F1_R2

C Number:	C99031
Lock Date:	05/21/2008
Cage Range:	ALL
Date Range:	ALL
Reasons For Removal:	ALL
Removal Date Range:	ALL
Treatment Groups:	Include ALL
Study Gender:	Both
TDMSE Version:	2.3.0
PWG Approval Date:	NONE

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Disposition Summary				
Animals Initially In Study	**50**	**50**	**50**	**50**
Early Deaths				
Moribund Sacrifice	**7**	**8**	**10**	**14**
Natural Death	**5**	**5**	**9**	**20**
Survivors				
Terminal Sacrifice	**38**	**37**	**31**	**16**
Animals Examined Microscopically	**50**	**50**	**50**	**50**
ALIMENTARY SYSTEM				
Esophagus	(50)	(50)	(50)	(50)
Intestine Large, Cecum	(50)	(49)	(50)	(50)
Inflammation, Chronic				1 [3.0]
Thrombosis	1 [4.0]			
Ulcer	1 [4.0]			
Intestine Large, Colon	(50)	(50)	(50)	(50)
Parasite Metazoan			2 [1.0]	1 [1.0]
Intestine Large, Rectum	(50)	(50)	(50)	(50)
Inflammation, Acute				1 [2.0]
Parasite Metazoan	3 [1.0]	8 [1.4]	5 [1.6]	5 [1.0]
Intestine Small, Duodenum	(50)	(50)	(50)	(50)
Intestine Small, Ileum	(50)	(49)	(50)	(50)
Intestine Small, Jejunum	(50)	(50)	(50)	(50)
Liver	(50)	(50)	(50)	(50)
Angiectasis	2 [1.0]	5 [1.0]	2 [1.0]	3 [1.0]
Basophilic Focus	30	37	23	22
Clear Cell Focus	31	30	18	11
Congestion	1 [3.0]			
Degeneration, Cystic	4 [1.0]	14 [1.1]	10 [1.0]	14 [1.1]
Eosinophilic Focus	14	21	19	21
Fatty Change, Focal	3			
Fibrosis		1 [1.0]		
Hepatodiaphragmatic Nodule	2	3	3	
Inflammation, Chronic	44 [1.0]	41 [1.0]	30 [1.1]	32 [1.2]

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)

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Mixed Cell Focus	19	32	24	17
Necrosis	1 [3.0]	4 [1.5]	6 [2.0]	7 [2.0]
Thrombosis				1 [3.0]
Bile Duct, Hyperplasia	32 [1.1]	43 [1.5]	46 [2.0]	46 [2.0]
Hepatocyte, Fatty Change	27 [1.4]	18 [1.3]	23 [1.4]	31 [1.5]
Hepatocyte, Hypertrophy	1 [1.0]	17 [1.4]	26 [2.1]	27 [2.7]
Oval Cell, Hyperplasia		1 [2.0]	1 [1.0]	10 [1.8]
Mesentery	(4)	(4)	(8)	(5)
Thrombosis				1 [3.0]
Fat, Necrosis	2 [3.5]	2 [2.0]	5 [3.2]	3 [3.0]
Pancreas	(50)	(50)	(50)	(50)
Inflammation, Chronic		1 [3.0]	1 [2.0]	
Mineralization	1 [2.0]			
Necrosis				1 [3.0]
Acinus, Atrophy	22 [1.3]	23 [1.4]	14 [1.2]	20 [1.2]
Acinus, Hyperplasia	11 [1.5]	7 [1.0]	5 [2.2]	12 [1.8]
Duct, Cyst	3 [1.3]	5 [1.0]	10 [1.0]	3 [1.0]
Salivary Glands	(50)	(50)	(50)	(50)
Stomach, Forestomach	(50)	(50)	(50)	(50)
Inflammation, Chronic	4 [3.0]	1 [3.0]	6 [3.2]	6 [2.8]
Mineralization			1 [4.0]	
Ulcer	2 [3.0]		4 [3.0]	4 [2.8]
Stomach, Glandular	(50)	(50)	(50)	(50)
Inflammation, Suppurative		1 [2.0]		
Inflammation, Chronic		1 [2.0]	1 [4.0]	
Mineralization	1 [4.0]	1 [3.0]	1 [3.0]	6 [2.3]
Ulcer	2 [2.5]			2 [2.0]
Epithelium, Hyperplasia	1 [1.0]			
Glands, Ectasia	17 [1.0]	30 [1.0]	19 [1.1]	12 [1.0]
Glands, Hyperplasia		2 [2.0]		1 [2.0]
CARDIOVASCULAR SYSTEM				
Blood Vessel	(50)	(50)	(50)	(50)
Thrombosis				1 [4.0]

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

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Aorta, Mineralization	1 [3.0]			2 [3.0]
Pulmonary Vein, Mineralization	1 [1.0]			
Heart	(50)	(50)	(50)	(50)
Cardiomyopathy	50 [1.9]	48 [2.0]	49 [1.8]	50 [1.8]
Atrium, Thrombosis	3 [2.7]		1 [3.0]	3 [3.0]
Myocardium, Mineralization	1 [3.0]			2 [2.0]
Valve, Inflammation, Chronic			1 [3.0]	
ENDOCRINE SYSTEM				
Adrenal Cortex	(50)	(50)	(50)	(50)
Cytoplasmic Alteration		1 [2.0]		
Degeneration, Cystic	2 [2.0]	4 [1.3]		
Hyperplasia, Focal	12 [1.5]	9 [1.3]	9 [1.2]	8 [1.5]
Hypertrophy, Focal	7 [1.9]	10 [1.8]	9 [1.9]	3 [1.7]
Hypertrophy, Diffuse	3 [1.7]	3 [2.0]	5 [1.8]	11 [2.1]
Infiltration Cellular, Mononuclear Cell		1 [2.0]		
Inflammation, Chronic				1 [2.0]
Necrosis		1 [2.0]	2 [2.5]	1 [2.0]
Vacuolization Cytoplasmic, Focal	17 [1.1]	17 [1.1]	16 [1.3]	13 [1.4]
Vacuolization Cytoplasmic, Diffuse	18 [1.2]	25 [1.2]	33 [1.3]	34 [1.3]
Adrenal Medulla	(50)	(50)	(50)	(50)
Hyperplasia	15 [1.8]	16 [1.6]	8 [1.9]	10 [1.9]
Necrosis			1 [4.0]	
Islets, Pancreatic	(50)	(50)	(50)	(50)
Hyperplasia	3 [1.0]	2 [1.0]	5 [2.0]	2 [2.5]
Parathyroid Gland	(47)	(48)	(48)	(47)
Hyperplasia	1 [2.0]	3 [2.3]	2 [2.5]	7 [2.3]
Pituitary Gland	(50)	(50)	(50)	(50)
Angiectasis			1 [2.0]	1 [2.0]
Cyst	5 [1.2]	4 [1.3]	7 [1.6]	8 [1.5]
Pars Distalis, Hyperplasia	19 [1.5]	18 [1.7]	18 [1.4]	16 [1.5]
Pars Intermedia, Hyperplasia	1 [2.0]			1 [2.0]
Thyroid Gland	(50)	(50)	(49)	(45)
C-cell, Hyperplasia	35 [1.4]	29 [1.4]	27 [1.5]	24 [1.0]

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

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Follicle, Hyperplasia		7 [1.3]	9 [2.3]	5 [2.8]
Follicular Cell, Hypertrophy	13 [1.0]	37 [1.2]	41 [1.3]	41 [1.8]
GENERAL BODY SYSTEM				
None				
GENITAL SYSTEM				
Coagulating Gland	(0)	(0)	(1)	(0)
Inflammation, Chronic			1 [2.0]	
Epididymis	(50)	(50)	(50)	(50)
Granuloma Sperm		1 [2.0]	3 [1.3]	
Inflammation, Chronic				1 [1.0]
Mineralization		1 [2.0]		
Preputial Gland	(50)	(50)	(50)	(50)
Cyst	2 [2.0]			
Hyperplasia	2 [2.0]	1 [1.0]		
Inflammation, Chronic	49 [1.7]	48 [1.6]	47 [1.7]	49 [1.9]
Metaplasia, Cartilagenous			1 [2.0]	
Prostate	(50)	(50)	(50)	(50)
Hyperplasia	6 [1.2]	3 [1.0]	3 [1.0]	4 [1.3]
Inflammation	36 [1.8]	27 [1.7]	25 [1.9]	32 [2.0]
Necrosis	1 [4.0]			
Seminal Vesicle	(50)	(50)	(50)	(50)
Inflammation	2 [2.5]	3 [2.0]	2 [4.0]	4 [2.3]
Testes	(50)	(50)	(50)	(50)
Germinal Epithelium, Atrophy	1 [4.0]		1 [4.0]	1 [4.0]
Germinal Epithelium, Necrosis			1 [1.0]	
Interstitial Cell, Hyperplasia				1 [3.0]
HEMATOPOIETIC SYSTEM				
Bone Marrow	(50)	(50)	(48)	(50)

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)

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Hyperplasia	14 [1.9]	15 [1.9]	21 [2.4]	26 [2.3]
Necrosis			1 [2.0]	
Lymph Node	(2)	(2)	(9)	(10)
Deep Cervical, Hyperplasia, Plasma Cell				1 [2.0]
Mediastinal, Ectasia			1 [4.0]	1 [3.0]
Mediastinal, Hyperplasia, Lymphoid		1 [2.0]	1 [2.0]	3 [2.7]
Mediastinal, Hyperplasia, Plasma Cell				1 [3.0]
Mediastinal, Infiltration Cellular, Histiocyte		1 [2.0]		
Mediastinal, Inflammation, Suppurative	1 [3.0]			
Pancreatic, Necrosis				1 [2.0]
Renal, Hyperplasia, Lymphoid				1 [3.0]
Lymph Node, Mandibular	(0)	(0)	(1)	(0)
Ectasia			1 [4.0]	
Lymph Node, Mesenteric	(50)	(50)	(50)	(50)
Ectasia			1 [2.0]	
Hyperplasia, Lymphoid		1 [3.0]		
Infiltration Cellular, Histiocyte	1 [2.0]	1 [2.0]		
Necrosis		1 [3.0]	1 [3.0]	1 [2.0]
Spleen	(50)	(50)	(50)	(50)
Fibrosis	1 [1.0]	1 [2.0]		
Hematopoietic Cell Proliferation	3 [1.7]	8 [1.8]	4 [2.5]	6 [2.8]
Hemorrhage		1 [2.0]		
Necrosis		1 [2.0]	1 [3.0]	1 [2.0]
Lymphoid Follicle, Atrophy	2 [2.5]	1 [4.0]	4 [1.8]	14 [2.8]
Thymus	(47)	(50)	(47)	(47)
Atrophy	47 [3.6]	48 [3.9]	46 [3.9]	46 [3.9]
Epithelial Cell, Hyperplasia	1 [2.0]	1 [3.0]		1 [2.0]
INTEGUMENTARY SYSTEM				
Mammary Gland	(50)	(50)	(50)	(50)
Hyperplasia				1 [4.0]
Inflammation, Chronic		1 [1.0]		
Duct, Cyst	12 [1.6]	19 [1.5]	13 [1.5]	12 [1.3]
Skin	(50)	(50)	(50)	(50)

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)

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Cyst Epithelial Inclusion		1		
Lymphatic, Subcutaneous Tissue, Cyst	1 [2.0]	1 [3.0]		
Sebaceous Gland, Cyst	1 [2.0]			
Subcutaneous Tissue, Hemorrhage				1 [3.0]
MUSCULOSKELETAL SYSTEM				
Bone	(50)	(50)	(50)	(50)
Hyperostosis		1 [2.0]	1 [2.0]	1 [2.0]
NERVOUS SYSTEM				
Brain	(50)	(50)	(50)	(50)
Necrosis		1 [2.0]		
Hippocampus, Necrosis				1 [1.0]
Peripheral Nerve	(0)	(0)	(0)	(1)
Axon, Degeneration				1 [2.0]
Spinal Cord	(0)	(0)	(0)	(1)
RESPIRATORY SYSTEM				
Lung	(50)	(50)	(50)	(50)
Inflammation, Acute				1 [3.0]
Inflammation, Chronic	19 [1.1]	14 [1.4]	12 [1.4]	8 [1.8]
Metaplasia, Osseous	1 [1.0]	1 [1.0]	2 [1.0]	
Thrombosis			1 [3.0]	2 [3.5]
Alveolar Epithelium, Hyperplasia	8 [1.9]	4 [1.0]	2 [2.0]	5 [1.4]
Bronchiole, Hyperplasia	1 [4.0]			
Bronchus, Hyperplasia	1 [2.0]			
Nose	(50)	(49)	(49)	(50)
Foreign Body	11	5	9	5
Hemorrhage			1 [2.0]	
Inflammation, Chronic Active	33 [1.2]	32 [1.3]	38 [1.9]	46 [2.2]
Respiratory Metaplasia	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)

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Synechia, Focal	1 [2.0]		1 [2.0]	
Thrombosis			1 [1.0]	
Glands, Hyperplasia	1 [2.0]			
Goblet Cell, Respiratory Epithelium, Hyperplasia	20 [1.5]	18 [1.2]	41 [1.7]	34 [2.1]
Goblet Cell, Transitional Epithelium, Hyperplasia			1 [2.0]	
Nasolacrimal Duct, Inflammation, Chronic Active	1 [2.0]			
Nerve, Olfactory Epithelium, Atrophy		17 [1.4]	14 [2.1]	23 [2.5]
Olfactory Epithelium, Accumulation, Hyaline Droplet	45 [1.9]	43 [1.4]	14 [1.1]	
Olfactory Epithelium, Atrophy	1 [1.0]	26 [1.3]	37 [1.6]	31 [2.2]
Olfactory Epithelium, Foreign Body			1	
Olfactory Epithelium, Hyperplasia				1 [2.0]
Olfactory Epithelium, Pigmentation		39 [1.5]	42 [1.7]	30 [2.1]
Olfactory Epithelium, Respiratory Metaplasia	9 [1.3]	30 [1.5]	40 [2.0]	32 [2.5]
Respiratory Epithelium, Hyperplasia	14 [1.6]	28 [1.4]	45 [1.8]	35 [2.2]
Respiratory Epithelium, Inflammation, Chronic	1 [1.0]			
Submucosa, Fibrosis				8 [1.6]
Transitional Epithelium, Hyperplasia	2 [1.5]	18 [1.2]	43 [1.9]	31 [2.5]
Pleura	(0)	(0)	(1)	(0)
Trachea	(50)	(50)	(50)	(50)
Inflammation, Chronic	1 [1.0]			2 [2.0]
SPECIAL SENSES SYSTEM				
Eye	(50)	(50)	(50)	(50)
Cataract	1 [3.0]	4 [2.5]		
Degeneration		1 [3.0]		
Anterior Chamber, Inflammation, Suppurative				2 [3.5]
Retina, Atrophy		1 [3.0]		
Harderian Gland	(50)	(50)	(50)	(50)
Atrophy		1 [1.0]		
Inflammation		2 [1.0]		
Zymbal's Gland	(1)	(0)	(0)	(1)

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

FISCHER 344 RATS MALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
URINARY SYSTEM				
Kidney	(50)	(50)	(50)	(50)
Hydronephrosis	1 [4.0]			1 [3.0]
Infarct			1 [2.0]	
Nephropathy	49 [1.7]	49 [2.0]	49 [2.4]	50 [2.9]
Papilla, Necrosis	3 [2.0]	1 [2.0]		1 [4.0]
Pelvis, Inflammation, Acute				1 [4.0]
Renal Tubule, Cyst	1 [2.0]	1 [2.0]	1 [2.0]	1 [2.0]
Renal Tubule, Hyperplasia	1 [1.0]			
Urinary Bladder	(50)	(50)	(50)	(50)
Inflammation, Acute				2 [2.5]
Inflammation, Chronic	2 [2.0]	1 [2.0]		2 [3.0]
Mineralization				1 [3.0]
Necrosis	1 [3.0]			
Ulcer				2 [3.0]

*** 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)

FISCHER 344 RATS FEMALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Disposition Summary				
Animals Initially In Study	**50**	**50**	**50**	**50**
Early Deaths				
Dosing Accident				**1**
Moribund Sacrifice	**9**	**13**	**7**	**4**
Natural Death	**4**	**10**	**6**	**13**
Survivors				
Natural Death			**1**	
Terminal Sacrifice	**37**	**27**	**36**	**32**
Animals Examined Microscopically	**50**	**50**	**50**	**50**
ALIMENTARY SYSTEM				
Esophagus	(50)	(50)	(50)	(50)
Intestine Large, Cecum	(50)	(50)	(50)	(49)
Inflammation, Chronic			2 [2.0]	
Intestine Large, Colon	(50)	(50)	(50)	(49)
Parasite Metazoan	3 [1.0]	2 [1.5]	2 [1.0]	
Intestine Large, Rectum	(50)	(50)	(50)	(50)
Parasite Metazoan	8 [1.5]	4 [1.3]	3 [1.3]	6 [1.2]
Intestine Small, Duodenum	(50)	(50)	(50)	(49)
Intestine Small, Ileum	(50)	(50)	(50)	(49)
Intestine Small, Jejunum	(50)	(50)	(50)	(49)
Liver	(50)	(50)	(50)	(50)
Angiectasis	7 [1.0]	2 [1.0]	4 [1.3]	4 [1.0]
Basophilic Focus	48	43	44	40
Clear Cell Focus	20	7	17	11
Degeneration, Cystic	1 [1.0]		2 [1.0]	
Eosinophilic Focus	24	30	38	30
Fatty Change, Focal	11	25	30	25
Hepatodiaphragmatic Nodule	6	7	1	3
Inflammation, Chronic	45 [1.0]	41 [1.0]	41 [1.1]	42 [1.1]
Mixed Cell Focus	24	12	17	10
Necrosis	5 [1.4]	2 [2.5]	3 [1.7]	3 [3.0]
Bile Duct, Hyperplasia	11 [1.0]	31 [1.1]	31 [1.1]	33 [1.1]

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

FISCHER 344 RATS FEMALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Hepatocyte, Fatty Change	14 [1.6]	7 [1.4]	7 [1.7]	9 [1.3]
Hepatocyte, Hypertrophy	7 [1.4]	15 [1.7]	27 [2.2]	33 [2.5]
Ito Cell, Hyperplasia		1 [1.0]		
Mesentery	(10)	(2)	(8)	(11)
Fat, Necrosis	10 [2.4]	2 [2.5]	8 [2.3]	11 [2.7]
Oral Mucosa	(0)	(0)	(1)	(0)
Pancreas	(50)	(50)	(50)	(48)
Hemorrhage				1 [2.0]
Acinus, Atrophy	17 [1.1]	13 [1.0]	12 [1.2]	20 [1.1]
Acinus, Hyperplasia	5 [1.4]		2 [1.5]	
Duct, Cyst	3 [1.0]		1 [1.0]	4 [1.0]
Duct, Hyperplasia		1 [1.0]		
Salivary Glands	(50)	(50)	(50)	(50)
Stomach, Forestomach	(50)	(50)	(50)	(50)
Inflammation, Chronic	1 [3.0]	1 [3.0]	1 [3.0]	1 [4.0]
Ulcer	1 [3.0]	1 [3.0]	2 [3.0]	2 [3.0]
Stomach, Glandular	(50)	(50)	(50)	(50)
Inflammation, Chronic		2 [2.0]		
Mineralization	1 [1.0]	2 [2.0]		1 [1.0]
Ulcer		2 [2.5]	3 [2.7]	
Glands, Ectasia	35 [1.0]	37 [1.0]	37 [1.1]	36 [1.1]
Glands, Hyperplasia	1 [2.0]		1 [1.0]	
CARDIOVASCULAR SYSTEM				
Blood Vessel	(50)	(50)	(50)	(50)
Aorta, Mineralization		1 [1.0]		
Heart	(50)	(50)	(50)	(50)
Cardiomyopathy	49 [1.6]	46 [1.5]	46 [1.5]	48 [1.5]
Atrium, Thrombosis		1 [4.0]		1 [2.0]
Myocardium, Mineralization				1 [1.0]
ENDOCRINE SYSTEM				
Adrenal Cortex	(50)	(50)	(50)	(50)

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)

FISCHER 344 RATS FEMALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Degeneration, Cystic	12 [1.5]	5 [1.6]	7 [1.0]	7 [1.6]
Hyperplasia, Focal	12 [1.6]	10 [1.6]	13 [1.5]	15 [1.4]
Hyperplasia, Diffuse				1 [1.0]
Hypertrophy, Focal	6 [1.7]	2 [2.5]	2 [1.5]	5 [1.8]
Hypertrophy, Diffuse	3 [1.0]	10 [1.4]	7 [1.0]	9 [1.3]
Vacuolization Cytoplasmic, Focal	23 [1.0]	14 [1.2]	22 [1.1]	21 [1.1]
Vacuolization Cytoplasmic, Diffuse	4 [1.3]	5 [1.4]	6 [1.5]	4 [1.5]
Capsule, Fibrosis	1 [1.0]			
Adrenal Medulla	(50)	(50)	(50)	(50)
Hyperplasia	3 [1.7]	3 [1.0]	5 [1.2]	2 [1.5]
Islets, Pancreatic	(50)	(50)	(50)	(50)
Hyperplasia	1 [1.0]			
Parathyroid Gland	(38)	(47)	(43)	(35)
Hyperplasia	1 [3.0]			
Pituitary Gland	(50)	(50)	(50)	(50)
Angiectasis		2 [2.0]		
Cyst	26 [1.6]	24 [1.4]	29 [1.7]	31 [1.5]
Pars Distalis, Cyst	1			
Pars Distalis, Hyperplasia	11 [2.2]	13 [1.9]	18 [1.8]	23 [1.9]
Thyroid Gland	(49)	(50)	(49)	(49)
C-cell, Hyperplasia	40 [1.2]	38 [1.3]	36 [1.3]	21 [1.3]
Follicle, Hyperplasia	3 [1.3]	3 [1.0]	1 [2.0]	5 [1.6]
Follicular Cell, Hypertrophy	15 [1.0]	41 [1.0]	45 [1.1]	48 [2.0]
GENERAL BODY SYSTEM				
None				
GENITAL SYSTEM				
Clitoral Gland	(50)	(50)	(50)	(50)
Atrophy			1 [3.0]	
Cyst			1 [1.0]	
Hyperplasia	3 [1.3]	4 [1.3]	3 [1.0]	2 [2.0]
Inflammation, Chronic	36 [1.2]	39 [1.3]	40 [1.4]	38 [1.3]

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)

FISCHER 344 RATS FEMALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Mineralization		1 [2.0]		
Ovary	(50)	(50)	(50)	(50)
Cyst	3 [2.3]	2 [2.0]	2 [2.0]	3 [2.0]
Uterus	(50)	(50)	(50)	(50)
Cyst	4 [2.0]			4 [2.0]
Decidual Reaction			1 [2.0]	
Inflammation, Suppurative	2 [2.5]	2 [1.0]		5 [1.6]
Necrosis		1 [3.0]		
Endometrium, Hyperplasia, Cystic	3 [2.0]	13 [1.6]	11 [1.5]	7 [1.3]
Vagina	(0)	(2)	(0)	(0)
HEMATOPOIETIC SYSTEM				
Bone Marrow	(50)	(50)	(50)	(50)
Hemorrhage	1 [2.0]			
Hyperplasia	7 [3.1]	11 [2.7]	9 [2.4]	12 [2.5]
Necrosis			1 [2.0]	
Lymph Node	(3)	(3)	(3)	(4)
Ectasia				1 [3.0]
Deep Cervical, Hyperplasia, Lymphoid		1 [2.0]		
Mediastinal, Ectasia				1 [2.0]
Mediastinal, Hemorrhage		1 [2.0]		
Mediastinal, Hyperplasia, Lymphoid			1 [3.0]	3 [2.3]
Pancreatic, Congestion	1 [3.0]			
Lymph Node, Mesenteric	(50)	(50)	(50)	(49)
Pigmentation, Hemosiderin				1 [2.0]
Spleen	(50)	(50)	(50)	(50)
Hematopoietic Cell Proliferation	14 [1.8]	18 [1.6]	13 [1.4]	13 [1.4]
Hyperplasia, Histiocytic	1 [4.0]	1 [4.0]	1 [4.0]	2 [4.0]
Infarct		1 [3.0]		
Necrosis			1 [2.0]	
Thrombosis		1 [3.0]		
Lymphoid Follicle, Atrophy	9 [2.1]	6 [2.2]	9 [2.1]	5 [2.6]
Thymus	(50)	(48)	(49)	(48)
Atrophy	48 [3.8]	47 [3.9]	46 [4.0]	47 [3.8]

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)

FISCHER 344 RATS FEMALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Ectopic Parathyroid Gland		1 [1.0]		
Ectopic Thyroid		1 [1.0]		1 [2.0]
INTEGUMENTARY SYSTEM				
Mammary Gland	(50)	(50)	(50)	(50)
Duct, Cyst	14 [1.5]	10 [1.6]	8 [1.5]	6 [1.3]
Skin	(50)	(50)	(50)	(50)
MUSCULOSKELETAL SYSTEM				
Bone	(50)	(50)	(50)	(50)
Hyperostosis				1 [2.0]
Skeletal Muscle	(0)	(0)	(1)	(0)
NERVOUS SYSTEM				
Brain	(50)	(50)	(50)	(50)
Congestion				1 [1.0]
Hydrocephalus	1 [4.0]			
Artery, Meninges, Inflammation, Chronic		1 [1.0]		
Hippocampus, Necrosis				1 [2.0]
Hypothalamus, Congestion		1 [1.0]		
Peripheral Nerve	(1)	(0)	(0)	(0)
Spinal Cord	(1)	(0)	(0)	(0)
Congestion	1 [2.0]			
RESPIRATORY SYSTEM				
Larynx	(0)	(0)	(0)	(1)
Foreign Body				1 [2.0]
Lung	(50)	(50)	(50)	(50)
Cyst	1			

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

FISCHER 344 RATS FEMALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Inflammation, Suppurative			1 [3.0]	
Inflammation, Chronic	7 [1.0]	3 [1.0]	11 [1.3]	6 [1.5]
Inflammation, Chronic Active				1 [2.0]
Metaplasia, Osseous	2 [1.0]	1 [1.0]		1 [1.0]
Thrombosis			1 [1.0]	
Alveolar Epithelium, Hyperplasia	3 [1.0]		3 [1.0]	2 [1.5]
Nose	(49)	(49)	(50)	(46)
Foreign Body	1	1	1	7
Inflammation, Chronic Active	22 [1.0]	16 [1.2]	26 [1.5]	38 [1.9]
Goblet Cell, Olfactory Epithelium, Hyperplasia		1 [1.0]		
Goblet Cell, Respiratory Epithelium, Hyperplasia	6 [1.2]	2 [1.0]	18 [1.6]	35 [1.8]
Nerve, Olfactory Epithelium, Atrophy		15 [1.1]	22 [1.6]	33 [2.2]
Olfactory Epithelium, Accumulation, Hyaline Droplet	44 [2.0]	39 [1.7]	25 [1.2]	
Olfactory Epithelium, Atrophy		18 [1.1]	25 [1.6]	37 [2.1]
Olfactory Epithelium, Degeneration		1 [1.0]		
Olfactory Epithelium, Hemorrhage	1 [1.0]			
Olfactory Epithelium, Hyperplasia				1 [1.0]
Olfactory Epithelium, Pigmentation		37 [1.5]	43 [2.0]	40 [1.9]
Olfactory Epithelium, Regeneration			1 [1.0]	2 [1.0]
Olfactory Epithelium, Respiratory Metaplasia	8 [1.3]	4 [1.3]	32 [2.0]	37 [2.5]
Respiratory Epithelium, Hyperplasia	9 [1.2]	9 [1.3]	19 [1.7]	34 [2.3]
Transitional Epithelium, Hyperplasia		6 [1.5]	32 [1.8]	36 [2.8]
Trachea	(50)	(50)	(50)	(50)
Inflammation, Chronic			1 [1.0]	1 [2.0]
SPECIAL SENSES SYSTEM				
Eye	(50)	(50)	(50)	(50)
Cataract		2 [3.0]	3 [2.3]	2 [2.0]
Degeneration			1 [3.0]	
Retinal Detachment				1 [3.0]
Retina, Atrophy		2 [3.0]	1 [3.0]	
Retina, Hemorrhage				1 [3.0]
Retrobulbar, Inflammation, Chronic			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)

FISCHER 344 RATS FEMALE	0 mg/kg	100 mg/kg	300 mg/kg	1000 mg/kg
Harderian Gland	(50)	(50)	(50)	(50)
Hyperplasia				1 [1.0]
Inflammation, Chronic	4 [1.0]	6 [1.3]	8 [1.0]	7 [1.1]
URINARY SYSTEM				
Kidney	(50)	(50)	(50)	(50)
Hydronephrosis			1 [2.0]	
Infarct	2 [3.0]	2 [1.5]	1 [2.0]	6 [2.0]
Inflammation, Chronic	2 [3.5]		1 [3.0]	
Nephropathy	41 [1.1]	42 [1.0]	43 [1.2]	42 [1.2]
Renal Tubule, Cyst			1 [2.0]	
Renal Tubule, Hyperplasia	3 [1.3]	1 [1.0]	3 [2.0]	
Renal Tubule, Mineralization	1 [2.0]	1 [1.0]		1 [2.0]
Renal Tubule, Pigmentation, Lipofuscin	1 [2.0]			
Urinary Bladder	(50)	(50)	(50)	(49)
Inflammation, Acute	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)