Route: DOSED FEED

Species/Strain: Rat/CD

P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH Date Report Requested: 10/23/2014 AVERAGE SEVERITY GRADES[b]

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:18:58

First Dose M/F: NA / NA

Lab: NCTR

C Number: MG96006

Lock Date: 12/02/2008

**Cage Range:** ΑII

**Date Range:** ΑII

**Reasons For Removal:** ΑII

**Removal Date Range:** ΑII

**Treatment Groups:** ΑII

**Study Gender:** Both

**PWG Approval Date NONE** 

Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:58

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:58

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Disposition Summary						
Animals Initially In Study	51	50	50	50	50	50
Early Deaths						
Moribund Sacrifice	9	13	17	12	15	7
Natural Death	7	6	3	4	4	10
Survivors						
Moribund Sacrifice				2		
Natural Death	1					
Terminal Sacrifice	34	31	30	32	31	33
Animals Examined Microscopically	51	50	50	50	50	50
ALIMENTARY SYSTEM						
Esophagus	(51)	(50)	(50)	(50)	(50)	(49)
Dilatation	1 [4.0]					
Hyperkeratosis			3 [2.0]	3 [1.3]	2 [2.0]	
Inflammation, Suppurative	1 [2.0]					
Intestine Large, Cecum	(44)	(45)	(47)	(46)	(48)	(43)
Dilatation				1 [3.0]	1 [4.0]	
Hyperplasia, Lymphoid			1 [1.0]			1 [2.0]
Inflammation, Chronic Active			1 [2.0]			1 [2.0]
Inflammation, Suppurative						1 [4.0]
Intestine Large, Colon	(45)	(47)	(48)	(47)	(49)	(43)
Dilatation					1 [4.0]	
Hyperplasia, Lymphoid	1 [3.0]					
Polyarteritis						
Intestine Large, Rectum	(45)	(45)	(47)	(46)	(49)	(42)
Dilatation					1 [4.0]	
Intestine Small, Duodenum	(44)	(45)	(48)	(47)	(49)	(42)
Intestine Small, Ileum	(43)	(45)	(47)	(46)	(48)	(41)

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:58

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:58

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Dilatation					1 [4.0]	
Intestine Small, Jejunum	(43)	(45)	(46)	(46)	(48)	(41)
Dilatation					1 [2.0]	
Inflammation, Chronic Active						1 [2.0]
Liver	(49)	(49)	(50)	(49)	(50)	(48)
Angiectasis	3 [2.3]	5 [1.8]	1 [2.0]	4 [1.8]		
Basophilic Focus	1	3	3	17	3	11
Bile Duct, Hyperplasia	2 [1.5]	9 [1.0]	7 [1.1]		3 [1.0]	7 [1.4]
Biliar Tract, Fibrosis	4 [1.3]	2 [1.0]	1 [1.0]	1 [1.0]	6 [1.0]	2 [1.0]
Capsule, Fibrosis					1 [1.0]	
Clear Cell Focus	1			1		
Cyst						
Cyst Multilocular			1			
Degeneration, Cystic	4 [1.0]	3 [1.0]	5 [1.2]	8 [1.1]	9 [1.1]	10 [1.3]
Eosinophilic Focus	3	5	8	15	11	5
Focal Cellular Change			1			
Hematopoietic Cell Proliferation	2 [1.5]	1 [2.0]	1 [1.0]	3 [1.3]		5 [1.0]
Hepatodiaphragmatic Nodule	6	1	4	6	2	2
Infiltration Cellular, Lymphocyte		3 [1.3]		1 [1.0]	1 [1.0]	1 [1.0]
Inflammation, Chronic			1 [1.0]			
Inflammation, Chronic Active	13 [1.1]	5 [1.2]	8 [1.1]	9 [1.1]	9 [1.1]	7 [1.1]
Inflammation, Suppurative						1 [4.0]
Left Lat Lobe, Developmental Malformation	1					
Mixed Cell Focus			1	1	1	
Necrosis		1 [2.0]	2 [2.0]	2 [1.0]	2 [1.5]	3 [2.0]
Oval Cell, Hyperplasia		1 [2.0]				
Tension Lipidosis			3 [3.0]	3 [2.0]		1 [4.0]
Vacuolization Cytoplasmic	12 [1.4]	4 [2.0]	10 [1.4]	13 [1.2]	8 [1.4]	16 [1.4]
Mesentery	(1)	(1)	(2)	(1)	(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)

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:58

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:58

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Fat, Necrosis		1 [4.0]	2 [3.5]	1 [4.0]	1 [4.0]	1 [4.0]
Oral Mucosa	(3)	(2)	(4)	(9)	(5)	(1)
Gingival, Inflammation, Chronic Active						1 [2.0]
Keratin Cyst		1		2	1	
Pancreas	(46)	(48)	(48)	(48)	(49)	(47)
Acinar Cell, Degeneration	40 [2.3]	42 [2.1]	40 [2.0]	39 [2.3]	46 [2.0]	39 [2.3]
Basophilic Focus				1		
Polyarteritis						
Salivary Glands	(48)	(50)	(49)	(50)	(50)	(48)
Acinus, Degeneration				1 [3.0]	1 [2.0]	
Duct, Dilatation				1 [4.0]		
Inflammation, Chronic Active				1 [1.0]		
Inflammation, Suppurative			1 [2.0]			
Stomach, Forestomach	(47)	(48)	(49)	(48)	(50)	(46)
Edema	1 [4.0]	1 [3.0]			1 [2.0]	1 [4.0]
Epithelium, Hyperplasia	2 [2.5]				4 [2.3]	1 [2.0]
Hyperplasia, Basal Cell					1 [4.0]	
Inflammation, Suppurative					1 [3.0]	1 [2.0]
Keratin Cyst	1		1			
Necrosis		1 [2.0]				
Ulcer					1 [2.0]	1 [3.0]
Stomach, Glandular	(45)	(47)	(48)	(46)	(50)	(45)
Cyst		1				
Edema				1 [3.0]	1 [2.0]	
Epithelium, Hyperplasia				1 [2.0]		
Tongue	(0)	(0)	(0)	(0)	(0)	(0)
Inflammation, Chronic Active						
ARDIOVASCULAR SYSTEM						
Blood Vessel	(51)	(50)	(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)

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:58

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:58

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Heart	(51)	(50)	(50)	(50)	(50)	(48)
Atrium, Dilatation		3 [3.3]			2 [2.5]	1 [2.0]
Atrium, Thrombosis					1 [4.0]	
Cardiomyopathy	42 [1.6]	34 [1.5]	40 [1.4]	37 [1.5]	40 [1.7]	41 [1.6]
Endocardium, Hyperplasia			1 [3.0]			1 [3.0]
Inflammation, Suppurative				1 [3.0]	1 [2.0]	
Metaplasia, Osseous	1 [2.0]				1 [2.0]	
Ventricle, Dilatation					1 [4.0]	1 [4.0]
ENDOCRINE SYSTEM						
Adrenal Cortex	(47)	(50)	(50)	(50)	(49)	(45)
Accessory Adrenal Cortical Nodule	3	1	2	1	2	2
Angiectasis	1 [1.0]	2 [2.0]	3 [2.0]	3 [1.7]	4 [1.8]	4 [2.0]
Atrophy				1 [4.0]	1 [3.0]	
Cyst				1 [4.0]		
Degeneration, Cystic	4 [1.5]		4 [1.8]	4 [2.5]	6 [1.3]	4 [1.8]
Hyperplasia	8 [1.4]	6 [2.2]	7 [1.7]	6 [1.5]	1 [1.0]	4 [1.5]
Hypertrophy	3 [1.7]	3 [1.3]	1 [1.0]	3 [1.7]	4 [2.0]	2 [1.0]
Metaplasia, Osseous		1 [4.0]				
Pigmentation				1 [4.0]		
Vacuolization Cytoplasmic	11 [1.2]	9 [2.0]	10 [1.6]	11 [1.3]	11 [1.4]	15 [1.3]
Adrenal Medulla	(46)	(48)	(49)	(49)	(49)	(45)
Angiectasis		1 [4.0]				
Atrophy					1 [3.0]	
Bilateral, Hyperplasia	1 [1.0]					
Cyst						1
Hyperplasia	7 [1.3]	4 [1.3]	5 [1.6]	13 [1.8]	12 [1.1]	16 [2.0]
Islets, Pancreatic	(46)	(48)	(49)	(49)	(49)	(47)
Hyperplasia	12 [1.3]	13 [1.2]	5 [1.0]	7 [1.0]	10 [1.0]	10 [1.3]
Parathyroid Gland	(49)	(46)	(46)	(46)	(50)	(44)

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:58

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:58

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Bilateral, Hyperplasia						
Hyperplasia	7 [2.3]	5 [1.8]	6 [1.3]	9 [1.9]	8 [2.1]	7 [1.9]
Pituitary Gland	(48)	(50)	(50)	(50)	(50)	(47)
Pars Distalis, Cyst	3	1	5	6	6	1
Pars Distalis, Hyperplasia	11 [1.5]	6 [2.3]	12 [1.8]	17 [2.1]	8 [2.0]	9 [1.9]
Pars Intermed, Cyst		1		1	1	1
Pars Intermed, Hyperplasia		1 [2.0]	1 [2.0]			
Thyroid Gland	(47)	(49)	(50)	(49)	(50)	(47)
C Cell, Hyperplasia	6 [1.2]	4 [1.0]	6 [1.2]	10 [1.5]	2 [1.0]	7 [1.3]
Cyst, Squamous	5		16	8	11	11
Follicular Cel, Hyperplasia			1 [2.0]			
Infiltration Cellular, Lymphocyte		,			1 [1.0]	
GENERAL BODY SYSTEM						
Tissue NOS	(1)	(0)	(1)	(0)	(1)	(0)
Degeneration, Cystic	1 [4.0]					
Infiltration Cellular, Plasma Cell	1 [4.0]					
GENITAL SYSTEM						
Coagulating Gland	(46)	(49)	(49)	(49)	(49)	(47)
Atrophy		2 [2.5]	2 [3.5]	1 [3.0]	1 [2.0]	
Degeneration, Cystic				1 [3.0]		
Developmental Malformation	9	10	11	14	14	8
Hemorrhage					1 [4.0]	
Inflammation, Chronic Active						1 [3.0]
Inflammation, Suppurative				2 [3.0]		2 [3.5]
Lumen, Dilatation		2 [2.5]		1 [1.0]		
Ductus Deferens	(0)	(0)	(0)	(0)	(0)	(1)
Inflammation, Suppurative						1 [4.0]
Epididymis	(51)	(50)	(50)	(50)	(50)	(49)

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:59

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:59

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Atrophy	3 [2.7]	5 [2.6]	5 [2.2]	2 [2.5]	4 [2.3]	4 [2.0]
Granuloma Sperm		1 [4.0]				
Hyperplasia					1 [4.0]	
Hypospermia	3 [4.0]	5 [4.0]	6 [4.0]	3 [4.0]	4 [3.5]	4 [4.0]
Infiltration Cellular, Lymphocyte					1 [1.0]	1 [1.0]
Polyarteritis						
Penis	(0)	(1)	(0)	(0)	(1)	(1)
Dilatation						1 [3.0]
Preputial Gland	(51)	(50)	(50)	(50)	(50)	(50)
Abscess	1 [4.0]	1 [4.0]	1 [4.0]	2 [4.0]		1 [4.0]
Atrophy		2 [2.0]	5 [1.8]	4 [2.5]	6 [1.7]	3 [2.7]
Cyst					1	
Duct, Dilatation	14 [2.5]	9 [2.0]	15 [2.5]	14 [2.4]	12 [2.8]	16 [2.4]
Hyperplasia, Basal Cell			1 [3.0]			
Infiltration Cellular, Lymphocyte	10 [1.8]	17 [1.6]	10 [1.7]	10 [1.3]	9 [1.2]	9 [1.2]
Inflammation, Chronic Active	2 [1.5]		2 [2.0]	2 [1.5]	2 [1.5]	
Inflammation, Suppurative	20 [2.7]	21 [2.4]	23 [2.7]	25 [2.6]	25 [2.6]	28 [2.4]
Keratin Cyst			1			
Polyarteritis						
Prostate, Dorsal/Lateral Lobe	(36)	(37)	(45)	(41)	(38)	(42)
Atrophy		1 [2.0]	1 [3.0]			
Cyst	1	1			1	
Degeneration						
Hemorrhage					1 [4.0]	
Infiltration Cellular, Lymphocyte			1 [1.0]			2 [1.5]
Inflammation, Chronic Active	2 [1.0]	2 [1.5]	4 [1.3]	2 [2.0]	4 [1.5]	2 [2.5]
Inflammation, Suppurative	34 [1.6]	33 [1.7]	39 [1.6]	38 [1.7]	34 [1.6]	37 [2.1]
Prostate, Dorsal/lateral Lobe	(13)	(13)	(5)	(9)	(12)	(7)
Prostate, Ventral Lobe	(49)	(50)	(50)	(50)	(50)	(49)

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:59

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:59

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Atrophy		1 [2.0]	1 [3.0]			,
Degeneration		1 [2.0]				
Hemorrhage					1 [4.0]	
Hyperplasia	7 [1.9]	4 [1.0]	3 [1.3]	3 [1.3]	7 [2.3]	5 [2.0]
Infiltration Cellular, Lymphocyte	2 [1.0]	1 [2.0]	4 [1.3]	3 [1.3]	2 [1.0]	4 [1.0]
Inflammation, Chronic Active	1 [1.0]	2 [2.0]	2 [2.5]	3 [1.7]	6 [2.2]	4 [2.3]
Inflammation, Suppurative	3 [1.3]	3 [2.3]	6 [1.2]	3 [1.7]	3 [2.7]	5 [2.0]
Rete Testes	(46)	(44)	(46)	(47)	(47)	(43)
Dilatation	2 [1.5]		2 [1.5]	3 [1.7]	2 [2.0]	2 [1.5]
Fibrosis	1 [1.0]		2 [2.5]	1 [2.0]	1 [3.0]	2 [1.5]
Seminal Vesicle	(43)	(46)	(47)	(47)	(49)	(43)
Atrophy	4 [1.8]	4 [2.0]	4 [2.3]	4 [2.3]	1 [3.0]	1 [1.0]
Dilatation					1 [3.0]	
Epithelium, Hyperplasia		1 [4.0]				
Hemorrhage					1 [4.0]	
Inflammation, Suppurative		1 [1.0]		1 [4.0]		1 [4.0]
Lumen, Dilatation	2 [4.0]	5 [2.8]	2 [2.5]	4 [3.0]	5 [3.2]	5 [3.4]
Testes	(48)	(50)	(50)	(50)	(50)	(48)
Granuloma Sperm					1 [2.0]	
Interstit Cell, Hyperplasia			1 [4.0]			
Malformation		1				
Polyarteritis						
Seminif Tub, Degeneration	19 [1.8]	19 [1.7]	19 [2.4]	20 [1.8]	23 [2.0]	29 [2.1]
EMATOPOIETIC SYSTEM						
Bone Marrow	(46)	(48)	(48)	(48)	(50)	(46)
Erythroid Cell, Hyperplasia	,	1 [2.0]	1 [1.0]	2 [2.0]	, ,	,
Hypocellularity	1 [3.0]		2 [3.0]	1 [2.0]	1 [2.0]	1 [2.0]
Myeloid Cell, Hyperplasia	6 [1.7]	7 [2.4]	8 [2.4]	7 [2.3]	6 [2.2]	7 [2.1]
Lymph Node	(14)	(16)	(16)	(15)	(19)	(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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:59

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:59

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Axillary, Hyperplasia, Lymphoid	1 [4.0]	1 [2.0]	1 [2.0]	1 [3.0]	1 [2.0]	
Axillary, Infiltration Cellular, Plasma Cell			1 [4.0]	2 [3.0]	2 [4.0]	1 [4.0]
Degeneration, Cystic	1 [4.0]	2 [3.5]				
Hemal, Degeneration, Cystic						1 [4.0]
Hemal, Inflammation, Chronic Active						1 [3.0]
Hemal, Necrosis						1 [2.0]
Inguinal, Degeneration, Cystic	1 [3.0]					
Inguinal, Hyperplasia, Lymphoid	1 [3.0]			1 [3.0]	1 [2.0]	1 [3.0]
Inguinal, Infiltration Cellular, Plasma Cell	1 [3.0]			2 [4.0]	1 [4.0]	1 [4.0]
Lumbar, Degeneration, Cystic	8 [3.5]	11 [3.3]	8 [2.8]	9 [3.0]	9 [2.3]	8 [3.1]
Lumbar, Hyperplasia, Lymphoid	3 [3.7]	2 [2.5]	2 [3.5]	1 [2.0]	3 [2.3]	6 [2.7]
Lumbar, Infiltration Cellular, Plasma Cell	7 [3.6]	5 [2.6]	7 [3.4]	5 [3.4]	9 [2.9]	10 [3.4]
Mediastinal, Degeneration, Cystic						
Mediastinal, Infiltration Cellular, Histiocyte						
Mediastinal, Infiltration Cellular, Plasma Cell	1 [3.0]	1 [2.0]	1 [2.0]			1 [2.0]
Pancreatic, Degeneration, Cystic			1 [3.0]			
Pancreatic, Hyperplasia, Lymphoid						
Popliteal, Degeneration, Cystic		1 [2.0]				
Popliteal, Infiltration Cellular, Plasma Cell		1 [4.0]		1 [4.0]		
Renal, Degeneration, Cystic	1 [3.0]	3 [2.7]	3 [4.0]	3 [2.7]	1 [3.0]	3 [3.3]
Renal, Hyperplasia, Lymphoid				1 [2.0]	1 [3.0]	1 [2.0]
Renal, Infiltration Cellular, Plasma Cell		1 [3.0]		2 [3.5]	2 [2.5]	3 [3.3]
Thoracic, Degeneration, Cystic			1 [4.0]			
Thoracic, Infiltration Cellular, Plasma Cell						
ymph Node, Mandibular	(49)	(50)	(50)	(48)	(50)	(47)
Degeneration, Cystic	10 [1.9]	11 [2.5]	6 [1.8]	11 [2.2]	6 [3.0]	4 [2.5]
Hyperplasia, Lymphoid	27 [2.1]	26 [2.2]	25 [1.9]	29 [2.1]	24 [2.0]	26 [2.0]
Infiltration Cellular, Plasma Cell	38 [2.4]	33 [2.4]	37 [2.4]	32 [2.4]	39 [2.2]	32 [2.6]
ymph Node, Mesenteric	(45)	(46)	(49)	(47)	(50)	(44)

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:59

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:59

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTI
Degeneration, Cystic	1 [3.0]	3 [1.7]	1 [1.0]		1 [2.0]	1 [4.0]
Hyperplasia, Lymphoid	3 [2.0]	1 [1.0]	1 [2.0]	3 [2.0]	1 [2.0]	5 [2.0]
Infiltration Cellular, Mast Cell						1 [3.0]
Infiltration Cellular, Plasma Cell			1 [2.0]			1 [2.0]
Inflammation, Granulomatous	9 [1.7]	15 [1.7]	14 [1.6]	9 [1.6]	14 [1.7]	10 [1.8]
Spleen	(48)	(49)	(49)	(49)	(50)	(47)
Capsule, Cyst			1			
Depletion Lymphoid	1 [2.0]	1 [4.0]	1 [3.0]	1 [4.0]	2 [3.0]	
Hematopoietic Cell Proliferation	29 [1.8]	14 [2.4]	23 [2.0]	24 [1.5]	16 [2.2]	19 [1.8]
Hyperplasia, Lymphoid	5 [1.8]	1 [1.0]	2 [2.5]		5 [1.8]	
Hyperplasia, Stromal			1 [4.0]	2 [4.0]		1 [4.0]
Infiltration Cellular, Polymorphnuclr	1 [3.0]	1 [4.0]			1 [4.0]	1 [2.0]
Inflammation, Chronic Active		1 [4.0]				
Necrosis		2 [3.0]		1 [4.0]		1 [2.0]
Pigmentation	20 [2.2]	21 [2.0]	22 [2.0]	30 [2.1]	18 [1.9]	18 [2.1]
Red Pulp, Hyperplasia			1 [4.0]			
Thymus	(45)	(44)	(46)	(44)	(46)	(40)
Atrophy	43 [3.9]	40 [3.9]	44 [3.9]	41 [4.0]	42 [3.9]	39 [3.9]
Cyst	1		1			
Cyst, Squamous					1	
Degeneration, Cystic					1 [3.0]	
Epithel Cell, Hyperplasia	2 [1.5]	2 [1.5]	2 [2.0]	4 [1.3]	2 [2.0]	2 [1.5]
TEGUMENTARY SYSTEM						
Mammary Gland	(44)	(45)	(47)	(44)	(45)	(47)
Acinus, Degeneration	29 [2.1]	19 [2.4]	24 [2.7]	11 [2.9]	24 [2.7]	17 [2.4]
Alveolus, Hyperplasia	1 [1.0]	4 [1.8]	6 [1.8]	18 [2.2]	2 [2.0]	6 [2.0]
Atypical Focus		1				1
Duct, Dilatation			1 [1.0]			
Duct, Ectasia			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)

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:59

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:18:59

First Dose M/F: NA / NA

Lab: NCTR

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Duct, Hyperplasia			2 [1.0]	3 [1.0]		1 [2.0]
Duct, Inflammation, Chronic Active			1 [3.0]			
Ectasia	1 [2.0]	3 [2.7]	7 [2.0]	4 [2.3]	3 [1.3]	12 [1.8]
Fibrosis						
Lactation	3 [1.7]	9 [1.9]	10 [2.0]	4 [2.8]	5 [2.0]	14 [2.4]
Skin	(51)	(50)	(50)	(50)	(50)	(49)
Abscess			1 [4.0]			1 [4.0]
Cyst Epithelial Inclusion	2	1	2	4	2	2
Dermis, Fibrosis					1 [3.0]	
Dermis, Inflammation, Chronic Active					1 [2.0]	2 [2.0]
Dermis, Necrosis					1 [3.0]	
Epidermis, Hyperplasia	1 [2.0]			1 [2.0]		1 [2.0]
Epidermis, Inflammation, Suppurative	1 [2.0]			1 [4.0]		
Epidermis, Necrosis		1 [2.0]				
Fibrosis	1 [3.0]		1 [2.0]			
Foreign Body			1			
Hair Follicle, Hemorrhage						1 [2.0]
Hyperkeratosis	1 [2.0]	1 [3.0]		1 [2.0]	2 [2.0]	1 [2.0]
Inflammation, Chronic Active	24 [4.0]	14 [3.8]	21 [3.9]	16 [4.0]	28 [4.0]	28 [3.9]
Inflammation, Pyogranulomat					1 [4.0]	
Inflammation, Suppurative				2 [3.0]	1 [4.0]	1 [4.0]
Keratin Cyst						
Ulcer				1 [2.0]		
MUSCULOSKELETAL SYSTEM						
Bone	(2)	(0)	(0)	(1)	(0)	(0)
Bone, Femur	(51)	(50)	(50)	(50)	(50)	(50)
Skeletal Muscle	(1)	(1)	(1)	(2)	(2)	(0)

**NERVOUS 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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:59

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:59

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Brain, Brain Stem	(49)	(50)	(50)	(50)	(50)	(48)
Compression	5 [2.2]	7 [2.0]	9 [2.7]	4 [2.3]	7 [2.6]	10 [2.5]
Gliosis						1 [1.0]
Hemorrhage			1 [1.0]		1 [2.0]	3 [1.3]
Brain, Cerebellum	(49)	(50)	(50)	(50)	(50)	(48)
Angiectasis			1 [3.0]			
Gliosis						
Hemorrhage						
Hydrocephalus					2 [2.0]	
Brain, Cerebrum	(49)	(50)	(50)	(50)	(50)	(48)
Hydrocephalus	1 [1.0]	3 [1.0]	1 [2.0]	1 [2.0]	3 [2.0]	4 [1.5]
Infiltration Cellular				1 [1.0]		
Meninges, Hyperplasia						
Peripheral Nerve	(0)	(1)	(1)	(2)	(2)	(0)
RESPIRATORY SYSTEM						
Lung	(46)	(46)	(48)	(48)	(50)	(45)
Alveolar Epith, Hyperplasia	7 [1.3]	2 [2.5]	4 [2.0]	3 [1.7]	6 [1.5]	9 [1.7]
Artery, Mineralization				2 [1.0]	2 [1.0]	3 [1.0]
Congestion		1 [4.0]				
Hemorrhage						
Infiltration Cellular, Histiocyte	18 [1.5]	18 [1.4]	13 [1.6]	11 [1.5]	23 [1.4]	17 [1.6]
Infiltration Cellular, Lymphocyte				1 [2.0]		
Inflammation, Chronic					2 [1.0]	
Inflammation, Chronic Active		1 [1.0]	2 [2.0]	1 [1.0]	2 [2.0]	1 [2.0]
Mediastinum, Hemorrhage	1 [4.0]					
Mediastinum, Inflammation, Suppurative	1 [4.0]					
Mediastinum, Necrosis	1 [4.0]					
Metaplasia, Osseous	1 [1.0]	1 [1.0]	3 [1.0]	1 [1.0]		2 [1.0]
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)

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:18:59

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:18:59

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Nose	(48)	(49)	(50)	(49)	(49)	(48)
Exudate		1 [2.0]			1 [2.0]	
Foreign Body				1		
Inflammation, Chronic Active	3 [2.0]		1 [2.0]		3 [2.0]	3 [1.7]
Inflammation, Suppurative	4 [1.5]	6 [1.8]	7 [2.4]	7 [2.4]	6 [2.0]	2 [1.5]
Metaplasia, Squamous		1 [3.0]				
Mucosa, Hyperkeratosis	2 [3.0]				2 [3.0]	1 [4.0]
Mucosa, Hyperplasia					1 [4.0]	1 [4.0]
Mucosa, Keratin Cyst			2	1	1	1
Polyp, Inflammatory		1 [4.0]				
Vomeronas Org, Dilatation		3 [3.3]	3 [3.0]		1 [3.0]	
Trachea	(49)	(50)	(50)	(50)	(50)	(46)
Epithelium, Hyperplasia				1 [2.0]		
Infiltration Cellular, Lymphocyte				1 [2.0]		
SPECIAL SENSES SYSTEM						
Eye	(46)	(46)	(47)	(46)	(49)	(45)
Bilateral, Cataract	2	, ,	, ,	, ,	, ,	, ,
Bilateral, Retina, Degeneration				1 [4.0]		
Cataract	1			1		
Cornea, Inflammation, Chronic Active				1 [2.0]		
Hemorrhage	1 [4.0]					
Inflammation, Chronic Active				1 [4.0]		
Inflammation, Suppurative	1 [4.0]			2 [4.0]	1 [4.0]	
Retina, Degeneration						1 [3.0]
Harderian Gland	(48)	(47)	(48)	(48)	(50)	(46)
Acinus, Degeneration	, ,	, ,	4 [1.3]	1 [1.0]	2 [1.0]	, ,
Degeneration			1 [3.0]			
Infiltration Cellular, Lymphocyte	3 [1.0]		1 [2.0]	5 [1.0]	3 [1.3]	1 [2.0]
Polyarteritis					• •	

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1 0PPB	F1 2PPB	F1 10PPB	F1 50PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Lacrimal Gland	(0)	(0)	(0)	(2)	(0)	(1)
Ectopic Harderian				1 [3.0]		1
Zymbal's Gland	(1)	(0)	(0)	(2)	(1)	(0)
Inflammation, Suppurative					1 [4.0]	
URINARY SYSTEM						
Kidney	(46)	(49)	(48)	(49)	(49)	(47)
Capsule, Inflammation, Chronic Active			1 [4.0]			
Cyst	20	18	21	19	25	24
Hydronephrosis			1 [1.0]		2 [2.5]	2 [4.0]
Inflammation, Suppurative					1 [3.0]	1 [3.0]
Medulla, Cyst	1		1		1	
Mineralization			1 [1.0]	4 [1.5]	1 [3.0]	
Nephropathy	42 [1.8]	41 [1.5]	46 [1.7]	40 [1.8]	47 [1.8]	44 [2.0]
Pelvis, Hemorrhage					1 [3.0]	
Pelvis, Inflammation, Suppurative		2 [2.0]	2 [1.5]	1 [3.0]	2 [3.5]	
Polycystic Kidney			1 [4.0]		1 [4.0]	
Renal Tubule, Hyperplasia			1 [1.0]	1 [1.0]	1 [2.0]	
Transit Epithe, Hyperplasia	3 [2.7]	2 [2.0]	5 [2.4]		1 [1.0]	
Urethra	(2)	(2)	(0)	(2)	(1)	(5)
Hemorrhage					1 [3.0]	
Urinary Bladder	(45)	(48)	(49)	(49)	(49)	(45)
Dilatation	1 [4.0]				2 [4.0]	3 [3.7]
Hemorrhage				2 [2.5]	1 [3.0]	1 [4.0]
Inflammation, Chronic Active					1 [2.0]	
Inflammation, Suppurative				2 [2.5]	1 [3.0]	1 [3.0]
Transit Epithe, Hyperplasia				1 [2.0]	2 [1.5]	

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)

Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Disposition Summary	
Animals Initially In Study	50
Early Deaths	
Moribund Sacrifice	14
Natural Death	3
Survivors	
Moribund Sacrifice	
Natural Death	
Terminal Sacrifice	33
Animals Examined Microscopically	50
ALIMENTARY SYSTEM	
Esophagus	(50)
Dilatation	
Hyperkeratosis	
Inflammation, Suppurative	
Intestine Large, Cecum	(46)
Dilatation	
Hyperplasia, Lymphoid	
Inflammation, Chronic Active	
Inflammation, Suppurative	
Intestine Large, Colon	(48)
Dilatation	
Hyperplasia, Lymphoid	
Polyarteritis	1 [2.0]
Intestine Large, Rectum Dilatation	(48)
Intestine Small, Duodenum	(47)
Intestine Small, Ileum	(45)

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Dilatation	
Intestine Small, Jejunum	(48)
Dilatation	
Inflammation, Chronic Active	
Liver	(49)
Angiectasis	1 [1.0]
Basophilic Focus	6
Bile Duct, Hyperplasia	4 [1.0]
Biliar Tract, Fibrosis	
Capsule, Fibrosis	
Clear Cell Focus	
Cyst	2
Cyst Multilocular	
Degeneration, Cystic	10 [1.4]
Eosinophilic Focus	10
Focal Cellular Change	
Hematopoietic Cell Proliferation	2 [1.0]
Hepatodiaphragmatic Nodule	3
Infiltration Cellular, Lymphocyte	2 [1.0]
Inflammation, Chronic	
Inflammation, Chronic Active	12 [1.1]
Inflammation, Suppurative	
Left Lat Lobe, Developmental Malformation	
Mixed Cell Focus	
Necrosis	1 [1.0]
Oval Cell, Hyperplasia	
Tension Lipidosis	2 [2.0]
Vacuolization Cytoplasmic	12 [1.6]
Mesentery	(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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Fat, Necrosis	
Oral Mucosa	(6)
Gingival, Inflammation, Chronic Active	
Keratin Cyst	1
Pancreas	(50)
Acinar Cell, Degeneration	46 [2.1]
Basophilic Focus	
Polyarteritis	1 [1.0]
Salivary Glands	(49)
Acinus, Degeneration	
Duct, Dilatation	
Inflammation, Chronic Active	
Inflammation, Suppurative	
Stomach, Forestomach	(49)
Edema	
Epithelium, Hyperplasia	2 [2.5]
Hyperplasia, Basal Cell	
Inflammation, Suppurative	
Keratin Cyst	
Necrosis	
Ulcer	
Stomach, Glandular	(49)
Cyst	
Edema	
Epithelium, Hyperplasia	
Tongue	(1)
Inflammation, Chronic Active	1 [2.0]
CARDIOVASCULAR SYSTEM	
Blood Vessel	(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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Heart	(50)
Atrium, Dilatation	2 [3.5]
Atrium, Thrombosis	
Cardiomyopathy	41 [1.6]
Endocardium, Hyperplasia	
Inflammation, Suppurative	
Metaplasia, Osseous	1 [2.0]
Ventricle, Dilatation	1 [3.0]
ENDOCRINE SYSTEM	
Adrenal Cortex	(50)
Accessory Adrenal Cortical Nodule	( )
Angiectasis	2 [2.0]
Atrophy	1 [4.0]
Cyst	
Degeneration, Cystic	6 [1.7]
Hyperplasia	14 [1.6]
Hypertrophy	6 [1.8]
Metaplasia, Osseous	
Pigmentation	
Vacuolization Cytoplasmic	15 [1.3]
Adrenal Medulla	(50)
Angiectasis	
Atrophy	
Bilateral, Hyperplasia	
Cyst	
Hyperplasia	7 [1.4]
Islets, Pancreatic	(50)
Hyperplasia	7 [1.0]
Parathyroid Gland	(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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Bilateral, Hyperplasia	1 [2.0]
Hyperplasia	6 [1.8]
Pituitary Gland	(49)
Pars Distalis, Cyst	1
Pars Distalis, Hyperplasia	10 [2.0]
Pars Intermed, Cyst	1
Pars Intermed, Hyperplasia	
Thyroid Gland	(49)
C Cell, Hyperplasia	7 [1.6]
Cyst, Squamous	8
Follicular Cel, Hyperplasia	
Infiltration Cellular, Lymphocyte	
GENERAL BODY SYSTEM	
Tissue NOS	(0)
Degeneration, Cystic	(-)
Infiltration Cellular, Plasma Cell	
GENITAL SYSTEM	
	(40)
Coagulating Gland	(48)
Atrophy  Degeneration Cyptic	2 [2.0]
Degeneration, Cystic  Developmental Malformation	8
·	O
Hemorrhage Inflammation, Chronic Active	
•	1 [1 0]
Inflammation, Suppurative Lumen, Dilatation	1 [1.0]
Ductus Deferens	3 [2.7]
	(0)
Inflammation, Suppurative Epididymis	(50)
Lpiuluyiiiis	(30)

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)

Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Atrophy	8 [2.1]
Granuloma Sperm	
Hyperplasia	
Hypospermia	8 [3.8]
Infiltration Cellular, Lymphocyte	
Polyarteritis	1 [2.0]
Penis	(0)
Dilatation	
Preputial Gland	(50)
Abscess	1 [4.0]
Atrophy	6 [2.8]
Cyst	
Duct, Dilatation	12 [2.2]
Hyperplasia, Basal Cell	
Infiltration Cellular, Lymphocyte	8 [1.6]
Inflammation, Chronic Active	3 [2.3]
Inflammation, Suppurative	23 [2.4]
Keratin Cyst	2
Polyarteritis	1 [2.0]
Prostate, Dorsal/Lateral Lobe	(38)
Atrophy	1 [3.0]
Cyst	
Degeneration	1 [2.0]
Hemorrhage	
Infiltration Cellular, Lymphocyte	
Inflammation, Chronic Active	3 [2.3]
Inflammation, Suppurative	35 [1.7]
Prostate, Dorsal/lateral Lobe	(12)
Prostate, Ventral Lobe	(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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Atrophy	
Degeneration	
Hemorrhage	
Hyperplasia	3 [1.3]
Infiltration Cellular, Lymphocyte	2 [1.0]
Inflammation, Chronic Active	4 [2.5]
Inflammation, Suppurative	3 [2.0]
Rete Testes	(46)
Dilatation	5 [1.2]
Fibrosis	
Seminal Vesicle	(48)
Atrophy	5 [2.4]
Dilatation	
Epithelium, Hyperplasia	1 [4.0]
Hemorrhage	
Inflammation, Suppurative	2 [3.0]
Lumen, Dilatation	
Testes	(50)
Granuloma Sperm	
Interstit Cell, Hyperplasia	
Malformation	
Polyarteritis	1 [2.0]
Seminif Tub, Degeneration	23 [2.0]
HEMATOPOIETIC SYSTEM	
Bone Marrow	(49)
Erythroid Cell, Hyperplasia	1 [4.0]
Hypocellularity	, [ ,,o]
Myeloid Cell, Hyperplasia	7 [2.3]
Lymph Node	(19)
-Jp 1000	()

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Axillary, Hyperplasia, Lymphoid	2 [4.0]
Axillary, Infiltration Cellular, Plasma Cell	2 [3.0]
Degeneration, Cystic	
Hemal, Degeneration, Cystic	
Hemal, Inflammation, Chronic Active	
Hemal, Necrosis	
Inguinal, Degeneration, Cystic	1 [4.0]
Inguinal, Hyperplasia, Lymphoid	1 [3.0]
Inguinal, Infiltration Cellular, Plasma Cell	
Lumbar, Degeneration, Cystic	17 [3.1]
Lumbar, Hyperplasia, Lymphoid	4 [2.8]
Lumbar, Infiltration Cellular, Plasma Cell	9 [3.0]
Mediastinal, Degeneration, Cystic	1 [1.0]
Mediastinal, Infiltration Cellular, Histiocyte	1 [3.0]
Mediastinal, Infiltration Cellular, Plasma Cell	
Pancreatic, Degeneration, Cystic	
Pancreatic, Hyperplasia, Lymphoid	1 [2.0]
Popliteal, Degeneration, Cystic	
Popliteal, Infiltration Cellular, Plasma Cell	1 [4.0]
Renal, Degeneration, Cystic	4 [3.8]
Renal, Hyperplasia, Lymphoid	
Renal, Infiltration Cellular, Plasma Cell	2 [3.0]
Thoracic, Degeneration, Cystic	
Thoracic, Infiltration Cellular, Plasma Cell	1 [3.0]
Lymph Node, Mandibular	(50)
Degeneration, Cystic	15 [1.9]
Hyperplasia, Lymphoid	29 [2.2]
Infiltration Cellular, Plasma Cell	33 [2.2]
Lymph Node, Mesenteric	(48)

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Degeneration, Cystic	2 [3.0]
Hyperplasia, Lymphoid	8 [2.0]
Infiltration Cellular, Mast Cell	
Infiltration Cellular, Plasma Cell	
Inflammation, Granulomatous	7 [2.0]
Spleen	(50)
Capsule, Cyst	
Depletion Lymphoid	2 [3.0]
Hematopoietic Cell Proliferation	21 [1.8]
Hyperplasia, Lymphoid	2 [2.0]
Hyperplasia, Stromal	
Infiltration Cellular, Polymorphnuclr	1 [2.0]
Inflammation, Chronic Active	
Necrosis	
Pigmentation	16 [1.9]
Red Pulp, Hyperplasia	
Thymus	(47)
Atrophy	47 [3.9]
Cyst	
Cyst, Squamous	
Degeneration, Cystic	
Epithel Cell, Hyperplasia	1 [2.0]
INTEGUMENTARY SYSTEM	
Mammary Gland	(48)
Acinus, Degeneration	11 [2.1]
Alveolus, Hyperplasia	14 [1.9]
Atypical Focus	1
Duct, Dilatation	1 [4.0]
Duct, Ectasia	

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

Lab: NCTR

CD Rat MALE	F1/50PPB TO CTL
Duct, Hyperplasia	3 [2.0]
Duct, Inflammation, Chronic Active	
Ectasia	4 [1.8]
Fibrosis	1 [4.0]
Lactation	9 [2.0]
Skin	(50)
Abscess	1 [4.0]
Cyst Epithelial Inclusion	4
Dermis, Fibrosis	
Dermis, Inflammation, Chronic Active	
Dermis, Necrosis	
Epidermis, Hyperplasia	
Epidermis, Inflammation, Suppurative	
Epidermis, Necrosis	
Fibrosis	
Foreign Body	
Hair Follicle, Hemorrhage	
Hyperkeratosis	
Inflammation, Chronic Active	29 [4.0]
Inflammation, Pyogranulomat	
Inflammation, Suppurative	1 [4.0]
Keratin Cyst	1
Ulcer	
MUSCULOSKELETAL SYSTEM	
Bone	(1)
Bone, Femur	(50)
Skeletal Muscle	(1)

**NERVOUS 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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Brain, Brain Stem	(50)
Compression	7 [3.1]
Gliosis	
Hemorrhage	
Brain, Cerebellum	(50)
Angiectasis	
Gliosis	1 [1.0]
Hemorrhage	1 [1.0]
Hydrocephalus	
Brain, Cerebrum	(50)
Hydrocephalus	4 [1.8]
Infiltration Cellular	
Meninges, Hyperplasia	1 [2.0]
Peripheral Nerve	(0)
RESPIRATORY SYSTEM	
Lung	(50)
Alveolar Epith, Hyperplasia	3 [1.3]
Artery, Mineralization	2 [1.0]
Congestion	
Hemorrhage	1 [2.0]
Infiltration Cellular, Histiocyte	11 [1.4]
Infiltration Cellular, Lymphocyte	
Inflammation, Chronic	
Inflammation, Chronic Active	1 [1.0]
Mediastinum, Hemorrhage	
Mediastinum, Inflammation, Suppurative	
Mediastinum, Necrosis	
Metaplasia, Osseous	1 [1.0]
Thrombosis	

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat MALE	F1/50PPB TO CTL
Nose	(50)
Exudate	
Foreign Body	
Inflammation, Chronic Active	
Inflammation, Suppurative	5 [2.6]
Metaplasia, Squamous	
Mucosa, Hyperkeratosis	
Mucosa, Hyperplasia	
Mucosa, Keratin Cyst	
Polyp, Inflammatory	
Vomeronas Org, Dilatation	
Trachea	(50)
Epithelium, Hyperplasia	
Infiltration Cellular, Lymphocyte	,
SPECIAL SENSES SYSTEM	
Eye	(48)
Bilateral, Cataract	
Bilateral, Retina, Degeneration	
Cataract	2
Cornea, Inflammation, Chronic Active	
Hemorrhage	
Inflammation, Chronic Active	1 [3.0]
Inflammation, Suppurative	
Retina, Degeneration	1 [4.0]
Harderian Gland	(50)
Acinus, Degeneration	5 [1.8]
Degeneration	
Infiltration Cellular, Lymphocyte	8 [1.4]
Polyarteritis	1 [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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:00

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

Lab: NCTR

CD Rat MALE	F1/50PPB TO CTL
Lacrimal Gland	(1)
Ectopic Harderian	
Zymbal's Gland	(0)
Inflammation, Suppurative	
URINARY SYSTEM	
Kidney	(49)
Capsule, Inflammation, Chronic Active	
Cyst	23
Hydronephrosis	
Inflammation, Suppurative	
Medulla, Cyst	
Mineralization	4 [1.3]
Nephropathy	46 [2.0]
Pelvis, Hemorrhage	
Pelvis, Inflammation, Suppurative	1 [2.0]
Polycystic Kidney	
Renal Tubule, Hyperplasia	1 [1.0]
Transit Epithe, Hyperplasia	2 [2.0]
Urethra	(1)
Hemorrhage	
Urinary Bladder	(49)
Dilatation	
Hemorrhage	
Inflammation, Chronic Active	
Inflammation, Suppurative	
Transit Epithe, Hyperplasia	

\*\*\*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)

Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:00

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:00

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Disposition Summary						
Animals Initially In Study	51	50	50	50	50	50
Early Deaths						
Moribund Sacrifice	24	22	27	17	26	27
Natural Death		5	3	8	7	1
Survivors						
Moribund Sacrifice	1		1		1	
Terminal Sacrifice	26	23	19	25	16	22
Animals Examined Microscopically	51	50	50	50	50	50
ALIMENTARY SYSTEM						
Esophagus	(51)	(50)	(49)	(50)	(50)	(50)
Hemorrhage				1 [2.0]		
Inflammation		2 [3.0]	1 [4.0]	1 [4.0]	1 [4.0]	2 [4.0]
Necrosis					1 [4.0]	
Intestine Large, Cecum	(51)	(50)	(49)	(49)	(50)	(50)
Dilatation					1 [2.0]	
Erosion					1 [3.0]	
Inflammation, Chronic		1 [2.0]			1 [2.0]	
Intestine Large, Colon	(51)	(50)	(49)	(49)	(50)	(50)
Developmental Malformation			1			
Inflammation		1 [1.0]				
Lymphoid Tiss, Inflammation		1 [3.0]				
Mineralization			1 [2.0]			
Muscularis, Inflammation, Chronic			1 [2.0]			
Intestine Large, Rectum	(51)	(50)	(49)	(50)	(50)	(50)
Inflammation, Chronic		1 [1.0]				
Intestine Small, Duodenum	(51)	(50)	(49)	(49)	(50)	(50)
Autolysis				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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:01

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:01

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Dilatation			1 [3.0]			
Intestine Small, Ileum	(51)	(50)	(48)	(49)	(48)	(50)
Autolysis				1 [3.0]		
Intestine Small, Jejunum	(51)	(50)	(47)	(47)	(48)	(50)
Autolysis				1 [3.0]		
Inflammation, Chronic		1 [4.0]				
Liver	(51)	(50)	(49)	(50)	(50)	(50)
Angiectasis	2 [1.5]	1 [1.0]	1 [2.0]	1 [2.0]	2 [1.5]	4 [2.0]
Basophilic Focus	1	1	1	1	1	1
Bile Duct, Hyperplasia	13 [1.4]	15 [1.3]	10 [1.3]	10 [1.2]	7 [1.7]	16 [1.4]
Capsule, Hemorrhage, Focal	2 [2.0]				1 [2.0]	
Caudate Lobe, Developmental Malformation						
Centrilobular, Necrosis			1 [3.0]			
Cyst		1			2	2
Cyst Multilocular	3		1		2	1
Degeneration, Cystic	1 [2.0]			2 [1.5]	3 [2.3]	5 [1.6]
Eosinophilic Focus	1	1	1	5	2	5
Eosinophilic Focus, Multiple						
Hematopoietic Cell Proliferation	2 [1.0]	4 [1.0]	3 [1.7]	1 [2.0]	2 [1.5]	1 [1.0]
Hepatodiaphragmatic Nodule	1			1	2	
Infiltration Cellular, Lymphocyte	3 [1.0]	2 [1.0]	1 [1.0]	1 [1.0]		1 [1.0]
Inflammation, Chronic Active	6 [1.0]	1 [1.0]	6 [1.2]	5 [1.0]	3 [2.0]	6 [1.0]
Left Lat Lobe, Developmental Malformation						
Median Lobe, Developmental Malformation					1	
Mixed Cell Focus						
Necrosis					2 [2.5]	
Oval Cell, Hyperplasia					1 [3.0]	
Periportal, Inflammation, Chronic	3 [1.0]	7 [1.3]	5 [1.0]	5 [1.0]	3 [1.7]	5 [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)

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:01

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:01

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Right Lat Lobe, Developmental Malformation		1				
Tension Lipidosis			1 [2.0]	1		
Vacuolization Cytoplasmic	12 [2.2]	8 [1.8]	4 [2.0]	2 [2.0]	9 [2.0]	11 [2.2]
Mesentery	(2)	(2)	(0)	(1)	(2)	(1)
Cyst					1	
Fat, Necrosis	2 [2.0]	1 [3.0]		1 [2.0]	2 [3.0]	
Oral Mucosa	(3)	(1)	(4)	(4)	(5)	(7)
Gingival, Inflammation				1 [3.0]		
Keratin Cyst			3	2	2	4
Pancreas	(51)	(50)	(49)	(49)	(50)	(50)
Acinar Cell, Degeneration	20 [2.0]	24 [1.8]	20 [2.0]	24 [1.8]	19 [1.6]	26 [1.9]
Acinar Cell, Hyperplasia				1 [1.0]		1 [2.0]
Artery, Inflammation, Chronic						1 [2.0]
Artery, Mineralization			1 [2.0]			
Inflammation, Chronic		7 [1.1]		1 [2.0]	1 [4.0]	
Lipomatosis	1 [2.0]	1 [2.0]	1 [2.0]	2 [3.0]	1 [2.0]	3 [2.0]
Salivary Glands	(51)	(49)	(49)	(50)	(50)	(50)
Artery, Mineralization			1 [2.0]			
Inflammation, Chronic	1 [3.0]	2 [3.0]			2 [3.0]	1 [1.0]
Stomach, Forestomach	(51)	(50)	(49)	(50)	(50)	(50)
Edema	1 [2.0]		1 [2.0]	2 [3.0]	3 [2.3]	
Hyperplasia	2 [2.5]	2 [2.0]		1 [2.0]	4 [2.3]	
Inflammation	1 [2.0]	4 [2.3]	1 [2.0]		2 [2.5]	
Keratin Cyst					1	
Ulcer				1 [3.0]		1 [3.0]
Stomach, Glandular	(51)	(50)	(49)	(50)	(50)	(50)
Glands, Dilatation			1 [3.0]			
Infiltration Cellular, Lymphocyte		1 [2.0]			1 [2.0]	
Inflammation				1 [1.0]	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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:01

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:01

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Mineralization			1 [3.0]			
Tooth	(0)	(0)	(0)	(0)	(1)	(0)
Dysplasia					1 [3.0]	
CARDIOVASCULAR SYSTEM						
Blood Vessel	(51)	(50)	(49)	(50)	(50)	(50)
Mineralization			1 [3.0]		1 [1.0]	
Heart	(51)	(50)	(49)	(50)	(50)	(50)
Artery, Mineralization			1 [2.0]			
Cardiomyopathy	23 [1.5]	18 [1.4]	10 [1.3]	12 [1.3]	22 [1.3]	16 [1.1]
Endocardium, Hyperplasia						
Myocardium, Necrosis		1 [2.0]				
ENDOCRINE SYSTEM						
Adrenal Cortex	(51)	(50)	(49)	(50)	(50)	(50)
Angiectasis	2 [2.0]	1 [2.0]		2 [2.0]	1 [1.0]	3 [2.0]
Atrophy	2 [2.5]		2 [2.5]		1 [2.0]	
Cyst				1 [4.0]		
Degeneration, Cystic	46 [2.8]	43 [2.6]	43 [2.5]	45 [2.5]	43 [2.7]	44 [2.6]
Hematopoietic Cell Proliferation				1 [2.0]		1 [1.0]
Hyperplasia	1 [3.0]	3 [2.0]	1 [1.0]			
Hypertrophy	5 [1.6]	11 [2.1]	7 [1.9]	7 [2.3]	2 [2.0]	2 [2.5]
Metaplasia, Osseous	1 [3.0]					
Vacuolization Cytoplasmic	1 [1.0]		1 [2.0]			
Adrenal Medulla	(48)	(50)	(49)	(50)	(49)	(48)
Hyperplasia					1 [1.0]	1 [2.0]
Islets, Pancreatic	(51)	(50)	(49)	(49)	(50)	(50)
Hyperplasia	1 [2.0]	1 [2.0]	3 [2.0]		1 [1.0]	
Parathyroid Gland	(48)	(47)	(46)	(43)	(46)	(44)
Hyperplasia		2 [2.0]	1 [2.0]	1 [2.0]	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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:01

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:01

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Pituitary Gland	(51)	(50)	(50)	(50)	(50)	(50)
Angiectasis	2 [2.0]					2 [2.0]
Cyst			1			
Pars Distalis, Cyst	2	2	1	1	1	1
Pars Distalis, Hyperplasia	4 [2.5]	3 [1.7]	3 [2.0]	2 [2.0]	2 [2.0]	4 [2.0]
Pars Intermed, Cyst	1	3			1	
Pigmentation						1 [3.0]
Thyroid Gland	(51)	(50)	(49)	(50)	(50)	(50)
C Cell, Hyperplasia	7 [1.4]	4 [1.5]	3 [2.0]	4 [2.0]		3 [2.0]
Follicle, Cyst						1 [2.0]
Follicular Cel, Hyperplasia				1 [1.0]	1 [3.0]	2 [1.5]
Infiltration Cellular, Lymphocyte	1 [1.0]				1 [1.0]	
Inflammation, Chronic	1 [1.0]	1 [3.0]		2 [1.0]		1 [1.0]
Ultimobranchial Cyst	2 [3.0]	4 [3.0]	1 [3.0]	5 [2.2]	1 [2.0]	2 [2.5]
GENERAL BODY SYSTEM						
Tissue NOS	(0)	(0)	(1)	(0)	(0)	(0)
GENITAL SYSTEM						
Clitoral Gland	(50)	(49)	(48)	(50)	(50)	(49)
Atrophy		6 [2.3]	4 [2.5]	2 [1.5]	3 [2.3]	1 [2.0]
Duct, Dilatation	11 [2.1]	12 [2.4]	10 [2.3]	9 [2.6]	17 [2.3]	9 [2.4]
Duct, Hyperplasia, Squamous	2 [3.0]	1 [3.0]	2 [2.0]		1 [1.0]	1 [2.0]
Hyperplasia		2 [1.5]	2 [2.0]			1 [2.0]
Infiltration Cellular, Lymphocyte					1 [2.0]	
Inflammation	34 [2.1]	33 [2.2]	32 [2.1]	28 [2.3]	32 [2.0]	29 [2.2]
Keratin Cyst						
Ovary	(51)	(50)	(49)	(50)	(50)	(50)
Angiectasis		1 [1.0]				
Atrophy	49 [2.7]	47 [3.1]	46 [3.2]	46 [3.3]	46 [3.2]	45 [2.9]

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:01

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:01

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CT
Corpus Luteum, Cyst	1	2	1	2	1	
Cyst	15	9	17	22	12	15
Granulosa Cell, Hyperplasia				1 [3.0]		
Hyperplasia, Sertoliform	23 [1.5]	17 [1.5]	13 [1.2]	23 [1.7]	20 [1.6]	19 [1.8]
Interstit Cell, Hyperplasia		1 [3.0]	2 [2.5]		2 [2.5]	2 [3.0]
Oviduct	(51)	(50)	(49)	(50)	(50)	(50)
Pigmentation					1 [2.0]	
Uterus	(51)	(50)	(49)	(50)	(50)	(50)
Atrophy	4 [2.3]	1 [2.0]	2 [2.0]		1 [2.0]	2 [2.0]
Cervix, Hyperplasia						1 [2.0]
Cervix, Muscularis, Hypertrophy						
Dilatation	1 [3.0]					
Endometrium, Hyperplasia	2 [1.5]	7 [1.7]	3 [2.3]	12 [2.5]	2 [1.5]	5 [1.8]
Endometrium, Hyperplasia, Cystic	15 [1.9]	14 [2.1]	20 [1.9]	17 [2.0]	23 [1.9]	12 [1.8]
Hemorrhage		1 [2.0]				
Hyperplasia, Atypical, Focal	6 [1.0]	14 [1.4]	16 [1.4]	20 [1.6]	10 [1.3]	7 [1.6]
Hypoplasia		1 [3.0]			1 [3.0]	
Inflammation	2 [1.5]	1 [1.0]		1 [1.0]	3 [1.3]	2 [1.5]
Metaplasia, Squamous	2 [1.0]	6 [2.0]	8 [1.4]	13 [1.6]	6 [1.5]	6 [1.0]
Pigmentation, Focal	1 [2.0]					
Vagina	(51)	(50)	(49)	(49)	(50)	(48)
Hemorrhage					1 [2.0]	
Inflammation	12 [1.7]	22 [1.8]	18 [1.6]	17 [1.8]	17 [2.0]	15 [2.0]
Mucocyte, Hyperplasia	43 [2.5]	33 [2.2]	35 [2.4]	33 [2.4]	33 [2.4]	33 [2.5]
IEMATOPOIETIC SYSTEM						
Bone Marrow	(51)	(50)	(49)	(50)	(50)	(50)
Hyperplasia	1 [2.0]	1 [2.0]				1 [2.0]
Myeloid Cell, Hyperplasia		1 [2.0]			2 [2.0]	
Lymph Node	(16)	(14)	(6)	(10)	(15)	(12)

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: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:01

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:01

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Axillary, Degeneration, Cystic	1 [2.0]					
Axillary, Hyperplasia, Lymphoid						1 [3.0]
Axillary, Infiltration Cellular, Plasma Cell	2 [3.0]					1 [4.0]
Deep Cervical, Hyperplasia, Lymphoid				1 [2.0]		
Deep Cervical, Infiltration Cellular, Plasma Cell	1 [4.0]			1 [2.0]		
Degeneration, Cystic					1 [2.0]	
Hyperplasia, Lymphoid	1 [4.0]					
Infiltration Cellular, Plasma Cell	1 [3.0]				3 [3.3]	
Inguinal, Degeneration, Cystic						
Inguinal, Hyperplasia, Lymphoid	1 [4.0]					1 [3.0]
Inguinal, Infiltration Cellular, Plasma Cell	1 [3.0]	1 [4.0]			1 [4.0]	1 [3.0]
Lumbar, Degeneration, Cystic	12 [3.0]	11 [2.9]	3 [2.7]	5 [2.4]	7 [3.0]	9 [3.3]
Lumbar, Hemorrhage					1 [3.0]	
Lumbar, Hyperplasia, Lymphoid	5 [2.4]	10 [2.4]	3 [2.7]	3 [2.0]	4 [2.8]	7 [2.3]
Lumbar, Infiltration Cellular, Plasma Cell	10 [2.7]	12 [2.9]	5 [2.8]	5 [3.0]	7 [2.7]	10 [2.6]
Mediastinal, Hemorrhage			1 [2.0]	1 [3.0]		1 [3.0]
Mediastinal, Hyperplasia, Lymphoid Mediastinal, Infiltration Cellular, Plasma Cell	1 [2.0]					1 [2.0]
Mediastinal, Pigmentation	1 [2.0]			1 [3.0]		
Mediastinal, Sinus, Dilatation						1 [3.0]
Pancreatic, Hemorrhage				1 [2.0]		
Pancreatic, Hyperplasia, Lymphoid					1 [2.0]	
Pancreatic, Pigmentation	1 [2.0]			1 [2.0]		
Popliteal, Degeneration, Cystic	1 [2.0]		1 [3.0]	1 [3.0]		
Popliteal, Hyperplasia, Lymphoid	2 [2.0]		2 [2.0]			
Popliteal, Infiltration Cellular, Plasma Cell	2 [3.0]	1 [4.0]	2 [3.5]	1 [4.0]		
Renal, Degeneration, Cystic	1 [2.0]	1 [4.0]				
Renal, Hyperplasia, Lymphoid		1 [2.0]			1 [3.0]	
Renal, Infiltration Cellular, Plasma Cell	1 [4.0]	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: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:01

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:01

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO C
Thoracic, Infiltration Cellular, Plasma Cell	1 [2.0]					
Lymph Node, Mandibular	(51)	(50)	(49)	(50)	(50)	(49)
Degeneration, Cystic	16 [2.1]	9 [2.2]	8 [2.0]	8 [2.4]	8 [1.9]	6 [2.0]
Hemorrhage	1 [2.0]					
Hyperplasia, Lymphoid	4 [2.0]	14 [2.1]	11 [2.0]	5 [2.2]	10 [2.1]	11 [2.1]
Infiltration Cellular, Plasma Cell	43 [2.3]	40 [2.6]	43 [2.3]	39 [2.3]	40 [2.3]	41 [2.3]
Inflammation, Suppurative				1 [2.0]	1 [2.0]	
Lymph Node, Mesenteric	(51)	(50)	(47)	(50)	(50)	(50)
Degeneration, Cystic	1 [2.0]		1 [2.0]	1 [2.0]	1 [2.0]	
Depletion Lymphoid	1 [2.0]	1 [2.0]		1 [3.0]	1 [3.0]	
Hemorrhage		1 [2.0]		2 [2.0]	1 [2.0]	1 [2.0]
Hyperplasia, Lymphoid	5 [2.0]	3 [2.3]	1 [2.0]	2 [2.0]	3 [2.0]	4 [2.0]
Infiltration Cellular, Plasma Cell	5 [2.0]	3 [2.3]	1 [3.0]	1 [2.0]	1 [2.0]	1 [2.0]
Pigmentation	2 [2.0]	1 [2.0]				
Sinus, Dilatation				1 [4.0]		
Spleen	(51)	(50)	(49)	(50)	(50)	(50)
Hematopoietic Cell Proliferation	15 [1.9]	22 [1.9]	10 [2.1]	6 [2.0]	13 [1.8]	17 [1.8]
Hyperplasia, Lymphoid						1 [2.0]
Hyperplasia, Stromal		1 [4.0]				
Lymphocyte, Atrophy		1 [2.0]				
Pigmentation	33 [1.9]	31 [2.0]	34 [2.1]	43 [2.2]	28 [1.8]	25 [1.9]
Red Pulp, Atrophy	1 [2.0]					
Thymus	(51)	(44)	(45)	(48)	(44)	(46)
Cyst	17	18	21	17	13	22
Hemorrhage				1 [3.0]	1 [3.0]	
Hyperplasia, Lymphoid					1 [2.0]	
Inflammation, Chronic		1 [3.0]			1 [2.0]	
ITEGUMENTARY SYSTEM	(54)	(40)	(40)	(50)	(40)	(50)
Mammary Gland	(51)	(49)	(49)	(50)	(49)	(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)

Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Alveolus, Hyperplasia	42 [2.1]	38 [2.4]	42 [2.2]	40 [2.2]	36 [2.3]	43 [2.3]
Atypical Focus	19 [1.5]	23 [1.8]	21 [1.3]	23 [1.6]	13 [2.0]	19 [1.7]
Duct, Dilatation	3 [2.0]	5 [2.8]		4 [2.8]		1 [2.0]
Duct, Inflammation						
Fibrosis						
Galactocele	2	2	4	6	3	4
Lactation	26 [2.0]	32 [2.1]	31 [1.9]	30 [2.0]	29 [1.9]	29 [1.8]
Lobules, Hyperplasia			1 [2.0]			
Skin	(51)	(50)	(49)	(50)	(50)	(50)
Abscess	1 [3.0]					
Cyst Epithelial Inclusion					1	
Hyperkeratosis					1 [2.0]	
Inflammation	23 [3.6]	16 [3.4]	9 [3.9]	7 [3.1]	23 [3.6]	22 [3.6]
MUSCULOSKELETAL SYSTEM						
Bone, Femur	(51)	(50)	(49)	(50)	(50)	(50)
Fibrous Osteodystrophy			1 [3.0]			
Osteopetrosis						1 [2.0]
Skeletal Muscle	(1)	(0)	(1)	(1)	(0)	(4)
Head, Hyalinization, Focal			1 [2.0]			
NERVOUS SYSTEM						
Brain, Brain Stem	(51)	(50)	(49)	(50)	(50)	(50)
Compression	26 [2.4]	26 [2.7]	23 [2.6]	26 [2.5]	23 [2.7]	18 [2.8]
Hemorrhage		1 [2.0]	1 [2.0]	1 [1.0]	2 [1.5]	1 [2.0]
Hydrocephalus			1 [2.0]			
Brain, Cerebellum	(51)	(50)	(49)	(50)	(50)	(50)
Hemorrhage					1 [1.0]	
Hydrocephalus	2 [1.5]			2 [1.5]	3 [1.7]	
Brain, Cerebrum	(51)	(50)	(49)	(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)

**Experiment Number: 99920-94** Test Type: SPECIAL STUDY Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Hemorrhage					1 [2.0]	
Hydrocephalus	2 [2.0]	2 [2.0]	6 [1.8]	5 [1.8]	3 [2.0]	2 [2.0]
Peripheral Nerve	(1)	(0)	(0)	(0)	(0)	(2)
RESPIRATORY SYSTEM						
Lung	(51)	(50)	(49)	(50)	(50)	(50)
Alveolar Epith, Hyperplasia	1 [2.0]	1 [1.0]	2 [1.5]	1 [2.0]	1 [1.0]	1 [1.0]
Artery, Mineralization		1 [1.0]		3 [1.7]	3 [1.3]	1 [2.0]
Autolysis					1 [2.0]	
Edema		1 [1.0]				
Hemorrhage	1 [3.0]			2 [2.0]	1 [1.0]	
Infiltration Cellular, Histiocyte	11 [1.5]	10 [1.9]	10 [1.6]	16 [1.8]	7 [1.9]	4 [1.5]
Infiltration Cellular, Lymphocyte			1 [1.0]	1 [1.0]		
Inflammation	3 [1.3]	1 [2.0]	4 [1.8]	3 [2.0]	3 [1.3]	2 [2.0]
Peribronchial, Inflammation, Chronic		1 [3.0]	1 [1.0]		1 [3.0]	
Perivascular, Inflammation		1 [2.0]	3 [1.0]	1 [1.0]	1 [2.0]	
Nose	(51)	(50)	(49)	(50)	(50)	(50)
Fibrous Osteodystrophy			1 [3.0]			
Foreign Body		1		1		1
Goblet Cell, Hyperplasia	1 [3.0]	2 [1.5]	2 [2.0]		1 [2.0]	1 [3.0]
Hemorrhage		1 [4.0]				
Inflammation	4 [1.8]	5 [2.0]	5 [1.8]	2 [2.5]	9 [2.1]	4 [1.8]
Nasolacrim Dct, Inflammation	29 [1.8]	29 [1.8]	31 [1.7]	30 [1.7]	25 [1.8]	29 [1.8]
Olfactory Epi, Hyaline Droplet	4 [1.8]	5 [1.2]	1 [1.0]	6 [1.7]		3 [1.3]
Osteopetrosis						1 [2.0]
Respirat Epith, Hyaline Droplet					1 [2.0]	
Respirat Epith, Hyperplasia	1 [2.0]				1 [1.0]	
Trachea	(51)	(50)	(49)	(50)	(50)	(50)
Inflammation	2 [1.5]		3 [1.0]	3 [1.0]	1 [1.0]	1 [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)

**Experiment Number: 99920-94** Test Type: SPECIAL STUDY Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
SPECIAL SENSES SYSTEM						
Ear	(0)	(1)	(0)	(0)	(0)	(0)
Eye	(51)	(49)	(48)	(49)	(50)	(50)
Bilateral, Cataract				1	2	
Bilateral, Retina, Degeneration	2 [2.0]	5 [2.0]	1 [2.0]	4 [2.0]	1 [2.0]	6 [2.3]
Cataract		1	1	1	2	2
Cornea, Inflammation	1 [2.0]					
Retina, Autolysis					2 [4.0]	
Retina, Degeneration	3 [2.0]	1 [2.0]	5 [1.8]		2 [1.5]	3 [2.3]
Harderian Gland	(51)	(49)	(49)	(50)	(50)	(50)
Atrophy, Focal	2 [1.5]					
Epithelium, Degeneration					1 [1.0]	
Hyperplasia	3 [1.3]	2 [2.0]	4 [1.5]	5 [1.0]	2 [1.5]	1 [1.0]
Infiltration Cellular, Lymphocyte	3 [1.7]	2 [1.5]	2 [1.0]	1 [1.0]	2 [1.5]	1 [2.0]
Inflammation	7 [1.1]	10 [1.3]	10 [1.1]	4 [1.5]	7 [1.1]	9 [1.2]
Zymbal's Gland	(0)	(0)	(0)	(0)	(0)	(0)
URINARY SYSTEM						
Kidney	(51)	(50)	(49)	(50)	(50)	(50)
Bilateral, Hydronephrosis						1 [3.0]
Cyst	15	16	8	7	11	16
Degeneration, Mucoid			1 [1.0]	1 [2.0]	1 [2.0]	1 [1.0]
Epithelium, Pelvis, Hyperplasia	1 [1.0]					1 [2.0]
Hemorrhage	1 [3.0]			1 [2.0]		
Hydronephrosis			1 [3.0]	1 [2.0]		1 [3.0]
Infiltration Cellular, Lymphocyte				1 [1.0]		
Inflammation	1 [2.0]	1 [3.0]			1 [3.0]	
Mineralization	31 [1.8]	33 [2.0]	27 [1.7]	34 [1.7]	30 [2.1]	36 [1.7]
Nephropathy	27 [1.2]	21 [1.4]	18 [1.3]	17 [1.1]	19 [1.3]	19 [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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1 0PPB	F1 2PPB	F1 10 PPB	F1 50 PPB	F1/2PPB TO CTL	F1/10PPB TO CTL
Polycystic Kidney			1 [4.0]	1 [2.0]		
Renal Tubule, Accumulation, Hyaline Droplet						1 [2.0]
Renal Tubule, Hyperplasia, Focal						
Urinary Bladder	(51)	(49)	(49)	(48)	(48)	(50)
Dilatation				1 [3.0]		
Hemorrhage				1 [3.0]		
Infiltration Cellular, Lymphocyte	2 [1.5]					
Inflammation						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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Disposition Summary	
Animals Initially In Study Early Deaths	50
Moribund Sacrifice	22
Natural Death	6
Survivors	
Moribund Sacrifice	
Terminal Sacrifice	22
Animals Examined Microscopically	50
ALIMENTARY SYSTEM	
Esophagus	(50)
Hemorrhage	
Inflammation	
Necrosis	
Intestine Large, Cecum	(50)
Dilatation	
Erosion	
Inflammation, Chronic	
Intestine Large, Colon	(50)
Developmental Malformation	
Inflammation	
Lymphoid Tiss, Inflammation	
Mineralization	
Muscularis, Inflammation, Chronic	
Intestine Large, Rectum	(50)
Inflammation, Chronic	(50)
Intestine Small, Duodenum	(50)
Autolysis	

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Dilatation	
Intestine Small, Ileum	(49)
Autolysis	
Intestine Small, Jejunum	(50)
Autolysis	
Inflammation, Chronic	
Liver	(50)
Angiectasis	3 [2.0]
Basophilic Focus	6
Bile Duct, Hyperplasia	8 [1.4]
Capsule, Hemorrhage, Focal	
Caudate Lobe, Developmental Malformation	1
Centrilobular, Necrosis	
Cyst	
Cyst Multilocular	1
Degeneration, Cystic	1 [2.0]
Eosinophilic Focus	3
Eosinophilic Focus, Multiple	1
Hematopoietic Cell Proliferation	3 [1.0]
Hepatodiaphragmatic Nodule	1
Infiltration Cellular, Lymphocyte	2 [1.0]
Inflammation, Chronic Active	4 [1.0]
Left Lat Lobe, Developmental Malformation	1
Median Lobe, Developmental Malformation	
Mixed Cell Focus	1
Necrosis	
Oval Cell, Hyperplasia	
Periportal, Inflammation, Chronic	2 [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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Right Lat Lobe, Developmental Malformation	
Tension Lipidosis	
Vacuolization Cytoplasmic	13 [2.0]
Mesentery	(0)
Cyst	
Fat, Necrosis	
Oral Mucosa	(4)
Gingival, Inflammation	
Keratin Cyst	3
Pancreas	(50)
Acinar Cell, Degeneration	23 [1.9]
Acinar Cell, Hyperplasia	
Artery, Inflammation, Chronic	
Artery, Mineralization	
Inflammation, Chronic	2 [1.0]
Lipomatosis	1 [2.0]
Salivary Glands	(50)
Artery, Mineralization	
Inflammation, Chronic	1 [3.0]
Stomach, Forestomach	(50)
Edema	1 [2.0]
Hyperplasia	
Inflammation	1 [3.0]
Keratin Cyst	1
Ulcer	
Stomach, Glandular	(50)
Glands, Dilatation	
Infiltration Cellular, Lymphocyte	
Inflammation	

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Mineralization	
Tooth	(0)
Dysplasia	
CARDIOVASCULAR SYSTEM	
Blood Vessel	(50)
Mineralization	
Heart	(50)
Artery, Mineralization	
Cardiomyopathy	26 [1.3]
Endocardium, Hyperplasia	1 [2.0]
Myocardium, Necrosis	
ENDOCRINE SYSTEM	
Adrenal Cortex	(50)
Angiectasis	1 [1.0]
Atrophy	1 [2.0]
Cyst	
Degeneration, Cystic	42 [2.2]
Hematopoietic Cell Proliferation	
Hyperplasia	1 [2.0]
Hypertrophy	5 [1.8]
Metaplasia, Osseous	
Vacuolization Cytoplasmic	
Adrenal Medulla	(49)
Hyperplasia	2 [1.5]
Islets, Pancreatic	(50)
Hyperplasia	
Parathyroid Gland	(46)
Hyperplasia	

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Pituitary Gland	(49)
Angiectasis	1 [1.0]
Cyst	
Pars Distalis, Cyst	2
Pars Distalis, Hyperplasia	1 [2.0]
Pars Intermed, Cyst	1
Pigmentation	
Thyroid Gland	(50)
C Cell, Hyperplasia	4 [3.0]
Follicle, Cyst	
Follicular Cel, Hyperplasia	
Infiltration Cellular, Lymphocyte	
Inflammation, Chronic	1 [2.0]
Ultimobranchial Cyst	2 [3.0]
GENERAL BODY SYSTEM	
Tissue NOS	(0)
GENITAL SYSTEM	
Clitoral Gland	(49)
Atrophy	2 [2.5]
Duct, Dilatation	9 [2.3]
Duct, Hyperplasia, Squamous	5 [2.4]
Hyperplasia	3 [2.0]
Infiltration Cellular, Lymphocyte	
Inflammation	34 [2.1]
Keratin Cyst	2
Ovary	(50)
Angiectasis	
Atrophy	42 [2.9]

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

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Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
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**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Corpus Luteum, Cyst	2
Cyst	17
Granulosa Cell, Hyperplasia	
Hyperplasia, Sertoliform	28 [1.7]
Interstit Cell, Hyperplasia	
Oviduct	(50)
Pigmentation	
Uterus	(50)
Atrophy	
Cervix, Hyperplasia	
Cervix, Muscularis, Hypertrophy	1
Dilatation	
Endometrium, Hyperplasia	4 [1.8]
Endometrium, Hyperplasia, Cystic	22 [2.2]
Hemorrhage	
Hyperplasia, Atypical, Focal	9 [1.7]
Hypoplasia	
Inflammation	3 [2.7]
Metaplasia, Squamous	1 [1.0]
Pigmentation, Focal	
Vagina	(50)
Hemorrhage	1 [2.0]
Inflammation	18 [1.6]
Mucocyte, Hyperplasia	39 [2.3]
HEMATOPOIETIC SYSTEM	
Bone Marrow	(50)
Hyperplasia	1 [2.0]
Myeloid Cell, Hyperplasia	1 [3.0]
Lymph Node	(13)
<b>7</b> 1	· -/

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Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

Lab: NCTR

### CD Rat FFMALE

## F1/50PPR TO CTI

CD Rat FEMALE	F1/50PPB TO CTL
Axillary, Degeneration, Cystic	
Axillary, Hyperplasia, Lymphoid	
Axillary, Infiltration Cellular, Plasma Cell	
Deep Cervical, Hyperplasia, Lymphoid	
Deep Cervical, Infiltration Cellular, Plasma Cell	
Degeneration, Cystic	
Hyperplasia, Lymphoid	
Infiltration Cellular, Plasma Cell	
Inguinal, Degeneration, Cystic	2 [2.0]
Inguinal, Hyperplasia, Lymphoid	2 [2.5]
Inguinal, Infiltration Cellular, Plasma Cell	2 [3.0]
Lumbar, Degeneration, Cystic	5 [3.0]
Lumbar, Hemorrhage	
Lumbar, Hyperplasia, Lymphoid	7 [2.3]
Lumbar, Infiltration Cellular, Plasma Cell	9 [2.8]
Mediastinal, Hemorrhage	1 [2.0]
Mediastinal, Hyperplasia, Lymphoid	
Mediastinal, Infiltration Cellular, Plasma Cell	2 [2.5]
Mediastinal, Pigmentation	
Mediastinal, Sinus, Dilatation	
Pancreatic, Hemorrhage	
Pancreatic, Hyperplasia, Lymphoid	
Pancreatic, Pigmentation	
Popliteal, Degeneration, Cystic	
Popliteal, Hyperplasia, Lymphoid	
Popliteal, Infiltration Cellular, Plasma Cell	1 [2.0]
Renal, Degeneration, Cystic	3 [2.7]
Renal, Hyperplasia, Lymphoid	2 [2.0]
Renal, Infiltration Cellular, Plasma Cell	4 [2.8]

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

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Thoracic, Infiltration Cellular, Plasma Cell	
Lymph Node, Mandibular	(50)
Degeneration, Cystic	12 [2.3]
Hemorrhage	1 [2.0]
Hyperplasia, Lymphoid	5 [2.0]
Infiltration Cellular, Plasma Cell	41 [2.4]
Inflammation, Suppurative	
Lymph Node, Mesenteric	(50)
Degeneration, Cystic	5 [2.0]
Depletion Lymphoid	1 [2.0]
Hemorrhage	
Hyperplasia, Lymphoid	7 [2.0]
Infiltration Cellular, Plasma Cell	5 [1.8]
Pigmentation	
Sinus, Dilatation	
Spleen	(50)
Hematopoietic Cell Proliferation	10 [1.6]
Hyperplasia, Lymphoid	
Hyperplasia, Stromal	
Lymphocyte, Atrophy	
Pigmentation	30 [1.8]
Red Pulp, Atrophy	
Thymus	(48)
Cyst	17
Hemorrhage	
Hyperplasia, Lymphoid	
Inflammation, Chronic	
INTEGUMENTARY SYSTEM	
Mammary Gland	(50)
•	` '

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Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Alveolus, Hyperplasia	45 [2.2]
Atypical Focus	18 [1.4]
Duct, Dilatation	4 [2.0]
Duct, Inflammation	1 [2.0]
Fibrosis	1 [3.0]
Galactocele	3
Lactation	20 [2.0]
Lobules, Hyperplasia	
Skin	(50)
Abscess	
Cyst Epithelial Inclusion	
Hyperkeratosis	
Inflammation	29 [3.8]
MUSCULOSKELETAL SYSTEM	
Bone, Femur	(50)
Fibrous Osteodystrophy	, ,
Osteopetrosis	
Skeletal Muscle	(1)
Head, Hyalinization, Focal	
NERVOUS SYSTEM	
Brain, Brain Stem	(50)
Compression	19 [2.4]
Hemorrhage	19 [2.4]
Hydrocephalus	1 [2.0]
Brain, Cerebellum	(50)
Hemorrhage	1 [3.0]
Hydrocephalus	2 [1.5]
Brain, Cerebrum	(50)
a,a	(00)

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)

Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:02

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:02

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
Hemorrhage	
Hydrocephalus	3 [2.0]
Peripheral Nerve	(0)
RESPIRATORY SYSTEM	
Lung	(50)
Alveolar Epith, Hyperplasia	
Artery, Mineralization	1 [1.0]
Autolysis	
Edema	1 [2.0]
Hemorrhage	1 [2.0]
Infiltration Cellular, Histiocyte	8 [2.0]
Infiltration Cellular, Lymphocyte	1 [1.0]
Inflammation	3 [1.7]
Peribronchial, Inflammation, Chronic	
Perivascular, Inflammation	
Nose	(50)
Fibrous Osteodystrophy	
Foreign Body	1
Goblet Cell, Hyperplasia	
Hemorrhage	
Inflammation	5 [2.2]
Nasolacrim Dct, Inflammation	33 [1.8]
Olfactory Epi, Hyaline Droplet	2 [1.5]
Osteopetrosis	
Respirat Epith, Hyaline Droplet	
Respirat Epith, Hyperplasia	
Trachea	(50)
Inflammation	6 [1.2]

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Route: DOSED FEED

Species/Strain: Rat/CD

# P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH AVERAGE SEVERITY GRADES[b] Date Report Requested: 10/23/2014 Time Report Requested: 13:19:03

**Test Compound:** ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number:** 57-63-6

Time Report Requested: 13:19:03

First Dose M/F: NA / NA

CD Rat FEMALE	F1/50PPB TO CTL
SPECIAL SENSES SYSTEM	
Ear	(0)
Eye	(47)
Bilateral, Cataract	
Bilateral, Retina, Degeneration	3 [2.0]
Cataract	
Cornea, Inflammation	2 [1.5]
Retina, Autolysis	
Retina, Degeneration	2 [1.0]
Harderian Gland	(50)
Atrophy, Focal	
Epithelium, Degeneration	
Hyperplasia	4 [1.0]
Infiltration Cellular, Lymphocyte	1 [1.0]
Inflammation	8 [1.0]
Zymbal's Gland	(1)
URINARY SYSTEM	
Kidney	(50)
Bilateral, Hydronephrosis	
Cyst	9
Degeneration, Mucoid	
Epithelium, Pelvis, Hyperplasia	
Hemorrhage	
Hydronephrosis	
Infiltration Cellular, Lymphocyte	1 [2.0]
Inflammation	
Mineralization	35 [1.9]
Nephropathy	20 [1.1]

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Route: DOSED FEED

Species/Strain: Rat/CD

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

Date Report Requested: 10/23/2014
Time Report Requested: 13:19:03

Test Compound: ENDOCRINE DISRUPTOR(Ethinyl Estradiol)

**CAS Number: 57-63-6** 

Time Report Requested: 13:19:03

First Dose M/F: NA / NA

Lab: NCTR

CD Rat FEMALE	F1/50PPB TO CTL
Polycystic Kidney	1 [2.0]
Renal Tubule, Accumulation, Hyaline Droplet	
Renal Tubule, Hyperplasia, Focal	1 [2.0]
Urinary Bladder	(49)
Dilatation	
Hemorrhage	
Infiltration Cellular, Lymphocyte	
Inflammation	

\*\* END OF REPORT \*\*

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