - * Liver
 - * Lung
 - * Nose
 - * Pancreas
 - [Adenoma TGLs = 1,2 - 17+18]
 - * Spleen
 - Testis
 - * Thymus
- | | | |
|--------|------------------------------|-------------------------|
| | Hyperplasia | Focal, Minimal |
| | Pheochromocytoma Benign | |
| | Hemorrhage | Minimal |
| Cornea | Inflammation | Acute, Minimal |
| | Hyperplasia | Moderate |
| | Nephropathy | Chronicprogr, Moderate |
| | Clear Cell Focus | |
| | Hyperplasia | Minimal |
| | Inflammation | Granulomatous, Minimal |
| | Foreign Body | |
| | Inflammation | Chronic Active, Mild |
| | Inflammation | Chronic Active, Minimal |
| | Squamous Metaplasia | Minimal |
| | Adenoma | Multiple |
| | Hyperplasia | Minimal |
| | Extramedullary Hematopoiesis | Mild |
| | Pigment | Minimal |
| | Polyarteritis Nodosa | Minimal |
| | Atrophy | Mild |
| | Polyarteritis Nodosa | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 96

TRT#: 3

SEX: Male

DAY ON TEST: 731

DOSE: 75 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402835

ORGAN AND ACCOUNTABLE SITE STATUS

* Thyroid Gland

C Cell

Hyperplasia

Moderate

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 97

TRT#: 3

SEX: Male

DAY ON TEST: 447

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402836

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Medulla | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose | * Pancreas |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

MISSING

- | | |
|-----------------|---------------------|
| * Mammary Gland | * Parathyroid Gland |
|-----------------|---------------------|

OBSERVATIONS

- | | | |
|------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------|------------------------|
| * Adrenal Cortex | Hyperplasia | Focal, Minimal |
| * Heart | Cardiomyopathy | Minimal |
| * Kidney | Nephropathy | Chronicprogr, Moderate |
| * Lung | Infiltration Cellular | Histiocyte, Minimal |
| Note: Processor problems led to one lung slide (and recut) being almost uninterpretable. Additional lung processed as recut is interpretable and has infiltrate. | | |
| * Spleen | Extramedullary Hematopoiesis | Mild |
| | Pigment | Minimal |
| * Thymus | Atrophy | Minimal |

PRIMARY CAUSE OF DEATH

- UNCERTAIN

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 98

TRT#: 3

SEX: Male

DAY ON TEST: 654

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402837

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Mammary Gland
- * Prostate
- * Stomach, Glandular
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Pancreas
- * Salivary Glands
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla [Pheochromocytoma Benign TGLs = 1 - 17] Bilateral Pheochromocytoma Benign
- * Bone Marrow Hypercellularity Mild
- * Heart Cardiomyopathy Mild
- * Islets, Pancreatic Hyperplasia Minimal
- * Kidney [Nephropathy TGLs = 2,3 - 8] Nephropathy Chronicprogr, Marked
- * Liver Bile Duct Cyst
- Bile Duct Dilation Mild
- Bile Duct Extramedullary Hematopoiesis Minimal
- * Lung Bile Duct Hyperplasia Minimal
- Hemorrhage Mild
- Infiltration Cellular Histiocyte, Moderate
- Inflammation Granulomatous, Minimal
- * Lymph Node, Mandibular [Infiltration Cellular TGLs = 5 - 12] Infiltration Cellular Plasma Cell, Mild
- * Nose Inflammation Suppurative, Mild
- * Parathyroid Gland Hyperplasia Diffuse, Marked
- * Spleen Extramedullary Hematopoiesis Moderate
- * Stomach, Forestomach Inflammation Acute, Minimal
- Testis Polyarteritis Nodosa Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 98

TRT#: 3

SEX: Male

DAY ON TEST: 654

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402837

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Marked

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 99

TRT#: 3

SEX: Male

DAY ON TEST: 557

DOSE: 75 mg/kg male

DISP: Natural Death

HISTO: 1402838

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Prostate
- * Stomach, Forestomach
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
Blood Vessel
Note: The aorta and cardiac, pancreatic, renal, and salivary arteries are mineralized.
 - * Bone Marrow
 - * Eye
 - * Heart
 - * Kidney
[Nephropathy TGLs = 3 - 8]
 - * Lung
 - * Parathyroid Gland
 - * Spleen
 - * Stomach, Glandular
 - Testis
[Degeneration TGLs = 1 - 14]
 - * Thymus
- | | | |
|--|----------------|----------------------|
| | Hyperplasia | Focal, Minimal |
| | Mineral | Moderate |
| | Hemorrhage | Marked |
| | Inflammation | Acute, Mild |
| | Cardiomyopathy | Marked |
| | Mineral | Moderate |
| | Nephropathy | Chronicprogr, Marked |
| | Mineral | Minimal |
| | Hyperplasia | Diffuse, Moderate |
| | Atrophy | Marked |
| | Pigment | Minimal |
| | Mineral | Mild |
| | Adenoma | |
| | Degeneration | Moderate |
| | Atrophy | Marked |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 100

TRT#: 3

SEX: Male

DAY ON TEST: 618

DOSE: 75 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402839

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lung | * Lymph Node, Mesenteric | * Mammary Gland |
| * Pancreas | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|--------------------------|--------------------------------|------------------------------|-------------------------|
| * Bone Marrow | | Hypercellularity | Moderate |
| * Kidney | Renal Tubule | Cyst | |
| | | Nephropathy | Chronicprogr, Marked |
| | [Cyst TGLs = 4 - 8] | | |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| | [Hyperplasia TGLs = 3 - 12] | | |
| * Lymph Node, Mandibular | | Inflammation | Moderate |
| | [Inflammation TGLs = 2 - 17] | | |
| * Nose | Nasolacrim Dct | Inflammation | Chronic Active, Minimal |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Skin | | Fibrosarcoma | |
| | [Fibrosarcoma TGLs = 5 - 18] | | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Marked |

PRIMARY CAUSE OF DEATH - Skin Fibrosarcoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 101

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402840

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Trachea
- * Bone Marrow
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Parathyroid Gland
- * Salivary Glands
- * Stomach, Glandular
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Seminal Vesicle
- Testis
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Cortex
- * Adrenal Medulla
- Blood Vessel
- Note: Gastric arteries are mineralized.
- * Heart
- * Kidney
- * Liver
- * Lung
- * Nose
- * Pancreas
- * Spleen
- * Thymus
- Bile Duct
- Nasolacrim Dct
- Hyperplasia
- Pheochromocytoma Benign
- Mineral
- Cardiomyopathy
- Nephropathy
- Clear Cell Focus
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Polyarteritis Nodosa
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Focal, Minimal
- Minimal
- Minimal
- Chronicprogr, Marked
- Mild
- Histiocyte, Minimal
- Chronic, Minimal
- Minimal
- Mild
- Minimal
- Minimal

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 102

TRT#: 5

SEX: Male

DAY ON TEST: 664

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402841

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

MISSING

- * Thymus

OBSERVATIONS

- | | | | |
|---------------------------------------------------------------------------------------|---------------------------------|-----------------------------------------|------------------------|
| * Adrenal Medulla | Bilateral | Hyperplasia
Pheochromocytoma Benign | Focal, Moderate |
| Blood Vessel | Aorta | Dilation
Mineral | Mild
Mild |
| Note: The aorta and pulmonary arteries are mineralized.
[Dilation TGLs = 2 - 10] | | | |
| * Bone | | Osteopetrosis | Minimal |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Moderate |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Inflammation | Granulomatous, Minimal |
| | [Inflammation TGLs = 3 - 6+7] | | |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Marked |
| | [Hyperplasia TGLs = 4 - 11] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | White Pulp | Atrophy
Extramedullary Hematopoiesis | Mild
Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 102

TRT#: 5

SEX: Male

DAY ON TEST: 664

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402841

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Glandular		Pigment	Minimal
Testis	Bilateral, Germinal Epith	Mineral	Mild
		Degeneration	Minimal
		Polyarteritis Nodosa	Minimal

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 103

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402842

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Epididymis | * Esophagus | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Preputial Gland | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Trachea | * Urinary Bladder | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------|------------------------------|-------------------------|
| * Adrenal Medulla | Bilateral | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Mild |
| * Eye | Cornea | Inflammation | Chronic Active, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Minimal |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Pancreas | Acinus | Adenoma | Multiple |
| | Acinus | Hyperplasia | Marked |
| | [Adenoma TGLs = 2 - 17] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Prostate | | Inflammation | Chronic, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Hyperplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 104

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402843

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | * Bone | * Brain | * Esophagus |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Pituitary Gland | * Preputial Gland |
| * Salivary Glands | * Stomach, Forestomach | * Stomach, Glandular | * Trachea |

MISSING

- | | |
|-----------------|---------------------|
| * Mammary Gland | * Parathyroid Gland |
|-----------------|---------------------|

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------|------------------------------|--------------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Mild |
| * Epididymis | | Polyarteritis Nodosa | Minimal |
| * Eye | Cornea | Inflammation | Chronic Active, Mild |
| * Kidney | Renal Tubule | Cyst | |
| | Pelvis | Inflammation | Chronic Active, Mild |
| | | Nephropathy | Chronicprogr, Marked |
| | [Cyst TGLs = 1 - 8] | | |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Prostate | | Inflammation | Chronic Active, Mild |
| | | Polyarteritis Nodosa | Minimal |
| * Seminal Vesicle | | Inflammation | Chronic Active, Minimal |
| | | Polyarteritis Nodosa | Minimal |
| * Skin | | Inflammation | Chronic Active, Moderate |
| | | Ulcer | Mild |
| | [Inflammation TGLs = 2 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | | Polyarteritis Nodosa | Mild |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 104

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402843

ORGAN AND ACCOUNTABLE SITE STATUS

* Thyroid Gland

C Cell

Adenoma

* Urinary Bladder

Infiltration Cellular

Lymphocyte, Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 105

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402844

ORGAN AND ACCOUNTABLE SITE STATUS

* Skin		Fibroma	
	[Fibroma TGLs = 3 - 17]		
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
Testis	Bilateral, Interstit Cell	Adenoma	Multiple
	Bilateral, Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Mild
* Thymus		Atrophy	Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 106

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402845

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Blood Vessel | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Parathyroid Gland | * Pituitary Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Trachea |
| * Urinary Bladder | | | |

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------------|------------------------------|----------------------|
| * Adrenal Cortex | | Amyloid Deposition | Focal, Minimal |
| * Adrenal Medulla | | Pheochromocytoma Benign | |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 2 - 8+18] | | |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Dilation | Marked |
| | | Necrosis | Minimal |
| | [Dilation TGLs = 1 - 17] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Mild |
| | [Inflammation TGLs = 4 - 7] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Adenoma | |
| | Note: Adenoma has a necrotic center. | | |
| | [Adenoma TGLs = 3 - 11] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 107

TRT#: 5

SEX: Male

DAY ON TEST: 711

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402846

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Lymph Node, Mandibular
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Stomach, Glandular
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Seminal Vesicle
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Preputial Gland
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
[Pheochromocytoma Benign TGLs = 5 - 11]
Blood Vessel
Note: The aorta and renal arteries are mineralized.
- * Bone Marrow
- * Eye
- * Heart
- * Intestine Small, Duodenum
- * Kidney
[Nephropathy TGLs = 1,2 - 8+18]
- * Liver
[Hepatodiaphragmatic Nodule TGLs = 4 - 17]
- * Lung
- * Nose
- * Pancreas
- * Parathyroid Gland
- Peripheral Nerve
- Axon, Sciatic
- Axon, Trigeminal
- Pheochromocytoma Benign
- Mineral
- Hypercellularity
- Inflammation
- Cardiomyopathy
- Thrombus
- Polyarteritis Nodosa
- Nephropathy
- Clear Cell Focus
- Extramedullary Hematopoiesis
- Hepatodiaphragmatic Nodule
- Infiltration Cellular
- Inflammation
- Inflammation
- Polyarteritis Nodosa
- Hyperplasia
- Degeneration
- Degeneration
- Minimal
- Mild
- Chronic Active, Moderate
- Minimal
- Moderate
- Chronicprogr, Marked
- Minimal
- Histiocyte, Moderate
- Acute, Minimal
- Chronic, Minimal
- Mild
- Diffuse, Moderate
- Minimal
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 107

TRT#: 5

SEX: Male

DAY ON TEST: 711

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402846

ORGAN AND ACCOUNTABLE SITE STATUS

* Pituitary Gland [Adenoma TGLs = 3 - 11]	Axon, Tibial Pars Distalis	Degeneration Adenoma	Minimal
Spinal Cord	Axon	Degeneration Gliosis	Mild Mild
* Spleen		Extramedullary Hematopoiesis Pigment	Moderate Minimal
Testis	Bilateral, Germinal Epith	Degeneration Polyarteritis Nodosa	Minimal Moderate
* Thymus		Atrophy	Marked

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 108

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402847

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Epididymis
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- Testis
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Medulla
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 1 - 6]
- * Spleen
- * Thymus
- * Thyroid Gland
- C Cell
- Hyperplasia
- Nephropathy
- Clear Cell Focus
- Infiltration Cellular
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Hyperplasia
- Focal, Marked
- Chronicprogr, Mild
- Histiocyte, Mild
- Mild
- Minimal
- Mild
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 109

TRT#: 5

SEX: Male

DAY ON TEST: 618

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402848

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Brain
- * Epididymis
- * Esophagus
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- Blood Vessel Mineral Minimal
Note: The aorta and cardiac arteries are mineralized
- * Bone Fibrous Osteodystrophy Minimal
Note: Osteodystrohy is in nasal section.
- * Bone Marrow Hemorrhage Minimal
Hypercellularity Mild
- * Eye Cornea Inflammation Acute, Mild
- * Heart Cardiomyopathy Moderate
Mineral Minimal
- * Kidney Atrium Thrombus
Nephropathy Chronicprogr, Marked
[Nephropathy TGLs = 1 - 8]
- * Liver Clear Cell Focus
Eosinophilic Focus
- * Lung Bronchiole Fibrosis Minimal
Infiltration Cellular Histiocyte, Marked
[Infiltration Cellular TGLs = 3 - 6+7]
- * Pancreas Acinus Adenoma Multiple
[Adenoma TGLs = 2 - 17]
- * Parathyroid Gland Hyperplasia Diffuse, Marked
- * Spleen White Pulp Atrophy Moderate
Extramedullary Hematopoiesis Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 109

TRT#: 5

SEX: Male

DAY ON TEST: 618

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402848

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Forestomach

Inflammation

Acute, Minimal

Ulcer

Minimal

* Stomach, Glandular

Mineral

Minimal

Testis

Germinal Epith

Degeneration

Minimal

* Thymus

Atrophy

Moderate

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 110

TRT#: 5

SEX: Male

DAY ON TEST: 435

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402849

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Liver
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Glandular
- Testis
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- * Bone Marrow
 - * Brain
 - * Heart
 - * Kidney
 - * Lung
 - * Mammary Gland
 - * Spleen
 - * Stomach, Forestomach
 - * Thymus
 - * Thyroid Gland
 - Zymbal's Gland
- Meninges
- [Amphophilic/Vacuolar Adenoma TGLs = 2,3 - 17]
- [Amphophilic/Vacuolar Carcinoma TGLs = 1 - 17]
- [Infiltration Cellular TGLs = 4 - 6+7]
- Epithelium
- [Carcinoma TGLs = 5 - 18]

- Hypercellularity
- Marked
- Carcinoma
- Metastatic (Kidney)
- Cardiomyopathy
- Minimal
- Amphophilic/Vacuolar Adenoma
- Multiple
- Amphophilic/Vacuolar Carcinoma
- Amphophilic/Vacuolar Hyperplasia
- Moderate
- Nephropathy
- Chronicprogr, Mild
- Infiltration Cellular
- Histiocyte, Mild
- Hyperplasia
- Mild
- Extramedullary Hematopoiesis
- Mild
- Pigment
- Minimal
- Hyperplasia
- Minimal
- Atrophy
- Moderate
- Carcinoma
- Metastatic (Kidney)
- Carcinoma

PRIMARY CAUSE OF DEATH - Zymbal's Gland Carcinoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 111

TRT#: 5

SEX: Male

DAY ON TEST: 694

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402850

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Preputial Gland
- * Trachea
- Blood Vessel
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Salivary Glands
- * Urinary Bladder
- * Bone
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Brain
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Epididymis
- * Eye
- * Heart
- * Kidney
- [Nephropathy TGLs = 3 - 8]
- * Lung
- * Pituitary Gland
- [Adenoma TGLs = 4 - 11]
- * Prostate
- * Skin
- [Squamous Cell Papilloma TGLs = 2 - 17]
- * Spleen
- * Stomach, Forestomach
- Testis
- [Degeneration TGLs = 1 - 14]
- * Thymus
- Degeneration
- Hypercellularity
- Inflammation
- Inflammation
- Cardiomyopathy
- Inflammation
- Nephropathy
- Infiltration Cellular
- Adenoma
- Inflammation
- Squamous Cell Papilloma
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Hyperplasia
- Inflammation
- Degeneration
- Atrophy
- Cystic, Minimal
- Minimal
- Chronic Active, Minimal
- Acute, Mild
- Minimal
- Acute, Minimal
- Chronicprogr, Marked
- Histiocyte, Mild
- Chronic Active, Mild
- Mild
- Minimal
- Mild
- Mild
- Chronic Active, Mild
- Mild
- Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 111

TRT#: 5

SEX: Male

DAY ON TEST: 694

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402850

ORGAN AND ACCOUNTABLE SITE STATUS

* Thyroid Gland

C Cell

Adenoma

C Cell

Hyperplasia

Moderate

PRIMARY CAUSE OF DEATH

- Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 112

TRT#: 5

SEX: Male

DAY ON TEST: 413

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402851

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Trachea | * Urinary Bladder |

MISSING

- * Brain

OBSERVATIONS

- | | | | |
|---------------------------------------------------------------------------------|---------------------------|--------------------------------|----------------------|
| Blood Vessel | | Mineral | Minimal |
| Note: The aorta is mineralized. | | | |
| * Bone Marrow | | Hemorrhage | Marked |
| * Heart | | Cardiomyopathy | Mild |
| * Intestine Large, Cecum | | Mineral | Minimal |
| * Kidney | | Amphophilic/Vacuolar Adenoma | Multiple |
| | | Amphophilic/Vacuolar Carcinoma | |
| | | Nephropathy | Chronicprogr, Marked |
| Note: Multiple renal cysts are due to carcinoma, hyperplasias, and nephropathy. | | | |
| [Amphophilic/Vacuolar Carcinoma TGLs = 1 - 8] | | | |
| * Pancreas | Acinus | Hyperplasia | Mild |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Moderate |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Pigment | Mild |
| * Stomach, Glandular | | Mineral | Mild |
| Testis | Bilateral, Germinal Epith | Degeneration | Marked |
| | | Polyarteritis Nodosa | Minimal |
| [Degeneration TGLs = 2 - 14] | | | |
| * Thymus | | Atrophy | Marked |
| * Thyroid Gland | | Carcinoma | Metastatic (Kidney) |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 112

TRT#: 5

SEX: Male

DAY ON TEST: 413

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402851

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 114

TRT#: 5

SEX: Male

DAY ON TEST: 694

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402853

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Glandular | Testis | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|------------------------|------------------------------|------------------------------|-------------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Mild |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Liver | | Degeneration | Cystic, Minimal |
| | Bile Duct | Hyperplasia | Minimal |
| | | Necrosis | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Pancreas | | Polyarteritis Nodosa | Minimal |
| * Pituitary Gland | | Adenoma | |
| | Pars Distalis | | |
| | [Adenoma TGLs = 2 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Minimal |
| | | Pigment | Mild |
| * Stomach, Forestomach | | Erosion | Minimal |
| | | Inflammation | Chronic Active, Minimal |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | | Hyperplasia | Marked |
| | C Cell | | |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

CONTRIBUTORY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 115

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402854

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Bone Marrow
* Brain	* Epididymis	* Esophagus	* Eye
* Harderian Gland	* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum
* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic
* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Mammary Gland	* Pancreas
* Parathyroid Gland	* Preputial Gland	* Prostate	* Salivary Glands
* Seminal Vesicle	* Skin	* Stomach, Forestomach	* Stomach, Glandular
Testis	* Trachea	* Urinary Bladder	

OBSERVATIONS

* Adrenal Cortex		Degeneration	Cystic, Minimal
* Heart		Cardiomyopathy	Minimal
* Kidney		Nephropathy	Chronicprogr, Mild
* Liver		Clear Cell Focus	
	Bile Duct	Hyperplasia	Minimal
* Lung		Inflammation	Granulomatous, Minimal
* Nose		Inflammation	Chronic, Minimal
	Respirat Epith	Squamous Metaplasia	Mild
* Pituitary Gland	Pars Distalis	Hyperplasia	Minimal
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
* Thymus		Atrophy	Mild
* Thyroid Gland	C Cell	Adenoma	

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 117

TRT#: 5

SEX: Male

DAY ON TEST: 595

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402856

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Preputial Gland
- * Stomach, Glandular
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Lung
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Trachea
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
 - * Bone Marrow
 - * Heart
 - * Intestine Large, Colon
 - * Kidney
 - Renal Tubule
 - [Cyst TGLs = 1 - 17]
 - [Nephropathy TGLs = 2 - 8]
 - * Liver
 - Bile Duct
 - * Parathyroid Gland
 - * Prostate
 - * Skin
 - [Cyst Epithelial Inclusion TGLs = 3 - 18]
 - * Spleen
 - Testis
 - * Thymus
- Hyperplasia
Hypercellularity
Cardiomyopathy
Polyarteritis Nodosa
Cyst
Nephropathy
Hyperplasia
Hyperplasia
Hyperplasia
Cyst Epithelial Inclusion
Extramedullary Hematopoiesis
Pigment
Polyarteritis Nodosa
Atrophy
Polyarteritis Nodosa
- Focal, Minimal
Minimal
Minimal
Minimal
Chronicprogr, Marked
Minimal
Diffuse, Mild
Minimal
Mild
Mild
Mild
Mild
Mild
Minimal

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 118

TRT#: 5

SEX: Male

DAY ON TEST: 402

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402857

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|--------------------------|---------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular |
| * Mammary Gland | * Pancreas | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Thyroid Gland | * Trachea | * Urinary Bladder |

MISSING

- | | |
|-----------------------------|--------------------------|
| * Intestine Small, Duodenum | * Lymph Node, Mesenteric |
|-----------------------------|--------------------------|

OBSERVATIONS

- | | | | |
|------------------------------------------------------------------|----------------|----------------|-------------------------|
| Blood Vessel | | Mineral | Minimal |
| Note: The aorta and gastric and renal areteries are mineralized. | | | |
| * Bone Marrow | | Hemorrhage | Marked |
| * Heart | | Cardiomyopathy | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Lung | | Mineral | Moderate |
| * Nose | Nasolacrim Dct | Inflammation | Chronic Active, Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Pigment | Mild |
| * Stomach, Glandular | | Mineral | Moderate |
| Testis | Germinal Epith | Degeneration | Minimal |
| * Thymus | | Atrophy | Marked |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 119

TRT#: 5

SEX: Male

DAY ON TEST: 589

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402858

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Parathyroid Gland | * Pituitary Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|---------------|------------------------------------------|------------------------------|----------------------|
| * Bone Marrow | | Hypercellularity | Moderate |
| * Heart | | Cardiomyopathy | Minimal |
| | Atrium | Thrombus | |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| | | Necrosis | Moderate |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Skin | | Schwannoma Malignant | |
| | [Schwannoma Malignant TGLs = 1 - 17] | | |
| * Spleen | White Pulp | Atrophy | Mild |
| | | Extramedullary Hematopoiesis | Moderate |
| Testis | Germinal Epith | Degeneration | Marked |
| | [Degeneration TGLs = 2 - 14] | | |
| * Thymus | | Atrophy | Marked |

PRIMARY CAUSE OF DEATH - Skin Schwannoma Malignant

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 120

TRT#: 5

SEX: Male

DAY ON TEST: 604

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402859

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|--------------------------|---------------------------|--------------------------|
| * Bone | * Brain | * Epididymis | * Esophagus |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Rectum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Stomach, Forestomach | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|-------------------------------------------------------|---------------------------|------------------------------|----------------------|
| * Adrenal Cortex | | Necrosis | Mild |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| Blood Vessel | | Mineral | Mild |
| Note: The aorta and cardiac arteries are mineralized. | | | |
| * Bone Marrow | | Hemorrhage | Mild |
| | | Hypercellularity | Moderate |
| * Eye | Cornea | Inflammation | Chronic Active, Mild |
| * Heart | | Cardiomyopathy | Moderate |
| * Intestine Large, Colon | | Mineral | Minimal |
| | | Ulcer | Mild |
| * Intestine Small, Duodenum | | Polyarteritis Nodosa | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| [Nephropathy TGLs = 4 - 8] | | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Pancreas | Acinus | Atrophy | Minimal |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Skin | | Fibroma | |
| [Fibroma TGLs = 3 - 17] | | | |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Extramedullary Hematopoiesis | Mild |
| * Stomach, Glandular | | Mineral | Moderate |
| Testis | Bilateral, Germinal Epith | Degeneration | Moderate |
| | | Polyarteritis Nodosa | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 120

TRT#: 5

SEX: Male

DAY ON TEST: 604

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402859

ORGAN AND ACCOUNTABLE SITE STATUS

[Degeneration TGLs = 1 - 14]

* Thymus

Atrophy

Marked

[Atrophy TGLs = 2 - 6]

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 121

TRT#: 5

SEX: Male

DAY ON TEST: 606

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402860

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Harderian Gland
- * Islets, Pancreatic
- * Pituitary Gland
- * Seminal Vesicle
- * Urinary Bladder
- * Bone
- * Intestine Large, Colon
- * Lymph Node, Mandibular
- * Preputial Gland
- * Skin
- * Epididymis
- * Intestine Large, Rectum
- * Lymph Node, Mesenteric
- * Prostate
- * Thyroid Gland
- * Esophagus
- * Intestine Small, Ileum
- * Mammary Gland
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
 - Degeneration
 - Necrosis
 - Mineral
 - Cystic, Minimal
 - Mild
 - Mild
- Blood Vessel
 - Note: The aorta and cardiac arteries are mineralized.
- * Bone Marrow
 - Hypercellularity
 - Minimal
- * Brain
 - Necrosis
 - Marked
- * Eye
 - Cornea
 - Inflammation
 - Chronic Active, Marked
- * Heart
 - Cardiomyopathy
 - Mild
- * Intestine Large, Cecum
 - Perivascular
 - Infiltration Cellular
 - Lymphocyte, Minimal
- * Intestine Small, Duodenum
 - Polyarteritis Nodosa
 - Mild
- * Intestine Small, Jejunum
 - Peyers Patch
 - Polyarteritis Nodosa
 - Minimal
- * Kidney
 - [Nephropathy TGLs = 1 - 8+17]
 - Hyperplasia
 - Marked
 - Nephropathy
 - Chronicprogr, Marked
- * Liver
 - Clear Cell Focus
 - Infiltration Cellular
 - Histiocyte, Minimal
- * Lung
 - Inflammation
 - Chronic Active, Minimal
- * Nose
 - Nasolacrim Dct
 - Polyarteritis Nodosa
 - Mild
- * Pancreas
 - Hyperplasia
 - Diffuse, Minimal
- * Parathyroid Gland
 - Atrophy
 - Minimal
- * Spleen
 - White Pulp
 - Pigment
 - Mild
- * Stomach, Forestomach
 - Ulcer
 - Minimal
- * Stomach, Glandular
 - Mineral
 - Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 121

TRT#: 5

SEX: Male

DAY ON TEST: 606

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402860

ORGAN AND ACCOUNTABLE SITE STATUS

Testis

Bilateral, Germinal Epith

Degeneration

Minimal

Polyarteritis Nodosa

Mild

* Thymus

Atrophy

Mild

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 122

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402861

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | * Bone | * Brain | * Epididymis |
| * Esophagus | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lung | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Parathyroid Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | |

MISSING

- * Thymus

OBSERVATIONS

- | | | | |
|-----------------------------------------|-------------------|------------------------------|------------------------|
| * Adrenal Cortex | Bilateral | Hyperplasia | Focal, Mild |
| Blood Vessel | | Mineral | Minimal |
| Note: Gastric arteries are mineralized. | | | |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Minimal |
| * Eye | Cornea | Inflammation | Chronic Active, Mild |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| * Lymph Node, Mandibular | | Ectasia | Marked |
| [Ectasia TGLs = 1 - 17] | | | |
| * Pancreas | | Polyarteritis Nodosa | Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Mild |
| | Interstit Cell | Hyperplasia | Mild |
| | | Polyarteritis Nodosa | Minimal |
| * Thyroid Gland | Bilateral, C Cell | Adenoma | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 123

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402862

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder
- Blood Vessel
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Bone
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Stomach, Glandular
- * Brain
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Cortex
 - * Bone Marrow
 - * Epididymis
 - * Kidney
 - [Nephropathy TGLs = 4 - 8]
 - * Liver
 - * Lung
 - [Inflammation TGLs = 2 - 6+7]
 - * Pituitary Gland
 - Skeletal Muscle
 - Pars Distalis
 - Note: Vascular smooth muscle tumor suspected but not available under skeletal muscle in TDMSE.
 - [Sarcoma TGLs = 1 - 17]
 - * Spleen
 - Testis
 - * Thymus
 - * Thyroid Gland
 - [Adenoma TGLs = 3 - 11]
 - C Cell
- Hyperplasia
 - Hypercellularity
 - Mesothelioma Malignant
 - Nephropathy
 - Clear Cell Focus
 - Infiltration Cellular
 - Inflammation
 - Hyperplasia
 - Sarcoma
 - Extramedullary Hematopoiesis
 - Pigment
 - Mesothelioma Malignant
 - Atrophy
 - Adenoma
- Focal, Minimal
 - Mild
 - Chronicprogr, Moderate
 - Histiocyte, Mild
 - Granulomatous, Mild
 - Minimal
 - Mild
 - Minimal
 - Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 124

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402863

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen		Extramedullary Hematopoiesis	Minimal
		Pigment	Minimal
* Stomach, Forestomach		Polyarteritis Nodosa	Minimal
Testis	Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Mild
* Thymus		Atrophy	Minimal
* Thyroid Gland	C Cell	Adenoma	Multiple
	C Cell	Hyperplasia	Moderate

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 125

TRT#: 5

SEX: Male

DAY ON TEST: 609

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402864

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Seminal Vesicle
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Glandular

MISSING

- * Spleen

OBSERVATIONS

- | | | |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <ul style="list-style-type: none"> * Adrenal Cortex * Bone Marrow * Heart * Intestine Large, Rectum * Kidney * Lung * Pancreas <li style="padding-left: 20px;">[Hyperplasia TGLs = 1 - 17] * Parathyroid Gland * Pituitary Gland * Stomach, Forestomach Testis * Thymus | <ul style="list-style-type: none"> Necrosis Hypercellularity Cardiomyopathy Polyarteritis Nodosa Nephropathy Infiltration Cellular Hyperplasia Hyperplasia Hyperplasia Polyarteritis Nodosa Degeneration Polyarteritis Nodosa Atrophy | <ul style="list-style-type: none"> Minimal Minimal Minimal Minimal Chronicprogr, Marked Histiocyte, Minimal Moderate Diffuse, Mild Moderate Minimal Mild Moderate Minimal |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 127

TRT#: 5

SEX: Male

DAY ON TEST: 725

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402866

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Seminal Vesicle
- * Trachea
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Forestomach
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla
 - * Bone Marrow
 - * Heart
 - * Intestine Large, Colon
 - [Schwannoma Malignant TGLs = 3 - 17]
 - * Kidney
 - [Nephropathy TGLs = 1 - 8]
 - * Liver
 - * Lung
 - * Parathyroid Gland
 - * Pituitary Gland
 - * Preputial Gland
 - * Spleen
 - * Stomach, Glandular
 - Testis
 - [Degeneration TGLs = 2 - 14]
 - * Thymus
- | | | |
|--|------------------------------|----------------------|
| | Hyperplasia | Focal, Mild |
| | Hemorrhage | Marked |
| | Hypercellularity | Moderate |
| | Cardiomyopathy | Minimal |
| | Schwannoma Malignant | |
| | Nephropathy | Chronicprogr, Marked |
| | Clear Cell Focus | |
| | Infiltration Cellular | Histiocyte, Marked |
| | Hyperplasia | Diffuse, Mild |
| | Hyperplasia | Minimal |
| | Hyperplasia | Squamous, Moderate |
| | Atrophy | Moderate |
| | Extramedullary Hematopoiesis | Minimal |
| | Pigment | Minimal |
| | Mineral | Minimal |
| | Degeneration | Mild |
| | Atrophy | Marked |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 128

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402867

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Bone Marrow
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Parathyroid Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Trachea
- * Urinary Bladder

MISSING

- * Thymus

OBSERVATIONS

- * Kidney [Nephropathy TGLs = 4 - 8+18] Nephropathy Chronicprogr, Marked
- * Liver Bile Duct Hyperplasia Minimal
- * Lung [Inflammation TGLs = 2 - 6+7] Infiltration Cellular Histiocyte, Minimal
Inflammation Granulomatous, Mild
- * Nose Nasolacrim Dct Inflammation Chronic, Minimal
- * Pancreas Acinus Adenoma Multiple
[Adenoma TGLs = 1 - 17]
- * Pituitary Gland Pars Distalis Adenoma
- * Spleen Extramedullary Hematopoiesis Mild
Pigment Mild
- Testis Polyarteritis Nodosa Minimal
- * Thyroid Gland C Cell Hyperplasia Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 129

TRT#: 5

SEX: Male

DAY ON TEST: 521

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402868

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|---------------------------|---------------|-----------------------|-----------------------|
| * Intestine Large, Colon | | Parasite Metazoan | |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| [Adenoma TGLs = 1 - 11] | | | |
| * Spleen | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 130

TRT#: 5

SEX: Male

DAY ON TEST: 498

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402869

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lung | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | |
|---------------------------|------------------------------|------------------------|
| * Bone Marrow | Hypercellularity | Minimal |
| * Heart | Cardiomyopathy | Minimal |
| * Kidney | Nephropathy | Chronicprogr, Moderate |
| * Lymph Node, Mandibular | Ectasia | Mild |
| [Ectasia TGLs = 1 - 12] | | |
| * Spleen | Extramedullary Hematopoiesis | Mild |
| | Pigment | Minimal |
| * Thymus | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH

- UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 131

TRT#: 5

SEX: Male

DAY ON TEST: 445

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402870

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|------------------------|------------|------------------------------|-----------------------|
| * Brain | | Necrosis | Moderate |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Extramedullary Hematopoiesis | Minimal |
| | | Pigment | Minimal |
| * Stomach, Forestomach | | Inflammation | Acute, Minimal |
| | | Mineral | Minimal |
| | | Necrosis | Minimal |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH

- Brain Necrosis

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 132

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402871

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Brain | * Epididymis |
| * Esophagus | * Eye | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|------------------------------------------------|-----------------------------------------|------------------------------|----------------------|
| * Bone | | Osteopetrosis | Minimal |
| * Bone Marrow | | Hemorrhage | Minimal |
| * Harderian Gland | | Atrophy | Minimal |
| | | Metaplasia | Mild |
| Note: Metaplasia to lacrimal gland morphology. | | | |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | Renal Tubule | Cyst | |
| | | Nephropathy | Chronicprogr, Marked |
| | [Cyst TGLs = 1 - 17] | | |
| * Liver | | Clear Cell Focus | |
| | Lymph Node | Infiltration Cellular | Plasma Cell, Mild |
| | [Infiltration Cellular TGLs = 2 - 18] | | |
| * Pancreas | Acinus | Adenoma | Multiple |
| | Acinus | Hyperplasia | Marked |
| | | Polyarteritis Nodosa | Minimal |
| | [Adenoma TGLs = 3,4 - 19+20] | | |
| | [Hyperplasia TGLs = 5 - 21] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 132

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402871

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Glandular

Testis

Bilateral, Germinal Epith

Polyarteritis Nodosa

Degeneration

Polyarteritis Nodosa

Ectopic Parathyroid Gland

Mild

Minimal

Moderate

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 133

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402872

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Preputial Gland | * Salivary Glands |
| * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|---------------------|---------------------------|------------------------------|---------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | | Degeneration | Cystic, Minimal |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Moderate |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Prostate | | Inflammation | Chronic, Minimal |
| * Skin | | Fibroma | |
| | [Fibroma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 134

TRT#: 5

SEX: Male

DAY ON TEST: 677

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402873

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|--------------------------|---------------------------------------------|------------------------------|----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Marked |
| * Bone Marrow | | Hypercellularity | Minimal |
| * Heart | | Cardiomyopathy | Mild |
| * Intestine Large, Colon | | Polyarteritis Nodosa | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| Testis | Bilateral, Germinal Epith
Interstit Cell | Degeneration | Marked |
| | | Hyperplasia | Moderate |
| | | Polyarteritis Nodosa | Moderate |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 135

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402874

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Bone Marrow
* Brain	* Epididymis	* Esophagus	* Harderian Gland
* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum	* Intestine Small, Duodenum
* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic	* Lymph Node, Mandibular
* Lymph Node, Mesenteric	* Mammary Gland	* Nose	* Parathyroid Gland
* Pituitary Gland	* Preputial Gland	* Prostate	* Salivary Glands
* Seminal Vesicle	* Skin	* Stomach, Forestomach	* Stomach, Glandular
Testis	* Thyroid Gland	* Trachea	* Urinary Bladder

OBSERVATIONS

* Adrenal Cortex		Hyperplasia	Focal, Minimal
* Eye	Retina	Degeneration	Minimal
* Heart		Cardiomyopathy	Minimal
* Kidney		Nephropathy	Chronicprogr, Moderate
* Liver		Clear Cell Focus	
	Bile Duct	Hyperplasia	Minimal
* Lung		Infiltration Cellular	Histiocyte, Minimal
* Pancreas	Acinus	Hyperplasia	Marked
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
* Thymus		Atrophy	Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 136

TRT#: 5

SEX: Male

DAY ON TEST: 688

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402875

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Forestomach

Epithelium

Hyperplasia

Marked

Testis

Bilateral, Germinal Epith

Degeneration

Mild

Polyarteritis Nodosa

Minimal

[Degeneration TGLs = 1 - 14]

* Thymus

Atrophy

Moderate

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 137

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402876

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Seminal Vesicle
- * Trachea
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Stomach, Glandular
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla
 - Blood Vessel
 - Note: Renal arteries are mineralized.
- * Bone Marrow
- * Eye
 - Cornea
- * Heart
- * Kidney
 - [Nephropathy TGLs = 2 - 8+23]
- * Liver
 - Bile Duct
- * Lung
- * Pancreas
 - Acinus
 - [Adenoma TGLs = 4,5,6,7,8 - 18+19+20+21+22]
- * Pituitary Gland
 - Pars Distalis
- * Prostate
- * Skin
 - [Keratoacanthoma TGLs = 3 - 17]
- * Spleen
- Testis
 - Bilateral, Germinal Epith
 - [Degeneration TGLs = 1 - 14]

- Hyperplasia
- Mineral
- Hypercellularity
- Inflammation
- Cardiomyopathy
- Nephropathy
- Clear Cell Focus
- Hyperplasia
- Inflammation
- Adenoma
- Adenoma
- Inflammation
- Keratoacanthoma
- Extramedullary Hematopoiesis
- Pigment
- Degeneration
- Polyarteritis Nodosa
- Focal, Mild
- Minimal
- Minimal
- Chronic Active, Mild
- Minimal
- Chronicprogr, Marked
- Minimal
- Granulomatous, Minimal
- Multiple
- Chronic, Minimal
- Mild
- Minimal
- Moderate
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 137

TRT#: 5

SEX: Male

DAY ON TEST: 730

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402876

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Minimal

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 138

TRT#: 5

SEX: Male

DAY ON TEST: 558

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402877

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Bone | * Bone Marrow | * Epididymis | * Esophagus |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Pancreas | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|----------------------------------------------------------------------------------|--|----------------------|-------------------------|
| * Adrenal Cortex
[Adenoma TGLs = 2 - 11] | | Adenoma | |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Brain | | Meningioma Malignant | |
| * Eye
Note: Tumor from cranial nerve region of B04 extends along optic nerve. | | Meningioma Malignant | Metastatic (Brain) |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney
[Nephropathy TGLs = 3 - 8] | | Nephropathy | Chronicprogr, Marked |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Nose
Nasolacrim Dct | | Inflammation | Chronic Active, Minimal |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Pituitary Gland
[Meningioma Malignant TGLs = 1 - 11] | | Meningioma Malignant | Metastatic (Brain) |
| * Spleen | | Pigment | Mild |
| * Thymus | | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH - Brain Meningioma Malignant

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 139

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402878

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Parathyroid Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|--------------------------------------|----------------|------------------------------|------------------------|
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Intestine Large, Colon | Lymphoid Tiss | Hyperplasia | Mild |
| * Islets, Pancreatic | | Adenoma | |
| Note: Islet adenoma was also in B17. | | | |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Nose | Respirat Epith | Hyperplasia | Minimal |
| * Pancreas | Acinus | Adenoma | Multiple |
| | Acinus | Hyperplasia | Marked |
| [Adenoma TGLs = 1 - 17] | | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Moderate |
| | | Polyarteritis Nodosa | Minimal |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Adenoma | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 140

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402879

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|-------------------|---------------------------------|------------------------------|----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Moderate |
| * Eye | Cornea | Inflammation | Chronic Active, Mild |
| * Heart | | Cardiomyopathy | Minimal |
| | Valve | Inflammation | Chronic, Minimal |
| | | Neuroendocrine Tumor, Benign | |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8+17] | | |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | | Polyarteritis Nodosa | Mild |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 141

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402880

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Cortex	Blood Vessel	* Bone	* Bone Marrow
* Brain	* Epididymis	* Esophagus	* Eye
* Harderian Gland	* Heart	* Intestine Large, Cecum	* Intestine Large, Colon
* Intestine Large, Rectum	* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum
* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Mammary Gland	* Pancreas
* Parathyroid Gland	* Preputial Gland	* Prostate	* Salivary Glands
* Seminal Vesicle	* Skin	* Stomach, Forestomach	* Stomach, Glandular
Testis	* Trachea	* Urinary Bladder	

OBSERVATIONS

* Adrenal Medulla	Bilateral	Pheochromocytoma Benign	
* Islets, Pancreatic		Hyperplasia	Minimal
* Kidney		Nephropathy	Chronicprogr, Moderate
* Liver		Clear Cell Focus	
	Bile Duct	Hyperplasia	Minimal
* Lung		Infiltration Cellular	Histiocyte, Mild
	[Infiltration Cellular TGLs = 1 - 6+7]	Inflammation	Granulomatous, Minimal
* Nose		Foreign Body	
		Inflammation	Chronic Active, Minimal
	Nasolacrim Dct	Inflammation	Chronic Active, Mild
* Pituitary Gland	Pars Distalis	Hyperplasia	Mild
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Mild
* Thymus		Atrophy	Moderate
* Thyroid Gland	C Cell	Hyperplasia	Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 142

TRT#: 5

SEX: Male

DAY ON TEST: 695

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402881

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Preputial Gland | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|-------------------------------------------------------|-------------------------------------------|-------------------------|----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| Blood Vessel | | Mineral | Minimal |
| Note: The aorta and cardiac arteries are mineralized. | | | |
| * Bone Marrow | | Hemorrhage | Moderate |
| | | Hypercellularity | Mild |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| Oral Mucosa | Buccal | Squamous Cell Carcinoma | |
| | [Squamous Cell Carcinoma TGLs = 4 - 17] | | |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Moderate |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 3 - 11] | | |
| * Prostate | | Inflammation | Chronic, Minimal |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Pigment | Mild |
| * Stomach, Forestomach | Epithelium | Hyperplasia | Moderate |
| | | Inflammation | Chronic, Minimal |
| Testis | Bilateral, Germinal Epith | Degeneration | Mild |
| | | Polyarteritis Nodosa | Minimal |
| | [Degeneration TGLs = 2 - 14] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 142

TRT#: 5

SEX: Male

DAY ON TEST: 695

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402881

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Marked

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

CONTRIBUTORY CAUSE OF DEATH

- Oral Mucosa Buccal Squamous Cell Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 143

TRT#: 5

SEX: Male

DAY ON TEST: 621

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402882

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Prostate
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Trachea
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
Blood Vessel
Note: The aorta is mineralized.
- * Bone Marrow
- * Heart
- * Kidney
[Nephropathy TGLs = 3 - 8]
- * Liver
- * Lung
[Infiltration Cellular TGLs = 4- 6+7]
- * Parathyroid Gland
- * Skin
[Keratoacanthoma TGLs = 1 - 17]
- * Spleen
- * Stomach, Glandular
Testis
- * Thymus
- Necrosis
- Mineral
- Hemorrhage
- Cardiomyopathy
- Nephropathy
- Eosinophilic Focus
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Hyperplasia
- Keratoacanthoma
- Atrophy
- Pigment
- Mineral
- Degeneration
- Polyarteritis Nodosa
- Atrophy
- Cyst
- Minimal
- Mild
- Marked
- Mild
- Chronicprogr, Marked
- Minimal
- Histiocyte, Minimal
- Granulomatous, Minimal
- Diffuse, Minimal
- Marked
- Mild
- Moderate
- Minimal
- Minimal
- Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 143

TRT#: 5

SEX: Male

DAY ON TEST: 621

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402882

ORGAN AND ACCOUNTABLE SITE STATUS

Ectopic Parathyroid Gland

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 144

TRT#: 5

SEX: Male

DAY ON TEST: 547

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402883

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lung
- * Nose
- * Salivary Glands
- Testis
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Seminal Vesicle
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta is mineralized.
 - * Bone Marrow
 - * Heart
 - * Kidney
[Nephropathy TGLs = 1 - 8]
 - * Liver
 - * Pancreas
 - * Parathyroid Gland
 - * Spleen
 - * Stomach, Glandular
 - * Thymus
- | | | |
|-----------|------------------------------|------------------------|
| | Hyperplasia | Focal, Marked |
| | Mineral | Minimal |
| | Hypercellularity | Mild |
| | Cardiomyopathy | Mild |
| | Nephropathy | Chronicprogr, Moderate |
| Bile Duct | Hyperplasia | Minimal |
| | Necrosis | Mild |
| Acinus | Hyperplasia | Moderate |
| | Hyperplasia | Diffuse, Minimal |
| | Extramedullary Hematopoiesis | Minimal |
| | Pigment | Minimal |
| | Mineral | Minimal |
| | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 145

TRT#: 5

SEX: Male

DAY ON TEST: 725

DOSE: 250 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402884

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- * Bone Marrow
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Preputial Gland
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Salivary Glands
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla [Pheochromocytoma Benign TGLs = 1- 17] Pheochromocytoma Benign
- * Heart Cardiomyopathy Minimal
- * Kidney [Nephropathy TGLs = 4 - 8+20] Nephropathy Chronicprogr, Marked
- * Lung Infiltration Cellular Histiocyte, Minimal
- * Pituitary Gland Pars Distalis Adenoma
- * Prostate [Adenoma TGLs = 3 - 19] Inflammation Chronic Active, Minimal
- * Skin [Fibroma TGLs = 2 - 18] Fibroma
- * Spleen Extramedullary Hematopoiesis Minimal
- * Stomach, Forestomach Epithelium Pigment Minimal
- Testis Bilateral, Germinal Epith Hyperplasia Minimal
- Inflammation Chronic Active, Mild
- Degeneration Mild
- Polyarteritis Nodosa Mild
- Atrophy Moderate
- * Thyroid Gland C Cell Adenoma

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 146

TRT#: 5

SEX: Male

DAY ON TEST: 682

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402885

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|---------------|-----------|------------------------------|---------------------|
| * Bone Marrow | | Hemorrhage | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH

- UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 147

TRT#: 5

SEX: Male

DAY ON TEST: 592

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402886

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | Testis | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|-----------------|--------|------------------------------|-----------------------|
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Hyperplasia | Moderate |

PRIMARY CAUSE OF DEATH

- UNCERTAIN

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 148

TRT#: 5

SEX: Male

DAY ON TEST: 679

DOSE: 250 mg/kg male

DISP: Natural Death

HISTO: 1402887

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Stomach, Glandular
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Prostate
- * Thyroid Gland
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Trachea
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
 - * Adrenal Medulla
 - * Bone Marrow
 - * Brain
 - * Heart
 - * Kidney
 - [Nephropathy TGLs = 1 - 8]
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 2 - 6+7]
 - * Nose
 - * Pancreas
 - * Pituitary Gland
 - * Seminal Vesicle
 - * Skin
 - * Spleen
 - Testis
 - * Thymus
- Bilateral
 - Pars Distalis
 - Subcut Tiss
- Hyperplasia
 - Hyperplasia
 - Hemorrhage
 - Hypercellularity
 - Gliosis
 - Cardiomyopathy
 - Nephropathy
 - Basophilic Focus
 - Infiltration Cellular
 - Inflammation
 - Inflammation
 - Polyarteritis Nodosa
 - Hyperplasia
 - Metaplasia
 - Polyarteritis Nodosa
 - Polyarteritis Nodosa
 - Extramedullary Hematopoiesis
 - Pigment
 - Polyarteritis Nodosa
 - Atrophy
- Focal, Minimal
 - Focal, Mild
 - Minimal
 - Mild
 - Minimal
 - Mild
 - Chronicprogr, Marked
 - Histiocyte, Mild
 - Granulomatous, Minimal
 - Chronic Active, Minimal
 - Minimal
 - Mild
 - Squamous, Mild
 - Minimal
 - Minimal
 - Mild
 - Minimal
 - Mild
 - Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 149

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402888

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------|------------------------------|--------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Minimal |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| * Thymus | | Atrophy | Mild |

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 150

TRT#: 5

SEX: Male

DAY ON TEST: 731

DOSE: 250 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402889

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| Blood Vessel | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|-------------------|---------------------------|------------------------------|-------------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Mild |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| * Nose | | Inflammation | Chronic Active, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| Testis | Bilateral, Germinal Epith | Degeneration | Moderate |
| * Thymus | | Atrophy | Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 151

TRT#: 7

SEX: Male

DAY ON TEST: 730

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402890

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Seminal Vesicle
- * Thyroid Gland
- * Bone Marrow
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Skin
- * Trachea
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
 - Hyperplasia
 - Pheochromocytoma Benign
- * Eye
 - Inflammation
- * Heart
 - Cardiomyopathy
- * Kidney
 - Nephropathy
- * Liver
 - Clear Cell Focus
 - Cyst
- Bile Duct
 - Note: Cholangiofibrosis/atypical biliary cyst.
 - [Cyst TGLs = 1 - 17]
- * Lung
 - Infiltration Cellular
- * Pancreas
 - Polyarteritis Nodosa
- * Parathyroid Gland
 - Hyperplasia
- * Pituitary Gland
 - Hyperplasia
- * Spleen
 - Extramedullary Hematopoiesis
 - Pigment
- Testis
 - Germinal Epith
 - Degeneration
 - Polyarteritis Nodosa
- * Thymus
 - Atrophy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 152

TRT#: 7

SEX: Male

DAY ON TEST: 613

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402891

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Seminal Vesicle
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
- Aorta
- Hyperplasia
- Dilation
- Mineral
- Focal, Marked
- Moderate
- Marked
- Note: The aorta and cardiac, colonic, pulmonary, renal, and salivary arteries are mineralized.
- [Dilation TGLs = 3 - 10]
- * Bone Marrow
- * Heart
- * Intestine Large, Cecum
- Hemorrhage
- Cardiomyopathy
- Erosion
- Inflammation
- Mineral
- Polyarteritis Nodosa
- Nephropathy
- Moderate
- Mild
- Moderate
- Chronic Active, Moderate
- Moderate
- Mild
- Chronicprogr, Marked
- * Kidney
- [Nephropathy TGLs = 2 - 8]
- * Liver
- Basophilic Focus
- Clear Cell Focus
- Hyperplasia
- Mineral
- Polyarteritis Nodosa
- Edema
- Minimal
- Minimal
- Minimal
- Mild
- * Lung
- * Pancreas
- * Skin
- [Edema TGLs = 1 - 12]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 152

TRT#: 7

SEX: Male

DAY ON TEST: 613

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402891

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen	White Pulp	Atrophy	Marked
		Pigment	Mild
* Stomach, Forestomach		Edema	Mild
* Stomach, Glandular		Mineral	Marked
Testis	Bilateral, Germinal Epith	Degeneration	Minimal
		Polyarteritis Nodosa	Mild
* Thymus		Atrophy	Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 154

TRT#: 7

SEX: Male

DAY ON TEST: 502

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402893

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta and cardiac and pulmonary arteries are mineralized.
 - * Bone Marrow
 - * Heart
 - * Intestine Large, Cecum
 - * Kidney
[Nephropathy TGLs = 3 - 8]
 - * Lung
 - * Pancreas
 - * Parathyroid Gland
 - * Spleen
 - * Stomach, Glandular
 - Testis
[Degeneration TGLs = 2 - 14]
 - * Thymus
- Hyperplasia
Mineral
Hemorrhage
Hypercellularity
Cardiomyopathy
Inflammation
Mineral
Nephropathy
Infiltration Cellular
Hyperplasia
Hyperplasia
Atrophy
Pigment
Mineral
Degeneration
Atrophy
- Focal, Minimal
Mild
Mild
Moderate
Mild
Chronic, Mild
Minimal
Chronicprogr, Marked
Histiocyte, Mild
Moderate
Diffuse, Marked
Marked
Minimal
Moderate
Marked
Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 155

TRT#: 7

SEX: Male

DAY ON TEST: 643

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402894

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Prostate
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Trachea
- * Epididymis
- * Intestine Large, Rectum
- * Liver
- * Parathyroid Gland
- * Seminal Vesicle
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
 - Blood Vessel
 - Note: The aorta and cardiac and pancreatic arteries are mineralized.
- * Bone Marrow
 - Thrombus
 - Mineral
 - Mild
- * Heart
 - Hemorrhage
 - Moderate
 - Hypercellularity
 - Mild
 - Cardiomyopathy
 - Moderate
- * Intestine Large, Cecum
 - Erosion
 - Minimal
 - Inflammation
 - Chronic Active, Mild
 - Mineral
 - Mild
 - Polyarteritis Nodosa
 - Mild
 - Polyarteritis Nodosa
 - Minimal
 - Polyarteritis Nodosa
 - Mild
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Kidney
 - Renal Tubule
 - Cyst
 - Nephropathy
 - Chronicprogr, Marked
 - [Cyst TGLs = 2 - 18]
 - [Nephropathy TGLs = 1 - 8]
- * Lung
 - Mineral
 - Minimal
- * Nose
 - Inflammation
 - Chronic Active, Mild
- * Pancreas
 - Nasolacrim Dct
 - Inflammation
 - Chronic Active, Moderate
 - Acinus
 - Hyperplasia
 - Minimal
- * Pituitary Gland
- * Skin
 - Pars Distalis
 - Adenoma
 - Foreign Body
 - Inflammation
 - Granulomatous, Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 155

TRT#: 7

SEX: Male

DAY ON TEST: 643

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402894

ORGAN AND ACCOUNTABLE SITE STATUS

Note: Foreign material is keratin from a previous (healing) lesion.

[Inflammation TGLs = 3 - 17]

* Spleen	White Pulp	Atrophy	Minimal
		Extramedullary Hematopoiesis	Mild
		Pigment	Mild
* Stomach, Glandular		Mineral	Moderate
Testis	Bilateral, Germinal Epith	Degeneration	Mild
		Polyarteritis Nodosa	Mild
* Thymus		Atrophy	Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 156

TRT#: 7

SEX: Male

DAY ON TEST: 725

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402895

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Parathyroid Gland
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Preputial Gland
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Mammary Gland
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Eye
- * Heart
- * Kidney
- [Nephropathy TGLs = 6 - 8+22]
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 8 - 6+7+23]
- Lymph Node
- [Hemorrhage TGLs = 1 - 18]
- [Hyperplasia TGLs = 2-19]
- * Lymph Node, Mandibular
- [Infiltration Cellular TGLs = 7 - 21]
- * Nose
- * Pancreas
- Hyperplasia
- Hemorrhage
- Hypercellularity
- Inflammation
- Cornea
- Cardiomyopathy
- Nephropathy
- Bile Duct
- Bronchiole
- Lumbar, Lymph Sinus
- Mediastinal
- Lumbar
- Mediastinal
- Ectasia
- Hemorrhage
- Hyperplasia
- Pigment
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Adenoma
- Focal, Mild
- Minimal
- Minimal
- Chronic Active, Minimal
- Minimal
- Chronicprogr, Marked
- Minimal
- Mild
- Histiocyte, Moderate
- Chronic Active, Moderate
- Mild
- Mild
- Plasma Cell, Mild
- Mild
- Lymphoid, Mild
- Plasma Cell, Mild
- Chronic Active, Minimal
- Multiple

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 156

TRT#: 7

SEX: Male

DAY ON TEST: 725

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402895

ORGAN AND ACCOUNTABLE SITE STATUS

	Acinus	Atrophy	Minimal
[Adenoma TGLs = 3 - 20]			
* Pituitary Gland	Pars Distalis	Hyperplasia	Minimal
* Prostate		Inflammation	Chronic Active, Moderate
[Inflammation TGLs = 4 - 16]			
* Seminal Vesicle		Inflammation	Chronic Active, Moderate
* Skin	Epidermis	Hyperplasia	Mild
		Inflammation	Chronic Active, Marked
		Ulcer	Marked
[Ulcer TGLs = 5 - 17]			
* Spleen		Extramedullary Hematopoiesis	Moderate
* Stomach, Forestomach	Epithelium	Hyperplasia	Mild
		Inflammation	Chronic Active, Minimal
Testis	Germinal Epith	Degeneration	Minimal
* Thymus		Atrophy	Moderate
* Thyroid Gland	C Cell	Adenoma	
* Urinary Bladder		Inflammation	Chronic, Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 157

TRT#: 7

SEX: Male

DAY ON TEST: 632

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402896

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Bone Marrow |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Parathyroid Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|------------------------|--------------------------------|------------------------------|------------------------|
| * Heart | | Cardiomyopathy | Minimal |
| * Islets, Pancreatic | | Adenoma | |
| | [Adenoma TGLs = 2 - 18] | | |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Fatty Change | Focal, Minimal |
| | [Fatty Change TGLs = 3 - 19] | | |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Pancreas | Acinus | Atrophy | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Skin | | Fibroma | |
| | [Fibroma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Minimal |
| | | Pigment | Minimal |
| * Stomach, Forestomach | Epithelium | Hyperplasia | Moderate |
| | | Inflammation | Chronic, Mild |
| Testis | Germinal Epith | Degeneration | Moderate |
| * Thymus | | Atrophy | Marked |
| * Thyroid Gland | C Cell | Hyperplasia | Minimal |

PRIMARY CAUSE OF DEATH - Islets, Pancreatic Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 158

TRT#: 7

SEX: Male

DAY ON TEST: 586

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402897

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Salivary Glands
- * Stomach, Glandular
- * Urinary Bladder
- Blood Vessel
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- Testis
- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Preputial Gland
- * Skin
- * Thyroid Gland
- * Bone Marrow
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Prostate
- * Stomach, Forestomach
- * Trachea

MISSING

- * Mammary Gland

OBSERVATIONS

- * Adrenal Cortex
- * Heart
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 1 - 6]
- [Inflammation TGLs = 2 - 7]
- * Spleen
- * Thymus
- Hyperplasia
- Cardiomyopathy
- Nephropathy
- Clear Cell Focus
- Infiltration Cellular
- Inflammation
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Focal, Mild
- Minimal
- Chronicprogr, Moderate
- Histiocyte, Minimal
- Granulomatous, Minimal
- Mild
- Mild
- Mild

PRIMARY CAUSE OF DEATH

- UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 160

TRT#: 7

SEX: Male

DAY ON TEST: 564

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402899

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Mammary Gland | * Pancreas | * Parathyroid Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|--------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------|----------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Brain | | Gliosis | Mild |
| | | Granular Cell Tumor Benign | |
| | Note: Possibly this started as an infarct caused by tumor embolus, although no tumor cells are visible in the section presented. | | |
| * Kidney | | Amphophilic/Vacuolar Adenoma | Multiple |
| | | Amphophilic/Vacuolar Carcinoma | Multiple |
| | | Amphophilic/Vacuolar Hyperplasia | Moderate |
| | | Nephropathy | Chronicprogr, Mild |
| | [Amphophilic/Vacuolar Carcinoma TGLs = 7, 8 - 19, 20] | | |
| * Liver | | Carcinoma | Metastatic (Kidney) |
| | | Extramedullary Hematopoiesis | Minimal |
| | [Carcinoma TGLs = 3,4,5 - 12+17] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| Lymph Node | Renal | Ectasia | Marked |
| | Note: Uncertain if this is a lymph node or simply a region of resolved hemorrhage leaving a cyst, however, "Cyst" is not a choice in any of the relevant organs. | | |
| | Note: Since this had no relationship with exposure and was not a cause of death, no attempt was made to add the term to "peritoneum" or "mesentery". | | |
| | [Ectasia TGLs = 9 - 21] | | |
| * Lymph Node, Mesenteric | | Carcinoma | Metastatic (Kidney) |
| | [Carcinoma TGLs = 6 - 18] | | |
| * Nose | | Inflammation | Suppurative, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 160

TRT#: 7

SEX: Male

DAY ON TEST: 564

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402899

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen

Extramedullary Hematopoiesis

Moderate

[Extramedullary Hematopoiesis TGLs = 2 - 8]

* Thymus

Atrophy

Moderate

PRIMARY CAUSE OF DEATH

- Kidney Amphophilic/Vacuolar Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 161

TRT#: 7

SEX: Male

DAY ON TEST: 633

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402900

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | * Bone | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Preputial Gland |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|----------------------|----------------|-------------------------------|------------------------|
| * Adrenal Cortex | | Necrosis | Minimal |
| * Bone Marrow | | Hemorrhage | Minimal |
| * Brain | | Inflammation | Chronic Active, Mild |
| * Heart | | Inflammation | Chronic Active, Marked |
| | Valve | Thrombus | |
| * Islets, Pancreatic | | Hyperplasia | Mild |
| * Kidney | Renal Tubule | Accumulation, Hyaline Droplet | Minimal |
| | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Necrosis | Minimal |
| | | Polyarteritis Nodosa | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Prostate | | Inflammation | Chronic, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Minimal |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Adenoma | |
| | C Cell | Hyperplasia | Minimal |

PRIMARY CAUSE OF DEATH - Heart Valve Thrombus

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 162

TRT#: 7

SEX: Male

DAY ON TEST: 598

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402901

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Glandular
- * Epididymis
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Esophagus
- * Intestine Small, Duodenum
- * Lung
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta is mineralized.
- * Bone Marrow
- * Brain
- * Eye
- * Heart
- * Intestine Large, Colon
- * Kidney
[Nephropathy TGLs = 1 - 8]
- * Liver
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Skin
[Keratoacanthoma TGLs = 2 - 17]
- * Spleen
- Testis
- * Thymus
- Hyperplasia
- Mineral
- Hypercellularity
- Necrosis
- Inflammation
- Cornea
- Cardiomyopathy
- Polyarteritis Nodosa
- Nephropathy
- Basophilic Focus
- Inflammation
- Polyarteritis Nodosa
- Hyperplasia
- Keratoacanthoma
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Degeneration
- Polyarteritis Nodosa
- Atrophy
- Focal, Marked
- Minimal
- Mild
- Mild
- Acute, Minimal
- Minimal
- Mild
- Chronicprogr, Marked
- Suppurative, Minimal
- Minimal
- Diffuse, Moderate
- Moderate
- Mild
- Minimal
- Minimal
- Moderate
- Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 162

TRT#: 7

SEX: Male

DAY ON TEST: 598

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402901

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 163

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402902

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Bone | * Brain | * Epididymis | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular |
| * Thymus | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|-------------------|----------------------------------------------|------------------------------|------------------------|
| * Adrenal Cortex | Bilateral | Hyperplasia | Focal, Mild |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Basophilic Focus | |
| | | Clear Cell Focus | |
| | | Hepatodiaphragmatic Nodule | |
| | Bile Duct | Hyperplasia | Minimal |
| | [Hepatodiaphragmatic Nodule TGLs = 2 - 18] | | |
| * Skin | | Fibroma | |
| | [Fibroma TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Minimal |
| | | Polyarteritis Nodosa | Mild |
| * Thyroid Gland | C Cell | Adenoma | |
| | | Polyarteritis Nodosa | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 164

TRT#: 7

SEX: Male

DAY ON TEST: 693

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402903

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Parathyroid Gland | * Pituitary Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|--------------------------|----------------|------------------------------|------------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Bone Marrow | | Hemorrhage | Marked |
| * Heart | | Cardiomyopathy | Mild |
| * Intestine Large, Cecum | | Inflammation | Acute, Minimal |
| * Kidney | Renal Tubule | Cyst | |
| | | Nephropathy | Chronicprogr, Marked |
| | | | [Cyst TGLs = 2,4 - 8] |
| | | | [Nephropathy TGLs = 3 - 8] |
| * Liver | Bile Duct | Hyperplasia | Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Marked |
| * Pancreas | | Polyarteritis Nodosa | Minimal |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Stomach, Glandular | | Mineral | Minimal |
| Testis | Germinal Epith | Degeneration | Minimal |
| | | Polyarteritis Nodosa | Minimal |
| * Thymus | | Atrophy | Marked |
| | | | [Atrophy TGLs = 1 - 6] |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 165

TRT#: 7

SEX: Male

DAY ON TEST: 569

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402904

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Epididymis |
| * Esophagus | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Pancreas |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|-----------------------------------------------------------------|----------------|----------------------|-----------------------|
| Blood Vessel | | Mineral | Mild |
| Note: The aorta and cardiac and renal arteries are mineralized. | | | |
| Note: Some pulmonary arteries are hyperplastic. | | | |
| * Bone Marrow | | Hemorrhage | Mild |
| * Brain | Meninges | Meningioma Malignant | |
| * Eye | Cornea | Inflammation | Chronic, Marked |
| Note: Perforation. | | | |
| [Inflammation TGLs = 2 - 13] | | | |
| * Harderian Gland | | Inflammation | Suppurative, Moderate |
| * Heart | | Mineral | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | Pleura | Fibrosis | Minimal |
| | Alveolar Epith | Hypertrophy | Minimal |
| | | Polyarteritis Nodosa | Mild |
| | | Thrombus | |
| [Polyarteritis Nodosa TGLs = 1-6+7] | | | |
| * Nose | Nasolacrim Dct | Inflammation | Chronic Active, Mild |
| * Parathyroid Gland | | Hyperplasia | Diffuse, Mild |
| * Spleen | White Pulp | Atrophy | Marked |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 165

TRT#: 7

SEX: Male

DAY ON TEST: 569

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402904

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- UNCERTAIN

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 166

TRT#: 7

SEX: Male

DAY ON TEST: 682

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402905

ORGAN AND ACCOUNTABLE SITE STATUS

Polyarteritis Nodosa

Mild

[Degeneration TGLs = 5 - 14]

Atrophy

Moderate

* Thymus

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 167

TRT#: 7

SEX: Male

DAY ON TEST: 607

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402906

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Preputial Gland |
| * Prostate | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|---------------------------|---------------|-----------------------|------------------------|
| * Adrenal Cortex | Bilateral | Hyperplasia | Focal, Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| [Adenoma TGLs = 1 - 11] | | | |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 168

TRT#: 7

SEX: Male

DAY ON TEST: 718

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402907

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|----------------------------------------------------------------------------------------|------------------|------------------------------|--------------------------|
| * Brain | | Necrosis | Moderate |
| * Heart | | Cardiomyopathy | Mild |
| * Islets, Pancreatic | | Carcinoma | |
| [Carcinoma TGLs = 1 - 17] | | | |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | Alveolar Epith | Hyperplasia | Minimal |
| * Lymph Node, Mandibular | | Infiltration Cellular | Plasma Cell, Mild |
| Peripheral Nerve | Axon, Trigeminal | Degeneration | Mild |
| | Axon, Sciatic | Degeneration | Mild |
| | Axon, Tibial | Degeneration | Moderate |
| | | Degeneration | Mild |
| | Trigeminal | Fibrosis | Mild |
| Note: Greater trigeminal nerve degeneration is evident in bone sections (B01 and B02). | | | |
| Skeletal Muscle | | Degeneration | Mild |
| * Skin | | Inflammation | Chronic Active, Moderate |
| | | Ulcer | Moderate |
| [Ulcer TGLs = 2 - 18] | | | |
| Spinal Cord | Axon | Degeneration | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 168

TRT#: 7

SEX: Male

DAY ON TEST: 718

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402907

ORGAN AND ACCOUNTABLE SITE STATUS

Testis
* Thymus

Germinal Epith

Degeneration
Atrophy

Minimal
Marked

PRIMARY CAUSE OF DEATH

- Islets, Pancreatic Carcinoma

CONTRIBUTORY CAUSE OF DEATH

- Brain Necrosis

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 169

TRT#: 7

SEX: Male

DAY ON TEST: 730

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402908

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Parathyroid Gland | * Pituitary Gland | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Stomach, Forestomach | Testis |
| * Thymus | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|----------------------|---------------------------|------------------------------|--------------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Marked |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| * Pancreas | Acinus | Adenoma | |
| | [Adenoma TGLs = 2 - 18] | | |
| * Skin | Epidermis | Hyperplasia | Minimal |
| | | Inflammation | Chronic Active, Moderate |
| | | Ulcer | Marked |
| | [Ulcer TGLs = 1 - 17] | | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Stomach, Glandular | | Mineral | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 170

TRT#: 7

SEX: Male

DAY ON TEST: 442

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402909

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Seminal Vesicle
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Preputial Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Prostate
- * Stomach, Glandular
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Cortex
 - Necrosis Mild
 - Thrombus
- * Bone Marrow
 - Hemorrhage Mild
 - Hypercellularity Mild
- * Heart
 - Cardiomyopathy Moderate
- * Kidney
 - Nephropathy Chronicprogr, Marked
- [Nephropathy TGLs = 2 - 8]
- * Liver
 - Bile Duct Hyperplasia Mild
 - Bile Duct Inflammation Chronic Active, Mild
- * Lung
 - Interstitial Fibrosis Mild
 - Inflammation Granulomatous, Minimal
- [Fibrosis TGLs = 3 - 6]
- * Nose
 - Nasolacrim Dct Inflammation Chronic Active, Minimal
- * Pancreas
 - Acinus Hyperplasia Minimal
- * Parathyroid Gland
 - Hyperplasia Diffuse, Moderate
- * Pituitary Gland
 - Pars Nervosa Thrombus
- * Skin
 - Hemorrhage Mild
 - Epidermis Hyperplasia Mild
 - Ulcer Mild
- [Ulcer TGLs = 1 - 17]
- * Spleen
 - White Pulp Atrophy Moderate
 - Extramedullary Hematopoiesis Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 170

TRT#: 7

SEX: Male

DAY ON TEST: 442

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402909

ORGAN AND ACCOUNTABLE SITE STATUS

Pigment

Minimal

Testis

Polyarteritis Nodosa

Minimal

* Thymus

Atrophy

Marked

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 171

TRT#: 7

SEX: Male

DAY ON TEST: 620

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402910

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Parathyroid Gland |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thymus | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|-------------------|----------------------------------------------|------------------------------|----------------------|
| * Adrenal Medulla | | Pheochromocytoma Malignant | |
| | [Pheochromocytoma Malignant TGLs = 3 - 11] | | |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Minimal |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Pancreas | | Polyarteritis Nodosa | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| Testis | | Polyarteritis Nodosa | Mild |
| * Thyroid Gland | C Cell | Hyperplasia | Marked |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 172

TRT#: 7

SEX: Male

DAY ON TEST: 650

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402911

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Bone | * Bone Marrow | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Preputial Gland | * Prostate | * Salivary Glands |
| * Seminal Vesicle | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| Testis | * Thyroid Gland | * Trachea | * Urinary Bladder |

OBSERVATIONS

- | | | | |
|-------------------|----------------|------------------------------|--------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Adrenal Medulla | | Pheochromocytoma Benign | |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| * Nose | Nasolacrim Dct | Inflammation | Chronic, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Adrenal Medulla Pheochromocytoma Benign

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 173

TRT#: 7

SEX: Male

DAY ON TEST: 730

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402912

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Bone Marrow
* Brain	* Epididymis	* Esophagus	* Eye
* Harderian Gland	* Heart	* Intestine Large, Cecum	* Intestine Large, Colon
* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum	* Lymph Node, Mandibular
* Lymph Node, Mesenteric	* Mammary Gland	* Nose	* Pancreas
* Pituitary Gland	* Preputial Gland	* Salivary Glands	* Seminal Vesicle
* Skin	* Stomach, Forestomach	* Stomach, Glandular	Testis
* Thymus	* Thyroid Gland	* Trachea	* Urinary Bladder

OBSERVATIONS

* Adrenal Cortex		Hyperplasia	Focal, Minimal
* Intestine Large, Rectum		Parasite Metazoan	
* Islets, Pancreatic		Hyperplasia	Moderate
* Kidney		Nephropathy	Chronicprogr, Moderate
* Liver		Clear Cell Focus	
	Bile Duct	Hyperplasia	Minimal
* Lung	Alveolar Epith	Hyperplasia	Minimal
		Infiltration Cellular	Histiocyte, Minimal
* Parathyroid Gland		Hyperplasia	Diffuse, Mild
* Prostate		Inflammation	Chronic, Minimal
* Spleen		Extramedullary Hematopoiesis	Minimal
		Pigment	Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 174

TRT#: 7

SEX: Male

DAY ON TEST: 663

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402913

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Epididymis | * Esophagus | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Pituitary Gland |
| * Preputial Gland | * Prostate | * Salivary Glands | * Seminal Vesicle |
| * Spleen | * Stomach, Forestomach | * Trachea | * Urinary Bladder |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|---------------------------------------|-----------------------------------|---------------------------|------------------------|
| Blood Vessel | | Mineral | Minimal |
| Note: Renal arteries are mineralized. | | | |
| * Bone Marrow | | Hypercellularity | Mild |
| * Eye | Cornea | Inflammation | Acute, Mild |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Minimal |
| * Skin | | Cyst Epithelial Inclusion | |
| | | Keratoacanthoma | |
| | [Keratoacanthoma TGLs = 1 - 17] | | |
| * Stomach, Glandular | | Mineral | Mild |
| Testis | | Polyarteritis Nodosa | Minimal |
| * Thymus | | Atrophy | Marked |
| * Thyroid Gland | C Cell | Hyperplasia | Moderate |

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 175

TRT#: 7

SEX: Male

DAY ON TEST: 730

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402914

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Forestomach
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Parathyroid Gland
- * Seminal Vesicle
- Testis
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Skin
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Heart
- * Kidney
- * Liver
- * Lung
 - [Infiltration Cellular TGLs = 2 - 6+7]
- * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 17]
- * Nose
- * Pituitary Gland
 - [Adenoma TGLs = 3 - 18]
- * Spleen
- * Thymus
- Hyperplasia
- Hypercellularity
- Cardiomyopathy
- Nephropathy
- Clear Cell Focus
- Infiltration Cellular
- Fibroadenoma
- Inflammation
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Focal, Minimal
- Minimal
- Minimal
- Chronicprogr, Mild
- Histiocyte, Mild
- Chronic, Minimal
- Mild
- Mild
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 177

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402916

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Eye
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Parathyroid Gland
- * Seminal Vesicle
- * Trachea
- * Epididymis
- * Intestine Large, Rectum
- * Liver
- * Preputial Gland
- * Skin
- * Esophagus
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Prostate
- * Stomach, Glandular

MISSING

- * Intestine Large, Cecum
- * Lymph Node, Mesenteric

OBSERVATIONS

- | | | | |
|---------------------------------------|------------------------------|------------------------------|----------------------|
| * Adrenal Cortex | | Angiectasis | Mild |
| | | Thrombus | |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| Blood Vessel | | Mineral | Minimal |
| Note: Renal arteries are mineralized. | | | |
| * Bone Marrow | | Hypercellularity | Mild |
| * Harderian Gland | | Atrophy | Mild |
| * Heart | | Cardiomyopathy | Mild |
| * Intestine Small, Duodenum | | Polyarteritis Nodosa | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 1 - 8] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | | Inflammation | Chronic Active, Mild |
| * Nose | Nasolacrim Dct | Inflammation | Chronic, Minimal |
| * Pancreas | | Polyarteritis Nodosa | Moderate |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Stomach, Forestomach | | Polyarteritis Nodosa | Mild |
| Testis | Germinal Epith | Degeneration | Mild |
| | | Polyarteritis Nodosa | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 177

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402916

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Moderate

* Urinary Bladder

Polyarteritis Nodosa

Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 178

TRT#: 7

SEX: Male

DAY ON TEST: 434

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402917

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Epididymis | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Preputial Gland | * Prostate |
| * Salivary Glands | * Seminal Vesicle | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | Testis | * Thyroid Gland | * Trachea |
| * Urinary Bladder | | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-----------------------|---------------|------------------------------|---------------------|
| * Adrenal Cortex | Bilateral | Hyperplasia | Focal, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | Renal Tubule | Amphophilic/Vacuolar Adenoma | Multiple |
| | | Cyst | |
| | | Nephropathy | Chronicprogr, Mild |
| [Cyst TGLs = 1 - 8] | | | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 179

TRT#: 7

SEX: Male

DAY ON TEST: 672

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402918

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lung
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pancreas
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Skin
- * Stomach, Forestomach
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

MISSING

- * Thymus

OBSERVATIONS

- Blood Vessel Mineral Mild
- Note: The aorta and cardiac arteries are mineralized.
- * Bone Marrow Hemorrhage Marked
- Hypercellularity Mild
- * Heart Cardiomyopathy Mild
- * Kidney Renal Tubule Cyst
- Nephropathy Chronicprogr, Marked
- [Cyst TGLs = 4 - 8]
- [Nephropathy TGLs = 2,3 - 8]
- * Liver Necrosis Marked
- [Necrosis TGLs = 1 - 17]
- * Parathyroid Gland Hyperplasia Diffuse, Moderate
- * Pituitary Gland Pars Distalis Hyperplasia Moderate
- * Spleen White Pulp Atrophy Marked
- Pigment Mild
- * Stomach, Glandular Mineral Mild
- Testis Bilateral, Germinal Epith Degeneration Mild
- Polyarteritis Nodosa Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 180

TRT#: 7

SEX: Male

DAY ON TEST: 713

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402919

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Trachea
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach

OBSERVATIONS

- * Adrenal Cortex
 - Bilateral
 - Degeneration
 - Hyperplasia
 - Mineral
 - Cystic, Minimal
 - Focal, Minimal
 - Mild
- Blood Vessel
 - Note: The aorta and cardiac, gastric, pancreatic, renal, and salivary arteries are mineralized.
- * Bone Marrow
 - Hemorrhage
 - Hypercellularity
 - Atrophy
 - Marked
 - Mild
 - Mild
- * Epididymis
 - [Atrophy TGLs = 5 - 14]
- * Heart
 - Cardiomyopathy
- * Kidney
 - Inflammation
 - Nephropathy
 - Chronic, Minimal
 - Chronicprogr, Marked
- * Liver
 - [Nephropathy TGLs = 1 - 8]
 - Bile Duct
 - Hepatocellular Adenoma
 - Hepatodiaphragmatic Nodule
 - Hyperplasia
 - Minimal
 - [Hepatocellular Adenoma TGLs = 3 - 18]
 - [Hepatodiaphragmatic Nodule TGLs = 2 - 17]
- * Lung
 - Mineral
- * Parathyroid Gland
 - Hyperplasia
- * Pituitary Gland
 - Pars Distalis
 - Adenoma
- * Spleen
 - White Pulp
 - Atrophy
 - Pigment
 - Marked
 - Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 180

TRT#: 7

SEX: Male

DAY ON TEST: 713

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402919

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Glandular

Mineral

Mild

Testis

Bilateral, Germinal Epith

Degeneration

Moderate

Polyarteritis Nodosa

Mild

[Degeneration TGLs = 4 - 14]

* Thymus

Atrophy

Marked

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 181

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402920

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Epididymis
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Mammary Gland
- * Salivary Glands
- * Stomach, Glandular
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Nose
- * Seminal Vesicle
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Lymph Node, Mandibular
- * Preputial Gland
- * Skin
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|----------------------------|----------------|------------------------------|------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| | | Hyperplasia | Focal, Moderate |
| * Intestine Small, Jejunum | Peyers Patch | Hyperplasia | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Basophilic Focus | |
| | | Clear Cell Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Pancreas | Acinus | Atrophy | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Moderate |
| | | Polyarteritis Nodosa | Minimal |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 182

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402921

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| Blood Vessel | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Pancreas |
| * Preputial Gland | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | Testis | * Trachea |
| * Urinary Bladder | | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|----------------|------------------------------------------|------------------------------------|
| * Adrenal Cortex | Bilateral | Hyperplasia
Vacuolization Cytoplasmic | Focal, Minimal
Mild |
| * Adrenal Medulla | | Hyperplasia | Focal, Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Nose | Respirat Epith | Hyperplasia
Inflammation | Minimal
Chronic Active, Minimal |
| | Respirat Epith | Squamous Metaplasia | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| * Prostate | | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis
Pigment | Mild
Mild |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Adenoma | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 184

TRT#: 7

SEX: Male

DAY ON TEST: 650

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402923

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Cortex
 - Blood Vessel
 - Note: Renal arteries are mineralized.
- * Bone Marrow
 - Thrombus
 - Mineral
- * Eye
 - Cornea
 - Hemorrhage
 - Hypercellularity
 - Inflammation
- * Heart
 - Atrium
 - Thrombus
- * Kidney
 - [Thrombus TGLs = 3 - 10]
 - Nephropathy
- * Liver
 - [Nephropathy TGLs = 4 - 8]
 - Clear Cell Focus
 - Hepatocellular Adenoma
- * Lung
 - [Hepatocellular Adenoma TGLs = 2 - 18]
 - Infiltration Cellular
- * Spleen
 - White Pulp
 - Atrophy
 - Extramedullary Hematopoiesis
- Testis
 - Germinal Epith
 - Degeneration
 - Polyarteritis Nodosa
- * Thymus
 - Atrophy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 184

TRT#: 7

SEX: Male

DAY ON TEST: 650

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402923

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

CONTRIBUTORY CAUSE OF DEATH

- Heart Atrium Thrombus

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 185

TRT#: 7

SEX: Male

DAY ON TEST: 730

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402924

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|--------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Bone | * Bone Marrow | * Brain |
| * Epididymis | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Preputial Gland | * Salivary Glands | * Seminal Vesicle | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland | * Trachea |
| * Urinary Bladder | | | |

MISSING

- | | |
|-----------------|---------------------|
| * Mammary Gland | * Parathyroid Gland |
|-----------------|---------------------|

OBSERVATIONS

- | | | | |
|---------------------------|----------------|------------------------------|-------------------------|
| * Adrenal Medulla | Bilateral | Hyperplasia | Focal, Moderate |
| Blood Vessel | Carotid Artery | Polyarteritis Nodosa | Mild |
| * Heart | | Cardiomyopathy | Minimal |
| * Intestine Large, Rectum | | Parasite Metazoan | |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| * Lung | | Inflammation | Chronic Active, Minimal |
| * Nose | | Inflammation | Chronic Active, Minimal |
| * Pancreas | Acinus | Hyperplasia | Marked |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Prostate | | Inflammation | Chronic Active, Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Testis | Germinal Epith | Degeneration | Minimal |
| | | Polyarteritis Nodosa | Mild |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 186

TRT#: 7

SEX: Male

DAY ON TEST: 449

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402925

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Salivary Glands
- * Thyroid Gland
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Mammary Gland
- * Pituitary Gland
- * Seminal Vesicle
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Liver
- * Nose
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Prostate
- * Stomach, Glandular

MISSING

- * Intestine Small, Duodenum

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Heart
- * Kidney
- * Lung
- * Spleen
- * Stomach, Forestomach
- Testis
- * Thymus
- Atrium
- White Pulp
- Epithelium
- Germinal Epith
- Necrosis
- Hemorrhage
- Hypercellularity
- Cardiomyopathy
- Thrombus
- Nephropathy
- Infiltration Cellular
- Inflammation
- Atrophy
- Hyperplasia
- Degeneration
- Atrophy
- Minimal
- Marked
- Mild
- Mild
- Chronicprogr, Marked
- Histiocyte, Marked
- Acute, Mild
- Marked
- Minimal
- Minimal
- Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 187

TRT#: 7

SEX: Male

DAY ON TEST: 730

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402926

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Epididymis |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Preputial Gland | * Salivary Glands | * Seminal Vesicle |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | Testis |
| * Thyroid Gland | * Trachea | * Urinary Bladder | |

OBSERVATIONS

- | | | | |
|------------------------------|-----------------|------------------------------|------------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Adrenal Medulla | Bilateral | Hyperplasia | Focal, Moderate |
| * Bone Marrow | | Hypercellularity | Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| | | Hepatodiaphragmatic Nodule | |
| | Bile Duct | Hyperplasia | Minimal |
| | | Mixed Cell Focus | |
| [Hepatodiaphragmatic Nodule | TGLs = 1 - 17] | | |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Prostate | | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Hyperplasia | Lymphoid, Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 188

TRT#: 7

SEX: Male

DAY ON TEST: 643

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402927

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Epididymis
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Seminal Vesicle
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
[Carcinoma TGLs = 2 - 18]
- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- * Pituitary Gland
- * Spleen
[Hemangiosarcoma TGLs = 1 - 17]
- Testis
- * Thymus
- Carcinoma
- Hypercellularity
- Accumulation, Hyaline Droplet
- Infarct
- Nephropathy
- Extramedullary Hematopoiesis
- Carcinoma
- Hyperplasia
- Hemangiosarcoma
- Degeneration
- Atrophy
- Moderate
- Moderate
- Minimal
- Chronicprogr, Mild
- Mild
- Metastatic (Adrenal Cortex)
- Minimal
- Minimal
- Moderate

PRIMARY CAUSE OF DEATH - Spleen Hemangiosarcoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 189

TRT#: 7

SEX: Male

DAY ON TEST: 669

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402928

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Parathyroid Gland
- * Stomach, Forestomach
- Blood Vessel
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Preputial Gland
- * Stomach, Glandular
- * Epididymis
- * Intestine Small, Duodenum
- * Lymph Node, Mesenteric
- * Seminal Vesicle
- Testis
- * Esophagus
- * Intestine Small, Ileum
- * Mammary Gland
- * Skin

OBSERVATIONS

- * Adrenal Medulla
- * Bone
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Colon
- * Kidney
- [Leukemia Granulocytic TGLs = 7 - 8+23]
- * Liver
- * Lung
- Lymph Node
- [Leukemia Granulocytic TGLs = 3 - 19]
- [Leukemia Granulocytic TGLs = 6 - 22]
- [Leukemia Granulocytic TGLs = 5 - 21]
- [Leukemia Granulocytic TGLs = 4 - 20]
- * Lymph Node, Mandibular
- Pineal Gland
- Mediastinal
- Inguinal
- Renal
- Lumbar
- Hyperplasia
- Osteopetrosis
- Leukemia Granulocytic
- Necrosis
- Leukemia Granulocytic
- Leukemia Granulocytic
- Leukemia Granulocytic
- Cardiomyopathy
- Inflammation
- Leukemia Granulocytic
- Nephropathy
- Degeneration
- Leukemia Granulocytic
- Leukemia Granulocytic
- Leukemia Granulocytic
- Leukemia Granulocytic
- Leukemia Granulocytic
- Focal, Mild
- Moderate
- Moderate
- Minimal
- Chronic Active, Minimal
- Chronicprogr, Moderate
- Cystic, Minimal
- Leukemia Granulocytic

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 189

TRT#: 7

SEX: Male

DAY ON TEST: 669

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402928

ORGAN AND ACCOUNTABLE SITE STATUS

[Leukemia Granulocytic TGLs = 2 - 18]

* Nose		Leukemia Granulocytic	
* Pancreas	Duct	Cyst	
Peripheral Nerve	Axon, Trigeminal	Leukemia Granulocytic Degeneration	Minimal
Note: Sciatic and tibial nerves are normal.			
* Pituitary Gland		Leukemia Granulocytic	
* Prostate		Leukemia Granulocytic	
* Salivary Glands		Leukemia Granulocytic	
Skeletal Muscle		Leukemia Granulocytic	
Spinal Cord		Leukemia Granulocytic	
* Spleen		Extramedullary Hematopoiesis Leukemia Granulocytic Necrosis	Mild Moderate

[Leukemia Granulocytic TGLs = 1 - 17]

[Necrosis TGLs = 8 - 17]

* Thymus		Atrophy	Mild
		Leukemia Granulocytic	
* Thyroid Gland	C Cell	Adenoma	
* Trachea		Leukemia Granulocytic	
* Urinary Bladder		Leukemia Granulocytic	

PRIMARY CAUSE OF DEATH - Spleen Leukemia Granulocytic

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 190

TRT#: 7

SEX: Male

DAY ON TEST: 520

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402929

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Seminal Vesicle
- * Thyroid Gland

MISSING

- * Lymph Node, Mesenteric
- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Medulla [Pheochromocytoma Benign TGLs = 4 - 11] Bilateral Pheochromocytoma Benign
- * Bone Marrow Hemorrhage Mild
- * Heart Hypercellularity Mild
- * Kidney [Nephropathy TGLs = 2 - 8] Atrium Cardiomyopathy Mild
- * Liver Nephropathy Thrombus
- * Lung Extramedullary Hematopoiesis Chronicprogr, Marked
- * Spleen [Thrombus TGLs = 3 - 6+7] White Pulp Infiltration Cellular Minimal
- * Testis Bilateral, Germinal Epith Inflammation Histiocyte, Moderate
- * Thymus Thrombus Degeneration Granulomatous, Minimal
- Atrophy Marked
- Extramedullary Hematopoiesis Moderate
- Pigment Minimal
- Degeneration Moderate
- Atrophy Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 190

TRT#: 7

SEX: Male

DAY ON TEST: 520

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402929

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 191

TRT#: 7

SEX: Male

DAY ON TEST: 705

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402930

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Blood Vessel	* Bone	* Bone Marrow	* Brain
* Epididymis	* Esophagus	* Eye	* Harderian Gland
* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum	* Intestine Small, Duodenum
* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic	* Liver
* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Mammary Gland	* Parathyroid Gland
* Preputial Gland	* Prostate	* Salivary Glands	* Seminal Vesicle
* Stomach, Glandular	Testis	* Trachea	* Urinary Bladder

OBSERVATIONS

* Adrenal Cortex		Hyperplasia	Focal, Mild
* Adrenal Medulla	Bilateral	Pheochromocytoma Benign	
* Heart		Cardiomyopathy	Mild
* Kidney		Nephropathy	Chronicprogr, Moderate
* Lung		Inflammation	Granulomatous, Minimal
	[Inflammation TGLs = 3 - 7]		
* Nose	Nasolacrim Dct	Inflammation	Chronic, Minimal
* Pancreas	Acinus	Hyperplasia	Minimal
* Pituitary Gland	Pars Distalis	Hyperplasia	Mild
* Skin		Inflammation	Chronic Active, Mild
	[Ulcer TGLs = 1 - 17]	Ulcer	Mild
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
* Stomach, Forestomach	Epithelium	Hyperplasia	Basal Cell, Mild
* Thymus		Atrophy	Minimal
* Thyroid Gland	C Cell	Adenoma	

PRIMARY CAUSE OF DEATH - Adrenal Medulla Bilateral Pheochromocytoma Benign

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 193

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402932

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Moderate

* Thyroid Gland

C Cell

Hyperplasia

Marked

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 194

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402933

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Eye
- * Intestine Large, Rectum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Seminal Vesicle
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Esophagus
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Heart
- * Intestine Small, Ileum
- * Kidney
- [Nephropathy TGLs = 1 - 8+17]
- * Liver
- * Lung
- * Pancreas
- * Pituitary Gland
- * Spleen
- Testis
- [Adenoma TGLs = 3 - 14]
- * Thymus
- * Thyroid Gland
- Acinus
- Pars Distalis
- Interstitial Cell
- Bilateral, Germinal Epith
- Follicular Cel
- C Cell
- Hyperplasia
- Hyperplasia
- Pheochromocytoma Benign
- Hypercellularity
- Cardiomyopathy
- Polyarteritis Nodosa
- Nephropathy
- Eosinophilic Focus
- Carcinoma
- Inflammation
- Hyperplasia
- Polyarteritis Nodosa
- Hyperplasia
- Extramedullary Hematopoiesis
- Adenoma
- Degeneration
- Polyarteritis Nodosa
- Atrophy
- Adenoma
- Carcinoma
- Focal, Mild
- Focal, Mild
- Minimal
- Mild
- Minimal
- Chronicprogr, Marked
- Metastatic (Thyroid Gland)
- Granulomatous, Mild
- Minimal
- Minimal
- Mild
- Mild
- Moderate
- Mild
- Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 194

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402933

ORGAN AND ACCOUNTABLE SITE STATUS

[Carcinoma TGLs = 2 - 11]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 195

TRT#: 7

SEX: Male

DAY ON TEST: 554

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402934

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Prostate
- * Stomach, Forestomach
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Seminal Vesicle
- * Trachea
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: The aorta and cardiac, gastric, mammary, pancreatic, prostatic, pulmonary, renal, salivary, and seminal vesicular arteries are mineralized.
- * Bone Marrow
- * Heart
- * Intestine Large, Cecum
- * Kidney
[Nephropathy TGLs = 1 - 8]
- * Lung
- * Parathyroid Gland
- * Pituitary Gland
- * Spleen
- * Stomach, Glandular
Testis
- Hyperplasia
- Mineral
- Hemorrhage
- Hypercellularity
- Cardiomyopathy
- Mineral
- Inflammation
- Mineral
- Polyarteritis Nodosa
- Ulcer
- Nephropathy
- Mineral
- Hyperplasia
- Hyperplasia
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Mineral
- Degeneration
- Polyarteritis Nodosa
- Focal, Minimal
- Marked
- Mild
- Mild
- Moderate
- Minimal
- Chronic Active, Moderate
- Minimal
- Mild
- Moderate
- Chronicprogr, Marked
- Moderate
- Diffuse, Mild
- Moderate
- Mild
- Mild
- Minimal
- Moderate
- Minimal
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 195

TRT#: 7

SEX: Male

DAY ON TEST: 554

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402934

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus

Atrophy

Moderate

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 196

TRT#: 7

SEX: Male

DAY ON TEST: 653

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402935

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Lymph Node, Mesenteric
- * Prostate
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Mammary Gland
- * Salivary Glands
- * Stomach, Glandular
- * Brain
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lung
- * Nose
- * Seminal Vesicle
- * Thyroid Gland
- * Epididymis
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Eye
- * Heart
- * Kidney
- [Nephropathy TGLs = 2 - 8+17]
- * Pancreas
- * Parathyroid Gland
- * Preputial Gland
- Skeletal Muscle
- * Spleen
- Testis
- * Thymus
- Cornea
- Optic Nerve
- Duct
- Hyperplasia
- Hemorrhage
- Hypercellularity
- Inflammation
- Polyarteritis Nodosa
- Cardiomyopathy
- Cyst
- Nephropathy
- Polyarteritis Nodosa
- Hyperplasia
- Hyperplasia
- Polyarteritis Nodosa
- Extramedullary Hematopoiesis
- Pigment
- Polyarteritis Nodosa
- Atrophy
- Focal, Minimal
- Minimal
- Mild
- Acute, Minimal
- Minimal
- Minimal
- Chronicprogr, Marked
- Minimal
- Diffuse, Moderate
- Squamous, Mild
- Minimal
- Mild
- Minimal
- Mild
- Mild

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 197

TRT#: 7

SEX: Male

DAY ON TEST: 614

DOSE: 750 mg/kg male

DISP: Natural Death

HISTO: 1402936

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Seminal Vesicle
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Preputial Gland
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Medulla
- Blood Vessel
- * Bone Marrow
- * Heart
- * Intestine Large, Colon
- * Kidney
 - [Nephropathy TGLs = 2 - 8]
- * Lung
 - [Infiltration Cellular TGLs = 3 - 6+7]
- * Pancreas
- * Parathyroid Gland
- * Prostate
- * Skin
 - [Inflammation TGLs = 1 - 17]
- * Spleen
- Testis
- * Thymus
- Pulmn Artery
- Pheochromocytoma Benign
- Thrombus
- Hypercellularity
- Cardiomyopathy
- Polyarteritis Nodosa
- Nephropathy
- Infiltration Cellular
- Polyarteritis Nodosa
- Hyperplasia
- Inflammation
- Inflammation
- Extramedullary Hematopoiesis
- Pigment
- Polyarteritis Nodosa
- Atrophy
- Moderate
- Mild
- Minimal
- Chronicprogr, Marked
- Histiocyte, Minimal
- Mild
- Diffuse, Minimal
- Chronic Active, Minimal
- Chronic Active, Marked
- Mild
- Minimal
- Mild
- Marked

PRIMARY CAUSE OF DEATH - Kidney Nephropathy

CONTRIBUTORY CAUSE OF DEATH - Skin Inflammation

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 198

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402937

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Salivary Glands
- * Stomach, Glandular
- * Urinary Bladder
- Blood Vessel
- * Epididymis
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Seminal Vesicle
- Testis
- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Preputial Gland
- * Skin
- * Thymus
- * Bone Marrow
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Prostate
- * Stomach, Forestomach
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Heart
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Nose
- * Spleen
- * Thyroid Gland
- Hyperplasia
- Cardiomyopathy
- Nephropathy
- Clear Cell Focus
- Infiltration Cellular
- Inflammation
- Extramedullary Hematopoiesis
- Pigment
- Adenoma
- Focal, Minimal
- Minimal
- Chronicprogr, Moderate
- Histiocyte, Minimal
- Chronic Active, Minimal
- Mild
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 199

TRT#: 7

SEX: Male

DAY ON TEST: 674

DOSE: 750 mg/kg male

DISP: Moribund Sacrifice

HISTO: 1402938

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Bone Marrow
- * Brain
- * Epididymis
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Pancreas
- * Parathyroid Gland
- * Pituitary Gland
- * Preputial Gland
- * Prostate
- * Salivary Glands
- * Seminal Vesicle
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea
- * Urinary Bladder

OBSERVATIONS

- * Kidney [Nephropathy TGLs = 2 - 8] Nephropathy Chronicprogr, Mild
- * Liver Bile Duct Hyperplasia Minimal
- * Lung [Infiltration Cellular TGLs = 3 - 6] Infiltration Cellular Histiocyte, Minimal
- * Nose Olfactory Epi Degeneration Mild
- Respirat Epith Hyperplasia Mild
- * Skin Inflammation Inflammation Suppurative, Marked
- [Inflammation TGLs = 1 - 17] Inflammation Chronic Active, Marked
- * Spleen Extramedullary Hematopoiesis Mild
- Pigment Minimal
- * Stomach, Forestomach Epithelium Hyperplasia Mild
- Testis Germinal Epith Degeneration Minimal
- * Thymus Atrophy Marked

PRIMARY CAUSE OF DEATH - Skin Inflammation

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 200

TRT#: 7

SEX: Male

DAY ON TEST: 731

DOSE: 750 mg/kg male

DISP: Terminal Sacrifice

HISTO: 1402939

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Cortex	* Adrenal Medulla	Blood Vessel	* Bone
* Brain	* Epididymis	* Esophagus	* Eye
* Harderian Gland	* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum
* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic
* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Mammary Gland	* Nose
* Parathyroid Gland	* Preputial Gland	* Prostate	* Salivary Glands
* Seminal Vesicle	* Skin	* Stomach, Forestomach	* Stomach, Glandular
* Thymus	* Trachea	* Urinary Bladder	

OBSERVATIONS

* Bone Marrow		Hypercellularity	Minimal
* Heart		Cardiomyopathy	Minimal
* Kidney		Nephropathy	Chronicprogr, Moderate
* Liver		Clear Cell Focus	
* Lung		Infiltration Cellular	Histiocyte, Minimal
		Inflammation	Granulomatous, Minimal
	[Inflammation TGLs = 2 - 6+7]		
* Pancreas	Acinus	Adenoma	Multiple
	Acinus	Atrophy	Mild
	[Adenoma TGLs = 1 - 17]		
* Pituitary Gland	Pars Distalis	Hyperplasia	Minimal
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Mild
Testis	Interstit Cell	Hyperplasia	Mild
* Thyroid Gland	C Cell	Hyperplasia	Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 201

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402960

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Nose
- * Pancreas
- * Skin
- * Stomach, Forestomach
- * Urinary Bladder
- Vagina
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Kidney Nephropathy Chronicprogr, Minimal
- * Liver Clear Cell Focus
- * Lung Infiltration Cellular Histiocyte, Mild
 - [Infiltration Cellular TGLs = 3 - 6+7]
- * Mammary Gland Hyperplasia Atypical, Moderate
- * Ovary Atrophy Mild
 - Follicle
 - Cyst
 - Pars Distalis Adenoma
- * Pituitary Gland Adenoma
 - [Adenoma TGLs = 4 - 11]
- * Spleen Extramedullary Hematopoiesis Mild
- * Thymus Pigment Minimal
- * Thyroid Gland Atrophy Minimal
- * Uterus Adenoma Cystic, Marked
 - Bilateral, C Cell
 - Endometrium Hyperplasia Atypical, Minimal
 - Hyperplasia Multiple
 - Polyp Stromal Multiple
 - Squamous Metaplasia Minimal
- [Polyp Stromal TGLs = 1,2 - 16+17]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 202

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402961

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- Vagina
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
- * Kidney
- * Liver
- * Lung
 - [Infiltration Cellular TGLs = 2 - 6+7]
- * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Thyroid Gland
- * Uterus
- Pars Distalis
- C Cell
- C Cell
- Endometrium
- Angiectasis
- Nephropathy
- Clear Cell Focus
- Infiltration Cellular
- Fibroadenoma
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Adenoma
- Hyperplasia
- Hyperplasia
- Squamous Metaplasia
- Mild
- Chronicprogr, Minimal
- Histiocyte, Mild
- Mild
- Moderate
- Mild
- Minimal
- Minimal
- Mild
- Cystic, Minimal
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 203

TRT#: 2

SEX: Female

DAY ON TEST: 578

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1402962

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Kidney | * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------------------|------------------------------|----------------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Heart | | Cardiomyopathy | Minimal |
| * Lung | | Adenocarcinoma | Metastatic (Mammary Gland) |
| | | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | | Adenocarcinoma | Multiple |
| | | Fibroadenoma | |
| | [Adenocarcinoma TGLs = 2,3,4 - 15+19+20] | | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Ovary | Follicle | Atrophy | Moderate |
| | Pars Distalis | Cyst | |
| * Pituitary Gland | | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Minimal |
| * Uterus | | Squamous Metaplasia | Mild |

PRIMARY CAUSE OF DEATH - Mammary Gland Adenocarcinoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 204

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402963

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Bone Marrow
* Brain	* Clitoral Gland	* Esophagus	* Eye
* Harderian Gland	* Heart	* Intestine Large, Cecum	* Intestine Large, Colon
* Intestine Large, Rectum	* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum
* Islets, Pancreatic	* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Mammary Gland
* Parathyroid Gland	* Salivary Glands	* Skin	* Stomach, Forestomach
* Stomach, Glandular	* Thyroid Gland	* Trachea	* Urinary Bladder
* Uterus	Vagina		

OBSERVATIONS

* Adrenal Cortex	Bilateral	Degeneration	Cystic, Minimal
* Kidney		Nephropathy	Chronicprogr, Minimal
* Liver		Clear Cell Focus	
* Lung		Infiltration Cellular	Histiocyte, Minimal
* Nose	Nasolacrim Dct	Inflammation	Chronic, Minimal
* Ovary	Follicle	Cyst	Multiple
[Cyst TGLs = 1 - 17]			
* Pancreas	Acinus	Atrophy	Minimal
* Pituitary Gland	Pars Distalis	Hyperplasia	Minimal
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
* Thymus		Atrophy	Minimal

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 205

TRT#: 2

SEX: Female

DAY ON TEST: 499

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1402964

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- * Uterus
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Liver
- [Angiectasis TGLs = 3 - 19]
- * Lung
- * Mammary Gland
- Note: Some of TGL-1 is also in B15.
- [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 2 - 11]
- * Spleen
- * Thymus
- Renal Tubule
- Follicle
- Pars Distalis
- Hyperplasia
- Hypercellularity
- Accumulation, Hyaline Droplet
- Angiectasis
- Extramedullary Hematopoiesis
- Infiltration Cellular
- Fibroadenoma
- Cyst
- Adenoma
- Extramedullary Hematopoiesis
- Atrophy
- Focal, Minimal
- Marked
- Mild
- Mild
- Minimal
- Histiocyte, Mild
- Moderate
- Mild

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 206

TRT#: 2

SEX: Female

DAY ON TEST: 606

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1402965

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Salivary Glands | * Skin | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

MISSING

- | | |
|------------------|----------|
| * Clitoral Gland | * Thymus |
|------------------|----------|

OBSERVATIONS

- | | | | |
|------------------------|------------------------------------------|----------------------------------------------|----------------------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | Renal Tubule | Accumulation, Hyaline Droplet
Nephropathy | Minimal
Chronicprogr, Minimal |
| * Liver | | Necrosis | Mild |
| * Lung | [Infiltration Cellular TGLs = 3 - 6+7] | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | [Fibroadenoma TGLs = 1,2 - 18+19] | Fibroadenoma | Multiple |
| * Nose | Nasolacrim Dct | Inflammation | Chronic, Minimal |
| * Ovary | | Atrophy | Mild |
| * Spleen | White Pulp | Atrophy | Minimal |
| | | Extramedullary Hematopoiesis | Moderate |
| * Stomach, Forestomach | | Inflammation | Chronic, Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 210

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402969

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Ovary | * Parathyroid Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Trachea | * Urinary Bladder |
| * Uterus | Vagina | | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|--------------------|
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | [Clear Cell Focus TGLs = 1 - 12] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Mammary Gland | | Hyperplasia | Mild |
| * Pancreas | | Polyarteritis Nodosa | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Hyperplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 211

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402970

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver |
| * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Trachea | * Urinary Bladder | * Uterus | Vagina |

OBSERVATIONS

- | | | | |
|-------------------|---------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Mild |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hemorrhage | Mild |
| | | Hypercellularity | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Mammary Gland | | Hyperplasia | Moderate |
| * Nose | | Inflammation | Chronic, Minimal |
| * Ovary | | Atrophy | Mild |
| * Pancreas | Acinus | Atrophy | Minimal |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 1 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Adenoma | |
| | C Cell | Hyperplasia | Mild |

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 212

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402971

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Uterus
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Pituitary Gland
- * Stomach, Glandular
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Ovary
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 1 - 6]
- * Mammary Gland
- * Spleen
- * Thymus
- * Thyroid Gland
- Bile Duct
- C Cell
- Pheochromocytoma Benign
- Hemorrhage
- Hypercellularity
- Nephropathy
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Fibroadenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Adenoma
- Minimal
- Minimal
- Chronicprogr, Minimal
- Mild
- Histiocyte, Minimal
- Granulomatous, Minimal
- Mild
- Mild
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 213

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402972

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thymus
- * Bone
- * Eye
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Bone Marrow
 - * Intestine Large, Rectum
 - * Kidney
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 2 - 6+7]
 - * Ovary
 - * Pituitary Gland
 - * Spleen
 - * Thyroid Gland
 - * Uterus
 - Bile Duct
 - Pars Distalis
 - C Cell
 - Endometrium
 - Cervix
 - Hypercellularity
 - Parasite Metazoan
 - Nephropathy
 - Hyperplasia
 - Infiltration Cellular
 - Atrophy
 - Adenoma
 - Extramedullary Hematopoiesis
 - Pigment
 - Hyperplasia
 - Hyperplasia
 - Hyperplasia
 - Polyp Stromal
 - Squamous Metaplasia
 - Thrombus
 - Moderate
 - Chronicprogr, Minimal
 - Minimal
 - Histiocyte, Moderate
 - Mild
 - Mild
 - Minimal
 - Minimal
 - Cystic, Mild
 - Stromal, Marked
 - Minimal
- [Hyperplasia TGLs = 1-16]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 214

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402973

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland | * Trachea |
| * Urinary Bladder | * Uterus | Vagina | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Mammary Gland | | Hyperplasia | Mild |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 2 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 215

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402974

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Brain
- * Clitoral Gland
- * Harderian Gland
- * Heart
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Nose
- * Pancreas
- * Skin
- * Stomach, Forestomach
- * Urinary Bladder
- * Vagina
- * Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Bone Marrow
 - Hemorrhage
 - Mild
 - Hypercellularity
 - Minimal
- * Kidney
 - Nephropathy
 - Chronicprogr, Mild
- * Liver
 - Clear Cell Focus
- * Lung
 - Bile Duct
 - Hyperplasia
 - Minimal
 - Infiltration Cellular
 - Histiocyte, Marked
 - Inflammation
 - Granulomatous, Minimal
- [Infiltration Cellular TGLs = 3 - 6+7]
- * Ovary
 - Atrophy
 - Moderate
 - Cyst
- [Cyst TGLs = 1 - 14]
- * Pituitary Gland
 - Periovarn Tiss
 - Hyperplasia
 - Mild
- * Spleen
 - Pars Distalis
 - Extramedullary Hematopoiesis
 - Mild
 - Pigment
 - Minimal
- * Thymus
 - Atrophy
 - Mild
- * Thyroid Gland
 - C Cell
 - Hyperplasia
 - Moderate
- * Uterus
 - Endometrium
 - Adenocarcinoma
 - Hyperplasia
 - Cystic, Mild
 - Inflammation
 - Chronic Active, Minimal
 - Squamous Metaplasia
 - Minimal
- [Hyperplasia TGLs = 2 - 18]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 216

TRT#: 2

SEX: Female

DAY ON TEST: 600

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1402975

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Ovary | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | * Uterus |
| Vagina | | | |

OBSERVATIONS

- | | | |
|--------------------------------|------------------------------|-----------------------|
| * Bone Marrow | Hypercellularity | Marked |
| * Kidney | Nephropathy | Chronicprogr, Minimal |
| * Liver | Extramedullary Hematopoiesis | Mild |
| * Lung | Infiltration Cellular | Histiocyte, Mild |
| * Mammary Gland | Fibroadenoma | |
| [Fibroadenoma TGLs = 1 - 18] | | |
| * Spleen | Extramedullary Hematopoiesis | Moderate |
| * Thymus | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 217

TRT#: 2

SEX: Female

DAY ON TEST: 504

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1402976

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- Vagina
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Kidney
- * Ovary
- * Skin
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lung
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Stomach, Glandular
- * Uterus

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Bone Marrow
- * Liver
- * Mammary Gland
- [Fibroadenoma TGLs = 1 - 18]
- * Nose
- * Spleen
- * Thymus
- Hypercellularity
- Necrosis
- Fibroadenoma
- Hyperplasia
- Inflammation
- Inflammation
- Atrophy
- Extramedullary Hematopoiesis
- Atrophy
- Marked
- Mild
- Minimal
- Chronic, Minimal
- Acute, Mild
- Mild
- Moderate
- Moderate

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 218

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402977

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Trachea |
| * Urinary Bladder | Vagina | | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Clear Cell Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Hyperplasia | Marked |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Inflammation | Acute, Minimal |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 219

TRT#: 2

SEX: Female

DAY ON TEST: 536

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1402978

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Parathyroid Gland
- * Salivary Glands
- * Stomach, Glandular
- * Trachea
- * Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Skin
- * Urinary Bladder
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Uterus

OBSERVATIONS

- * Kidney
- * Lung
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Thyroid Gland
- Nephropathy
- Infiltration Cellular
- Inflammation
- Atrophy
- Cyst
- Adenoma
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Hyperplasia
- Chronicprogr, Minimal
- Histiocyte, Minimal
- Granulomatous, Minimal
- Minimal
- Moderate
- Minimal
- Mild
- Mild
- Mild

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 220

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402979

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Stomach, Forestomach
- * Urinary Bladder
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Skin
- * Trachea

OBSERVATIONS

- * Bone Marrow
 - Hemorrhage
 - Hypercellularity
 - Nephropathy
 - Clear Cell Focus
 - Cyst
- * Kidney
- * Liver
 - Bile Duct
 - [Cyst TGLs = 1 - 18]
- * Lung
 - Infiltration Cellular
 - [Infiltration Cellular TGLs = 2 - 6+7]
- * Ovary
- * Pancreas
- * Pituitary Gland
 - Pars Distalis
 - [Adenoma TGLs = 4 - 11]
- * Spleen
 - Extramedullary Hematopoiesis
 - Pigment
- * Thymus
 - Atrophy
- * Uterus
 - Hyperplasia
 - Papilloma
 - Squamous Metaplasia

Note: Squamous cell papillomas. One as TGL in B19, another in routine section B16.

[Papilloma TGLs = 3 - 19]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 222

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402981

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose | * Pancreas |
| * Parathyroid Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|-------------------|-------------------------------------------|------------------------------|-------------------------|
| * Adrenal Medulla | | Pheochromocytoma Benign | |
| | [Pheochromocytoma Benign TGLs = 1 - 11] | | |
| * Bone Marrow | | Hemorrhage | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | | Eosinophilic Focus | |
| * Lung | | Inflammation | Chronic Active, Minimal |
| * Mammary Gland | | Hyperplasia | Mild |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Adenoma | |
| | [Adenoma TGLs = 2 - 11] | | |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 223

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402982

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|------------------|--------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Hypertrophy | Focal, Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Clear Cell Focus | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 2 - 18] | | |
| * Ovary | | Atrophy | Mild |
| | Follicle | Cyst | |
| | [Cyst TGLs = 1 - 14] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| * Uterus | | Polyp Stromal | |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 224

TRT#: 2

SEX: Female

DAY ON TEST: 677

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1402983

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lung
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Kidney
- Lacrimal Gland
- * Liver
- * Mammary Gland
- [Fibroadenoma TGLs = 1,2 - 18+19]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- Bile Duct
- Pars Distalis
- White Pulp
- Pheochromocytoma Benign
- Hypercellularity
- Nephropathy
- Inflammation
- Basophilic Focus
- Hyperplasia
- Fibroadenoma
- Atrophy
- Hyperplasia
- Atrophy
- Extramedullary Hematopoiesis
- Atrophy
- Dilation
- Hemorrhage
- Inflammation
- Moderate
- Chronicprogr, Minimal
- Granulomatous, Moderate
- Minimal
- Multiple
- Mild
- Mild
- Mild
- Mild
- Moderate
- Marked
- Moderate
- Chronic Active, Mild

[Dilation TGLs = 3 - 20]

PRIMARY CAUSE OF DEATH - Uterus Hemorrhage

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 225

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402984

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Thyroid Gland | * Trachea |
| * Urinary Bladder | * Uterus | Vagina | |

OBSERVATIONS

- | | | | |
|----------------------|---------------|------------------------------|-----------------------|
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | | Hyperplasia | Mild |
| * Ovary | | Atrophy | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Stomach, Glandular | | Mineral | Minimal |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 226

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402985

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Cortex	Blood Vessel	* Bone	* Bone Marrow
* Brain	* Clitoral Gland	* Esophagus	* Eye
* Harderian Gland	* Heart	* Intestine Large, Cecum	* Intestine Large, Colon
* Intestine Large, Rectum	* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum
* Islets, Pancreatic	* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Pancreas
* Parathyroid Gland	* Salivary Glands	* Skin	* Stomach, Forestomach
* Stomach, Glandular	* Thymus	* Thyroid Gland	* Trachea
* Urinary Bladder	Vagina		

OBSERVATIONS

* Adrenal Medulla		Hyperplasia	Focal, Minimal
* Kidney		Nephropathy	Chronicprogr, Minimal
* Liver		Clear Cell Focus	
* Lung	Alveolar Epith	Hyperplasia	Mild
	[Infiltration Cellular TGLs = 2 - 6+7]	Infiltration Cellular	Histiocyte, Mild
* Mammary Gland		Fibroadenoma	
	[Fibroadenoma TGLs = 1 - 18]		
* Nose	Nasolacrim Dct	Inflammation	Chronic, Minimal
* Ovary		Atrophy	Mild
* Pituitary Gland	Pars Distalis	Hyperplasia	Marked
* Spleen		Extramedullary Hematopoiesis	Moderate
		Pigment	Mild
* Uterus	Endometrium	Hyperplasia	Cystic, Minimal
		Squamous Metaplasia	Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 227

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402986

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|--------------------------------------------------------------------------------------------|------------------------------------------|------------------------------|------------------------|
| * Heart | | Cardiomyopathy | Minimal |
| * Int Lg Rectum | | | |
| Note: There is mammary tissue in the retroperitoneum surrounding the rectal section (B09). | | | |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Eosinophilic Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Minimal |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 1 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Uterus | | Polyp Stromal | |
| | | Squamous Metaplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 229

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402988

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Lung
- [Infiltration Cellular TGLs = 3 - 6+7]
- * Mammary Gland
- [Fibroadenoma TGLs = 1 - 19]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- Angiectasis
- Thrombus
- Hypercellularity
- Nephropathy
- Infiltration Cellular
- Inflammation
- Fibroadenoma
- Atrophy
- Cyst
- Hyperplasia
- Extramedullary Hematopoiesis
- Atrophy
- Inflammation
- Polyp Stromal
- Squamous Metaplasia
- Mild
- Moderate
- Chronicprogr, Minimal
- Histiocyte, Mild
- Granulomatous, Minimal
- Mild
- Minimal
- Mild
- Minimal
- Acute, Mild
- Multiple
- Mild

[Polyp Stromal TGLs = 2 - 17]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 230

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402989

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
 - Adenoma
 - Angiectasis
 - Degeneration
 - Hypercellularity
 - Nephropathy
 - Clear Cell Focus
 - Eosinophilic Focus
 - Infiltration Cellular
- * Bone Marrow
 - Mild
 - Cystic, Minimal
- * Kidney
 - Mild
 - Chronicprogr, Minimal
- * Liver
 - Chronicprogr, Minimal
- * Lung
 - [Infiltration Cellular TGLs = 3 - 6+7]
 - Histiocyte, Mild
- * Mammary Gland
 - [Fibroadenoma TGLs = 1,2 - 19+20]
 - Fibroadenoma
 - Multiple
- * Ovary
 - Atrophy
 - Mild
- * Pituitary Gland
 - [Adenoma TGLs = 5 - 11]
 - Pars Distalis
 - Adenoma
- * Spleen
 - Extramedullary Hematopoiesis
 - Mild
 - Pigment
 - Minimal
- * Thymus
 - Atrophy
 - Minimal
- * Uterus
 - Endometrium
 - Hyperplasia
 - Cystic, Mild
 - Cervix
 - Hyperplasia
 - Squamous, Minimal
 - Polyp Stromal
 - Squamous Metaplasia
 - Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 230

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402989

ORGAN AND ACCOUNTABLE SITE STATUS

[Polyp Stromal TGLs = 4 - 18]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 232

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402991

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Pancreas | * Parathyroid Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Trachea | * Urinary Bladder |
| Vagina | | | |

OBSERVATIONS

- | | | | |
|-------------------|-------------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Adenoma | |
| | | Infiltration Cellular | Lipocyte, Mild |
| | | Metaplasia | Osseous, Minimal |
| | [Adenoma TGLs = 3 - 18] | | |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Eosinophilic Focus | |
| * Lung | Bronchiole | Hyperplasia | Minimal |
| | | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | | Fibroadenoma | Multiple |
| | [Fibroadenoma TGLs = 1,2 - 19+20] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Hyperplasia | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Mild |
| | Bilateral | Polyp Stromal | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 233

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402992

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Ovary | * Pancreas |
| * Parathyroid Gland | * Salivary Glands | * Stomach, Glandular | * Trachea |
| * Urinary Bladder | * Uterus | Vagina | |

OBSERVATIONS

- | | | | |
|------------------------|--------------------------------|------------------------------|---------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Acute, Minimal |
| * Mammary Gland | | Fibroadenoma | |
| | | Hyperplasia | Mild |
| | [Fibroadenoma TGLs = 2 - 19] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Skin | | Lipoma | |
| | [Lipoma TGLs = 1 - 18] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Stomach, Forestomach | Epithelium | Hyperplasia | Moderate |
| | | Inflammation | Acute, Mild |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Hyperplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 234

TRT#: 2

SEX: Female

DAY ON TEST: 655

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1402993

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Skin
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Forestomach
- * Uterus
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Stomach, Glandular
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Lung
 - [Infiltration Cellular TGLs = 3 - 6+7]
- * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Spleen
 - [Extramedullary Hematopoiesis TGLs = 2 - 8]
- * Thymus
- * Thyroid Gland
- Degeneration
- Hypercellularity
- Nephropathy
- Infiltration Cellular
- Fibroadenoma
- Cyst
- Extramedullary Hematopoiesis
- Atrophy
- Adenoma
- Cystic, Minimal
- Marked
- Chronicprogr, Minimal
- Histiocyte, Mild
- Marked
- Mild

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 235

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402994

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Trachea |
| * Urinary Bladder | Vagina | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|---------------------------------------|------------------------------|-----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Clitoral Gland | | Squamous Metaplasia | Moderate |
| | [Squamous Metaplasia TGLs = 1 - 15] | | |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 2 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Adenoma | |
| | C Cell | Hyperplasia | Moderate |
| * Uterus | Endometrium | Hyperplasia | Cystic, Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 236

TRT#: 2

SEX: Female

DAY ON TEST: 715

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1402995

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Brain
* Clitoral Gland	* Esophagus	* Eye	* Harderian Gland
* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum	* Intestine Small, Duodenum
* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic	* Lymph Node, Mandibular
* Lymph Node, Mesenteric	* Nose	* Pancreas	* Parathyroid Gland
* Salivary Glands	* Skin	* Stomach, Forestomach	* Stomach, Glandular
* Thyroid Gland	* Trachea	* Urinary Bladder	Vagina

OBSERVATIONS

* Adrenal Cortex		Angiectasis	Moderate
	Bilateral	Degeneration	Cystic, Minimal
	[Angiectasis TGLs = 1 - 18]		
* Bone Marrow		Hemorrhage	Minimal
		Hypercellularity	Moderate
* Heart		Cardiomyopathy	Minimal
* Kidney		Nephropathy	Chronicprogr, Mild
* Liver		Eosinophilic Focus	
	Hepatocyte	Hyperplasia	Mild
	Oval Cell	Hyperplasia	Minimal
	Bile Duct	Hyperplasia	Minimal
* Lung		Infiltration Cellular	Histiocyte, Minimal
	[Infiltration Cellular TGLs = 3,4 - 6+7]		
* Mammary Gland		Fibroadenoma	
* Ovary		Atrophy	Mild
* Pituitary Gland	Pars Distalis	Adenoma	
	[Adenoma TGLs = 2 - 19]		
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Mild
* Thymus		Atrophy	Minimal
* Uterus	Endometrium	Hyperplasia	Cystic, Mild
	Cervix	Hyperplasia	Squamous, Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 236

TRT#: 2

SEX: Female

DAY ON TEST: 715

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1402995

ORGAN AND ACCOUNTABLE SITE STATUS

Squamous Metaplasia

Minimal

PRIMARY CAUSE OF DEATH

- Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 239

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402998

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thymus | * Thyroid Gland | * Trachea | * Urinary Bladder |
| Vagina | | | |

OBSERVATIONS

- | | | | |
|-------------------|-------------------------------|------------------------------|------------------------|
| * Bone Marrow | | Hypercellularity | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | | Inflammation | Granulomatous, Minimal |
| | [Inflammation TGLs = 1 - 6] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Uterus | | Polyp Stromal | Multiple |
| | | Squamous Metaplasia | Marked |

Note: TGLS = 2 - NCL

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 240

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1402999

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|----------------------|------------------------------------------|------------------------------|---------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Minimal |
| * Islets, Pancreatic | | Hyperplasia | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Mammary Gland | | Adenocarcinoma | |
| | | Fibroadenoma | |
| | Note: Fibroadenoma in routine section. | | |
| | [Adenocarcinoma TGLs = 1 - 18] | | |
| * Ovary | | Atrophy | Moderate |
| | Rete Ovarii | Cyst | |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 3 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Mild |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 241

TRT#: 2

SEX: Female

DAY ON TEST: 670

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1403000

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 4 - 6+7]
- * Mammary Gland
- [Fibroadenoma TGLs = 1 - 20]
- Mesentery
- Bile Duct
- Degeneration
- Hypercellularity
- Nephropathy
- Hyperplasia
- Infiltration Cellular
- Fibroadenoma
- Adenocarcinoma
- Hemorrhage
- Cystic, Minimal
- Moderate
- Chronicprogr, Minimal
- Minimal
- Histiocyte, Mild
- Metastatic (Uterus)
- Moderate

Note: The crystals in B21 are either formed from zymogen, or hemoglobin, or both.

Note: The cause of TGL-6 nodules was considered hemorrhage and/or rupture of the pancreatic duct, probably secondary to metastases of uterine carcinoma (coded).

[Hemorrhage TGLs = 6 - 21]

- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 5 - 11]
- * Spleen
- * Stomach, Forestomach
- * Stomach, Glandular
- * Thymus
- * Uterus
- Pars Distalis
- Endometrium
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Mineral
- Mineral
- Atrophy
- Adenocarcinoma
- Dilation
- Mild
- Moderate
- Minimal
- Minimal
- Mild
- Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 241

TRT#: 2

SEX: Female

DAY ON TEST: 670

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1403000

ORGAN AND ACCOUNTABLE SITE STATUS

Squamous Metaplasia

Minimal

[Adenocarcinoma TGLs = 3 - 18]

[Dilation TGLs = 2 - 19]

PRIMARY CAUSE OF DEATH

- Uterus Endometrium Adenocarcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 242

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403001

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina

OBSERVATIONS

- * Adrenal Cortex
 - Bilateral
 - Angiectasis
 - Hyperplasia
 - Thrombus
- * Bone Marrow
 - Hypercellularity
- * Heart
 - Schwannoma Malignant
- * Kidney
 - Nephropathy
- * Liver
 - Basophilic Focus
 - Clear Cell Focus
 - Eosinophilic Focus
 - Hepatocellular Adenoma
- Note: TGL-5 was in B12 (eosinophilic focus) rather than B23 as suggested at trim.
 - [Eosinophilic Focus TGLs = 5 - 12+23]
 - [Hepatocellular Adenoma TGLs = 4 - 22]
- * Lung
 - [Infiltration Cellular TGLs = 7 - 6+7]
 - Infiltration Cellular
- * Mammary Gland
 - [Fibroadenoma TGLs = 1,2,3,6 - 18+19+20+21]
 - Fibroadenoma
- * Ovary
 - Atrophy
- * Pituitary Gland
 - Pars Distalis
 - Hyperplasia
- * Spleen
 - Extramedullary Hematopoiesis
 - Pigment
- * Thymus
 - Atrophy

Minimal
Focal, Mild
Mild
Chronicprogr, Minimal
Histiocyte, Moderate
Multiple
Moderate
Marked
Mild
Minimal
Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 242

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403001

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Endometrium

Hyperplasia

Cystic, Minimal

Squamous Cell Carcinoma

Squamous Metaplasia

Moderate

Note: Squamous cell carcinoma in situ.

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 243

TRT#: 2

SEX: Female

DAY ON TEST: 719

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1403002

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- Vagina
- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- Blood Vessel
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Trachea
- * Bone
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Bone Marrow
 - Hemorrhage
 - Hypercellularity
- * Clitoral Gland
 - Squamous Cell Carcinoma

Note: Squamous cell carcinoma of the clitoral gland, one tumor incorporating both glands.
 [Squamous Cell Carcinoma TGLs = 1 - 18]
- * Kidney
 - Pelvis
 - Dilation
 - Nephropathy
- * Liver
 - Basophilic Focus
- * Lung
 - Infiltration Cellular
- * Ovary
 - Atrophy
- * Pituitary Gland
 - Pars Distalis
 - Adenoma

[Adenoma TGLs = 2 - 11]
- * Spleen
 - White Pulp
 - Atrophy
 - Pigment
- * Thymus
 - Atrophy
- * Thyroid Gland
 - C Cell
 - Adenoma

[Adenoma TGLs = 3 - 11]
- * Uterus
 - Inflammation

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 244

TRT#: 2

SEX: Female

DAY ON TEST: 681

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403003

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Liver
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Trachea
- * Urinary Bladder
- Vagina

OBSERVATIONS

- * Bone Marrow Hemorrhage Minimal
- * Islets, Pancreatic Hyperplasia Minimal
- * Kidney Nephropathy Chronicprogr, Minimal
- * Lung Alveolar Epith Hyperplasia Minimal
[Hyperplasia TGLs = 2 - 6+7]
- * Ovary Atrophy Mild
- * Pituitary Gland Pars Distalis Adenoma
[Adenoma TGLs = 1 - 11]
- * Spleen Extramedullary Hematopoiesis Mild
- * Thymus Pigment Minimal
- * Thyroid Gland Atrophy Minimal
- * Uterus C Cell Hyperplasia Moderate
- Endometrium Hyperplasia Cystic, Mild
- Squamous Metaplasia Mild
[Hyperplasia TGLs = 3 - 17]

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 245

TRT#: 2

SEX: Female

DAY ON TEST: 729

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403004

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Lung
- * Nose
- * Pancreas
- * Skin
- * Stomach, Forestomach
- * Trachea
- * Urinary Bladder
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland

OBSERVATIONS

- * Islets, Pancreatic
 - [Adenoma TGLs = 3 - 20]
- * Kidney
 - Adenoma
 - Nephropathy
 - Clear Cell Focus
 - Hepatocellular Adenoma
 - Chronicprogr, Minimal
- * Liver
 - [Hepatocellular Adenoma TGLs = 2 - 19]
- * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 18]
 - Fibroadenoma
 - Hyperplasia
 - Mild
- * Ovary
 - Follicle
 - Cyst
- * Pituitary Gland
 - Pars Distalis
 - Hyperplasia
 - Minimal
- * Spleen
 - Extramedullary Hematopoiesis
 - Mild
 - Pigment
 - Minimal
- * Thymus
 - Atrophy
 - Minimal
- * Uterus
 - Cervix
 - Hypertrophy
 - Polyp Stromal
 - Stromal, Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 246

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403005

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Urinary Bladder
- * Bone Marrow
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- Vagina

OBSERVATIONS

- * Adrenal Cortex
- * Brain
- Note: Mild glial cell hyperplasia is present; but could not be coded due to lexicon restriction.
- * Kidney
- * Liver
- [Cyst TGLs = 4 - 12]
- * Lung
- [Infiltration Cellular TGLs = 5 - 6+7]
- * Mammary Gland
- [Fibroadenoma TGLs = 1 - 18]
- [Galactocele TGLs = 2,3 - 19+20]
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 6 - 11]
- * Spleen
- * Thymus
- * Thyroid Gland
- * Uterus
- Glial Cell
- Bile Duct
- Pars Distalis
- Endometrium
- Hyperplasia
- Hyperplasia
- Nephropathy
- Cyst
- Eosinophilic Focus
- Infiltration Cellular
- Fibroadenoma
- Galactocele
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Cyst
- Hyperplasia
- Hyperplasia
- Focal, Minimal
- Mild
- Chronicprogr, Mild
-
-
-
- Marked
- Mild
-
-
- Mild
- Mild
- Moderate
- Moderate
- Cystic, Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 246

TRT#: 2

SEX: Female

DAY ON TEST: 730

DOSE: 0 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403005

ORGAN AND ACCOUNTABLE SITE STATUS

Squamous Metaplasia

Minimal

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 247

TRT#: 2

SEX: Female

DAY ON TEST: 556

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1403006

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea
- * Urinary Bladder
- * Vagina

OBSERVATIONS

- * Bone Marrow Hemorrhage Minimal
- * Kidney Nephropathy Chronicprogr, Minimal
- * Liver Bile Duct Hyperplasia Minimal
- * Lung Infiltration Cellular [Infiltration Cellular TGLs = 2 - 6+7] Infiltration Cellular Histiocyte, Mild
- * Ovary Atrophy Mild
- * Pituitary Gland Pars Distalis Adenoma
- * Spleen White Pulp Atrophy Mild
- * Thymus Pigment Mild
- * Uterus Atrophy Moderate
- Hyperplasia Atypical, Mild
- Squamous Metaplasia Minimal

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 249

TRT#: 2

SEX: Female

DAY ON TEST: 524

DOSE: 0 mg/kg female

DISP: Natural Death

HISTO: 1403008

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Kidney
- * Nose
- * Pituitary Gland
- * Stomach, Glandular
- * Uterus
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Liver
- * Ovary
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Trachea

OBSERVATIONS

- * Bone Marrow
- * Lung
 - [Infiltration Cellular TGLs = 2 - 6+7]
- * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 18]
- * Spleen
- * Thymus
- Vagina
- Hypercellularity
- Infiltration Cellular
- Fibroadenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Inflammation
- Moderate
- Histiocyte, Minimal
- Moderate
- Minimal
- Moderate
- Acute, Minimal

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 250

TRT#: 2

SEX: Female

DAY ON TEST: 521

DOSE: 0 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403009

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Kidney
- * Nose
- * Skin
- * Urinary Bladder
- * Adrenal Medulla
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lung
- * Pancreas
- * Stomach, Forestomach
- * Uterus
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Bone Marrow
- * Heart
- * Liver
- * Mammary Gland
[Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Pituitary Gland
[Adenoma TGLs = 2 - 11]
- * Spleen
- * Thymus
- * Thyroid Gland
- Hypercellularity
- Cardiomyopathy
- Eosinophilic Focus
- Extramedullary Hematopoiesis
- Fibroadenoma
- Cyst
- Adenoma
- Extramedullary Hematopoiesis
- Atrophy
- Hyperplasia
- Marked
- Minimal
- Minimal
- Moderate
- Mild
- Mild

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 251

TRT#: 4

SEX: Female

DAY ON TEST: 634

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403010

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Stomach, Forestomach
- * Uterus
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Kidney
- [Amphophilic/Vacuolar Adenoma TGLs = 3 - 19]
- [Amphophilic/Vacuolar Carcinoma TGLs = 1, 2 - 18]
- * Lung
- [Carcinoma TGLs = 6 - 7]
- Lymph Node
- [Carcinoma TGLs = 5 - 20]
- * Mammary Gland
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 7 - 11]
- * Spleen
- * Thymus
- Renal Tubule
- Deep Cervical
- Pars Distalis
- White Pulp
- Hyperplasia
- Hypercellularity
- Adenoma
- Amphophilic/Vacuolar Adenoma
- Amphophilic/Vacuolar Carcinoma
- Nephropathy
- Carcinoma
- Carcinoma
- Fibroadenoma
- Atrophy
- Adenoma
- Atrophy
- Extramedullary Hematopoiesis
- Hemorrhage
- Pigment
- Atrophy
- Focal, Mild
- Marked
- Multiple
- Multiple
- Chronicprogr, Marked
- Metastatic (Kidney)
- Metastatic (Thyroid Gland)
- Marked
- Mild
- Mild
- Mild
- Mild
- Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 251

TRT#: 4

SEX: Female

DAY ON TEST: 634

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403010

ORGAN AND ACCOUNTABLE SITE STATUS

* Thyroid Gland

C Cell

Adenoma

C Cell

Carcinoma

Note: One thyroid gland has C-cell adenoma, other has C-cell carcinoma extending into muscle.

[Carcinoma TGLs = 4 - 11]

PRIMARY CAUSE OF DEATH

- Kidney Amphophilic/Vacuolar Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 254

TRT#: 4

SEX: Female

DAY ON TEST: 691

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403013

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Bone Marrow
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lung
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea
- * Urinary Bladder
- * Vagina

OBSERVATIONS

- * Kidney [Nephropathy TGLs = 2 - 8] Nephropathy Chronicprogr, Mild
- * Liver [Hepatocellular Adenoma TGLs = 3 - 19] Hepatocellular Adenoma
- * Mammary Gland [Fibroadenoma TGLs = 1 - 18] Fibroadenoma
- * Ovary Atrophy Mild
- * Pituitary Gland Pars Distalis Adenoma
- * Spleen [Adenoma TGLs = 4 - 11] White Pulp Atrophy Mild
- * Thymus Extramedullary Hematopoiesis Mild
- * Uterus Pigment Mild
- Adenomyosis Mild
- Squamous Metaplasia Minimal

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 255

TRT#: 4

SEX: Female

DAY ON TEST: 605

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403014

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Kidney | * Liver | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|----------------------------------------|------------------------------|------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 4 - 7] | | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Ovary | | Atrophy | Mild |
| | Periovarn Tiss | Cyst | |
| | Follicle | Cyst | |
| | [Cyst TGLs = 2 - 14] | | |
| * Pituitary Gland | | Adenoma | |
| | Pars Distalis | | |
| | [Adenoma TGLs = 3 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Mild |

PRIMARY CAUSE OF DEATH

- Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 256

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403015

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Pancreas
- * Parathyroid Gland
- * Stomach, Forestomach
- * Stomach, Glandular
- * Urinary Bladder
- Vagina
- Blood Vessel
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thymus
- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Heart
- * Intestine Large, Rectum
- * Kidney
- * Liver
- * Lung
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thyroid Gland
- * Uterus
- Pars Distalis
- [Adenoma TGLs = 2 - 11]
- C Cell
- C Cell
- Endometrium
- Cardiomyopathy
- Parasite Metazoan
- Nephropathy
- Basophilic Focus
- Infiltration Cellular
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Adenoma
- Hyperplasia
- Hyperplasia
- Squamous Metaplasia
- Minimal
- Chronicprogr, Minimal
- Histiocyte, Minimal
- Mild
- Mild
- Minimal
- Minimal
- Cystic, Minimal
- Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 257

TRT#: 4

SEX: Female

DAY ON TEST: 712

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403016

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Mammary Gland
- * Stomach, Forestomach
- Vagina
- Blood Vessel
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Glandular
- * Bone
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Salivary Glands
- * Trachea
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mesenteric
- * Skin
- * Urinary Bladder

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Brain
- * Kidney
- * Liver
- * Lung
- * Lymph Node, Mandibular
- * Ovary
- Meninges
- [Amphophilic/Vacuolar Carcinoma TGLs = 1, 2 - 18, 19]
- Hepatocyte
- Hepatocyte
- Note: Mitotic alteration was used for the presence of multinucleate hepatocytes.
- [Eosinophilic Focus TGLs = 3 - 20]
- [Hepatocellular Adenoma TGLs = 4,5,6 - 21+22+23]
- Note: Mandibular lymph node with salivary gland submitted as pancreas in B09 has metastatic carcinoma.
- Necrosis
- Hypercellularity
- Carcinoma
- Amphophilic/Vacuolar Carcinoma
- Eosinophilic Focus
- Extramedullary Hematopoiesis
- Hepatocellular Adenoma
- Hypertrophy
- Multinucleated
- Necrosis
- Pigment
- Carcinoma
- Carcinoma
- Atrophy
- Carcinoma
- Mild
- Marked
- Metastatic (Kidney)
- Multiple
- Mild
- Multiple
- Moderate
- Moderate
- Mild
- Marked
- Metastatic (Thyroid Gland)
- Metastatic (Thyroid Gland)
- Marked
- Metastatic (Kidney)

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 257

TRT#: 4

SEX: Female

DAY ON TEST: 712

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403016

ORGAN AND ACCOUNTABLE SITE STATUS

Bursa

Cyst

Note: Second ovary overrun by metastatic renal carcinoma.

* Pancreas

Acinus

Atrophy

Moderate

* Pituitary Gland

Pars Distalis

Adenoma

[Adenoma TGLs = 7 - 11]

* Spleen

Extramedullary Hematopoiesis

Moderate

* Thymus

Atrophy

Mild

* Thyroid Gland

C Cell

Carcinoma

Note: Routine section glands are essentially replaced by metastatic carcinoma, as is the thyroid isthmus in B06.

* Uterus

Endometrium

Adenoma

PRIMARY CAUSE OF DEATH

- Kidney Amphophilic/Vacuolar Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 258

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403017

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Ovary | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |
| * Uterus | Vagina | | |

OBSERVATIONS

- | | | |
|------------------------------------------|------------------------------|-----------------------|
| * Kidney | Nephropathy | Chronicprogr, Minimal |
| * Liver | Pigment | |
| * Lung | Infiltration Cellular | Histiocyte, Minimal |
| [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Mammary Gland | Hyperplasia | Minimal |
| * Spleen | Extramedullary Hematopoiesis | Mild |
| | Pigment | Mild |
| * Thymus | Atrophy | Mild |

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 259

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403018

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Ovary | * Pancreas | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Trachea | * Urinary Bladder |
| Vagina | | | |

MISSING

- | | |
|---------------------|-----------------|
| * Parathyroid Gland | * Thyroid Gland |
|---------------------|-----------------|

OBSERVATIONS

- | | | | |
|-------------------|----------------------------------------------|------------------------------|----------------------|
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Cyst | |
| | Bile Duct | Hepatodiaphragmatic Nodule | |
| | [Cyst TGLs = 2 - 19] | Hyperplasia | Minimal |
| | [Hepatodiaphragmatic Nodule TGLs = 1 - 18] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Mammary Gland | | Hyperplasia | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Salivary Glands | | Atrophy | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Uterus | | Hyperplasia | Atypical, Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 260

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403019

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Lung
- * Pancreas
- * Stomach, Forestomach
- Vagina
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Islets, Pancreatic
- * Kidney
- * Liver
- * Mammary Gland
- [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Urinary Bladder
- * Uterus
- Pars Distalis
- Endometrium
- Pheochromocytoma Benign
- Hyperplasia
- Nephropathy
- Clear Cell Focus
- Fibroadenoma
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Inflammation
- Hyperplasia
- Squamous Metaplasia
- Mild
- Chronicprogr, Minimal
- Mild
- Minimal
- Minimal
- Chronic Active, Mild
- Cystic, Minimal
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 261

TRT#: 4

SEX: Female

DAY ON TEST: 600

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403020

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Islets, Pancreatic
- * Liver
- * Mammary Gland
- * Nose
- * Salivary Glands
- * Skin
- * Thyroid Gland
- * Trachea
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina

OBSERVATIONS

- * Intestine Large, Colon
- * Kidney
- * Lung
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 1 - 11]
- * Spleen
- * Thymus
- * Uterus
- Parasite Metazoan
- Nephropathy
- Infiltration Cellular
- Chronicprogr, Minimal
- Histiocyte, Mild
- Atrophy
- Adenoma
- Moderate
- White Pulp
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Hyperplasia
- Squamous Metaplasia
- Minimal
- Minimal
- Mild
- Minimal
- Cystic, Minimal
- Minimal

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 262

TRT#: 4

SEX: Female

DAY ON TEST: 332

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403021

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thymus | * Thyroid Gland | * Trachea | * Urinary Bladder |
| * Uterus | Vagina | | |

OBSERVATIONS

- | | | | |
|---------------------------|---------------|-----------------------|-----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Ovary | | Atrophy | Moderate |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| [Adenoma TGLs = 1 - 11] | | | |
| * Spleen | White Pulp | Atrophy | Minimal |
| | | Pigment | Mild |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 263

TRT#: 4

SEX: Female

DAY ON TEST: 397

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403022

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Ovary | * Pancreas | * Pituitary Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | * Uterus | Vagina |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | |
|--------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | Degeneration | Cystic, Minimal |
| * Bone Marrow | Hypercellularity | Marked |
| * Kidney | Nephropathy | Chronicprogr, Minimal |
| * Lung | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | Fibroadenoma | |
| [Fibroadenoma TGLs = 1 - 18] | | |
| * Spleen | Extramedullary Hematopoiesis | Moderate |
| * Thymus | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 264

TRT#: 4

SEX: Female

DAY ON TEST: 588

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403023

ORGAN AND ACCOUNTABLE SITE STATUS

[Sarcoma TGLs = 2 - 19]

PRIMARY CAUSE OF DEATH

- Uterus Cervix Sarcoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 265

TRT#: 4

SEX: Female

DAY ON TEST: 648

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403024

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Trachea |
| * Urinary Bladder | * Uterus | Vagina | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | [Infiltration Cellular TGLs = 1 - 6+7] | Infiltration Cellular | Histiocyte, Minimal |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 2 - 11] | | |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Extramedullary Hematopoiesis | Minimal |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | Bilateral, C Cell | Adenoma | |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 266

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403025

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Pancreas
- * Parathyroid Gland
- * Stomach, Forestomach
- * Stomach, Glandular
- * Urinary Bladder
- * Vagina
- * Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Trachea

OBSERVATIONS

- * Kidney
- * Liver
- * Lung
- * Mammary Gland
- * Nose
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- Bile Duct
- [Eosinophilic Focus TGLs = 3 - 19]
- Alveolar Epith
- [Infiltration Cellular TGLs = 2 - 6+7]
- [Fibroadenoma TGLs = 1- 18]
- Respirat Epith
- Pars Distalis
- Endometrium
- Endometrium
- Nephropathy
- Cyst
- Eosinophilic Focus
- Infiltration Cellular
- Metaplasia
- Fibroadenoma
- Hyperplasia
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Adenocarcinoma
- Hyperplasia
- Squamous Metaplasia
- Chronicprogr, Minimal
- Histiocyte, Mild
- Respiratory, Minimal
- Minimal
- Mild
- Minimal
- Mild
- Minimal
- Mild
- Cystic, Minimal
- Minimal

Note: Early carcinoma arising in a polypoid uterine adenoma in B17.

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 267

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403026

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Bone Marrow
* Brain	* Clitoral Gland	* Esophagus	* Eye
* Harderian Gland	* Heart	* Intestine Large, Cecum	* Intestine Large, Colon
* Intestine Large, Rectum	* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum
* Islets, Pancreatic	* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Nose
* Parathyroid Gland	* Salivary Glands	* Skin	* Stomach, Forestomach
* Stomach, Glandular	* Trachea	* Urinary Bladder	Vagina

OBSERVATIONS

* Adrenal Cortex		Degeneration	Cystic, Minimal
* Kidney		Nephropathy	Chronicprogr, Mild
* Liver		Eosinophilic Focus	
* Lung		Infiltration Cellular	Histiocyte, Minimal
	[Infiltration Cellular TGLs = 2 - 6+7]		
* Mammary Gland		Fibroadenoma	
	[Fibroadenoma TGLs = 1 - 18]		
* Ovary		Atrophy	Mild
* Pancreas	Acinus	Hyperplasia	Focal, Mild
* Pituitary Gland	Pars Distalis	Hyperplasia	Marked
	[Hyperplasia TGLs = 3 - 11]		
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Mild
* Thymus		Atrophy	Mild
* Thyroid Gland	C Cell	Adenoma	
	C Cell	Hyperplasia	Marked
* Uterus	Endometrium	Hyperplasia	Cystic, Minimal
		Hyperplasia	Atypical, Minimal
		Squamous Metaplasia	Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 268

TRT#: 4

SEX: Female

DAY ON TEST: 712

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403027

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Parathyroid Gland | * Pituitary Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | | |

OBSERVATIONS

- | | | | |
|------------------|-------------------------------|------------------------------|---------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | Renal Tubule | Dilation | Mild |
| | Pelvis | Dilation | Mild |
| * Liver | Bile Duct | Cholangioma | |
| | [Cholangioma TGLs = 2 - 19] | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Ovary | Follicle | Cyst | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Moderate |
| * Uterus | | Hyperplasia | Atypical, Minimal |
| | | Leiomyosarcoma | |

Note: Adenomyosis in B17.

[Leiomyosarcoma TGLs = 1 - 18]

Vagina

Schwannoma Malignant

PRIMARY CAUSE OF DEATH

- Uterus Leiomyosarcoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 269

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403028

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Pancreas | * Parathyroid Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Thymus | * Thyroid Gland | * Trachea |
| * Urinary Bladder | Vagina | | |

OBSERVATIONS

- | | | | |
|----------------------|------------------------------------------|------------------------------|-----------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | [Infiltration Cellular TGLs = 2 - 6+7] | Infiltration Cellular | Histiocyte, Moderate |
| * Mammary Gland | [Fibroadenoma TGLs = 1 - 18] | Fibroadenoma | |
| * Ovary | | Atrophy | Moderate |
| | [Cyst TGLs = 3 - 14] | Cyst | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Stomach, Glandular | | Mineral | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 270

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403029

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

Blood Vessel	* Bone	* Bone Marrow	* Brain
* Clitoral Gland	* Esophagus	* Eye	* Harderian Gland
* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum	* Intestine Small, Duodenum
* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic	* Kidney
* Lung	* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Nose
* Pancreas	* Parathyroid Gland	* Pituitary Gland	* Salivary Glands
* Skin	* Stomach, Forestomach	* Stomach, Glandular	* Thyroid Gland
* Trachea	* Urinary Bladder	Vagina	

OBSERVATIONS

* Adrenal Cortex		Angiectasis	Mild
	Bilateral	Degeneration	Cystic, Mild
* Adrenal Medulla		Hyperplasia	Focal, Mild
* Heart		Schwannoma Malignant	
* Liver	Bile Duct	Cyst	
		Mixed Cell Focus	
	[Cyst TGLs = 4 - 20]		
	[Mixed Cell Focus TGLs = 3 - 21]		
* Mammary Gland		Fibroadenoma	Multiple
	[Fibroadenoma TGLs = 1,2- 18 + 19]		
* Ovary		Atrophy	Mild
* Spleen		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
* Thymus		Atrophy	Mild
* Uterus	Endometrium	Hyperplasia	Cystic, Minimal
		Inflammation	Chronic Active, Minimal
		Squamous Metaplasia	Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 271

TRT#: 4

SEX: Female

DAY ON TEST: 726

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403030

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Glandular
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lung
- * Nose
- * Stomach, Forestomach
- Vagina

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Intestine Large, Rectum
- * Kidney
- * Liver
- * Ovary
- * Pituitary Gland
- * Skin
- * Spleen
- * Thymus
- * Thyroid Gland
- * Uterus
- Bile Duct
- Pars Distalis
- [Adenoma TGLs = 3 - 11]
- [Fibroma TGLs = 1 - 18]
- C Cell
- Endometrium
- Degeneration
- Hyperplasia
- Hypercellularity
- Parasite Metazoan
- Nephropathy
- Clear Cell Focus
- Hyperplasia
- Atrophy
- Adenoma
- Fibroma
- Extramedullary Hematopoiesis
- Atrophy
- Adenoma
- Hyperplasia
- Cystic, Minimal
- Focal, Minimal
- Marked
- Chronicprogr, Minimal
- Minimal
- Marked
- Moderate
- Minimal
- Cystic, Minimal

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 272

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403031

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Esophagus
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Urinary Bladder
- * Bone Marrow
- * Eye
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Glandular
- * Clitoral Gland
- * Heart
- * Islets, Pancreatic
- * Parathyroid Gland
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Adrenal Medulla
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Kidney
- * Liver
- * Lung
- * Mammary Gland
- [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Pancreas
- * Pituitary Gland
- * Spleen
- * Stomach, Forestomach
- * Thymus
- * Thyroid Gland
- * Uterus
- Bile Duct
- Pars Distalis
- C Cell
- Endometrium
- Degeneration
- Hyperplasia
- Pheochromocytoma Benign
- Polyarteritis Nodosa
- Polyarteritis Nodosa
- Polyarteritis Nodosa
- Nephropathy
- Hyperplasia
- Infiltration Cellular
- Fibroadenoma
- Hyperplasia
- Atrophy
- Polyarteritis Nodosa
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Polyarteritis Nodosa
- Atrophy
- Hyperplasia
- Hyperplasia
- Squamous Metaplasia
- Cystic, Minimal
- Focal, Moderate
- Mild
- Mild
- Mild
- Chronicprogr, Moderate
- Minimal
- Histiocyte, Minimal
- Minimal
- Mild
- Mild
- Marked
- Mild
- Minimal
- Mild
- Mild
- Moderate
- Cystic, Mild
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 273

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403032

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| Blood Vessel | * Bone | * Bone Marrow | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|------------------------|
| * Adrenal Cortex | Bilateral | Degeneration | Cystic, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| | | Pheochromocytoma Benign | |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Cyst | |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 274

TRT#: 4

SEX: Female

DAY ON TEST: 614

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403033

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|---------------------------|---------------|-----------------------|------------------------|
| * Heart | | Cardiomyopathy | Minimal |
| | | Infiltration Cellular | Histiocyte, Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| [Adenoma TGLs = 1 - 11] | | | |
| * Spleen | White Pulp | Atrophy | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| * Uterus | | Squamous Metaplasia | Mild |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 275

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403034

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thymus |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|------------------------------------------|---------------|------------------------------|------------------------|
| * Islets, Pancreatic | | Adenoma | |
| [Adenoma TGLs = 2 - 18] | | | |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Eosinophilic Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| [Infiltration Cellular TGLs = 1 - 6+7] | | Inflammation | Granulomatous, Minimal |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Uterus | | Adenomyosis | Minimal |
| | Endometrium | Hyperplasia | Cystic, Mild |
| | | Inflammation | Chronic Active, Mild |
| | | Polyp Stromal | |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 278

TRT#: 4

SEX: Female

DAY ON TEST: 386

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403037

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Pancreas | * Parathyroid Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland | * Trachea |
| * Urinary Bladder | Vagina | | |

OBSERVATIONS

- | | | | |
|--------------------------------|----------------|------------------------------|-----------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Extramedullary Hematopoiesis | Minimal |
| * Lung | Alveolar Epith | Hyperplasia | Minimal |
| | | Infiltration Cellular | Histiocyte, Minimal |
| | | Fibroadenoma | |
| * Mammary Gland | | | |
| [Fibroadenoma TGLs = 1 - 18] | | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Minimal |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 279

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403038

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Small, Duodenum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Pancreas
- * Salivary Glands
- * Stomach, Glandular
- * Thyroid Gland
- Vagina
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Trachea
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Forestomach
- * Urinary Bladder

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Intestine Large, Rectum
- * Kidney
- * Liver
- Bile Duct
- Bile Duct
- [Clear Cell Focus TGLs = 3 - 20]
- * Lung
- [Infiltration Cellular TGLs = 4 - 6+7]
- * Mammary Gland
- [Fibroadenoma TGLs = 1,2 - 18+19]
- * Ovary
- Follicle
- * Pituitary Gland
- Pars Distalis
- * Spleen
- Atrophy
- Cyst
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- * Thymus
- Atrophy
- * Uterus
- Endometrium
- Hyperplasia
- Squamous Metaplasia
- Parasite Metazoan
- Nephropathy
- Basophilic Focus
- Clear Cell Focus
- Cyst
- Hyperplasia
- Infiltration Cellular
- Fibroadenoma
- Chronicprogr, Mild
- Minimal
- Histiocyte, Minimal
- Multiple
- Mild
- Mild
- Mild
- Minimal
- Cystic, Minimal
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 280

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403039

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|--------------------------|-----------------------------|--------------------------|
| * Adrenal Medulla | * Bone | * Bone Marrow | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | * Uterus | Vagina |

MISSING

- * Thymus

OBSERVATIONS

- | | | | |
|---------------------------|----------------------------------------|------------------------------|---------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Intestine Large, Rectum | | Polyarteritis Nodosa | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 2 - 7] | | |
| * Mammary Gland | | Hyperplasia | Mild |
| * Ovary | Periovarn Tiss | Cyst | |
| | [Cyst TGLs = 1 - 14] | | |
| * Pancreas | | Polyarteritis Nodosa | Minimal |
| * Pituitary Gland | | Cyst | |
| | Pars Distalis | Hyperplasia | Minimal |
| | [Cyst TGLs = 3 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 281

TRT#: 4

SEX: Female

DAY ON TEST: 499

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403040

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Kidney | * Liver | * Lung | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Thymus | * Thyroid Gland | * Trachea | * Urinary Bladder |
| Vagina | | | |

OBSERVATIONS

- | | | | |
|--------------------------------|-------------|------------------------------|-----------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Islets, Pancreatic | | Hyperplasia | Minimal |
| * Mammary Gland | | Fibroadenoma | |
| [Fibroadenoma TGLs = 1 - 18] | | | |
| * Ovary | | Atrophy | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Stomach, Glandular | | Mineral | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Minimal |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 282

TRT#: 4

SEX: Female

DAY ON TEST: 512

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403041

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Ovary | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|-----------------------------------------------------------------------------------------|---------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Necrosis | Marked |
| * Adrenal Medulla | | Necrosis | Marked |
| * Bone Marrow | | Hypercellularity | Marked |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Necrosis | Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | | Fibroadenoma | |
| Note: This fibroadenoma appears to be necrotic, with granulation tissue surrounding it. | | | |
| [Fibroadenoma TGLs = 1 - 18] | | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Moderate |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Minimal |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 283

TRT#: 4

SEX: Female

DAY ON TEST: 615

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403042

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Kidney |
| * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Ovary | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | * Uterus | Vagina |

OBSERVATIONS

- | | | | |
|--------------------------------|---------------|------------------------------|----------------|
| * Adrenal Cortex | | Adenoma | |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Marked |
| * Liver | | Necrosis | Mild |
| * Mammary Gland | | Fibroadenoma | |
| | | Hyperplasia | Mild |
| [Fibroadenoma TGLs = 1 - 18] | | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Moderate |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 284

TRT#: 4

SEX: Female

DAY ON TEST: 730

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403043

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Pancreas | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | Vagina |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------|------------------------------|---------------------|
| * Adrenal Medulla | | Pheochromocytoma Benign | |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Cyst | |
| | [Cyst TGLs = 4 - 21] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 20] | | |
| * Ovary | | Atrophy | Moderate |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 5 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Adenoma | |
| * Uterus | Cervix | Cyst | Squamous |
| | Endometrium | Hyperplasia | Cystic, Mild |
| | Cervix | Hyperplasia | Squamous, Mild |
| | [Cyst TGLs = 3 - 16+19] | | |
| | [Hyperplasia TGLs = 2 - 18] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 285

TRT#: 4

SEX: Female

DAY ON TEST: 712

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403044

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose | * Pituitary Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | * Uterus |

OBSERVATIONS

- | | | | |
|---------------------|-----------------------------------------------|------------------------------|----------------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Extramedullary Hematopoiesis | Minimal |
| | Centrilobular | Necrosis | Mild |
| | | Pigment | Minimal |
| | | Regeneration | Mild |
| | [Regeneration TGLs = 4 - 20] | | |
| * Lung | | Adenocarcinoma | Metastatic (Mammary Gland) |
| * Mammary Gland | | Adenocarcinoma | |
| | | Fibroadenoma | |
| | [Adenocarcinoma TGLs = 2 - 19] | | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Ovary | | Atrophy | Mild |
| * Pancreas | Acinus | Atrophy | Mild |
| * Parathyroid Gland | | Hyperplasia | Focal, Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Marked |
| | [Extramedullary Hematopoiesis TGLs = 3 - 8] | | |
| * Thymus | | Atrophy | Mild |
| Vagina | | Inflammation | Chronic Active, Mild |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 286

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403045

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Trachea
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Urinary Bladder
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin

OBSERVATIONS

- * Adrenal Cortex
 - Angiectasis
 - Degeneration
 - Thrombus
- * Kidney
 - Nephropathy
- * Liver
 - Bile Duct
 - Hyperplasia
- * Lung
 - Infiltration Cellular TGLs = 3 - 6+7]
 - Infiltration Cellular TGLs = 3 - 6+7]
- * Mammary Gland
 - Hyperplasia
- * Ovary
 - Atrophy
 - Bursa
 - Cyst
 - [Cyst TGLs = 2 - 14]
- * Pituitary Gland
 - Pars Distalis
 - Hyperplasia
- * Spleen
 - Extramedullary Hematopoiesis
 - Pigment
- * Stomach, Glandular
 - Mineral
 - Atrophy
- * Thymus
 - Adenoma
- * Thyroid Gland
 - C Cell
 - Dilation
 - Inflammation
- * Uterus
 - Squamous Metaplasia
 - Ulcer

[Inflammation TGLs = 1 - 17+18]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 286

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403045

ORGAN AND ACCOUNTABLE SITE STATUS

Vagina

Hyperplasia

Squamous, Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 287

TRT#: 4

SEX: Female

DAY ON TEST: 302

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403046

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Salivary Glands | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | * Uterus |
| Vagina | | | |

OBSERVATIONS

- | | | |
|---------------------------|------------------------------|-----------------------|
| * Bone Marrow | Hypercellularity | Marked |
| * Kidney | Nephropathy | Chronicprogr, Minimal |
| * Liver | Eosinophilic Focus | |
| * Ovary | Atrophy | Mild |
| * Skin | Fibroma | |
| [Fibroma TGLs = 1 - 18] | | |
| * Spleen | Extramedullary Hematopoiesis | Moderate |
| * Thymus | Atrophy | Minimal |

PRIMARY CAUSE OF DEATH - Skin Fibroma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 288

TRT#: 4

SEX: Female

DAY ON TEST: 672

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403047

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Glandular
- * Uterus
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Thyroid Gland
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Trachea
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
Blood Vessel
Note: An adrenal cortical artery is involved.
- * Bone Marrow
- * Heart
- * Kidney
[Nephropathy TGLs = 6 - 8]
- * Liver
- * Lung
- Lymph Node
[Hemorrhage TGLs = 3 - 19]
- * Mammary Gland
[Fibroadenoma TGLs = 1 - 18]
- * Ovary
[Cyst TGLs = 5 - 14]
- * Parathyroid Gland
- * Pituitary Gland
[Adenoma TGLs = 7 - 11]
- * Spleen
- Media
Hyperplasia
Hypertrophy
Focal, Minimal
Moderate
- Atrium
Hypercellularity
Thrombus
Nephropathy
Moderate
- Peribronchiolr
Necrosis
Fibrosis
Infiltration Cellular
Hemorrhage
Moderate
- Mediastinal
Fibroadenoma
- Periovarn Tiss
Atrophy
Cyst
Moderate
- Pars Distalis
Hyperplasia
Adenoma
Diffuse, Mild
- White Pulp
Atrophy
Extramedullary Hematopoiesis
Moderate
Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 288

TRT#: 4

SEX: Female

DAY ON TEST: 672

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403047

ORGAN AND ACCOUNTABLE SITE STATUS

* Stomach, Forestomach	Epithelium	Pigment	Mild
		Hyperplasia	Mild
		Inflammation	Acute, Mild
* Thymus		Ulcer	Mild
		Atrophy	Moderate
[Atrophy TGLs = 2 - 6]			

PRIMARY CAUSE OF DEATH - Heart Atrium Thrombus

CONTRIBUTORY CAUSE OF DEATH - Kidney Nephropathy

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 289

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403048

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Nose
- * Salivary Glands
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Ovary
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Glandular
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Cortex
 - * Bone Marrow
 - * Heart
 - * Intestine Small, Duodenum
 - [Leiomyoma TGLs = 8 - 21]
 - * Kidney
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 5 - 6+7]
 - * Mammary Gland
 - [Fibroadenoma TGLs = 1,6 - 18+20]
 - * Pituitary Gland
 - [Adenoma TGLs = 7 - 11]
 - * Skin
 - [Keratoacanthoma TGLs = 2 - 19]
 - * Spleen
 - * Thymus
 - * Uterus
 - [Polyp Stromal TGLs = 4 - 16]
- Degeneration
 Hypercellularity
 Cardiomyopathy
 Leiomyoma
 Nephropathy
 Necrosis
 Infiltration Cellular
 Fibroadenoma
 Hyperplasia
 Adenoma
 Keratoacanthoma
 Extramedullary Hematopoiesis
 Pigment
 Atrophy
 Hyperplasia
 Polyp Stromal
- Cystic, Minimal
 Moderate
 Minimal
 Chronicprogr, Minimal
 Minimal
 Histiocyte, Marked
 Multiple
 Minimal
 Mild
 Minimal
 Mild
 Cystic, Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 290

TRT#: 4

SEX: Female

DAY ON TEST: 629

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403049

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Brain
* Clitoral Gland	* Esophagus	* Eye	* Harderian Gland
* Heart	* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum
* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic
* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Mammary Gland	* Nose
* Parathyroid Gland	* Pituitary Gland	* Salivary Glands	* Skin
* Stomach, Forestomach	* Stomach, Glandular	* Trachea	Vagina

OBSERVATIONS

* Adrenal Cortex		Degeneration	Cystic, Minimal
* Bone Marrow		Hypercellularity	Marked
* Kidney		Nephropathy	Chronicprogr, Minimal
* Liver		Adenocarcinoma	Metastatic (Uterus)
	Bile Duct	Hyperplasia	Mild
		Necrosis	Mild
* Lung		Infiltration Cellular	Histiocyte, Mild
		Inflammation	Granulomatous, Minimal
	[Infiltration Cellular TGLs = 3 - 6+7]		
Lymph Node	Pancreatic	Adenocarcinoma	Metastatic (Uterus)
* Ovary		Adenocarcinoma	Metastatic (Uterus)
		Atrophy	Moderate
* Pancreas		Adenocarcinoma	Metastatic (Uterus)
	[Adenocarcinoma TGLs = 2 - 19]		
Peritoneum		Adenocarcinoma	Metastatic (Uterus)
* Spleen	White Pulp	Atrophy	Moderate
		Extramedullary Hematopoiesis	Moderate
		Pigment	Minimal
* Thymus		Atrophy	Marked
* Thyroid Gland	C Cell	Adenoma	
* Urinary Bladder		Adenocarcinoma	Metastatic (Uterus)
* Uterus	Endometrium	Adenocarcinoma	

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 290

TRT#: 4

SEX: Female

DAY ON TEST: 629

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403049

ORGAN AND ACCOUNTABLE SITE STATUS

Squamous Metaplasia

Mild

[Adenocarcinoma TGLs = 1 - 18]

PRIMARY CAUSE OF DEATH

- Uterus Endometrium Adenocarcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 294

TRT#: 4

SEX: Female

DAY ON TEST: 570

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403053

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- Blood Vessel
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Stomach, Forestomach
- Vagina
- * Bone
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Glandular
- * Brain
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Thyroid Gland

MISSING

- * Clitoral Gland

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 4 - 6+7]
- Lymph Node
- [Ectasia TGLs = 3 - 20]
- * Nose
- * Ovary
- * Pituitary Gland
- * Skin
- [Schwannoma Malignant TGLs = 1 - 18]
- * Spleen
- [Extramedullary Hematopoiesis TGLs = 2 - 19]
- * Thymus
- * Urinary Bladder
- * Uterus
- Extramedullary Hematopoiesis
- Hypercellularity
- Inflammation
- Necrosis
- Infiltration Cellular
- Ectasia
- Inflammation
- Atrophy
- Hyperplasia
- Schwannoma Malignant
- Extramedullary Hematopoiesis
- Atrophy
- Inflammation
- Squamous Metaplasia
- Minimal
- Marked
- Acute, Minimal
- Mild
- Histiocyte, Mild
- Moderate
- Suppurative, Minimal
- Mild
- Minimal
- Marked
- Moderate
- Chronic Active, Mild
- Minimal

PRIMARY CAUSE OF DEATH - Skin Schwannoma Malignant

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 296

TRT#: 4

SEX: Female

DAY ON TEST: 482

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403055

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|--------------------------|-----------------------------|--------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Kidney | * Lung |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose | * Ovary |
| * Pancreas | * Parathyroid Gland | * Pituitary Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | * Uterus | Vagina |

MISSING

- * Clitoral Gland

OBSERVATIONS

- | | | |
|--------------------------------------------------------------|------------------------------|----------|
| * Adrenal Cortex | Extramedullary Hematopoiesis | Mild |
| * Bone Marrow | Hypercellularity | Marked |
| * Harderian GI | | |
| Note: Some mammary fibroadenoma is on slide 13 as a floater. | | |
| * Intestine Large, Rectum | Parasite Metazoan | |
| * Liver | Necrosis | Moderate |
| * Mammary Gland | Fibroadenoma | |
| [Fibroadenoma TGLs = 1 - 18] | | |
| * Spleen | Extramedullary Hematopoiesis | Moderate |
| * Thymus | Atrophy | Mild |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 297

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403056

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Pancreas
- * Parathyroid Gland
- * Pituitary Gland
- * Salivary Glands
- * Skin
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea
- * Urinary Bladder
- Vagina

OBSERVATIONS

- * Bone Marrow: Hypercellularity, Moderate
- * Kidney: Nephropathy, Chronicprogr, Minimal
- * Liver: Bile Duct, Hyperplasia, Minimal
- * Lung: Infiltration Cellular, Histocyte, Minimal; Inflammation, Granulomatous, Minimal
- * Mammary Gland: Fibroadenoma
 - Note: The center of the fibroadenoma is necrotic. liquefying, and surrounded by granulation tissue.
 - [Fibroadenoma TGLs = 1 - 18]
- * Nose: Inflammation, Suppurative, Minimal
- * Ovary: Atrophy, Mild
- * Spleen: Extramedullary Hematopoiesis, Mild; Pigment, Minimal
- * Stomach, Forestomach: Epithelium, Hyperplasia, Minimal
- * Thymus: Atrophy, Minimal
- * Uterus: Endometrium, Hyperplasia, Cystic, Minimal; Polyp Stromal, Squamous Metaplasia, Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 298

TRT#: 4

SEX: Female

DAY ON TEST: 633

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403057

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pituitary Gland
- * Stomach, Glandular
- Vagina
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Thyroid Gland
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Forestomach
- * Urinary Bladder

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 4 - 6+7]
- * Mammary Gland
- [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- [Malignant Mixed Sex Cord Stromal Tumor TGLs = 3 - 19]
- * Pancreas
- Peritoneum
- * Spleen
- White Pulp
- * Thymus
- Hypercellularity
- Nephropathy
- Necrosis
- Infiltration Cellular
- Inflammation
- Malignant Mixed Sex Cord Stromal Tumor
- Fibroadenoma
- Atrophy
- Malignant Mixed Sex Cord Stromal Tumor
- Malignant Mixed Sex Cord Stromal Tumor
- Malignant Mixed Sex Cord Stromal Tumor
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Malignant Mixed Sex Cord Stromal Tumor
- Moderate
- Chronicprogr, Minimal
- Minimal
- Histiocyte, Minimal
- Granulomatous, Minimal
- Metastatic (Ovary)
- Marked
- Metastatic (Ovary)
- Metastatic (Ovary)
- Marked
- Moderate
- Minimal
- Marked
- Metastatic (Ovary)

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 298

TRT#: 4

SEX: Female

DAY ON TEST: 633

DOSE: 75 mg/kg female

DISP: Natural Death

HISTO: 1403057

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Malignant Mixed Sex Cord Stromal Tumor Metastatic (Ovary)

PRIMARY CAUSE OF DEATH

- Ovary Malignant Mixed Sex Cord Stromal Tumor

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 299

TRT#: 4

SEX: Female

DAY ON TEST: 729

DOSE: 75 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403058

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|------------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | | Inflammation | Granulomatous, Minimal |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Mild |
| | | Squamous Metaplasia | Minimal |

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 300

TRT#: 4

SEX: Female

DAY ON TEST: 675

DOSE: 75 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403059

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

* Adrenal Medulla	Blood Vessel	* Bone	* Brain
* Clitoral Gland	* Esophagus	* Eye	* Harderian Gland
* Heart	* Intestine Large, Cecum	* Intestine Large, Colon	* Intestine Large, Rectum
* Intestine Small, Duodenum	* Intestine Small, Ileum	* Intestine Small, Jejunum	* Islets, Pancreatic
* Kidney	* Lymph Node, Mandibular	* Lymph Node, Mesenteric	* Nose
* Ovary	* Pancreas	* Parathyroid Gland	* Pituitary Gland
* Salivary Glands	* Skin	* Stomach, Forestomach	* Stomach, Glandular
* Trachea	* Urinary Bladder	* Uterus	Vagina

OBSERVATIONS

* Adrenal Cortex		Degeneration	Cystic, Minimal
* Bone Marrow		Hypercellularity	Marked
* Liver		Extramedullary Hematopoiesis	Minimal
	[Hepatodiaphragmatic Nodule TGLs = 2 - 19]	Hepatodiaphragmatic Nodule	
* Lung		Infiltration Cellular	Histiocyte, Minimal
* Mammary Gland		Fibroadenoma	
	[Fibroadenoma TGLs = 1 - 18]		
* Spleen		Extramedullary Hematopoiesis	Moderate
* Thymus		Atrophy	Mild
* Thyroid Gland	C Cell	Adenoma	
	C Cell	Hyperplasia	Mild

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 301

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403060

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Skin
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Parathyroid Gland
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Heart
- * Kidney
- * Lung
- [Infiltration Cellular TGLs = 4 - 6+7]
- * Mammary Gland
- [Fibroadenoma TGLs = 1,2,3 - 18+19+20]
- * Nose
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 5 - 11]
- * Spleen
- * Thymus
- * Thyroid Gland
- [Adenoma TGLs = 6 - 11]
- * Uterus
- Pars Distalis
- C Cell
- Endometrium
- Hyperplasia
- Hypercellularity
- Cardiomyopathy
- Nephropathy
- Infiltration Cellular
- Inflammation
- Fibroadenoma
- Inflammation
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Adenoma
- Hyperplasia
- Hyperplasia
- Squamous Metaplasia
- Focal, Marked
- Mild
- Mild
- Chronicprogr, Minimal
- Histiocyte, Minimal
- Granulomatous, Minimal
- Multiple
- Chronic, Minimal
- Moderate
- Mild
- Minimal
- Minimal
- Cystic, Minimal
- Atypical, Marked
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 302

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403061

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Thymus |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|----------------------|-------------------------------------|------------------------------|------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Cyst | |
| | [Cyst TGLs = 3 - 20] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Minimal |
| * Mammary Gland | | Fibroadenoma | Multiple |
| | [Fibroadenoma TGLs = 1,2 - 18+19] | Hyperplasia | Mild |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Stomach, Glandular | Submucosa | Inflammation | Chronic, Minimal |
| * Thyroid Gland | C Cell | Adenoma | |
| | C Cell | Hyperplasia | Moderate |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 303

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403062

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Nose
- * Pancreas
- * Skin
- * Stomach, Forestomach
- * Trachea
- * Urinary Bladder
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland

OBSERVATIONS

- * Kidney Nephropathy Chronicprogr, Mild
- * Liver Hepatocellular Adenoma
- Bile Duct Hyperplasia Minimal
- [Hepatocellular Adenoma TGLs = 1 - 18]
- * Lung Infiltration Cellular Histiocyte, Mild
- [Infiltration Cellular TGLs = 4 - 6]
- * Mammary Gland Galactocele Mild
- [Galactocele TGLs = 2 - 19]
- * Ovary Atrophy Mild
- Follicle Cyst
- [Cyst TGLs = 3 - 14]
- * Pituitary Gland Pars Distalis Hyperplasia Marked
- [Hyperplasia TGLs = 5 - 11]
- * Spleen Extramedullary Hematopoiesis Mild
- Pigment Minimal
- * Thymus Atrophy Minimal
- * Uterus Endometrium Hyperplasia Cystic, Minimal
- Polyp Stromal
- Squamous Metaplasia Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 304

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403063

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Blood Vessel | * Bone | * Bone Marrow | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|--------------------------|------------------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Intestine Large, Colon | Lymphoid Tiss | Hyperplasia | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 2 - 18] | | |
| * Ovary | | Atrophy | Moderate |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | Bilateral, C Cell | Adenoma | |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 306

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403065

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Thyroid Gland
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- Vagina

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- * Mammary Gland
- * Ovary
- * Pituitary Gland
- * Spleen
- * Stomach, Glandular
- * Thymus
- * Uterus
- Bile Duct
- Pars Distalis
- Cervix
- Hyperplasia
- Hypercellularity
- Nephropathy
- Extramedullary Hematopoiesis
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Fibroadenoma
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Mineral
- Atrophy
- Hyperplasia
- Squamous Metaplasia
- Focal, Minimal
- Moderate
- Chronicprogr, Mild
- Minimal
- Minimal
- Histiocyte, Minimal
- Granulomatous, Minimal
- Mild
- Moderate
- Minimal
- Minimal
- Mild
- Squamous, Mild
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 307

TRT#: 6

SEX: Female

DAY ON TEST: 676

DOSE: 250 mg/kg female

DISP: Natural Death

HISTO: 1403066

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Skin
- * Urinary Bladder
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Mammary Gland
- * Thyroid Gland
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Pancreas
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Intestine Large, Rectum
 - * Kidney [Nephropathy TGLs = 1 - 8]
 - * Liver
 - * Lung
 - * Nose
 - * Ovary
 - * Pituitary Gland [Adenoma TGLs = 3 - 11]
 - * Salivary Glands
 - * Spleen
 - * Stomach, Forestomach
 - * Thymus
 - * Uterus
 - Vagina
- | | | |
|-----------|------------------------------|----------------------|
| | Polyarteritis Nodosa | Mild |
| | Nephropathy | Chronicprogr, Marked |
| Bile Duct | Cyst | |
| | Infiltration Cellular | Histiocyte, Minimal |
| | Inflammation | Acute, Minimal |
| | Atrophy | Minimal |
| | Polyarteritis Nodosa | Minimal |
| | Adenoma | |
| | Polyarteritis Nodosa | Mild |
| | Extramedullary Hematopoiesis | Mild |
| | Pigment | Minimal |
| | Hyperplasia | Moderate |
| | Inflammation | Chronic, Mild |
| | Polyarteritis Nodosa | Mild |
| | Atrophy | Mild |
| | Polyarteritis Nodosa | Mild |
| | Inflammation | Acute, Mild |

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 308

TRT#: 6

SEX: Female

DAY ON TEST: 658

DOSE: 250 mg/kg female

DISP: Natural Death

HISTO: 1403067

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Skin
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Heart
- * Kidney
- [Cyst TGLs = 2 - 18]
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 3 - 6+7]
- * Nose
- * Ovary
- * Pituitary Gland
- * Spleen
- [Leukemia Mononuclear TGLs = 1- 8]
- * Thymus
- Leukemia Mononuclear
- Hypercellularity
- Cardiomyopathy
- Leukemia Mononuclear
- Cyst
- Leukemia Mononuclear
- Cyst
- Leukemia Mononuclear
- Infiltration Cellular
- Inflammation
- Leukemia Mononuclear
- Inflammation
- Atrophy
- Hyperplasia
- Atrophy
- Extramedullary Hematopoiesis
- Leukemia Mononuclear
- Pigment
- Atrophy
- Moderate
- Minimal
- Histiocyte, Mild
- Granulomatous, Minimal
- Chronic, Mild
- Mild
- Minimal
- Marked
- Mild
- Mild
- Moderate

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 308

TRT#: 6

SEX: Female

DAY ON TEST: 658

DOSE: 250 mg/kg female

DISP: Natural Death

HISTO: 1403067

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Endometrium

Hyperplasia

Cystic, Minimal

Polyp Stromal

Squamous Metaplasia

Minimal

Thrombus

PRIMARY CAUSE OF DEATH

- Spleen Leukemia Mononuclear

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 309

TRT#: 6

SEX: Female

DAY ON TEST: 704

DOSE: 250 mg/kg female

DISP: Natural Death

HISTO: 1403068

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | |
|------------------------------------------|------------------------------|-------------------------|
| Blood Vessel | Mineral | Minimal |
| Note: The aorta is mineralized. | | |
| * Bone Marrow | Hypercellularity | Moderate |
| * Heart | Cardiomyopathy | Minimal |
| * Kidney | Nephropathy | Chronicprogr, Mild |
| * Liver | Necrosis | Minimal |
| * Lung | Infiltration Cellular | Histiocyte, Mild |
| | Inflammation | Granulomatous, Minimal |
| [Infiltration Cellular TGLs = 5 - 6+7] | | |
| * Mammary Gland | Fibroadenoma | Multiple |
| [Fibroadenoma TGLs = 1,2 - 18+19] | | |
| * Ovary | Atrophy | Moderate |
| * Pituitary Gland | Adenoma | |
| [Adenoma TGLs = 4 - 11] | | |
| * Spleen | Extramedullary Hematopoiesis | Moderate |
| | Pigment | Minimal |
| * Thymus | Atrophy | Mild |
| * Uterus | Inflammation | Chronic Active, Minimal |
| | Polyp Stromal | |
| | Squamous Metaplasia | Mild |
| [Polyp Stromal TGLs = 3 - 17] | | |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 310

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403069

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Kidney | * Liver | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

MISSING

- * Thymus

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|----------------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Lung | | Adenocarcinoma | Metastatic (Mammary Gland) |
| | [Infiltration Cellular TGLs = 3 - 6+7] | Infiltration Cellular | Histiocyte, Minimal |
| * Mammary Gland | | Adenocarcinoma | |
| | [Adenocarcinoma TGLs = 1 - 18] | Fibroadenoma | |
| | [Fibroadenoma TGLs = 2 - 19] | | |
| * Ovary | | Atrophy | Mild |
| | Follicle | Cyst | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Thrombus | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 312

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403071

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Parathyroid Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|----------------------|------------------------------------------|------------------------------|-------------------------|
| * Bone Marrow | | Hypercellularity | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | | Adenocarcinoma | Metastatic (Uterus) |
| | | Infiltration Cellular | Histiocyte, Moderate |
| | | Inflammation | Chronic Active, Minimal |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| * Stomach, Glandular | | Mineral | Minimal |
| * Thymus | | Atrophy | Moderate |
| * Uterus | Endometrium | Adenocarcinoma | Multiple |
| | Endometrium | Adenoma | |
| | | Dilation | Marked |
| | | Papilloma | Squamous |
| | | Polyp Stromal | |
| | | Squamous Metaplasia | Marked |

Note: Inflammatory polyp in uterine horn (B17) adjacent to second probable early endometrial carcinoma.

Note: B19 has abundant keratin but no tumor in that slide, so correlated only with the squamous metaplasia.

Note: Squamous Cell Papilloma within uterine body (B17). Small (3 mm long) endometrial carcinoma in B17 uterine body adjacent to the papilloma.

Note: B18 = large uterine carcinoma involving the oviduct; it appeared grossly distinct from that in B17 and 19.

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 312

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403071

ORGAN AND ACCOUNTABLE SITE STATUS

[Adenocarcinoma TGLs = 1 - 18]

[Papilloma TGLs = 2 - 17+19]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 313

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403072

ORGAN AND ACCOUNTABLE SITE STATUS

Thrombus

[Hyperplasia TGLs = 2 - 16+17+19]

[Thrombus TGLs = 3 - 19]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 314

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403073

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|-------------------------------------|------------------------------|-------------------------|
| * Adrenal Cortex | | Angiectasis | Moderate |
| | | Degeneration | Cystic, Minimal |
| | | Hyperplasia | Focal, Mild |
| | | Thrombus | |
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Minimal |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Basophilic Focus | |
| * Lung | | Inflammation | Chronic Active, Minimal |
| * Mammary Gland | | Fibroadenoma | Multiple |
| | [Fibroadenoma TGLs = 1,2 - 18+19] | | |
| * Ovary | | Atrophy | Mild |
| | Periovarn Tiss | Cyst | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| | [Hyperplasia TGLs = 3 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 315

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403074

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Kidney |
| * Lymph Node, Mesenteric | * Mammary Gland | * Ovary | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Thymus | * Thyroid Gland |
| * Trachea | * Urinary Bladder | * Uterus | Vagina |

OBSERVATIONS

- | | | | |
|--------------------------|--------------------------------------------------------------------|------------------------------|------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Bone Marrow | | Hypercellularity | Moderate |
| * Harderian Gland | | Fibrosis | Mild |
| * Liver | | Eosinophilic Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Lymph Node, Mandibular | | Infiltration Cellular | Plasma Cell, Mild |
| | [Infiltration Cellular TGLs = 1 - 12] | | |
| * Nose | | Foreign Body | |
| | Nasolacrim Dct | Inflammation | Suppurative, Moderate |
| | | Inflammation | Chronic Active, Mild |
| | Respirat Epith | Osteosarcoma | |
| | Note: One incisor is displaced into the anterior/mid nasal cavity. | Squamous Metaplasia | Mild |
| | [Osteosarcoma TGLs = 3 - 19] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| Tooth | | Malformation | |
| | Note: Malocclusion (gross diagnosis). | | |
| | [Malformation TGLs = 2 - 18+20] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 316

TRT#: 6

SEX: Female

DAY ON TEST: 648

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403075

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Kidney
- * Liver
- * Lung
- * Mammary Gland
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- Bile Duct
- Pars Distalis
- [Adenoma TGLs = 1 - 11]
- Hyperplasia
- Nephropathy
- Cyst
- Infiltration Cellular
- Fibroadenoma
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Squamous Metaplasia
- Focal, Minimal
- Chronicprogr, Minimal
- Histiocyte, Minimal
- Moderate
- Mild
- Mild
- Mild
- Mild

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 317

TRT#: 6

SEX: Female

DAY ON TEST: 608

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403076

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Liver | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Pancreas | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|--------------------------|-----------------------------------------|------------------------------|----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Mild |
| * Heart | | Cardiomyopathy | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Marked |
| | [Nephropathy TGLs = 2 - 8] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| * Lymph Node, Mandibular | | Infiltration Cellular | Plasma Cell, Mild |
| | [Infiltration Cellular TGLs = 1 - 12] | | |
| * Ovary | | Atrophy | Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Uterus | | Adenomyosis | Mild |
| | | Dilation | Moderate |
| | | Inflammation | Chronic Active, Mild |
| | | Squamous Metaplasia | Minimal |
| | | Thrombus | |

PRIMARY CAUSE OF DEATH

- Kidney Nephropathy

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 318

TRT#: 6

SEX: Female

DAY ON TEST: 697

DOSE: 250 mg/kg female

DISP: Natural Death

HISTO: 1403077

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Kidney | * Liver | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Pituitary Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland | * Trachea |
| * Urinary Bladder | Vagina | | |

OBSERVATIONS

- | | | | |
|------------------|---------------------------------------------|------------------------------|------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| Oral Mucosa | Gingival | Squamous Cell Papilloma | |
| | [Squamous Cell Papilloma TGLs = 2 - 1+18] | | |
| * Ovary | | Atrophy | Marked |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Moderate |
| * Uterus | | Squamous Metaplasia | Minimal |

PRIMARY CAUSE OF DEATH - Oral Mucosa Gingival Squamous Cell Papilloma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 319

TRT#: 6

SEX: Female

DAY ON TEST: 715

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403078

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose | * Pancreas |
| * Parathyroid Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |
| * Uterus | | | |

INSUFFICIENT TISSUE

- * Thymus

OBSERVATIONS

- | | | | |
|------------------------------------------|---------------|------------------------------|----------------------|
| * Adrenal Cortex | Bilateral | Degeneration
Necrosis | Cystic, Mild
Mild |
| [Degeneration TGLs = 4 - 20] | | | |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Marked |
| * Heart | | Cardiomyopathy | Mild |
| * Intestine Large, Colon | | Parasite Metazoan | |
| * Intestine Large, Rectum | | Parasite Metazoan | |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Eosinophilic Focus | |
| | | Extramedullary Hematopoiesis | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| [Infiltration Cellular TGLs = 3 - 6+7] | | | |
| * Mammary Gland | | Fibroadenoma | |
| [Fibroadenoma TGLs = 1 - 18] | | | |
| * Ovary | | Atrophy | Moderate |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Extramedullary Hematopoiesis | Marked |

Note: Processor issues made spleen section very difficult to interpret, however, edges processed properly to confirm diagnoses.

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 319

TRT#: 6

SEX: Female

DAY ON TEST: 715

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403078

ORGAN AND ACCOUNTABLE SITE STATUS

[Extramedullary Hematopoiesis TGLs = 2 - 19]

Vagina

Inflammation

Acute, Minimal

PRIMARY CAUSE OF DEATH

- Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 320

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403079

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Trachea
- * Urinary Bladder
- Vagina

OBSERVATIONS

- Blood Vessel
 - Mineral
 - Minimal
 - Note: Gastric arteries are mineralized.
- * Bone Marrow
 - Hypercellularity
 - Mild
- * Heart
 - Cardiomyopathy
 - Minimal
- * Kidney
 - Nephropathy
 - Chronicprogr, Mild
- * Liver
 - Eosinophilic Focus
- * Lung
 - Infiltration Cellular
 - Histiocyte, Moderate
 - [Infiltration Cellular TGLs = 3 - 6+7]
- * Mammary Gland
 - Fibroadenoma
 - Multiple
 - [Fibroadenoma TGLs = 1,2 - 18+19]
- * Ovary
 - Atrophy
 - Mild
- * Pituitary Gland
 - Pars Distalis
 - Adenoma
 - [Adenoma TGLs = 4 - 11]
- * Spleen
 - Extramedullary Hematopoiesis
 - Mild
 - Pigment
 - Minimal
- * Stomach, Glandular
 - Mineral
 - Minimal
- * Thymus
 - Atrophy
 - Minimal
- * Thyroid Gland
 - C Cell
 - Adenoma
- * Uterus
 - Endometrium
 - Hyperplasia
 - Cystic, Minimal
 - Squamous Metaplasia
 - Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 321

TRT#: 6

SEX: Female

DAY ON TEST: 597

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403080

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|------------------------|------------------------------------------|-----------------------|--------------------|
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| | Follicle | Cyst | |
| | [Cyst TGLs = 1 - 14] | | |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 4 - 19] | | |
| * Spleen | | Pigment | Mild |
| * Stomach, Forestomach | | Inflammation | Chronic, Minimal |
| * Stomach, Glandular | | Erosion | Mild |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Hyperplasia | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | [Hyperplasia TGLs = 2 - 18] | | |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 323

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403082

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Pancreas | * Parathyroid Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland | * Trachea |
| * Urinary Bladder | Vagina | | |

OBSERVATIONS

- | | | | |
|-------------------|-------------------------------------|------------------------------|-------------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Extramedullary Hematopoiesis | Minimal |
| * Lung | Bronchiole | Foreign Body | Multiple |
| | | Infiltration Cellular | Histiocyte, Mild |
| | | Inflammation | Granulomatous, Moderate |
| | [Inflammation TGLs = 4 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | Multiple |
| | [Fibroadenoma TGLs = 1,2 - 18+19] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Mild |
| * Uterus | | Dilation | Mild |
| | | Hyperplasia | Atypical, Moderate |
| | | Inflammation | Chronic Active, Mild |
| | | Thrombus | |
| | [Dilation TGLs = 3 - 16] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 324

TRT#: 6

SEX: Female

DAY ON TEST: 690

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403083

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Kidney
- * Liver
- [Cyst TGLs = 2 - 12]
- * Lung
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 1 - 18]
- * Spleen
- * Thymus
- * Uterus
- Bile Duct
- Bile Duct
- Pars Distalis
- Endometrium
- Hyperplasia
- Hypercellularity
- Nephropathy
- Cyst
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Atrophy
- Adenoma
- Pigment
- Atrophy
- Hyperplasia
- Squamous Metaplasia
- Focal, Mild
- Mild
- Chronicprogr, Minimal
- Minimal
- Histiocyte, Minimal
- Granulomatous, Minimal
- Mild
- Mild
- Minimal
- Cystic, Minimal
- Minimal

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 325

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403084

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Trachea | * Urinary Bladder | Vagina |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|----------------------|------------------------------------------|------------------------------|------------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | Bile Duct | Hyperplasia | Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | | Inflammation | Granulomatous, Minimal |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| | | Cyst | |
| | [Cyst TGLs = 3 - 14] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Stomach, Glandular | | Mineral | Mild |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Adenoma | |
| | [Adenoma TGLs = 1 - 18] | | |
| * Uterus | | Hyperplasia | Atypical, Marked |
| | Endometrium | Hyperplasia | Cystic, Mild |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 327

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403086

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Salivary Glands | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | |
|-----------------------------------------------------------------|------------------------------|------------------------|
| * Adrenal Medulla
[Pheochromocytoma Benign TGLs = 2 - 11] | Pheochromocytoma Benign | |
| * Bone Marrow | Hypercellularity | Moderate |
| * Heart | Cardiomyopathy | Minimal |
| * Kidney
[Nephropathy TGLs = 5 - 8] | Nephropathy | Chronicprogr, Moderate |
| * Liver | Clear Cell Focus | |
| | Extramedullary Hematopoiesis | Minimal |
| * Lung
[Infiltration Cellular TGLs = 3 - 6+7] | Infiltration Cellular | Histiocyte, Mild |
| * Ovary | Atrophy | Mild |
| * Pituitary Gland
Pars Distalis
[Adenoma TGLs = 4 - 11] | Adenoma | |
| * Skin
[Hemangiosarcoma TGLs = 1 - 18] | Hemangiosarcoma | |
| * Spleen | Extramedullary Hematopoiesis | Moderate |
| | Pigment | Minimal |
| * Thymus | Atrophy | Mild |
| * Uterus | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 328

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403087

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Forestomach
- Vagina
- * Brain
- * Harderian Gland
- * Islets, Pancreatic
- * Pituitary Gland
- * Stomach, Glandular
- * Clitoral Gland
- * Heart
- * Lymph Node, Mandibular
- * Salivary Glands
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Cortex
 - Angiectasis Mild
 - Hypertrophy Focal, Mild
 - Hypercellularity Moderate
- * Bone Marrow
 - Bilateral
 - Polyarteritis Nodosa Mild
 - Polyarteritis Nodosa Mild
 - Polyarteritis Nodosa Mild
 - Polyarteritis Nodosa Mild
 - Nephropathy Chronicprogr, Moderate
- * Kidney
 - Eosinophilic Focus
- * Liver
 - Infiltration Cellular TGLs = 3 - 6+7] Histiocyte, Mild
- * Lung
 - Fibroadenoma
- * Mammary Gland
 - [Fibroadenoma TGLs = 4 - 19]
- * Ovary
 - Atrophy Moderate
- * Oviduct
 - Inflammation Acute, Mild
 - [Inflammation TGLs = 2 - 14]
- * Pancreas
 - Polyarteritis Nodosa Mild
- * Spleen
 - Extramedullary Hematopoiesis Mild
 - Pigment Mild
- * Thymus
 - Atrophy Mild
- * Thyroid Gland
 - C Cell
 - Hyperplasia Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 328

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403087

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Polyarteritis Nodosa

Mild

Dilation

Marked

Papilloma

Squamous

Squamous Metaplasia

Marked

[Dilation TGLs = 1 - 17, 18]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 329

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403088

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Pituitary Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | Vagina |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|----------------------|------------------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Angiectasis | Minimal |
| | | Degeneration | Cystic, Minimal |
| * Bone Marrow | | Hypercellularity | Marked |
| * Islets, Pancreatic | | Adenoma | |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Alveolar/Bronchiolar Adenoma | |
| | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Ovary | Follicle | Cyst | |
| | [Cyst TGLs = 5 - 14] | | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Adenoma | |
| | [Adenoma TGLs = 2 - 11] | | |
| * Uterus | | Dilation | Marked |
| | | Hemangiosarcoma | |
| | | Polyp Stromal | |
| | | Squamous Metaplasia | Minimal |
| | [Dilation TGLs = 4 - 19] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 329

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403088

ORGAN AND ACCOUNTABLE SITE STATUS

[Hemangiosarcoma TGLs = 1 - 18]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 330

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403089

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Ovary | * Pancreas | * Parathyroid Gland |
| * Pituitary Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |
| * Uterus | Vagina | | |

OBSERVATIONS

- | | | | |
|-----------------|--------|------------------------------|------------------------|
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Angiectasis | Mild |
| | | Clear Cell Focus | |
| | | Eosinophilic Focus | |
| | Serosa | Fibrosis | Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Inflammation | Granulomatous, Minimal |
| * Mammary Gland | | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 331

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403090

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Trachea | * Urinary Bladder | Vagina | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|--------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Basophilic Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Marked |
| | | Inflammation | Chronic Active, Mild |
| | [Infiltration Cellular TGLs = 4 - 6+7] | | |
| * Ovary | Follicle | Cyst | |
| | Bursa | Cyst | |
| | [Cyst TGLs = 3- 14] | | |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 5 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Minimal |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Hyperplasia | Marked |
| * Uterus | | Adenomyosis | Moderate |
| | | Dilation | Moderate |
| | Endometrium | Hyperplasia | Cystic, Marked |
| | | Inflammation | Chronic Active, Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 331

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403090

ORGAN AND ACCOUNTABLE SITE STATUS

Polyarteritis Nodosa

Minimal

Squamous Metaplasia

Mild

[Adenomyosis TGLs = 2 - 16+17]

[Hyperplasia TGLs = 1 - 16+17+18]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 332

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403091

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|------------------|------------------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Hyperplasia | Focal, Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Dilation | Minimal |
| | | Eosinophilic Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Squamous Metaplasia | Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 333

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403092

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Trachea
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Urinary Bladder
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- Vagina
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Pituitary Gland
- * Thyroid Gland

OBSERVATIONS

- * Adrenal Medulla
 - * Kidney
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 2 - 7]
 - * Ovary
 - * Spleen
 - * Stomach, Glandular
 - * Thymus
 - * Uterus
 - Endometrium
- [Adenocarcinoma TGLs = 1 - 16+17]

Pheochromocytoma Benign
 Nephropathy
 Angiectasis
 Infiltration Cellular
 Atrophy
 Extramedullary Hematopoiesis
 Pigment
 Mineral
 Atrophy
 Adenocarcinoma
 Inflammation
 Squamous Metaplasia
 Ulcer

Chronicprogr, Minimal
 Minimal
 Histiocyte, Minimal
 Mild
 Mild
 Mild
 Minimal
 Mild
 Chronic Active, Moderate
 Marked
 Minimal

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 334

TRT#: 6

SEX: Female

DAY ON TEST: 360

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403093

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Ovary
- * Salivary Glands
- * Thyroid Gland
- Vagina
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Pituitary Gland
- * Stomach, Glandular
- * Uterus

OBSERVATIONS

- * Bone Marrow
 - * Kidney
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 3 - 6+7]
 - * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 18]
 - * Spleen
 - [Accessory Spleen TGLs = 2 - 19]
 - * Thymus
- Hypercellularity Moderate
- Nephropathy Chronicprogr, Minimal
- Necrosis Mild
- Infiltration Cellular Histiocyte, Minimal
- Fibroadenoma
- Accessory Spleen
- Extramedullary Hematopoiesis Moderate
- Atrophy Mild

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 335

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403094

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- * Urinary Bladder
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thymus
- Vagina
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Thyroid Gland
- * Bone Marrow
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Intestine Large, Colon
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Mammary Gland
- [Adenoma TGLs = 1 - 18]
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 3 - 11]
- * Spleen
- * Uterus
- Lymphoid Tiss
- Bile Duct
- Pars Distalis
- Endometrium
- Degeneration
- Hyperplasia
- Nephropathy
- Dilation
- Infiltration Cellular
- Adenoma
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Hyperplasia
- Cystic, Minimal
- Mild
- Chronicprogr, Minimal
- Minimal
- Histiocyte, Moderate
- Mild
- Mild
- Cystic, Minimal

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 336

TRT#: 6

SEX: Female

DAY ON TEST: 488

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403095

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Ovary
- * Salivary Glands
- * Thymus
- Vagina
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Pituitary Gland
- * Stomach, Glandular
- * Uterus

OBSERVATIONS

- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Mammary Gland
- [Fibroadenoma TGLs = 1- 18]
- * Spleen
- * Thyroid Gland
- C Cell
- Hypercellularity
- Nephropathy
- Clear Cell Focus
- Infiltration Cellular
- Fibroadenoma
- Extramedullary Hematopoiesis
- Hyperplasia
- Marked
- Chronicprogr, Minimal
- Histiocyte, Minimal
- Moderate
- Moderate

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 337

TRT#: 6

SEX: Female

DAY ON TEST: 726

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403096

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Stomach, Forestomach | * Stomach, Glandular | * Trachea | * Urinary Bladder |
| Vagina | | | |

MISSING

- * Clitoral Gland

OBSERVATIONS

- | | | | |
|-------------------|--------------------------------------------------|--------------------------------|-----------------------|
| * Adrenal Cortex | | Hypertrophy | Focal, Mild |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Extramedullary Hematopoiesis | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 5 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Ovary | | Atrophy | Moderate |
| | | Granulosa Cell Tumor Malignant | |
| | | Hyperplasia | Moderate |
| | Interstit Cell | | |
| | [Granulosa Cell Tumor Malignant TGLs = 2 - 14] | | |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 4 - 11] | | |
| * Skin | | Inflammation | Chronic, Mild |
| * Spleen | | Extramedullary Hematopoiesis | Marked |
| | [Extramedullary Hematopoiesis TGLs = 3 - 8] | | |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Adenoma | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 337

TRT#: 6

SEX: Female

DAY ON TEST: 726

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403096

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Endometrium

Hyperplasia

Cystic, Minimal

PRIMARY CAUSE OF DEATH

- Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 338

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403097

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Nose
- * Parathyroid Gland
- * Stomach, Forestomach
- * Stomach, Glandular
- * Uterus
- * Vagina
- * Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Trachea
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Urinary Bladder

OBSERVATIONS

- * Kidney Nephropathy Chronicprogr, Mild
- * Liver Clear Cell Focus
- Bile Duct Cyst
- Bile Duct Eosinophilic Focus
- [Clear Cell Focus TGLs = 3 - 19] Hyperplasia Minimal
- [Cyst TGLs = 2,4 - 18+20]
- * Lung Infiltration Cellular Histiocyte, Moderate
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Mammary Gland Hyperplasia Minimal
- * Ovary Atrophy Mild
- * Pancreas Basophilic Focus
- * Pituitary Gland Pars Distalis Adenoma
- * Spleen Extramedullary Hematopoiesis Mild
- Pigment Minimal
- * Thymus Atrophy Mild
- * Thyroid Gland Follicular Cel Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 339

TRT#: 6

SEX: Female

DAY ON TEST: 538

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403098

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Nose
- * Pancreas
- * Pituitary Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea
- * Urinary Bladder
- Vagina

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Bone Marrow
- Hypercellularity
- Moderate
- * Kidney
- Nephropathy
- Chronicprogr, Minimal
- * Liver
- Eosinophilic Focus
- Bile Duct
- Hyperplasia
- Minimal
- * Lung
- Infiltration Cellular
- Histiocyte, Minimal
- * Mammary Gland
- Fibroadenoma
- [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- Follicle
- Cyst
- * Spleen
- Extramedullary Hematopoiesis
- Moderate
- Pigment
- Minimal
- * Thymus
- Atrophy
- Moderate
- * Uterus
- Endometrium
- Hyperplasia
- Cystic, Mild
- Inflammation
- Acute, Mild
- Squamous Metaplasia
- Mild

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 340

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403099

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|---------------|------------------------------|------------------------|
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Clear Cell Focus | |
| | Bile Duct | Dilation | Minimal |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Inflammation | Granulomatous, Minimal |
| * Mammary Gland | | Fibroadenoma | |
| * Ovary | | Atrophy | Mild |
| | Follicle | Cyst | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Mild |
| | | Inflammation | Acute, Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 341

TRT#: 6

SEX: Female

DAY ON TEST: 668

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403100

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Mammary Gland
- * Nose
- * Salivary Glands
- * Skin
- * Thyroid Gland
- * Trachea
- Vagina
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- * Uterus

OBSERVATIONS

- * Kidney
- * Liver
- * Lung
- * Ovary
- [Cyst TGLs = 1 - 14]
- * Pituitary Gland
- [Adenoma TGLs = 2 - 11]
- * Spleen
- * Thymus
- Bile Duct
- Bursa
- Corpus Luteum
- Pars Distalis
- Nephropathy
- Cyst
- Infiltration Cellular
- Cyst
- Cyst
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Chronicprogr, Minimal
- Histiocyte, Minimal
- Minimal
- Mild
- Mild

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 342

TRT#: 6

SEX: Female

DAY ON TEST: 538

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403101

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Skin
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Heart
- * Kidney
- [Amphophilic/Vacuolar Adenoma TGLs = 2,3,4 - 8+19+20]
- * Liver
- * Lung
- Mesentery
- [Adenocarcinoma TGLs = 6- 22]
- * Nose
- Respirat Epith
- * Ovary
- Peritoneum
- * Pituitary Gland
- [Adenoma TGLs = 7 - 11]
- * Spleen
- Adenocarcinoma
- Hypercellularity
- Cardiomyopathy
- Amphophilic/Vacuolar Adenoma
- Amphophilic/Vacuolar Carcinoma
- Amphophilic/Vacuolar Hyperplasia
- Nephropathy
- Extramedullary Hematopoiesis
- Infiltration Cellular
- Adenocarcinoma
- Foreign Body
- Hyperplasia
- Inflammation
- Atrophy
- Adenocarcinoma
- Adenoma
- Adenocarcinoma
- Extramedullary Hematopoiesis
- Hemorrhage
- Metastatic (Uterus)
- Marked
- Minimal
- Multiple
- Multiple
- Mild
- Chronicprogr, Minimal
- Minimal
- Histiocyte, Mild
- Metastatic (Uterus)
- Mild
- Acute, Marked
- Mild
- Metastatic (Uterus)
- Metastatic (Uterus)
- Moderate
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 342

TRT#: 6

SEX: Female

DAY ON TEST: 538

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403101

ORGAN AND ACCOUNTABLE SITE STATUS

[Hemorrhage TGLs = 5 - 8+21]

* Thymus

Atrophy

Moderate

* Thyroid Gland

C Cell

Adenoma

* Uterus

Endometrium

Adenocarcinoma

Note: Little uterus present on slide 18 (all = tumor), however, a tiny portion is present in B16, possibly the origin).

[Adenocarcinoma TGLs = 1 - 18]

PRIMARY CAUSE OF DEATH - Kidney Amphophilic/Vacuolar Carcinoma

CONTRIBUTORY CAUSE OF DEATH - Uterus Endometrium Adenocarcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 343

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403102

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |
| Vagina | | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|------------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| | | Inflammation | Granulomatous, Minimal |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Ovary | | Inflammation | Chronic Active, Marked |
| | [Inflammation TGLs = 1 - 14] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Hyperplasia | Atypical, Moderate |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 344

TRT#: 6

SEX: Female

DAY ON TEST: 729

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403103

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | Vagina |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|---------------------------|------------------------------------------|------------------------------|------------------------|
| * Intestine Large, Rectum | | Parasite Metazoan | |
| * Islets, Pancreatic | | Hyperplasia | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Cyst | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | | Inflammation | Granulomatous, Minimal |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Adenoma | |
| * Uterus | | Squamous Metaplasia | Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 345

TRT#: 6

SEX: Female

DAY ON TEST: 638

DOSE: 250 mg/kg female

DISP: Natural Death

HISTO: 1403104

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Skin
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Stomach, Forestomach
- Vagina

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Heart
- * Kidney
- * Lung
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Nose
- * Ovary
- [Cyst TGLs = 2 - 17]
- * Pituitary Gland
- * Spleen
- * Thymus
- * Thyroid Gland
- * Uterus
- Hyperplasia
- Hypercellularity
- Cardiomyopathy
- Nephropathy
- Infiltration Cellular
- Inflammation
- Focal, Mild
- Mild
- Minimal
- Chronicprogr, Mild
- Histiocyte, Moderate
- Granulomatous, Minimal
- Respirat Epith
- Periovarn Tiss
- Pars Distalis
- C Cell
- Cervix
- Hyperplasia
- Atrophy
- Cyst
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Adenoma
- Papilloma
- Squamous Cell Carcinoma
- Squamous Cell Carcinoma
- Squamous Metaplasia
- Ulcer
- Mild
- Mild
- Moderate
- Mild
- Minimal
- Mild
- Squamous
- Marked
- Mild

Note: Second squamous cell carcinoma arising in cervix.

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 345

TRT#: 6

SEX: Female

DAY ON TEST: 638

DOSE: 250 mg/kg female

DISP: Natural Death

HISTO: 1403104

ORGAN AND ACCOUNTABLE SITE STATUS

Note: Squamous cell carcinoma in uterine horn.

Note: Small squamous cell papilloma also arising along uterine horn, distinct from other tumors.

[Squamous Cell Carcinoma TGLs = 3 - 18]

PRIMARY CAUSE OF DEATH - Uterus Squamous Cell Carcinoma

CONTRIBUTORY CAUSE OF DEATH - Uterus Cervix Squamous Cell Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 346

TRT#: 6

SEX: Female

DAY ON TEST: 677

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403105

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Ovary | * Pancreas | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Trachea | * Urinary Bladder |
| * Uterus | Vagina | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Clear Cell Focus | |
| | | Eosinophilic Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | Multiple |
| | [Fibroadenoma TGLs = 1,2 - 18+19] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Hyperplasia | Minimal |

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 347

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403106

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Salivary Glands
- * Stomach, Glandular
- * Thyroid Gland
- Vagina
- Blood Vessel
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Trachea
- * Bone
- * Esophagus
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Heart
- * Intestine Large, Rectum
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Mammary Gland
- * Ovary
- * Pituitary Gland
- Note: Extension of hyperplastic pars distalis into pars intermedia.
- * Spleen
- * Thymus
- * Uterus
- Bile Duct
- Pars Distalis
- Endometrium
- Cardiomyopathy
- Parasite Metazoan
- Nephropathy
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Galactocele
- Hyperplasia
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Hyperplasia
- Squamous Metaplasia
- Minimal
- Chronicprogr, Mild
- Minimal
- Histiocyte, Mild
- Granulomatous, Minimal
- Minimal
- Minimal
- Mild
- Mild
- Minimal
- Mild
- Minimal
- Cystic, Minimal
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 348

TRT#: 6

SEX: Female

DAY ON TEST: 605

DOSE: 250 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403107

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Brain
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Nose
- * Pancreas
- * Skin
- * Stomach, Forestomach
- * Urinary Bladder
- * Vagina
- * Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Kidney Nephropathy Chronicprogr, Minimal
- * Liver Bile Duct Hyperplasia Minimal
- * Lung Infiltration Cellular TGLs = 2 - 6+7 Infiltration Cellular Histocyte, Mild
- * Mammary Gland Galactocele Moderate
[Galactocele TGLs = 1 - 18]
- * Ovary Atrophy Moderate
- * Pituitary Gland Pars Distalis Adenoma
- * Spleen White Pulp Atrophy Mild
Extramedullary Hematopoiesis Mild
Pigment Mild
- * Thymus Atrophy Mild
- * Thyroid Gland C Cell Hyperplasia Marked
- * Uterus Endometrium Hyperplasia Cystic, Minimal

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 349

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403108

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thymus |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-------------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Extramedullary Hematopoiesis | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | | Inflammation | Chronic Active, Minimal |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 3 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | Cervix | Hyperplasia | Stromal, Moderate |
| | | Squamous Metaplasia | Minimal |
| | [Hyperplasia TGLs = 1 - 16] | | |

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 350

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403109

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lung
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Liver
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
 - * Bone Marrow
 - * Islets, Pancreatic
 - * Kidney
 - [Nephropathy TGLs = 4 - 8]
 - * Mammary Gland
 - [Fibroadenoma TGLs = 2 - 21]
 - [Galactocele TGLs = 1,3 - 20+22]
 - * Ovary
 - [Cyst TGLs = 7 - 14]
 - * Pituitary Gland
 - * Spleen
 - * Thymus
 - * Uterus
 - [Adenocarcinoma TGLs = 5 - 18]
- | | | |
|----------------|------------------------------|----------------------|
| | Hyperplasia | Focal, Minimal |
| | Hypercellularity | Mild |
| | Adenoma | |
| | Nephropathy | Chronicprogr, Marked |
| | Fibroadenoma | |
| | Galactocele | Marked |
| | Atrophy | Mild |
| Periovarn Tiss | Cyst | |
| | Hyperplasia | Moderate |
| Pars Distalis | Extramedullary Hematopoiesis | Mild |
| | Pigment | Minimal |
| | Atrophy | Minimal |
| Endometrium | Adenocarcinoma | |
| | Hyperplasia | Stromal, Mild |
| Endometrium | Hyperplasia | Cystic, Minimal |
| | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 350

TRT#: 6

SEX: Female

DAY ON TEST: 730

DOSE: 250 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403109

ORGAN AND ACCOUNTABLE SITE STATUS

[Hyperplasia TGLs = 6 - 19]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 351

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403110

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Pancreas |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Trachea | * Urinary Bladder | Vagina | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Clitoral Gland | | Squamous Metaplasia | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Nose | | Inflammation | Acute, Moderate |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Moderate |
| * Thyroid Gland | C Cell | Adenoma | |
| * Uterus | | Squamous Metaplasia | Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 352

TRT#: 8

SEX: Female

DAY ON TEST: 600

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403111

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Salivary Glands | * Skin | * Stomach, Glandular | * Trachea |
| * Urinary Bladder | Vagina | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|------------------------------------------|----------------|------------------------------|----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Mild |
| [Degeneration TGLs = 3 - 11] | | | |
| * Bone Marrow | | Hypercellularity | Moderate |
| * Intestine Large, Cecum | | Erosion | Mild |
| * Kidney | Renal Tubule | Apoptosis | Mild |
| | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Necrosis | Mild |
| [Necrosis TGLs = 2 - 19] | | | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| [Infiltration Cellular TGLs = 5 - 6+7] | | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| | | Pigment | Minimal |
| * Stomach, Forestomach | | Inflammation | Chronic Active, Mild |
| | | Mineral | Minimal |
| | | Ulcer | Mild |
| [Ulcer TGLs = 4 - 9] | | | |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | Follicular Cel | Hyperplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 352

TRT#: 8

SEX: Female

DAY ON TEST: 600

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403111

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Dilation

Marked

Hemorrhage

Marked

Inflammation

Acute, Marked

Squamous Metaplasia

Minimal

Ulcer

Moderate

[Dilation TGLs = 1 - 16+17+18]

PRIMARY CAUSE OF DEATH

- Uterus Hemorrhage

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 353

TRT#: 8

SEX: Female

DAY ON TEST: 629

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403112

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Liver |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Pancreas |
| * Parathyroid Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |
| * Uterus | Vagina | | |

OBSERVATIONS

- | | | | |
|----------------------|---------------------------|------------------------------|-----------------------|
| * Bone Marrow | | Hypercellularity | Mild |
| * Heart | | Cardiomyopathy | Minimal |
| * Islets, Pancreatic | | Hyperplasia | Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | | Squamous Metaplasia | Mild |
| * Nose | Nasolacrim Dct | Inflammation | Chronic, Minimal |
| * Ovary | | Atrophy | Moderate |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 1 - 11] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Minimal |

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 354

TRT#: 8

SEX: Female

DAY ON TEST: 475

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403113

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|---------------|------------------------------------------|-----------------------|------------------------|
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Moderate |
| * Kidney | Renal Tubule | Apoptosis | Moderate |
| | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Necrosis | Mild |
| * Lung | [Infiltration Cellular TGLs = 2 - 6+7] | Infiltration Cellular | Histiocyte, Mild |
| * Ovary | | Atrophy | Mild |
| * Spleen | White Pulp | Atrophy | Moderate |
| * Thymus | | Atrophy | Moderate |
| * Uterus | | Dilation | Moderate |
| | | Inflammation | Chronic Active, Marked |
| | | Squamous Metaplasia | Mild |
| | | Ulcer | Marked |
| | [Inflammation TGLs = 1 - 16+17+18] | | |

PRIMARY CAUSE OF DEATH - Uterus Inflammation

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 355

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403114

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Nose | * Pancreas | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Trachea |
| * Urinary Bladder | Vagina | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Mild |
| * Clitoral Gland | | Squamous Cell Carcinoma | |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Clear Cell Focus | |
| | | Eosinophilic Focus | |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| | [Infiltration Cellular TGLs = 4 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | Multiple |
| | [Fibroadenoma TGLs = 1,2,3 - 18+19+20] | | |
| * Ovary | | Atrophy | Moderate |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Adenoma | |
| | | Hyperplasia | Atypical, Mild |
| * Uterus | | Hyperplasia | Cystic, Minimal |
| | Endometrium | Hyperplasia | Squamous, Mild |
| | Cervix | Hyperplasia | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 355

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403114

ORGAN AND ACCOUNTABLE SITE STATUS

Squamous Metaplasia

Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 356

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403115

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Medulla
- * Bone Marrow
- * Kidney
- [Dilation TGLs = 2 - 18]
- * Liver
- [Cholangioma TGLs = 3 - 19]
- * Lung
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 5 - 21]
- * Spleen
- * Thymus
- * Uterus
- [Dilation TGLs = 4 - 17+20]
- Vagina
- [Squamous Cell Carcinoma TGLs = 6 - 16]
- Renal Tubule
- Pelvis
- Pars Distalis
- Hyperplasia
- Hypercellularity
- Dilation
- Dilation
- Edema
- Nephropathy
- Cholangioma
- Infiltration Cellular
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Atrophy
- Dilation
- Ulcer
- Squamous Cell Carcinoma
- Focal, Minimal
- Marked
- Mild
- Mild
- Focal, Mild
- Chronicprogr, Minimal
- Histiocyte, Minimal
- Mild
- Moderate
- Moderate
- Marked
- Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 357

TRT#: 8

SEX: Female

DAY ON TEST: 650

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403116

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- Vagina
- * Bone
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Glandular
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Trachea
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Brain
- * Heart
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 5 - 6+7]
- * Ovary
- [Cyst TGLs = 1 - 14]
- * Pituitary Gland
- * Spleen
- * Stomach, Forestomach
- [Cyst TGLs = 4 - 19]
- * Thymus
- * Thyroid Gland
- Pars Distalis
- White Pulp
- C Cell
- C Cell
- Degeneration
- Hypercellularity
- Gliosis
- Cardiomyopathy
- Nephropathy
- Polyarteritis Nodosa
- Necrosis
- Infiltration Cellular
- Squamous Metaplasia
- Atrophy
- Cyst
- Polyarteritis Nodosa
- Hyperplasia
- Atrophy
- Extramedullary Hematopoiesis
- Cyst
- Atrophy
- Adenoma
- Hyperplasia
- Cystic, Mild
- Moderate
- Moderate
- Minimal
- Chronicprogr, Moderate
- Mild
- Mild
- Histiocyte, Moderate
- Mild
- Minimal
- Mild
- Mild
- Mild
- Moderate
- Squamous
- Moderate
- Moderate
- Marked

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 357

TRT#: 8

SEX: Female

DAY ON TEST: 650

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403116

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Angiectasis

Marked

Dilation

Marked

Polyarteritis Nodosa

Mild

Squamous Metaplasia

Marked

Thrombus

Ulcer

Moderate

[Dilation TGLs = 2 - 17]

[Thrombus TGLs = 3 - 16+17+18]

PRIMARY CAUSE OF DEATH

- Uterus Thrombus

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 359

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403118

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Glandular
- Vagina
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Thyroid Gland

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Intestine Large, Rectum
- * Kidney
- * Liver
- * Lung
- * Mammary Gland
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- Angiectasis
- Hypercellularity
- Parasite Metazoan
- Nephropathy
- Clear Cell Focus
- Cyst
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Fibroadenoma
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Hyperplasia
- Squamous Metaplasia
- Minimal
- Mild
- Chronicprogr, Moderate
- Minimal
- Histiocyte, Moderate
- Granulomatous, Minimal
- Multiple
- Mild
- Moderate
- Mild
- Minimal
- Mild
- Atypical, Minimal
- Minimal

[Cyst TGLs = 3 - 20]

[Infiltration Cellular TGLs = 4 - 6+7]

[Fibroadenoma TGLs = 1,2 - 18+19]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 360

TRT#: 8

SEX: Female

DAY ON TEST: 4

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403119

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Kidney | * Liver |
| * Lung | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Nose |
| * Ovary | * Pancreas | * Parathyroid Gland | * Pituitary Gland |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Thymus | * Thyroid Gland | * Trachea | * Urinary Bladder |
| * Uterus | Vagina | | |

MISSING

- * Mammary Gland

OBSERVATIONS

- * Bone
Note: Apparent bone proliferation is due to age at death (4 days).
- * Bone Marrow
Note: Apparent marrow hyperplasia is due to age at death (4 days).
- * Spleen

Extramedullary Hematopoiesis

Mild

PRIMARY CAUSE OF DEATH

- UNCERTAIN

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 361

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403120

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Bone
- * Eye
- * Intestine Large, Colon
- * Islets, Pancreatic
- * Parathyroid Gland
- * Stomach, Glandular
Vagina
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Thyroid Gland
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Trachea
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Jejunum
- * Nose
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone Marrow
- * Intestine Large, Rectum
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Mammary Gland
- * Ovary
- * Pancreas
- * Pituitary Gland
- [Adenoma TGLs = 2 - 18]
- * Spleen
- * Thymus
- * Uterus
- Bilateral
- Follicle
- Pars Distalis
- Endometrium
- Angiectasis
- Hyperplasia
- Hypercellularity
- Polyarteritis Nodosa
- Nephropathy
- Eosinophilic Focus
- Infiltration Cellular
- Inflammation
- Squamous Metaplasia
- Hyperplasia
- Atrophy
- Cyst
- Polyarteritis Nodosa
- Adenoma
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Dilation
- Hyperplasia
- Squamous Metaplasia
- Mild
- Focal, Minimal
- Mild
- Mild
- Chronicprogr, Moderate
- Histiocyte, Moderate
- Granulomatous, Mild
- Minimal
- Mild
- Mild
- Mild
- Mild
- Moderate
- Moderate
- Moderate
- Cystic, Moderate
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 362

TRT#: 8

SEX: Female

DAY ON TEST: 501

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403121

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Urinary Bladder | Vagina | | |

MISSING

- | | | | |
|--------------|------------|----------------------|-------------------|
| Blood Vessel | * Heart | * Islets, Pancreatic | * Liver |
| * Lung | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Spleen | * Thymus | * Thyroid Gland | * Trachea |

OBSERVATIONS

- | | | | |
|--------------------------|---------------|---------------------|-----------------------|
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Marked |
| * Kidney | Pelvis | Inflammation | Chronic, Mild |
| | | Nephropathy | Chronicprogr, Minimal |
| * Lymph Node, Mandibular | | Atrophy | Moderate |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Uterus | | Squamous Metaplasia | Mild |

PRIMARY CAUSE OF DEATH

- Kidney Pelvis Inflammation

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 363

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403122

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Liver
- * Lung
 - [Infiltration Cellular TGLs = 1 - 6+7]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- Hyperplasia
- Hypercellularity
- Nephropathy
- Angiectasis
- Infiltration Cellular
- Atrophy
- Cyst
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Hyperplasia
- Squamous Metaplasia
- Focal, Mild
- Mild
- Chronicprogr, Mild
- Minimal
- Histiocyte, Marked
- Mild
- Mild
- Mild
- Minimal
- Moderate
- Cystic, Mild
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 364

TRT#: 8

SEX: Female

DAY ON TEST: 637

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403123

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Salivary Glands
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Stomach, Glandular
- * Bone Marrow
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Trachea
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Parathyroid Gland
- * Urinary Bladder

OBSERVATIONS

- * Adrenal Medulla
 - * Brain
 - Note: Astrocytosis in B04 probably secondary to damage from pituitary tumor.
 - * Kidney
 - * Lung
 - [Infiltration Cellular TGLs = 3 - 6+7]
 - * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 18]
 - * Ovary
 - Follicle
 - Acinus
 - * Pancreas
 - Pars Distalis
 - * Pituitary Gland
 - [Adenoma TGLs = 4 - 11]
 - * Skin
 - [Squamous Cell Carcinoma TGLs = 2 - 19]
 - * Spleen
 - * Stomach, Forestomach
 - Epithelium
- Hyperplasia
Giosis
Nephropathy
Infiltration Cellular
Inflammation
Squamous Metaplasia
Fibroadenoma
Atrophy
Cyst
Atrophy
Polyarteritis Nodosa
Adenoma
Squamous Cell Carcinoma
Extramedullary Hematopoiesis
Pigment
Hyperplasia
Inflammation
- Focal, Mild
Minimal
Chronicprogr, Moderate
Histiocyte, Mild
Granulomatous, Minimal
Minimal
Mild
Marked
Mild
Minimal
Minimal
Moderate
Chronic, Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 364

TRT#: 8

SEX: Female

DAY ON TEST: 637

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403123

ORGAN AND ACCOUNTABLE SITE STATUS

* Thymus		Polyarteritis Nodosa	Mild
* Thyroid Gland	C Cell	Atrophy	Minimal
* Uterus		Hyperplasia	Mild
		Inflammation	Chronic Active, Mild

PRIMARY CAUSE OF DEATH - Pituitary Gland Pars Distalis Adenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 365

TRT#: 8

SEX: Female

DAY ON TEST: 617

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403124

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Heart |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lung |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Parathyroid Gland | * Salivary Glands | * Skin |
| * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland | * Trachea |
| * Urinary Bladder | Vagina | | |

OBSERVATIONS

- | | | | |
|-------------------|---------------------------------------|-------------------------------|-----------------------|
| * Adrenal Cortex | | Leukemia Mononuclear | |
| * Adrenal Medulla | | Leukemia Mononuclear | |
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | Renal Tubule | Accumulation, Hyaline Droplet | Minimal |
| | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| | | Leukemia Mononuclear | |
| | | Necrosis | Minimal |
| * Ovary | Follicle | Atrophy | Mild |
| | | Cyst | |
| | [Cyst TGLs = 2 - 14] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Leukemia Mononuclear | |
| | [Leukemia Mononuclear TGLs = 1 - 8] | | |
| * Thymus | | Atrophy | Moderate |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | Bilateral | Polyp Stromal | |
| | | Squamous Metaplasia | Minimal |

PRIMARY CAUSE OF DEATH - Spleen Leukemia Mononuclear

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 366

TRT#: 8

SEX: Female

DAY ON TEST: 578

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403125

ORGAN AND ACCOUNTABLE SITE STATUS

* Ovary		Carcinoma	Metastatic (Kidney)
	Follicle	Cyst	
	[Carcinoma TGLs = 5 - 14]		
	[Cyst TGLs = 6 - 14]		
* Spleen		Extramedullary Hematopoiesis	Moderate
* Thymus		Atrophy	Mild
* Thyroid Gland	Bilateral, C Cell	Adenoma	

PRIMARY CAUSE OF DEATH - Kidney Amphophilic/Vacuolar Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 367

TRT#: 8

SEX: Female

DAY ON TEST: 635

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403126

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|--------------------------|----------------------------|---------------------------|-----------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum |
| * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular |
| * Lymph Node, Mesenteric | * Mammary Gland | * Nose | * Pancreas |
| * Parathyroid Gland | * Salivary Glands | * Stomach, Forestomach | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | Bilateral | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Marked |
| * Heart | Endocardium | Hyperplasia | Schwann Cell, Mild |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Skin | | Fibroma | |
| | [Fibroma TGLs = 1 - 18] | | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Moderate |
| * Uterus | | Inflammation | Acute, Mild |
| | | Squamous Metaplasia | Mild |

PRIMARY CAUSE OF DEATH

- Skin Fibroma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 368

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403127

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Bone Marrow |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|------------------------|
| * Adrenal Medulla | | Hyperplasia | Focal, Minimal |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Degeneration | Cystic, Minimal |
| | | Eosinophilic Focus | |
| | Bile Duct | Hyperplasia | Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| | | Inflammation | Granulomatous, Minimal |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 2 - 18] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| * Uterus | | Adenomyosis | Moderate |
| | Endometrium | Hyperplasia | Cystic, Mild |
| | | Squamous Metaplasia | Mild |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 369

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403128

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Skin
- * Trachea

OBSERVATIONS

- * Bone Marrow
- * Kidney
- * Liver
- [Cyst TGLs = 3 - 19]
- * Lung
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Nose
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- [Dilation TGLs = 1 - 16+17+18]
- Bile Duct
- Pars Distalis
- Hypercellularity
- Nephropathy
- Cyst
- Infiltration Cellular
- Inflammation
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Atrophy
- Dilation
- Squamous Metaplasia
- Moderate
- Chronicprogr, Minimal
- Histiocyte, Marked
- Acute, Minimal
- Mild
- Moderate
- Mild
- Mild
- Marked
- Marked

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 372

TRT#: 8

SEX: Female

DAY ON TEST: 551

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403131

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland | * Nose |
| * Pancreas | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Trachea | * Urinary Bladder | Vagina |

INSUFFICIENT TISSUE

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|--------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| | | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Liver | | Eosinophilic Focus | |
| | [Eosinophilic Focus TGLs = 3 - 12] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Mild |
| * Thyroid Gland | C Cell | Adenoma | |
| * Uterus | | Dilation | Marked |
| | | Squamous Metaplasia | Marked |
| | | Ulcer | Mild |
| | [Dilation TGLs = 2 - 16+17+18+19] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 373

TRT#: 8

SEX: Female

DAY ON TEST: 570

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403132

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Skin
- Vagina

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Heart
- * Kidney
- [Apoptosis TGLs = 2 - 8]
- * Lung
- [Infiltration Cellular TGLs = 3 - 6+7]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Stomach, Glandular
- * Thymus
- * Thyroid Gland
- * Uterus
- Renal Tubule
- Pars Distalis
- C Cell
- Degeneration
- Hypercellularity
- Cardiomyopathy
- Apoptosis
- Nephropathy
- Infiltration Cellular
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Mineral
- Atrophy
- Hyperplasia
- Dilation
- Hemorrhage
- Inflammation
- Squamous Metaplasia
- Ulcer
- Cystic, Minimal
- Marked
- Minimal
- Mild
- Chronicprogr, Mild
- Histiocyte, Mild
- Mild
- Minimal
- Moderate
- Minimal
- Marked
- Moderate
- Marked
- Minimal
- Acute, Mild
- Marked
- Moderate

[Dilation TGLs = 1 - 18]

PRIMARY CAUSE OF DEATH - Uterus Ulcer

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 374

TRT#: 8

SEX: Female

DAY ON TEST: 478

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403133

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Urinary Bladder
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- Vagina
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Thyroid Gland
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Salivary Glands
- * Trachea

OBSERVATIONS

- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Stomach, Glandular
- * Thymus
- * Uterus
- Renal Tubule
- Bile Duct
- Bile Duct
- Pars Distalis
- Hypercellularity
- Apoptosis
- Nephropathy
- Cyst
- Extramedullary Hematopoiesis
- Hyperplasia
- Infiltration Cellular
- Inflammation
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Erosion
- Atrophy
- Dilation
- Hemorrhage
- Inflammation
- Squamous Metaplasia
- Thrombus
- Marked
- Mild
- Chronicprogr, Mild
- Minimal
- Minimal
- Histiocyte, Mild
- Acute, Mild
- Moderate
- Mild
- Moderate
- Mild
- Minimal
- Mild
- Moderate
- Acute, Moderate
- Minimal

PRIMARY CAUSE OF DEATH

- Uterus Inflammation

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 375

TRT#: 8

SEX: Female

DAY ON TEST: 532

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403134

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Pancreas | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Stomach, Glandular | * Thyroid Gland | * Trachea | * Urinary Bladder |
| Vagina | | | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-------------------|----------------|------------------------------|----------------------|
| * Bone Marrow | | Hypercellularity | Marked |
| * Heart | | Cardiomyopathy | Minimal |
| * Kidney | | Infarct | Mild |
| | | Inflammation | Chronic, Mild |
| | | Nephropathy | Chronicprogr, Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| * Nose | Respirat Epith | Hyperplasia | Minimal |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Minimal |
| * Uterus | | Dilation | Marked |
| | | Hemorrhage | Marked |
| | | Inflammation | Chronic Active, Mild |
| | | Squamous Metaplasia | Moderate |
| | | Ulcer | Minimal |

[Hemorrhage TGLs = 1 - 18]

PRIMARY CAUSE OF DEATH - Uterus Hemorrhage

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 376

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403135

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Parathyroid Gland
- * Thyroid Gland
- Blood Vessel
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Salivary Glands
- * Trachea
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Skin
- * Urinary Bladder
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Pancreas
- * Stomach, Glandular
- Vagina

OBSERVATIONS

- * Adrenal Cortex
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Mammary Gland
- [Galactocele TGLs = 1 - 19]
- * Nose
- * Ovary
- * Pituitary Gland
- * Spleen
- * Stomach, Forestomach
- * Thymus
- * Uterus
- Thrombus
- Nephropathy
- Eosinophilic Focus
- Infiltration Cellular
- Inflammation
- Galactocele
- Inflammation
- Atrophy
- Cyst
- Follicle
- Pars Distalis
- Epithelium
- Endometrium
- Chronicprogr, Moderate
- Histiocyte, Moderate
- Granulomatous, Minimal
- Marked
- Chronic, Minimal
- Mild
- Minimal
- Mild
- Minimal
- Minimal
- Minimal
- Mild
- Moderate
- Cystic, Minimal
- Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 377

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403136

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|------------------------|
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Mild |
| * Heart | | Schwannoma Malignant | |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | | Inflammation | Granulomatous, Minimal |
| | [Infiltration Cellular TGLs = 1 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| | | Cyst | |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| | [Adenoma TGLs = 2 - 18] | | |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Minimal |
| * Thymus | | Atrophy | Mild |
| | | Ectopic Parathyroid Gland | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Minimal |
| | | Hyperplasia | Atypical, Minimal |
| | | Polyp Stromal | |
| | | Squamous Metaplasia | Minimal |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 379

TRT#: 8

SEX: Female

DAY ON TEST: 571

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403138

ORGAN AND ACCOUNTABLE SITE STATUS

PRIMARY CAUSE OF DEATH

- Mammary Gland Fibroadenoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 380

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403139

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Brain | * Clitoral Gland | * Esophagus | * Eye |
| * Harderian Gland | * Heart | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|----------------------|
| * Bone Marrow | | Hypercellularity | Moderate |
| * Kidney | | Nephropathy | Chronicprogr, Mild |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Mammary Gland | | Fibroadenoma | |
| | [Fibroadenoma TGLs = 1 - 18] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Marked |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| * Thymus | | Atrophy | Moderate |
| * Uterus | | Dilation | Marked |
| | | Hemorrhage | Mild |
| | | Inflammation | Chronic Active, Mild |
| | | Squamous Metaplasia | Minimal |
| | [Dilation TGLs = 2 - 16+17+19] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 381

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403140

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Skin
- * Urinary Bladder
- * Bone
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Stomach, Glandular
- Vagina
- * Bone Marrow
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Salivary Glands
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Medulla
- * Kidney
- * Liver
- * Lung
- * Nose
- * Ovary
- * Pancreas
- * Pituitary Gland
- * Spleen
- * Stomach, Forestomach
- * Thymus
- * Uterus
- Pheochromocytoma Benign
- Nephropathy
- Angiectasis
- Clear Cell Focus
- Cyst
- Eosinophilic Focus
- Infiltration Cellular
- Inflammation
- Atrophy
- Cyst
- Polyarteritis Nodosa
- Hyperplasia
- Extramedullary Hematopoiesis
- Pigment
- Polyarteritis Nodosa
- Atrophy
- Hyperplasia
- Hyperplasia
- Chronicprogr, Moderate
- Minimal
- Histiocyte, Mild
- Chronic, Minimal
- Mild
- Mild
- Minimal
- Mild
- Mild
- Mild
- Mild
- Cystic, Minimal
- Squamous, Minimal

[Cyst TGLs = 2 - 18]

[Infiltration Cellular TGLs = 1 - 6+7]

Bile Duct

Nasolacrim Dct

Follicle

Pars Distalis

Endometrium

Cervix

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 381

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403140

ORGAN AND ACCOUNTABLE SITE STATUS

Squamous Metaplasia

Mild

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 382

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403141

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Intestine Large, Colon
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Trachea
- * Urinary Bladder
- Vagina

OBSERVATIONS

- Blood Vessel Carotid Artery Polyarteritis Nodosa Mild
- * Bone Marrow Hypercellularity Mild
- * Heart Cardiomyopathy Minimal
- Intestine Large, Cecum Polyarteritis Nodosa Mild
- * Intestine Large, Rectum Polyarteritis Nodosa Minimal
- * Kidney Renal Tubule Adenoma Moderate
- Nephropathy Chronicprogr, Marked
- * Liver Polyarteritis Nodosa Mild
- Bile Duct Angiectasis Minimal
- Bile Duct Cyst Cystic, Minimal
- Bile Duct Degeneration Mild
- Bile Duct Dilation Mild
- Bile Duct Eosinophilic Focus Mild
- [Cyst TGLs = 2 - 19; 4 - 12+20] Hyperplasia
- * Lung Infiltration Cellular Histiocyte, Mild
- Polyarteritis Nodosa Minimal
- * Mammary Gland Fibroadenoma
- [Fibroadenoma TGLs = 1 - 18]
- * Ovary Atrophy Marked
- * Pituitary Gland Pars Distalis Adenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 382

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403141

ORGAN AND ACCOUNTABLE SITE STATUS

Polyarteritis Nodosa

Mild

[Adenoma TGLs = 3 - 11]

Extramedullary Hematopoiesis

Mild

Pigment

Minimal

Atrophy

Moderate

Polyarteritis Nodosa

Minimal

Adenoma

Squamous Metaplasia

Minimal

* Spleen

* Thymus

* Thyroid Gland

* Uterus

C Cell

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 383

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403142

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Intestine Large, Cecum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Thyroid Gland
- Blood Vessel
- * Esophagus
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Skin
- * Trachea
- * Bone
- * Eye
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder
- * Brain
- * Harderian Gland
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Pituitary Gland
- * Stomach, Glandular

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Heart
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Kidney
- * Lung
 - [Infiltration Cellular TGLs = 3 - 6+7]
- * Mammary Gland
 - [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Spleen
- * Thymus
- * Uterus
 - Cervix
 - [Squamous Metaplasia TGLs = 2 - 19]
 - Vagina
- Angiectasis
- Hypercellularity
- Cardiomyopathy
- Parasite Metazoan
- Parasite Metazoan
- Nephropathy
- Infiltration Cellular
- Fibroadenoma
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Hyperplasia
- Squamous Metaplasia
- Hyperplasia
- Minimal
- Mild
- Minimal
- Chronicprogr, Moderate
- Histiocyte, Mild
- Moderate
- Mild
- Minimal
- Mild
- Squamous, Marked
- Marked
- Squamous, Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 384

TRT#: 8

SEX: Female

DAY ON TEST: 641

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403143

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Trachea
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Pituitary Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Thyroid Gland

MISSING

- * Intestine Large, Cecum

OBSERVATIONS

- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- * Ovary
- * Spleen
- * Thymus
- * Uterus
- Bile Duct
- [Infiltration Cellular TGLs = 2 - 6+7]
- [Adenocarcinoma TGLs = 1 - 16+17+18]
- [Dilatation TGLs = 3 - 18]
- Hemorrhage
- Hypercellularity
- Nephropathy
- Hyperplasia
- Infiltration Cellular
- Atrophy
- Extramedullary Hematopoiesis
- Pigment
- Atrophy
- Adenocarcinoma
- Dilatation
- Ulcer
- Moderate
- Mild
- Chronicprogr, Minimal
- Minimal
- Histiocyte, Mild
- Minimal
- Mild
- Minimal
- Mild
- Marked
- Moderate

PRIMARY CAUSE OF DEATH - Uterus Endometrium Adenocarcinoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 385

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403144

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Eosinophilic Focus | |
| | | Hepatocellular Adenoma | |
| | [Hepatocellular Adenoma TGLs = 1 - 18] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Moderate |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pancreas | Acinus | Adenoma | |
| | [Adenoma TGLs = 3 - 19] | | |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Moderate |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Mild |
| * Uterus | Endometrium | Hyperplasia | Cystic, Mild |
| | | Squamous Metaplasia | Minimal |

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 386

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403145

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Eye
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Mammary Gland
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Salivary Glands
- * Skin
- * Stomach, Forestomach
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea
- * Urinary Bladder
- * Vagina

OBSERVATIONS

- * Bone Marrow Hemorrhage Mild
- * Kidney Nephropathy Chronicprogr, Mild
- * Liver Bile Duct Cyst Eosinophilic Focus
- [Cyst TGLs = 1 - 18]
- [Eosinophilic Focus TGLs = 3 - 19]
- * Lung Infiltration Cellular Histiocyte, Mild
Inflammation Granulomatous, Minimal
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Ovary Atrophy Mild
- * Pituitary Gland Pars Distalis Adenoma
- * Spleen Extramedullary Hematopoiesis Mild
Pigment Mild
- * Thymus Atrophy Minimal
- * Uterus Endometrium Hyperplasia Cystic, Minimal
Hyperplasia Atypical, Minimal
Squamous Metaplasia Minimal

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 387

TRT#: 8

SEX: Female

DAY ON TEST: 729

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403146

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Bone |
| * Bone Marrow | * Brain | * Clitoral Gland | * Esophagus |
| * Eye | * Harderian Gland | * Heart | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric |
| * Mammary Gland | * Nose | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Trachea | * Urinary Bladder | Vagina | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Kidney | | Infarct | Minimal |
| | | Nephropathy | Chronicprogr, Minimal |
| * Liver | | Eosinophilic Focus | |
| | [Eosinophilic Focus TGLs = 1 - 18] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Mild |
| | [Infiltration Cellular TGLs = 2 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pancreas | | Inflammation | Chronic, Minimal |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Mild |
| * Spleen | | Extramedullary Hematopoiesis | Mild |
| | | Pigment | Mild |
| * Thymus | | Atrophy | Minimal |
| * Uterus | | Squamous Metaplasia | Moderate |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 388

TRT#: 8

SEX: Female

DAY ON TEST: 625

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403147

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Adrenal Medulla
- * Blood Vessel
- * Bone
- * Brain
- * Clitoral Gland
- * Esophagus
- * Harderian Gland
- * Heart
- * Intestine Large, Cecum
- * Intestine Large, Colon
- * Intestine Large, Rectum
- * Intestine Small, Duodenum
- * Intestine Small, Ileum
- * Intestine Small, Jejunum
- * Islets, Pancreatic
- * Lymph Node, Mandibular
- * Lymph Node, Mesenteric
- * Nose
- * Pancreas
- * Parathyroid Gland
- * Pituitary Gland
- * Salivary Glands
- * Skin
- * Stomach, Glandular
- * Thyroid Gland
- * Trachea
- * Urinary Bladder
- Vagina

OBSERVATIONS

- * Bone Marrow
 - * Eye
 - * Kidney
 - * Liver
 - * Lung
 - [Infiltration Cellular TGLs = 4 - 6+7]
 - * Mammary Gland
 - [Fibroadenoma TGLs = 1,2 - 18+19]
 - * Ovary
 - * Spleen
 - * Stomach, Forestomach
 - * Thymus
 - * Uterus
 - [Dilation TGLs = 3 - 16+17+20]
- Hypercellularity Marked
- Inflammation Acute, Minimal
- Nephropathy Chronicprogr, Minimal
- Necrosis Mild
- Infiltration Cellular Histiocyte, Moderate
- Thrombus
- Fibroadenoma Multiple
- Atrophy Mild
- Extramedullary Hematopoiesis Moderate
- Mineral Minimal
- Atrophy Moderate
- Dilation Marked
- Hemorrhage Moderate
- Thrombus

PRIMARY CAUSE OF DEATH - Uterus Hemorrhage

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 389

TRT#: 8

SEX: Female

DAY ON TEST: 509

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403148

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Salivary Glands
- * Thyroid Gland
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Skin
- * Trachea
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina

OBSERVATIONS

- * Bone Marrow
- * Kidney
- * Lung
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Ovary
- * Pituitary Gland
- * Spleen
- * Thymus
- * Uterus
- [Dilation TGLs = 1 - 16+17+18+19]
- Capsule
- Pars Distalis
- Hypercellularity
- Inflammation
- Nephropathy
- Infiltration Cellular
- Thrombus
- Atrophy
- Hyperplasia
- Extramedullary Hematopoiesis
- Atrophy
- Dilation
- Squamous Metaplasia
- Marked
- Chronic, Minimal
- Chronicprogr, Minimal
- Histiocyte, Mild
- Mild
- Minimal
- Mild
- Mild
- Marked
- Marked

PRIMARY CAUSE OF DEATH - UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 390

TRT#: 8

SEX: Female

DAY ON TEST: 661

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403149

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|---------------------------|-----------------------------|--------------------------|----------------------------|
| Blood Vessel | * Bone | * Brain | * Esophagus |
| * Eye | * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon |
| * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum |
| * Islets, Pancreatic | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Parathyroid Gland |
| * Pituitary Gland | * Salivary Glands | * Skin | * Stomach, Forestomach |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

OBSERVATIONS

- | | | |
|--------------------------------------------------------|-------------------------|------------------------|
| * Adrenal Cortex | Necrosis | Minimal |
| * Adrenal Medulla | Hyperplasia | Focal, Minimal |
| * Bone Marrow | Hypercellularity | Moderate |
| * Clitoral Gland | Squamous Cell Carcinoma | |
| * Esophagus | | |
| Note: Esophagus contains feed material (not a lesion). | | |
| * Heart | Cardiomyopathy | Moderate |
| * Kidney | Nephropathy | Chronicprogr, Moderate |
| [Nephropathy TGLs = 5 - 8] | | |
| * Liver | Adenocarcinoma | Metastatic (Uterus) |
| [Adenocarcinoma TGLs = 6 - 21] | Eosinophilic Focus | |
| * Lung | Infiltration Cellular | Histiocyte, Mild |
| [Inflammation TGLs = 7 - 6+7] | Inflammation | Acute, Marked |
| * Mammary Gland | Hyperplasia | Marked |
| * Nose | Foreign Body | |
| | Inflammation | Suppurative, Marked |
| * Ovary | Adenocarcinoma | Metastatic (Uterus) |
| [Adenocarcinoma TGLs = 4 - 14] | Atrophy | Mild |
| * Pancreas | Adenocarcinoma | Metastatic (Uterus) |
| [Adenocarcinoma TGLs = 1 - 18] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 390

TRT#: 8

SEX: Female

DAY ON TEST: 661

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403149

ORGAN AND ACCOUNTABLE SITE STATUS

* Spleen		Adenocarcinoma	Metastatic (Uterus)
	White Pulp	Atrophy	Moderate
		Extramedullary Hematopoiesis	Mild
		Pigment	Minimal
	[Adenocarcinoma TGLs = 3 - 20]		
* Stomach, Glandular		Adenocarcinoma	Metastatic (Uterus)
	[Adenocarcinoma TGLs = 2 - 19]		
* Thymus		Atrophy	Moderate
* Uterus	Endometrium	Adenocarcinoma	
		Polyp Stromal	
		Squamous Metaplasia	Mild

PRIMARY CAUSE OF DEATH - Uterus Endometrium Adenocarcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 391

TRT#: 8

SEX: Female

DAY ON TEST: 702

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403150

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Salivary Glands
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Nose
- * Skin
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Pancreas
- * Trachea

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Liver
- * Lung
- [Infiltration Cellular TGLs = 5 - 6+7]
- * Mammary Gland
- [Adenocarcinoma TGLs = 1 - 18]
- * Ovary
- * Pituitary Gland
- [Adenoma TGLs = 2 - 11]
- * Spleen
- [Extramedullary Hematopoiesis TGLs = 4 - 8]
- * Stomach, Forestomach
- Note: Peculiar section has isolated epithelium apparently within muscularis: sectioning artifact!
- * Stomach, Glandular
- * Thymus
- * Thyroid Gland
- * Uterus
- Pars Distalis
- Atrophy
- Adenoma
- Extramedullary Hematopoiesis
- Mineral
- Mineral
- Atrophy
- Hyperplasia
- Cyst
- Hyperplasia
- Squamous Metaplasia
- Cystic, Minimal
- Marked
- Chronicprogr, Mild
- Mild
- Histiocyte, Mild
- Granulomatous, Minimal
- Mild
- Marked
- Minimal
- Mild
- Mild
- Minimal
- Squamous
- Cystic, Mild
- Mild

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 391

TRT#: 8

SEX: Female

DAY ON TEST: 702

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403150

ORGAN AND ACCOUNTABLE SITE STATUS

Note: B19 = uterine cyst grossly - although no uterine mucosa is present on slide 19, the content of the cyst indicates it was squamous.

[Cyst TGLs = 3 - 19]

PRIMARY CAUSE OF DEATH

- Mammary Gland Adenocarcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 393

TRT#: 8

SEX: Female

DAY ON TEST: 586

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403152

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Cortex | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Kidney | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Pancreas |
| * Salivary Glands | * Skin | * Stomach, Forestomach | * Stomach, Glandular |
| * Trachea | * Urinary Bladder | Vagina | |

MISSING

- * Parathyroid Gland

OBSERVATIONS

- | | | | |
|-----------------------------------------------------------------------------------------------------------------|-------------------------------------------------|------------------------------|----------------------------|
| * Adrenal Medulla | Bilateral | Hyperplasia | Focal, Minimal |
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Marked |
| * Liver | | Necrosis | Mild |
| * Lung | | Adenocarcinoma | Metastatic (Mammary Gland) |
| | | Infiltration Cellular | Histiocyte, Mild |
| Note: Several metastases have squamous differentiation as well as fat globules characteristic of mammary gland. | | | |
| Note: The malignant thymoma is also invading into the lung (B19). | | | |
| | [Adenocarcinoma TGLs = 2 - 19; 8 - 6+7] | | |
| * Mammary Gland | | Adenocarcinoma | |
| | [Adenocarcinoma TGLs = 1 - 18] | | |
| * Nose | | Foreign Body | |
| | | Inflammation | Chronic Active, Minimal |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Hyperplasia | Minimal |
| * Spleen | | Extramedullary Hematopoiesis | Marked |
| * Thymus | | Thymoma Malignant | |
| | [Thymoma Malignant TGLs = 3,4,5,6 - 19+20+21] | | |
| * Thyroid Gland | C Cell | Adenoma | |
| | [Adenoma TGLs = 7 - 11] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 393

TRT#: 8

SEX: Female

DAY ON TEST: 586

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403152

ORGAN AND ACCOUNTABLE SITE STATUS

* Uterus

Endometrium

Hyperplasia

Cystic, Minimal

Squamous Metaplasia

Minimal

PRIMARY CAUSE OF DEATH

- Thymus Thymoma Malignant

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 394

TRT#: 8

SEX: Female

DAY ON TEST: 540

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403153

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pituitary Gland
- * Stomach, Glandular
- Vagina
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Skin
- * Trachea

MISSING

- * Parathyroid Gland

OBSERVATIONS

- * Adrenal Cortex
 - * Bone Marrow
 - * Kidney
 - * Lung
 - [Infiltration Cellular TGLs = 1 - 6+7]
 - * Nose
 - * Ovary
 - * Spleen
 - * Thymus
 - * Uterus
 - [Dilation TGLs = 2 - 16+17+18]
- Nasolacrim Dct
 - White Pulp
- Necrosis
 - Hemorrhage
 - Hypercellularity
 - Nephropathy
 - Infiltration Cellular
 - Inflammation
 - Atrophy
 - Atrophy
 - Extramedullary Hematopoiesis
 - Atrophy
 - Dilation
 - Squamous Metaplasia
- Moderate
 - Minimal
 - Marked
 - Chronicprogr, Minimal
 - Histiocyte, Mild
 - Chronic, Minimal
 - Mild
 - Mild
 - Moderate
 - Mild
 - Marked
 - Moderate

PRIMARY CAUSE OF DEATH - UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 395

TRT#: 8

SEX: Female

DAY ON TEST: 702

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403154

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|----------------------------|---------------------------|-----------------------------|--------------------------|
| Blood Vessel | * Bone | * Brain | * Clitoral Gland |
| * Esophagus | * Eye | * Harderian Gland | * Intestine Large, Cecum |
| * Intestine Large, Colon | * Intestine Large, Rectum | * Intestine Small, Duodenum | * Intestine Small, Ileum |
| * Intestine Small, Jejunum | * Islets, Pancreatic | * Lymph Node, Mandibular | * Parathyroid Gland |
| * Pituitary Gland | * Salivary Glands | * Skin | * Stomach, Glandular |
| * Thyroid Gland | * Trachea | * Urinary Bladder | Vagina |

MISSING

- * Lymph Node, Mesenteric

OBSERVATIONS

- | | | | |
|-----------------------------------------------------------------|---------------------------|-------------------------|------------------------|
| * Adrenal Cortex | Capsule | Squamous Cell Carcinoma | Metastatic (Uterus) |
| * Adrenal Medulla | | Hyperplasia | Focal, Mild |
| * Bone Marrow | | Hypercellularity | Marked |
| * Heart | | Squamous Cell Carcinoma | Metastatic (Uterus) |
| Note: Free-floating metastatic cell cluster in right ventricle. | | | |
| * Kidney | | Nephropathy | Chronicprogr, Moderate |
| * Liver | Bile Duct | Hyperplasia | Moderate |
| | | Necrosis | Mild |
| | | Squamous Cell Carcinoma | Metastatic (Uterus) |
| [Squamous Cell Carcinoma | TGLs = 10,11,12 - 12+27] | | |
| * Lung | | Infiltration Cellular | Histiocyte, Minimal |
| | | Squamous Cell Carcinoma | Metastatic (Uterus) |
| [Squamous Cell Carcinoma | TGLs = 13 - 6+7] | | |
| Lymph Node | Renal | Squamous Cell Carcinoma | Metastatic (Uterus) |
| | Mediastinal | Squamous Cell Carcinoma | Metastatic (Uterus) |
| | Lumbar | Squamous Cell Carcinoma | Metastatic (Uterus) |
| [Squamous Cell Carcinoma | TGLs = 6 - 23] | | |
| [Squamous Cell Carcinoma | TGLs = 5 - 22] | | |
| [Squamous Cell Carcinoma | TGLs = 4 - 21] | | |
| * Mammary Gland | | Fibroadenoma | |
| [Fibroadenoma | TGLs = 9 - 26] | | |

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 395

TRT#: 8

SEX: Female

DAY ON TEST: 702

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403154

ORGAN AND ACCOUNTABLE SITE STATUS

Mesentery		Squamous Cell Carcinoma	Metastatic (Uterus)
Note: B09, mesojejunum.			
* Nose	Turbinate	Atrophy Inflammation	Minimal Chronic, Mild
Note: Changes appear secondary to tooth (pulp) inflammation and penetration into nasal cavity. Septum also perforated.			
* Ovary		Atrophy Squamous Cell Carcinoma	Marked Metastatic (Uterus)
* Pancreas		Squamous Cell Carcinoma	Metastatic (Uterus)
[Squamous Cell Carcinoma TGLs = 1- 18; 7 - 24]			
Skeletal Muscle		Squamous Cell Carcinoma	Metastatic (Uterus)
[Squamous Cell Carcinoma TGLs = 8 - 25]			
* Spleen		Extramedullary Hematopoiesis	Moderate
* Stomach, Forestomach	Epithelium	Hyperplasia	Mild
* Thymus		Squamous Cell Carcinoma	Metastatic (Uterus)
Tooth		Dysplasia Inflammation	Moderate Chronic, Mild
Note: Dysplastic incisor resulting in septal perforation and nasal inflammation.			
* Uterus	Endometrium	Dilation Hyperplasia Squamous Cell Carcinoma Squamous Metaplasia	Marked Cystic, Moderate Moderate
Note: Poorly-differentiated squamous cell carcinoma in B16, 17, 19, and 20 (B16 and 20 best diagnostically).			
[Dilation TGLs = 3 - 20]			
[Squamous Cell Carcinoma TGLs = 2 - 19]			

PRIMARY CAUSE OF DEATH - Uterus Squamous Cell Carcinoma

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 396

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403155

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Esophagus
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Mammary Gland
- * Pituitary Gland
- * Stomach, Glandular
- * Bone
- * Eye
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Salivary Glands
- * Trachea
- * Brain
- * Harderian Gland
- * Intestine Small, Duodenum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Urinary Bladder
- * Clitoral Gland
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- Vagina

OBSERVATIONS

- * Adrenal Medulla Bilateral Hyperplasia Focal, Moderate
- Blood Vessel Mineral Minimal
- Note: Gastric arteries are mineralized.
- * Bone Marrow Hypercellularity Marked
- * Heart Cardiomyopathy Minimal
- * Kidney Nephropathy Chronicprogr, Mild
- * Liver Bile Duct Cyst Minimal
- Bile Duct Hyperplasia Minimal
- * Lung Adenocarcinoma Metastatic (Uterus)
- Infiltration Cellular Histiocyte, Mild
- [Infiltration Cellular TGLs = 2 - 6+7]
- * Ovary Atrophy Mild
- * Spleen Extramedullary Hematopoiesis Moderate
- * Thymus Atrophy Minimal
- * Thyroid Gland C Cell Adenoma
- * Uterus Endometrium Adenocarcinoma Multiple
- Dilation Marked
- Hemangiosarcoma

Note: Hemangiosarcoma in B17. Endometrial carcinoma in B16 and B17. Endometrial adenomas in B18.

[Dilation TGLs = 1 - 16+17+18]

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 397

TRT#: 8

SEX: Female

DAY ON TEST: 498

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403156

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Cortex
- * Brain
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Mammary Gland
- * Pituitary Gland
- * Stomach, Glandular
- Vagina
- * Adrenal Medulla
- * Clitoral Gland
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Nose
- * Salivary Glands
- * Thyroid Gland
- Blood Vessel
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Pancreas
- * Skin
- * Trachea
- * Bone
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Parathyroid Gland
- * Stomach, Forestomach
- * Urinary Bladder

OBSERVATIONS

- * Bone Marrow
 - Hemorrhage
 - Hypercellularity
- * Kidney
 - Nephropathy
- * Lung
 - Infiltration Cellular
- [Infiltration Cellular TGLs = 1 - 6+7]
- * Ovary
 - Atrophy
- * Spleen
 - White Pulp
 - Atrophy
 - Extramedullary Hematopoiesis
 - Pigment
- * Thymus
 - Atrophy
- * Uterus
 - Squamous Metaplasia

PRIMARY CAUSE OF DEATH - UNCERTAIN

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 398

TRT#: 8

SEX: Female

DAY ON TEST: 730

DOSE: 750 mg/kg female

DISP: Terminal Sacrifice

HISTO: 1403157

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Adrenal Medulla | Blood Vessel | * Bone | * Brain |
| * Clitoral Gland | * Esophagus | * Eye | * Harderian Gland |
| * Heart | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Liver | * Lymph Node, Mandibular | * Lymph Node, Mesenteric | * Mammary Gland |
| * Nose | * Pancreas | * Parathyroid Gland | * Salivary Glands |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Trachea |
| * Urinary Bladder | Vagina | | |

OBSERVATIONS

- | | | | |
|-------------------|------------------------------------------|------------------------------|-----------------------|
| * Adrenal Cortex | | Degeneration | Cystic, Minimal |
| * Bone Marrow | | Hypercellularity | Marked |
| * Kidney | | Nephropathy | Chronicprogr, Minimal |
| * Lung | | Infiltration Cellular | Histiocyte, Marked |
| | [Infiltration Cellular TGLs = 3 - 6+7] | | |
| * Ovary | | Atrophy | Mild |
| * Pituitary Gland | Pars Distalis | Adenoma | |
| * Spleen | | Extramedullary Hematopoiesis | Moderate |
| * Thymus | | Atrophy | Minimal |
| * Thyroid Gland | C Cell | Adenoma | |
| * Uterus | | Dilation | Marked |
| | | Hyperplasia | Atypical, Minimal |
| | | Thrombus | |
| | | Ulcer | Minimal |

Note: One nodule of TGL-2 was inflammation in uterine wall, there were several thromboses in B16 and B19.

[Dilation TGLs = 1 - 16+17+18]

[Thrombus TGLs = 2 - 19]

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 399

TRT#: 8

SEX: Female

DAY ON TEST: 600

DOSE: 750 mg/kg female

DISP: Moribund Sacrifice

HISTO: 1403158

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- * Adrenal Medulla
- * Esophagus
- * Intestine Large, Cecum
- * Intestine Small, Ileum
- * Lymph Node, Mandibular
- * Parathyroid Gland
- * Stomach, Glandular
- Vagina
- Blood Vessel
- * Eye
- * Intestine Large, Colon
- * Intestine Small, Jejunum
- * Lymph Node, Mesenteric
- * Salivary Glands
- * Thyroid Gland
- * Bone
- * Harderian Gland
- * Intestine Large, Rectum
- * Islets, Pancreatic
- * Nose
- * Skin
- * Trachea
- * Brain
- * Heart
- * Intestine Small, Duodenum
- * Liver
- * Pancreas
- * Stomach, Forestomach
- * Urinary Bladder

MISSING

- * Clitoral Gland

OBSERVATIONS

- * Adrenal Cortex
- * Bone Marrow
- * Kidney
- * Lung
- [Infiltration Cellular TGLs = 5 - 6+7]
- * Mammary Gland
- [Adenocarcinoma TGLs = 2,3 - 15]
- [Fibroadenoma TGLs = 1 - 18]
- * Ovary
- * Pituitary Gland
- * Spleen
- [Extramedullary Hematopoiesis TGLs = 4 - 8]
- * Thymus
- * Uterus
- Hyperplasia
- Hypercellularity
- Nephropathy
- Infiltration Cellular
- Inflammation
- Adenocarcinoma
- Fibroadenoma
- Atrophy
- Cyst
- Hyperplasia
- Extramedullary Hematopoiesis
- Atrophy
- Squamous Metaplasia
- Focal, Moderate
- Marked
- Chronicprogr, Minimal
- Histiocyte, Moderate
- Granulomatous, Minimal
- Multiple
- Mild
- Mild
- Marked
- Marked
- Mild

PRIMARY CAUSE OF DEATH - Mammary Gland Fibroadenoma

* PROTOCOL REQUIRED TISSUE

Experiment Number: 00058 - 03

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/HSD

P14: INDIVIDUAL ANIMAL PATHOLOGY DATA

Black Cohosh

CAS Number: 84776-26-1

Date Report Requested: 11/13/2020

Time Report Requested: 09:33:45

First Dose M/F: 07/03/12 / 07/02/12

Lab: BAT

ANIMAL ID: 400

TRT#: 8

SEX: Female

DAY ON TEST: 719

DOSE: 750 mg/kg female

DISP: Natural Death

HISTO: 1403159

ORGAN AND ACCOUNTABLE SITE STATUS

NORMAL

- | | | | |
|-----------------------------|--------------------------|----------------------------|---------------------------|
| * Bone | * Brain | * Clitoral Gland | * Eye |
| * Harderian Gland | * Intestine Large, Cecum | * Intestine Large, Colon | * Intestine Large, Rectum |
| * Intestine Small, Duodenum | * Intestine Small, Ileum | * Intestine Small, Jejunum | * Islets, Pancreatic |
| * Lymph Node, Mesenteric | * Mammary Gland | * Pancreas | * Pituitary Gland |
| * Skin | * Stomach, Forestomach | * Stomach, Glandular | * Thyroid Gland |
| * Urinary Bladder | Vagina | | |

MISSING

- | | | | |
|--------------------------|-------------------|---------------------|-------------------|
| * Adrenal Cortex | * Adrenal Medulla | Blood Vessel | * Esophagus |
| * Heart | * Kidney | * Liver | * Lung |
| * Lymph Node, Mandibular | * Ovary | * Parathyroid Gland | * Salivary Glands |
| * Thymus | * Trachea | | |

AUTO PRECLUDES DIAG.

- * Uterus

OBSERVATIONS

- | | | | |
|---------------|------------|------------------------------|----------------|
| * Bone Marrow | | Hemorrhage | Minimal |
| | | Hypercellularity | Marked |
| * Nose | | Inflammation | Acute, Minimal |
| * Spleen | White Pulp | Atrophy | Moderate |
| | | Extramedullary Hematopoiesis | Moderate |

PRIMARY CAUSE OF DEATH

- UNCERTAIN

*** END OF REPORT ***