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## **Appendix 1**

## Clinical pathology

Reference values (WIL Research Laboratories)

Haematology:

Heinz bodies 0–150  $10^3/\mu$ l Erythrocyte count 7.64–11.6  $10^6/\mu$ l Haemoglobin (PCV) 28.8–50.6%

Haemoglobin 10.1-16.0 g/dl

MCH 12.0-15.0 pg MCHC 29.6-35.3 g/dl MCV 34.4-46.9 fl

Platelet count 134–698 10<sup>3</sup>/µl Reticulocyte count 8.3–69.8 10<sup>3</sup>/µl Total leukocyte count 8.2–22.7 10<sup>3</sup>/µl

Differential leukocyte count
Basophil 0-0.05 10³/μl
Eosinophil 0.14-1.71 10³/μl
Lymphohocyte 2.45-10.91 10³/μl
Neutrophil 2.98-16.34 10³/μl
Monocyte 0.7-4.0 10³/μl

Coagulation factors:

Activated partial thromboplastin (APTT) 11.2-17.7 s

Prothrombin time (PT) 8.0–11.2 s

Fibrinogen 109-168 mg/dl

Urinalysis:

Colour and appearance descriptive

Glucose ± Leukocytes ± Ketones ±

Microscopy of sediment descriptive

Nitrites ±
Occult blood±
pH 6.0–8.5
Protein ±

Specific gravity 1.0380–1.0920 Urobilinogen 0.2–1.0 mg/dl

Volume 1-42 ml

Serum chemistry: A/G ratio 0.81–1.50

Alanine aminotransferase 51-150 U/I

Albumin 3.2-4.0 g/dl

Alkaline phosphatase 28-98 U/I

Amylase 695-1050 U/I

Aspartate aminotransferase 14-59 U/I

Bilirubin

Calcium 9.7–11.5 mg/dl Chloride 114–126 mEq/l Cholesterol 59–179 mg/dl Creatinine 0.8–1.6 mg/dl Creatine kinase 112–3023 U/l

Gamma glutamyltransferase 0.1-1.5 U/I

Globulin 2.5–4.2 g/dl Glucose 4.8–9.0 mmol/l

Lactate dehydrogenase 60–300 U/l Magnesium 2.4–3.0 mg/dl Phosphorus 4.5–8.5 mg/dl Potassium 3.93–6.59 mEq/l

Sodium 152-163 mEq/l

Sorbitol dehydrogenase 3.0–11.0 U/l Total bilirubin 0.03–0.13 mg/dl Total bile acids 0.3–13.8 Umol/l Total protein 6.1–7.8 g/dl Triglyceride 20–148 mg/dl Urea nitrogen 17.1–31.6 mg/dl

Blood gases (venous): pH 7.11-7.39 PCO<sub>2</sub>3.7-8.1 kPa

PCV = packed cell volume (haematorcirit), MCH = mean corpuscular haemoglobin, MCHC = mean corpuscular harmoglobin concentration, MCV = mean corpuscular volume

## Data collected from 16 cats during phase 1 and phase 2 study

Comparison between groups and with time — yes = P < 0.05, no = no significant difference

рН							
PC							
Day	-5	4	5	18/28	19/29	25/35	
Mean	7.31	7.35	7.27	7.30	7.31	7.28	
SD	0.08	0.04	0.13	0.05	0.06	0.13	
PB							
day	<b>-</b> 5	4	5	18/28	19/29	25/35	
mean	7.33	7.34	7.24	7.31	7.31	7.23	
SD	0.06	0.04	0.06	0.04	0.05	0.17	

Haematocr	it %						
PC							
Day	<b>-</b> 5	4	5	18/28	19/29	25/35	
Mean	31.5	27.3	29.2	26.5	29.2	32.5	
SD	3.7	2.4	3.1	4.7	3.4	3.3	
PB							
Day	<b>-</b> 5	4	5	18/28	19/29	25/35	
Mean	32.6	27.4	30.6	26.8	29.2	31.8	
SD	1.9	2.7	1.9	4.5	2.4	4.0	

Between groups: no With time: yes

Activated p	artial thrombopla	astin (s)					
PC							
Day	-5	4	5	18/29	19/29	25/35	
Mean	12.5	12.3	12.2	13.3	12.9	11.6	
SD	1.3	0.7	1.2	1.3	1.5	0.9	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	13.1	12.6	12.2	12.4	12.5	12.2	
SD	0.6	0.6	1.1	1.2	0.7	0.9	

Between groups: no With time: yes

Platelets (1	0 <sup>3</sup> /μl)						
<i>PC</i> Day	<b>–</b> 5	4	5	18/28	19/29	25/35	
Mean SD	444 157	476 112	520 102	407 182	386 122	456 184	
PB Day	<b>–</b> 5	4	5	18/28	19/29	25/35	
Mean SD	491 114	423 206	592 90	567 62	445 241	591 132	

Between groups: no With time: yes (? Bio)

Prothrombi	n time (s)						
PC							
Day	<b>-</b> 5	4	5	18/28	19/29	25/35	
Mean	9.5	8.6	8.3	8.5	7.7	8.6	
SD	0.5	0.2	0.2	0.4	0.3	0.7	
РВ							
Day	<b>-</b> 5	4	5	18/28	19/29	25/35	
Mean	9.6	8.3	8.2	8.2	7.6	8.5	
SD	0.6	0.2	0.2	0.2	0.3	0.3	

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Basophils (	10 <sup>3</sup> /μl)						
PC Day Mean	-5 0.02	4 0.01	5 0.12	18/28 0.03	19/29 0.07	25/35 0.07	
SD PB Day Mean SD	1.67 -5 0.05 0.06	1.33 4 0.05 0.08	1.63 5 0.02 0.02	0.06 18/28 0.04 0.07	0.07 19/29 0.07 0.08	0.10 25/35 0.13 0.22	

Between groups: no With time: no

Eosinophils	$(10^3/\mu l)$						
PC							
Day	-5	4	5	18/29	19/29	25/35	
Mean	0.67	0.92	1.01	0.81	1.15	0.84	
SD	0.21	0.45	0.34	0.35	0.42	0.39	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	0.77	0.94	1.23	0.65	0.95	0.91	
SD	0.40	0.37	0.66	0.23	0.17	0.65	

Between groups: no With time: yes

Fibrinogen	(mg/dl)						
PC							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	151	186	237	163	220	158	
SD	15	20	38	33	30	34	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	147	199	260	178	230	147	
SD	17	15	35	18	22	16	

Between groups: no With time: yes

Lymphocyte	es (10³/μl)						
PC Day Mean SD	–5 6.96 3.70	4 6.33 3.00	5 7.81 3.12	18/29 5.68 4.02	19/29 6.24 3.48	25/35 8.36 4.16	
PB Day Mean SD	–5 6.90 2.51	4 6.48 3.01	5 9.30 4.68	18/29 5.72 3.02	19/29 6.26 3.26	25/35 10.82 4.93	

Total leukoo	cytes (10³/μl)						
PC							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	15.55	14.20	17.15	13.49	15.87	16.45	
SD	8.51	5.56	6.43	5.77	4.93	6.54	
PB							
Day	-5	4	5	18/29	19/29	25/35	
Mean	14.68	16.09	19.71	13.07	15.27	18.59	
SD	3.80	5.10	8.64	4.56	5.14	7.55	

Between groups: no With time: yes

Alanine am	inotransferase (	U/I)					
PC Day	-5	4	5	18/29	19/29	25/35	
Day Mean	=5 81	68	66	65	70	90	
SD	37	26	26	16	17	18	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	80	66	61	65	75	88	
SD	11	6	8	11	12	15	

Between groups: no With time: yes

Albumin (g	/dl)						
PC							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	3.7	3.3	3.3	3.0	3.4	3.4	
SD	0.2	0.1	0.2	0.2	0.2	0.3	
РВ							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	3.6	3.4	3.4	3.0	3.4	3.5	
SD	0.1	0.2	0.1	0.2	0.2	0.2	

Between groups: no With time: yes

Alkaline ph	osphatase (U/I)						
PC							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	100	84	84	90	79	105	
SD	14	11	21	17	9	15	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	89	77	72	73	71	104	
SD	21	18	19	18	20	25	

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Aspartate t	ransaminae (U/I)	)					
PC							
Day	-5	4	5	18/29	19/29	25/35	
Mean	36	31	19	34	19	40	
SD	17	23	7	33	3	16	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	31	19	17	25	20	35	
SD	11	2	4	6	6	9	

Between groups: yes With time: yes

Calcium (m	ng/dl)						
PC							
Day	<b>–</b> 5	4	5	18/29	19/29	25/35	
Mean	10.9	10.7	10.8	10.6	10.9	11.2	
SD	0.3	0.3	0.3	0.4	0.4	0.7	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	10.6	10.8	10.7	10.3	11.0	11.1	
SD	0.3	0.3	0.4	0.4	0.5	0.7	

Between groups: no With time: yes

Creatine ki	nase (U/I)						
PC							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	1816	1483	475	722	447	1329	
SD	2562	1855	573	969	280	1385	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	840	284	337	914	332	1008	
SD	525	178	236	1017	421	723	

Between groups: no With time: no

Blood urea	nitrogen (mg/dl)	)					
PC							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	26.0	22.4	27.4	22.1	24.0	26.6	
SD	3.0	3.1	3.4	2.7	4.0	3.1	
PB							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	25.7	23.0	27.1	23.6	27.2	26.8	
SD	3.6	1.7	3.6	1.4	4.8	2.3	

Creatinine (mg/dl)								
PC Day Mean SD	-5 0.8 0.2	4 0.8 0.2	5 0.8 0.1	18/29 0.7 0.1	19/29 0.8 0.2	25/35 0.9 0.2		
PB Day Mean SD	-5 0.8 0.2	4 0.8 0.1	5 0.8 0.2	18/29 0.7 0.2	19/29 0.8 0.2	25/35 0.9 0.2		

Between groups: no With time: yes

Amylase (U	I/I)						
PC							
Day	<b>-</b> 5	4	5	18/29	19/29	25/35	
Mean	831	851	887	759	851	891	
SD	140	104	119	81	111	127	
PB							
Day	<b>–</b> 5	4	5	18/29	19/29	25/35	
Mean	880	879	941	843	913	855	
SD	136	96	102	71	141	107	