

TDMS No. 20007 - 06
Test Type: CHRONIC
Route: GAVAGE
Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
Time Report Requested: 10:57:30
First Dose M/F: 08/24/04 / 08/23/04
Lab: BAT

F_M3

C Number: C20007
Lock Date: 05/10/2007
Cage Range: ALL
Date Range: ALL
Reasons For Removal: ALL
Removal Date Range: ALL
Treatment Groups: Include ALL
Study Gender: Both
TDMSE Version: 2.1.0

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	DAY ON TEST																									
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
	7	7	7	5	5	6	6	7	6	7	7	5	7	7	4	6	6	7	7	7	7	3	6	7	7	
	3	3	3	7	7	6	5	3	6	3	3	8	3	3	5	4	4	3	0	3	3	6	4	3	3	
	1	2	2	9	0	8	2	2	8	1	1	7	1	0	5	4	7	2	3	1	0	9	8	2	1	
B6C3F1 MICE MALE	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
0 G/KG	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
	0	0	0	0	0	0	0	0	0	1	1	1	1	1	1	1	1	1	1	2	2	2	2	2	2	males (cont...)
	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	

ALIMENTARY SYSTEM

Esophagus	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Gallbladder	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Infiltration Cellular, Mononuclear Cell																								
Intestine Large, Cecum	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Amyloid Deposition																								
Serosa, Inflammation, Granulomatous					2																			
Intestine Large, Colon	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Intestine Large, Rectum	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Intestine Small, Duodenum	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Intestine Small, Ileum	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Inflammation, Chronic										2														
Epithelium, Hyperplasia										2														
Intestine Small, Jejunum	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Liver	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Angiectasis																								
Basophilic Focus					X	X																		
Clear Cell Focus			X		X		X		X								X	X						X

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	DAY ON TEST																									males (cont...)			
	0 7 3 1	0 7 3 2	0 7 3 2	0 5 7 9	0 5 7 0	0 6 6 8	0 6 5 2	0 7 3 2	0 6 5 2	0 7 3 8	0 6 3 1	0 7 3 1	0 5 8 7	0 7 3 1	0 4 5 0	0 6 4 4	0 6 4 7	0 7 3 2	0 7 0 3	0 7 3 1	0 7 3 0	0 3 6 9	0 6 4 8	0 7 3 2	0 7 3 1				
B6C3F1 MICE MALE 0 G/KG	ANIMAL ID																												
	0 0 0 0 1	0 0 0 0 2	0 0 0 0 3	0 0 0 0 4	0 0 0 0 5	0 0 0 0 6	0 0 0 0 7	0 0 0 0 8	0 0 0 0 9	0 0 0 0 0	0 0 0 0 1	0 0 0 0 1	0 0 0 0 2	0 0 0 0 3	0 0 0 0 4	0 0 0 0 5	0 0 0 0 6	0 0 0 0 7	0 0 0 0 8	0 0 0 0 9	0 0 0 0 0	0 0 0 0 1	0 0 0 0 2	0 0 0 0 3	0 0 0 0 4	0 0 0 0 5			
Degeneration, Cystic Eosinophilic Focus																													1
Fatty Change	2		1	2	2		2	1				1			1		2	1		1	1	1	1	1	2	1	1	2	1
Fibrosis																													
Hematopoietic Cell Proliferation																													
Infiltration Cellular, Mononuclear Cell	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	2	1	1	1	1	
Mineralization													1																
Mixed Cell Focus	X	X	X							X	X								X	X	X								
Necrosis						2					2																		
Tension Lipidosis																								X				X	
Mesentery																													
Inflammation, Chronic																													
Fat, Necrosis	2																												
Pancreas																													
Acinus, Atrophy																													
Acinus, Hypertrophy																													
Salivary Glands																													
Infiltration Cellular, Mononuclear Cell	1	1	2		2	1	1	1	1	1		1	1	1	1	1	1		1	1	2			1	1	1			
Parotid Gland, Atrophy																													
Stomach, Forestomach																													
Inflammation, Chronic																													
Necrosis																													
Epithelium, Hyperplasia																													
Stomach, Glandular																													

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 1) Minimal 3) Moderate
 2) Mild 4) Marked

DAY ON TEST		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
DAY ON TEST		7	7	7	5	5	6	6	7	6	7	7	5	7	7	4	6	6	7	7	7	7	3	6	7	7
DAY ON TEST		3	3	3	7	7	6	5	3	6	3	3	8	3	3	5	4	4	3	0	3	3	6	4	3	3
DAY ON TEST		1	2	2	9	0	8	2	2	8	1	1	7	1	0	5	4	7	2	3	1	0	9	8	2	1

ANIMAL ID		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANIMAL ID		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANIMAL ID		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANIMAL ID		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANIMAL ID		0	0	0	0	0	0	0	0	0	1	1	1	1	1	1	1	1	1	1	2	2	2	2	2	2
ANIMAL ID		1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5

males (cont...)

Infiltration Cellular, Mononuclear Cell																										
Ulcer												2														
Epithelium, Hyperplasia												2														
Glands, Ectasia														2												
Tooth																										
Dysplasia																										

CARDIOVASCULAR SYSTEM

Blood Vessel																										
Heart																										
Cardiomyopathy																										
Artery, Infiltration Cellular, Mononuclear Cell																										
Mycocardium, Mineralization																										

ENDOCRINE SYSTEM

Adrenal Cortex																										
Hypertrophy																										
Infiltration Cellular, Mononuclear Cell																										
Vacuolization Cytoplasmic																										
Subcapsular, Hyperplasia																										
Zona Reticularis, Hyperplasia																										

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DAY ON TEST	0 0																								
	7	7	7	5	5	6	6	7	6	7	7	5	7	7	4	6	6	7	7	7	7	3	6	7	7
ANIMAL ID	3 3 3 7 7 6 5 3 6 3 3 8 3 3 5 4 4 3 0 3 3 6 4 3 3																								
	1	2	2	9	0	8	2	2	8	1	1	7	1	0	5	4	7	2	3	1	0	9	8	2	1
B6C3F1 MICE MALE 0 G/KG	0 0																								
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANIMAL ID	0 0																								
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0 G/KG	0 0																								
	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5

males (cont...)

Adrenal Medulla Hyperplasia	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Islets, Pancreatic Hyperplasia	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Parathyroid Gland Cyst Infiltration Cellular, Mononuclear Cell	M	+	M	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Pituitary Gland Cyst Pars Distalis, Hyperplasia	+	+	+	+	+	+	+	+	+	+	+	M	+	+	+	+	+	+	+	+	+	+	+	+
Thyroid Gland Atrophy Cyst Follicle, Hyperplasia	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

Epididymis Granuloma Sperm Infiltration Cellular, Mononuclear Cell	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
	1		1			1	1	1		1		1			1	1	1	1	1			1		

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DAY ON TEST	0 0																								
	7 7 7 5 5 6 6 7 6 7 7 5 7 7 4 6 6 7 7 7 7 3 6 7 7																								
ANIMAL ID	3 3 3 7 7 6 5 3 6 3 3 8 3 3 5 4 4 3 0 3 3 6 4 3 3																								
	1 2 2 9 0 8 2 2 8 1 1 7 1 0 5 4 7 2 3 1 0 9 8 2 1																								
B6C3F1 MICE MALE	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0 G/KG	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1	2	3	4	5	6	7	8	9	0	1	1	1	1	1	1	1	1	1	2	2	2	2	2	
	1	2	3	4	5	6	7	8	9	0	1	1	1	1	1	1	1	1	1	2	2	2	2	2	

males (cont...)

Preputial Gland	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Cyst																								
Inflammation								1		1				1	2			1			2			
Prostate	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Hyperplasia																								
Infiltration Cellular, Mononuclear Cell			1					1		1		1		1	1			1	1	1		1		1
Inflammation				1	2	3																		
Seminal Vesicle	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Fibrosis	2		1																					
Infiltration Cellular, Mononuclear Cell				1																				
Inflammation	1		1			1																		
Testes	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Inflammation																								
Mineralization																								
Germinal Epithelium, Atrophy																								

HEMATOPOIETIC SYSTEM

Bone Marrow	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Lymph Node, Mandibular	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Lymph Node, Mesenteric	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Atrophy						3																		

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DAY ON TEST	0 0																								
	7 7 7 5 5 6 6 7 6 7 7 5 7 7 4 6 6 7 7 7 7 3 6 7 7																								
ANIMAL ID	3 3 3 7 7 6 5 3 6 3 3 8 3 3 5 4 4 3 0 3 3 6 4 3 3																								
	1 2 2 9 0 8 2 2 8 1 1 7 1 0 5 4 7 2 3 1 0 9 8 2 1																								
B6C3F1 MICE MALE 0 G/KG	0 0																								
	0 0																								
	0 0																								
	0 0																								
	1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 males (cont...)																								

Spleen	+ +																								
Hematopoietic Cell Proliferation	2 3 2 3 1 3																								
Hyperplasia, Lymphoid																									
Lymphoid Follicle, Atrophy	3 2 4 2																								

Thymus	M + M + +																								
Atrophy	3 1 3 4 4 2 1 4 2 4 4 3 4 1 1 2 2																								
Hyperplasia, Histiocytic																									

INTEGUMENTARY SYSTEM

Mammary Gland	M M M M M + M M M M M M M M M M M M M M M M M M M																								
Skin	+ +																								
Inflammation, Chronic																									
Ulcer																									

MUSCULOSKELETAL SYSTEM

Bone	+ +																								
Cranium, Hyperostosis																									
Skeletal Muscle	+																								

NERVOUS SYSTEM

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	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	7	7	7	5	5	6	6	7	6	7	7	5	7	7	4	6	6	7	7	7	7	3	6	7	7
	3	3	3	7	7	6	5	3	6	3	3	8	3	3	5	4	4	3	0	3	3	6	4	3	3
	1	2	2	9	0	8	2	2	8	1	1	7	1	0	5	4	7	2	3	1	0	9	8	2	1
B6C3F1 MICE MALE 0 G/KG	ANIMAL ID																								
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	1	1	1	1	1	1	1	1	1	1	2	2	2	2	2	2
	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5

males
(cont...)

Brain +

RESPIRATORY SYSTEM

Lung	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Alveolar Epithelium, Hyperplasia																									1
Alveolus, Infiltration Cellular, Histiocyte			1			2																			
Bronchus, Hyperplasia							2																		
Nose	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Inflammation, Suppurative																									2
Inflammation, Chronic			1										1		1					1				1	
Nasolacrimal Duct, Inflammation, Suppurative			2																						
Respiratory Epithelium, Hyperplasia												2													
Trachea	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+

SPECIAL SENSES SYSTEM

Eye	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Harderian Gland	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Fibrosis																									
Hyperplasia			2									1													
Infiltration Cellular, Mononuclear Cell	1	1		1	1	1	1		1	1	1	1	1	1	1	1	1	1	1				1	1	

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	7 7 7 5 5 6 6 7 6 7 7 5 7 7 4 6 6 7 7 7 7 3 6 7 7																								
ANIMAL ID	3 3 3 7 7 6 5 3 6 3 3 8 3 3 5 4 4 3 0 3 3 6 4 3 3																								
	1 2 2 9 0 8 2 2 8 1 1 7 1 0 5 4 7 2 3 1 0 9 8 2 1																								
B6C3F1 MICE MALE 0 G/KG	0 0																								
	0 0																								
	0 0																								
	0 0																								
	1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5																								

males
(cont...)

URINARY SYSTEM

Kidney	+ +																								
Hydronephrosis																									
Inflammation	2 2 3																								
Mineralization	3																								
Nephropathy	2 1 1 1 1 1 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1																								
Cortex, Medulla, Necrosis	3 3 3																								
Papilla, Necrosis	3																								
Renal Tubule, Hyperplasia	1																								
Renal Tubule, Mineralization	1 1 2 2 1																								
Renal Tubule, Pigmentation, Lipofuscin																									
Urethra	+																								
Inflammation	3																								
Urinary Bladder	+ +																								
Infiltration Cellular, Mononuclear Cell	1 1																								
Inflammation	1 1 3																								

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DAY ON TEST	0 0																								* TOTALS
	7 7 7 7 7 7 6 7 7 7 7 7 7 7 7 5 7 7 7 6 7 7 7 7 6																								
B6C3F1 MICE MALE 0 G/KG	3 3 3 3 3 3 1 3 3 3 3 3 3 3 3 3 3 3 3 2 3 3 3 3 8																								
	2 2 0 1 0 2 7 2 2 1 1 1 2 1 0 3 2 2 2 6 2 1 0 0 3																								
ANIMAL ID	0 0																								
	0 0																								
	2 2 2 2 3 3 3 3 3 3 3 3 3 3 3 4 4 4 4 4 4 4 4 4 5																								
	6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0																								

ALIMENTARY SYSTEM

Esophagus	+ +																								50			
Gallbladder	+ M + + +																								49			
Infiltration Cellular, Mononuclear Cell																									1	1 1.0		
Intestine Large, Cecum	+ +																								50			
Amyloid Deposition																									2	1 2.0		
Serosa, Inflammation, Granulomatous																									1	2.0		
Intestine Large, Colon	+ +																								50			
Intestine Large, Rectum	+ +																								50			
Intestine Small, Duodenum	+ +																								50			
Intestine Small, Ileum	+ +																								50			
Inflammation, Chronic																										1 2.0		
Epithelium, Hyperplasia																										1 2.0		
Intestine Small, Jejunum	+ +																								50			
Liver	+ +																								50			
Angiectasis	1																								1	1	3 1.0	
Basophilic Focus																										2		
Clear Cell Focus	X X X																								X X	X X	X X X X	18

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 M .. Missing tissue
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 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

	DAY ON TEST																								* TOTALS
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
B6C3F1 MICE MALE 0 G/KG	ANIMAL ID																								* TOTALS
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
Degeneration, Cystic Eosinophilic Focus		X		X	X	X	X	X		X		X	X	X			X	X		X	X	X		1 1.0 28	
Fatty Change	2	1	1	1	1	1			1	2	1	1	1		1	1	1	2	1					32 1.3	
Fibrosis														2										1 2.0	
Hematopoietic Cell Proliferation						2																		1 2.0	
Infiltration Cellular, Mononuclear Cell	1	2	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	49 1.0	
Mineralization																								1 1.0	
Mixed Cell Focus	X				X						X		X				X	X	X					15	
Necrosis														1										3 1.7	
Tension Lipidosis	X											X				X								5	
Mesentery																								3	
Inflammation, Chronic																								1 3.0	
Fat, Necrosis														2										2 2.0	
Pancreas	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	
Acinus, Atrophy																								1 4.0	
Acinus, Hypertrophy	2									1														2 1.5	
Salivary Glands	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	
Infiltration Cellular, Mononuclear Cell	1	1	1		1		2	1	1	1	1	1	1	1	1		1	1	1	1	1	1	1	43 1.1	
Parotid Gland, Atrophy																								1 2.0	
Stomach, Forestomach	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	
Inflammation, Chronic		2			1	2		2				1					2	1		1			2	19 1.7	
Necrosis						2		1									2						2	8 2.0	
Epithelium, Hyperplasia		3			1	2		3			2						2	3		3			2	18 2.3	
Stomach, Glandular	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	

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BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
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2) Mild 4) Marked

DAY ON TEST	0 0																								* TOTALS
	7 7 7 7 7 6 7 7 7 7 7 7 7 7 5 7 7 7 6 7 7 7 7 6																								
B6C3F1 MICE MALE 0 G/KG	3 3 3 3 3 1 3 3 3 3 3 3 3 3 1 3 3 3 2 3 3 3 3 8																								
	2 2 0 1 0 2 7 2 2 1 1 1 2 1 0 3 2 2 2 6 2 1 0 0 3																								
ANIMAL ID	0 0																								
	0 0																								
	2 2 2 2 3 3 3 3 3 3 3 3 3 3 4 4 4 4 4 4 4 4 4 5																								
	6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0																								

Adrenal Medulla Hyperplasia	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	1	2.0
Islets, Pancreatic Hyperplasia	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	37	1.9
Parathyroid Gland Cyst	+	+	+	M	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	47	1	1.0
Infiltration Cellular, Mononuclear Cell															2											1	2.0
Pituitary Gland Cyst	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	49	1	1.0
Pars Distalis, Hyperplasia																										2	2.0
Thyroid Gland Atrophy	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		3.0
Cyst																										3	1.8
Follicle, Hyperplasia																										2	2.0

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

Epididymis	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Granuloma Sperm																										3	2.5
Infiltration Cellular, Mononuclear Cell																										1	1.0

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 2) Mild 4) Marked
 Page 13

DAY ON TEST	0																								* TOTALS
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
	7	7	7	7	7	7	6	7	7	7	7	7	7	7	7	5	7	7	7	6	7	7	7	6	0
	3	3	3	3	3	3	1	3	3	3	3	3	3	3	3	1	3	3	3	2	3	3	3	3	8
	2	2	0	1	0	2	7	2	2	1	1	1	2	1	0	3	2	2	2	6	2	1	0	0	3
B6C3F1 MICE MALE 0 G/KG	ANIMAL ID																								
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2	2	2	2	3	3	3	3	3	3	3	3	3	3	3	4	4	4	4	4	4	4	4	4	5
	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0

Preputial Gland	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50			
Cyst					2																				4	2.0		
Inflammation						3														4				3	3	10	2.1	
Prostate	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Hyperplasia																										1	1.0	
Infiltration Cellular, Mononuclear Cell	1	1	1	2	1			1	1	1		1		1	1		1	1	1				1	1	1	1	29	1.0
Inflammation																			2								4	2.0
Seminal Vesicle	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Fibrosis																											2	1.5
Infiltration Cellular, Mononuclear Cell											1			1									1				4	1.0
Inflammation				1										1									1				6	1.0
Testes	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Inflammation																2											1	2.0
Mineralization									1																		1	1.0
Germinal Epithelium, Atrophy																								2	2		2	2.0

HEMATOPOIETIC SYSTEM

Bone Marrow	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Lymph Node, Mandibular	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	
Lymph Node, Mesenteric	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	
Atrophy																											1	3.0

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TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL
 Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

	DAY ON TEST	0																				* TOTALS				
		7	7	7	7	7	7	6	7	7	7	7	7	7	7	5	7	7	7	6	7		7	7	6	
		3	3	3	3	3	3	1	3	3	3	3	3	3	3	3	1	3	3	3	2	3	3	3	8	
		2	2	0	1	0	2	7	2	2	1	1	1	2	1	0	3	2	2	2	6	2	1	0	3	
B6C3F1 MICE MALE 0 G/KG	ANIMAL ID	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
		2	2	2	2	3	3	3	3	3	3	3	3	3	3	3	4	4	4	4	4	4	4	4	4	5
		6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0

Spleen	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50			
Hematopoietic Cell Proliferation			1				4	3				2	3				1	3					2		2	16	2.4	
Hyperplasia, Lymphoid									2						2											3	2.0	
Lymphoid Follicle, Atrophy															3							2				6	2.7	
Thymus	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	48		
Atrophy	1	4			2	1	4	4	2	4	2	2	2	2	4	3	2	3	3		3	2	4	3	2	4	41	2.7
Hyperplasia, Histiocytic																										1	3.0	

INTEGUMENTARY SYSTEM

Mammary Gland	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	M	1	
Skin	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50	
Inflammation, Chronic																											1	2.0
Ulcer																											1	3.0

MUSCULOSKELETAL SYSTEM

Bone	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Cranium, Hyperostosis															2													1	2.0
Skeletal Muscle																												2	

NERVOUS SYSTEM

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DAY ON TEST	0																								* TOTALS
	7	7	7	7	7	7	6	7	7	7	7	7	7	7	5	7	7	7	6	7	7	7	6	0	
	3	3	3	3	3	3	1	3	3	3	3	3	3	3	3	1	3	3	3	2	3	3	3	8	
	2	2	0	1	0	2	7	2	2	1	1	1	2	1	0	3	2	2	2	6	2	1	0	3	
B6C3F1 MICE MALE 0 G/KG	ANIMAL ID																								
	0																								
	0																								
	2																								
6																									

Brain + 50

RESPIRATORY SYSTEM

Lung	+ +	50
Alveolar Epithelium, Hyperplasia		2 1.5
Alveolus, Infiltration Cellular, Histiocyte		2 1.5
Bronchus, Hyperplasia		2 2.0
Nose	+ +	50
Inflammation, Suppurative		1 2.0
Inflammation, Chronic		9 1.0
Nasolacrimal Duct, Inflammation, Suppurative		1 2.0
Respiratory Epithelium, Hyperplasia		1 2.0
Trachea	+ +	50

SPECIAL SENSES SYSTEM

Eye	+ +	50
Harderian Gland	+ +	50
Fibrosis		2 1.0
Hyperplasia		4 2.0
Infiltration Cellular, Mononuclear Cell		36 1.1

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DAY ON TEST	0 0																								
	7 7 7 7 7 7 6 7 7 7 7 7 7 7 7 5 7 7 7 6 7 7 7 7 6																								
ANIMAL ID	3 3 3 3 3 3 1 3 3 3 3 3 3 3 3 1 3 3 3 2 3 3 3 3 8																								
	2 2 0 1 0 2 7 2 2 1 1 1 2 1 0 3 2 2 2 6 2 1 0 0 3																								
B6C3F1 MICE MALE	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0 G/KG	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2	2	2	2	3	3	3	3	3	3	3	3	3	3	3	4	4	4	4	4	4	4	4	4	5
	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0
																								* TOTALS	

URINARY SYSTEM

Kidney	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Hydronephrosis					2					2													2		6 2.2		
Inflammation													3												2 3.0		
Mineralization																					1				3 1.0		
Nephropathy		1	1	1	3	2	2		1	1			1		1	1		1	1	1	1	4	3	2	2	32 1.4	
Cortex, Medulla, Necrosis													3								4					6 3.2	
Papilla, Necrosis					2																		2	2		3 2.0	
Renal Tubule, Hyperplasia							2						1								2					6 1.3	
Renal Tubule, Mineralization	1		1	1						1	1		1	1	1		1		1	1	1	1	1	1		31 1.1	
Renal Tubule, Pigmentation, Lipofuscin						2																				1 2.0	
Urethra																									2		
Inflammation																						2				2 2.5	
Urinary Bladder	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	50		
Infiltration Cellular, Mononuclear Cell						1		1				1			1	1		1	1				1	1	1	1	20 1.0
Inflammation													3														4 2.0

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	DAY ON TEST	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
		5	4	7	6	7	7	7	7	7	7	4	6	7	7	7	7	5	6	7	7	7	7	7	3	
		4	3	3	9	3	3	0	2	3	3	3	6	3	3	3	3	5	2	3	3	3	3	3	1	
		6	7	1	3	2	2	7	0	1	1	2	1	4	1	1	2	2	3	3	2	2	1	1	2	9
B6C3F1 MICE MALE	ANIMAL ID	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
		5	5	5	5	5	5	5	5	5	6	6	6	6	6	6	6	6	6	6	7	7	7	7	7	
		1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5
0.25 G/KG																									males (cont...)	

ALIMENTARY SYSTEM

Esophagus	+ +																									
Gallbladder	+ +																									
Intestine Large, Cecum	+ +																									
Intestine Large, Colon	+ +																									
Intestine Large, Rectum	+ +																									
Intestine Small, Duodenum	+ +																									
Intestine Small, Ileum	+ +																									
Intestine Small, Jejunum Epithelium, Hyperplasia	+ +																									
Liver	+ +																									
Angiectasis																										
Basophilic Focus						1	1																			
Clear Cell Focus			X			X						X			X	X							X	X		
Eosinophilic Focus	X		X			X			X	X	X			X	X	X	X			X		X	X	X	X	
Fatty Change	1	2			1	1	1	2		1	1			1	1	1	2	1	2							
Hematopoietic Cell Proliferation									4																	
Infarct																										

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	DAY ON TEST																									males (cont...)
	0 5 4 6	0 4 3 7	0 7 3 1	0 6 9 3	0 7 3 2	0 7 3 2	0 7 0 7	0 7 2 0	0 7 3 1	0 7 3 1	0 7 3 2	0 4 6 1	0 7 3 1	0 7 3 1	0 7 3 2	0 5 2 3	0 6 3 3	0 7 3 2	0 7 3 2	0 7 3 1	0 7 3 1	0 7 3 2	0 7 3 1	0 7 3 2		
B6C3F1 MICE MALE	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
0.25 G/KG	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
	5	5	5	5	5	5	5	5	5	6	6	6	6	6	6	6	6	6	6	6	6	6	6	6	6	
	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	
Infiltration Cellular, Mononuclear Cell	1	2	1	1	1	1		1	1	1	1	1		1	1	1		1	1	1	1	1	1	1	1	
Mineralization																										
Mixed Cell Focus				X	X	X		X		X		X	X	X						X		X	X			
Necrosis				1																						
Tension Lipidosis																										
Vacuolization Cytoplasmic																										
Centrilobular, Fatty Change																										
Centrilobular, Hypertrophy		1	1		1	1				1		1	1	1		1	1	1	1		1	1	1	1	1	
Mesentery				+						+						+									+	
Fat, Necrosis				2						2						2									2	
Pancreas	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	
Salivary Glands	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	
Infiltration Cellular, Mononuclear Cell				1						1	1	1	1	1	1	1	1			1	1	1	1	1	1	
Parotid Gland, Mineralization																										
Submandibular Gland, Atrophy										2																
Stomach, Forestomach	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	
Inflammation, Chronic	2						1							1			1		1		1	2	1		2	
Necrosis	1																						2			
Epithelium, Hyperplasia	1						2					1			1		1		1		1	3	2		1	
Stomach, Glandular	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	
Mineralization																										
Glands, Ectasia																										
Tooth				+	+	+	+	+		+	+			+	+	+	+	+		+	+			+	+	

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

DAY ON TEST	0 0																								
	5 4 7 6 7 7 7 7 7 7 7 4 6 7 7 7 7 5 6 7 7 7 7 7 3																								
ANIMAL ID	4 3 3 9 3 3 0 2 3 3 3 3 6 3 3 3 3 5 2 3 3 3 3 3 1																								
	6 7 1 3 2 2 7 0 1 1 2 1 4 1 1 2 2 3 3 2 2 1 1 2 9																								
B6C3F1 MICE MALE	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0.25 G/KG	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	5	5	5	5	5	5	5	5	5	6	6	6	6	6	6	6	6	6	6	7	7	7	7	7	
	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5

males (cont...)

Dysplasia	2	2	4	3	3	1	2	4	2	3	1	1	3	2	1	2
Inflammation, Suppurative														3		

CARDIOVASCULAR SYSTEM

Blood Vessel	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Heart	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Cardiomyopathy			1			1	3								1	1			1	1				
Inflammation, Suppurative			2																					
Inflammation, Chronic							3																	
Artery, Infiltration Cellular, Mononuclear Cell								1																
Myocardium, Mineralization								1				1												

ENDOCRINE SYSTEM

Adrenal Cortex	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Hypertrophy																								
Vacuolization Cytoplasmic																								
Subcapsular, Hyperplasia	1	2	2	1	2	1		1	1	1	1	1	2	1	1	1	2		1		1	2	2	2
Zona Reticularis, Hyperplasia						2																2		
Adrenal Medulla	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Islets, Pancreatic	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
Hyperplasia	3		2	2	1	2	1		3	2	2		2	2		3	1		2		1	2	1	1

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 20

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

DAY ON TEST	0																								
	5	4	7	6	7	7	7	7	7	7	7	7	4	6	7	7	7	7	5	6	7	7	7	7	3
4	3	3	9	3	3	0	2	3	3	3	3	6	3	3	3	3	5	2	3	3	3	3	3	1	
6	7	1	3	2	2	7	0	1	1	2	1	4	1	1	2	2	3	3	2	2	1	1	2	9	
ANIMAL ID	0																								
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
5	5	5	5	5	5	5	5	5	5	6	6	6	6	6	6	6	6	6	6	7	7	7	7	7	
1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	

B6C3F1 MICE MALE
0.25 G/KG

males
(cont...)

Parathyroid Gland
Cyst

+ + + M + + + + + + + M M + + + + + + + M + + + +

Pituitary Gland

+ +

Thyroid Gland

+ +

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

Coagulating Gland
Hyperplasia

+
2

Epididymis
Infiltration Cellular, Mononuclear Cell
Inflammation, Chronic

+
 1 2 1 1 1 1 1 1

Penis
Infiltration Cellular, Polymorphonuclear

+
1

Preputial Gland
Cyst
Inflammation

+
 2 2
 1

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ .. Tissue examined microscopically

x .. Lesion present

I .. Insufficient tissue

M .. Missing tissue

A .. Autolysis precludes evaluation

BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 5 4 7 6 7 7 7 7 7 7 7 4 6 7 7 7 7 5 6 7 7 7 7 7 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 4 3 3 9 3 3 0 2 3 3 3 3 6 3 3 3 3 5 2 3 3 3 3 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 1 3 2 2 7 0 1 1 2 1 4 1 1 2 2 3 3 2 2 1 1 2 9 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

males (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Prostate | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Infiltration Cellular, Mononuclear Cell | | | | 1 | | | 1 | | | 1 | 1 | 1 | | | 1 | 1 | | 1 | | 1 | 1 | 1 | 1 | 1 | 1 |
| Inflammation | 3 | | 1 | 3 | | | 2 | | | | | | | | | | | 2 | 2 | | | | | | |
| Mineralization | | | | | | | | | | | | | 1 | | | | | | | | | | | | |
| Seminal Vesicle | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | 1 | | | | | | | | | | | |
| Inflammation | 2 | | | | | | | | | | | | | | | | | | 1 | 2 | | | | | |
| Testes | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Germinal Epithelium, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | 1 |

HEMATOPOIETIC SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Bone Marrow | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Lymph Node | | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mandibular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | 3 |
| Lymph Node, Mesenteric | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hyperplasia, Lymphoid | | | | 3 | | | | | | | | | | | | | | | | | | | | | |
| Spleen | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hematopoietic Cell Proliferation | | | | | | | 3 | 3 | | | | | | 2 | | 1 | | | 2 | | 2 | 2 | | 1 | |
| Hyperplasia, Lymphoid | | | | | | | | | | 2 | | | | | | | | | | | | | | | |
| Lymphoid Follicle, Atrophy | | | | 2 | | | | | | | | | | | | | | | | | 4 | | | | 1 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
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 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
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TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| | DAY ON TEST | Animals | | | | | | | | | | | | | | | | | | | | | | | | | males
(cont...) |
|-------------------------|-------------|---------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------------------|---|--------------------|
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 5 | 4 | 7 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 4 | 6 | 7 | 7 | 7 | 7 | 5 | 6 | 7 | 7 | 7 | 7 | 7 | 3 | | |
| | 4 | 3 | 3 | 9 | 3 | 3 | 0 | 2 | 3 | 3 | 3 | 3 | 6 | 3 | 3 | 3 | 3 | 5 | 2 | 3 | 3 | 3 | 3 | 3 | 1 | | |
| | 6 | 7 | 1 | 3 | 2 | 2 | 7 | 0 | 1 | 1 | 2 | 1 | 4 | 1 | 1 | 2 | 2 | 3 | 3 | 2 | 2 | 1 | 1 | 2 | 9 | | |
| B6C3F1 MICE MALE | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 0.25 G/KG | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | males
(cont...) | | |
| | | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Thymus | + | + | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Atrophy | | | 3 | 3 | | 3 | | 4 | 2 | 2 | 2 | | 4 | 2 | 4 | 2 | 1 | | 3 | 2 | 4 | 2 | 2 | 1 | 4 |
| Ectopic Thyroid | | | | | | | | | | | | | | 1 | | | | | | | | | | | |
| Hyperplasia, Histiocytic | | | | | | | | | | | | | | | | | | | | | | | | | |
| Epithelial Cell, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 2 |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Mammary Gland | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M |
| Skin | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | |
| Ulcer | | | | | | | | | | | | | | | | | | | | | | | | | |

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
|------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hemorrhage | | | | | | | | | | | | | | | | | | | | | | | | | |

RESPIRATORY SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 5 4 7 6 7 7 7 7 7 7 7 4 6 7 7 7 7 5 6 7 7 7 7 7 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 4 3 3 9 3 3 0 2 3 3 3 3 6 3 3 3 3 5 2 3 3 3 3 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 1 3 2 2 7 0 1 1 2 1 4 1 1 2 2 3 3 2 2 1 1 2 9 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

males (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | 1 | | | | | | | | | | | | | 1 | | |
| Alveolus, Infiltration Cellular, Histiocyte | | | | | | | | | | | 1 | 1 | | | | | | | | | | | | 3 |
| Bronchus, Hyperplasia | | | | | | | | | | | 1 | | | | | | | | | | | 1 | | |
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Chronic | | | | 1 | | | | | | | | | | 1 | | | | | 1 | | 1 | 1 | | |
| Polyp, Inflammatory | | | | | 2 | | | | | | | | | | | | | | | | | | | |
| Nasolacrimal Duct, Inflammation, Suppurative | | | | | | | | 2 | | | | | | | | | | | | | | | | |
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | 2 | |

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Eye | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Degeneration | | | | | | | | | | | | | | | | | | | | | | | | |
| Cornea, Inflammation, Chronic | | | | 3 | | | | | | | | | | | | | | | | | | | | |
| Harderian Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | 3 | | |
| Infiltration Cellular, Mononuclear Cell | | | | | | 1 | 1 | 1 | 1 | | | 1 | | 1 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 5 4 7 6 7 7 7 7 7 7 7 4 6 7 7 7 7 5 6 7 7 7 7 7 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 4 3 3 9 3 3 0 2 3 3 3 3 6 3 3 3 3 5 2 3 3 3 3 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 1 3 2 2 7 0 1 1 2 1 4 1 1 2 2 3 3 2 2 1 1 2 9 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
0.25 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 5 5 5 5 5 5 5 5 5 6 6 6 6 6 6 6 6 6 6 6 6 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |

males (cont...)

URINARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Kidney | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hydronephrosis | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Metaplasia, Osseous | | | | | | | | | | | | | | | | | | | | | | | | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | |
| Nephropathy | 2 1 1 1 1 1 1 1 1 2 1 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Thrombosis | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Renal Tubule, Dilatation | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Renal Tubule, Hyperplasia | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Renal Tubule, Mineralization | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Urethra | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Cyst | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Urinary Bladder | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | 1 4 4 | | | | | | | | | | | | | | | | | | | | | | | | |
| Transitional Epithelium, Hyperplasia | 4 | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
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 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 5 7 7 6 7 4 7 7 7 7 6 4 7 7 7 7 6 7 5 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 9 3 3 9 3 2 3 3 3 3 6 8 3 3 3 3 8 3 5 3 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 2 1 5 2 3 1 2 2 2 4 4 2 1 2 1 7 1 3 2 1 0 2 2 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
0.25 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 9 | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | * TOTALS | | | | | | | | | | | | | | | | | | | | | | | |

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|-----------|---------------|
| Esophagus | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Gallbladder | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Large, Cecum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Large, Colon | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Large, Rectum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Duodenum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Ileum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Jejunum Epithelium, Hyperplasia | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| | 3 | | | | | | | | | | | | | | | | | | | | | | | | 3 | 2 3.0 |
| Liver | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Angiectasis | 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 | 6 1.0 |
| Basophilic Focus | X | | | | | | | | | | | | | | | | | | | | | | | | 3 | 18 |
| Clear Cell Focus | X X | | | | | | | | | | | | | | | | | | | | | | | | 2 | 32 |
| Eosinophilic Focus | X | | | | | | | | | | | | | | | | | | | | | | | | 32 | 34 1.3 |
| Fatty Change | 1 1 1 2 1 2 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2 2.5 |
| Hematopoietic Cell Proliferation | 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 3.0 |
| Infarct | | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|---|-------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|---------------|
| | 0731 | 0736 | 0742 | 0747 | 0753 | 0758 | 0804 | 0809 | 0815 | 0820 | 0826 | 0831 | 0837 | 0843 | 0848 | 0854 | 0900 | 0905 | 0911 | 0916 | 0922 | 0927 | 0933 | 0938 | |
| B6C3F1 MICE MALE | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | 0000 | |
| 0.25 G/KG | 7761 | 7766 | 7772 | 7777 | 7783 | 7788 | 7794 | 7800 | 7805 | 7811 | 7816 | 7822 | 7827 | 7833 | 7838 | 7844 | 7850 | 7855 | 7861 | 7866 | 7872 | 7877 | 7883 | 7888 | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 2 | 47 1.1 |
| Mineralization | | | | | | 2 | | | | | | | | | | | | 1 | | | | | | | 2 1.5 |
| Mixed Cell Focus | X | | X | X | | | | X | X | X | | | X | X | X | X | | X | X | X | X | X | X | | 28 |
| Necrosis | | 2 | | 2 | 3 | | | | 1 | | | | | | | | | 2 | | | | | | 2 | 10 2.0 |
| Tension Lipidosis | | | | | | | | | | | | | | | | | | | X | | | | | | 2 |
| Vacuolization Cytoplasmic | | | | | 2 | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Centrilobular, Fatty Change | | | | | | | | | | | | | | | | | | 3 | | | | | | | 1 3.0 |
| Centrilobular, Hypertrophy | 1 | 1 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | | | 1 | 1 | | | | 2 | 1 | | 1 | 1 | 1 | | 34 1.0 |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | 6 |
| Fat, Necrosis | | 3 | | | 2 | | | | | | | | | | | | | | | | | | | | 6 2.2 |
| Pancreas | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Infiltration Cellular, Mononuclear Cell | 1 | | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 3 | 1 | 2 | 1 | | 1 | 1 | | 1 | 1 | 1 | 36 1.1 |
| Parotid Gland, Mineralization | | | | | | | | | | | | | | 1 | | | | | | | | | | | 1 1.0 |
| Submandibular Gland, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation, Chronic | 2 | 2 | | | | 2 | | 1 | 2 | | | 2 | 2 | | | 2 | | | 1 | 2 | | 1 | | | 22 1.6 |
| Necrosis | 2 | | | 2 | | 2 | | | | | | 2 | | | | 2 | | | 2 | 2 | | | | 2 | 11 1.9 |
| Epithelium, Hyperplasia | 2 | 1 | | 2 | | 2 | | 2 | | | | 2 | 2 | | | 2 | | | 1 | 2 | | 1 | | | 22 1.6 |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1.0 |
| Glands, Ectasia | | | | | | 1 | | | | | | | | | | | | | | | | | | | 1 1.0 |
| Tooth | | + | + | + | | | | | | + | + | + | + | | | | | + | + | + | + | + | + | + | 32 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
|-------------------------------|---|--|--|-------|--|--|---|--|--|-------|--|--|-------|--|--|---|--|--|---|--|--|--------|--|--|----------|--|
| | 7 5 7 7 6 7 4 7 7 7 7 6 4 7 7 7 7 6 7 5 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
0.25 G/KG | 3 9 3 3 9 3 2 3 3 3 3 6 8 3 3 3 3 8 3 5 3 3 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
| | 1 6 2 1 5 2 3 1 2 2 2 4 4 2 1 2 1 7 1 3 2 1 0 2 2 | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
| 0.25 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 9 9 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
| Dysplasia | 1 2 3 | | | 2 1 3 | | | 1 | | | 3 3 1 | | | 2 1 1 | | | 4 | | | 2 | | | 32 2.1 | | | | |
| Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 | |

CARDIOVASCULAR SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------|--|
| Blood Vessel | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Heart | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Cardiomyopathy | 1 | | | | | | | | | | | | | | | | | | | | | | | | 16 1.1 | |
| Inflammation, Suppurative | 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | |
| Inflammation, Chronic | 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 | |
| Artery, Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Myocardium, Mineralization | 1 | | | | | | | | | | | | | | | | | | | | | | | | 4 1.0 | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------|--|
| Adrenal Cortex | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Hypertrophy | 1 | | | | | | | | | | | | | | | | | | | | | | | | 2 1.0 | |
| Vacuolization Cytoplasmic | 1 | | | | | | | | | | | | | | | | | | | | | | | | 2 1.0 | |
| Subcapsular, Hyperplasia | 1 2 1 1 1 1 1 2 1 1 1 1 1 1 1 1 2 2 2 1 2 2 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 45 1.3 | |
| Zona Reticularis, Hyperplasia | 1 | | | | | | | | | | | | | | | | | | | | | | | | 3 1.7 | |
| Adrenal Medulla | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Islets, Pancreatic | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Hyperplasia | 2 1 2 1 2 2 2 2 2 1 1 3 2 1 1 2 1 2 | | | | | | | | | | | | | | | | | | | | | | | | 35 1.7 | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL
 Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
|------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|--|
| DAY ON TEST | 7 | 5 | 7 | 7 | 6 | 7 | 4 | 7 | 7 | 7 | 7 | 6 | 4 | 7 | 7 | 7 | 7 | 6 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 7 | | |
| | 3 | 9 | 3 | 3 | 9 | 3 | 2 | 3 | 3 | 3 | 3 | 6 | 8 | 3 | 3 | 3 | 3 | 8 | 3 | 5 | 3 | 3 | 3 | 3 | 3 | 3 | | |
| | 1 | 6 | 2 | 1 | 5 | 2 | 3 | 1 | 2 | 2 | 2 | 4 | 4 | 2 | 1 | 2 | 1 | 7 | 1 | 3 | 2 | 1 | 0 | 2 | 2 | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | | * TOTALS | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|------------|-----------|--|
| Parathyroid Gland
Cyst | + | + | + | + | + | + | M | + | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | 44 | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|--|
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
|-----------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|--|

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|--|
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
|---------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|--|

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|----------|------------|
| Coagulating Gland
Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 | 2.0 |
|----------------------------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|----------|------------|

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|------------|---|-----------|--|
| Epididymis
Infiltration Cellular, Mononuclear Cell
Inflammation, Chronic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 22 | 1.2 | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|----------|------------|
| Penis
Infiltration Cellular, Polymorphonuclear | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 | 1.0 |
|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|----------|------------|

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|------------|---|-----------|--|
| Preputial Gland
Cyst
Inflammation | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 5 | 2.0 | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 4 | 1.0 | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ .. Tissue examined microscopically

x .. Lesion present

l .. Insufficient tissue

M .. Missing tissue

A .. Autolysis precludes evaluation

BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|-------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|
| | 7 5 7 7 6 7 4 7 7 7 7 6 4 7 7 7 7 6 7 5 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 9 3 3 9 3 2 3 3 3 3 6 8 3 3 3 3 8 3 5 3 3 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 2 1 5 2 3 1 2 2 2 4 4 2 1 2 1 7 1 3 2 1 0 2 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| 0.25 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------|
| Prostate | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | 1 1 1 1 1 1 1 1 1 1 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 31 1.0 |
| Mineralization | 2 2 2 | | | | | | | | | | | | | | | | | | | | | | | | 9 2.1 |
| Seminal Vesicle | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Infiltration Cellular, Mononuclear Cell | 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 4 1.0 |
| Inflammation | 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 5 1.4 |
| Testes | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Germinal Epithelium, Atrophy | 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | | 4 2.0 |

HEMATOPOIETIC SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------|
| Bone Marrow | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Lymph Node | + | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Lymph Node, Mandibular | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Hyperplasia, Lymphoid | 3 4 3 4 | | | | | | | | | | | | | | | | | | | | | | | | 3 3.3 |
| Lymph Node, Mesenteric | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Hyperplasia, Lymphoid | 1 1 1 4 3 1 1 2 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| Spleen | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Hematopoietic Cell Proliferation | 3 2 1 1 1 4 3 1 1 2 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | | 19 1.9 |
| Hyperplasia, Lymphoid | 2 2 2 | | | | | | | | | | | | | | | | | | | | | | | | 3 2.0 |
| Lymphoid Follicle, Atrophy | 3 2 | | | | | | | | | | | | | | | | | | | | | | | | 5 2.4 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|---|--|
| | 7 5 7 7 6 7 4 7 7 7 7 6 4 7 7 7 7 6 7 5 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 9 3 3 9 3 2 3 3 3 3 6 8 3 3 3 3 8 3 5 3 3 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 2 1 5 2 3 1 2 2 2 4 4 2 1 2 1 7 1 3 2 1 0 2 2 | | | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation | 2 | | | | | | | | | | | | | | | | | | | | | | | | | 2 1.5 |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 3 1.0 |
| Alveolus, Infiltration Cellular, Histiocyte | 3 | | | | | 2 | | | | | | | | | | | | | | | | | | | | 6 1.8 |
| Bronchus, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 2 1.0 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | 8 1.0 |
| Polyp, Inflammatory | | | | | | | | | | | | | | | | | | | | | | | | | | 2 2.0 |
| Nasolacrimal Duct, Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | | 2 2.0 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------|
| Eye | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Degeneration | | | | | | | | | | | | | | | | | | | | | | | | | | 1 4.0 |
| Cornea, Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| Harderian Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 3 2.3 |
| Infiltration Cellular, Mononuclear Cell | 1 | | 1 | 1 | | | | | | | | | | | | | | | | | | | | | | 30 1.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 7 7 7 7 7 7 7 7 6 7 6 6 7 7 7 6 5 7 7 7 5 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 3 3 3 3 3 1 3 2 7 3 3 3 7 3 3 3 3 8 0 3 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 1 2 2 1 1 1 1 6 2 3 0 0 1 2 9 4 0 1 1 0 5 1 2 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | |

males (cont...)

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Esophagus | + | | | | | | | | | | | | | | | | | | | | | | | |
| Gallbladder
Infiltration Cellular, Mononuclear Cell
Inflammation, Granulomatous | + + M +
1 | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Large, Cecum | + | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Large, Colon | + | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Large, Rectum | + | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Small, Duodenum | + | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Small, Ileum
Inflammation, Chronic | + | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Small, Jejunum
Peyer's Patch, Hyperplasia, Lymphoid | + | | | | | | | | | | | | | | | | | | | | | | | |
| Liver
Amyloid Deposition | + | | | | | | | | | | | | | | | | | | | | | | | |
| Angiectasis | 1 1 2 1 | | | | | | | | | | | | | | | | | | | | | | | |
| Basophilic Focus | X | | | | | | | | | | | | | | | | | | | | | | | |
| Clear Cell Focus | X | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | males
(cont...) | |
|---|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|--------------------|--|
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1 | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | |
| Degeneration, Cystic Eosinophilic Focus | | | | | | | | | | | | | | | 1 | | | | | | | | | | | |
| Fatty Change | X | X | X | X | X | X | X | X | X | X | | X | X | X | X | | | X | X | X | X | | X | X | | |
| Hepatodiaphragmatic Nodule | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | | | 1 | 1 | 1 | | 2 | 1 | | | 1 | 1 | | | | 2 | | |
| Infarct | | | | 3 | | | | | | | | | | | | | | | 2 | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | 2 | | |
| Mixed Cell Focus | X | | | | X | | | X | | | | | | X | X | | | X | X | | | | | | | |
| Necrosis | | | | | | | 2 | | | | | | | | | 2 | | | | 2 | | | | 2 | | |
| Tension Lipidosis | | | | | | | | | | | | X | | | | | | | | | | | | | | |
| Centrilobular, Hypertrophy | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | | | | 2 | | 2 | | 2 | | | | |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Fat, Necrosis | | | | | | | + | | | | | | | | | | | | | | | | | | | |
| Pancreas | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Granulomatous | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Acinus, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Salivary Glands | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | | 1 | 1 | 1 | 1 | | | 1 | 1 | | 1 | 1 | 1 | | |
| Parotid Gland, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | 2 | | |
| Stomach, Forestomach | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------------------------|
| | 7 7 7 7 7 7 7 7 7 6 7 6 6 7 7 7 6 5 7 7 7 5 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 3 3 3 3 3 1 3 2 7 3 3 3 7 3 3 3 3 8 0 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 1 2 2 1 1 1 1 6 2 3 0 0 1 2 9 4 0 1 1 0 5 1 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | males
(cont...) |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | |

Stomach, Glandular Infiltration Cellular, Mononuclear Cell Glands, Ectasia +

Tooth Dysplasia +
 3 1 2 4 3 4 2 2 3 2 1 3 1 4

CARDIOVASCULAR SYSTEM

Blood Vessel +

Heart +
 Cardiomyopathy 2 1 1 1 1 3 1
 Artery, Infiltration Cellular, Mononuclear Cell

ENDOCRINE SYSTEM

Adrenal Cortex +
 Vacuolization Cytoplasmic 1 2
 Subcapsular, Hyperplasia 1 1 1 1 1 2 2 2 1 1 1 1 2 1 2 1 2 2 1 1 1 1 1
 Zona Reticularis, Hyperplasia

Adrenal Medulla +
 Hyperplasia 2

Islets, Pancreatic +

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue

M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | | | | | | | |
|----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 6 | 7 | 6 | 6 | 7 | 7 | 7 | 6 | 5 | 7 | 7 | 7 | 5 | 6 | 7 | 7 | | | | | | | | |
| ANIMAL ID | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 1 | 3 | 2 | 7 | 3 | 3 | 3 | 7 | 3 | 3 | 3 | 3 | 8 | 0 | 3 | 3 | | | | | | | | |
| | 1 | 2 | 1 | 2 | 2 | 1 | 1 | 1 | 1 | 6 | 2 | 3 | 0 | 0 | 1 | 2 | 9 | 4 | 0 | 1 | 1 | 0 | 5 | 1 | 2 | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | | | | | | | |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | | | | | | | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | | | | | | | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | | | | | | | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | | | | | | | | |
| | | | | | | | | | | | | | | males (cont...) | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | 2 | 2 | 1 | 2 | 3 | 1 | 3 | 2 | | | | | | | | | | | | | | 1 | 1 | 1 | 1 | 1 | 2 | 4 | 2 | 3 | 2 | 2 | 2 |
| Parathyroid Gland | + | + | + | + | + | + | + | + | + | M | + | + | + | + | + | + | + | + | + | M | + | + | + | + | + | | | | | | | | |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | | | | | | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Pars Distalis, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | | | | | | |
| Cyst | | | | | | | | | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Follicle, Hyperplasia | | | | | | | | | | | | | | | | | 1 | | | | | | | | | | | | | | | | |

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

Coagulating Gland

+

Epididymis

Granuloma Sperm

Infiltration Cellular, Mononuclear Cell

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--|--|---|---|
| + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | | | | |
| 1 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | | | | | | | | | | | | | | 1 | 1 | | | 1 | 1 | | | 1 | 1 |

Preputial Gland

Cyst

Inflammation

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| | 3 | | | | | | | | | | | | | | | | | | | 2 | | | | | |
| | | | | | | | | | | 2 | 2 | | | | | 1 | | | | | | | | | 2 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ .. Tissue examined microscopically

x .. Lesion present

I .. Insufficient tissue

M .. Missing tissue

A .. Autolysis precludes evaluation

BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 7 7 7 7 7 7 7 7 7 6 7 6 6 7 7 7 6 5 7 7 7 5 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 3 3 3 3 3 3 1 3 2 7 3 3 3 7 3 3 3 3 8 0 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 1 2 2 1 1 1 1 1 6 2 3 0 0 1 2 9 4 0 1 1 0 5 1 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

males (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Prostate | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Fibrosis | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 1 1 1 1 1 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | 1 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Seminal Vesicle | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Testes | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Germinal Epithelium, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | |

HEMATOPOIETIC SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Bone Marrow | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node Renal, Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mandibular Hyperplasia, Lymphoid | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mesenteric Hyperplasia, Lymphoid | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Spleen | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Angiectasis | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Hematopoietic Cell Proliferation | 1 2 2 2 2 1 2 1 1 2 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia, Lymphoid | 2 4 | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 7 7 7 7 7 7 7 7 6 7 6 6 7 7 7 6 5 7 7 7 5 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 3 3 3 3 3 1 3 2 7 3 3 3 7 3 3 3 3 8 0 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 1 2 2 1 1 1 1 6 2 3 0 0 1 2 9 4 0 1 1 0 5 1 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

males
(cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Lymphoid Follicle, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | |
| Thymus | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Atrophy | 2 | 2 | 2 | 3 | 3 | 4 | 2 | 2 | 3 | 3 | 2 | 4 | 4 | 2 | | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 3 | |
| Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | | | | |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--|--|---|
| Mammary Gland | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | | | |
| Skin | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | |
| Cyst Epithelial Inclusion | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Ulcer | | | | | | | | | | 3 | | | | | | | 3 | | | | | | | 3 | | | | 3 |

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Skeletal Muscle | | | | | | | | | | | | | | | | | | | | | | | | + | |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hypothalamus, Compression | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 39

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 7 7 7 7 7 7 7 7 6 7 6 6 7 7 7 6 5 7 7 7 5 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 3 3 3 3 3 1 3 2 7 3 3 3 7 3 3 3 3 8 0 3 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 1 2 2 1 1 1 1 6 2 3 0 0 1 2 9 4 0 1 1 0 5 1 2 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | |

males
(cont...)

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|---|--|--|--|--|--|--|--|--|--|--|--|---|--|
| Lung | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Metaplasia, Osseous | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Alveolus, Infiltration Cellular, Histiocyte | 2 | | | | | | | | | | | | 1 | | | | | | | | | | | | 3 | |
| Bronchus, Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Nose | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Suppurative | 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Inflammation, Chronic | | | | | | | | | | | | | 1 | | | | | | | | | | | | | |
| Nasolacrimal Duct, Inflammation, Suppurative | | | | | | | | | | | | | 1 | | | | | | | | | | | | 2 | |
| Respiratory Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Trachea | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Chronic | | | | | | | | | | | | | 2 | | | | | | | | | | | | | |
| Epithelium, Cytoplasmic Alteration | | | | | | | | | | | | | 2 | | | | | | | | | | | | | |

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Eye | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Cornea, Inflammation, Chronic | | | | | | | | | | | | | 2 | | | | | | | | | | | | | |
| Harderian Gland | + | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
|---|-------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|----------------|------|
| | 0732 | 0733 | 0734 | 0735 | 0736 | 0737 | 0738 | 0739 | 0740 | 0741 | 0742 | 0743 | 0744 | 0745 | 0746 | 0747 | 0748 | 0749 | 0750 | 0751 | 0752 | 0753 | 0754 | 0755 | | 0756 |
| B6C3F1 MICE MALE | 0011 | 0012 | 0013 | 0014 | 0015 | 0016 | 0017 | 0018 | 0019 | 0020 | 0021 | 0022 | 0023 | 0024 | 0025 | 0026 | 0027 | 0028 | 0029 | 0030 | 0031 | 0032 | 0033 | 0034 | 0035 | |
| 0.5 G/KG | 0036 | 0037 | 0038 | 0039 | 0040 | 0041 | 0042 | 0043 | 0044 | 0045 | 0046 | 0047 | 0048 | 0049 | 0050 | 0051 | 0052 | 0053 | 0054 | 0055 | 0056 | 0057 | 0058 | 0059 | 0060 | |
| Degeneration, Cystic Eosinophilic Focus | X | X | X | | X | X | X | X | X | | X | X | X | X | | X | X | X | | X | X | X | X | | 1 1.0
42 | |
| Fatty Change | | 1 | | | | | | 1 | 1 | | | | 2 | 1 | 1 | | 1 | | | 3 | | 1 | 1 | | 27 1.2
1 | |
| Hepatodiaphragmatic Nodule Infarct | | | | | | | | | | | | | | | | | | | | | | | | | 2 2.5 | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 47 1.0 | |
| Mineralization | | | | | | | | | | | | | | | | | | | | 1 | | | | | 2 1.5 | |
| Mixed Cell Focus | | X | | | | | | | X | X | X | X | X | | X | | | | X | | | | | | 15 | |
| Necrosis | | | | | | | | | | | | | | 1 | | | | 3 | | | | | | 2 | 7 2.0 | |
| Tension Lipidosis | | | | | | | | | X | | | | | X | | | | | | | | | | | 3 | |
| Centrilobular, Hypertrophy | 2 | 2 | | 2 | 2 | | | 2 | 2 | | 2 | | 2 | 2 | | 2 | 2 | | | 2 | | 2 | | | 30 2.0 | |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | 2 | |
| Fat, Necrosis | | | | | | | | | 2 | | | | | | | | | | | | | | | | 2 2.0 | |
| Pancreas | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Inflammation, Granulomatous | | | | | | | | | | | | | | | | | | | | | 1 | | | | 1 1.0 | |
| Inflammation, Chronic Acinus, Atrophy | | | | 2 | | | | | | | | | 3 | | | | | | | | | | | | 1 2.0
1 3.0 | |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 40 1.0 | |
| Parotid Gland, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Inflammation, Chronic | | | 2 | | | | | | | | | | 1 | | 1 | | 2 | | 1 | 2 | 1 | 1 | 1 | 2 | 24 1.6 | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | 1 | | | 1 1.0 | |
| Necrosis | | | | | | | | | | | | | | | 2 | | | | | | 3 | | | | 12 2.3 | |
| Epithelium, Hyperplasia | | | 2 | | | | | | | | | 2 | | 1 | | 2 | | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 25 2.0 | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 7 7 7 6 7 7 7 6 6 7 7 7 7 6 7 4 7 7 7 4 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 6 2 3 3 5 8 3 3 3 1 8 3 5 3 3 3 9 3 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 1 0 8 9 1 2 3 3 1 1 0 1 3 1 5 1 2 1 4 0 0 0 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |
| | * TOTALS | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|-----|
| Stomach, Glandular Infiltration Cellular, Mononuclear Cell Glands, Ectasia | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| | | | | | | | | | 1 | | | | | | | | | | | | | | | | | 2 | 1.0 | |
| | | | | | | | | | | | | | | | 1 | | | | | | | | | | | | 2 | 1.0 |
| Tooth Dysplasia | + | | | | + | + | | | | | | + | | + | + | + | + | | | + | + | + | + | | + | 27 | | |
| | 3 | | | | 3 | 3 | | | | | | 1 | | 2 | 3 | 4 | 4 | | | 2 | 2 | 1 | 2 | | 1 | 27 | 2.4 | |

CARDIOVASCULAR SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|
| Blood Vessel | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Cardiomyopathy | | | | | | | | 1 | | | | | | | | | | | | | | | | | | 11 | 1.3 |
| Artery, Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1.0 |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|-----|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Vacuolization Cytoplasmic Subcapsular, Hyperplasia | 2 | 1 | 1 | 1 | 2 | 2 | 1 | 1 | 2 | 1 | | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 48 | 1.3 | |
| Zona Reticularis, Hyperplasia | | | | | 1 | | | | | | | | | | | | | | | | | | | | | 2 | 2 | 1.5 |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 | |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
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TDMS No. 20007 - 06
 Test Type: CHRONIC
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 Species/Strain: MICE/B6C3F1

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Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---------------|
| | 7 7 7 7 6 7 7 7 6 6 7 7 7 7 6 7 4 7 7 7 4 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 6 2 3 3 5 8 3 3 3 1 8 3 5 3 3 3 9 3 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 1 0 8 9 1 2 3 3 1 1 0 1 3 1 5 1 2 1 4 0 0 0 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| 0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 2 3 3 3 3 3 3 3 3 3 3 4 4 4 4 4 4 4 4 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | 2 | 1 | | 2 | 2 | 2 | 1 | 1 | | 2 | 2 | 2 | 2 | | 2 | 1 | | 1 | 2 | 1 | | | | | 36 1.8 |
| Parathyroid Gland | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | M | + | + | + | + | + | + | + | 46 |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Cyst | | | | | | | | | | | | | | | | | | | | | | 1 | | | 1 1.0 |
| Pars Distalis, Hyperplasia | | | | | | | | | 2 | | | | | | | | | | | | | | | | 1 2.0 |
| Thyroid Gland | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 49 |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |
| Follicle, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---------------|
| Coagulating Gland | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Epididymis | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Granuloma Sperm | | | | | | 2 | | | | | | | | | | | | | | | | 3 | 2 | | 3 2.3 |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | 1 | 1 | | 33 1.0 |
| Preputial Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Cyst | | | | | | | | | | | | | | | | | 2 | | | | | | | 2 | 5 2.2 |
| Inflammation | | | | 2 | | | | | | | 1 | 1 | 2 | | | | | | | 2 | | | | 2 | 9 1.7 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ .. Tissue examined microscopically

x .. Lesion present

I .. Insufficient tissue

M .. Missing tissue

A .. Autolysis precludes evaluation

BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 7 | 7 | 7 | 7 | 6 | 7 | 7 | 7 | 6 | 6 | 7 | 7 | 7 | 7 | 6 | 7 | 4 | 7 | 7 | 7 | 4 | 7 | 6 | 7 | 7 |
| | 3 | 3 | 3 | 3 | 6 | 2 | 3 | 3 | 5 | 8 | 3 | 3 | 3 | 1 | 8 | 3 | 5 | 3 | 3 | 3 | 9 | 3 | 6 | 3 | 3 |
| | 2 | 2 | 1 | 0 | 8 | 9 | 1 | 2 | 3 | 3 | 1 | 1 | 0 | 1 | 3 | 1 | 5 | 1 | 2 | 1 | 4 | 0 | 0 | 0 | 2 |
| B6C3F1 MICE MALE
0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------|
| Prostate | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Fibrosis | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | 1 | 1 | 1 | 1 | | | | | 1 | 1 | 1 | 1 | 38 1.0 |
| Inflammation | | | | | | | | | | | | | | 2 | | | | | | 2 | | | | | 4 1.8 |
| Seminal Vesicle | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | | | | | | | | | | | | | | | | 1 | | | | | 1 | | 6 1.0 |
| Testes | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Germinal Epithelium, Atrophy | | | | | | | | | | | | | | | | | | | | | | | 2 | | 1 2.0 |

HEMATOPOIETIC SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------|
| Bone Marrow | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Lymph Node | | | | + | | | | | | | | | | | | | | | | | | | | | 3 |
| Renal, Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | 4 | | 1 4.0 |
| Lymph Node, Mandibular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Hyperplasia, Lymphoid | | | | 2 | | | | | | | | | | | | | | | | | | | | | 2 3.0 |
| Lymph Node, Mesenteric | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Hyperplasia, Lymphoid | | | | 2 | | | | | | | | | | | | | | | | | | | 4 | | 2 3.0 |
| Spleen | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 49 |
| Angiectasis | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Hematopoietic Cell Proliferation | | | | 2 | 2 | 3 | | | | 3 | | | | 2 | | | | | | | 2 | 2 | 2 | 2 | 21 2.0 |
| Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | 2 3.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| | 7 | 7 | 7 | 7 | 6 | 7 | 7 | 7 | 6 | 6 | 7 | 7 | 7 | 7 | 6 | 7 | 4 | 7 | 7 | 7 | 4 | 7 | 6 | 7 | |
| B6C3F1 MICE MALE
0.5 G/KG | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 3 | 3 | 3 | 3 | 6 | 2 | 3 | 3 | 5 | 8 | 3 | 3 | 3 | 1 | 8 | 3 | 5 | 3 | 3 | 3 | 9 | 3 | 6 | 3 | 3 |
| ANIMAL ID | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 | 2 | 1 | 0 | 8 | 9 | 1 | 2 | 3 | 3 | 1 | 1 | 0 | 1 | 3 | 1 | 5 | 1 | 2 | 1 | 4 | 0 | 0 | 0 | 2 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------------------|---|---|---|---|---|--|---|---|---|---|---|-----|---|---|--|---|---|---|---|---|---|---|---|---|----|-----|---|
| Lymphoid Follicle, Atrophy | 1 | | | | | | | | | | | 1 2 | | | | | | | | | | | | | 3 | 1.3 | |
| Thymus | + | | | | | | | | | | | M | | + | | | | | | | | | | | 47 | | |
| Atrophy | 4 | 2 | 3 | 4 | 4 | | 4 | 2 | 3 | 4 | 1 | 3 | 2 | 4 | | 2 | 2 | 3 | 4 | 2 | 3 | 4 | 4 | 4 | M | + | 4 |
| Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | 2 | | | | | | | | | | | 2 | 2.0 | |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|--|--|--|---|--|---|--|--|--|--|---|--|--|--|--|--|--|--|---|--|---|---|-----|----|---|-----|
| Mammary Gland | M | | | | | | | | | | | | | | | | | | | | | | | | 0 | | |
| Skin | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Cyst Epithelial Inclusion | | | | | | | | | | | | 2 | | | | | | | | | | | | | 1 | 2 | 1.5 |
| Ulcer | 3 | | | | 4 | | 4 | | | | | | | | | | | | | 3 | | 3 | 9 | 3.2 | | | |

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----|--|
| Bone | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Skeletal Muscle | | | | | | | | | | | | | | | | | | | | | | | | | 1 | |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|--|--|--|----|---|-----|
| Brain | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Hypothalamus, Compression | | | | | | | | | | | | | | | | | | | | | 4 | | | | 4 | 1 | 4.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 47

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|-----------------|
| | 7 7 7 7 6 7 7 7 6 6 7 7 7 7 6 7 4 7 7 7 4 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 3 6 2 3 3 5 8 3 3 3 1 8 3 5 3 3 3 9 3 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 1 0 8 9 1 2 3 3 1 1 0 1 3 1 5 1 2 1 4 0 0 0 2 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 2 3 3 3 3 3 3 3 3 3 3 3 3 4 4 4 4 4 4 4 5 | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation | | | | | | | 1 | | | | | 1 | | | | | | | | | 2 | | | | 3 1.3 |
| Metaplasia, Osseous | | | | | | | 1 | | | | | | | | | | | | | | | | | | 1 1.0 |
| Mineralization | | | | | | | 1 | | | 1 | | | | | | | | | | | | | | | 2 1.0 |
| Alveolar Epithelium, Hyperplasia | | | | | | | 1 | | | | | | | | | | | | | | | | | | 1 1.0 |
| Alveolus, Infiltration Cellular, Histiocyte | | | | | | | 3 | | | | | | | | | | | | | | | | 2 | | 5 2.2 |
| Bronchus, Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | | 1 | | 1 1.0 |
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation, Suppurative | | | | | | | | | | | | | | | 2 | 2 | | | | | | | | | 4 1.5 |
| Inflammation, Chronic | | | | | | | 1 | | | | 1 | | | | | | 1 | | | | | | | | 7 1.0 |
| Nasolacrimal Duct, Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | 2 1.5 |
| Respiratory Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | 1 | | 1 1.0 |
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Epithelium, Cytoplasmic Alteration | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Eye | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Cornea, Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Harderian Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | males
(cont...) | |
|---|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|--------------------|--|
| | 0
7
3
1 | 0
5
9
6 | 0
5
4
7 | 0
7
3
1 | 0
4
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5 | 0
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9
6 | 0
7
3
1 | 0
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3
2 | 0
7
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0 | 0
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3
2 | 0
7
3
1 | 0
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2 | 0
7
7
2 | 0
7
7
2 | 0
7
7
2 | 0
7
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1 | 0
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1 | 0
5
6
4 | 0
6
7
8 | 0
7
3
1 | 0
7
3
2 | 0
5
3
2 | 0
7
8
2 | 0
3
9
5 | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 1.0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | | |
| Fatty Change | 1 | 2 | 1 | 1 | | 1 | | 1 | | | | | | | | | | 1 | | 1 | | | | | 1 | 2 | |
| Hematopoietic Cell Proliferation | | | | | | | | | | | | 2 | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 2 | 1 | 1 | | |
| Inflammation, Suppurative | | | | | | | | | | | | 3 | | | | | | | | | | | | | | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | 2 | | | | |
| Mixed Cell Focus | | | | | | | X | | | | | | X | | | | | | | | | | | | | X | |
| Necrosis | | | | | 3 | | | | | 2 | | 1 | | 1 | | 1 | | | 3 | 1 | | | 3 | 3 | | | |
| Tension Lipidosis | | | X | X | | | | | | | | | | | | | | | | | | | | | | | |
| Vacuolization Cytoplasmic | | | | | | | | | | | | | 2 | | | | | | | | | | | | | | |
| Bile Duct, Cyst | | | | | | | | | | 2 | | | | | | | | | | | | | | | | | |
| Centrilobular, Fatty Change | | | | | | | | | | | | | | | | | | | | | | | 4 | | | | |
| Centrilobular, Hypertrophy | 2 | 2 | 2 | 2 | 2 | 2 | 2 | | 2 | | 2 | | | | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | | |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Fat, Necrosis | | | | | | + | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | 2 | | | | | | | | | | | | | | | | | | | | | |
| Pancreas | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Duct, Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | X | |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | | | 1 | | 1 | 1 | 1 | 1 | | 1 | 1 | 2 | 1 | 1 | | 1 | 1 | | 1 | 1 | | 1 | | |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Inflammation, Chronic | | | | | | | | | 2 | 1 | | 2 | 1 | 1 | 1 | 1 | 1 | | 2 | 2 | | | 2 | 1 | 1 | | |
| Necrosis | | | | | | | | | | | | 1 | | | | | | | 1 | 1 | | | 1 | 3 | | | |
| Epithelium, Hyperplasia | | | | | | | | 2 | 2 | | 2 | 2 | 2 | 2 | 1 | 3 | | 2 | 2 | | | | 2 | | | | |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 5 5 7 4 5 7 7 7 7 7 6 7 7 7 7 7 5 6 7 7 5 3 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 9 4 3 6 9 3 3 3 3 3 7 3 3 3 3 3 7 6 3 3 8 9 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 7 1 5 6 1 2 0 2 1 3 2 2 2 1 1 0 4 8 1 2 2 2 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
1.0 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 5 5 5 5 5 5 5 6 6 6 6 6 6 6 6 6 6 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |

males
(cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Glands, Ectasia | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Tooth Dysplasia | + | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 1 1 4 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |

CARDIOVASCULAR SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Blood Vessel | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Heart | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Cardiomyopathy | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Suppurative Artery, Infiltration Cellular, Mononuclear Cell | 2 2 2 2 1 1 2 2 1 1 1 1 1 1 1 1 1 1 2 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Myocardium, Mineralization Valve, Inflammation | 4 | | | | | | | | | | | | | | | | | | | | | | | | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Adrenal Cortex | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hypertrophy | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Vacuolization Cytoplasmic Subcapsular, Hyperplasia | 2 2 2 2 1 1 2 2 1 1 1 1 1 1 1 1 1 1 2 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Zona Reticularis, Hyperplasia | 2 2 2 2 1 1 2 2 1 1 1 1 1 1 1 1 1 1 2 2 3 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Adrenal Medulla | + | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 5 5 7 4 5 7 7 7 7 7 6 7 7 7 7 7 5 6 7 7 5 3 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 9 4 3 6 9 3 3 3 3 3 7 3 3 3 3 3 7 6 3 3 8 9 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 7 1 5 6 1 2 0 2 1 3 2 2 2 1 1 0 4 8 1 2 2 2 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
1.0 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 5 5 5 5 5 5 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |

males (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|--|---|--|---|---|---|---|--|--|--|--|---|---|---|---|---|---|---|---|--|---|---|---|
| Atrophy | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | | 1 | | 1 | 1 | 1 | 1 | | | | | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | | 1 | 1 | 1 |
| Inflammation | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Seminal Vesicle | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Atrophy | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Testes | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Germinal Epithelium, Atrophy | 2 | | | | | | | | | | | | | | | | | | | | | | | | |

HEMATOPOIETIC SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Bone Marrow | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mandibular | + M + + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia, Lymphoid | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mesenteric | + + + + + + + + + + + + + M + + + + + + + + + + + | | | | | | | | | | | | | | | | | | | | | | | | |
| Spleen | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hematopoietic Cell Proliferation | 4 2 2 3 1 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia, Lymphoid | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymphoid Follicle, Atrophy | 1 4 4 | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 54

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 5 5 7 4 5 7 7 7 7 7 6 7 7 7 7 7 5 6 7 7 5 3 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 9 4 3 6 9 3 3 3 3 3 7 3 3 3 3 3 7 6 3 3 8 9 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 7 1 5 6 1 2 0 2 1 3 2 2 2 1 1 0 4 8 1 2 2 2 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 1.0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

males
(cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Thymus | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Atrophy | 1 | | | 2 | 4 | | 4 | 2 | 4 | 4 | 4 | 4 | 4 | 2 | 2 | 4 | 4 | 4 | 2 | 2 | 2 | M | + | + |
| Inflammation, Chronic Active Epithelial Cell, Hyperplasia | | | | | | | | | | 3 | | | | | | | | | | | | | | |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Mammary Gland | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M |
| Skin | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hemorrhage | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Chronic Ulcer | | | | | | | | | | | 3 | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | |

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
|------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|

RESPIRATORY SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | | males
(cont...) |
|------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------------------|--------------------|
| | 7 5 5 7 4 5 7 7 7 7 7 6 7 7 7 7 7 5 6 7 7 5 3 7 | | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
1.0 G/KG | 3 9 4 3 6 9 3 3 3 3 3 7 3 3 3 3 3 7 6 3 3 8 9 3 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 7 1 5 6 1 2 0 2 1 3 2 2 2 1 1 0 4 8 1 2 2 2 1 | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
| 1.0 G/KG | 1 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 5 5 5 5 5 5 5 5 6 6 6 6 6 6 6 6 6 6 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | males
(cont...) | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|---|--|--|--|--|--|--|---|--|--|--|--|--|--|--|---|---|--|--|--|---|
| Lung | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | 2 | | | | | | | | | | | | | 1 |
| Alveolus, Infiltration Cellular, Histiocyte | | | | | | | | | | | | | 3 | | | | | | | | | | | | | |
| Bronchus, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Nose | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Suppurative | | | | | | | | | | | | | 2 | | | | | | | | | | | | | |
| Inflammation, Chronic | 2 | | | | | 1 | | | | | | | | | | | | | | | 1 | 1 | | | | |
| Polyp, Inflammatory | | | | | | | | | | | | | 2 | | | | | | | | | | | | | |
| Nasolacrimal Duct, Inflammation, Suppurative | 2 | | | | | 1 | | | | | | | | | | | | | | | | | | | | |
| Trachea | + | | | | | | | | | | | | | | | | | | | | | | | | | |

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|---|---|---|---|---|---|---|---|--|--|---|--|---|---|---|---|---|---|---|---|---|
| Eye | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Anterior Chamber, Inflammation, Suppurative | | | | | | | | | | | | | 3 | | | | | | | | | | | | | |
| Cornea, Inflammation, Chronic | 2 | | | | | | | | | | | | | | | 1 | | | | | | | | | | |
| Retrolbulbar, Inflammation, Chronic | | | | | | | | | | | | | 2 | | | | | | | | | | | | | |
| Harderian Gland | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | | | | | | | | | | | | | 2 | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | | | | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |

URINARY SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
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Kava kava extract
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 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------------------|
| | 7 5 5 7 4 5 7 7 7 7 7 6 7 7 7 7 7 5 6 7 7 5 3 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
1.0 G/KG | 3 9 4 3 6 9 3 3 3 3 3 7 3 3 3 3 3 7 6 3 3 8 9 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 6 7 1 5 6 1 2 0 2 1 3 2 2 2 1 1 0 4 8 1 2 2 2 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| 1.0 G/KG | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 5 5 5 5 5 5 5 5 6 6 6 6 6 6 6 6 6 6 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | males
(cont...) |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|---|---|---|---|---|---|---|---|---|---|---|---|--|---|---|---|---|---|--|--|---|---|---|---|
| Kidney | + | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Hydronephrosis | | | | | | | | | | | | | | | | | | | | | | | | | | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Metaplasia, Osseous | 1 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Nephropathy | 1 | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2 | 1 | 1 | 2 | 1 | 2 | 2 | 1 | 1 | 1 | 1 | 2 | | 1 | 1 | 2 | 1 | 1 | | | | | | |
| Papilla, Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | | | |
| Renal Tubule, Hyperplasia | 1 | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | | | | | | | | | | | | | | | | | | | | | 1 | | |
| Renal Tubule, Mineralization | 1 | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Renal Tubule, Pigmentation, Lipofuscin | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 3 |

Urethra
 Inflammation
 Necrosis

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|
| Urinary Bladder | + | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Transitional Epithelium, Cytoplasmic Alteration | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
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 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
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 1-4 .. Lesion qualified as:
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TDMS No. 20007 - 06
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Kava kava extract
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| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|---|
| | 5 7 6 7 7 7 7 7 7 7 7 7 7 7 7 7 5 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 8 3 9 3 3 1 3 3 3 3 3 3 3 3 3 3 9 3 3 3 3 0 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 1 2 2 0 8 2 2 1 1 0 1 2 0 1 1 1 5 1 2 2 1 7 2 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 1.0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |
| | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------------|---------------|
| Esophagus | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Gallbladder | + | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 47 |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Intestine Small, Duodenum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Intestine Small, Ileum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Intestine Small, Jejunum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Peyer's Patch, Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | 1 4.0 | |
| Liver | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Amyloid Deposition | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 | |
| Angiectasis | | | 1 | | 1 | | 2 | 2 | | | | | | 2 | 1 | | | | 2 | | | | | | 10 1.7 |
| Basophilic Focus | | | X | | | | | | | | | | | | | | | | | | | | | | 1 |
| Clear Cell Focus | | | | | X | | X | X | X | | | X | X | X | X | X | | X | X | X | X | | | | 21 |
| Degeneration, Cystic | | | | | X | | X | X | X | X | X | X | X | X | X | X | | X | X | X | X | X | X | X | 1 1.0 |
| Eosinophilic Focus | X | X | X | X | X | X | X | X | X | X | X | X | X | X | X | X | | X | X | X | X | X | X | X | 43 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
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 1-4 .. Lesion qualified as:
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| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|---|-------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|----------|
| | 0582 | 0576 | 0567 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | 0577 | |
| B6C3F1 MICE MALE | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | 0011 | |
| 1.0 G/KG | 76 | 77 | 78 | 79 | 80 | 81 | 82 | 83 | 84 | 85 | 86 | 87 | 88 | 88 | 88 | 88 | 88 | 88 | 88 | 88 | 88 | 88 | 88 | 88 | |
| Fatty Change | 1 | | | | 1 | 2 | | | 1 | 2 | 2 | 2 | 1 | | 2 | | | | 1 | 1 | 2 | 1 | | | |
| Hematopoietic Cell Proliferation | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | |
| Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | |
| Mixed Cell Focus | | | | | X | | | | X | X | X | X | | | X | | | | X | X | | X | | | |
| Necrosis | 2 | | 1 | | | | | | | | | | | | | | | 2 | | | | 3 | | | |
| Tension Lipidosis | | | | | | | | | | | | | | | | | | | | | | | | | |
| Vacuolization Cytoplasmic | | | | | | | | | | | | | | | | | | | | | | | | | |
| Bile Duct, Cyst | | | | | | | | | | | | | | | | | | | | | | | | | |
| Centrilobular, Fatty Change | | | | | | | | | | | | | | | | | | | | | | | | | |
| Centrilobular, Hypertrophy | 2 | 2 | 2 | 2 | | 2 | 2 | 2 | 2 | | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | | |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | |
| Fat, Necrosis | | | | | | | | | | | | | | | | | | | | | | | 3 | | |
| Pancreas | | | | | | | | | | | | | | | | | | | | | | | | | |
| Duct, Cyst | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Salivary Glands | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | |
| Stomach, Forestomach | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Chronic | 1 | | 2 | 1 | 2 | | | 1 | 1 | 2 | 2 | | | | | | | | 2 | | 2 | | 1 | | |
| Necrosis | 3 | | 2 | | 2 | | | | | 2 | | | | | | | | | | | 2 | | | | |
| Epithelium, Hyperplasia | 1 | | 1 | 1 | | | | 1 | 1 | 2 | 2 | | | | | | | | 3 | | 2 | 2 | 2 | | |
| Stomach, Glandular | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | 50 | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
|-------------------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 5 | 7 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 7 | 0 | |
| 8 | 3 | 9 | 3 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 9 | 3 | 3 | 3 | 3 | 0 | 3 | 3 | 0 |
| 2 | 1 | 2 | 2 | 0 | 8 | 2 | 2 | 1 | 1 | 0 | 1 | 2 | 0 | 1 | 1 | 1 | 1 | 5 | 1 | 2 | 2 | 1 | 7 | 2 | 2 | 0 |
| B6C3F1 MICE MALE | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 1.0 G/KG | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 |
| 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 |
| 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 0 | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|----|-----|
| Islets, Pancreatic
Hyperplasia | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 33 | 1.3 |
| Parathyroid Gland
Cyst | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | M | + | + | + | + | + | + | 44 | 1 | 1.0 |
| Pituitary Gland
Cyst | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 2 | 1.0 |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |

GENERAL BODY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|---|-----|
| Peritoneum
Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2 | 2.5 |
|-------------------------------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|---|-----|

GENITAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|----|-----|
| Epididymis
Granuloma Sperm
Infiltration Cellular, Mononuclear Cell | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 1 | 2.0 |
| | 1 | | | | 1 | 1 | | | | 1 | 1 | 1 | | | | 1 | 1 | 1 | | | | 1 | 1 | | | | 28 | 1.0 |
| Preputial Gland
Cyst
Inflammation | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 3 | 2.3 |
| | | | 1 | | | | | | | 1 | 1 | 2 | | | | 2 | 3 | | | | 2 | | | | | | 13 | 1.3 |
| Prostate | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 61

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
|------------------------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | 0 |
| 5 | 7 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | |
| 8 | 3 | 9 | 3 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 9 | 3 | 3 | 3 | 3 | |
| 2 | 1 | 2 | 2 | 0 | 8 | 2 | 2 | 1 | 1 | 0 | 1 | 2 | 0 | 1 | 1 | 1 | 1 | 5 | 1 | 2 | 2 | 1 | 7 | 2 | 2 | |
| B6C3F1 MICE MALE
1.0 G/KG | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | 0 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | 0 |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | 1 |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | | 9 |
| 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|-----|
| Thymus | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 49 | |
| Atrophy | 4 | 4 | 3 | 4 | 2 | 4 | | 1 | 4 | 1 | 2 | 2 | 3 | 1 | 3 | 2 | 2 | 4 | 2 | 2 | 3 | 1 | 4 | | 2 | 43 | 2.8 | |
| Inflammation, Chronic Active | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 3.0 |
| Epithelial Cell, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|
| Mammary Gland | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | M | 0 | |
| Skin | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Hemorrhage | | | 3 | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 3.0 |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 |
| Ulcer | | | 3 | 3 | | | | | | | | | | | | | | | | | | | | | | | | 4 | 3.0 |

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
|------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|

RESPIRATORY SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue

M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

Page 63

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 5 | 7 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 7 | | |
| | 8 | 3 | 9 | 3 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 9 | 3 | 3 | 3 | 3 | 0 | 3 | 3 | |
| | 2 | 1 | 2 | 2 | 0 | 8 | 2 | 2 | 1 | 1 | 0 | 1 | 2 | 0 | 1 | 1 | 1 | 5 | 1 | 2 | 2 | 1 | 7 | 2 | 2 | |
| | | | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | |
| 1.0 G/KG | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | * TOTALS |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|---|-----|-----|-----|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | | |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 1.0 | | |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 1.5 | |
| Alveolus, Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 3.0 |
| Bronchus, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|--|--|---|-----|-----|
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | | | |
| Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 5 | 1.2 |
| Polyp, Inflammatory | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 |
| Nasolacrimal Duct, Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 1.7 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|--|
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
|---------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|--|

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|--|--|--|---|-----|
| Eye | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | | | |
| Anterior Chamber, Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 3.0 |
| Cornea, Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 1.5 |
| Retrolbulbar, Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|--|--|--|----|-----|
| Harderian Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | | | |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 8 | 1.9 |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 39 | 1.0 |

URINARY SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 64

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|
| | 5 7 6 7 7 7 7 7 7 7 7 7 7 7 7 7 5 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE MALE
1.0 G/KG | 8 3 9 3 3 1 3 3 3 3 3 3 3 3 3 3 9 3 3 3 3 0 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 1 2 2 0 8 2 2 1 1 0 1 2 0 1 1 1 5 1 2 2 1 7 2 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 9 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------|
| Kidney | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Hydronephrosis | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| Metaplasia, Osseous | | | | | | | | | | | | | | | | | | | | | | | | | 2 1.5 |
| Nephropathy | 2 2 2 2 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 2 2 1 | | | | | | | | | | | | | | | | | | | | | | | | 38 1.3 |
| Papilla, Necrosis | 1 | | | | | | | | | | | | | | | | | | | | | | | | 3 1.0 |
| Renal Tubule, Hyperplasia | 1 1 2 1 | | | | | | | | | | | | | | | | | | | | | | | | 10 1.1 |
| Renal Tubule, Mineralization | 1 | | | | | | | | | | | | | | | | | | | | | | | | 31 1.0 |
| Renal Tubule, Pigmentation, Lipofuscin | | | | | | | | | | | | | | | | | | | | | | | | | 2 2.5 |
| Urethra | + | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Inflammation | 3 | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| Necrosis | 3 | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| Urinary Bladder | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | 20 1.0 |
| Transitional Epithelium, Cytoplasmic Alteration | 2 | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |

*** END OF MALE DATA ***

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 65

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 7 7 7 5 7 7 7 6 7 7 7 7 7 7 7 5 7 7 7 5 7 7 6 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 2 4 2 2 3 8 3 2 3 2 3 3 3 2 6 0 3 2 8 2 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 1 0 9 4 9 9 1 4 0 9 1 9 1 1 0 9 0 1 0 9 1 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 0 0 0 0 0 0 0 0 0 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 females (cont...) | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Mixed Cell Focus | X | | | | | | | | X | | | | | | | | | | X | X | | | | | |
| Necrosis | | | | 1 | | | | | | | | | | | | | | | | 2 | | | | | |
| Tension Lipidosis | | | | X | | | | | | | | | | | | | | | | X | | | | | |
| Mesentery | | | | + | | | | | | | | | | | | | | | | + | | | | | + |
| Fat, Necrosis | | | | 3 | | | | | | | | | | | | | | | | 2 | | | | | 3 |
| Pancreas | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Inflammation, Chronic | | | | | 1 | | | | | | | | | | | | | | | | | | | | |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Infiltration Cellular, Mononuclear Cell | 1 | | | 1 | | 1 | 1 | 1 | | | 1 | 1 | | 1 | 1 | | 1 | | | 1 | 1 | 3 | 2 | 1 | |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | |
| Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Infiltration Cellular, Mononuclear Cell | | | | | 1 | | | | | | | | | | | | | | | | | | | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | |
| Glands, Ectasia | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Tooth | | | | | | | | | | | | | | | | | | | | | | | | | |
| Dysplasia | | | | | | | | | | | | | | | | | | | | | | | | | |

CARDIOVASCULAR SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Blood Vessel | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
|--------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 67

TDMS No. 20007 - 06

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract

CAS Number: 9000-38-8

Date Report Requested: 04/13/2009

Time Report Requested: 10:57:30

First Dose M/F: 08/24/04 / 08/23/04

Lab: BAT

| DAY ON TEST | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
|---------------------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| DAY ON TEST | | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 5 | 7 | 7 | 6 | |
| DAY ON TEST | | 3 | 3 | 3 | 2 | 4 | 2 | 2 | 3 | 8 | 3 | 2 | 3 | 2 | 3 | 3 | 3 | 2 | 6 | 0 | 3 | 2 | 8 | 2 | 3 | 1 |
| DAY ON TEST | | 0 | 1 | 0 | 9 | 4 | 9 | 9 | 1 | 4 | 0 | 9 | 1 | 9 | 1 | 1 | 0 | 9 | 0 | 1 | 0 | 9 | 1 | 9 | 0 | 0 |
| B6C3F1 MICE FEMALE | ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | ANIMAL ID | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 |
| | ANIMAL ID | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

females (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--|
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Cardiomyopathy | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Artery, Infiltration Cellular, Mononuclear Cell | | 1 | | | | | | | | | | | | | | | | | | | | | | | | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Vacuolization Cytoplasmic Subcapsular, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Subcapsular, Hyperplasia | 2 | 2 | 2 | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 2 | 2 | 2 |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Parathyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Pars Distalis, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 4 |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Ectopic Thymus | | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked
Page 68

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------------------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 5 | 7 | 7 | 6 | |
| | 3 | 3 | 3 | 2 | 4 | 2 | 2 | 3 | 8 | 3 | 2 | 3 | 2 | 3 | 3 | 3 | 2 | 6 | 0 | 3 | 2 | 8 | 2 | 3 | 1 | |
| | 0 | 1 | 0 | 9 | 4 | 9 | 9 | 1 | 4 | 0 | 9 | 1 | 9 | 1 | 1 | 0 | 9 | 0 | 1 | 0 | 9 | 1 | 9 | 0 | 0 | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0 G/KG | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | females (cont...) |

Infiltration Cellular, Mononuclear Cell
 Follicle, Hyperplasia

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

Clitoral Gland
 Inflammation

+ + + + M +

Ovary
 Angiectasis
 Cyst
 Hemorrhage

+ + M +
 2
 3

Uterus
 Inflammation, Suppurative
 Metaplasia, Squamous
 Endometrium, Hyperplasia, Cystic

+
 2
 2 2 2 3 3 2 1 3 1 3 1 2 1 1 3 2 1 2 3 3 2

HEMATOPOIETIC SYSTEM

Bone Marrow

+ +

Lymph Node

+ +

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ .. Tissue examined microscopically

x .. Lesion present

I .. Insufficient tissue

M .. Missing tissue

A .. Autolysis precludes evaluation

BLANK .. Not examined microscopically

Page 69

1-4 .. Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 7 7 7 5 7 7 7 6 7 7 7 7 7 7 7 5 7 7 7 5 7 7 6 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 3 3 2 4 2 2 3 8 3 2 3 2 3 3 3 2 6 0 3 2 8 2 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 1 0 9 4 9 9 1 4 0 9 1 9 1 1 0 9 0 1 0 9 1 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

females (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Mediastinal, Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mandibular Hyperplasia, Lymphoid | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | M | + | + | + |
| Lymph Node, Mesenteric | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Spleen Hematopoietic Cell Proliferation | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Hyperplasia, Lymphoid | | 2 | 2 | | | | | | | | | | | | | | | | | | | | | | | |
| Lymphoid Follicle, Atrophy | | | | | | 1 | | | | | | | | | | | | | | | | | | | | |
| Thymus Atrophy | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Hyperplasia, Histiocytic Infiltration Cellular, Histiocyte | | | 1 | | | 2 | 2 | 2 | 3 | 1 | 2 | 2 | | 2 | 3 | | 2 | 3 | 3 | 1 | 2 | 4 | | 1 | 4 | |
| Epithelial Cell, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Mammary Gland Hyperplasia Inflammation, Chronic Active Metaplasia, Squamous | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Skin Inflammation, Chronic Ulcer | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 7 7 7 5 7 7 7 6 7 7 7 7 7 7 7 5 7 7 7 5 7 7 6 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0 G/KG | 3 3 3 2 4 2 2 3 8 3 2 3 2 3 3 3 2 6 0 3 2 8 2 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 1 0 9 4 9 9 1 4 0 9 1 9 1 1 0 9 0 1 0 9 1 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 0 0 0 0 0 0 0 0 0 1 1 1 1 1 1 1 1 1 1 2 2 2 2 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |

females (cont...)

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Bone | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Fibrosis | 2 3 2 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Skeletal Muscle | + | | | | | | | | | | | | | | | | | | | | | | | | |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Brain | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Choroid Plexus, Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Hypothalamus, Compression | 4 | | | | | | | | | | | | | | | | | | | | | | | | |
| Peripheral Nerve | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Axon, Degeneration | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Spinal Cord | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Axon, Degeneration | 3 | | | | | | | | | | | | | | | | | | | | | | | | |

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Lung | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hemorrhage | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
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 2) Mild 4) Marked

TDMS No. 20007 - 06

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: MICE/B6C3F1

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Kava kava extract

CAS Number: 9000-38-8

Date Report Requested: 04/13/2009

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First Dose M/F: 08/24/04 / 08/23/04

Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | females
(cont...) | | |
|---|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|---|----------------------|---|--|
| | 0
7
3
0 | 0
7
3
1 | 0
7
3
0 | 0
7
2
9 | 0
5
4
4 | 0
7
2
9 | 0
7
2
9 | 0
7
3
1 | 0
6
8
4 | 0
7
3
0 | 0
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2
9 | 0
7
3
1 | 0
7
2
9 | 0
7
3
1 | 0
7
3
1 | 0
7
3
0 | 0
5
6
0 | 0
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1 | 0
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0 | 0
7
2
9 | 0
5
8
1 | 0
7
2
9 | 0
7
3
0 | 0
7
2
0 | | | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | | | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | | | |
| Metaplasia, Osseous Mineralization | | | | | | | | | | | | | | | | | 1 | | | | | | | | 2 | | | |
| Nephropathy | 1 | | | 1 | 1 | | 1 | 1 | 1 | | 1 | 1 | 1 | | 1 | | | | | 2 | | 2 | 1 | 1 | | 2 | 1 | |
| Papilla, Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Renal Tubule, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Urinary Bladder Infiltration Cellular, Mononuclear Cell | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | |
| | | 1 | 1 | 1 | | 1 | 1 | | 1 | 1 | | 1 | | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | | |

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+ .. Tissue examined microscopically

x .. Lesion present

I .. Insufficient tissue

M .. Missing tissue

A .. Autolysis precludes evaluation

BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|
| | 7 4 7 6 7 7 7 5 6 7 7 7 7 7 7 7 3 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0 G/KG | 2 2 2 3 2 3 3 2 8 3 3 2 2 3 3 2 3 2 9 3 3 0 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 9 1 9 9 9 0 0 3 8 0 1 9 9 0 0 9 1 9 4 0 1 0 0 1 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----|--------|
| Esophagus | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Gallbladder Cyst | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | 1 1.0 |
| Intestine Large, Cecum Inflammation, Chronic | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | 1 1.0 |
| Intestine Large, Colon | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Large, Rectum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Duodenum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Ileum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Jejunum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Liver | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Angiectasis | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Clear Cell Focus | X | | | | | | | | | | | | | | | | | | | | | | | | 1 | 3 1.0 |
| Eosinophilic Focus | X | | | | | | | | | | | | | | | | | | | | | | | | 1 | 3 |
| Fatty Change | X | | | | | | | | | | | | | | | | | | | | | | | | 1 | 9 |
| Hematopoietic Cell Proliferation | 2 2 1 2 | | | | | | | | | | | | | | | | | | | | | | | | 2 | 39 1.6 |
| Infiltration Cellular, Mononuclear Cell | 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 2.0 |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | 25 | 45 1.1 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract

CAS Number: 9000-38-8

Date Report Requested: 04/13/2009

Time Report Requested: 10:57:30

First Dose M/F: 08/24/04 / 08/23/04

Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|---------------------------|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 7 | 4 | 7 | 6 | 7 | 7 | 7 | 5 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 3 | 7 | 7 | 7 | 7 | 7 | 7 | |
| | 2 | 2 | 2 | 3 | 2 | 3 | 3 | 2 | 8 | 3 | 3 | 2 | 2 | 3 | 3 | 2 | 3 | 2 | 9 | 3 | 3 | 0 | 3 | 3 | |
| | 9 | 1 | 9 | 9 | 9 | 0 | 0 | 3 | 8 | 0 | 1 | 9 | 9 | 0 | 0 | 9 | 1 | 9 | 4 | 0 | 1 | 0 | 0 | 1 | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | |
| | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-------|
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Cardiomyopathy | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2 1.0 |
| Artery, Infiltration Cellular, Mononuclear Cell | | | | | | | | 1 | | | | | | | | | | | | | | | | | | 1 1.0 |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|--------|-------|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Atrophy | | | | | | | | | | | | | 1 | | | | | | | | | | | | | 1 1.0 | |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Vacuolization Cytoplasmic Subcapsular, Hyperplasia | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | | 49 1.9 | |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 1 2 | 4 1.3 |
| Parathyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | M | + | + | + | + | M | + | 48 | |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1.0 |
| Pars Distalis, Hyperplasia | 2 | | | | | | | | | | | | | | | | | | | | | | | | | 2 3 | 4 2.8 |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Ectopic Thymus | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

TDMS No. 20007 - 06

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract

CAS Number: 9000-38-8

Date Report Requested: 04/13/2009

Time Report Requested: 10:57:30

First Dose M/F: 08/24/04 / 08/23/04

Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|
| | 7 4 7 6 7 7 7 5 6 7 7 7 7 7 7 7 3 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0 G/KG | 2 2 2 3 2 3 3 2 8 3 3 2 2 3 3 2 3 2 9 3 3 0 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 9 1 9 9 9 0 0 3 8 0 1 9 9 0 0 9 1 9 4 0 1 0 0 1 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 2 3 3 3 3 3 3 3 3 3 3 3 3 4 4 4 4 4 4 4 4 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |

| | | |
|---|---|-------|
| Infiltration Cellular, Mononuclear Cell | 1 | 1 1.0 |
| Follicle, Hyperplasia | 2 | 1 2.0 |

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

| | | | | |
|---|---|-----------------------|-------|--------|
| Clitoral Gland Inflammation | + | 1 | 49 | 1 1.0 |
| Ovary Angiectasis | + | | 49 | 1 2.0 |
| Ovary Cyst | | X | 2 | 4 1.7 |
| Ovary Hemorrhage | | | 1 | 1 3.0 |
| Uterus Inflammation, Suppurative | + | 2 | 50 | 3 1.3 |
| Uterus Metaplasia, Squamous | | | 1 | 1 2.0 |
| Uterus Endometrium, Hyperplasia, Cystic | 3 2 2 3 1 3 2 | 3 2 3 3 1 2 3 2 4 1 2 | 1 2 1 | 42 2.1 |

HEMATOPOIETIC SYSTEM

| | | | | |
|-------------|---|--|----|---|
| Bone Marrow | + | | 50 | |
| Lymph Node | | | | 3 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|
| | 7 4 7 6 7 7 7 5 6 7 7 7 7 7 7 7 3 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0 G/KG | 2 2 2 3 2 3 3 2 8 3 3 2 2 3 3 2 3 2 9 3 3 0 3 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 9 1 9 9 9 0 0 3 8 0 1 9 9 0 0 9 1 9 4 0 1 0 0 1 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 3 3 3 3 3 3 3 3 3 3 3 3 3 4 4 4 4 4 4 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|
| Mediastinal, Hyperplasia, Lymphoid | 4 | | | | | | | | | | | | | | | | | | | | | | | | 1 4.0 |
| Lymph Node, Mandibular
Hyperplasia, Lymphoid | + M + + + | | | | | | | | | | | | | | | | | | | | | | | | 48
4 3.3 |
| Lymph Node, Mesenteric | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Spleen
Hematopoietic Cell Proliferation
Hyperplasia, Lymphoid
Lymphoid Follicle, Atrophy | + | | | | | | | | | | | | | | | | | | | | | | | | 49
26 1.8
7 1.6
4 2.8 |
| Thymus
Atrophy
Hyperplasia, Histiocytic
Infiltration Cellular, Histiocyte
Epithelial Cell, Hyperplasia | + | | | | | | | | | | | | | | | | | | | | | | | | 50
39 2.6
1 3.0
5 2.4
1 1.0 |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|-------------------------------|
| Mammary Gland
Hyperplasia
Inflammation, Chronic Active
Metaplasia, Squamous | + | | | | | | | | | | | | | | | | | | | | | | | | 50
1 2.0
1 2.0
1 2.0 |
| Skin
Inflammation, Chronic
Ulcer | + | | | | | | | | | | | | | | | | | | | | | | | | 50
1 3.0
1 4.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|---------------------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 7 | 4 | 7 | 6 | 7 | 7 | 7 | 5 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 3 | 7 | 7 | 7 | 7 | 7 | 7 | 0 |
| 2 | 2 | 2 | 3 | 2 | 3 | 3 | 2 | 8 | 3 | 3 | 2 | 2 | 3 | 3 | 2 | 3 | 2 | 9 | 3 | 3 | 0 | 3 | 3 | 3 | 0 |
| 9 | 1 | 9 | 9 | 9 | 0 | 0 | 3 | 8 | 0 | 1 | 9 | 9 | 0 | 0 | 9 | 1 | 9 | 4 | 0 | 1 | 0 | 0 | 1 | 0 | 0 |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 | 5 |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|-----|
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1.0 | |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 3 | 1.0 | |
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 4 | 1.3 |
| Olfactory Epithelium, Degeneration | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 | 2.0 |
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|---|-----|-----|
| Ear | | | | | | | | | | | | | | | | | | | | | | | | | 1 | | | |
| Eye | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | |
| Anterior Chamber, Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | | 3 | 1 | 3.0 | |
| Harderian Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | |
| Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1.0 | |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 3 | 1.7 | |
| Infiltration Cellular, Mononuclear Cell | 1 | | 1 | 1 | 1 | 1 | | 1 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | | 1 | 1 | | 2 | 1 | 1 | 1 | 1 | 1 | 38 | 1.0 |

URINARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|---|---|-----|
| Kidney | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | |
| Amyloid Deposition | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 | 1.0 |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 | 1.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 |
| 8 | 3 | 2 | 2 | 3 | 2 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 2 | 2 | 3 | 2 | 6 | 2 | 3 | 6 | 2 | 0 | 3 | 2 |
| 1 | 1 | 9 | 9 | 0 | 9 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 4 | 9 | 0 | 3 | 5 | 1 | 0 | 6 | 9 | 1 | 0 | 9 |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 |
| 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 |

B6C3F1 MICE FEMALE
0.25 G/KG

females (cont...)

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Esophagus
Inflammation, Chronic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Gallbladder
Cyst | + | + | + | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Small, Duodenum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Small, Ileum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Small, Jejunum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Liver | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Basophilic Focus | | | | | | | | | | | | | | | | | | | | | | | | | |
| Eosinophilic Focus | | X | | | | | | | | | | | | X | | | | | | | | | | | X |
| Fatty Change | 2 | 1 | 1 | 2 | | | 2 | | 2 | 2 | 2 | 2 | | | 1 | | 2 | | 1 | 4 | | | 1 | 1 | |
| Hematopoietic Cell Proliferation | | | | 1 | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| Mixed Cell Focus | X | | | | | | | | | | X | | | | | | | | | | X | X | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue

M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically

1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

Page 82

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | females
(cont...) |
|---|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 5 | 7 | 6 | 7 | 7 | 7 | 7 | |
| 8 | 3 | 2 | 2 | 3 | 2 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 2 | 2 | 3 | 2 | 6 | 2 | 3 | 6 | 2 | 0 | 3 | 2 | |
| 1 | 1 | 9 | 9 | 0 | 9 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 4 | 9 | 0 | 3 | 5 | 1 | 0 | 6 | 9 | 1 | 0 | 9 | |
| <hr/> | | | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | |

| | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Blood Vessel Mineralization | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Cardiomyopathy | | | | | 2 | | | | | | | | | | | | | | | | | | | |
| Thrombosis | | | | | | | | | | | | | | | | | | | | | | 4 | | |
| Artery, Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Myocardium, Mineralization | | | | | | | | | | | | | | | | | | | | | | | | |
| Valve, Inflammation | | | | | | | | | | | | | | | | | | | | | | | | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Subcapsular, Hyperplasia | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | |
| Parathyroid Gland | + | + | + | + | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | |
| Pars Distalis, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract

CAS Number: 9000-38-8

Date Report Requested: 04/13/2009

Time Report Requested: 10:57:30

First Dose M/F: 08/24/04 / 08/23/04

Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 5 | 7 | 6 | 7 | 7 | 7 | 7 |
| 8 | 3 | 2 | 2 | 3 | 2 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 2 | 2 | 3 | 2 | 6 | 2 | 3 | 6 | 2 | 0 | 3 | 2 | | | |
| 1 | 1 | 9 | 9 | 0 | 9 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 4 | 9 | 0 | 3 | 5 | 1 | 0 | 6 | 9 | 1 | 0 | 9 | | |
| B6C3F1 MICE FEMALE
0.25 G/KG | ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 |
| | | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 |

females (cont...)

Thyroid Gland Cyst

+ +

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

Clitoral Gland

+ +

Ovary
Angiectasis
Cyst
Thrombosis

+
1 1 1 1 1

Uterus
Inflammation, Suppurative
Metaplasia, Squamous
Endometrium, Hyperplasia, Cystic

+
1 3 3 1 1 2 3 1 1 2 2 1 2 1 2 2 1 3 1 1 2 1 1

HEMATOPOIETIC SYSTEM

Bone Marrow
Fibrosis

+
2

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
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TDMS No. 20007 - 06
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Kava kava extract
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 Lab: BAT

| DAY ON TEST | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | |
| 8 | 3 | 2 | 2 | 3 | 2 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 2 | 2 | 3 | 2 | 6 | 2 | 3 | 6 | 2 | 0 | 3 | 2 |
| 1 | 1 | 9 | 9 | 0 | 9 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 4 | 9 | 0 | 3 | 5 | 1 | 0 | 6 | 9 | 1 | 0 | 9 |

| B6C3F1 MICE FEMALE
0.25 G/KG | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | females
(cont...) |
|---------------------------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | |
| 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | |
| 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Lymph Node
Mediastinal, Hyperplasia, Plasma Cell | | | | | | | | | | | | | | | | | | | | | | | | | | + | 3 |
| Lymph Node, Mandibular
Hyperplasia, Lymphoid | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | |
| Lymph Node, Mesenteric
Hyperplasia, Plasma Cell | | | | | | | | | | | | | | | | | | | | | | | | | | + | 3 |
| Spleen
Hematopoietic Cell Proliferation
Hyperplasia, Lymphoid
Lymphoid Follicle, Atrophy | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | |
| Thymus
Atrophy | 4 | 2 | 1 | 1 | 1 | 2 | 1 | 3 | 1 | 2 | 4 | 1 | 2 | 3 | 1 | 1 | 2 | 4 | 4 | 2 | 4 | 1 | 3 | 1 | + | | |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Mammary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Skin
Ulcer
Subcutaneous Tissue, Fibrosis | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |

MUSCULOSKELETAL SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 86

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
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| DAY ON TEST | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | females
(cont...) |
|-------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 5 | 7 | 6 | 7 | 7 | 7 | 0 |
| 8 | 3 | 2 | 2 | 3 | 2 | 3 | 1 | 3 | 3 | 3 | 3 | 3 | 3 | 2 | 2 | 3 | 2 | 6 | 2 | 3 | 6 | 2 | 0 | 3 | 2 | 0 |
| 1 | 1 | 9 | 9 | 0 | 9 | 1 | 1 | 1 | 1 | 1 | 1 | 0 | 0 | 4 | 9 | 0 | 3 | 5 | 1 | 0 | 6 | 9 | 1 | 0 | 9 | 0 |

| B6C3F1 MICE FEMALE
0.25 G/KG | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | females
(cont...) |
|---------------------------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 7 |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Fibrosis | | | | 2 | | | | | | | | | | | | | | | | | | | | | | |
| Joint, Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Skeletal Muscle | | | | | | | | | | | | | | | | | | | | | | | | | | |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--|

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Inflammation, Acute | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Metaplasia, Osseous | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Alveolus, Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Serosa, Fibrosis | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | 1 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
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 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
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 Page 87

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | |
|---|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---------------|-----------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 7 | 7 | 7 | 7 | 6 | 6 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 6 | 3 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | |
| | 3 | 0 | 3 | 1 | 9 | 6 | 3 | 3 | 3 | 2 | 7 | 2 | 3 | 6 | 7 | 3 | 3 | 2 | 2 | 2 | 2 | 2 | 3 | 3 | |
| | 0 | 8 | 0 | 5 | 7 | 9 | 1 | 1 | 0 | 9 | 5 | 9 | 1 | 9 | 2 | 0 | 0 | 9 | 9 | 9 | 9 | 9 | 0 | 0 | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 3 | |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | * TOTALS |
| Necrosis | | | | 2 | | 1 | | | | | | | | 1 | | | | | | | | | | | 4 1.3 |
| Tension Lipidosis | | | | | | | | X | | X | | | | X | | | | | | | | | | | 8 |
| Centrilobular, Hypertrophy | | | | 1 | | | | | | | | | 1 | 1 | | | 1 | 1 | | 1 | 1 | | 1 | 20 1.0 | |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | |
| Fat, Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 10 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 10 2.9 |
| Pancreas | | | | | | | | | | | | | | | | | | | | | | | | | |
| Duct, Cyst | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 4.0 |
| Salivary Glands | | | | | | | | | | | | | | | | | | | | | | | | | |
| Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 49 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 33 1.1 |
| Stomach, Forestomach | | | | | | | | | | | | | | | | | | | | | | | | | |
| Erosion | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | |
| Ulcer | | | | | | | | | | | | | | | | | | | | | | | | | |
| Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 6 1.5 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 2 2.5 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 6 1.3 |
| Stomach, Glandular | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | |
| Glands, Ectasia | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |
| Tooth | | | | | | | | | | | | | | | | | | | | | | | | | |
| Dysplasia | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |

CARDIOVASCULAR SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
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Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|---------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|
| | 7 7 7 7 6 6 7 7 7 7 5 7 7 6 3 7 7 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0.25 G/KG | 3 0 3 1 9 6 3 3 3 2 7 2 3 6 7 3 3 2 2 2 2 3 3 1 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 8 0 5 7 9 1 1 0 9 5 9 1 9 2 0 0 9 9 9 9 0 0 8 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-------|
| Blood Vessel Mineralization | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 1 2.0 |
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 1 2.0 |
| Cardiomyopathy | | | | | | | | | | | | | | | | | | | | | | | | | | 1 4.0 |
| Thrombosis | | | | | | | | | | | | | | | | | | | | | | | | | | 4 1.0 |
| Artery, Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | 1 | 1 | | | | | | 2 2.0 |
| Myocardium, Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | | 2 4.0 |
| Valve, Inflammation | | | | | | | | | | | | | 4 | | | | | | | | | | | | | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|--------|-------|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 50 2.0 | |
| Subcapsular, Hyperplasia | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | | | |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 4 1.0 | |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | 1 | | | | |
| Parathyroid Gland | M | + | + | + | M | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | M | + | 44 | 1 1.0 |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | 1 1.0 | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | | 4 1.3 |
| Pars Distalis, Hyperplasia | 1 | | | | | | | | | | | | 2 | | | | | | | | | 1 | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|---|
| | 7 7 7 7 6 6 7 7 7 7 5 7 7 6 3 7 7 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 0 3 1 9 6 3 3 3 2 7 2 3 6 7 3 3 2 2 2 2 3 3 1 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 8 0 5 7 9 1 1 0 9 5 9 1 9 2 0 0 9 9 9 9 0 0 8 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 3 | |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |
| | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|----|-----|-----|
| Lymph Node
Mediastinal, Hyperplasia, Plasma Cell | | | | | | | | | | | | | | | | | | | | | | | | + | 3 | 1 | 3.0 |
| Lymph Node, Mandibular
Hyperplasia, Lymphoid | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | |
| | 2 | | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | 3 | M | 4 | 2.0 |
| Lymph Node, Mesenteric
Hyperplasia, Plasma Cell | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | 50 | 1 | 3.0 | |
| Spleen
Hematopoietic Cell Proliferation | + | A | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | |
| | 2 | | | 3 | 2 | | | | | | | | | | | | | | | | | | | | | | |
| Spleen
Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | | 24 | 1.8 |
| Spleen
Lymphoid Follicle, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | | 6 | 1.8 |
| | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2.0 |
| Thymus
Atrophy | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | | |
| | 1 | | 4 | 1 | 2 | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | 50 | 44 | 2.1 | |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|---|-----|
| Mammary Gland | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | |
| | | | | | | | | | | | | | | | | | | | | | | | | 49 | | |
| Skin
Ulcer | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | | |
| | | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Skin
Subcutaneous Tissue, Fibrosis | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 4.0 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 2.0 |

MUSCULOSKELETAL SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----------------|--------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 7 | 7 | 7 | 7 | 6 | 6 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 6 | 3 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | | |
| | 3 | 0 | 3 | 1 | 9 | 6 | 3 | 3 | 3 | 2 | 7 | 2 | 3 | 6 | 7 | 3 | 3 | 2 | 2 | 2 | 2 | 2 | 3 | 3 | | |
| | 0 | 8 | 0 | 5 | 7 | 9 | 1 | 1 | 0 | 9 | 5 | 9 | 1 | 9 | 2 | 0 | 0 | 9 | 9 | 9 | 9 | 9 | 0 | 0 | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 0.25 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 3 | | |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 | | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | * TOTALS | |
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | |
| Fibrosis | | | | | | | | | | | | | 2 | | | 2 | | 1 | | 2 | | | 1 | 3 | 2 | 8 1.9 |
| Joint, Inflammation, Chronic | | | | | | | | | | 1 | | | | | | | | | | | | | | | 1 1.0 | |
| Skeletal Muscle | | | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| NERVOUS SYSTEM | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| RESPIRATORY SYSTEM | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Inflammation, Acute | | 2 | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | |
| Metaplasia, Osseous | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Mineralization | | 1 | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Alveolus, Infiltration Cellular, Histiocyte | | | 2 | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | |
| Serosa, Fibrosis | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Inflammation, Chronic | | | | | | | | | | | | | 1 | | | | | | | | | | | | 2 1.0 | |
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 94

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|-------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|---|------------------------------|
| | 0730 | 0753 | 0772 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | 0777 | | 0777 |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | females
(cont...) |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| ANIMAL ID | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 3 | 4 | |

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Esophagus | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Gallbladder | + | + | + | + | + | + | + | + | + | + | + | M | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Small, Duodenum
Epithelium, Hyperplasia | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Small, Ileum
Inflammation, Chronic Active
Ulcer | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Intestine Small, Jejunum
Peyer's Patch, Hyperplasia | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Liver
Angiectasis
Basophilic Focus
Clear Cell Focus
Cyst | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | females
(cont...) |
|---|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|------------------|---|----------------------|
| | 0
7
3
0 | 0
5
3
4 | 0
7
2
9 | 0
7
3
0 | 0
7
3
0 | 0
7
2
9 | 0
7
3
0 | 0
7
3
1 | 0
6
3
8 | 0
7
3
0 | 0
7
3
9 | 0
7
2
9 | 0
7
2
9 | 0
7
3
1 | 0
7
2
9 | 0
7
3
0 | 0
7
3
1 | 0
7
3
1 | 0
7
3
0 | 0
7
3
0 | 0
7
2
9 | 0
7
2
9 | 0
7
3
0 | 0
7
3
0 | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | |
| Eosinophilic Focus | | | | X | X | | | | | | | | X | X | X | X | | | X | | | | X | X | X | |
| Fatty Change | 2 | 1 | 2 | | 2 | 2 | 1 | 2 | 1 | 1 | 2 | | 2 | 1 | 1 | 1 | 1 | | 1 | 1 | | 1 | 1 | 1 | 1 | |
| Hematopoietic Cell Proliferation | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| Mixed Cell Focus | X | | | | | | | X | X | | | | | X | | X | X | | X | X | | X | | X | X | |
| Necrosis | | | | 1 | | | | | | | | | | | | | | | | | | | | 1 | 1 | |
| Tension Lipidosis | | | X | | | | | | | | | | X | | | | | X | X | | X | | | | | |
| Centrilobular, Hypertrophy | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Fat, Necrosis | | | | + | | | + | | | | | | | | | | | | | | | | + | | | |
| Pancreas | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Acinus, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Acinus, Vacuolization Cytoplasmic | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Duct, Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | | 1 | 1 | 1 | 1 | | 1 | | | 1 | | 1 | 1 | 1 | | |
| Duct, Submandibular Gland, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | 2 | | | |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Erosion | | | | | | | | | | | | | | | | | | | | | | | | | 3 | |
| Inflammation, Chronic | | | | 1 | | | | | | | | | | | 1 | | | | | | | 3 | | 3 | | |
| Ulcer | | | | | | | | | | | | | | | | | | | | | | | 3 | | | |
| Epithelium, Hyperplasia | | | | 1 | | | | | | | | | | | 1 | | | 2 | | | | 3 | | 3 | 2 | |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 1 | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|-------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|
| | 0730 | 0733 | 0732 | 0733 | 0733 | 0732 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 | 0733 |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| ANIMAL ID | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

females (cont...)

| | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|--|--|--|--|--|--|--|--|--|--|---|--|--|--|--|--|--|--|--|--|--|---|
| Glands, Ectasia | | | | | | | | | | | 1 | | | | | | | | | | | 1 |
| Tooth Dysplasia | | | | | | | | | | | | | | | | | | | | | | + |
| | | | | | | | | | | | | | | | | | | | | | | 1 |

CARDIOVASCULAR SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Blood Vessel | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Heart | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Cardiomyopathy | | | | | | | | | | | | | | | | | | | | | | | | | |
| Artery, Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Adrenal Cortex | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Vacuolization Cytoplasmic Subcapsular, Hyperplasia | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Adrenal Medulla | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | 2 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Islets, Pancreatic | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Parathyroid Gland | + + + + + + + M + + + + + + + + + + + + + + + + + | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract

CAS Number: 9000-38-8

Date Report Requested: 04/13/2009

Time Report Requested: 10:57:30

First Dose M/F: 08/24/04 / 08/23/04

Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|--------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 5 7 7 7 7 7 7 6 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0.5 G/KG | 3 3 2 3 3 2 3 3 3 3 2 2 3 2 3 3 2 3 3 3 3 2 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 4 9 0 0 9 0 1 8 0 9 9 1 9 0 0 9 1 1 0 0 9 9 0 9 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 0 0 0 0 0 0 0 0 0 1 1 1 1 1 1 1 1 1 1 2 2 2 2 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |

females (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Pituitary Gland | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Angiectasis | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Cyst | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Pars Distalis, Hyperplasia | 1 3 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Thyroid Gland | + | | | | | | | | | | | | | | | | | | | | | | | | |

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Clitoral Gland | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Ovary | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Angiectasis | | | | | | | | | | | | | | | | | | | | | | | | | |
| Cyst | 1 1 2 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Thrombosis | 3 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Uterus | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Endometrium, Hyperplasia, Cystic | 2 2 1 2 1 3 2 3 1 2 3 2 3 2 1 3 1 2 2 2 | | | | | | | | | | | | | | | | | | | | | | | | |

HEMATOPOIETIC SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Bone Marrow | + | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
BLANK .. Not examined microscopically
1-4 .. Lesion qualified as:
1) Minimal 3) Moderate
2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
|--------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|-------------------|
| | 7 5 7 7 7 7 7 7 6 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0.5 G/KG | 3 3 2 3 3 2 3 3 3 3 2 2 3 2 3 3 2 3 3 3 3 2 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 4 9 0 0 9 0 1 8 0 9 9 1 9 0 0 9 1 1 0 0 9 9 0 9 | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | | |
| 0.5 G/KG | 3 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 0 0 0 0 0 0 0 0 0 1 1 1 1 1 1 1 1 1 1 2 2 2 2 2 | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 2 3 4 5 6 7 8 9 0 1 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | females (cont...) |

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|
| Fibrosis | 2 | | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mandibular
Hyperplasia, Lymphoid | + | | | | | | | | | | | | | | | | | | | | | | | | | 2 |
| Lymph Node, Mesenteric | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Spleen
Hematopoietic Cell Proliferation
Hyperplasia, Lymphoid
Lymphoid Follicle, Atrophy | + | | | | | | | | | | | | | | | | | | | | | | | | | 2 1 1 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 2 2 1 1 2 2 |
| Thymus
Atrophy
Hyperplasia, Histiocytic | + | | | | | | | | | | | | | | | | | | | | | | | | | 2 2 2 2 2 1 1 3 1 2 1 1 2 2 1 2 2 2 1 2 2 2 2 2 1 |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|-------------|
| Mammary Gland | + | | | | | | | | | | | | | | | | | | | | | | | | | |
| Skin
Fibrosis
Inflammation, Chronic
Subcutaneous Tissue, Necrosis | + | | | | | | | | | | | | | | | | | | | | | | | | | 3
3
3 |

MUSCULOSKELETAL SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|-------------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|------|---|
| | 0730 | 0733 | 0739 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0736 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | 0737 | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| ANIMAL ID | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

females
(cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Bone Fibrosis | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| | 2 | | 1 | | | | | | | | | | | | | | | | | | | | | | |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Lung Inflammation | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| | | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Nose Glands, Dilatation, Nasolacrimal Duct, Inflammation, Suppurative | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| | | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
|---------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Eye | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
|-----|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Harderian Gland Atrophy Hyperplasia | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
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Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|--------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 5 7 7 7 7 7 7 6 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0.5 G/KG | 3 3 2 3 3 2 3 3 3 3 2 2 3 2 3 3 2 3 3 3 2 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 4 9 0 0 9 0 1 8 0 9 9 1 9 0 0 9 1 1 0 0 9 9 0 9 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| females (cont...) | 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | |

Infiltration Cellular, Mononuclear Cell 1

URINARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Kidney | + | | | | | | | | | | | | | | | | | | | | | | | |
| Amyloid Deposition | | | | | | | | | | | | | | | | | | | | | | | | |
| Metaplasia, Osseous | | | | | | | | | | | | | | | | | | | | | | | | |
| Mineralization | 1 | | | | | | | | | | | | | | | | | | | | | | | |
| Nephropathy | 1 1 2 1 1 2 1 1 1 1 2 1 1 1 2 1 1 1 1 1 2 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | |
| Papilla, Necrosis | | | | | | | | | | | | | | | | | | | | | | | | |
| Pelvis, Cyst | 2 | | | | | | | | | | | | | | | | | | | | | | | |
| Renal Tubule, Hyperplasia | 1 | | | | | | | | | | | | | | | | | | | | | | | |
| Urinary Bladder | + | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
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 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 102

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 7 7 7 7 7 5 7 7 7 7 7 7 7 7 7 7 7 7 7 5 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 2 2 2 3 2 5 2 3 3 2 2 2 3 2 3 2 2 3 2 5 1 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | |
| | 9 9 9 9 0 9 4 9 0 1 9 9 9 1 9 1 9 9 0 9 6 2 9 1 9 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 4 4 4 4 4 5 | | | | | | | | | | | | | | | | | | | | | | | |
| | * TOTALS | | | | | | | | | | | | | | | | | | | | | | | |

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----|----|---|-----|---|-----|---|-----|---|---|-----|
| Esophagus | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | | | | | | | | | |
| Gallbladder | + | | | | | | | | | | | | | | | | | | | | | | | | 49 | | | | | | | | | | |
| Intestine Large, Cecum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | | | | | | | | | |
| Intestine Large, Colon | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | | | | | | | | | |
| Intestine Large, Rectum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | | | | | | | | | |
| Intestine Small, Duodenum
Epithelium, Hyperplasia | + | | | | | | | | | | | | | | | | | | | | | | | | 2 | 50 | 1 | 2.0 | | | | | | | |
| Intestine Small, Ileum
Inflammation, Chronic Active
Ulcer | + | | | | | | | | | | | | | | | | | | | | | | | | | 50 | 1 | 2.0 | 1 | 1.0 | | | | | |
| Intestine Small, Jejunum
Peyer's Patch, Hyperplasia | + | | | | | | | | | | | | | | | | | | | | | | | | 4 | 50 | 1 | 4.0 | | | | | | | |
| Liver
Angiectasis
Basophilic Focus
Clear Cell Focus
Cyst | + | | | | | | | | | | | | | | | | | | | | | | | | | 50 | | | | 1 | 2 | 1.0 | 4 | 1 | 1.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
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Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|---|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | |
| | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | |
| Eosinophilic Focus | | | | X | | | X | X | X | | X | | | | | | | | | X | | | | | |
| Fatty Change | 1 | | 1 | | | 1 | | 1 | 1 | 1 | 1 | 2 | 2 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | |
| Hematopoietic Cell Proliferation | 1 | 2 | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | 2 | 1 | 1 | 1 | 2 | | 1 | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| Mixed Cell Focus | X | | | | | | | | | | | X | | X | | | | | | | | X | | | |
| Necrosis | | 1 | | | | | | | | | | | | | | | | | | | | | 2 | | |
| Tension Lipidosis | | | | | X | | | | | | | | | X | | | | | | | | | | | |
| Centrilobular, Hypertrophy | 2 | 2 | | 1 | 2 | 1 | 2 | 1 | 2 | 1 | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 2 | | 2 | 2 | |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | |
| Fat, Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | |
| Pancreas | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Acinus, Atrophy | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Acinus, Vacuolization Cytoplasmic | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Duct, Cyst | 4 | | | | | | | | | | | | | | | | | | | | | | | | |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | 1 | 1 | | 1 | | | 1 | 1 | 2 | | | 2 | | | | 1 | 1 | | 1 | 1 | |
| Duct, Submandibular Gland, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Erosion | | 3 | | | | | 2 | 3 | 4 | 3 | | | | | 4 | | 3 | 3 | 4 | 3 | 2 | | 4 | 3 | |
| Inflammation, Chronic | | 3 | | | 3 | 3 | 2 | 3 | 3 | 3 | | | | | 3 | 1 | 3 | 4 | 3 | 3 | 2 | 1 | 4 | 3 | |
| Ulcer | | | | | 3 | 3 | | | | | | | | | | | | | | | | | | | |
| Epithelium, Hyperplasia | | 3 | | | 4 | 3 | 3 | 3 | 3 | 4 | | | | | 4 | 1 | 3 | 4 | 4 | 3 | 1 | 2 | 4 | 3 | |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|
| | 7 7 7 7 7 7 5 7 7 7 7 7 7 7 7 7 7 7 7 7 5 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 2 2 2 3 2 5 2 3 3 2 2 2 3 2 3 2 2 3 2 5 1 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | |
| | 9 9 9 9 0 9 4 9 0 1 9 9 9 1 9 1 9 9 0 9 6 2 9 1 9 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 |
| | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|--|--|--|--|--|--|--------------|--------------|
| Glands, Ectasia | | | | | | | | | | | | | | | | | | | | | | | | 2 1.0 | |
| Tooth Dysplasia | | | | | | | | | | | | | | | | | + | | | | | | | 2 | 2 1.0 |

CARDIOVASCULAR SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|---|--|--|--|--|--|--|--------------|-----------|-----|--|--------------|
| Blood Vessel | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | | |
| Heart | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | | | |
| Cardiomyopathy | 1 | | 1 | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1 | | 4 1.0 |
| Artery, Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | 1 | | | | | | | 3 1.0 | | | | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---------------|-----------|
| Adrenal Cortex | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Vacuolization Cytoplasmic Subcapsular, Hyperplasia | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 2 | 2 | 2 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 50 2.0 | |
| Adrenal Medulla | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | 2 2.0 | |
| Islets, Pancreatic | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | |
| Parathyroid Gland | M | + | | | | | | | | | | | | | | | | | | | | | | | 48 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 105

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
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 Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|--|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | | |
| | 2 | 2 | 2 | 2 | 3 | 2 | 5 | 2 | 3 | 3 | 2 | 2 | 2 | 3 | 2 | 3 | 2 | 2 | 3 | 2 | 5 | 1 | 2 | 3 | 2 | |
| | 9 | 9 | 9 | 9 | 0 | 9 | 4 | 9 | 0 | 1 | 9 | 9 | 9 | 1 | 9 | 1 | 9 | 9 | 0 | 9 | 6 | 2 | 9 | 1 | 9 | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | | |
| | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | * TOTALS | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|--------------|
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | M | + | 49 | |
| Angiectasis | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Cyst | | | | | | | | | | | | | | | | | 1 | | | | | | | | | | 2 1.0 |
| Pars Distalis, Hyperplasia | | | 1 | 1 | | | | | | | | | | | | | | | | | 1 | | | | | | 7 1.3 |
| Thyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |

GENERAL BODY SYSTEM

NONE

GENITAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---------------|
| Clitoral Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Ovary | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Angiectasis | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 2.0 |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | | 6 1.3 |
| Thrombosis | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 3.5 |
| Uterus | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Endometrium, Hyperplasia, Cystic | 1 | 2 | 3 | 2 | 3 | 2 | 2 | 1 | 2 | | 2 | 1 | 3 | 4 | 3 | 2 | 3 | 3 | 3 | 1 | 2 | | | | 1 | 4 | 42 2.1 |

HEMATOPOIETIC SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|
| Bone Marrow | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 7 7 7 7 7 5 7 7 7 7 7 7 7 7 7 7 7 5 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 2 2 2 3 2 5 2 3 3 2 2 2 3 2 3 2 2 3 2 5 1 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | |
| | 9 9 9 9 0 9 4 9 0 1 9 9 9 1 9 1 9 9 0 9 6 2 9 1 9 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| 0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | * TOTALS | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---------------|
| Fibrosis | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Lymph Node, Mandibular
Hyperplasia, Lymphoid | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| | 2 | | | | | | | | | | | | | | | | | | | | | | | | 4 2.0 |
| Lymph Node, Mesenteric | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Spleen
Hematopoietic Cell Proliferation
Hyperplasia, Lymphoid
Lymphoid Follicle, Atrophy | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| | 2 4 1 1 4 1 2 1 1 1 1 M 2 1 1 2 1 3 4 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 20 1.5 |
| | 3 | | | | | | | | | | | | | | | | | | | | | | | | 6 2.0 |
| | 4 2 2 | | | | | | | | | | | | | | | | | | | | | | | | 4 2.5 |
| Thymus
Atrophy
Hyperplasia, Histiocytic | + | | | | | | | | | | | | | | | | | | | | | | | | 49 |
| | 4 2 2 1 1 1 4 1 2 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 46 1.7 |
| | 2 | | | | | | | | | | | | | | | | | | | | | | | | 2 2.0 |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------------|
| Mammary Gland | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Skin
Fibrosis
Inflammation, Chronic
Subcutaneous Tissue, Necrosis | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |

MUSCULOSKELETAL SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
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TDMS No. 20007 - 06
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Kava kava extract
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 Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|--|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | | |
| | 2 | 2 | 2 | 2 | 3 | 2 | 5 | 2 | 3 | 3 | 2 | 2 | 2 | 3 | 2 | 3 | 2 | 2 | 3 | 2 | 5 | 1 | 2 | 3 | 2 | |
| | 9 | 9 | 9 | 9 | 0 | 9 | 4 | 9 | 0 | 1 | 9 | 9 | 9 | 1 | 9 | 1 | 9 | 9 | 0 | 9 | 6 | 2 | 9 | 1 | 9 | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| 0.5 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | | |
| | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 5 | | |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | * TOTALS | |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------------|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Fibrosis | | | 1 | | | | | | | | 3 | | | | 1 | | | | | | | | | | 7 1.6 |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------------|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation | | | | | | | | | | | | | | | | | | | | | 1 | | | | 2 1.0 |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------------|
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Glands, Dilatation | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Nasolacrimal Duct, Inflammation, Suppurative | | | | | | | 2 | | | | | | | | | | | | | | | | | | 1 2.0 |

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
|---------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|
| Eye | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
|-----|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------|

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|--------------|
| Harderian Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Atrophy | | | | | | | | | | | | | | | 1 | | | | | | | | | | 1 1.0 |
| Hyperplasia | | | 1 | | 2 | | | | | | | | | | | | | | | | 1 | | | | 5 1.6 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
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 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL
 Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---------------|
| | 7 7 7 7 7 7 5 7 7 7 7 7 7 7 7 7 7 7 7 7 5 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 2 2 2 3 2 5 2 3 3 2 2 2 3 2 3 2 2 3 2 5 1 2 3 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 9 9 9 9 0 9 4 9 0 1 9 9 9 1 9 1 9 9 0 9 6 2 9 1 9 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| 0.5 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 2 2 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | 42 1.0 |

URINARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---------------|
| Kidney | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Amyloid Deposition | | | | | | | | | | | | | | | | | | | | | | | | | 2 1.0 |
| Metaplasia, Osseous | 1 | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 |
| Mineralization | 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 7 1.0 |
| Nephropathy | 2 1 1 1 1 1 4 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 26 1.3 |
| Papilla, Necrosis | 2 | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Pelvis, Cyst | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |
| Renal Tubule, Hyperplasia | 1 | | | | | | | | | | | | | | | | | | | | | | | | 2 1.0 |
| Urinary Bladder | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | 40 1.0 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
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 Species/Strain: MICE/B6C3F1

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|--|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 7 6 7 6 7 7 6 5 7 7 7 5 7 7 7 7 7 5 7 7 6 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 5 3 7 2 3 8 2 3 3 3 7 3 3 3 3 3 4 2 2 0 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 4 0 8 9 0 4 3 1 0 1 6 1 0 0 1 1 1 8 9 9 4 3 9 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
1.0 G/KG | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 5 5 5 5 5 5 5 6 6 6 6 6 6 6 6 6 6 7 7 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 | | | | | | | | | | | | | | | | | | | | | | | | |

females
(cont...)

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Esophagus | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Gallbladder
Cyst
Epithelium, Cytoplasmic Alteration | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Large, Cecum | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Large, Colon | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Large, Rectum | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Small, Duodenum
Metaplasia, Squamous | +
1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Small, Ileum | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Intestine Small, Jejunum
Epithelium, Hyperplasia | +
3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Liver
Basophilic Focus
Clear Cell Focus
Eosinophilic Focus
Fatty Change | +
X
X
X X X X X X X X
2 4 2 2 1 1 1 2 1 2 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
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|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 6 7 6 7 7 6 5 7 7 7 5 7 7 7 7 7 5 7 7 6 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 5 3 7 2 3 8 2 3 3 3 7 3 3 3 3 3 4 2 2 0 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 4 0 8 9 0 4 3 1 0 1 6 1 0 0 1 1 1 8 9 9 4 3 9 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 1.0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | 7 |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

females (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Hematopoietic Cell Proliferation | | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | | | 1 | 1 | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | 1 | 2 | 1 | 1 | 1 | 1 | 1 | | | | 1 | 1 | |
| Mixed Cell Focus | X | | | | | | | | | | X | | | | | | | X | X | | | | | | |
| Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | |
| Tension Lipidosis | | | | | | | | | | | X | | | | | | | X | | | | | | | X |
| Vacuolization Cytoplasmic | | | | | | | | | | | | | | | | | | | | | | 2 | | | |
| Centrilobular, Hypertrophy | | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| Mesentery | | | + | | | | | | | | | | | | | | | | | | | | | | |
| Fat, Necrosis | | | 2 | | | | | | | | | | | | | | | | | | | | | | |
| Pancreas | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | |
| Acinus, Hypertrophy | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Duct, Cyst | | | | | | | | | | | | | | | | | | | | | | | | | |
| Salivary Glands | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Infiltration Cellular, Mononuclear Cell | 1 | | 1 | 1 | | | | | | | | | | 1 | | 1 | | 1 | | 1 | | | 1 | 1 | 1 |
| Parotid Gland, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Erosion | | | | | | 4 | | 3 | 3 | 2 | | 3 | 2 | | 3 | | 3 | 2 | | | | | | 2 | |
| Inflammation, Chronic | | 2 | | | 2 | 3 | 3 | 3 | 3 | 2 | 3 | 3 | 3 | 1 | 3 | | 3 | 3 | 3 | | | | | 3 | |
| Ulcer | | 2 | | | 2 | | 3 | | | | 3 | | | | | | | | 2 | | | | | | |
| Epithelium, Hyperplasia | | 1 | | | 3 | 3 | 3 | 3 | 3 | 1 | 3 | 3 | 3 | 1 | 3 | | 3 | 3 | 3 | | | | | 3 | |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | 2 | | | | | | | | | | | |
| Ulcer | | 2 | | | | | | | 2 | | | | | | | | | | | | | | | | |

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|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 7 6 7 6 7 7 6 5 7 7 7 5 7 7 7 7 7 5 7 7 6 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 2 5 3 7 2 3 8 2 3 3 3 7 3 3 3 3 3 4 2 2 0 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 4 0 8 9 0 4 3 1 0 1 6 1 0 0 1 1 1 8 9 9 4 3 9 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 1.0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

females (cont...)

| | | |
|-------------------------|---|---|
| Epithelium, Hyperplasia | 2 | 2 |
| Tooth Dysplasia | + | 1 |

CARDIOVASCULAR SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Blood Vessel | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Cardiomyopathy | | | | | | | | | | | | | | | | | | | | | | | | |
| Myocardium, Mineralization | 2 | | | | | | | | | | | | | | | | | | | | | | | |

ENDOCRINE SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|-------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Subcapsular, Hyperplasia | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 2 | 2 |
| Zona Reticularis, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Hyperplasia | | | | | | | | | | | 2 | | | | | | | | | | 1 | | | |
| Vacuolization Cytoplasmic | 1 | | | | | | | | | | | | | | | | | | | | | | | |
| Islets, Pancreatic | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Parathyroid Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
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 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
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| DAY ON TEST | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | females (cont...) |
|-------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 7 | 6 | 7 | 6 | 7 | 7 | 6 | 5 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 6 | 7 | 7 | 7 | 0 | |
| 2 | 5 | 3 | 7 | 2 | 3 | 8 | 2 | 3 | 3 | 3 | 7 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 2 | 2 | 0 | 2 | 2 | 3 | 0 | |
| 5 | 4 | 0 | 8 | 9 | 0 | 4 | 3 | 1 | 0 | 1 | 6 | 1 | 0 | 0 | 1 | 1 | 1 | 8 | 9 | 9 | 4 | 3 | 9 | 1 | 0 | |

| B6C3F1 MICE FEMALE
1.0 G/KG | | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | females (cont...) |
|--------------------------------|--|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------------------|
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| Pituitary Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Pars Distalis, Hyperplasia | | | | | | | | | | | 2 | | | | | | | | | | 2 | 1 | | | | | |
| Thyroid Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | |

GENERAL BODY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|
| Tissue NOS | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | 3 |

GENITAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------------------------------|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Clitoral Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Ovary | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Angiectasis | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Uterus | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Endometrium, Hyperplasia, Cystic | | 2 | | 2 | 1 | 1 | 2 | 1 | | 3 | | 2 | 3 | | 1 | 3 | 1 | 2 | 1 | 1 | | 1 | 2 | 2 | 3 | |
| Vagina | | | | | | | | | | | | | | | | | | | | | | | | | | + |
| Vacuolization Cytoplasmic | | | | | | | | | | | | | | | | | | | | | | | | | | 2 |

HEMATOPOIETIC SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
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Page 113

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|--------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|---|
| | 7 6 7 6 7 7 6 5 7 7 7 5 7 7 7 7 7 5 7 7 6 7 7 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
1.0 G/KG | 2 5 3 7 2 3 8 2 3 3 3 7 3 3 3 3 3 4 2 2 0 2 2 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 4 0 8 9 0 4 3 1 0 1 6 1 0 0 1 1 1 8 9 9 4 3 9 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| 1.0 G/KG | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 5 5 5 5 5 5 5 5 5 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 |

females (cont...)

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Bone Marrow
Fibrosis
Necrosis | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mandibular
Hyperplasia, Lymphoid | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Lymph Node, Mesenteric
Hyperplasia, Lymphoid
Inflammation, Granulomatous | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Spleen
Hematopoietic Cell Proliferation
Hyperplasia, Lymphoid
Lymphoid Follicle, Atrophy | A +
1 2 2 1 2 2
4 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| Thymus
Atrophy
Hyperplasia, Histiocytic
Infiltration Cellular, Histiocyte | + M + + + + M + + + + + M + + + + + + + + + + + + +
4 4 4 4 1 2 2 1 2 1 2 2 2 3 1 4 4 1 1 | | | | | | | | | | | | | | | | | | | | | | | | |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| Mammary Gland | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Skin | + | | | | | | | | | | | | | | | | | | | | | | | | |

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 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| | DAY ON TEST | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 7 | 6 | 7 | 6 | 7 | 7 | 6 | 5 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 6 | 7 | 7 | 7 | |
| | 2 | 5 | 3 | 7 | 2 | 3 | 8 | 2 | 3 | 3 | 3 | 7 | 3 | 3 | 3 | 3 | 3 | 4 | 2 | 2 | 0 | 2 | 2 | 3 | |
| | 5 | 4 | 0 | 8 | 9 | 0 | 4 | 3 | 1 | 0 | 1 | 6 | 1 | 0 | 0 | 1 | 1 | 1 | 8 | 9 | 9 | 4 | 3 | 9 | 1 |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 1.0 G/KG | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 6 | 7 | 7 | 7 | 7 | 7 | 7 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

females
(cont...)

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Fibrosis | | | | | 1 | | | | | | | | | | 2 | | | | | | | | | |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|--|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Peripheral Nerve
Axon, Degeneration | | | | | | | | | | | | | | | | | | | | | | | | |
| Spinal Cord | | | | | | | | | | | | | | | | | | | | | | | | |

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | 2 | | |
| Alveolus, Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | 1 | | |
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |
| Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | 2 | | |
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | females
(cont...) |
|-------------|-----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------------------|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 7 | 6 | 7 | 6 | 7 | 7 | 6 | 5 | 7 | 7 | 7 | 5 | 7 | 7 | 7 | 7 | 7 | 7 | 5 | 7 | 7 | 6 | 7 | 7 | 7 | 0 | |
| 2 | 5 | 3 | 7 | 2 | 3 | 8 | 2 | 3 | 3 | 3 | 7 | 3 | 3 | 3 | 3 | 3 | 3 | 4 | 2 | 2 | 0 | 2 | 2 | 3 | 0 | |
| 5 | 4 | 0 | 8 | 9 | 0 | 4 | 3 | 1 | 0 | 1 | 6 | 1 | 0 | 0 | 1 | 1 | 1 | 8 | 9 | 9 | 4 | 3 | 9 | 1 | 0 | |

B6C3F1 MICE FEMALE
1.0 G/KG

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|---|---|---|--|--|--|---|---|--|---|---|---|---|---|--|---|---|--|--|---|---|---|--|
| Eye | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Harderian Gland | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Hyperplasia | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | 1 | 1 | 1 | | | | 1 | 1 | | 1 | 1 | 1 | 1 | 1 | | 1 | 1 | | | 1 | 1 | 1 | |

URINARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|--|---|---|---|---|---|---|---|---|---|---|---|--|---|--|--|--|---|---|---|--|
| Kidney | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Amyloid Deposition | | | | | | | | | | | | | | | | | | | | | | | | | |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | |
| Metaplasia, Osseous | 1 | | | | | | | | | | | | | | | | | | | | | | | | |
| Mineralization | 1 | | | | | | | | 1 | 1 | 1 | | | 1 | 1 | 1 | | 2 | | | | | 1 | 1 | |
| Nephropathy | | | 1 | | | 1 | 4 | 4 | 1 | 3 | 1 | | 4 | | 1 | | | 1 | | | | 3 | 2 | 1 | |
| Papilla, Necrosis | 2 | | | | | | | | | | | | | | | | | | | | | | | | |
| Urinary Bladder | + | | | | | | | | | | | | | | | | | | | | | | | | |
| Infiltration Cellular, Mononuclear Cell | | 1 | 1 | 1 | | 1 | | | | | | 1 | 1 | 1 | 1 | | | 1 | | | | 1 | 1 | 1 | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
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 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 5 7 1 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
1.0 G/KG | 4 1 4 3 3 3 3 3 3 3 3 3 2 2 3 3 3 3 3 3 2 3 2 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 8 9 8 0 0 1 0 0 1 0 0 0 9 9 1 0 0 1 1 1 9 0 9 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| * TOTALS | 3 4 | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 0 | | | | | | | | | | | | | | | | | | | | | | | |
| 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |

ALIMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|------------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----|-----|
| Esophagus | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Gallbladder | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1.0 |
| Epithelium, Cytoplasmic Alteration | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2.0 |
| Intestine Large, Cecum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Large, Colon | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Large, Rectum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Duodenum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Metaplasia, Squamous | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1.0 |
| Intestine Small, Ileum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Intestine Small, Jejunum | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 3.0 |
| Liver | + | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Basophilic Focus | | | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Clear Cell Focus | | | | | | | | | | | | | | | | | | | | | | | | | X | 5 |
| Eosinophilic Focus | X | | | | | | | | | | | | | | | | | | | | | | | | X | 26 |
| Fatty Change | 1 | | | | | | | | | | | | | | | | | | | | | | | | 23 | 1.3 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
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TDMS No. 20007 - 06

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First Dose M/F: 08/24/04 / 08/23/04

Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|-------|--------|--------|
| | 5 7 1 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
1.0 G/KG | 4 1 4 3 3 3 3 3 3 3 3 3 2 2 3 3 3 3 3 3 2 3 2 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | | | |
| | 8 9 8 0 0 1 0 0 1 0 0 0 9 9 1 0 0 1 1 1 9 0 9 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 3 4 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | | | |
| ANIMAL ID | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 9 9 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | | | |
| Hematopoietic Cell Proliferation | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 1 2.0 | |
| Infiltration Cellular, Mononuclear Cell | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 44 | 1.1 |
| Mixed Cell Focus | | | | | | | | | | | | | | | | | | | | | | | | | X | 7 | | |
| Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1.0 | | |
| Tension Lipidosis | | | | | | | | | | | | | | | | | | | | | | | | | X | 4 | | |
| Vacuolization Cytoplasmic | | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | | |
| Centrilobular, Hypertrophy | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 49 | 2.0 |
| Mesentery | | | | | | | | | | | | | | | | | | | | | | | | | | 6 | | |
| Fat, Necrosis | 2 | 2 | | | 4 | | | | | | 2 | | | | | | | | | | | | | | | 2 | 6 | 2.3 |
| Pancreas | | | | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1.0 | | |
| Acinus, Hypertrophy | | | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | | |
| Duct, Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | 3 | 1 3.0 | |
| Salivary Glands | | | | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Infiltration Cellular, Mononuclear Cell | 1 | | | 1 | 1 | 1 | | | | 1 | 1 | 1 | | | | 1 | 1 | | | 1 | 1 | 1 | | | | 1 | 26 | 1.0 |
| Parotid Gland, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 1 2.0 | | |
| Stomach, Forestomach | | | | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Erosion | | | | | | | | | | | | | | | | | | | | | | | | | | 3 | 11 2.7 | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 4 2 | 22 2.5 |
| Ulcer | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 6 2.3 | |
| Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 3 3 | 24 2.3 |
| Stomach, Glandular | | | | | | | | | | | | | | | | | | | | | | | | | | 50 | | |
| Infiltration Cellular, Mononuclear Cell | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | |
| Ulcer | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 2.0 | |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
+ .. Tissue examined microscopically
x .. Lesion present
I .. Insufficient tissue
M .. Missing tissue
A .. Autolysis precludes evaluation
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|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 5 7 1 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
1.0 G/KG | 4 1 4 3 3 3 3 3 3 3 3 3 2 2 3 3 3 3 3 3 2 3 2 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 8 9 8 0 0 1 0 0 1 0 0 0 9 9 1 0 0 1 1 1 9 0 9 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| 1.0 G/KG | 3 4 | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 0 | | | | | | | | | | | | | | | | | | | | | | | |
| 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| * TOTALS | | | | | | | | | | | | | | | | | | | | | | | | |

| | |
|-------------------------|--------------|
| Epithelium, Hyperplasia | 2 2.0 |
| Tooth Dysplasia | 2 1.0 |

CARDIOVASCULAR SYSTEM

| | |
|----------------------------|--------------|
| Blood Vessel | 50 |
| Heart | 50 |
| Cardiomyopathy | 1 1.0 |
| Myocardium, Mineralization | 1 2.0 |

ENDOCRINE SYSTEM

| | |
|-------------------------------|---------------|
| Adrenal Cortex | 50 |
| Subcapsular, Hyperplasia | 50 1.9 |
| Zona Reticularis, Hyperplasia | 1 1.0 |
| Adrenal Medulla | 50 |
| Hyperplasia | 2 1.5 |
| Vacuolization Cytoplasmic | 1 1.0 |
| Islets, Pancreatic | 50 |
| Parathyroid Gland | 48 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
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|----------------------------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|--------|
| | 5 7 1 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 6 7 7
4 1 4 3 3 3 3 3 3 3 3 3 2 2 3 3 3 3 3 2 3 2 6 3 3
8 9 8 0 0 1 0 0 1 0 0 9 9 1 0 0 1 1 1 9 0 9 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |
| | 0
3 4
7 7 7 7 8 8 8 8 8 8 8 8 8 8 9 9 9 9 9 9 9 9 9 0
6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Pituitary Gland | + | | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1.0 |
| Pars Distalis, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 5 1.8 |
| Thyroid Gland | + | | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| GENERAL BODY SYSTEM | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Tissue NOS | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Inflammation, Chronic | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 3.0 |
| GENITAL SYSTEM | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Clitoral Gland | + | | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Ovary | + | | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Angiectasis | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1 1.0 |
| Cyst | | | | | | | | | | | | | | | | | | | | | | | | | | 1 1 | 1 1.0 |
| Uterus | + | | | | | | | | | | | | | | | | | | | | | | | | | 50 | |
| Endometrium, Hyperplasia, Cystic | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 36 1.8 |
| Vagina | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Vacuolization Cytoplasmic | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 |

HEMATOPOIETIC SYSTEM

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
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 1-4 .. Lesion qualified as:
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| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS |
|--------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|----------|
| | 5 7 1 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
1.0 G/KG | 4 1 4 3 3 3 3 3 3 3 3 3 2 2 3 3 3 3 3 3 2 3 2 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 8 9 8 0 0 1 0 0 1 0 0 0 9 9 1 0 0 1 1 1 9 0 9 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| 1.0 G/KG | 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 | | | | | | | | | | | | | | | | | | | | | | | | |
| 1.0 G/KG | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| | | | | | | | | | | | | | | | | | | | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|-----------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|-----|-----|-----|
| Bone Marrow | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | | |
| Fibrosis | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1.0 | | | |
| Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2.0 | | |
| Lymph Node, Mandibular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | | |
| Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | | | 3 | 2.5 | | |
| Lymph Node, Mesenteric | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 | | | | |
| Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2.0 | | |
| Inflammation, Granulomatous | | | | | | | | | | | | | | | | | | | | | | | | | | | | 3 | 3.0 | |
| Spleen | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 49 | | | | |
| Hematopoietic Cell Proliferation | | | | | | | | | | | | | | | | | | | | | | | | | | | | 1 | 1.8 | |
| Hyperplasia, Lymphoid | | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2.0 | |
| Lymphoid Follicle, Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 3.5 | |
| Thymus | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 47 | | | | |
| Atrophy | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2.1 |
| Hyperplasia, Histiocytic | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 4 | 2.0 |
| Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | | | | | | | | 2 | 2.5 |

INTEGUMENTARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|
| Mammary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Skin | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----------------|---|
| | 5 7 1 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 4 1 4 3 3 3 3 3 3 3 3 3 2 2 3 3 3 3 3 3 2 3 2 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | | |
| | 8 9 8 0 0 1 0 0 1 0 0 9 9 1 0 0 1 1 1 9 0 9 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 1.0 G/KG | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 3 | 4 |
| | 7 | 7 | 7 | 7 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | 0 |
| | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 |
| | | | | | | | | | | | | | | | | | | | | | | | | * TOTALS | |

MUSCULOSKELETAL SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|----------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Bone | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Fibrosis | 1 | | | 1 | | | | | | 2 | | | | 2 | | | | | | | 2 | 3 | 2 | | | 9 1.8 |

NERVOUS SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|--------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|----|
| Brain | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Peripheral Nerve | + | | | | | | | | | | | | | | | | | | | | | | | | 1 | |
| Axon, Degeneration | 4 | | | | | | | | | | | | | | | | | | | | | | | | 1 4.0 | |
| Spinal Cord | + | | | | | | | | | | | | | | | | | | | | | | | | 1 | |

RESPIRATORY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|---|----|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Alveolar Epithelium, Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | | |
| Alveolus, Infiltration Cellular, Histiocyte | | | | | | | | | | | | | | | | | | | | | | | | 1 1.0 | | |
| Nose | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |
| Inflammation, Suppurative | | | | | | | | | | | | | | | | | | | | | | | | 1 2.0 | | |
| Trachea | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 50 |

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
 BLANK .. Not examined microscopically
 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
 2) Mild 4) Marked
 Page 122

TDMS No. 20007 - 06
 Test Type: CHRONIC
 Route: GAVAGE
 Species/Strain: MICE/B6C3F1

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Kava kava extract
 CAS Number: 9000-38-8

Date Report Requested: 04/13/2009
 Time Report Requested: 10:57:30
 First Dose M/F: 08/24/04 / 08/23/04
 Lab: BAT

| DAY ON TEST | 0 | | | | | | | | | | | | | | | | | | | | | | | |
|--------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | 5 7 1 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 6 7 7 | | | | | | | | | | | | | | | | | | | | | | | |
| B6C3F1 MICE FEMALE
1.0 G/KG | 4 1 4 3 3 3 3 3 3 3 3 3 2 2 3 3 3 3 3 3 2 3 2 6 3 3 | | | | | | | | | | | | | | | | | | | | | | | |
| | 8 9 8 0 0 1 0 0 1 0 0 9 9 1 0 0 1 1 1 9 0 9 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 3 4 | | | | | | | | | | | | | | | | | | | | | | | |
| | 7 7 7 7 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 0 | | | | | | | | | | | | | | | | | | | | | | | |
| | 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 | | | | | | | | | | | | | | | | | | | | | | | |
| * TOTALS | | | | | | | | | | | | | | | | | | | | | | | | |

SPECIAL SENSES SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------|
| Eye | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Harderian Gland | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Hyperplasia | | | | | | | | | | | | | | | | | | | | | | | | | 2 |
| Infiltration Cellular, Mononuclear Cell | 1 | | | | | | | | | | | | | | | | | | | | | | | | 38 1.0 |

URINARY SYSTEM

| | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--------|
| Kidney | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Amyloid Deposition | | | | | | | | | | | | | | | | | | | | | | | | | 2 |
| Inflammation | | | | | | | | | | | | | | | | | | | | | | | | | 3 |
| Metaplasia, Osseous | | | | | | | | | | | | | | | | | | | | | | | | | 2 |
| Mineralization | | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Nephropathy | 2 | | | | | | | | | | | | | | | | | | | | | | | | 1 |
| Papilla, Necrosis | | | | | | | | | | | | | | | | | | | | | | | | | 2 |
| Urinary Bladder | + | | | | | | | | | | | | | | | | | | | | | | | | 50 |
| Infiltration Cellular, Mononuclear Cell | 1 1 1 1 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | 34 1.0 |

*** END OF REPORT ***

* .. Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade
 + .. Tissue examined microscopically
 x .. Lesion present
 I .. Insufficient tissue
 M .. Missing tissue
 A .. Autolysis precludes evaluation
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 1-4 .. Lesion qualified as:
 1) Minimal 3) Moderate
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