Test Type: CHRONIC

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

Metal working fluids (Trim VX)

Route: RESPIRATORY EXPOSURE WHOLE BODY Species/Strain: RATS/Wistar Han

**CAS Number:** TRIMVX

Lab: BNW

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41

First Dose M/F: 07/20/09 / 07/20/09

F1\_RE

**NTP Study Number:** C20523

Lock Date: 06/04/2012

**Cage Range:** ALL

**Date Range:** ALL

**Reasons For Removal:** ALL

**Removal Date Range:** ALL

Include ALL **Treatment Groups:** 

**Study Gender:** Both

3.0.2.2\_002 **TDMSE Version:** 

**PWG Approval Date:** NONE

Species/Strain: RATS/Wistar Han

Route: RESPIRATORY EXPOSURE WHOLE BODY

Test Type: CHRONIC

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

Metal working fluids (Trim VX)

**CAS Number:** TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS MALE	Control	10 mg/m3	30 mg/m3	100 mg/m3
Disposition Summary				
Animals Initially In Study	50	50	50	50
Early Deaths				
Moribund Sacrifice	12	10	14	15
Natural Death	2	1	3	1
Survivors				
Terminal Sacrifice	36	39	33	34
Animals Examined Microscopically	50	50	50	50
ALIMENTARY SYSTEM				
Esophagus	(50)	(50)	(50)	(50)
Intestine Large, Cecum	(50)	(50)	(50)	(50)
Inflammation	1 (2%)			1 (2%)
Artery, Inflammation	1 (2%)			
Lymphoid Tissue, Hyperplasia				1 (2%)
Intestine Large, Colon	(50)	(50)	(50)	(50)
Lymphoid Tissue, Hyperplasia			1 (2%)	
Intestine Large, Rectum	(50)	(50)	(50)	(50)
Intestine Small, Duodenum	(50)	(50)	(50)	(50)
Artery, Inflammation	1 (2%)			
Intestine Small, Ileum	(50)	(50)	(50)	(50)
Intestine Small, Jejunum	(50)	(50)	(50)	(50)
Liver	(50)	(50)	(50)	(50)
Angiectasis	5 (10%)	2 (4%)		4 (8%)
Basophilic Focus	5 (10%)	10 (20%)	15 (30%)	9 (18%)
Clear Cell Focus	14 (28%)	23 (46%)	17 (34%)	23 (46%)
Eosinophilic Focus	2 (4%)			2 (4%)
Fatty Change			1 (2%)	1 (2%)
Hepatodiaphragmatic Nodule	1 (2%)	2 (4%)		1 (2%)
Infiltration Cellular, Lymphocyte		1 (2%)		
Inflammation, Granulomatous		1 (2%)		2 (4%)
Mixed Cell Focus		1 (2%)	2 (4%)	1 (2%)
Bile Duct, Cyst			2 (4%)	1 (2%)
Bile Duct, Dilatation	1 (2%)			2 (4%)

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

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS MALE	Control	10 mg/m3	30 mg/m3	100 mg/m3	
Bile Duct, Hyperplasia	2 (4%)	2 (4%)	1 (2%)	1 (2%)	
Serosa, Fibrosis			1 (2%)		
Mesentery	(2)	(2)	(3)	(5)	
Fat, Hemorrhage			1 (33%)		
Fat, Necrosis	2 (100%)	2 (100%)	2 (67%)	4 (80%)	
Pancreas	(50)	(50)	(50)	(50)	
Inflammation	1 (2%)	1 (2%)			
Acinus, Atrophy	7 (14%)	2 (4%)	9 (18%)	2 (4%)	
Acinus, Hyperplasia			1 (2%)		
Artery, Inflammation	1 (2%)				
Salivary Glands	(50)	(50)	(50)	(50)	
Stomach, Forestomach	(50)	(50)	(50)	(50)	
Cyst	1 (2%)	, ,	, ,	, ,	
Inflammation	1 (2%)				
Stomach, Glandular	(50)	(50)	(50)	(50)	
Mineralization	,	,	,	1 (2%)	
Artery, Inflammation	1 (2%)			, ,	
CARDIOVASCULAR SYSTEM					
Blood Vessel	(50)	(50)	(50)	(50)	
Heart	(50)	(50)	(50)	(50)	
Cardiomyopathy	21 (42%)	17 (34%)	22 (44%)	18 (36%)	
Endocardium, Hyperplasia	,	,	2 (4%)	,	
ENDOCRINE SYSTEM					
Adrenal Cortex	(50)	(50)	(50)	(50)	
Angiectasis	4 (8%)	1 (2%)	1 (2%)	3 (6%)	
Hypertrophy	13 (26%)	18 (36%)	13 (26%)	15 (30%)	
Necrosis	2 (4%)	,	,	,	
Thrombosis	, ,		1 (2%)		
Vacuolization Cytoplasmic	19 (38%)	23 (46%)	21 (42%)	21 (42%)	
Adrenal Medulla	(50)	(50)	(49)	(49)	
Hyperplasia	1 (2%)	1 (2%)	` '	1 (2%)	

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Lab: BNW

Wistar Han RATS MALE	Control	10 mg/m3	30 mg/m3	100 mg/m3
Islets, Pancreatic	(50)	(50)	(50)	(50)
Parathyroid Gland	(39)	(44)	(46)	(49)
Fibrosis	1 (3%)			
Hyperplasia		1 (2%)	1 (2%)	
Pituitary Gland	(50)	(50)	(50)	(50)
Angiectasis	1 (2%)			
Cyst	2 (4%)	2 (4%)	4 (8%)	1 (2%)
Pars Distalis, Hyperplasia	12 (24%)	15 (30%)	8 (16%)	11 (22%)
Pars Intermedia, Hyperplasia			3 (6%)	
Thyroid Gland	(50)	(50)	(50)	(50)
C-cell, Hyperplasia	36 (72%)	39 (78%)	34 (68%)	40 (80%)
Follicle, Hyperplasia	4 (8%)	7 (14%)	4 (8%)	1 (2%)

#### **GENERAL BODY SYSTEM**

None

GENITAL SYSTEM				
Epididymis	(50)	(50)	(50)	(50)
Granuloma Sperm		1 (2%)		
Preputial Gland	(49)	(50)	(50)	(50)
Cyst	1 (2%)			
Fibrosis				1 (2%)
Inflammation	17 (35%)	13 (26%)	11 (22%)	16 (32%)
Prostate	(50)	(50)	(50)	(50)
Atrophy		1 (2%)		
Fibrosis		1 (2%)		
Inflammation	17 (34%)	12 (24%)	10 (20%)	10 (20%)
Seminal Vesicle	(50)	(50)	(50)	(50)
Atrophy				1 (2%)
Fibrosis	1 (2%)			
Inflammation	4 (8%)		1 (2%)	
Testes	(50)	(50)	(50)	(50)
Cyst	, ,		. ,	1 (2%)

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

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS MALE	Control	10 mg/m3	30 mg/m3	100 mg/m3
Edema	3 (6%)	2 (4%)	1 (2%)	2 (4%)
Artery, Inflammation, Chronic	3 (6%)	,	,	1 (2%)
Germinal Epithelium, Atrophy	3 (6%)	8 (16%)		6 (12%)
Germinal Epithelium, Hypoplasia	,	,	1 (2%)	,
Interstitial Cell, Hyperplasia	2 (4%)		, ,	1 (2%)
HEMATOPOIETIC SYSTEM				
Bone Marrow	(50)	(50)	(50)	(50)
Lymph Node	(11)	(7)	(7)	(8)
Axillary, Hyperplasia, Plasma Cell		1 (14%)		
Iliac, Hyperplasia, Plasma Cell	1 (9%)	2 (29%)	4 (57%)	1 (13%)
Lumbar, Hyperplasia, Lymphoid				1 (13%)
Lumbar, Hyperplasia, Plasma Cell	7 (64%)	5 (71%)	3 (43%)	4 (50%)
Renal, Hyperplasia, Plasma Cell	2 (18%)			2 (25%)
Lymph Node, Bronchial	(39)	(43)	(42)	(42)
Hyperplasia, Lymphohistiocytic		17 (40%)	29 (69%)	35 (83%)
Lymph Node, Mandibular	(46)	(46)	(46)	(48)
Hyperplasia, Plasma Cell	1 (2%)	1 (2%)		
Lymph Node, Mediastinal	(43)	(48)	(44)	(43)
Congestion				1 (2%)
Hyperplasia, Lymphohistiocytic		20 (42%)	22 (50%)	32 (74%)
Lymph Node, Mesenteric	(50)	(50)	(50)	(50)
Hemorrhage			1 (2%)	
Hyperplasia, Lymphohistiocytic			. ,	1 (2%)
Spleen	(50)	(50)	(50)	(50)
Congestion	,	` '	1 (2%)	, ,
Cyst		1 (2%)	, ,	
Hematopoietic Cell Proliferation	7 (14%)	9 (18%)	8 (16%)	8 (16%)
Hemorrhage	, ,	1 (2%)	, ,	, ,
Lymphoid Follicle, Atrophy	3 (6%)	, ,	2 (4%)	
Thymus	(48)	(49)	(50)	(50)
Atrophy	35 (73%)	36 (73%)	26 (52%)	26 (52%)
Cyst	( /	( /	1 (2%)	(/

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS MALE	Control	10 mg/m3	30 mg/m3	100 mg/m3	
INTEGUMENTARY SYSTEM					
Mammary Gland	(4)	(5)	(1)	(2)	
Galactocele	1 (25%)	, ,	, ,	. ,	
Hyperplasia	1 (25%)				
Skin	(50)	(50)	(50)	(50)	
Cyst Epithelial Inclusion	2 (4%)	2 (4%)	1 (2%)	1 (2%)	
Hyperkeratosis	1 (2%)	,	,	,	
Inflammation	2 (4%)	1 (2%)		4 (8%)	
Ulcer	16 (32%)	14 (28%)	13 (26%)	13 (26%)	
MUSCULOSKELETAL SYSTEM					
Bone	(50)	(50)	(50)	(50)	
Hyperostosis	, ,	, ,	` ,	1 (2%)	
Skeletal Muscle	(4)	(4)	(8)	(3)	
NERVOUS SYSTEM					
Brain	(50)	(50)	(50)	(50)	
Compression	8 (16%)	6 (12%)	8 (16%)	7 (14%)	
Degeneration	1 (2%)	,	,	,	
Gliosis	1 (2%)				
Hemorrhage	1 (2%)				
Hydrocephalus	5 (10%)	3 (6%)	8 (16%)	4 (8%)	
Necrosis	,	,	1 (2%)	,	
Meninges, Inflammation	1 (2%)		,		
Peripheral Nerve	(4)	(2)	(8)	(2)	
Axon, Degeneration	\ /	( )	2 (25%)	<b>、</b> /	
Spinal Cord	(4)	(2)	(7)	(2)	
RESPIRATORY SYSTEM					
Larynx	(50)	(50)	(50)	(50)	

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

Experiment Number: 20523 - 03

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS MALE	Control	10 mg/m3	30 mg/m3	100 mg/m3
Foreign Body	1 (2%)		1 (2%)	
Infiltration Cellular, Mixed Cell	1 (2%)	9 (18%)	27 (54%)	31 (62%)
Inflammation, Chronic Active	1 (2%)	1 (2%)		
Epiglottis, Hyperplasia, Squamous		26 (52%)	48 (96%)	50 (100%)
Epiglottis, Metaplasia, Squamous	3 (6%)	50 (100%)	50 (100%)	50 (100%)
Lung	(50)	(50)	(50)	(50)
Fibrosis	4 (8%)	43 (86%)	45 (90%)	49 (98%)
Foreign Body			1 (2%)	
Hyperplasia, Lymphohistiocytic		1 (2%)	1 (2%)	
Infiltration Cellular, Histiocyte	14 (28%)	50 (100%)	50 (100%)	50 (100%)
Inflammation, Chronic Active	7 (14%)	46 (92%)	46 (92%)	48 (96%)
Alveolar/bronchiolar Epithelium, Hyperplasia	4 (8%)	22 (44%)	39 (78%)	46 (92%)
Alveolar Epithelium, Hyperplasia	11 (22%)	43 (86%)	45 (90%)	49 (98%)
Alveolar Epithelium, Metaplasia, Squamous				5 (10%)
Alveolus, Proteinosis		1 (2%)	31 (62%)	45 (90%)
Bronchus-associated Lymphoid Tissue, Hyperplasia, Lymphohistiocytic		1 (2%)	4 (8%)	6 (12%)
Mediastinum, Inflammation, Chronic		1 (2%)		
Nose	(50)	(50)	(50)	(50)
Foreign Body	1 (2%)		1 (2%)	1 (2%)
Inflammation, Suppurative	7 (14%)	46 (92%)	47 (94%)	46 (92%)
Glands, Olfactory Epithelium, Hyperplasia		43 (86%)	41 (82%)	49 (98%)
Goblet Cell, Hyperplasia	5 (10%)	36 (72%)	37 (74%)	42 (84%)
Goblet Cell, Nasolacrimal Duct, Metaplasia		1 (2%)		
Goblet Cell, Olfactory Epithelium, Hyperplasia			1 (2%)	
Nasolacrimal Duct, Inflammation, Chronic		1 (2%)		
Nasolacrimal Duct, Metaplasia, Squamous		1 (2%)		
Olfactory Epithelium, Accumulation, Hyaline Droplet	20 (40%)	50 (100%)	50 (100%)	50 (100%)
Olfactory Epithelium, Atrophy		1 (2%)	1 (2%)	
Olfactory Epithelium, Metaplasia, Squamous		1 (2%)		1 (2%)
Respiratory Epithelium, Accumulation, Hyaline Droplet	10 (20%)	50 (100%)	48 (96%)	50 (100%)
Respiratory Epithelium, Hyperplasia	2 (4%)	7 (14%)	7 (14%)	19 (38%)
Transitional Epithelium, Hyperplasia	7 (14%)	7 (14%)	8 (16%)	13 (26%)
Transitional Epithelium, Metaplasia, Squamous	2 (4%)	2 (4%)	3 (6%)	6 (12%)
Trachea	(50)	(50)	(50)	(50)

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Vistar Han RATS MALE	Control	10 mg/m3	30 mg/m3	100 mg/m3	
SPECIAL SENSES SYSTEM					
Eye	(50)	(50)	(50)	(50)	
Cataract			1 (2%)	1 (2%)	
Inflammation			1 (2%)		
Retina, Atrophy	5 (10%)	5 (10%)	1 (2%)	3 (6%)	
Harderian Gland	(50)	(50)	(50)	(50)	
Cyst		1 (2%)			
Hyperplasia	1 (2%)	1 (2%)		1 (2%)	
Inflammation	1 (2%)	1 (2%)	1 (2%)	2 (4%)	
Lacrimal Gland	(1)	(0)	(0)	(1)	
Atrophy	1 (100%)	, ,	, ,	1 (100%)	
Zymbal's Gland	(1)	(0)	(0)	(0)	
JRINARY SYSTEM					
Kidney	(50)	(50)	(50)	(50)	
Cyst	(00)	(00)	1 (2%)	(55)	
Hydronephrosis	1 (2%)		1 (2%)	3 (6%)	
Infarct	(=75)	1 (2%)	2 (4%)	2 (273)	
Inflammation, Granulomatous		(=75)	_ ( · · · · )	1 (2%)	
Metaplasia, Lipocyte			1 (2%)	1 (2%)	
Necrosis		1 (2%)	(=)	(/	
Nephropathy	27 (54%)	31 (62%)	27 (54%)	28 (56%)	
Pelvis, Inflammation	19 (38%)	17 (34%)	5 (10%)	9 (18%)	
Renal Tubule, Cyst	- ()	1 (2%)	1 (2%)	(,	
Renal Tubule, Hyperplasia		(/	1 (2%)		
Urinary Bladder	(50)	(50)	(50)	(50)	
Hemorrhage	(/	(/	1 (2%)	(/	
•		. ()			
Inflammation		1 (2%)	1 (2%)		

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

**Species/Strain:** RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Lab: BNW

Wistar Han RATS MALE Control 10 mg/m3 30 mg/m3 100 mg/m3

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

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

Experiment Number: 20523 - 03

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS FEMALE	Control	10 mg/m3	30 mg/m3	100 mg/m3
Disposition Summary				
Animals Initially In Study	50	50	50	50
Early Deaths				
Moribund Sacrifice	19	15	15	16
Natural Death	1	2	2	4
Survivors				
Moribund Sacrifice	1	1		
Terminal Sacrifice	29	32	33	30
Animals Examined Microscopically	50	50	50	50
ALIMENTARY SYSTEM				
Esophagus	(50)	(50)	(50)	(50)
Intestine Large, Cecum	(50)	(50)	(50)	(50)
Intestine Large, Colon	(50)	(50)	(50)	(50)
Intestine Large, Rectum	(50)	(50)	(50)	(50)
Intestine Small, Duodenum	(50)	(50)	(50)	(50)
Intestine Small, Ileum	(50)	(50)	(50)	(50)
Intestine Small, Jejunum	(50)	(50)	(50)	(50)
Liver	(50)	(50)	(50)	(50)
Angiectasis		4 (8%)	1 (2%)	
Basophilic Focus	21 (42%)	30 (60%)	24 (48%)	25 (50%)
Clear Cell Focus	6 (12%)	16 (32%)	4 (8%)	8 (16%)
Eosinophilic Focus	2 (4%)	3 (6%)		2 (4%)
Fatty Change	2 (4%)		3 (6%)	
Fibrosis		1 (2%)		
Hematopoietic Cell Proliferation	1 (2%)			
Hepatodiaphragmatic Nodule			1 (2%)	1 (2%)
Inflammation, Granulomatous	1 (2%)	3 (6%)		2 (4%)
Mixed Cell Focus	3 (6%)	5 (10%)		
Necrosis			1 (2%)	
Bile Duct, Cyst			2 (4%)	1 (2%)
Bile Duct, Dilatation		1 (2%)	1 (2%)	
Bile Duct, Hyperplasia	2 (4%)	8 (16%)	6 (12%)	4 (8%)
Centrilobular, Degeneration	1 (2%)			

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

Species/Strain: RATS/Wistar Han

Route: RESPIRATORY EXPOSURE WHOLE BODY

Test Type: CHRONIC

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

Metal working fluids (Trim VX) **CAS Number:** TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS FEMALE	Control	10 mg/m3	30 mg/m3	100 mg/m3	
Serosa, Fibrosis				1 (2%)	
Sinusoid, Dilatation	1 (2%)			,	
Mesentery	(4)	(5)	(9)	(4)	
Fat, Necrosis	4 (100%)	4 (80%)	7 (78%)	4 (100%)	
Pancreas	(50)	(50)	(50)	(50)	
Hyperplasia	1 (2%)	, ,	, ,	, ,	
Acinus, Atrophy	4 (8%)	2 (4%)	2 (4%)	1 (2%)	
Salivary Glands	(50)	(50)	(50)	(50)	
Sublingual Gland, Inflammation	` '	1 (2%)	,	,	
Stomach, Forestomach	(50)	(50)	(50)	(50)	
Inflammation	2 (4%)	` '	, ,	,	
Epithelium, Hyperplasia	2 (4%)				
Stomach, Glandular	(50)	(50)	(50)	(50)	
Tongue	(0)	(0)	(0)	(1)	
Hyperkeratosis	(-)	(-)	(-)	1 (100%)	
CARDIOVASCULAR SYSTEM			,		
Blood Vessel	(50)	(50)	(50)	(50)	
Heart	(50)	(50)	(50)	(50)	
Cardiomyopathy	7 (14%)	7 (14%)	8 (16%)	1 (2%)	
ENDOCRINE SYSTEM					
Adrenal Cortex	(50)	(50)	(50)	(50)	
Angiectasis	28 (56%)	25 (50%)	28 (56%)	27 (54%)	
Hematopoietic Cell Proliferation	3 (6%)				
Hyperplasia		1 (2%)		1 (2%)	
Hypertrophy	10 (20%)	5 (10%)	15 (30%)	10 (20%)	
Necrosis		·	2 (4%)	1 (2%)	
Vacuolization Cytoplasmic	9 (18%)	8 (16%)	14 (28%)	11 (22%)	
Zona Fasciculata, Atrophy		1 (2%)			
Adrenal Medulla	(50)	(50)	(50)	(49)	
, taronar modana					
Hyperplasia	3 (6%)	1 (2%)	, ,	1 (2%)	

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Lab: BNW

Vistar Han RATS FEMALE	Control	10 mg/m3	30 mg/m3	100 mg/m3
Hyperplasia	1 (2%)			
Parathyroid Gland	(42)	(43)	(47)	(47)
Fibrosis		2 (5%)	1 (2%)	
Hyperplasia	1 (2%)			
Pituitary Gland	(50)	(50)	(50)	(50)
Angiectasis				1 (2%)
Cyst		1 (2%)		
Pars Distalis, Hyperplasia	4 (8%)	13 (26%)	12 (24%)	16 (32%)
Pars Intermedia, Hyperplasia		1 (2%)		
Thyroid Gland	(49)	(50)	(50)	(50)
C-cell, Hyperplasia	40 (82%)	44 (88%)	45 (90%)	42 (84%)
Follicle, Hyperplasia	2 (4%)	4 (8%)	7 (14%)	3 (6%)

#### **GENERAL BODY SYSTEM**

None

GENITAL SYSTEM				
Clitoral Gland	(50)	(48)	(49)	(50)
Cyst				2 (4%)
Fibrosis	1 (2%)			
Inflammation	11 (22%)	12 (25%)	13 (27%)	12 (24%)
Ovary	(50)	(50)	(50)	(50)
Cyst	9 (18%)	10 (20%)	12 (24%)	10 (20%)
Uterus	(50)	(50)	(50)	(50)
Inflammation, Suppurative	1 (2%)			
Thrombosis				1 (2%)
Cervix, Hyperplasia		1 (2%)		
Endometrium, Hyperplasia, Cystic	12 (24%)	11 (22%)	12 (24%)	10 (20%)
Epithelium, Metaplasia, Squamous				1 (2%)
Vagina	(1)	(0)	(0)	(0)

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

Species/Strain: RATS/Wistar Han

Route: RESPIRATORY EXPOSURE WHOLE BODY

Test Type: CHRONIC

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

Metal working fluids (Trim VX)

**CAS Number:** TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS FEMALE	Control	10 mg/m3	30 mg/m3	100 mg/m3	
HEMATOPOIETIC SYSTEM					
Bone Marrow	(50)	(50)	(50)	(50)	
Lymph Node	(4)	(1)	(2)	(1)	
Lumbar, Hyperplasia, Plasma Cell	3 (75%)				
Popliteal, Hyperplasia, Plasma Cell	1 (25%)				
Renal, Hyperplasia, Plasma Cell	1 (25%)				
Lymph Node, Bronchial	(37)	(37)	(44)	(36)	
Hematopoietic Cell Proliferation	1 (3%)				
Hyperplasia, Lymphohistiocytic	1 (3%)	18 (49%)	37 (84%)	31 (86%)	
Hyperplasia, Plasma Cell	1 (3%)				
Lymph Node, Mandibular	(40)	(46)	(47)	(50)	
Hematopoietic Cell Proliferation	1 (3%)				
Hyperplasia, Plasma Cell				1 (2%)	
Lymph Node, Mediastinal	(46)	(49)	(46)	(45)	
Hematopoietic Cell Proliferation	1 (2%)				
Hyperplasia, Lymphohistiocytic		11 (22%)	14 (30%)	28 (62%)	
Hyperplasia, Plasma Cell	5 (11%)		1 (2%)	1 (2%)	
Pigmentation, Hemosiderin				1 (2%)	
Lymph Node, Mesenteric	(50)	(50)	(50)	(50)	
Hyperplasia, Plasma Cell				1 (2%)	
Spleen	(50)	(50)	(50)	(50)	
Cyst				1 (2%)	
Hematopoietic Cell Proliferation	13 (26%)	7 (14%)	11 (22%)	9 (18%)	
Lymphoid Follicle, Atrophy	2 (4%)	2 (4%)	6 (12%)	7 (14%)	
Thymus	(48)	(49)	(49)	(49)	
Atrophy	14 (29%)	12 (24%)	21 (43%)	15 (31%)	
Cyst		1 (2%)	2 (4%)		
Ectopic Thyroid	1 (2%)				
Hyperplasia, Lymphohistiocytic			1 (2%)		
INTEGUMENTARY SYSTEM					
Mammary Gland	(50)	(50)	(50)	(50)	
Hyperplasia	3 (6%)	(/	4 (8%)	3 (6%)	

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Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Wistar Han RATS FEMALE	Control	10 mg/m3	30 mg/m3	100 mg/m3	
Duct, Cyst	4 (8%)	7 (14%)	6 (12%)	2 (4%)	-
Skin	(50)	(50)	(50)	(50)	
Cyst Epithelial Inclusion	1 (2%)		1 (2%)		
Inflammation		1 (2%)	1 (2%)	1 (2%)	
Ulcer	6 (12%)		3 (6%)	2 (4%)	
MUSCULOSKELETAL SYSTEM					
Bone	(50)	(50)	(50)	(50)	
Osteopetrosis	1 (2%)				
Joint, Inflammation	1 (2%)				
Skeletal Muscle	(4)	(3)	(3)	(4)	
NERVOUS SYSTEM					
Brain	(50)	(50)	(50)	(50)	
Compression	17 (34%)	16 (32%)	21 (42%)	17 (34%)	
Degeneration	1 (2%)				
Hydrocephalus	6 (12%)	9 (18%)	4 (8%)	8 (16%)	
Peripheral Nerve	(4)	(3)	(2)	(3)	
Spinal Cord	(4)	(3)	(2)	(3)	
RESPIRATORY SYSTEM					
Larynx	(50)	(50)	(50)	(50)	
Infiltration Cellular, Mixed Cell		9 (18%)	18 (36%)	22 (44%)	
Inflammation, Chronic Active		2 (4%)		1 (2%)	
Epiglottis, Hyperplasia, Squamous	1 (2%)	24 (48%)	41 (82%)	50 (100%)	
Epiglottis, Metaplasia, Squamous		49 (98%)	50 (100%)	50 (100%)	
Lung	(50)	(50)	(50)	(50)	
Fibrosis	5 (10%)	35 (70%)	49 (98%)	50 (100%)	
Hyperplasia, Lymphohistiocytic				1 (2%)	
Infiltration Cellular, Histiocyte	16 (32%)	48 (96%)	50 (100%)	50 (100%)	
Inflammation, Chronic	1 (2%)				

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Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Lab: BNW

Wistar Han RATS FEMALE	Control	10 mg/m3	30 mg/m3	100 mg/m3	
Inflammation, Chronic Active	5 (10%)	46 (92%)	50 (100%)	50 (100%)	
Alveolar/bronchiolar Epithelium, Hyperplasia	2 (4%)	9 (18%)	31 (62%)	50 (100%)	
Alveolar Epithelium, Hyperplasia	8 (16%)	43 (86%)	49 (98%)	50 (100%)	
Alveolar Epithelium, Metaplasia, Squamous		3 (6%)	9 (18%)	21 (42%)	
Alveolus, Proteinosis	1 (2%)	15 (30%)	41 (82%)	48 (96%)	
Bronchus-associated Lymphoid Tissue, Hyperplasia, Lymphohistiocytic		2 (4%)	7 (14%)	10 (20%)	
Nose	(50)	(50)	(49)	(50)	
Inflammation, Suppurative	1 (2%)	46 (92%)	47 (96%)	48 (96%)	
Glands, Olfactory Epithelium, Hyperplasia		32 (64%)	43 (88%)	46 (92%)	
Glands, Olfactory Epithelium, Hyperplasia, Squamous			1 (2%)		
Goblet Cell, Hyperplasia	3 (6%)	36 (72%)	40 (82%)	47 (94%)	
Olfactory Epithelium, Accumulation, Hyaline Droplet	14 (28%)	50 (100%)	49 (100%)	50 (100%)	
Olfactory Epithelium, Metaplasia, Respiratory	1 (2%)				
Olfactory Epithelium, Metaplasia, Squamous				2 (4%)	
Respiratory Epithelium, Accumulation, Hyaline Droplet	5 (10%)	50 (100%)	48 (98%)	50 (100%)	
Respiratory Epithelium, Hyperplasia		7 (14%)	8 (16%)	25 (50%)	
Transitional Epithelium, Hyperplasia	5 (10%)	11 (22%)	4 (8%)	21 (42%)	
Transitional Epithelium, Metaplasia, Squamous	1 (2%)	5 (10%)	1 (2%)	2 (4%)	
Trachea	(49)	(50)	(50)	(50)	
SPECIAL SENSES SYSTEM					
Eye	(50)	(50)	(50)	(50)	
Cataract	,	,	1 (2%)	1 (2%)	
Retina, Atrophy	3 (6%)	2 (4%)	1 (2%)	2 (4%)	
Harderian Gland	(50)	(50)	(50)	(50)	
Atrophy	1 (2%)	ζ/	(/	()	
Inflammation	1 (2%)	3 (6%)	3 (6%)	2 (4%)	
Zymbal's Gland	(0)	(2)	(1)	(1)	

#### **URINARY SYSTEM**

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

Test Type: CHRONIC

Route: RESPIRATORY EXPOSURE WHOLE BODY

Species/Strain: RATS/Wistar Han

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

Metal working fluids (Trim VX)

CAS Number: TRIMVX

Date Report Requested: 10/29/2014 Time Report Requested: 10:15:41 First Dose M/F: 07/20/09 / 07/20/09

Nistar Han RATS FEMALE	Control	10 mg/m3	30 mg/m3	100 mg/m3
Kidney	(50)	(50)	(50)	(50)
Hydronephrosis		2 (4%)		1 (2%)
Infarct	1 (2%)	2 (4%)		
Metaplasia, Lipocyte		1 (2%)		
Nephropathy	4 (8%)	5 (10%)	5 (10%)	7 (14%)
Pelvis, Inflammation	28 (56%)	32 (64%)	41 (82%)	30 (60%)
Renal Tubule, Cyst		1 (2%)	1 (2%)	3 (6%)
Renal Tubule, Hyperplasia		1 (2%)		
Ureter	(0)	(0)	(0)	(1)
Cyst				1 (100%)
Urinary Bladder	(50)	(50)	(50)	(50)

<sup>\*\*\*</sup> END OF REPORT \*\*\*