Test Type: CHRONIC

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH

AVERAGE SEVERITY GRADES[b]

Pulegone

CAS Number: 89-82-7

Time Report Requested: 09:18:35 First Dose M/F: 04/15/03 / 04/14/03

Date Report Requested: 07/23/2008

Lab: BAT

F1__M3

C Number: C99020

Lock Date: 12/21/2005

Cage Range: ALL

Date Range: ALL

Reasons For Removal: 25022 ACCK 25021 TSAC 25020 NATD

> 25019 MSAC 25018 DACC

Removal Date Range: ALL

Treatment Groups: Include ALL

Study Gender: Both

TDMSE Version: 2.0.0

Species/Strain: MICE/B6C3F1

Route: GAVAGE

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Lab: BAT

B6C3F1 MICE MALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
isposition Summary					
Animals Initially in Study	50	50	50	50	
Early Deaths					
Accidently Killed		1			
Dosing Accident	_	1	_	_	
Moribund Sacrifice	7	8	7	8	
Natural Death	5	4	1	1	
Survivors	20	26	40	44	
Terminal Sacrifice Animals Examined Microscopically	38 50	36 50	42 50	41 50	
Animais Examined microscopically	50	30	30	30	
LIMENTARY SYSTEM					
Esophagus	(50)	(50)	(50)	(50)	
Inflammation	2 [1.5]		1 [2.0]	1 [1.0]	
Gallbladder	(49)	(49)	(50)	(50)	
Infiltration Cellular, Mononuclear Cell	1 [1.0]	(==)	(==)	(==)	
Intestine Large, Cecum	(50)	(50)	(50)	(50)	
Hemorrhage	(50)	1 [3.0]	(50)	(50)	
Intestine Small, Ileum	(50)	(50)	(50)	(50)	
Hyperplasia, Lymphoid Inflammation	1 [2.0]	1 [3.0]			
Epithelium, Hyperplasia	1 [2.0]	ا [۵.0]			
Intestine Small, Jejunum	(50)	(50)	(50)	(50)	
Hyperplasia, Lymphoid	1 [3.0]	(30)	(30)	(00)	
Liver	(50)	(50)	(50)	(50)	
Angiectasis	, ,	2 [1.0]	,	,	
Basophilic Focus	4	5	4	6	
Clear Cell Focus	15	27	28	34	
Eosinophilic Focus	7	12	20	36	
Fatty Change, Focal	3 [2.0]	8 [1.4]	20 [1.2]	23 [1.2]	
Fatty Change, Diffuse	38 [1.3]	27 [1.1]	21 [1.0]	3 [3.3]	
Hematopoietic Cell Proliferation	3 [1.0]		1 [1.0]	2 [1.0]	
Hemorrhage	4			1 [1.0]	
Hepatodiaphragmatic Nodule	1	24 [4 0]	20 [4 4]	20 [4 4]	
Inflammation	24 [1.0]	21 [1.0]	20 [1.1]	29 [1.1]	

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

18

Mixed Cell Focus

19

34

20

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

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

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Time Report Requested: 09:18:35 First Dose M/F: 04/15/03 / 04/14/03

3F1 MICE MALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG
ecrosis	1 [1.0]	8 [1.6]	5 [1.6]	26 [1.6]
gmentation	3 [1.3]	0 [0]	2 [1.0]	1 [1.0]
ension Lipidosis	5	6	7	2
le Duct, Cyst	·	· ·	3 [2.0]	_ 14 [1.9]
le Duct, Fibrosis			0 [=.0]	2 [1.5]
le Duct, Hyperplasia			1 [1.0]	35 [1.5]
entrilobular, Degeneration		1 [2.0]	. [0]	55 [5]
entrilobular, Vacuolization Cytoplasmic		1 [2.0]	1 [2.0]	
entrilobular, Hepatocyte, Hypertrophy		10 [1.1]	22 [1.1]	46 [1.7]
epatocyte, Hypertrophy		1 [1.0]	1 [2.0]	
val Cell, Hyperplasia	1 [1.0]	. []	1 [1.0]	36 [1.4]
erosa, Inflammation, Chronic Active	. []	1 [3.0]	. []	
ein, Intravascular Hepatocyte	3 [1.0]	1 [1.0]	15 [1.0]	47 [2.1]
sentery	(4)	(2)	(3)	(O)
at, Necrosis	3 [3.0]	1 [3.0]	3 [3.0]	()
creas	(50)	(50)	(50)	(50)
yst	1 [2.0]	, ,	, ,	, ,
ytoplasmic Alteration, Focal		2 [1.0]	1 [2.0]	1 [1.0]
filtration Cellular, Mononuclear Cell	5 [1.0]	5 [1.2]	4 [1.0]	7 [1.3]
cinus, Atrophy	1 [3.0]	1 [2.0]		1 [3.0]
vary Glands	(50)	(50)	(50)	(50)
trophy			1 [2.0]	
filtration Cellular, Mononuclear Cell	40 [1.4]	32 [1.2]	30 [1.2]	33 [1.1]
mach, Forestomach	(50)	(50)	(50)	(50)
yperplasia, Squamous	7 [2.0]	10 [2.3]	27 [2.1]	41 [2.6]
flammation	3 [1.7]	9 [2.0]	24 [1.7]	39 [2.1]
ineralization		1 [1.0]		
lcer		3 [1.3]	9 [1.9]	22 [2.0]
rtery, Inflammation, Chronic Active	1 [2.0]	4	<i>(</i>)	()
mach, Glandular	(50)	(50)	(50)	(50)
rosion	0.54.03	4.54.03		1 [2.0]
ineralization	2 [1.0]	1 [1.0]		
pithelium, Hyperplasia	4.54.03	1 [2.0]	0.14.01	5.14.03
lands, Cyst	4 [1.0]	4 [1.0]	3 [1.0]	5 [1.0]
lands, Hyperplasia	1 [2.0]	1 [2.0]	(25)	(20)
th	(37)	(37)	(35)	(20)
ysplasia	37 [2.2]	37 [1.9]	35 [1.8]	20 [1.1]
eridontal Tissue, Inflammation ulp, Inflammation	1 [2 0]	1 [2 0]	1 [3.0]	1 [2 0]
ap, irinamination	1 [2.0]	1 [3.0]	1 [2.0]	1 [3.0]

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

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

Test Type: CHRONIC

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

Pulegone

CAS Number: 89-82-7

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B6C3F1 MICE MALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
CARDIOVASCULAR SYSTEM					
Blood Vessel	(50)	(50)	(50)	(50)	
Heart Cardiomyopathy	(50) 1 [1.0]	(50)	(50)	(50)	
Infiltration Cellular, Mononuclear Cell	4 [1.0]		1 [1.0]	1 [1.0]	
Inflammation	1 [1.0]		1		
Mineralization	1 [2.0]	1 [1.0]	1 [1.0]	4 [2.8]	
NDOCRINE SYSTEM					
Adrenal Cortex	(50)	(50)	(50)	(50)	
Degeneration, Cystic	47 [4 6]	0.[4.6]	1 [2.0]	4 [4 2]	
Hypertrophy Vacuolization Cytoplasmic	17 [1.6]	9 [1.6] 1 [2.0]	8 [1.3]	4 [1.3] 1 [1.0]	
Subcapsular, Hyperplasia	39 [1.3]	40 [1.4]	47 [1.2]	44 [1.4]	
Zona Fasciculata, Hyperplasia	1 [2.0]		[=]		
Adrenal Medulla	(50)	(50)	(50)	(50)	
Hyperplasia		2 [1.0]	1 [3.0]	2 [1.0]	
Islets, Pancreatic	(50)	(50)	(50)	(50)	
Hyperplasia	20 [1.4]	17 [1.5]	21 [1.2]	2 [1.0]	
Pituitary Gland Cyst	(50) 1 [1.0]	(50)	(50)	(50) 1 [1.0]	
Pars Distalis, Hyperplasia	1 [1.0]		1 [3.0]	1 [1.0]	
Thyroid Gland	(50)	(50)	(50)	(50)	
Infiltration Cellular, Mononuclear Cell	1 [1.0]	(55)	(00)	1 [2.0]	
Follicular Cell, Hyperplasia				1 [1.0]	
GENERAL BODY SYSTEM					
None					
GENITAL SYSTEM					
Coagulating Gland	(1)	(1)	(0)	(1)	
Hyperplasia	1 [2.0]	(' /	(0)	(· /	
Inflammation				1 [3.0]	
Epididymis	(50)	(50)	(50)	(50)	

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Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

Pulegone

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B6C3F1 MICE MALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
Cyst		1 [3.0]			
Granuloma Sperm			1 [2.0]		
Infiltration Cellular, Mononuclear Cell	23 [1.2]	27 [1.1]	20 [1.1]	20 [1.0]	
Inflammation	2 [1.5]				
Mineralization	1 -1	1 [1.0]			
Preputial Gland	(50)	(50)	(50)	(50)	
Cyst	(/	()	()	2 [2.0]	
Infiltration Cellular, Mononuclear Cell	10 [1.1]	15 [1.3]	12 [1.3]	7 [1.1]	
Inflammation	4 [1.0]	4 [3.0]	3 [2.3]	4 [1.3]	
Duct, Ectasia	1 [2.0]	1 [2.0]	3 [2.0]	3 [1.7]	
Prostate	(50)	(50)	(50)	(50)	
Infiltration Cellular, Mononuclear Cell	23 [1.0]	30 [1.0]	23 [1.0]	14 [1.1]	
Inflammation	1 [1.0]	30 [1.0]	1 [2.0]	1 [3.0]	
Epithelium, Hyperplasia	6 [1.0]	3 [1.3]	1 [2.0]	1 [3.0]	
Seminal Vesicle	(50)	(50)	(50)	(50)	
Atrophy	(30)	1 [3.0]	(30)	(30)	
Dilatation		1 [2.0]			
Testes	(50)	(50)	(50)	(50)	
	(50)		(50)	(50)	
Cyst		1 [3.0]	4 [0.0]		
Mineralization	0.00.01	2 [1.5]	1 [2.0]		
Germinal Epithelium, Atrophy	2 [2.0]	2 [1.5]	3 [2.0]		
MATOPOIETIC SYSTEM					
Bone Marrow	(50)	(50)	(50)	(50)	
Atrophy, Focal				1 [2.0]	
Hemorrhage				1 [2.0]	
Myelofibrosis	2 [2.0]		3 [1.3]	1 [1.0]	
Lymph Node	(1)	(3)	(0)	(2)	
Lymph Node, Mandibular	(50)	(50)	(50)	(50)	
Atrophy	1 [2.0]	4 [2.0]	3 [2.3]	2 [2.5]	
Hyperplasia, Lymphoid	1 [2.0]	1 [3.0]			
Hyperplasia, Plasma Cell	2 [1.5]	1 [2.0]			
Lymph Node, Mesenteric	(47)	(49)	(45)	(44)	
Atrophy	\ /	1 [2.0]	2 [2.5]	3 [2.0]	
Hyperplasia, Lymphoid	1 [2.0]	. []	- []	1 [3.0]	
Spleen	(50)	(50)	(50)	(50)	
Atrophy	(00)	(33)	(00)	1 [3.0]	
Hematopoietic Cell Proliferation	29 [1.3]	32 [1.7]	23 [1.9]	23 [1.3]	
riomatopolotio con rifoliloration	20 [1.0]	ο <u>-</u> [· · ·]	20[1.0]	_0 [1.0]	

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Test Type: CHRONIC

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B6C3F1 MICE MALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
Hyperplasia, Lymphoid Lymphoid Follicle, Atrophy Red Pulp, Atrophy Thymus Atrophy Hyperplasia, Lymphoid Necrosis, Lymphoid	2 [2.5] 1 [2.0] 1 [2.0] (49) 36 [2.4]	3 [1.7] (46) 35 [2.6] 2 [2.0] 2 [2.5]	1 [3.0] 1 [2.0] (45) 32 [2.7]	3 [2.3] 3 [2.3] (42) 30 [2.3]	
NTEGUMENTARY SYSTEM					
Skin Inflammation Ulcer Epidermis, Hyperplasia Subcutaneous Tissue, Hemorrhage	(50)	(50) 1 [3.0] 1 [3.0] 1 [2.0]	(50) 2 [2.5] 2 [2.5] 1 [2.0]	(50)	
MUSCULOSKELETAL SYSTEM					
Bone Fracture	(50)	(50) 1	(50)	(50)	
Osteosclerosis Skeletal Muscle	(1)	1 [2.0]	(0)	1 [3.0] (0)	
NERVOUS SYSTEM					
Brain Arteriole, Infiltration Cellular, Lymphoid Meninges, Infiltration Cellular, Lymphoid	(50)	(50) 1 [1.0] 1 [2.0]	(50)	(50)	
RESPIRATORY SYSTEM					
Lung Hemorrhage	(50) 1 [2.0]	(50)	(50)	(50)	
Inflammation Alveolar Epithelium, Hyperplasia	2 [3.0]	5 [1.8] 7 [2.0]	2 [2.5] 2 [2.5]	1 [2.0] 9 [1.7]	
Alveolus, Infiltration Cellular, Histiocyte Serosa, Inflammation	2 [1.0]	4 [1.5] 2 [2.5]	3 [2.0]	3 [1.0]	
Nose	(50)	(50)	(50)	(50)	

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B6C3F1 MICE MALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG
Inflammation	2 [1.5]	3 [1.7]	2 [2.0]	22 [1.4]
Polyp, Inflammatory		2 [2.5]		
Glands, Cyst	4 [4 0]	1 [2.0]	0.14.01	45 [4 0]
Nerve, Atrophy Olfactory Epithelium, Degeneration	1 [1.0] 3 [2.0]	3 [1.7] 3 [2.0]	3 [1.0] 11 [1.4]	45 [1.8] 46 [1.7]
Olfactory Epithelium, Erosion	3 [2.0]	3 [2.0]	11[1.4]	1 [2.0]
Olfactory Epithelium, Metaplasia	1 [2.0]	5 [1.4]	3 [1.0]	44 [1.9]
Olfactory Epithelium, Necrosis				1 [3.0]
Respiratory Epithelium, Hyperplasia	39 [1.1]	38 [1.2]	42 [1.2]	40 [1.0]
SPECIAL SENSES SYSTEM				
Eye	(50)	(50)	(50)	(50)
Atrophy	, ,	, ,		1 [4.0]
Cornea, Inflammation	1 [2.0]	4 [0 0]	1 [2.0]	3 [1.3]
Optic Nerve, Degeneration Harderian Gland	(50)	1 [2.0] (50)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	34 [1.0]	27 [1.0]	23 [1.0]	17 [1.0]
Inflammation, Chronic Active	01[1.0]	1 [3.0]	20 [1.0]	17 [1.0]
Epithelium, Hyperplasia	1 [1.0]	3 [2.0]	1 [2.0]	3 [2.0]
JRINARY SYSTEM				
Kidney	(50)	(50)	(50)	(50)
Accumulation, Hyaline Droplet	1 [2.0]			
Cyst	9 [1.4]	3 [1.0]	4 [1.3]	44 [4 0]
Glomerulopathy, Hyaline Infarct	1 [1.0] 1 [3.0]	19 [1.1]	30 [1.1]	44 [1.9] 1 [1.0]
Metaplasia, Osseous	3 [2.0]	2 [2.0]	2 [1.5]	4 [2.3]
Mineralization	30 [1.0]	34 [1.0]	41 [1.1]	38 [1.0]
Necrosis				1 [2.0]
Nephropathy	45 [1.2]	45 [1.3]	49 [1.4]	49 [1.9]
Pigmentation Glomerulus, Congestion	9 [1.6]	14 [1 4]	2 [3.0] 17 [1.4]	10 01 11
Papilla, Necrosis	9 [1.6] 1 [3.0]	14 [1.4]	17 [1. 4]	44 [2.0]
Pelvis, Inflammation	1 [2.0]			2 [2.5]
Renal Tubule, Hyperplasia	1 [2.0]		1 [2.0]	1 [1.0]
Urethra	(0)	(0)	(1)	(0)
Urinary Bladder	(50)	(50)	(50)	(50)

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TDMS No. 99020 - 06
Test Type: CHRONIC

Species/Strain: MICE/B6C3F1

Route: GAVAGE

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Lab: BAT

B6C3F1 MICE MALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
Infiltration Cellular, Mononuclear Cell Transitional Epithelium, Hyperplasia	28 [1.0]	19 [1.0]	19 [1.0]	31 [1.1] 1 [2.0]	

*** END OF MALE ***

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

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

Species/Strain: MICE/B6C3F1

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Disposition Summary Animals Initially in Study Early Deaths	50 9	50	50	50
Early Deaths	9	50	50	50
				50
				4
Dosing Accident Moribund Sacrifice		3	5	1 9
Natural Death	5	6		3
Survivors	9	O	,	J
Terminal Sacrifice	35	41	38	37
Animals Examined Microscopically	49	50	50	50
ALIMENTARY SYSTEM				
	(42)	(==)	(==)	(==)
Esophagus	(49)	(50)	(50)	(50)
Perforation		4 [2 0]		1 [3.0]
Periesophageal Tissue, Inflammation Gallbladder	(48)	1 [3.0] (50)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	1 [1.0]	2 [1.5]	(30)	1 [2.0]
Intestine Large, Cecum	(49)	(50)	(50)	(50)
Intestine Large, Celon	(49)	(50)	(50)	(50)
Intestine Large, Colon Intestine Large, Rectum	(49)	(50)	(50)	(50)
Artery, Inflammation, Chronic Active	(40)	1 [3.0]	(00)	(00)
Intestine Small, Duodenum	(49)	(50)	(50)	(50)
Artery, Inflammation, Chronic Active	(/	1 [2.0]	(30)	()
Intestine Small, Ileum	(49)	(50)	(50)	(50)
Artery, Inflammation, Chronic Active	` '	1 [3.0]	` '	` '
Intestine Small, Jejunum	(49)	(50)	(50)	(50)
Artery, Inflammation, Chronic Active		1 [2.0]		
Liver	(49)	(50)	(50)	(50)
Angiectasis		1 [3.0]	3 [1.3]	1 [2.0]
Basophilic Focus	1	3		1
Clear Cell Focus	0	6	23	32
Eosinophilic Focus	3	7	10	31
Fatty Change, Focal	1 [1.0]	2 [1.5]	20 [1.2]	12 [1.4]
Fatty Change, Diffuse Fibrosis	36 [1.5]	31 [1.1]	34 [1.4]	3 [2.7]
Hematopoietic Cell Proliferation	1 [1.0]			2 [2.0] 1 [1.0]
Inflammation	40 [1.2]	40 [1.1]	14 [1.0]	31 [1.1]

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Species/Strain: MICE/B6C3F1

Route: GAVAGE

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B6C3F1 MICE FEMALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
Mineralization				1 [3.0]	
Mixed Cell Focus	4	8	16	20	
Necrosis	5 [2.2]	2 [2.0]	4 [2.5]	27 [1.3]	
Pigmentation	0 [2.2]	_ [=.0]	. [2.0]	46 [1.5]	
Tension Lipidosis	5	5	4	10 [110]	
Bile Duct, Cyst	Ŭ	· ·	4 [2.0]	38 [2.6]	
Bile Duct, Hyperplasia			2 [1.0]	47 [2.2]	
Centrilobular, Hepatocyte, Hypertrophy		3 [1.3]	7 [1.0]	18 [1.1]	
Hepatocyte, Hypertrophy		1 [1.0]	5 [1.2]	11 [1.4]	
Oval Cell, Hyperplasia		1 [1.0]	3 [1.3]	46 [1.5]	
Portal, Fibrosis			O [1.0]	1 [2.0]	
Vein, Hypertrophy				1 [2.0]	
Vein, Intravascular Hepatocyte		2 [1.0]	20 [1.1]	46 [2.2]	
Mesentery	(6)	(12)	(11)	(2)	
Necrosis	2 [3.0]	()	(,	(-)	
Artery, Inflammation, Chronic Active	2 [0.0]	1 [3.0]			
Fat, Necrosis	4 [3.0]	11 [3.0]	10 [3.0]	1 [2.0]	
Pancreas	(49)	(50)	(50)	(50)	
Atrophy	(10)	(33)	(00)	1 [2.0]	
Cytoplasmic Alteration, Focal	1 [2.0]	2 [1.5]		2 [2.0]	
Infiltration Cellular, Mononuclear Cell	12 [1.2]	16 [1.2]	15 [1.1]	11 [1.4]	
Acinus, Atrophy	1 [2.0]	1 [2.0]	10[111]	[]	
Artery, Inflammation, Chronic Active	1 [2.0]	1 [3.0]			
Salivary Glands	(49)	(50)	(49)	(50)	
Infiltration Cellular, Mononuclear Cell	39 [1.2]	29 [1.3]	28 [1.2]	26 [1.1]	
Stomach, Forestomach	(49)	(50)	(50)	(50)	
Hyperplasia, Squamous	13 [2.1]	1 [1.0]	10 [1.9]	26 [2.1]	
Inflammation	10 [1.9]	. [0]	7 [1.9]	20 [2.1]	
Mineralization	.0[0]		. [0]	1 [1.0]	
Ulcer	8 [2.1]		4 [2.0]	10 [2.0]	
Stomach, Glandular	(49)	(50)	(50)	(50)	
Mineralization	(10)	(00)	1 [2.0]	(55)	
Artery, Inflammation, Chronic Active		1 [2.0]	. [=.0]		
Glands, Cyst	2 [1.0]	3 [1.0]	3 [1.0]	2 [1.0]	
Tongue	(0)	(0)	(0)	(1)	
Tooth	(3)	(1)	(2)	(1)	
Dysplasia	3 [1.3]	1 [3.0]	2 [2.5]	1 [1.0]	
Peridontal Tissue, Inflammation	0[0]	1 [3.0]	2 [2.0]	. []	

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

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

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

Pulegone

CAS Number: 89-82-7

Date Report Requested: 07/23/2008

Time Report Requested: 09:18:35 First Dose M/F: 04/15/03 / 04/14/03

Lab: BAT

B6C3F1 MICE FEMALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG
CARDIOVASCULAR SYSTEM				
Blood Vessel	(49)	(50)	(50)	(50)
Mineralization			1 [3.0]	
Heart	(49)	(50)	(50)	(50)
Cardiomyopathy	1 [1.0]	2 [1.5]	1 [1.0]	5 [1.2]
Infiltration Cellular, Mononuclear Cell	1 [2.0]			
Inflammation	1 [2.0]	1 [2.0]	1 [1.0]	1 [2.0]
Mineralization	2 [3.0]		2 [3.0]	8 [1.9]
Necrosis				2 [2.5]
Ventricle, Thrombosis				1 [1.0]
ENDOCRINE SYSTEM				
Adrenal Cortex	(49)	(50)	(50)	(50)
Hypertrophy	, ,	1 [2.0]	1 [1.0]	1 [2.0]
Inflammation				1 [1.0]
Vacuolization Cytoplasmic	1 [2.0]		1 [2.0]	
Subcapsular, Hyperplasia	49 [2.0]	50 [2.1]	49 [2.0]	49 [2.1]
Adrenal Medulla	(49)	(50)	(50)	(50)
Hyperplasia	1 [3.0]			
Islets, Pancreatic	(49)	(50)	(50)	(50)
Hyperplasia	5 [1.4]	2 [1.0]	3 [1.0]	1 [1.0]
Parathyroid Gland	(45)	(47)	(39)	(39)
Infiltration Cellular, Lymphocyte		1 [3.0]		
Pituitary Gland	(49)	(50)	(49)	(49)
Angiectasis	3 [1.7]	3 [1.7]	1 [2.0]	2 [1.0]
Cyst				1 [2.0]
Degeneration	1 [3.0]			
Pars Distalis, Cyst	- r- ·	2 [2.0]	# FG 63	4 10
Pars Distalis, Hyperplasia	7 [2.4]	4 [1.8]	5 [2.0]	4 [2.3]
Thyroid Gland	(49)	(50)	(49)	(50)
Infiltration Cellular, Mononuclear Cell	1 [1.0]	4 [4 0]	1 [1.0]	
C-cell, Hyperplasia	4 [0 0]	1 [1.0]		
Follicle, Cyst	1 [3.0]			4 [0.0]
Follicular Cell, Hyperplasia	1 [3.0]			1 [3.0]

GENERAL BODY SYSTEM

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

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

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

Pulegone

CAS Number: 89-82-7

Date Report Requested: 07/23/2008

Time Report Requested: 09:18:35 First Dose M/F: 04/15/03 / 04/14/03

B6C3F1 MICE FEMALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
None					
ENITAL SYSTEM					
Clitoral Gland	(47)	(50)	(49)	(49)	
Infiltration Cellular, Mononuclear Cell	. ,		1 [2.0]	1 [1.0]	
Inflammation		1 [3.0]	1 [2.0]	3 [1.7]	
Ovary	(49)	(49)	(50)	(50)	
Angiectasis	1 [2.0]	1 [3.0]	1 [3.0]		
Atrophy	11 [3.3]	10 [3.3]	24 [3.5]	39 [3.9]	
Cyst	6 [2.3]	7 [2.4]	9 [2.1]	7 [2.4]	
Uterus	(49)	(49)	(50)	(50)	
Angiectasis	3 [2.3]				
Decidual Reaction		1 [3.0]			
Infiltration Cellular, Mononuclear Cell	4 [0 0]	4 [0.0]		1 [1.0]	
Inflammation	1 [2.0]	1 [2.0]			
Metaplasia Endometrium, Hyperplasia, Cystic	32 [2.8]	1 [2.0] 32 [2.6]	40 [2.6]	35 [3.0]	
Endometham, rryperplasia, Cystic	32 [2.0]	32 [2.0]	40 [2.0]	30 [3.0]	
EMATOPOIETIC SYSTEM					
Bone Marrow	(49)	(50)	(50)	(50)	
Atrophy			1 [2.0]	1 [2.0]	
Hyperplasia	2 [2.5]	9 [2.6]	4 [2.5]	5 [2.6]	
Myelofibrosis	20 [1.6]	17 [1.8]	17 [1.6]	14 [1.6]	
Myelofibrosis Necrosis		17 [1.8]	17 [1.6] 1 [2.0]	14 [1.6]	
Myelofibrosis Necrosis Lymph Node	(1)	17 [1.8] (2)	17 [1.6] 1 [2.0] (5)	14 [1.6] (2)	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular		17 [1.8] (2) (50)	17 [1.6] 1 [2.0]	14 [1.6] (2) (50)	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy	(1) (49)	17 [1.8] (2)	17 [1.6] 1 [2.0] (5) (49)	14 [1.6] (2) (50) 2 [3.0]	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid	(1)	17 [1.8] (2) (50)	17 [1.6] 1 [2.0] (5) (49) 1 [3.0]	14 [1.6] (2) (50)	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid Pigmentation	(1) (49) 2 [2.5]	17 [1.8] (2) (50) 1 [3.0]	17 [1.6] 1 [2.0] (5) (49) 1 [3.0] 1 [2.0]	14 [1.6] (2) (50) 2 [3.0] 2 [2.5]	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid Pigmentation Lymph Node, Mesenteric	(1) (49)	17 [1.8] (2) (50)	17 [1.6] 1 [2.0] (5) (49) 1 [3.0] 1 [2.0] (50)	14 [1.6] (2) (50) 2 [3.0] 2 [2.5] (49)	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid Pigmentation Lymph Node, Mesenteric Atrophy	(1) (49) 2 [2.5]	17 [1.8] (2) (50) 1 [3.0]	17 [1.6] 1 [2.0] (5) (49) 1 [3.0] 1 [2.0] (50) 2 [2.5]	14 [1.6] (2) (50) 2 [3.0] 2 [2.5]	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid Pigmentation Lymph Node, Mesenteric Atrophy Hyperplasia, Reticulum Cell	(1) (49) 2 [2.5] (47)	17 [1.8] (2) (50) 1 [3.0] (49)	17 [1.6] 1 [2.0] (5) (49) 1 [3.0] 1 [2.0] (50) 2 [2.5] 1 [3.0]	14 [1.6] (2) (50) 2 [3.0] 2 [2.5] (49) 1 [3.0]	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid Pigmentation Lymph Node, Mesenteric Atrophy Hyperplasia, Reticulum Cell Spleen	(1) (49) 2 [2.5] (47)	17 [1.8] (2) (50) 1 [3.0] (49)	17 [1.6] 1 [2.0] (5) (49) 1 [3.0] 1 [2.0] (50) 2 [2.5] 1 [3.0] (50)	14 [1.6] (2) (50) 2 [3.0] 2 [2.5] (49) 1 [3.0]	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid Pigmentation Lymph Node, Mesenteric Atrophy Hyperplasia, Reticulum Cell Spleen Hematopoietic Cell Proliferation	(1) (49) 2 [2.5] (47) (49) 24 [1.6]	17 [1.8] (2) (50) 1 [3.0] (49) (50) 29 [1.4]	17 [1.6] 1 [2.0] (5) (49) 1 [3.0] 1 [2.0] (50) 2 [2.5] 1 [3.0] (50) 25 [1.4]	14 [1.6] (2) (50) 2 [3.0] 2 [2.5] (49) 1 [3.0] (50) 40 [1.5]	
Myelofibrosis Necrosis Lymph Node Lymph Node, Mandibular Atrophy Hyperplasia, Lymphoid Pigmentation Lymph Node, Mesenteric Atrophy Hyperplasia, Reticulum Cell Spleen	(1) (49) 2 [2.5] (47)	17 [1.8] (2) (50) 1 [3.0] (49)	17 [1.6] 1 [2.0] (5) (49) 1 [3.0] 1 [2.0] (50) 2 [2.5] 1 [3.0] (50)	14 [1.6] (2) (50) 2 [3.0] 2 [2.5] (49) 1 [3.0]	

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

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

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

Pulegone

CAS Number: 89-82-7

Date Report Requested: 07/23/2008

Time Report Requested: 09:18:35 First Dose M/F: 04/15/03 / 04/14/03

B6C3F1 MICE FEMALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
Lymphoid Follicle, Atrophy Red Pulp, Atrophy Thymus Atrophy Hyperplasia, Lymphoid	1 [2.0] (48) 28 [2.5] 4 [2.0]	(49) 21 [2.5] 5 [2.0]	(49) 20 [2.7] 2 [2.5]	1 [3.0] 1 [2.0] (46) 13 [2.5] 1 [2.0]	
INTEGUMENTARY SYSTEM					
Mammary Gland Skin	(49) (49)	(50) (50)	(50) (50)	(50) (50)	
MUSCULOSKELETAL SYSTEM					
Bone Osteosclerosis Cranium, Myelofibrosis	(49)	(50)	(50) 1 [2.0] 1 [3.0]	(50) 1 [2.0]	
Skeletal Muscle	(0)	(0)	(1)	(1)	
NERVOUS SYSTEM					
Brain Hemorrhage	(49)	(50)	(50)	(50) 1 [3.0]	
Infiltration Cellular, Mononuclear Cell	1 [1.0]		4 70 01	1 [0.0]	
Necrosis Hippocampus, Gliosis			1 [3.0] 1 [2.0]		
RESPIRATORY SYSTEM					
Lung Inflammation	(49) 2 [1.5]	(50) 4 [1.0]	(50) 2 [2.5]	(50)	
Mineralization Pigmentation	2 [1.0]	4 [1.0]	2 [2.5] 1 [2.0]	1 [2.0]	
Alveolar Epithelium, Hyperplasia Mediastinum, Serosa, Inflammation	1 [2.0]	3 [1.3]	2 [1.5]	7 [2:0] 7 [1.7] 1 [2.0]	
Nose	(49)	(50)	(50)	(50)	
Inflammation Ulcer	2 [1.5]	1 [1.0]	4 [1.8]	27 [1.8] 1 [1.0]	
Glands, Cyst	1 [2.0]				

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

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

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

Pulegone

CAS Number: 89-82-7

Date Report Requested: 07/23/2008

Time Report Requested: 09:18:35 First Dose M/F: 04/15/03 / 04/14/03

B6C3F1 MICE FEMALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG
Nerve, Atrophy		1 [2.0]	2 [1.0]	49 [2.7]
Olfactory Epithelium, Degeneration		5 [1.8]	22 [1.6]	48 [1.9]
Olfactory Epithelium, Metaplasia	1 [1.0]	2 [1.0]	4 [1.8]	49 [2.8]
Respiratory Epithelium, Metaplasia	1 [1.0]	,	,	
PECIAL SENSES SYSTEM				
Eye	(49)	(50)	(49)	(50)
Atrophy		2 [3.0]	1 [3.0]	3 [3.7]
Cataract	1 [3.0]			1 [2.0]
Anterior Chamber, Inflammation				2 [2.0]
Choroid, Sclera, Inflammation			1 [3.0]	1 [3.0]
Cornea, Inflammation	2 [2.0]			9 [2.2]
Cornea, Pigmentation	4 [0 0]			1 [1.0]
Retina, Degeneration Harderian Gland	1 [3.0] (49)	(50)	(40)	(50)
Cyst	1 [2.0]	(50)	(49)	(50)
Infiltration Cellular, Mononuclear Cell	30 [1.0]	22 [1.0]	17 [1.1]	15 [1.1]
Inflammation, Chronic Active	30 [1.0]	22 [1.0]	17 [1.1]	1 [3.0]
Epithelium, Hyperplasia	3 [2.3]			4 [1.8]
RINARY SYSTEM				
Kidney	(49)	(50)	(50)	(50)
Accumulation, Hyaline Droplet	2 [2.5]			
Cyst			2 [2.0]	
Glomerulopathy, Hyaline		3 [1.0]	15 [1.3]	41 [1.5]
Infarct	3 [2.0]	2 [1.5]	1 [3.0]	1 [2.0]
Metaplasia, Osseous	1 [2.0]	2 [2.0]	2 [1.5]	4 [1.5]
Mineralization Nephropathy	1 [1.0] 13 [1.4]	19 [1.2]	3 [1.0] 12 [1.2]	20 [1.0]
Pigmentation	13 [1.4] 1 [1.0]	19 [1.2]	12 [1.2]	25 [1.2]
Vacuolization Cytoplasmic	1 [1.0]	1 [3.0]	2 [2.0]	
Glomerulus, Congestion	5 [1.2]	2 [1.0]	12 [1.1]	37 [1.3]
Papilla, Mineralization	1 [3.0]	-[]	[]	o. [o]
Papilla, Necrosis	3 [3.0]	1 [3.0]	1 [2.0]	
Pelvis, Dilatation	- []	1 [2.0]	r -1	
Urinary Bladder	(49)	(49)	(50)	(50)
Infiltration Cellular, Mononuclear Cell	37 [1.2]	33 [1.1]	30 [1.2]	30 [1.4]

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

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

Test Type: CHRONIC

Species/Strain: MICE/B6C3F1

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b]

Pulegone

CAS Number: 89-82-7

Date Report Requested: 07/23/2008

First Dose M/F: 04/15/03 / 04/14/03

Time Report Requested: 09:18:35

Lab: BAT

B6C3F1 MICE FEMALE	0 MG/KG	37.5 MG/KG	75 MG/KG	150 MG/KG	
Artery, Inflammation, Chronic Active		1 [3.0]			

*** END OF REPORT ***

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

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