

Supplementary Table S1. Hematological and biochemical parameters for safety evaluation

Parameters	Control (H ₂ O Gavage)						100 mg/day DOX equiv.						200 mg/day DOX equiv.						300 mg/day DOX equiv.					
	Pre-Surgical		28 dpi		Pre-Surgical		28 dpi		Pre-Surgical		28 dpi		Pre-Surgical		28 dpi		Pre-Surgical		28 dpi		Pre-Surgical		28 dpi	
	Mean	Range	Mean	Range	Mean	Range	Mean	Range	Mean	Range	Mean	Range	Mean	Range	Mean	Range	Mean	Range	Mean	Range	Mean	Range		
RBC (M/ μ l)	9.1	8.55-9.6	9.7	9.41-10.1	9.1	8.83-9.34	9.4	9.01-9.61	9.1	8.65-9.51	9.3	8.8-9.81	8.9	8.66-9.22	9.8	9.63-10.05								
Hemoglobin (g/dL)	16.6	15.7-17.5	17.1	16.6-17.9	16.6	16.1-17.2	16.6	15.8-17.3	16.4	15.5-16.9	16.3	15.5-17.1	16.2	15.5-16.7	16.1	15.8-16.4								
Hematocrit (HCT %) (PCV)	46.4	43.8-49.6	49.3	47.8-52.1	46.2	45.1-48.2	47.3	45.6-49.1	46.4	43.8-48.4	47.1	44.9-49.4	45.4	43.8-46.4	48.5	47.1-51.1								
Mean Corpuscular Volume (MCV fL)	51.0	48.9-51.7	50.9	50.3-51.6	51.1	50.3-51.6	50.5	50.1-51.1	51.0	50.6-51.2	50.7	50.4-51	50.8	50.3-51.5	49.5	48.5-50.8								
MCH (pg)	18.2	17.7-18.4	17.7	17.6-17.8	18.3	18.1-18.6	17.8	17.5-18	18.0	17.8-18.1	17.5	17.2-17.9	18.1	17.9-18.3	16.4	16.3-16.5								
Mean Cell Hemoglobin Concentration (MCHC g/dL)	35.7	35.3-36.1	34.8	34.4-35.3	35.9	35.6-36.4	35.2	34.6-35.6	35.3	34.9-35.4	34.6	34.2-35.2	35.7	35.4-36	33.2	32.1-34								
RDW-SD (fL)	27.0	26.3-27.8	28.5	27.5-30.4	27.1	26.6-27.5	28.9	28.7-29.3	27.7	26.9-29.4	28.3	27.7-28.7	27.6	26.4-29.4	28.6	28.4-28.7								
RDW-CV (%)	22.1	21.4-22.9	23.3	22.6-24.3	22.1	21.5-22.3	23.1	22.6-23.6	22.4	21.9-22.9	22.7	22.3-23.4	22.3	21.7-22.9	23.1	22.7-23.8								
Reticulocytes (K/uL)	400.3	273.6-456	252.3	226.6-280.4	349.1	305.2-382.3	232.8	219.1-245.1	431.4	355-535.2	249.6	225.6-275.7	414.4	305.2-535.2	225.1	213.8-233.2								
Reticulocytes (%)	4.4	2.88-5.06	2.6	2.38-2.9	3.9	3.31-4.33	2.5	2.28-2.65	4.8	3.85-6.18	2.7	2.3-3.06	4.7	3.31-6.18	2.3	2.22-2.35								
Platelets (K/uL)	456.6	288-667	561.4	501-631	539.0	351-639	587.3	574-611	610.8	333-727	575.5	552-610	558.7	312-727	601.0	576-641								
PDW (Platelet Distribution Width fL)	10.5	10-11.1	9.7	8.8-10.1	11.7	9.5-13.1	9.7	9.3-9.9	11.1	9.9-14.3	9.4	9-9.9	10.5	10-11.1	10.5	10-11.3								
MPV (Mean Platelet Volume fL)	8.2	7.9-8.5	7.9	7.7-8.1	8.5	7.9-9.1	7.8	7.6-7.9	8.2	7.8-8.8	7.5	7.5-7.5	8.2	7.8-8.4	7.7	7.4-8.3								
P-LCR (Platelet Larger Cell Ratio %)	16.5	14-19.3	13.0	11.4-14.2	18.6	12.9-22.9	12.4	10.6-13.6	16.0	13.1-21.1	10.1	9.7-10.2	15.9	13.5-17.7	13.2	10.8-17.6								
PCT (Plateletcrit %)	0.4	0.25-0.54	0.4	0.4-0.5	0.5	0.31-0.53	0.5	0.44-0.48	0.5	0.29-0.58	0.4	0.41-0.46	0.5	0.26-0.57	0.5	0.43-0.49								
WBC (K/uL)	15.7	12.28-21.1	6.2	5.67-6.93	15.4	13.31-17.31	6.0	4.94-6.75	12.9	11.87-14.03	6.6	6.13-7.39	15.0	13.41-17.31	7.1	5.77-9.12								
Neutrophils (K/uL)	3.8	1.76-6.48	1.0	0.75-1.37	3.0	2.06-3.77	0.7	0.55-0.92	2.2	1.98-2.43	0.6	0.43-0.82	2.9	2.43-3.52	0.7	0.58-0.85								
Lymphocytes (K/uL)	10.3	9.44-12.47	4.8	4.04-5.39	10.7	9.42-12.2	5.1	4.14-5.66	9.6	9.11-10.48	5.5	4.93-6.51	10.4	9.44-12.2	5.8	4.82-7.84								
Monocytes (K/uL)	1.3	0.93-1.68	0.4	0.29-0.47	1.6	1.33-1.81	0.2	0.08-0.26	0.9	0.7-1.22	0.4	0.36-0.44	1.3	1.22-1.53	0.4	0.32-0.58								
Eosinophils (K/uL)	0.4	0.04-0.76	0.0	0.01-0.05	0.1	0.04-0.09	0.0	0-0.02	0.0	0.04-0.06	0.0	0.01-0.01	0.3	0.04-0.76	0.0	0.01-0.04								
Basophils (K/uL)	0.1	0.01-0.18	0.0	0-0.02	0.1	0.02-0.09	0.0	0-0.02	0.1	0.02-0.08	0.0	0.01-0.01	0.0	0.01-0.06	0.0	0-0.01								
Neutrophils (%)	22.5	14.3-31.3	15.6	12.4-23.4	19.1	15.4-25.1	11.4	9.8-13.7	17.1	16-18.2	10.0	9.9-13.3	19.5	18.2-20.4	10.7	9.3-12.6								
Lymphocytes (%)	66.7	58.7-77.3	77.6	68.7-81.1	69.7	62.6-73.6	84.8	82.5-88.7	75.0	72-76.7	83.7	80.4-88.1	69.5	66.1-72	82.3	77.4-86								
Monocytes (%)	8.2	6-10.8	6.3	5.1-7.1	10.4	8.8-11.5	3.5	1.3-4.7	7.3	5.8-9.1	6.1	5.6-6.9	8.8	8.5-9.1	6.4	4.5-9.3								
Eosinophils (%)	2.3	0.3-5.3	0.4	0.2-0.7	0.5	0.2-0.6	0.2	0-0.3	0.4	0.3-0.4	0.2	0.1-0.2	1.9	0.2-5.3	0.4	0.1-0.7								
Basophils (%)	0.3	0.1-0.9	0.2	0-0.3	0.4	0.1-0.6	0.2	0-0.3	0.4	0.2-0.6	0.2	0.1-0.2	0.2	0.1-0.4	0.2	0.1-0.2								
Glucose (mg/dL)	138.8	130-145	238.6	145-309	138.5	126-157	206.3	153-269	136.0	128-142	219.8	165-253	139.3	131-145	354.0	212-558								
BUN (Blood Urea Nitrogen mg/dL)	47.7	36-58	20.0	19-21	47.8	30-81	19.3	17-22	35.0	30-41	17.8	15-19	39.1	35-43.33	15.3	14-16								
CRE (Creatinine mg/dL)	1.1	0.7-1.4	0.4	0.4-0.4	1.1	0.5-2.1	0.4	0.3-0.4	0.7	0.5-0.9	0.4	0.4-0.4	0.9	0.7-1.07	0.3	0.3-0.4								
PHOS (Phosphorus mg/dL)	10.1	8.3-11.9	8.5	7.5-9.9	10.4	8.7-13.9	7.3	6.8-8	9.6	8.9-10.1	7.6	6.4-8.4	9.4	8.9-9.87	11.1	7.8-15.1								
CA (Calcium mg/dL)	10.4	10.2-10.5	10.7	10.4-11.1	10.1	9.5-10.3	10.2	10-10.5	10.0	9.8-10.3	10.2	9.6-10.5	10.4	10.3-10.5	11.3	9.9-13.3								
TP (Total Protein g/dL)	6.4	6.1-6.7	5.4	5.3-5.6	6.0	5.9-6.1	5.3	5.2-5.4	6.2	6-6.4	5.3	5-5.4	6.2	5.9-6.6	6.0	5.9-6.1								
ALB (Albumin g/dL)	3.4	3.3-3.5	2.7	2.6-2.9	3.3	3.2-3.3	2.7	2.6-2.7	3.3	3.2-3.4	2.6	2.5-2.7	3.3	3.2-3.5	3.3	3-3.6								
GLOB (Globulin g/dL)	3.0	2.8-3.2	2.7	2.6-2.8	2.8	2.7-2.8	2.6	2.6-2.7	2.9	2.8-3	2.7	2.5-2.8	2.9	2.7-3.1	2.8	2.5-3								
ALT (Alanine Aminotransferase U/L)	52.7	47.67-65	50.4	13-74	51.5	38-78	59.5	54-69	39.3	25-50	54.0	38-69	45.9	44-47.67	65.3	58-72								
ALP (Alkaline Phosphatase U/L)	370.1	316.33-445	904.0	311-3187	329.8	250-380	298.5	243-345	341.0	292-379	280.0	261-310	354.1	316.33-380	257.67	172-302								
TBL (Total Bilirubin mg/dL)	1.0	0.4-1.3	0.1	0.1-0.1	1.0	0.6-1.8	0.1	0.1-0.1	1.0	0.6-1.2	0.1	0.1-0.1	0.9	0.6-1.27	0.1	0.1-0.1								
CHOL (Cholesterol mg/dL)	67.7	58.67-73	72.8	71-75	73.0	73-73	79.3	75-81	58.0	58-58	77.8	66-86	63.2	58-73	85.0	84-87								
AMY (Amylase U/L)	1,636.9	1,459.67-1,736	1,278.0	1,088-1,454	1,444.5	1,197-1,567	1,405.3	1,279-1,569	1,415.5	1,382-1,478	1,194.3	1,107-1,361	1,458.6	1,382-1,534	1,386.0	1,327-1,457								
AST (Aspartate Aminotransferase U/L)	109.9	93-134	86.6	62-136	110.3	86-163	64.8	60-73	80.3	68-87	99.5	59-211	88.1	68-110.33	94.7	68-132								

Blood and sera were evaluated to determine the safety of allometrically scaled doses of DOX in rats for 1 month.