TDMS No. 99037 - 05 Test Type: CHRONIC Route: GAVAGE Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture **CAS Number:** THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

F2\_R2

C99037B
11/20/2006
ALL
ALL
ALL
ALL
Include ALL
Both
2.2.0

Route: GAVAGE

## Test Type: CHRONIC

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS MALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG
Disposition Summary				
Animals Initially In Study	50	50	50	50
Early Deaths	50	50		
Dosing Accident	2		1	
Moribund Sacrifice	14	21	13	1
Natural Death	7	4	19	49
Survivors				
Natural Death	1			
Terminal Sacrifice	26	25	17	
Animals Examined Microscopically	50	50	50	50
ALIMENTARY SYSTEM				
Esophagus	(50)	(50)	(50)	(50)
Hemorrhage	1 (2%)	· · /		
Hyperkeratosis				1 (2%)
Inflammation, Acute	2 (4%)			
Perforation	1 (2%)			
Intestine Large, Cecum	(48)	(48)	(40)	(39)
Edema	1 (2%)			
Inflammation, Acute	1 (2%)			
Necrosis, Lymphoid				1 (3%)
Intestine Large, Colon	(44)	(47)	(33)	(15)
Edema	2 (5%)	1 (2%)	1 (3%)	
Hemorrhage			1 (3%)	
Necrosis			1 (3%)	
Intestine Large, Rectum	(46)	(48)	(41)	(42)
Inflammation, Chronic				1 (2%)
Intestine Small, Duodenum	(48)	(48)	(37)	(33)
Necrosis, Lymphoid				1 (3%)
Intestine Small, Ileum	(43)	(47)	(30)	(16)
Inflammation, Acute	4 (00()		1 (3%)	1 (6%)
Inflammation, Chronic Active	1 (2%)	4 (00/)		
Necrosis	1 (2%)	1 (2%)	(22)	(2E)
Intestine Small, Jejunum	(43)	(47)	(33)	(25)

## Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture **CAS Number:** THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

1.	(50)	(50)	(50)		
Liver	(50)	(50)	(50)	(50)	
Basophilic Focus	29 (58%)	26 (52%)	25 (50%)	4 (8%)	
Clear Cell Focus	19 (38%)	23 (46%)	11 (22%)	4 (8%)	
Degeneration, Cystic	13 (26%)	7 (14%)	4 (8%)	2 (4%)	
Eosinophilic Focus	6 (12%)	4 (8%)	4 (8%)	1 (2%)	
Fibrosis		. ()		1 (2%)	
Hematopoietic Cell Proliferation		1 (2%)			
Hepatodiaphragmatic Nodule	8 (16%)	3 (6%)	5 (10%)	3 (6%)	
Infiltration Cellular, Mixed Cell	21 (42%)	22 (44%)	18 (36%)	9 (18%)	
Mixed Cell Focus	3 (6%)	6 (12%)	3 (6%)	2 (4%)	
Necrosis, Focal		1 (2%)		3 (6%)	
Thrombosis		1 (2%)			
Bile Duct, Hyperplasia	41 (82%)	45 (90%)	40 (80%)	20 (40%)	
Centrilobular, Necrosis		1 (2%)	1 (2%)	5 (10%)	
Hepatocyte, Vacuolization Cytoplasmic	7 (14%)	10 (20%)	4 (8%)	1 (2%)	
Hepatocyte, Vacuolization Cytoplasmic, Focal		2 (4%)	1 (2%)	1 (2%)	
Kupffer Cell, Pigmentation		4 (8%)			
Mesentery	(11)	(10)	(11)	(4)	
Accessory Spleen		1 (10%)	1 (9%)	2 (50%)	
Hemorrhage			1 (9%)	1 (25%)	
Fat, Necrosis	10 (91%)	9 (90%)	8 (73%)	1 (25%)	
Oral Mucosa	(0)	(1)	(0)	(1)	
Pancreas	(50)	(50)	(50)	(50)	
Atrophy	29 (58%)	35 (70%)	30 (60%)	18 (36%)	
Cyst	21 (42%)	25 (50%)	15 (30%)	6 (12%)	
Hemorrhage	( )		· · · · ·	1 (2%)	
Necrosis		1 (2%)		1 (2%)	
Acinus, Hyperplasia, Focal	8 (16%)	1 (2%)	6 (12%)		
Salivary Glands	(50)	(50)	(50)	(50)	
Atrophy	7 (14%)	4 (8%)	9 (18%)	(~~)	
Hyperplasia	. ( , . , . ,	. (0,0)		1 (2%)	
Inflammation, Acute	1 (2%)		1 (2%)	. (_,)	
Inflammation, Chronic	1 (270)	2 (4%)	· (2,0)		
Necrosis		2 (170)		1 (2%)	
Duct, Hyperplasia	1 (2%)			· (270)	
Stomach, Forestomach	(50)	(50)	(50)	(49)	

## Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS MALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Edema	2 (4%)	4 (8%)	1 (2%)		
Erosion	1 (2%)	1 (070)	1 (270)		
Inflammation, Chronic Active	( ( 2 / 0 )		1 (2%)		
Stromal Hyperplasia			1 (270)	1 (2%)	
Ulcer	1 (2%)	3 (6%)	2 (4%)	. (273)	
Epithelium, Hyperplasia	9 (18%)	12 (24%)	13 (26%)	3 (6%)	
Stomach, Glandular	(50)	(50)	(49)	(48)	
Cyst	(00)	1 (2%)	(10)	(10)	
Edema		2 (4%)	1 (2%)		
Erosion	3 (6%)	6 (12%)	6 (12%)	1 (2%)	
Hemorrhage	0 (070)	0 (1270)	0 (1270)	2 (4%)	
Inflammation		1 (2%)		2 (170)	
Necrosis, Lymphoid		1 (2 /0)		2 (4%)	
Ulcer	2 (4%)	5 (10%)	2 (4%)	2 (470)	
	2 (470)	0 (1070)	2 (470)		
CARDIOVASCULAR SYSTEM					
Heart	(50)	(50)	(50)	(50)	
Cardiomyopathy	44 (88%)	41 (82%)	47 (94%)	45 (90%)	
Hemorrhage				2 (4%)	
Hypertrophy		1 (2%)			
Thrombosis	3 (6%)	4 (8%)	2 (4%)		
Epicardium, Inflammation		1 (2%)			
ENDOCRINE SYSTEM					
Adrenal Cortex	(50)	(50)	(49)	(50)	
Accessory Adrenal Cortical Nodule	21 (42%)	19 (38%)	21 (43%)	4 (8%)	
Degeneration, Fatty	26 (52%)	28 (56%)	31 (63%)	34 (68%)	
Hematopoietic Cell Proliferation	2 (4%)	2 (4%)			
Hyperplasia, Focal	17 (34%)	11 (22%)	9 (18%)	4 (8%)	
Hypertrophy, Focal	6 (12%)	2 (4%)		· · ·	
Necrosis	1 (2%)	3 (6%)	1 (2%)	2 (4%)	
Bilateral, Necrosis	1 (2%)	· · /		· · /	
Capsule, Hyperplasia	1 (2%)		1 (2%)		

## Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

# alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

	(50) 25 (50%) 2 (4%) (50)	(49) 13 (27%)	(49) 1 (2%)	
50) (6%)	2 (4%)	13 (27%)	1 (2%)	
(6%)				
(6%)				
	(~~)	(50)	(50)	
			1 (2%)	
48)	(49)	(47)	(49)	
	1 (2%)	1 (2%)		
50)			(49)	
12%)			2 (4%)	
			2 (4%)	
(32%)	13 (26%)	3 (6%)		
			1 (2%)	
	4 (8%)	1 (2%)		
(8%)			, , , , , , , , , , , , , , , , , , ,	
. ,	1 (2%)			
<b>`</b> ,		1 (2%)	3 (6%)	
46)				
			. ,	
<b>`</b>	2 (4%)	1 (2%)	1 (2%)	
(1)	(1)	(2)	(0)	
50)	(50)	(50)	(50)	
			、 <i>·</i>	
(0)		(0)	(1)	
	(6%) (48) (50) (14%) (4%) (12%) (32%) (32%) (8%) (4%) (4%) (4%) (4%) (4%) (4%) (15%) (4%) (4%) (10) (50) (0)	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	(6%) 3 (6%)   (48) (49) (47)   1 (2%) 1 (2%)   (50) (50) (50)   (14%) 6 (12%) 9 (18%)   (4%) 2 (4%) 1 (2%)   1 (2%) 1 (2%) 1 (2%)   (12%) 3 (6%) 4 (8%)   (32%) 13 (26%) 3 (6%)   4 (8%) 1 (2%)   (32%) 13 (26%) 3 (6%)   4 (8%) 1 (2%)   (4%) 1 (2%)   (4%) 1 (2%)   (4%) 1 (2%)   (46) (48) (42)   (15%) 6 (13%) 4 (10%)   (4%) 2 (4%) 1 (2%)   (4%) 7 (15%) 2 (5%)   2 (4%) 1 (2%) 1 (2%)   (1) (1) (2)   (50) (50) (50)   1 (2%) (50) (50)	$\begin{array}{cccccccccccccccccccccccccccccccccccc$

Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS MALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Inflammation, Acute		1 (100%)			
Preputial Gland	(49)	(49)	(50)	(50)	
Cyst	1 (2%)	(10)	(00)	(00)	
Hyperplasia	2 (4%)		1 (2%)		
Inflammation, Chronic	13 (27%)	19 (39%)	18 (36%)	5 (10%)	
Necrosis		1 (2%)			
Prostate	(50)	(50)	(50)	(50)	
Fibrosis	1 (2%)	()	1 (2%)	()	
Inflammation, Chronic	32 (64%)	20 (40%)	23 (46%)	9 (18%)	
Mineralization		1 (2%)			
Necrosis		1 (2%)	1 (2%)		
Epithelium, Hyperplasia	23 (46%)	17 (34%)	18 (36%)	4 (8%)	
Seminal Vesicle	(50)	(50)	(50)	(50)	
Hyperplasia	(00)	1 (2%)	(00)	(00)	
Infiltration Cellular		. (= /0)		1 (2%)	
Inflammation, Acute		1 (2%)	1 (2%)	1 (2%)	
Inflammation, Chronic		1 (2%)	. (= / )	(_/0)	
Necrosis		1 (2%)			
Testes	(50)	(50)	(50)	(50)	
Germinal Epithelium, Atrophy	5 (10%)	2 (4%)	3 (6%)	4 (8%)	
Interstitial Cell, Hyperplasia	9 (18%)	5 (10%)	6 (12%)	13 (26%)	
HEMATOPOIETIC SYSTEM					
Bone Marrow	(50)	(50)	(50)	(50)	
Hyperplasia	7 (14%)	7 (14%)	12 (24%)	4 (8%)	
Myelofibrosis	5 (10%)	4 (8%)			
Necrosis		1 (2%)			
Thrombosis		1 (2%)			
Lymph Node	(12)	(14)	(12)	(5)	
Mediastinal, Ectasia	1 (8%)				
Mediastinal, Hemorrhage	2 (17%)			1 (20%)	
Mediastinal, Hyperplasia, Lymphoid	1 (8%)	1 (7%)	3 (25%)	1 (20%)	
Mediastinal, Pigmentation			1 (8%)		
Pancreatic, Hemorrhage	1 (8%)			1 (20%)	

## Test Type: CHRONIC

Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS MALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Pancreatic, Hyperplasia, Lymphoid		1 (7%)		3 (60%)	
Pancreatic, Necrosis, Lymphoid				1 (20%)	
Pancreatic, Pigmentation	1 (8%)			1 (20%)	
Lymph Node, Mandibular	(2)	(0)	(0)	(3)	
Congestion				1 (33%)	
Hyperplasia, Lymphoid				1 (33%)	
Lymph Node, Mesenteric	(50)	(49)	(50)	(50)	
Ectasia	1 (2%)	1 (2%)	1 (2%)		
Hemorrhage	4 (8%)	2 (4%)	10 (20%)	10 (20%)	
Hyperplasia, Lymphoid	1 (2%)	2 (4%)	4 (8%)	3 (6%)	
Necrosis		1 (2%)	2 (4%)	1 (2%)	
Spleen	(50)	(50)	(49)	(48)	
Accessory Spleen				1 (2%)	
Angiectasis	1 (2%)				
Fibrosis	2 (4%)	3 (6%)	1 (2%)		
Hematopoietic Cell Proliferation	6 (12%)	7 (14%)	1 (2%)	1 (2%)	
Hemorrhage	1 (2%)				
Infarct	1 (2%)	1 (2%)	1 (2%)		
Necrosis		1 (2%)	2 (4%)	3 (6%)	
Pigmentation	19 (38%)	24 (48%)	30 (61%)	46 (96%)	
Lymphoid Follicle, Atrophy		1 (2%)	2 (4%)	1 (2%)	
Lymphoid Follicle, Hyperplasia	10 (20%)	12 (24%)	8 (16%)	3 (6%)	
Thymus	(44)	(45)	(45)	(47)	
Cyst		1 (2%)			
Ectopic Parathyroid Gland			1 (2%)		
Fibrosis		1 (2%)			
INTEGUMENTARY SYSTEM				·	
Mammary Gland	(48)	(47)	(49)	(42)	
Angiectasis		1 (2%)			
Fibrosis	1 (2%)				
Hyperplasia	2 (4%)	1 (2%)			
Duct, Ectasia	28 (58%)	36 (77%)	34 (69%)	15 (36%)	
Skin	(50)	(50)	(50)	(50)	

Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
	2 (4%)	4 (8%)		
	2 (170)	1 (070)	1 (2%)	
1 (2%)	3 (6%)	5 (10%)		
. (_,,,,		4 (8%)	3 (6%)	
			- ( )	
			2 (4%)	
1 (2%)				
	1 (2%)	3 (6%)		
1 (2%)		( )		
	1 (2%)			
		1 (2%)		
			· · · · · · · · · · · · · · · · · · ·	
(50)	(50)	(50)	(50)	
(00)			(00)	
(2)			(0)	
(-)	2 (50%)	(-)		
(50)	(EO)	(50)	(50)	
9(10%)	4 (0%)	. ,		
	1 (20/)	I (∠%)		
	I (Z%)		3 (0%)	
(50)	(50)	(50)	(50)	
	1 (2%) 1 (2%) 1 (2%) 1 (2%) (50) (2) (50) 9 (18%)	$\begin{array}{c} 2 (4\%) \\ 1 (2\%) & 3 (6\%) \\ 1 (2\%) & 1 (2\%) \\ 1 (2\%) & 1 (2\%) \\ 1 (2\%) & 1 (2\%) \\ (50) & (50) \\ 2 (4\%) \\ (2) & (4) \\ 2 (50\%) \\ \end{array}$ $\begin{array}{c} (50) & (50) \\ 2 (4\%) \\ (4) \\ 2 (50\%) \\ 1 (2\%) \\ \end{array}$	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	$\begin{array}{cccccccccccccccccccccccccccccccccccc$

## Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

# alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

ISCHER 344 RATS MALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Infiltration Cellular, Histiocyte	31 (62%)	43 (86%)	33 (66%)	18 (36%)	
Inflammation, Chronic	1 (2%)	6 (12%)	2 (4%)	4 (8%)	
Metaplasia, Osseous	2 (4%)	5 (10%)	6 (12%)	2 (4%)	
Thrombosis	_ ( ,			1 (2%)	
Alveolar Epithelium, Hyperplasia	11 (22%)	17 (34%)	19 (38%)	3 (6%)	
Nose	(50)	(50)	(50)	(50)	
Congestion				1 (2%)	
Foreign Body	11 (22%)	8 (16%)	9 (18%)	1 (2%)	
Inflammation, Chronic	13 (26%)	15 (30%)	13 (26%)	2 (4%)	
Goblet Cell, Hyperplasia	2 (4%)	1 (2%)	3 (6%)	1 (2%)	
Respiratory Epithelium, Hyperplasia	13 (26%)	7 (14%)	10 (20%)	1 (2%)	
Trachea	(50)	(50)	(50)	(50)	
Inflammation, Acute	1 (2%)				
SPECIAL SENSES SYSTEM					
Eye	(49)	(50)	(46)	(48)	
Cataract	6 (12%)	9 (18%)	10 (22%)	2 (4%)	
Developmental Malformation				1 (2%)	
Edema	1 (2%)	1 (2%)			
Hemorrhage		1 (2%)			
Inflammation, Chronic		2 (4%)			
Bilateral, Cataract	1 (2%)	2 (4%)	2 (4%)		
Retina, Degeneration	1 (2%)	2 (4%)			
Harderian Gland	(50)	(50)	(50)	(50)	
Hyperplasia, Focal	1 (2%)	1 (2%)		1 (2%)	
Inflammation, Chronic	3 (6%)	4 (8%)	4 (8%)	5 (10%)	
Pigmentation		1 (2%)			
Zymbal's Gland	(0)	(1)	(1)	(0)	
JRINARY SYSTEM					
Kidney	(48)	(48)	(44)	(49)	
	(48) 1 (2%)	(48)	(44)	(49)	

Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

ISCHER 344 RATS MALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG
Cyst	1 (2%)	3 (6%)		
Hydronephrosis	1 (2%)	1 (2%)		
Infarct	3 (6%)	1 (2%)		
Inflammation, Suppurative		2 (4%)		1 (2%)
Mineralization	17 (35%)	33 (69%)	41 (93%)	38 (78%)
Nephropathy	45 (94%)	44 (92%)	41 (93%)	46 (94%)
Artery, Thrombosis	1 (2%)			
Papilla, Necrosis	1 (2%)			
Renal Tubule, Accumulation, Hyaline Droplet	1 (2%)			
Renal Tubule, Necrosis				2 (4%)
Transitional Epithelium, Hyperplasia		1 (2%)	4 (9%)	2 (4%)
Urinary Bladder	(50)	(50)	(49)	(50)
Atrophy		1 (2%)		
Hemorrhage				1 (2%)
Inflammation, Acute		1 (2%)		
Inflammation, Chronic		1 (2%)		1 (2%)
Necrosis		1 (2%)		
Transitional Epithelium, Hyperplasia		1 (2%)		1 (2%)

\*\*\* END OF MALE \*\*\*

Route: GAVAGE

Test Type: CHRONIC

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS FEMALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Disposition Summary					
Animals Initially In Study	50	50	50	50	
Early Deaths					
Moribund Sacrifice	6	9	10	5	
Natural Death	9	8	21	45	
Survivors					
Terminal Sacrifice	35	33	19		
Animals Examined Microscopically	50	50	50	50	
ALIMENTARY SYSTEM					
Esophagus	(50)	(50)	(50)	(50)	
Diverticulum	1 (2%)	, , , , , , , , , , , , , , , , , , ,		· · /	
Intestine Large, Cecum	(41)	(44)	(39)	(47)	
Hemorrhage		1 (2%)			
Intestine Large, Colon	(41)	(43)	(33)	(25)	
Intestine Large, Rectum	(46)	(45)	(41)	(47)	
Hemorrhage				1 (2%)	
Intestine Small, Duodenum	(44)	(45)	(40)	(47)	
Necrosis			1 (3%)		
Necrosis, Lymphoid				1 (2%)	
Intestine Small, Jejunum	(41)	(42)	(33)	(31)	
Necrosis, Lymphoid				1 (3%)	
Liver	(50)	(50)	(49)	(50)	
Angiectasis	1 (2%)		1 (2%)		
Basophilic Focus	46 (92%)	46 (92%)	40 (82%)	12 (24%)	
Clear Cell Focus	17 (34%)	21 (42%)	11 (22%)		
Eosinophilic Focus	8 (16%)	2 (4%)	2 (4%)		
Hematopoietic Cell Proliferation	1 (2%)	1 (2%)	4 (8%)		
Hepatodiaphragmatic Nodule	5 (10%)	8 (16%)	6 (12%)	4 (8%)	
Infiltration Cellular, Histiocyte				1 (2%)	
Infiltration Cellular, Mixed Cell	30 (60%)	30 (60%)	32 (65%)	19 (38%)	
Inflammation, Chronic			1 (2%)		
Mixed Cell Focus	10 (20%)	4 (8%)	4 (8%)		
Necrosis, Focal	1 (2%)	1 (2%)	5 (10%)	4 (8%)	

## Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS FEMALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Artery, Inflammation		1 (2%)	1 (2%)		
Bile Duct, Hyperplasia	4 (8%)	2 (4%)	7 (14%)		
Capsule, Fibrosis	1 (2%)	= ( : / 3)	. ( , .)		
Centrilobular, Necrosis			3 (6%)		
Hepatocyte, Vacuolization Cytoplasmic	5 (10%)	4 (8%)	3 (6%)		
Kupffer Cell, Pigmentation		1 (2%)	1 (2%)		
Mesentery	(13)	(11)	(9)	(2)	
Accessory Spleen	( - /	1 (9%)	2 (22%)	1 (50%)	
Necrosis		()	()	1 (50%)	
Fat, Necrosis	13 (100%)	10 (91%)	8 (89%)	1 (50%)	
Oral Mucosa	(0)	(2)	(0)	(0)	
Inflammation, Chronic Active		1 (50%)			
Pancreas	(49)	(49)	(50)	(50)	
Atrophy	25 (51%)	24 (49%)	23 (46%)	5 (10%)	
Cyst	14 (29%)	11 (22%)	9 (18%)	1 (2%)	
Acinus, Hyperplasia, Focal	2 (4%)	1 (2%)	2 (4%)	· · · · · · · · · · · · · · · · · · ·	
Salivary Glands	(50)	(50)	(50)	(50)	
Atrophy	9 (18%)	10 (20%)	5 (10%)	2 (4%)	
Hyperplasia			. ,	1 (2%)	
Inflammation, Chronic		1 (2%)			
Mineralization		2 (4%)		1 (2%)	
Necrosis				1 (2%)	
Vacuolization Cytoplasmic		1 (2%)		1 (2%)	
Stomach, Forestomach	(50)	(50)	(50)	(50)	
Edema	2 (4%)	1 (2%)	. ,		
Inflammation, Chronic Active			1 (2%)		
Ulcer	1 (2%)	2 (4%)			
Epithelium, Hyperplasia	7 (14%)	8 (16%)	2 (4%)		
Epithelium, Hyperplasia, Atypical			1 (2%)		
Stomach, Glandular	(48)	(47)	(48)	(50)	
Cyst		1 (2%)			
Edema	1 (2%)				
Erosion	1 (2%)	1 (2%)	1 (2%)	3 (6%)	
Ulcer	1 (2%)		3 (6%)	1 (2%)	
Tongue	(1)	(3)	(0)	(0)	
Hyperplasia	1 (100%)	2 (67%)			

TDMS No. 99037 - 05 Test Type: CHRONIC Route: GAVAGE Species/Strain: RATS/F 344/N FISCHER 344 RATS FEMALE	P03: INCIDENCE F	Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI			
	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Ulcer		1 (33%)			
Tooth Degeneration	(0)	(1) 1 (100%)	(0)	(0)	
CARDIOVASCULAR SYSTEM		·			
Heart	(50)	(50)	(50)	(50)	
Cardiomyopathy	40 (80%)	30 (60%)	30 (60%)	8 (16%)	
Hemorrhage				1 (2%)	
Artery, Necrosis				1 (2%)	
ENDOCRINE SYSTEM			· · · · · · · · · · · · · · · · · · ·		
Adrenal Cortex	(47)	(49)	(46)	(50)	
Accessory Adrenal Cortical Nodule	7 (15%)	12 (24%)	13 (28%)	9 (18%)	
Degeneration, Fatty	23 (49%)	18 (37%)	19 (41%)	9 (18%)	
Hematopoietic Cell Proliferation	1 (2%)	2 (4%)	4 (9%)		
Hemorrhage			1 (2%)	3 (6%)	
Hyperplasia, Focal	22 (47%)	14 (29%)	9 (20%)	1 (2%)	
Hypertrophy, Focal	7 (15%)	6 (12%)	2 (4%)	1 (2%)	
Inflammation, Acute				1 (2%)	
Necrosis	2 (4%)	1 (2%)	6 (13%)	4 (8%)	
Adrenal Medulla	(48)	(50)	(48)	(50)	
Angiectasis			. ,	1 (2%)	
Hyperplasia	7 (15%)	5 (10%)	3 (6%)	. ,	
Infiltration Cellular, Lymphoid	· · ·	1 (2%)	· · /		
Islets, Pancreatic	(49)	(49)	(50)	(50)	
Hyperplasia	1 (2%)	·	1 (2%)		
Pituitary Gland	(50)	(49)	(49)	(48)	
Pigmentation	5 (10%)	5 (10%)	9 (18%)	. *	
Pars Distalis, Angiectasis	9 (18%)	4 (8%)	5 (10%)	2 (4%)	
Pars Distalis, Atrophy	· · ·	. ,	2 (4%)	12 (25%)	
Pars Distalis, Cyst	18 (36%)	20 (41%)	17 (35%)	8 (17%)	
Pars Distalis, Dilatation	· · · ·	. ,	1 (2%)	. ,	
Pars Distalis, Hyperplasia, Focal	5 (10%)	13 (27%)	6 (12%)	1 (2%)	

TDMS No. 99037 - 05 Test Type: CHRONIC Route: GAVAGE Species/Strain: RATS/F 344/N	P03: INCIDENCE F	Date Report Requested: 08/18/201 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI			
FISCHER 344 RATS FEMALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Pars Intermedia, Angiectasis	2 (4%)				
Pars Intermedia, Cyst	1 (2%)	2 (4%)	2 (4%)	2 (4%)	
Pars Intermedia, Hyperplasia	1 (2%)	, , ,	· · /	, , , , , , , , , , , , , , , , , , ,	
Rathke's Cleft, Dilatation	7 (14%)	1 (2%)	13 (27%)	26 (54%)	
Thyroid Gland	(42)	(46)	(44)	(50)	
Degeneration, Cystic	1 (2%)	2 (4%)	5 (11%)	( )	
Ectopic Thymus		· /	· /	1 (2%)	
Ultimobranchial Cyst	2 (5%)	1 (2%)	3 (7%)	2 (4%)	
C-cell, Hyperplasia	1 (2%)	11 (24%)	2 (5%)		
Follicular Cell, Hyperplasia, Focal	()	1 (2%)	1 (2%)		
GENERAL BODY SYSTEM Tissue NOS Fat, Mediastinum, Necrosis	(1) 1 (100%)	(1)	(0)	(0)	
GENITAL SYSTEM					
Clitoral Gland	(50)	(50)	(50)	(50)	
Cyst	2 (4%)	(00)	1 (2%)	(00)	
Hyperplasia	9 (18%)	7 (14%)	11 (22%)		
Infiltration Cellular, Lymphocyte	0 (10,0)	1 (11/0)	11 (22,0)	1 (2%)	
Inflammation, Chronic			1 (2%)	1 (2%)	
Ovary	(50)	(50)	(50)	(50)	
Cyst	3 (6%)	6 (12%)	2 (4%)	1 (2%)	
Bilateral, Cyst	0 (070)	0 (12/0)	1 (2%)	· (2/0)	
Uterus	(50)	(50)	(49)	(50)	
Cyst	1 (2%)	(00)	()	(00)	
Hyperplasia, Cystic	22 (44%)	17 (34%)	19 (39%)	9 (18%)	
Cervix, Cyst	22 (7770)	17 (07-70)	1 (2%)	3 (1070)	
Fat, Necrosis		1 (2%)	1 (2/0)		
Vagina	(2)		(1)	(0)	
vayına	(2)	(2)	(1)	(0)	

## Test Type: CHRONIC

## Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS FEMALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
HEMATOPOIETIC SYSTEM					
Bone Marrow	(50)	(50)	(50)	(50)	
Hyperplasia	4 (8%)	6 (12%)	6 (12%)		
Infiltration Cellular, Histiocyte	1 (2%)	1 (2%)	2 (4%)	4 (8%)	
Myelofibrosis		1 (2%)			
Necrosis		, , ,		1 (2%)	
Lymph Node	(7)	(8)	(6)	(7)	
Deep Cervical, Hyperplasia		1 (13%)			
Deep Cervical, Pigmentation		1 (13%)			
Iliac, Hemorrhage				1 (14%)	
Iliac, Hyperplasia, Lymphoid				1 (14%)	
Iliac, Pigmentation				1 (14%)	
Mediastinal, Hemorrhage		2 (25%)	2 (33%)	2 (29%)	
Mediastinal, Hyperplasia, Lymphoid		6 (75%)	3 (50%)	2 (29%)	
Mediastinal, Pigmentation		5 (63%)	1 (17%)	1 (14%)	
Pancreatic, Hemorrhage		1 (13%)	2 (33%)	1 (14%)	
Pancreatic, Hyperplasia, Lymphoid			1 (17%)	4 (57%)	
Pancreatic, Pigmentation		1 (13%)	1 (17%)	1 (14%)	
Lymph Node, Mandibular	(1)	(4)	(1)	(4)	
Hyperplasia, Lymphoid	1 (100%)	4 (100%)		2 (50%)	
Pigmentation		1 (25%)		× ,	
Lymph Node, Mesenteric	(50)	(50)	(47)	(50)	
Atrophy	1 (2%)	1 (2%)			
Ectasia	1 (2%)				
Hemorrhage	8 (16%)	9 (18%)	6 (13%)	3 (6%)	
Hyperplasia, Lymphoid	4 (8%)	5 (10%)	4 (9%)	5 (10%)	
Necrosis		( ),	1 (2%)	1 (2%)	
Pigmentation			1 (2%)	, , , , , , , , , , , , , , , , , , ,	
Spleen	(48)	(49)	(48)	(50)	
Ectopic Spleen, Multiple		1 (2%)	· · /	· · /	
Fibrosis	1 (2%)	· · /			
Hematopoietic Cell Proliferation	21 (44%)	31 (63%)	21 (44%)		
Hemorrhage	1 (2%)		. ,		
Inflammation		1 (2%)			
Necrosis		1 (2%)	3 (6%)	4 (8%)	

<b>FDMS No.</b> 99037 - 05	P03: INCIDENCE R	Date Report Requested: 08/18/2010			
Test Type: CHRONIC		Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI			
Route: GAVAGE					
Species/Strain: RATS/F 344/N FISCHER 344 RATS FEMALE					
	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Pigmentation	39 (81%)	40 (82%)	39 (81%)	45 (90%)	
Lymphoid Follicle, Atrophy	2 (4%)	3 (6%)	2 (4%)	2 (4%)	
Lymphoid Follicle, Hyperplasia	3 (6%)	3 (6%)	2 (4%)	8 (16%)	
Lymphoid Follicle, Hyperplasia, Focal	,		1 (2%)		
Thymus	(48)	(45)	(50)	(50)	
Cyst	(10)	2 (4%)	(00)	(00)	
Ectopic Parathyroid Gland	1 (2%)	2 (170)			
Necrosis	. (_70)		1 (2%)	3 (6%)	
INTEGUMENTARY SYSTEM					
Mammary Gland	(50)	(50)	(50)	(50)	
Fibrosis	1 (2%)				
Hyperplasia	19 (38%)	21 (42%)	10 (20%)		
Hyperplasia, Focal	3 (6%)	1 (2%)			
Malformation		. ,	1 (2%)		
Duct, Ectasia	47 (94%)	45 (90%)	39 (78%)		
Skin	(50)	(50)	(50)	(50)	
Cyst Epithelial Inclusion	2 (4%)	()	1 (2%)	()	
Hyperkeratosis	1 (2%)	1 (2%)	1 (2%)		
Inflammation, Chronic	1 (2%)	( ( ) )	( ( ) )		
Ulcer	1 (270)	1 (2%)			
Epidermis, Hyperplasia	2 (4%)	1 (2%)			
Subcutaneous Tissue, Fibrosis	1 (2%)	1 (270)			
Subcularieous rissue, Fibrosis	1 (278)				
MUSCULOSKELETAL SYSTEM					
Bone	(50)	(50)	(50)	(50)	
Femur, Osteopetrosis	1 (2%)	1 (2%)			
Skeletal Muscle	(0)	(0)	(1)	(0)	
NERVOUS SYSTEM				<u>.</u>	
Brain	(50)	(50)	(50)	(50)	

TDMS No. 99037 - 05 Test Type: CHRONIC Route: GAVAGE Species/Strain: RATS/F 344/N FISCHER 344 RATS FEMALE	P03: INCIDENCE F	Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI			
	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Compression	9 (18%)	10 (20%)	6 (12%)		
Gliosis			1 (2%)		
Mineralization	1 (2%)		1 (2%)		
Necrosis			3 (6%)		
Pigmentation	1 (2%)	3 (6%)	5 (10%)	19 (38%)	
RESPIRATORY SYSTEM					
Lung	(50)	(50)	(50)	(50)	
Hemorrhage	· · · ·			1 (2%)	
Hyperplasia		1 (2%)			
Infiltration Cellular, Histiocyte	46 (92%)	43 (86%)	36 (72%)	8 (16%)	
Inflammation, Suppurative		1 (2%)		- ( /	
Inflammation, Chronic	10 (20%)	3 (6%)	5 (10%)	1 (2%)	
Metaplasia, Osseous		1 (2%)	2 (4%)	. (_,,,,	
Necrosis		(=,,,)	- ( ,	1 (2%)	
Alveolar Epithelium, Hyperplasia	23 (46%)	25 (50%)	17 (34%)	4 (8%)	
Nose	(50)	(50)	(50)	(50)	
Congestion	(00)	(00)	(00)	2 (4%)	
Foreign Body	7 (14%)	5 (10%)	3 (6%)	1 (2%)	
Inflammation, Chronic	11 (22%)	11 (22%)	9 (18%)	1 (2%)	
Thrombosis	1 (2%)	11 (2270)	0 (10,0)	1 (270)	
Goblet Cell, Hyperplasia	1 (2%)				
Nasolacrimal Duct, Cyst	1 (270)			1 (2%)	
Respiratory Epithelium, Hyperplasia	5 (10%)	4 (8%)	2 (4%)	1 (270)	
SPECIAL SENSES SYSTEM					
Eye	(44)	(47)	(41)	(49)	
Cataract	( /	5 (11%)	2 (5%)	( - <i>)</i>	
Edema		1 (2%)	2 (5%)		
Hemorrhage	1 (2%)	1 (2%)	(***)		
Inflammation, Acute	. (= / )	1 (2%)			
Inflammation, Chronic	1 (2%)	1 (2%)			
Synechia	( ( ) )	1 (2%)			

Test Type: CHRONIC

#### Route: GAVAGE

Species/Strain: RATS/F 344/N

## P03: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE(a)

alpha/beta Thujone mixture

CAS Number: THUJONEMIXAB

Date Report Requested: 08/18/2010 Time Report Requested: 08:14:26 First Dose M/F: 06/09/03 / 06/09/03 Lab: SRI

FISCHER 344 RATS FEMALE	0 MG/KG	12.5 MG/KG	25 MG/KG	50 MG/KG	
Bilateral, Cataract			1 (2%)		
Bilateral, Edema		1 (2%)	· · · ·		
Bilateral, Retinal Detachment		1 (2%)			
Retina, Degeneration		1 (2%)	2 (5%)	1 (2%)	
Harderian Gland	(50)	(50)	(49)	(50)	
Hemorrhage	. ,	. ,	4 (8%)		
Hyperplasia			1 (2%)		
Inflammation, Chronic	9 (18%)	11 (22%)	9 (18%)	6 (12%)	
URINARY SYSTEM					
Kidney	(44)	(45)	(40)	(50)	
Casts Protein	14 (32%)	8 (18%)	12 (30%)		
Cyst	1 (2%)	1 (2%)			
Infarct	2 (5%)	1 (2%)	1 (3%)		
Inflammation, Suppurative	1 (2%)				
Inflammation, Chronic		1 (2%)	3 (8%)		
Mineralization	16 (36%)	24 (53%)	17 (43%)	8 (16%)	
Nephropathy	33 (75%)	36 (80%)	26 (65%)	4 (8%)	
Renal Tubule, Accumulation, Hyaline Droplet	2 (5%)	5 (11%)	4 (10%)		
Renal Tubule, Necrosis			1 (3%)		
Transitional Epithelium, Hyperplasia	2 (5%)	1 (2%)			
Urinary Bladder	(50)	(50)	(50)	(50)	
Inflammation, Chronic				1 (2%)	
Transitional Epithelium, Hyperplasia		1 (2%)			

## \*\*\* END OF REPORT \*\*\*