

**Supplementary Table S1.** Blood chemistry results by time point and treatment.

Item	Normal Reference Range <sup>1</sup>	Baseline (D0)	SEM	Treatment			SEM	P - Value		
				Con (D63)	Low (D63)	High (D63)		Day	Sex	Trt × Time
Alb, g/dL	2.5 – 4.4	3.9	0.05	3.7	3.7	3.7	0.06	< 0.01	0.01	0.64
Alp, U/L	20 – 150	61	4.1	78	82	76	7.3	< 0.01	0.86	0.74
Alt, U/L	10 – 118	53	1.2	47	53	51	5.3	0.48	0.41	0.73
Amy, U/L	200 – 1200	711	34.4	681	698	607	43	0.05	0.07	0.19
TBil, mg/dL <sup>2</sup>	0.1 – 0.6	0.28	0.04	0.30	0.30	0.30	-	-	-	-
BUN, mg/dL	7 – 25	15	0.40	17	16	16	0.68	< 0.01	0.01	0.30
Ca, mg/dL	8.6 – 11.8	10.4	0.07	10.0	9.95	9.95	0.10	< 0.01	0.50	0.75
Phos, mg/dL	2.9 – 6.6	4.3	0.09	4.3	4.6	4.4	0.14	0.10	0.04	0.33
Cre, mg/dL	0.3 – 1.4	0.98	0.04	0.93	0.87	0.93	0.06	0.07	0.66	0.52
Glu, mg/dL	60 – 110	88	1.1	86	86	88	1.5	0.09	< 0.01	0.56
K <sup>+</sup> , nmol/L	3.7 – 5.8	4.9	0.04	4.6	4.7	4.6	0.07	< 0.01	0.80	0.24
Na <sup>+</sup> , nmol/L	138 – 160	146	0.32	147	147	147	0.64	0.05	0.53	0.77
TP, g/dL	5.4 – 8.2	5.9	0.05	6.0	6.0	5.9	0.07	0.60	0.85	0.23
Glob, g/dL	2.3 – 5.2	2.0	0.05	2.3	2.2	2.2	0.07	< 0.01	0.03	0.32

<sup>1</sup> Zoetis VETSCAN HM5 Operator's Manual; 2018

<sup>2</sup> Statistical analysis for total bilirubin could not be performed as machine reported values were 0.3 for all dogs on day 63 of the study.

Note: Abbreviations: Alb, albumin; Alp, alkaline phosphatase; Alt, alanine aminotransferase; Amy, amylase; TBil, total bilirubin;

BUN, blood urea nitrogen; Ca, calcium; Phos, phosphorus; Cre, creatinine; Glu, glucose; Na<sup>+</sup>, sodium; K<sup>+</sup>, potassium; TP, total protein; Glob, globulin.

**Supplementary Table S2.** Complete blood count results by time point and treatment.

Item	Normal Reference Range <sup>1</sup>	Baseline (D0)	SEM	Treatment			SEM	P - Value		
				Con (D63)	Low (D63)	High (D63)		Day	Sex	Trt × Time
WBC, 10 <sup>9</sup> /L	6 – 17	8.54	0.30	9.52	8.19	8.73	0.42	0.36	0.62	0.08
Lym, 10 <sup>9</sup> /L	1 – 4.8	1.42	0.08	1.59	1.26	1.38	0.10	0.91	0.43	0.08
Mon, 10 <sup>9</sup> /L	0.2 – 1.5	0.38	0.03	0.32	0.32	0.36	0.04	0.19	0.35	0.62
Neu, 10 <sup>9</sup> /L	3 – 12	6.64	0.25	7.43	6.48	6.93	0.36	0.19	0.69	0.17
Eos, 10 <sup>9</sup> /L	0.00 – 0.80	0.07 <sup>a</sup>	0.01	0.13 <sup>b</sup>	0.07 <sup>a</sup>	0.08 <sup>a</sup>	0.01	0.05	0.88	0.01
Bas, 10 <sup>9</sup> /L	0.00 – 0.40	0.02	0.004	0.04	0.02	0.03	0.01	0.06	0.78	0.09
RBC, 10 <sup>12</sup> /L	0 – 100	7.44	0.18	7.32	7.54	7.37	0.24	0.84	0.23	0.38
HGB, g/dL	12 – 18	15.8	0.16	15.3	15.6	15.5	0.24	0.05	0.29	0.69
HCT, %	37 – 55	54.82	1.11	47.27	48.03	47.57	0.69	< 0.01	0.10	0.74
MCV, fL	60 – 77	74	0.83	65	64	65	0.61	< 0.01	0.43	0.28
MCH, pg	19.5 – 24.5	22.5	1.65	20.9	20.7	21.1	0.24	0.32	0.89	0.52
MCHC, g/dL	31 – 39	28.3	0.13	32.3	32.5	32.5	0.20	< 0.01	0.20	0.63
RDW, %	14 – 20	15.6	0.11	17.0	17.0	16.7	0.18	< 0.01	0.95	0.61
PLT, 10 <sup>9</sup> /L	165 – 500	250	8.0	240	272	257	15	0.53	0.10	0.31
MPV, fL	3.9 – 11.1	9.4	0.11	9.8	9.4	9.5	0.18	0.07	0.38	0.13
PCT, %	N/A	0.23	0.01	0.24	0.26	0.24	0.02	0.20	0.08	0.71
PDW, %	N/A	36.8	0.25	38.0	37.6	37.3	0.37	< 0.01	0.97	0.42

<sup>1</sup> Zoetis VETSCAN HM5 Operator's Manual; 2018

Note: Abbreviations: WBC, white blood cells; Lym, lymphocytes; Mon, monocytes; Neu, neutrophils; Eos, eosinophils; Bas, basophils; RBC, red blood cells; HGB, hemoglobin; HCT, hematocrit; MCV, mean corpuscular volume; MCH, mean corpuscular hemoglobin; MCHC, mean corpuscular hemoglobin concentration; RDW, red blood cell distribution; PLT, platelets; MPV, mean platelet volume; PCT, plateletcrit; PDW, platelet distribution width.