

S9 Table. Histopathological Findings of Male and Female Rabbits After Intranasal Treatment with Iota-Carrageenan

Tissue ^a	Vehicle		Low Dose		High Dose	
	M	F	M	F	M	F
Liver						
Interstitial fibrosis, diffuse						
Minimal	0	1	0	1	1	0
Mild	0	0	0	1	0	0
Periportal infiltrate(s), mononuclear cell (minimal)	0	2	1	1	0	2
Macrovacuolation, hepatocellular, centrilobular						
Minimal	1	0	0	0	0	0
Mild	0	1	0	0	0	0
Congestion, diffuse						
Minimal	0	2	2	2	1	2
Mild	3	0	0	0	2	1
Kidney						
Basophilic tubules, cortex						
Minimal	1	2	0	0	1	0
Mild	1	0	0	0	0	0
Infiltrate(s), mononuclear cell (minimal)	0	0	0	0	1	0
Mineralized focus(i), papilla/medulla (minimal)	1	1	0	0	0	1
Aorta						
Mineralized focus(i), media (minimal)	0	0	1	1	0	0
Lung with bronchus						
Focus(i) of pneumonitis						
Minimal	1	1	1	1	1	0
Mild	0	0	0	1	0	0
Perivascular infiltrate(s), mononuclear cell (minimal)	0	1	1	0	1	1
Alveolar macrophages, (multi)focal						
Minimal	0	0	2	1	0	1
Mild	0	1	0	0	0	0
Osseous metaplasia, focal (minimal)	0	0	2	0	0	0
Alveolar edema, multifocal						
Minimal	1	0	0	1	0	1
Mild	0	1	0	0	0	0
Congestion, diffuse						
Minimal	1	0	0	1	1	0
Mild	1	0	1	1	1	0
Moderate	1	0	0	1	1	1
Marked	0	1	0	0	0	0
Thymus						
Atrophy / regression (minimal)	1	0	0	0	1	0
Parenchymal haemorrhage(s) (minimal)	0	0	1	1	1	1
Spleen						
Presence of germinal centers (minimal)	1	0	0	0	2	0
Pancreas						
Accessory spleen (present)	1	0	0	1	0	0

Tissue ^a	Vehicle		Low Dose		High Dose	
	M	F	M	F	M	F
Peyer's patch						
Presence of germinal centers						
Mild	2	0	0	1	1	2
Moderate	1	2	3	1	2	0
Marked	0	1	0	1	0	1
Colon						
Mucosal infiltration, mononuclear cell						
Minimal	0	0	1	0	0	0
Mild	0	1	0	0	0	0
Mesenteric lymph node						
Presence of germinal centers						
Minimal	0	1	0	0	1	2
Mild	2	2	3	2	1	1
Moderate	1	0	0	1	0	0
Sinusoidal erythrocytes (minimal)	0	0	0	1	0	0
Adrenal gland						
Cortical haemorrhage(s)						
Minimal	0	0	1	0	0	0
Mild	1	0	0	0	1	0
Accessory cortical tissue, (multi)focal (minimal)	0	0	1	1	0	0
Epididymis						
Intraluminal cellular debris, bilateral						
Minimal	2	n.a.	0	n.a.	1	n.a.
Mild	1	n.a.	1	n.a.	1	n.a.
Moderate	0	n.a.	2	n.a.	1	n.a.
Oligospermia, bilateral						
Minimal	0	n.a.	0	n.a.	1	n.a.
Mild	2	n.a.	1	n.a.	0	n.a.
Moderate	0	n.a.	1	n.a.	0	n.a.
Severe	0	n.a.	1	n.a.	0	n.a.
Seminal vesicle						
Decreased secretory content, diffuse						
Minimal	0	n.a.	0	n.a.	1	n.a.
Mild	2	n.a.	2	n.a.	0	n.a.
Moderate	0	n.a.	1	n.a.	1	n.a.
Decreased secretory content, diffuse						
Minimal	0	n.a.	1	n.a.	1	n.a.
Mild	2	n.a.	1	n.a.	0	n.a.
Moderate	2	n.a.	3	n.a.	2	n.a.
Cervix						
Subepithelial haemorrhage(s) (minimal)	n.a.	1	n.a.	0	n.a.	0
Ovary						
Mesothelial proliferation, multifocal						
Minimal	n.a.	1	n.a.	1	n.a.	1
Mild	n.a.	0	n.a.	0	n.a.	1

Tissue ^a	Vehicle		Low Dose		High Dose	
	M	F	M	F	M	F
Oviduct						
Cyst(s) (mild)	n.a.	1	n.a.	0	n.a.	0
Thyroid gland						
Infiltrate(s), mononuclear cell (minimal)	0	1	0	0	0	0
Cyst(s)						
Minimal	0	1	0	1	1	1
Mild	2	1	3	2	2	2
Moderate	0	1	0	0	0	0
Ectopic thymic tissue						
Minimal	1	1	0	0	0	0
Mild	0	0	2	0	1	0
Moderate	0	0	0	1	0	0
Parathyroid gland						
Cyst(s) (minimal)	0	1	0	0	0	0
Bone marrow, sternum						
Bone marrow fat						
Mild	1	1	1	0	1	2
Moderate	2	2	2	3	2	1
Femoral bone						
Bone marrow fat						
Mild	2	3	1	3	1	2
Moderate	1	0	2	0	2	1
Submandibular gland						
Lobular atrophy/degeneration, focal (minimal)	0	1	0	0	0	0
Mandibular lymph node						
Presence of germinal centers						
Minimal	1	3	0	1	0	3
Mild	0	0	0	0	2	0
Sinusoidal erythrocytes						
Minimal	1	0	0	0	0	0
Mild	0	1	0	0	0	0
Moderate	0	1	1	1	0	0
Cystic dilated sinusoid(s) (minimal)	0	1	0	0	0	0
Black pigment, (multi)focal (minimal)	0	0	0	1	0	0
Mammary gland						
Increased secretory content, diffuse						
Minimal	n.a.	0	n.a.	1	n.a.	2
Mild	n.a.	1	n.a.	0	n.a.	0
Acinar development, diffuse						
Minimal	n.a.	1	n.a.	0	n.a.	2
Mild	n.a.	1	n.a.	0	n.a.	0
Eye						
Conjunctival infiltrate(s), mononuclear cell (minimal)	0	0	0	0	0	1
Retinal rosette(s) (minimal)	0	0	0	1	0	0
Harderian gland						

Tissue ^a	Vehicle		Low Dose		High Dose	
	M	F	M	F	M	F
Acinar metaplasia, focal (minimal)	0	0	1	0	0	0
Tongue						
Muscular infiltrate(s), mixed cell (minimal)	0	0	0	0	0	1
Muscular fibrosis, (multi)focal (mild)	0	0	0	1	0	0
Axillary lymph node						
Sinusoidal erythrocytes (minimal)	0	1/2	0	1	0	0
Nasal cavity, left						
Purulent exudate (minimal)	0	0	1	0	0	0
Crust, squamous epithelium, focal (minimal)	0	0	0	0	1	0
Nasal cavity, right						
Crust, squamous epithelium, focal (minimal)	0	0	0	0	1	0
Nostril, left						
Dermatitis, (multi)focal						
Minimal	2	1	2	0	3	1
Mild	0	1	1	0	0	1
Epidermal hyperplasia, (multi)focal						
Minimal	0	0	1	0	1	0
Mild	0	2	0	0	0	2
Nostril, right						
Dermatitis, (multi)focal						
Minimal	1	1	0	2	1	2
Mild	0	1	1	0	0	0
Epidermal hyperplasia, (multi)focal						
Minimal	0	0	1	1	1	0
Mild	1	0	0	0	0	0

^a only tissues with histopathological findings shown

Vehicle = 0.5% NaCl; Low Dose = 112 µg/kg/day; High Dose = 448 µg/kg/day.