

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

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

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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 Lab: BAT

B6C3F1 MICE MALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
<b>Disposition Summary</b>				
Animals Initially in Study	50	50	50	50
Early Deaths				
Moribund Sacrifice	3	10	10	7
Natural Death	13	7	5	7
Survivors				
Moribund Sacrifice			1	
Natural Death	1			
Terminal Sacrifice	33	33	34	36
Animals Examined Microscopically	50	50	50	50

**ALIMENTARY SYSTEM**

Gallbladder	(49)	(50)	(49)	(47)
Infiltration Cellular, Mononuclear Cell	1 (2%)		1 (2%)	1 (2%)
Inflammation, Granulomatous			1 (2%)	
Intestine Large, Cecum	(50)	(50)	(50)	(50)
Amyloid Deposition	1 (2%)			
Serosa, Inflammation, Granulomatous	1 (2%)			
Intestine Small, Duodenum	(50)	(50)	(50)	(50)
Intestine Small, Ileum	(50)	(50)	(50)	(50)
Inflammation, Chronic	1 (2%)		1 (2%)	
Epithelium, Hyperplasia	1 (2%)			
Intestine Small, Jejunum	(50)	(50)	(50)	(50)
Epithelium, Hyperplasia		2 (4%)		
Peyer's Patch, Hyperplasia, Lymphoid			1 (2%)	1 (2%)
Liver	(50)	(50)	(50)	(50)
Amyloid Deposition			1 (2%)	1 (2%)
Angiectasis	3 (6%)	6 (12%)	7 (14%)	10 (20%)
Basophilic Focus	2 (4%)	3 (6%)	3 (6%)	1 (2%)
Clear Cell Focus	18 (36%)	18 (36%)	19 (38%)	21 (42%)
Degeneration, Cystic	1 (2%)		1 (2%)	1 (2%)
Eosinophilic Focus	28 (56%)	32 (64%)	42 (84%)	43 (86%)
Fatty Change	32 (64%)	34 (68%)	27 (54%)	24 (48%)
Fibrosis	1 (2%)			
Hematopoietic Cell Proliferation	1 (2%)	2 (4%)		1 (2%)
Hepatodiaphragmatic Nodule			1 (2%)	
Infarct		1 (2%)	2 (4%)	
Infiltration Cellular, Mononuclear Cell	49 (98%)	47 (94%)	47 (94%)	48 (96%)
Inflammation, Suppurative				1 (2%)
Mineralization	1 (2%)	2 (4%)	2 (4%)	1 (2%)
Mixed Cell Focus	15 (30%)	28 (56%)	15 (30%)	12 (24%)

a - Number of animals examined microscopically at site and number of animals with lesion

B6C3F1 MICE MALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Necrosis	3 (6%)	10 (20%)	7 (14%)	13 (26%)
Tension Lipidosis	5 (10%)	2 (4%)	3 (6%)	2 (4%)
Vacuolization Cytoplasmic		1 (2%)		1 (2%)
Bile Duct, Cyst				1 (2%)
Centrilobular, Fatty Change		1 (2%)		1 (2%)
Centrilobular, Hypertrophy		34 (68%)	30 (60%)	39 (78%)
Mesentery	(3)	(6)	(2)	(3)
Inflammation, Chronic	1 (33%)			
Fat, Necrosis	2 (67%)	6 (100%)	2 (100%)	3 (100%)
Pancreas	(50)	(50)	(50)	(50)
Inflammation, Granulomatous			1 (2%)	
Inflammation, Chronic			1 (2%)	
Acinus, Atrophy	1 (2%)		1 (2%)	
Acinus, Hypertrophy	2 (4%)			
Duct, Cyst				1 (2%)
Salivary Glands	(50)	(50)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	43 (86%)	36 (72%)	40 (80%)	36 (72%)
Parotid Gland, Atrophy	1 (2%)			
Parotid Gland, Hyperplasia			1 (2%)	
Parotid Gland, Mineralization		1 (2%)		
Submandibular Gland, Atrophy		1 (2%)		
Stomach, Forestomach	(50)	(50)	(50)	(50)
Inflammation, Chronic	19 (38%)	22 (44%)	24 (48%)	24 (48%)
Mineralization			1 (2%)	
Necrosis	8 (16%)	11 (22%)	12 (24%)	10 (20%)
Epithelium, Hyperplasia	18 (36%)	22 (44%)	25 (50%)	22 (44%)
Stomach, Glandular	(50)	(50)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	2 (4%)		2 (4%)	1 (2%)
Mineralization		1 (2%)		
Ulcer	1 (2%)			
Epithelium, Hyperplasia	2 (4%)			
Glands, Ectasia	2 (4%)	1 (2%)	2 (4%)	2 (4%)
Tooth	(33)	(32)	(27)	(21)
Dysplasia	33 (100%)	32 (100%)	27 (100%)	21 (100%)
Inflammation, Suppurative		1 (3%)		

## CARDIOVASCULAR SYSTEM

Heart	(50)	(50)	(50)	(50)
Cardiomyopathy	14 (28%)	16 (32%)	11 (22%)	14 (28%)
Inflammation, Suppurative		1 (2%)		1 (2%)
Inflammation, Chronic		1 (2%)		
Artery, Infiltration Cellular, Mononuclear Cell	2 (4%)	1 (2%)	1 (2%)	1 (2%)
Myocardium, Mineralization	1 (2%)	4 (8%)		1 (2%)
Valve, Inflammation				1 (2%)

B6C3F1 MICE MALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
<b>ENDOCRINE SYSTEM</b>				
Adrenal Cortex	(50)	(50)	(50)	(50)
Hypertrophy	1 (2%)	2 (4%)		2 (4%)
Infiltration Cellular, Mononuclear Cell	1 (2%)			
Vacuolization Cytoplasmic	2 (4%)	2 (4%)	2 (4%)	3 (6%)
Subcapsular, Hyperplasia	44 (88%)	45 (90%)	48 (96%)	49 (98%)
Zona Reticularis, Hyperplasia	5 (10%)	3 (6%)	2 (4%)	4 (8%)
Adrenal Medulla	(50)	(50)	(50)	(50)
Hyperplasia	1 (2%)		1 (2%)	
Islets, Pancreatic	(50)	(50)	(50)	(50)
Hyperplasia	37 (74%)	35 (70%)	36 (72%)	33 (66%)
Parathyroid Gland	(47)	(44)	(46)	(44)
Cyst	1 (2%)	1 (2%)		1 (2%)
Infiltration Cellular, Mononuclear Cell	1 (2%)			
Pituitary Gland	(49)	(50)	(50)	(50)
Cyst	1 (2%)		1 (2%)	2 (4%)
Pars Distalis, Hyperplasia	1 (2%)		1 (2%)	
Thyroid Gland	(50)	(50)	(49)	(50)
Atrophy	1 (2%)			
Cyst	4 (8%)		1 (2%)	
Follicle, Hyperplasia	1 (2%)		1 (2%)	
<b>GENERAL BODY SYSTEM</b>				
Peritoneum	(0)	(0)	(0)	(2)
Inflammation, Chronic				2 (100%)
<b>GENITAL SYSTEM</b>				
Coagulating Gland	(0)	(1)	(1)	(0)
Hyperplasia		1 (100%)		
Epididymis	(50)	(50)	(50)	(50)
Granuloma Sperm	2 (4%)		3 (6%)	1 (2%)
Infiltration Cellular, Mononuclear Cell	29 (58%)	22 (44%)	33 (66%)	28 (56%)
Inflammation, Chronic		1 (2%)		
Penis	(0)	(1)	(0)	(0)
Infiltration Cellular, Polymorphonuclear		1 (100%)		
Preputial Gland	(50)	(50)	(50)	(50)
Cyst	4 (8%)	5 (10%)	5 (10%)	3 (6%)
Inflammation	10 (20%)	4 (8%)	9 (18%)	13 (26%)
Prostate	(50)	(50)	(50)	(50)

B6C3F1 MICE MALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Atrophy				1 (2%)
Fibrosis			1 (2%)	
Hyperplasia	1 (2%)			1 (2%)
Infiltration Cellular, Mononuclear Cell	29 (58%)	31 (62%)	38 (76%)	29 (58%)
Inflammation	4 (8%)	9 (18%)	4 (8%)	2 (4%)
Mineralization		1 (2%)		
Seminal Vesicle	(50)	(50)	(50)	(50)
Atrophy				1 (2%)
Fibrosis	2 (4%)			
Hyperplasia				1 (2%)
Infiltration Cellular, Mononuclear Cell	4 (8%)	4 (8%)	6 (12%)	
Inflammation	6 (12%)	5 (10%)		2 (4%)
Testes	(50)	(50)	(50)	(50)
Inflammation	1 (2%)			
Mineralization	1 (2%)			
Germinal Epithelium, Atrophy	2 (4%)	4 (8%)	1 (2%)	1 (2%)
<b>HEMATOPOIETIC SYSTEM</b>				
Bone Marrow	(50)	(50)	(50)	(50)
Lymph Node	(0)	(1)	(3)	(3)
Renal, Hyperplasia, Lymphoid			1 (33%)	
Lymph Node, Mandibular	(50)	(50)	(50)	(49)
Hyperplasia, Lymphoid		3 (6%)	2 (4%)	1 (2%)
Lymph Node, Mesenteric	(50)	(50)	(50)	(49)
Atrophy	1 (2%)			
Hyperplasia, Lymphoid		1 (2%)	2 (4%)	
Spleen	(50)	(50)	(49)	(50)
Angiectasis			1 (2%)	
Hematopoietic Cell Proliferation	16 (32%)	19 (38%)	21 (43%)	15 (30%)
Hyperplasia, Lymphoid	3 (6%)	3 (6%)	2 (4%)	3 (6%)
Lymphoid Follicle, Atrophy	6 (12%)	5 (10%)	3 (6%)	6 (12%)
Thymus	(48)	(47)	(47)	(49)
Atrophy	41 (85%)	41 (87%)	46 (98%)	43 (88%)
Ectopic Thyroid		1 (2%)		
Hyperplasia, Histiocytic	1 (2%)	1 (2%)		
Infiltration Cellular, Histiocyte			2 (4%)	
Inflammation, Chronic Active				1 (2%)
Epithelial Cell, Hyperplasia		1 (2%)		1 (2%)
<b>INTEGUMENTARY SYSTEM</b>				
Skin	(50)	(50)	(50)	(50)
Cyst Epithelial Inclusion			2 (4%)	

B6C3F1 MICE MALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Hemorrhage				1 (2%)
Inflammation, Chronic	1 (2%)	1 (2%)		1 (2%)
Ulcer	1 (2%)	1 (2%)	9 (18%)	4 (8%)
<b>MUSCULOSKELETAL SYSTEM</b>				
Bone	(50)	(50)	(50)	(50)
Cranium, Hyperostosis	1 (2%)			
Skeletal Muscle	(2)	(0)	(1)	(0)
<b>NERVOUS SYSTEM</b>				
Brain	(50)	(50)	(50)	(50)
Hemorrhage		1 (2%)		
Hypothalamus, Compression			1 (2%)	
<b>RESPIRATORY SYSTEM</b>				
Lung	(50)	(50)	(50)	(50)
Inflammation		2 (4%)	3 (6%)	2 (4%)
Metaplasia, Osseous			1 (2%)	
Mineralization			2 (4%)	
Alveolar Epithelium, Hyperplasia	2 (4%)	3 (6%)	1 (2%)	2 (4%)
Alveolus, Infiltration Cellular, Histiocyte	2 (4%)	6 (12%)	5 (10%)	1 (2%)
Bronchus, Hyperplasia	2 (4%)	2 (4%)		1 (2%)
Bronchus, Infiltration Cellular, Histiocyte			1 (2%)	
Nose	(50)	(50)	(50)	(50)
Inflammation, Suppurative	1 (2%)	1 (2%)	4 (8%)	1 (2%)
Inflammation, Chronic	9 (18%)	8 (16%)	7 (14%)	5 (10%)
Polyp, Inflammatory		2 (4%)		1 (2%)
Nasolacrimal Duct, Inflammation, Suppurative	1 (2%)	2 (4%)	2 (4%)	3 (6%)
Respiratory Epithelium, Hyperplasia	1 (2%)		1 (2%)	
Trachea	(50)	(50)	(50)	(50)
Inflammation, Chronic		1 (2%)	1 (2%)	
Epithelium, Cytoplasmic Alteration			1 (2%)	
<b>SPECIAL SENSES SYSTEM</b>				
Eye	(50)	(50)	(50)	(50)
Degeneration		1 (2%)		
Anterior Chamber, Inflammation, Suppurative				1 (2%)
Cornea, Inflammation, Chronic		1 (2%)	1 (2%)	2 (4%)

B6C3F1 MICE MALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Retrobulbar, Inflammation, Chronic				1 (2%)
Harderian Gland	(50)	(50)	(50)	(50)
Fibrosis	1 (2%)			
Hyperplasia	4 (8%)	3 (6%)	1 (2%)	8 (16%)
Infiltration Cellular, Mononuclear Cell	36 (72%)	30 (60%)	34 (68%)	39 (78%)
Inflammation, Granulomatous			1 (2%)	
<b>URINARY SYSTEM</b>				
Kidney	(50)	(50)	(50)	(50)
Amyloid Deposition			1 (2%)	
Hydronephrosis	6 (12%)	1 (2%)		1 (2%)
Inflammation	2 (4%)	2 (4%)	1 (2%)	1 (2%)
Metaplasia, Osseous		2 (4%)		2 (4%)
Mineralization	3 (6%)	1 (2%)		
Nephropathy	32 (64%)	40 (80%)	37 (74%)	38 (76%)
Thrombosis		1 (2%)		
Cortex, Medulla, Necrosis	6 (12%)			
Papilla, Necrosis	3 (6%)			3 (6%)
Renal Tubule, Cyst			3 (6%)	
Renal Tubule, Dilatation		1 (2%)		
Renal Tubule, Hyperplasia	6 (12%)	20 (40%)	13 (26%)	10 (20%)
Renal Tubule, Mineralization	31 (62%)	42 (84%)	38 (76%)	31 (62%)
Renal Tubule, Pigmentation, Lipofuscin	1 (2%)		1 (2%)	2 (4%)
Urethra	(2)	(4)	(0)	(1)
Cyst		1 (25%)		
Inflammation	2 (100%)	4 (100%)		1 (100%)
Necrosis				1 (100%)
Urinary Bladder	(50)	(50)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	20 (40%)	25 (50%)	26 (52%)	20 (40%)
Inflammation	4 (8%)	2 (4%)		
Transitional Epithelium, Cytoplasmic Alteration				1 (2%)
Transitional Epithelium, Hyperplasia		1 (2%)		

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

B6C3F1 MICE FEMALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
<b>Disposition Summary</b>				
Animals Initially in Study	50	50	50	50
Early Deaths				
Accidently Killed	1			1
Moribund Sacrifice	4	4	1	7
Natural Death	7	12	4	5
Survivors				
Terminal Sacrifice	38	34	45	37
Animals Examined Microscopically	50	50	50	50

## ALIMENTARY SYSTEM

Esophagus	(50)	(50)	(50)	(50)
Inflammation, Chronic		1 (2%)		
Gallbladder	(50)	(48)	(49)	(50)
Cyst	1 (2%)	1 (2%)		1 (2%)
Epithelium, Cytoplasmic Alteration				1 (2%)
Intestine Large, Cecum	(50)	(50)	(50)	(50)
Inflammation, Chronic	1 (2%)			
Intestine Small, Duodenum	(50)	(50)	(50)	(50)
Metaplasia, Squamous				1 (2%)
Epithelium, Hyperplasia			1 (2%)	
Intestine Small, Ileum	(50)	(50)	(50)	(50)
Inflammation, Chronic Active			1 (2%)	
Ulcer			1 (2%)	
Intestine Small, Jejunum	(50)	(50)	(50)	(50)
Epithelium, Hyperplasia				1 (2%)
Peyer's Patch, Hyperplasia			1 (2%)	
Liver	(50)	(50)	(50)	(50)
Angiectasis	3 (6%)		2 (4%)	
Basophilic Focus		1 (2%)	4 (8%)	1 (2%)
Clear Cell Focus	3 (6%)		1 (2%)	5 (10%)
Cyst			1 (2%)	
Eosinophilic Focus	9 (18%)	7 (14%)	16 (32%)	26 (52%)
Fatty Change	39 (78%)	33 (66%)	42 (84%)	23 (46%)
Hematopoietic Cell Proliferation	1 (2%)	4 (8%)	4 (8%)	1 (2%)
Infiltration Cellular, Mononuclear Cell	45 (90%)	47 (94%)	49 (98%)	44 (88%)
Mixed Cell Focus	12 (24%)	13 (26%)	14 (28%)	7 (14%)
Necrosis	5 (10%)	4 (8%)	5 (10%)	1 (2%)
Tension Lipidosis	3 (6%)	8 (16%)	7 (14%)	4 (8%)
Vacuolization Cytoplasmic				1 (2%)
Centriobular, Hypertrophy		20 (40%)	48 (96%)	49 (98%)



B6C3F1 MICE FEMALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Mesentery	(8)	(10)	(5)	(6)
Fat, Necrosis	8 (100%)	10 (100%)	5 (100%)	6 (100%)
Pancreas	(49)	(50)	(50)	(50)
Infiltration Cellular, Mononuclear Cell				1 (2%)
Inflammation, Chronic	2 (4%)			
Acinus, Atrophy			1 (2%)	
Acinus, Hypertrophy				1 (2%)
Acinus, Vacuolization Cytoplasmic			1 (2%)	
Duct, Cyst		1 (2%)	1 (2%)	1 (2%)
Salivary Glands	(49)	(49)	(50)	(50)
Atrophy		1 (2%)		
Infiltration Cellular, Mononuclear Cell	34 (69%)	33 (67%)	31 (62%)	26 (52%)
Duct, Submandibular Gland, Hyperplasia			1 (2%)	
Parotid Gland, Hyperplasia				1 (2%)
Stomach, Forestomach	(50)	(50)	(50)	(50)
Erosion		1 (2%)	14 (28%)	11 (22%)
Inflammation, Chronic	3 (6%)	6 (12%)	21 (42%)	22 (44%)
Ulcer		2 (4%)	3 (6%)	6 (12%)
Epithelium, Hyperplasia	3 (6%)	6 (12%)	23 (46%)	24 (48%)
Stomach, Glandular	(50)	(50)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	2 (4%)	1 (2%)		1 (2%)
Mineralization	1 (2%)			
Ulcer				2 (4%)
Epithelium, Hyperplasia			1 (2%)	2 (4%)
Glands, Ectasia	1 (2%)	1 (2%)	2 (4%)	
Tooth	(1)	(1)	(2)	(2)
Dysplasia	1 (100%)	1 (100%)	2 (100%)	2 (100%)
<b>CARDIOVASCULAR SYSTEM</b>				
Blood Vessel	(50)	(50)	(50)	(50)
Mineralization		1 (2%)		
Heart	(50)	(50)	(50)	(50)
Cardiomyopathy	2 (4%)	1 (2%)	4 (8%)	1 (2%)
Thrombosis		1 (2%)		
Artery, Infiltration Cellular, Mononuclear Cell	1 (2%)	4 (8%)	3 (6%)	
Myocardium, Mineralization		2 (4%)		1 (2%)
Valve, Inflammation		2 (4%)		
<b>ENDOCRINE SYSTEM</b>				
Adrenal Cortex	(50)	(50)	(50)	(50)
Atrophy	1 (2%)			
Mineralization	1 (2%)			

B6C3F1 MICE FEMALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Vacuolization Cytoplasmic Subcapsular, Hyperplasia Zona Reticularis, Hyperplasia	1 (2%) 49 (98%)		4 (8%) 50 (100%)	50 (100%) 1 (2%)
Adrenal Medulla Hyperplasia Vacuolization Cytoplasmic	(50) 1 (2%)	(50)	(50) 2 (4%)	(50) 2 (4%) 1 (2%)
Islets, Pancreatic Hyperplasia	(50) 4 (8%)	(50) 4 (8%)	(50) 1 (2%)	(50)
Parathyroid Gland Cyst	(48)	(44)	(48)	(48)
Pituitary Gland Angiectasis Cyst	(50) 1 (2%)	(50) 1 (2%)	(49) 1 (2%) 2 (4%)	(50) 1 (2%)
Pars Distalis, Hyperplasia Thyroid Gland Cyst	4 (8%) (50)	4 (8%) (50)	7 (14%) (50)	5 (10%) (50)
Ectopic Thymus Infiltration Cellular, Mononuclear Cell Follicle, Hyperplasia	1 (2%) 1 (2%) 1 (2%)			
<b>GENERAL BODY SYSTEM</b>				
Tissue NOS Inflammation, Chronic	(0)	(0)	(0)	(1) 1 (100%)
<b>GENITAL SYSTEM</b>				
Clitoral Gland Inflammation	(49) 1 (2%)	(48)	(50)	(50)
Ovary Angiectasis Cyst Hemorrhage Thrombosis	(49) 1 (2%) 4 (8%) 1 (2%)	(50) 1 (2%) 10 (20%)	(50) 2 (4%) 6 (12%)	(50) 1 (2%) 4 (8%)
Uterus Inflammation, Suppurative Metaplasia, Squamous Endometrium, Hyperplasia, Cystic	(50) 3 (6%) 1 (2%) 42 (84%)	(50) 1 (2%) 2 (4%) 44 (88%)	(50) 2 (4%) 42 (84%)	(50) 36 (72%)
Vagina Vacuolization Cytoplasmic	(0)	(0)	(0)	(1) 1 (100%)
<b>HEMATOPOIETIC SYSTEM</b>				

B6C3F1 MICE FEMALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Bone Marrow	(50)	(50)	(50)	(50)
Fibrosis		2 (4%)	1 (2%)	1 (2%)
Necrosis				1 (2%)
Lymph Node	(3)	(3)	(0)	(0)
Mediastinal, Hyperplasia, Lymphoid	1 (33%)			
Mediastinal, Hyperplasia, Plasma Cell		1 (33%)		
Lymph Node, Mandibular	(48)	(49)	(50)	(50)
Hyperplasia, Lymphoid	4 (8%)	4 (8%)	4 (8%)	2 (4%)
Lymph Node, Mesenteric	(50)	(50)	(50)	(50)
Hyperplasia, Lymphoid				1 (2%)
Hyperplasia, Plasma Cell		1 (2%)		
Inflammation, Granulomatous				1 (2%)
Spleen	(49)	(48)	(50)	(49)
Hematopoietic Cell Proliferation	26 (53%)	24 (50%)	20 (40%)	10 (20%)
Hyperplasia, Lymphoid	7 (14%)	6 (13%)	6 (12%)	5 (10%)
Lymphoid Follicle, Atrophy	4 (8%)	2 (4%)	4 (8%)	2 (4%)
Thymus	(50)	(50)	(49)	(47)
Atrophy	39 (78%)	44 (88%)	46 (94%)	40 (85%)
Hyperplasia, Histiocytic	1 (2%)		2 (4%)	1 (2%)
Infiltration Cellular, Histiocyte	5 (10%)			2 (4%)
Epithelial Cell, Hyperplasia	1 (2%)			
<b>INTEGUMENTARY SYSTEM</b>				
Mammary Gland	(50)	(49)	(50)	(50)
Hyperplasia	1 (2%)			
Inflammation, Chronic Active	1 (2%)			
Metaplasia, Squamous	1 (2%)			
Skin	(50)	(50)	(50)	(50)
Fibrosis			1 (2%)	
Inflammation, Chronic	1 (2%)		1 (2%)	
Ulcer	1 (2%)	1 (2%)		
Subcutaneous Tissue, Fibrosis		1 (2%)		
Subcutaneous Tissue, Necrosis			1 (2%)	
<b>MUSCULOSKELETAL SYSTEM</b>				
Bone	(50)	(50)	(50)	(50)
Fibrosis	15 (30%)	8 (16%)	7 (14%)	9 (18%)
Joint, Inflammation, Chronic		1 (2%)		
Skeletal Muscle	(3)	(1)	(0)	(0)
<b>NERVOUS SYSTEM</b>				

B6C3F1 MICE FEMALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Brain	(50)	(50)	(50)	(50)
Choroid Plexus, Infiltration Cellular, Mononuclear Cell	1 (2%)			
Hypothalamus, Compression	2 (4%)			
Peripheral Nerve	(1)	(0)	(0)	(1)
Infiltration Cellular, Mononuclear Cell	1 (100%)			
Axon, Degeneration	1 (100%)			1 (100%)
Spinal Cord	(1)	(0)	(0)	(1)
Axon, Degeneration	1 (100%)			
<b>RESPIRATORY SYSTEM</b>				
Lung	(50)	(50)	(50)	(50)
Hemorrhage	1 (2%)			
Inflammation	1 (2%)		2 (4%)	
Inflammation, Acute		1 (2%)		
Metaplasia, Osseous		1 (2%)		
Mineralization		1 (2%)		
Alveolar Epithelium, Hyperplasia	3 (6%)	1 (2%)		1 (2%)
Alveolus, Infiltration Cellular, Histiocyte		1 (2%)		1 (2%)
Serosa, Fibrosis		1 (2%)		
Nose	(50)	(50)	(50)	(50)
Inflammation, Suppurative				1 (2%)
Inflammation, Chronic	4 (8%)	2 (4%)		
Glands, Dilatation			1 (2%)	
Nasolacrimal Duct, Inflammation, Suppurative			1 (2%)	
Olfactory Epithelium, Degeneration	1 (2%)			
Trachea	(50)	(50)	(50)	(50)
Inflammation, Chronic		1 (2%)		
<b>SPECIAL SENSES SYSTEM</b>				
Eye	(50)	(50)	(50)	(50)
Anterior Chamber, Inflammation, Suppurative	1 (2%)			
Harderian Gland	(50)	(49)	(50)	(50)
Atrophy	1 (2%)		1 (2%)	
Hyperplasia	3 (6%)	4 (8%)	5 (10%)	5 (10%)
Infiltration Cellular, Mononuclear Cell	38 (76%)	42 (86%)	42 (84%)	38 (76%)
<b>URINARY SYSTEM</b>				
Kidney	(50)	(50)	(50)	(50)
Amyloid Deposition	1 (2%)	2 (4%)	2 (4%)	1 (2%)

TDMS No. 20007 - 06

Test Type: CHRONIC

Route: GAVAGE

Species/Strain: MICE/B6C3F1

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

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

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B6C3F1 MICE FEMALE	0 G/KG	0.25 G/KG	0.5 G/KG	1.0 G/KG
Hydronephrosis		1 (2%)		
Inflammation	1 (2%)	1 (2%)		1 (2%)
Metaplasia, Osseous	3 (6%)	1 (2%)	1 (2%)	5 (10%)
Mineralization	12 (24%)	20 (40%)	7 (14%)	13 (26%)
Nephropathy	21 (42%)	25 (50%)	26 (52%)	20 (40%)
Papilla, Necrosis	1 (2%)	3 (6%)	1 (2%)	3 (6%)
Pelvis, Cyst			1 (2%)	
Renal Tubule, Hyperplasia	3 (6%)	1 (2%)	2 (4%)	
Urinary Bladder	(50)	(48)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	34 (68%)	34 (71%)	40 (80%)	34 (68%)
Inflammation		1 (2%)		

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