

Supplement Table 1-1. Mortality/Cage Side Observation, male, Phase A

Group No.	Day 1		Day 2		Day 3		Day4
	a.m.	p.m.	a.m.	p.m.	a.m.	p.m.	a.m.
Inauhzin-C Group 1/ ♂ 50 mg/kg	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	-
Inauhzin-C Group 2/ ♂ 150 mg/kg	NAD (3/3)	NAD (3/3)	Piloerection (3/3)	Piloerection (2/3); NAD (1/3)	NAD (3/3)	NAD (3/3)	-
Inauhzin-C Group 3/ ♂ 250 mg/kg	DMA (3/3)	DMA (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	-
Inauhzin-C Group 4/ ♂ 200 mg/kg	DMA (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)

Supplement Table 1-2. Mortality/Cage Side Observation, female, Phase A

Group No.	Day 1		Day 2		Day 3		Day4
	a.m.	p.m.	a.m.	p.m.	a.m.	p.m.	a.m.
Inauhzin-C Group 1/ ♀ 50 mg/kg	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)
Inauhzin-C Group 2/ ♀ 150 mg/kg	NAD (3/3)	NAD (3/3)	Piloerection (3/3)	Mattered hair (1/3); NAD (2/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)
Inauhzin-C Group 3/ ♀ 250 mg/kg	DMA (3/3)	DMA (3/3)	DMA (3/3)	DMA (1/3); piloerection (1/3); Found dead (2/3)	Found dead (1/1)	-	-
Inauhzin-C Group 4/ ♀ 200 mg/kg	DMA (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)

NAD: No Abnormalities Detected

DMA: Decreased Motor Activity

Supplement Table 2-1. Detailed clinical observation, male, Phase A

Group No.	Day 1		Day 2		Day 3		Day 4
	a.m.	p.m.	a.m.	p.m.	a.m.	p.m.	a.m.
Group 1/♂ 50 mg/kg	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)
Group 2/♂ 150 mg/kg	NAD (3/3)	NAD (3/3)	Piloerection (3/3)	Piloerection (2/3); NAD (1/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)
Group 3/♂ 250 mg/kg	NAD (3/3)	DMA (3/3)	DMA (3/3); discharge, both eyes (1/3); both eyes partly closed (1/3)	DMA (3/3); piloerection (3/3); discharge, both eyes (1/3); cold to touch (1/3)	NAD (2/3); discharge, right eye (1/3)	NAD (1/3); staining around anus (1/3); soft stool (1/3)	NAD (3/3)
Group 4/♂ 200 mg/kg	NAD (3/3)	DMA (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)

Supplement Table 2-2. Detailed clinical observation, female, Phase A

Group No.	Day 1		Day 2		Day 3		Day 4
	a.m.	p.m.	a.m.	p.m.	a.m.	p.m.	a.m.
Group 1/♀ 50 mg/kg	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)
Group 2/♀ 150 mg/kg	NAD (3/3)	NAD (3/3)	Piloerection (3/3)	Hair staining, around anus (1/3); NAD (2/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)
Group 3/♀ 250 mg/kg	NAD (3/3)	DMA (3/3)	DMA (3/3); discharge, both eyes (1/3); both eyes partly closed (1/3)	DMA (3/3); Piloerection (3/3); discharge, both eyes (3/3); both eyes partly closed (2/3); cold to touch (3/3)	DMA (3/3); Piloerection (1/3); discharge, both eyes (2/3); eye(s) partly closed (2/3); cold to touch (3/3)	Found dead (2/3); DMA (1/3); piloerection (1/3); deep respiration (1/3)	Found dead (1/1)
Group 4/♀ 200 mg/kg	NAD (3/3)	DMA (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)	NAD (3/3)

NAD: No Abnormalities Detected

DMA: Decreased Motor Activity

Supplement Table 3-1. Necropsy Findings, male, Phase A

Group No	Animal No	External Examination	Internal Examination	Necropsy Date
Inauhzin-C ♂ 50 mg/kg	1101	Normal	Normal	Day 4
	1102	Normal	Normal	Day 4
	1103	Normal	Normal	Day 4
Inauhzin-C ♂ 150 mg/kg	1201	Normal	Normal	Day 4
	1202	Normal	Normal	Day 4
	1203	Normal	Normal	Day 4
Inauhzin-C ♂ 250 mg/kg	1301	Normal	White substance adhered to the surface of liver and diaphragm, large quantity	Day 4
	1301	Normal	White substance adhered to the surface of liver and diaphragm, large quantity	Day 4
	1301	Normal	White substance adhered to the surface of liver and diaphragm, large quantity	Day 4
Inauhzin-C ♂ 200 mg/kg	1401	Normal	White substance adhered to the surface of liver, spleen and diaphragm	Day 5
	1402	Normal	White substance adhered to the surface of liver, spleen and diaphragm	Day 5
	1403	Normal	White substance adhered to the surface of liver and diaphragm; adhesion-stomach and liver	Day 5

Supplement Table 3-2. Necropsy Findings, female, Phase A

Group No	Animal No	External Examination	Internal Examination	Necropsy Date
Inauhzin-C ♀ 50 mg/kg	2101	Normal	Normal	Day 4
	2102	Normal	Normal	Day 4
	2103	Normal	Normal	Day 4
Inauhzin-C ♀ 150 mg/kg	2201	Normal	Normal	Day 4
	2202	Normal	Normal	Day 4
	2203	Normal	Normal	Day 4
Inauhzin-C ♀ 250 mg/kg	2301	Normal	Duodenum-partly autolysis, with content in, dark red; spleen-small, ~1/3 of normal size; liver-discoloration, light; white substance adhered to the surface of liver and spleen; liquid observed in the thoracic cavity, large volume	Day 3
	2301	Normal	Duodenum-with content in, dark red; spleen-small, ~1/3 of normal size; stomach-gaseous distension; liver-partly discoloration, light; white substance adhered to the surface of liver and spleen; liquid observed in the thoracic cavity, large volume	Day 2
	2301	Normal	Duodenum-with content in, dark red; spleen-small, ~1/2 of normal size; liver-discoloration, light with white substance adhered to the surface; liquid observed in the thoracic cavity, large volume	Day 2
Inauhzin-C ♀ 200 mg/kg	2401	Normal	Adhesion-stomach and liver; white substance adhered to liver, diaphragm and mesentery	Day 5
	2402	Normal	White substance adhered to liver and diaphragm	Day 5
	2403	Normal	White substance adhered to liver and diaphragm	Day 5

Supplement Table 4(5). Necropsy Findings, female, Phase B

Group No	Animal No	External findings	Internal findings	Necropsy Date
Control/♀ 0 mg/kg	21001	NAD	NAD	Day 7
	21002	NAD	NAD	Day 7
	21003	NAD	NAD	Day 7
	21004	NAD	NAD	Day 7
	21005	NAD	NAD	Day 7
	21006	NAD	NAD	Day 7
Inauhzin-C Low/♀ 30 mg/kg	22001	NAD	Wall of duodenum- thin	Day 7
	22002	NAD	NAD	Day 7
	22003	NAD	Discoloration at head of spleen, dark	Day 7
	22004	NAD	NAD	Day 7
	22005	NAD	NAD	Day 7
	22006	NAD	Spleen enlargement	Day 7
Inauhzin-C Mid/♀ 60 mg/kg	23001	NAD	NAD	Day 7
	23002	NAD	Spleen enlargement	Day 7
	23003	NAD	NAD	Day 7
	23004	NAD	Discoloration at head of spleen, dark	Day 7
	23005	NAD	NAD	Day 7
	23006	NAD	NAD	Day 7
Inauhzin-C High/♀ 120 mg/kg	24001	NAD	White particles on the abdominal wall, mesentery, liver and diaphragma; edge of liver lobes-blunt; wall of duodenum-thin; liver-adhered to stomach and duodenum; spleen-large	Day 7
	24002	NAD	White particles on the abdominal wall, mesentery, liver and diaphragma; edge of liver lobes-blunt; wall of duodeneum-thin; liver-adhered to stomach, right kidney and duodenum; spleen-large and blunt at the edge	Day 7
	24003	NAD	Liquid observed in abdominal cavity, large volume; white particles on the abdominal wall, mesentery, liver and diaphragma; edge of liver lobes-blunt; wall of duodeneum-thin; liver-adhered to stomach and duodenum; spleen-large; liquid in the duodenum, yellow, thick	Day 7
	24004	NAD	White particles on the abdominal wall, mesentery, liver and diaphragma; edge of liver lobes-blunt; wall of duodenum-thin; liver-adhered to stomach, spleen, right kidney and duodenum	Day 7
	24005	NAD	Liquid observed in abdominal cavity, small volume; white particles on the abdominal wall, mesentery, liver and diaphragma; edge of liver lobes-blunt; wall of duodenum-thin; liver-adhered to stomach and duodenum; spleen-large and blunt at the edge; liquid in the duodenum, light brown thick	Day 7
	24006	NAD	White particles on the abdominal wall, mesentery, liver and diaphragma; edge of liver lobes-blunt; liver-adhered to abdominal wall	Day 7

Supplement Table 5 (6). Histopathology

Group No	Animal NO.	Brain	Heart	Kidneys	Ovaries	Adrenal glands	Liver	Lung	Spleen	Uterus
Control/♀ 0 mg/kg	21001	Brainstem hemorrhage	Normal	Cortex renal tubular epithelial cells undergoing degeneration	Normal	Normal	Infiltration of focal inflammatory cells mainly macrophage	Normal	Normal	Normal
	21002	Hemorrhage in parenchyma	Normal	Cortex renal tubular epithelial cells undergoing degeneration	Normal	Normal	Infiltration of focal inflammatory cells mainly macrophages and neutrophils	Normal	Normal	Normal
	21003	Hemorrhage in parenchyma	Normal	Cortex renal tubular epithelial cells undergoing degeneration	Normal	Normal	Normal	Normal	Normal	Normal
Low/♀ 30 mg/kg	24001	Hemorrhage in parenchyma	Normal	Cortex renal tubular epithelial cells undergoing degeneration	Normal	Normal	Normal	Normal	Normal	Normal
Mid/♀ 60 mg/kg	24002	Hemorrhage in parenchyma	Normal	Cortex renal tubular epithelial cells undergoing degeneration	Normal	Normal	The hepatic capsule thickening within inflammatory cell infiltration mainly neutrophils	Normal	Normal	Normal
High/♀ 120 mg/kg	24003	Parenchyma and subarachnoid hemorrhage	Normal	Cortex renal tubular epithelial cells undergoing degeneration	Normal	Normal	Normal	Normal	Normal	Normal