Species/Strain: Mouse/P53(C57BL/6)

Test Type: 25-WEEK

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH Date Report Requested: 10/20/2014 AVERAGE SEVERITY GRADES[b]

Test Compound: Transgenic model evaluation (Cyclosporin A)

CAS Number: 59865-13-3

Time Report Requested: 17:43:33

First Dose M/F: NA / NA

Lab: MBA

C Number: C95007D

Lock Date: 06/02/1997

Cage Range: ΑII

Date Range: ΑII

Reasons For Removal: ΑII

Removal Date Range: ΑII

Treatment Groups: ΑII

Study Gender: Both

PWG Approval Date NONE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/20/2014 Time Report Requested: 17:43:33

Test Type: 25-WEEK Route: GAVAGE

Species/Strain: Mouse/P53(C57BL/6)

Test Compound: Transgenic model evaluation (Cyclosporin A)

CAS Number: 59865-13-3

Time Report Requested: 17:43:33

First Dose M/F: NA / NA

P53(C57BL/6) Mouse MALE	0 MG/KG	10 MG/KG	25 MG/KG
Disposition Summary			
Animals Initially In Study	15	15	15
Early Deaths			
Moribund Sacrifice	1		
Survivors			
Terminal Sacrifice	14	15	15
Animals Examined Microscopically	15	15	15
ALIMENTARY SYSTEM			
Liver	(15)	(15)	(15)
Centrilobular, Hepatocyte, Hypertrophy	2 [2.0]	, ,	, ,
Hepatocyte, Necrosis, Focal			1 [1.0]
Hepatocyte, Vacuolization Cytoplasmic		4 [3.3]	
Hepatocyte, Vacuolization Cytoplasmic, Diffuse	1 [3.0]	7 [3.3]	6 [2.8]
Hepatocyte, Vacuolization Cytoplasmic, Focal	4 [1.8]	4 [1.8]	4 [3.0]
Infiltration Cellular, Lymphocyte, Focal	2 [1.0]	1 [1.0]	
Necrosis, Focal	3 [1.0]		1 [1.0]
Salivary Glands	(15)	(15)	(15)
Infiltration Cellular, Lymphocyte, Focal	10 [1.0]	11 [1.2]	7 [1.0]
Stomach, Forestomach	(15)	(15)	(15)
Stomach, Glandular	(15)	(15)	(15)
Tooth	(1)	(0)	(0)
CARDIOVASCULAR SYSTEM			
None			
ENDOCRINE SYSTEM			
Adrenal Cortex	(15)	(15)	(15)
Adrenal Medulla	(15)	(15)	(15)

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: 25-WEEK

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/20/2014 Time Report Requested: 17:43:33

Test Compound: Transgenic model evaluation (Cyclosporin A)

Species/Strain: Mouse/P53(C57BL/6)

CAS Number: 59865-13-3

Time Report Requested: 17:43:33

First Dose M/F: NA / NA

Lab: MBA

P53(C57BL/6) Mouse MALE	0 MG/KG	10 MG/KG	25 MG/KG
Pituitary Gland	(12)	(15)	(12)
Thyroid Gland	(15)	(15)	(14)
GENERAL BODY SYSTEM			
None			
GENITAL SYSTEM			
Epididymis	(15)	(15)	(15)
Testes	(15)	(15)	(15)
Degeneration, Diffuse	1 [3.0]		
HEMATOPOIETIC SYSTEM			
Bone Marrow	(15)	(15)	(15)
Lymph Node, Mandibular	(15)	(15)	(14)
Hyperplasia, Lymphoid		2 [1.5]	1 [3.0]
Lymph Node, Mediastinal	(12)	(12)	(12)
Lymph Node, Mesenteric	(15)	(15)	(15)
Spleen	(15)	(15)	(15)
Hematopoietic Cell Proliferation	7 [1.4]	13 [1.2]	14 [1.4]
Thymus	(15)	(15)	(14)
INTEGUMENTARY SYSTEM			
Skin	(15)	(15)	(15)
Epidermis, Hyperplasia, Focal			2 [3.0]
Epidermis, Ulcer, Focal	1 [3.0]		1 [3.0]
MUSCULOSKELETAL SYSTEM			
Bone	(1)	(0)	(0)
NERVOUS SYSTEM			
None			

RESPIRATORY 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)

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/20/2014 Time Report Requested: 17:43:33

Test Type: 25-WEEK Route: GAVAGE

Test Compound: Transgenic model evaluation (Cyclosporin A)

Species/Strain: Mouse/P53(C57BL/6)

CAS Number: 59865-13-3

Time Report Requested: 17:43:33

First Dose M/F: NA / NA

P53(C57BL/6) Mouse MALE	0 MG/KG	10 MG/KG	25 MG/KG
Lung	(15)	(15)	(15)
Hyperplasia, Lymphoid, Focal	1 [1.0]	1 [1.0]	
Perivascular, Inflammation, Chronic Active	2 [1.0]		
SPECIAL SENSES SYSTEM None URINARY SYSTEM			,
Kidney	(15)	(15)	(15)
Infiltration Cellular, Lymphocyte, Focal			2 [1.5]
Renal Tubule, Degeneration, Focal			1 [1.0]
Renal Tubule, Regeneration, Focal			1 [3.0]

^{***}END OF MALE DATA***

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: 25-WEEK

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/20/2014 Time Report Requested: 17:43:33

Test Compound: Transgenic model evaluation (Cyclosporin A)

Species/Strain: Mouse/P53(C57BL/6)

CAS Number: 59865-13-3

Time Report Requested: 17:43:33

First Dose M/F: NA / NA

P53(C57BL/6) Mouse FEMALE	0 MG/KG	10 MG/KG	25 MG/KG
Disposition Summary			
Animals Initially In Study	15	15	15
Early Deaths			
Survivors			
Terminal Sacrifice	15	15	15
Animals Examined Microscopically	15	15	15
ALIMENTARY SYSTEM			
Liver	(15)	(15)	(15)
Hepatocyte, Vacuolization Cytoplasmic, Focal	2 [2.0]	2 [2.0]	6 [2.2]
Infiltration Cellular, Lymphocyte, Focal	3 [1.0]	6 [1.2]	2 [1.5]
Necrosis, Focal	2 [1.0]		1 [1.0]
Pancreas	(0)	(1)	(0)
Cyst		1 [3.0]	
Salivary Glands	(15)	(15)	(15)
Infiltration Cellular, Lymphocyte, Focal	6 [2.0]	8 [1.0]	12 [1.2]
Stomach, Forestomach	(15)	(15)	(15)
Stomach, Glandular	(15)	(15)	(15)
CARDIOVASCULAR SYSTEM None			
ENDOCRINE SYSTEM			
Adrenal Cortex	(15)	(15)	(15)
Necrosis, Diffuse		1 [4.0]	
Adrenal Medulla	(15)	(15)	(15)
Necrosis, Diffuse	•	1 [4.0]	
Pituitary Gland	(14)	(14)	(14)
Thyroid Gland	(14)	(15)	(14)

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: Mouse/P53(C57BL/6)

Test Type: 25-WEEK

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/20/2014 Time Report Requested: 17:43:33

Test Compound: Transgenic model evaluation (Cyclosporin A)

CAS Number: 59865-13-3

Time Report Requested: 17:43:33

First Dose M/F: NA / NA

Lab: MBA

P53(C57BL/6) Mouse FEMALE	0 MG/KG	10 MG/KG	25 MG/KG
GENERAL BODY SYSTEM None			
GENITAL SYSTEM			
Ovary	(15)	(15)	(15)
Uterus	(15)	(15)	(15)
Hydrometra		1 [2.0]	
Hyperplasia, Cystic	1 [2.0]		
HEMATOPOIETIC SYSTEM			
Bone Marrow	(15)	(15)	(15)
Myeloid Cell, Hyperplasia		1 [2.0]	
Lymph Node	(2)	(2)	(1)
Iliac, Hyperplasia, Lymphoid		1 [2.0]	
Lumbar, Hyperplasia, Lymphoid	1 [2.0]	1 [1.0]	
Mediastinal, Pigmentation	1 [2.0]		
Renal, Hyperplasia, Lymphoid	1 [2.0]	1 [2.0]	
Lymph Node, Mandibular	(15)	(15)	(15)
Hyperplasia, Lymphoid		3 [1.7]	
Lymph Node, Mediastinal	(15)	(12)	(15)
Pigmentation	1 [2.0]		
Lymph Node, Mesenteric	(15)	(15)	(15)
Hyperplasia, Lymphoid		1 [2.0]	
Inflammation, Chronic, Focal			1 [2.0]
Spleen	(15)	(15)	(15)
Hematopoietic Cell Proliferation	15 [1.5]	15 [1.4]	14 [1.4]
Pigmentation	3 [1.0]	3 [1.0]	2 [1.0]
Thymus	(15)	(15)	(15)

INTEGUMENTARY 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)

Test Type: 25-WEEK

Route: GAVAGE

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/20/2014 Time Report Requested: 17:43:33

Test Compound: Transgenic model evaluation (Cyclosporin A)

Species/Strain: Mouse/P53(C57BL/6)

CAS Number: 59865-13-3

Time Report Requested: 17:43:33

First Dose M/F: NA / NA

P53(C57BL/6) Mouse FEMALE	0 MG/KG	10 MG/KG	25 MG/KG
Skin	(15)	(15)	(15)
MUSCULOSKELETAL SYSTEM			
None			
NERVOUS SYSTEM			
None			
RESPIRATORY SYSTEM			
Lung	(15)	(15)	(15)
Hemorrhage, Focal		1 [1.0]	
Hyperplasia, Lymphoid	1 [1.0]	2 [2.0]	
Hyperplasia, Lymphoid, Focal		1 [1.0]	2 [1.0]
Infiltration Cellular, Lymphocyte, Focal			5 [1.2]
Perivascular, Hyperplasia, Lymphoid		1 [2.0]	
Perivascular, Inflammation, Chronic Active		1 [2.0]	
SPECIAL SENSES SYSTEM			
Eye	(0)	(0)	(1)
Developmental Malformation			1
URINARY SYSTEM			
Kidney	(15)	(15)	(15)
Infiltration Cellular, Lymphocyte, Focal			1 [1.0]
Nephropathy			1 [2.0]
Pelvis, Dilatation		1 [4.0]	
Renal Tubule, Degeneration, Focal			1 [1.0]

^{**} 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)