Experiment Number: 00058 - 04

Species/Strain: MICE/B6C3F1/N

Test Type: CHRONIC

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH Date Report Requested: 12/10/2020 AVERAGE SEVERITY GRADES[b]

Black Cohosh

CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

Lab: BAT

2-year Core Mice_Final 1

NTP Study Number: C00058B

Lock Date: 10/02/2018

Cage Range: ALL

Date Range: ALL

Reasons For Removal: 25022 ACCK 25021 TSAC 25020 NATD

25019 MSAC

Removal Date Range: ALL

Treatment Groups: Include ALL

Study Gender: Female

TDMSE Version: 3.0.2.3_002

PWG Approval Date: NONE

Test Type: CHRONIC Route: GAVAGE

Species/Strain: MICE/B6C3F1/N

Experiment Number: 00058 - 04

Black Cohosh CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

B6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Disposition Summary						
Animals Initially In Study	70	70	70	70	70	
Early Deaths						
Accidentally Killed				1		
Moribund Sacrifice	4	2	3	3	6	
Natural Death	6	8	3	7	6	
Survivors						
Natural Death		1				
Terminal Sacrifice	40	39	44	39	38	
Animals Examined Microscopically	50	50	50	50	50	
ALIMENTARY SYSTEM						
Esophagus	(50)	(50)	(50)	(50)	(50)	
Gallbladder	(46)	(50)	(50)	(49)	(50)	
Intestine Large, Cecum	(50)	(50)	(50)	(50)	(50)	
Intestine Large, Colon	(50)	(50)	(50)	(50)	(50)	
Inflammation, Acute	1 [2.0]	(2-)	()	(2-)	ζ/	
Ulcer	1 [2.0]					
Intestine Large, Rectum	(50)	(50)	(50)	(50)	(50)	
Intestine Small, Duodenum	(50)	(50)	(50)	(50)	(50)	
Perforation	()	()	1	(/	ζ/	
Polyarteritis Nodosa	1 [2.0]					
Ulcer			1 [4.0]			
Intestine Small, Ileum	(50)	(50)	(50)	(50)	(50)	
Intestine Small, Jejunum	(50)	(50)	(50)	(50)	(50)	
Epithelium, Hyperplasia	1 [2.0]	, ,	2 [1.5]	, ,	,	
Peyer's Patch, Hyperplasia, Lymphocyte	4 [3.3]	5 [2.2]	3 [3.3]	2 [2.0]	4 [2.3]	
Peyer's Patch, Inflammation, Chronic Active		- ·	- ·	- -	1 [2.0]	
Liver	(50)	(50)	(50)	(50)	(50)	
Amyloid			1			
Angiectasis	1 [1.0]				1 [2.0]	
Basophilic Focus	6	1			- -	
Clear Cell Focus	3	1	3	1	4	
Eosinophilic Focus	7	5	2	1	4	

a - Number of animals examined microscopically at site and number of animals with lesion b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

Test Type: CHRONIC Route: GAVAGE

Species/Strain: MICE/B6C3F1/N

Experiment Number: 00058 - 04

Black Cohosh **CAS Number:** 84776-26-1

Time Report Requested: 11:32:38
First Dose M/F: NA / 04/09/12

6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Extramedullary Hematopoiesis			1 [2.0]	1 [1.0]	2 [1.0]	
Fatty Change	2 [1.0]		1 [1.0]		1 [1.0]	
Hemorrhage					1 [1.0]	
Hepatodiaphragmatic Nodule					1	
Infiltration Cellular, Lymphocyte			3 [1.7]			
Inflammation, Chronic Active		1 [1.0]	1 [1.0]			
Intrahepatocellular Erythrocytes	2	1	1	2		
Mixed Cell Focus			3		1	
Necrosis	2 [1.0]		1 [1.0]	1 [1.0]	8 [1.1]	
Pigment	1 [1.0]					
Tension Lipidosis	2 [1.5]	2 [1.0]	4 [1.0]	1 [2.0]		
Bile Duct, Cyst	1	-	-	-		
Hepatocyte, Atrophy	1 [2.0]		1 [2.0]			
Mesentery	(6)	(2)	(0)	(2)	(1)	
Inflammation, Granulomatous	1 [2.0]				• •	
Fat, Necrosis	4 [3.0]	1 [2.0]				
Pancreas	(50)	(50)	(50)	(49)	(50)	
Infiltration Cellular, Adipocyte		1 [2.0]				
Acinar Cell, Vacuolation, Cytoplasmic	1 [1.0]					
Acinus, Atrophy	3 [2.3]	1 [2.0]	1 [1.0]	1 [2.0]		
Acinus, Hypertrophy		1 [1.0]				
Duct, Cyst				1		
Salivary Glands	(50)	(50)	(50)	(50)	(50)	
Atrophy	, ,	1 [3.0]	. ,	. ,	• •	
Infiltration Cellular, Lymphocyte		• •	1 [2.0]			
Polyarteritis Nodosa	1 [2.0]				1 [1.0]	
Duct, Hyperplasia	• •				1 [4.0]	
Stomach, Forestomach	(50)	(50)	(50)	(50)	(50)	
Diverticulum	1	. ,	, ,	. ,	• •	
Ulcer		1 [2.0]				
Epithelium, Hyperplasia, Focal		2 [2.5]	1 [1.0]			
Stomach, Glandular	(50)	(50)	(50)	(50)	(50)	
Mineral	1	1	1	1	1	
Polyarteritis Nodosa	1 [2.0]					
Epithelium, Hyperplasia		1 [3.0]				
Tongue	(0)	(0)	(1)	(0)	(0)	

a - Number of animals examined microscopically at site and number of animals with lesion b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

Test Type: CHRONIC Route: GAVAGE

Species/Strain: MICE/B6C3F1/N

Experiment Number: 00058 - 04

Black Cohosh **CAS Number:** 84776-26-1

Time Report Requested: 11:32:38
First Dose M/F: NA / 04/09/12

B6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Tooth	(0)	(3)	(0)	(2)	(0)	
Inflammation, Acute				1 [2.0]		
Necrosis		1 [1.0]		1 [1.0]		
CARDIOVASCULAR SYSTEM						
Aorta	(1)	(0)	(0)	(0)	(1)	
Polyarteritis Nodosa	1 [2.0]					
Blood Vessel	(50)	(48)	(49)	(49)	(49)	
Infiltration Cellular, Lymphocyte		1 [2.0]				
Heart	(50)	(50)	(50)	(50)	(50)	
Cardiomyopathy	8 [1.0]	5 [1.2]	8 [1.0]	11 [1.1]	12 [1.0]	
Mineral	1					
Polyarteritis Nodosa	2 [1.5]					
Valve, Degeneration, Myxomatous	1 [1.0]					
Valve, Inflammation, Acute	1 [4.0]					
Valve, Inflammation, Chronic Active					1 [3.0]	
ENDOCRINE SYSTEM						
Adrenal Cortex	(50)	(50)	(50)	(50)	(50)	
Hyperplasia, Focal	1 [1.0]					
Hypertrophy, Focal	1 [1.0]			1 [1.0]	1 [1.0]	
Infiltration Cellular, Lymphocyte		1 [2.0]				
Vacuolation, Cytoplasmic, Focal				1 [1.0]		
Bilateral, Vacuolation, Cytoplasmic, Focal	2 [2.5]					
Subcapsular, Hyperplasia			1 [1.0]			
Adrenal Medulla	(50)	(50)	(50)	(50)	(50)	
Hyperplasia, Focal	1 [1.0]				1 [1.0]	
Islets, Pancreatic	(50)	(50)	(50)	(50)	(50)	
Atrophy				1 [3.0]		
Hyperplasia	7 [1.6]		3 [1.0]		1 [1.0]	
Parathyroid Gland	(41)	(45)	(42)	(41)	(42)	
Pituitary Gland	(49)	(49)	(50)	(50)	(50)	
Hemorrhage				1 [1.0]		

a - Number of animals examined microscopically at site and number of animals with lesion b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

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Experiment Number: 00058 - 04

Black Cohosh CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

B6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Pars Distalis, Atypia Cellular	1 [1.0]					
Pars Distalis, Hyperplasia, Focal	8 [1.4]	8 [1.9]	10 [1.4]	7 [1.0]	8 [1.6]	
Pars Distalis, Hypertrophy	2 [1.0]	1 [1.0]		3 [1.0]	4 [1.0]	
Pars Intermedia, Hyperplasia, Focal	1 [1.0]					
Thyroid Gland	(50)	(50)	(49)	(49)	(50)	
Cyst, Congenital					1	
Infiltration Cellular, Lymphocyte		1 [1.0]				
Bilateral, Follicle, Dilation					1 [1.0]	
Follicle, Dilation		1 [2.0]	1 [1.0]	1 [1.0]	5 [1.4]	
Follicular Cell, Hyperplasia			1 [2.0]			
GENERAL BODY SYSTEM						
Peritoneum	(0)	(0)	(0)	(1)	(0)	
Tissue NOS	(0)	(0)	(1)	(0)	(0)	
Fat, Necrosis	`,	. ,	1 [4.0]	,	. ,	
GENITAL SYSTEM						
Clitoral Gland	(50)	(49)	(50)	(50)	(49)	
Ovary	(49)	(50)	(50)	(50)	(50)	
Angiectasis			2 [2.5]		1 [1.0]	
Atrophy	35 [3.1]	38 [3.4]	34 [2.6]	39 [3.2]	30 [2.6]	
Cyst, Epithelial, Multiple		2	1			
Cyst, Epithelial	3	4	2	3	6	
Fatty Change			1 [2.0]			
Hemorrhage			1 [3.0]	1 [3.0]		
Infiltration Cellular, Lymphocyte	2 [2.0]		1 [2.0]			
Pigment			1 [1.0]			
Polyarteritis Nodosa					1 [1.0]	
Thrombus	1	1	1		1	
Bilateral, Follicle, Cyst			1		1	
Bursa, Cyst			2		1	
Follicle, Cyst	10	3	4	5	8	
Follicle, Cyst, Multiple		1				

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Species/Strain: MICE/B6C3F1/N

Experiment Number: 00058 - 04

Black Cohosh CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

B6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Germinal Epithelium, Hyperplasia	1 [2.0]		1 [2.0]		1 [1.0]	
Interstitial Cell, Hyperplasia		1 [2.0]	3 [2.0]	2 [1.5]	1 [2.0]	
Paraovarian Tissue, Cyst				1		
Rete Ovarii, Cyst	1			2	1	
Uterus	(50)	(50)	(50)	(50)	(50)	
Adenomyosis		1 [2.0]				
Angiectasis	2 [2.0]	2 [2.0]	1 [1.0]		3 [2.0]	
Dilation		1 [2.0]				
Hemorrhage					1 [2.0]	
Inflammation, Acute	5 [1.8]	1 [2.0]	2 [1.5]	5 [1.4]		
Polyarteritis Nodosa				• •	1 [1.0]	
Thrombus				2		
Endometrium, Hyperplasia, Cystic	41 [2.2]	31 [1.9]	45 [1.9]	41 [2.1]	45 [2.3]	
Endometrium, Metaplasia, Squamous	2 [1.0]			1 [1.0]	1 [1.0]	
Vagina	(50)	(50)	(49)	(49)	(50)	
Congestion	,	,	(/	,	1 [3.0]	
Inflammation, Acute	21 [1.2]	6 [1.0]	8 [1.4]	6 [1.0]	3 [1.3]	
Polyarteritis Nodosa					1 [1.0]	
Epithelium, Hyperplasia, Cystic		1 [2.0]				
Epithelium, Hyperplasia, Squamous	31 [2.1]	38 [1.9]	41 [2.1]	37 [2.0]	33 [2.4]	
HEMATOPOIETIC SYSTEM						
Bone Marrow	(50)	(50)	(50)	(50)	(50)	
Angiectasis					1 [3.0]	
Fibrosis				1 [1.0]		
Hypercellularity	1 [2.0]	2 [1.5]	5 [2.8]	5 [1.8]	3 [3.0]	
Necrosis			1 [4.0]			
Lymph Node	(6)	(1)	(2)	(5)	(7)	
Lumbar, Angiectasis			1 [3.0]			
Lumbar, Hyperplasia, Lymphocyte					1 [4.0]	
Lymph Node, Mandibular	(50)	(49)	(50)	(50)	(50)	
Atrophy	1 [2.0]	1 [3.0]		1 [3.0]		
Hyperplasia, Lymphocyte		4 [3.3]	5 [2.0]	-	7 [2.6]	
Infiltration Cellular, Plasma Cell			1 [2.0]	1 [3.0]	1 [4.0]	

a - Number of animals examined microscopically at site and number of animals with lesion b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

Test Type: CHRONIC Route: GAVAGE

Species/Strain: MICE/B6C3F1/N

b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

Experiment Number: 00058 - 04

Black Cohosh CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

Anglectasis Atrophy At	0 mg/	F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Sinus, Infiltration Cellular, Neutrophil Lymph Node, Mesenteric (50) (49) (50) (49) (50) (49) (50) (49) (50) (49) (50) (49) (50) (49) (50) (49) (50) (49) (49) (40) (4		Pigment		2 [1.0]				
Lymph Node, Mesenteric (50) (49) (50) (49) Angiectasis 1[2.0] 1[2.0] 1[2.0] 1 [2.0] 1		Sinus, Infiltration Cellular, Ne			1 [2.0]			
Anglectasis Atrophy At	(50		(50)	(49)		(49)	(50)	
Atrophy 1 [2.0] 1 [2.0	•		, ,			, ,	, ,	
Hemorrhage	1 [2.	Atrophy	1 [2.0]			1 [2.0]	1 [4.0]	
Hemorrhage	1 [1.	Extramedullary Hematopoies	1 [1.0]	1 [2.0]	1 [2.0]	1 [1.0]		
Hyperplasia, Lymphocyte		Hemorrhage						
Infiltration Cellular, Histiocyte	_			10 [2.3]	11 [2.4]	4 [3.0]	7 [2.4]	
Infiltration Cellular, Plasma Cell 1 [2.0] 1 [2.0] 3 [2.3] Inflammation, Chronic Active	•						1 [4.0]	
Inflammation, Chronic Active Spleen (50) (50) (50) (50) (50) Extramedullary Hematopoiesis, Increased 17 21 22 14 Hyperplasia, Lymphocyte 10 [1.9] 18 [2.5] 10 [2.0] 14 [2.3] 7 Pigment 2 [1.0] 2 [1.0] 11 White Pulp, Atrophy 2 [2.0] 4 [3.8] 3 [3.3] 5 [3.2] 2 Thymus (48) (46) (48) (49) Hyperplasia, Epithelial 1 [4.0] INTEGUMENTARY SYSTEM Mammary Gland (50) (50) (50) (50) (47) Galactocele Duct, Hyperplasia 1 [1.0] Epithelium, Hyperplasia 1 [2.0] 1 [1.0] 1 Skin (50) (50) (50) (50) (50) (50) Erosion 1 [2.0] Infiltration Cellular, Lymphocyte 1 [2.0] Ulcer 2 [3.0] Epidermis, Hyperplasia 1 [2.0] Subcutaneous Tissue, Fibrosis 1 [2.0]	1 [2.	Infiltration Cellular, Plasma C	1 [2.0]		1 [2.0]		• •	
Spleen	•						1 [2.0]	
Extramedullary Hematopoiesis, Increased 17 21 22 14 Hyperplasia, Lymphocyte 10[1.9] 18 [2.5] 10 [2.0] 14 [2.3] 7 Pigment 2 [1.0] 2 [1.0] 11 [2.0] 1 White Pulp, Atrophy 2 [2.0] 4 [3.8] 3 [3.3] 5 [3.2] 2 Thymus (48) (46) (48) (49) 1 Hyperplasia, Epithelial 1 [4.0] INTEGUMENTARY SYSTEM Mammary Gland (50) (50) (50) (50) (47) Galactocele Duct, Hyperplasia 1 [1.0] Epithelium, Hyperplasia 1 [2.0] 1 [1.0] 1 Skin (50) (50) (50) (50) (50) (50) Infiltration Cellular, Lymphocyte 1 [2.0] Ulcer 2 2 [3.0] 2 Epidermis, Hyperplasia 1 [2.0] MUSCULOSKELETAL SYSTEM	(50		(50)	(50)	(50)	(50)	(50)	
Hyperplasia, Lymphocyte							11	
Pigment 2 [1.0] 2 [1.0]			10 [1.9]		10 [2.0]	14 [2.3]	7 [1.7]	
White Pulp, Atrophy 2 [2.0] 4 [3.8] 3 [3.3] 5 [3.2] 2 Thymus (48) (46) (48) (49) Hyperplasia, Epithelial 1 [4.0] (49) (49) INTEGUMENTARY SYSTEM Mammary Gland (50) (50) (50) (50) (47) Galactocele Duct, Hyperplasia 1 [1.0] 1 [1.0] 1 Epithelium, Hyperplasia 1 [2.0] 1 [1.0] 1 Skin (50) (50) (50) (50) (50) Erosion 1 [2.0] 1 [2.0] 1 Ulcer 2 [3.0] 2 Epidermis, Hyperplasia 1 [3.0] 2 Subcutaneous Tissue, Fibrosis 1 [2.0]					• •		1 [1.0]	
Thymus (48) (46) (48) (49) (49) Hyperplasia, Epithelial (50) (50) (50) (50) (47) (47) (48) (49) (49) (49) (49) (49) (49) (49) (49		~			3 [3.3]	5 [3.2]	2 [4.0]	
Hyperplasia, Epithelial							(46)	
Mammary Gland (50) (50) (50) (47) Galactocele Duct, Hyperplasia 1 [1.0] 1 [1.0] 1 [1.0] 1 [2.0] 1 [2.0] 1 [2.0] 1 [2.0] 1 [2.0] 1 [2.0] MUSCULOSKELETAL SYSTEM	`	-	, ,	, ,		,	,	
Galactocele 1 [1.0] Duct, Hyperplasia 1 [2.0] 1 [1.0] Epithelium, Hyperplasia 1 [2.0] 1 [1.0] 1 Skin (50) (50) (50) (50) 1 [2.0] Erosion 1 [2.0] 1 [2.0] 1 [2.0] 2 [3.0] 2 Ulcer 2 [3.0] 2 [3.0] 2 Epidermis, Hyperplasia 1 [3.0] 1 [3.0] Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM		GUMENTARY SYSTEM						
Galactocele 1 [1.0] Duct, Hyperplasia 1 [2.0] 1 [1.0] Epithelium, Hyperplasia 1 [2.0] 1 [1.0] 1 Skin (50) (50) (50) (50) 1 [2.0] Erosion 1 [2.0] 1 [2.0] 1 [2.0] 2 [3.0] 2 Ulcer 2 [3.0] 2 [3.0] 2 Epidermis, Hyperplasia 1 [3.0] 1 [3.0] Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM	(50	mmary Gland	(50)	(50)	(50)	(47)	(50)	
Duct, Hyperplasia 1 [1.0] Epithelium, Hyperplasia 1 [2.0] 1 [1.0] Skin (50) (50) (50) Erosion 1 [2.0] Infiltration Cellular, Lymphocyte 1 [2.0] Ulcer 2 [3.0] 2 Epidermis, Hyperplasia 1 [3.0] Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM	(00		(00)	(00)	(00)	(,	1	
Epithelium, Hyperplasia 1 [2.0] 1 [1.0] 1 Skin (50) (50) (50) (50) (50) Erosion 1 [2.0] Infiltration Cellular, Lymphocyte 1 [2.0] Ulcer 2 2 [3.0] 2 Epidermis, Hyperplasia 1 [3.0] Subcutaneous Tissue, Fibrosis 1 [2.0]					1 [1.0]		·	
Skin (50) (50) (50) (50) Erosion 1 [2.0] Infiltration Cellular, Lymphocyte 1 [2.0] Ulcer 2 [3.0] 2 Epidermis, Hyperplasia 1 [3.0] Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM				1 [2 0]			1 [1.0]	
Erosion Infiltration Cellular, Lymphocyte Ulcer Epidermis, Hyperplasia Subcutaneous Tissue, Fibrosis 1 [2.0] 1 [2.0] 2 [3.0] 2 [3.0] MUSCULOSKELETAL SYSTEM	(50		(50)			(50)	(50)	
Infiltration Cellular, Lymphocyte Ulcer Epidermis, Hyperplasia Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM	(00		(00)	(00)	(00)		(00)	
Ulcer Epidermis, Hyperplasia Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM				1 [2 0]		. [2.0]		
Epidermis, Hyperplasia Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM				. [2.0]	2 [3 0]		2 [3.5]	
Subcutaneous Tissue, Fibrosis 1 [2.0] MUSCULOSKELETAL SYSTEM					_ [0.0]	1 [3 0]	- [0.0]	
MUSCULOSKELETAL SYSTEM	1 [2		1 [2 0]			1 [0.0]		
	۱ اک	Odbodianoodo 1100de, 1 1b10t	1 [2.0]					
		CULOSKELETAL SYSTE						
Bone (50) (50) (50)	(50	ne	(50)	(50)	(50)	(50)	(50)	

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Test Type: CHRONIC Route: GAVAGE

Species/Strain: MICE/B6C3F1/N

Experiment Number: 00058 - 04

Black Cohosh CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

B6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Fibro-Osseous Lesion	8 [1.3]	3 [2.0]	2 [2.5]	2 [1.5]	7 [1.1]	
Increased Bone			1		1	
Joint, Degeneration			1 [3.0]	3 [2.3]		
Skeletal Muscle	(1)	(2)	(0)	(1)	(1)	
NERVOUS SYSTEM						
Brain	(50)	(50)	(50)	(50)	(50)	
Gliosis					1 [2.0]	
Hemorrhage				1 [2.0]		
Hydrocephalus			2		1	
Infiltration Cellular, Lymphocyte		2 [1.5]			1 [2.0]	
Inflammation, Chronic Active	1 [1.0]					
Necrosis			1 [2.0]			
Polyarteritis Nodosa	2 [1.5]			1 [2.0]	3 [1.7]	
Medulla, Necrosis	1 [1.0]					
Peripheral Nerve	(2)	(0)	(0)	(1)	(1)	
Axon, Sciatic, Degeneration					1 [2.0]	
Spinal Cord	(2)	(0)	(0)	(1)	(1)	
Polyarteritis Nodosa				1 [1.0]		
Axon, Degeneration	1 [1.0]			1 [1.0]	1 [1.0]	
RESPIRATORY SYSTEM						
Lung	(50)	(50)	(50)	(50)	(50)	
Hemorrhage	1 [2.0]	1 [3.0]	1 [1.0]			
Infiltration Cellular, Histiocyte	1 [2.0]			1 [2.0]		
Infiltration Cellular, Lymphocyte		3 [2.0]	1 [2.0]			
Inflammation, Chronic Active					1 [1.0]	
Pigment		1 [1.0]				
Polyarteritis Nodosa				1 [1.0]		
Alveolus, Epithelium, Hyperplasia	2 [2.5]	2 [1.0]	2 [1.5]	1 [3.0]	1 [3.0]	
Nose	(50)	(50)	(50)	(50)	(50)	
Olfactory Epithelium, Accumulation, Hyaline Droplet	` '	, ,	1 [2.0]	. ,	, ,	

a - Number of animals examined microscopically at site and number of animals with lesion b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

Test Type: CHRONIC Route: GAVAGE

Species/Strain: MICE/B6C3F1/N

Experiment Number: 00058 - 04

Black Cohosh
CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

B6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Olfactory Epithelium, Atrophy	1 [1.0]		1 [3.0]		3 [1.3]	
Olfactory Epithelium, Metaplasia, Respiratory	4 [1.0]	7 [1.0]	5 [1.0]	7 [1.0]	3 [1.0]	
Respiratory Epithelium, Accumulation, Hyaline Droplet	13 [1.2]	14 [1.1]	15 [1.3]	13 [1.2]	10 [1.3]	
Respiratory Epithelium, Hyperplasia	17 [1.0]	19 [1.0]	9 [1.0]	13 [1.0]	18 [1.1]	
Respiratory Epithelium, Inflammation, Acute	2 [1.0]	1 [1.0]	1 [1.0]		1 [2.0]	
Respiratory Epithelium, Inflammation, Chronic				1 [1.0]		
Respiratory Epithelium, Inflammation, Chronic Active	1 [1.0]		1 [1.0]	1 [1.0]		
Trachea	(50)	(50)	(50)	(50)	(50)	
SPECIAL SENSES SYSTEM						
Ear	(0)	(0)	(0)	(0)	(1)	
Eye	(50)	(50)	(50)	(50)	(50)	
Phthisis Bulbi	1	2	,	,	,	
Anterior Chamber, Inflammation, Acute		1 [2.0]		1 [2.0]		
Ciliary Body, Inflammation, Acute		1 [2.0]		1 [2.0]		
Cornea, Inflammation, Acute		1 [2.0]		1 [2.0]		
Cornea, Inflammation, Chronic Active	1 [1.0]					
Lens, Cataract	3 [1.3]				1 [1.0]	
Harderian Gland	(50)	(50)	(50)	(50)	(50)	
Fibrosis	1 [4.0]	,	,	,	,	
Hyperplasia	1 [4.0]		3 [2.7]	4 [1.8]	3 [2.3]	
URINARY SYSTEM						
Kidney	(50)	(50)	(50)	(50)	(50)	
Glomerulosclerosis	,	,	,	,	1 [1.0]	
Infarct, Chronic	3 [1.0]	2 [1.5]				
Infiltration Cellular, Lymphocyte	1 [1.0]	2 [2.0]	1 [2.0]	1 [2.0]	1 [2.0]	
Nephropathy, Chronic Progressive	13 [1.2]	14 [1.1]	10 [1.3]	20 [1.1]	17 [1.4]	
Polyarteritis Nodosa	1 [1.0]				• •	
Glomerulus, Amyloid				1		
Glomerulus, Cyst		1				

a - Number of animals examined microscopically at site and number of animals with lesion b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

Experiment Number: 00058 - 04

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH Date Report Requested: 12/10/2020 AVERAGE SEVERITY GRADES[b]

Black Cohosh

CAS Number: 84776-26-1

Time Report Requested: 11:32:38 First Dose M/F: NA / 04/09/12

Lab: BAT

Test Type: CHRONIC Route: GAVAGE

Species/Strain: MICE/B6C3F1/N

6C3F1/N MICE FEMALE	0 mg/kg	30 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg	
Renal Tubule, Accumulation, Hyaline Droplet		3 [2.0]	1 [2.0]			
Renal Tubule, Cyst	2			1		
Urinary Bladder	(50)	(50)	(50)	(50)	(49)	
Fibrosis				1 [4.0]		
Infiltration Cellular, Lymphocyte	2 [1.0]		1 [2.0]			
Polyarteritis Nodosa	1					
Ulcer				1 [4.0]		

*** END OF REPORT ***