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Pectus Excavatum: Computed Tomography and Medium Term Surgical Outcome in a Prospective Cohort of 10 Kittens

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Keywords:	feline, pectus excavatum, soft tissue surgery, computed tomography, thoracic
	Objectives To report the use of computed tomography (CT) in conjunction with clinical signs to assess severity of pectus excavatum (PE) in kittens and to guide surgical decision making. To report medium term outcome in a prospective cohort of kittens undergoing surgical correction. Methods Prospective study of ten, 10-15 week old kittens diagnosed with moderate/severe pectus excavatum
Abstract:	Results CT provides additional information useful for selecting patients for surgical correction and for planning that surgery. Traditional radiographic indices (vertebral, frontosagittal) provide reasonable approximations of the CT determined dimensions but these seem to correlate poorly with the severity of clinical signs. Kittens commonly have lateralised deformities which are associated with less severe clinical symptoms, whilst those with midline deformities are associated with more severe clinical signs. 6/7 kittens with severe PE which had a ventral splint applied for 4 weeks had excellent medium term outcomes.
	Clinical Significance Restriction of diastolic filling by midline sternal deviation may be an important cause of exercise intolerance in cats with pectus excavatum. CT can be used to assess affected kittens and to plan surgery when indicated.

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- **1** Pectus Excavatum: Computed Tomography and Medium Term Surgical Outcome
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- 17 Abstract
- 18 Objectives

To report the use of computed tomography (CT) in conjunction with clinical signs to assess severity of pectus excavatum (PE) in kittens and to guide surgical decision making. To report medium term outcome in a prospective cohort of kittens undergoing surgical correction.

23

24 Methods

25	Prospective study of ten, 10-15 week old kittens diagnosed with moderate/severe					
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44	Keywords: Feline, Pectus Excavatum, Computed Tomography, Soft Tissue, Thoracic					
45						

46 Introduction

47

48 Pectus Excavatum (PE) is an uncommon, congenital thoracic wall deformity that has 49 been previously documented in a variety of species including man, dogs and cats.¹⁻⁴ PE 50 is characterised by a palpable dorsal deviation of the caudal sternebrae resulting in a 51 loss of thoracic volume and potential respiratory compromise.

52

53 In man, PE is the most commonly observed thoracic wall abnormality occurring 54 between 1:400 and 1:1000 live births and is commonly associated with connective 55 tissue disorders such as Marfan and Ehlers Danlos syndromes.^{6,7} Although familial 56 tendencies have been demonstrated, it may well be a phenotypic response to a variety 57 of underlying conditions and its aetiology is incompletely understood.⁷ The incidence of 58 PE in kittens is unknown although the defect seems to be more commonly seen in 59 Bengal cats than domestic short hair (DSH) cats which is suggestive of there being a 60 familial component to its expression.⁸ The presence of PE is also positively correlated 61 with flat-chested kitten syndrome in Burmese cats.⁹

62

In cats, the severity of the deformity is traditionally graded using the vertebral (VI) and
frontosagittal (FSI) indices as measured from orthogonal view thoracic radiographs
(table 1).¹ In man, however, computed tomography (CT) is commonly employed to
assess both the severity of the deformity and to assist with preoperative surgical
planning.^{5,10}

68

69 The authors had noted an apparent discrepancy between the severity of clinical70 symptoms and radiographically determined vertebral and frontosagittal indices. This

study describes the use of CT in assessing severity of PE in cats with the hypotheses that standard radiography accurately approximates CT determined indices but that CT will provide additional information to explain the discrepancy between the radiographic and clinical severity of PE. The authors also provide short and medium term follow up for a cohort of 10 kittens who underwent CT +/- surgical correction.

76

77 Materials and Methods:

78

79 Kittens seen by the primary author in the period 2012-2014 that were between ten and 80 fifteen weeks of age diagnosed with moderate/severe PE (using published VI/FSI 81 ranges) were eligible for inclusion in this study. Full patient data (age breed, weight, 82 history, results of clinical examination) were recorded and cases were allocated a 83 Clinical Severity Score (CSS) (table 2). Kittens who had not had thoracic radiographs 84 taken by the referring veterinarian were radiographed by the primary author during 85 the initial patient assessments. All radiographs were reviewed and VI/FSI calculated by 86 the primary author.

87

88 After a full clinical examination, kittens were premedicated with a standard protocol of 89 0.01mg/kg acepromazine (ACP, Novartis) and 0.02mg/kg buprenorphine (Vetergesic, 90 Alstoe) and preoxygenated in an oxygen cage before induction with propofol (PropoFlo, 91 Abbott Animal Health). Anaesthesia was maintained with isoflurane (IsoFlo, Abbott 92 Animal Health) and supportive intravenous fluid therapy (Hartmann's solution) was 93 administered at 10ml/kg/hr. CT images were acquired using a 2 slice GE Lightspeed 94 scanner and standard helical thoracic protocol with 2.5mm slice thickness with 1.25mm 95 slice interval with the kittens placed in dorsal recumbency. Post contrast series were

acquired using 2ml/kg ioversol 64% (Optiray 300, Covidien) given as an intravenous
bolus immediately prior to scanning. The kitten's lungs were not hyperinflated prior to
scanning and no attempt was made to induce a respiratory pause.

99

All CT scans were assessed at a later date by a board certified diagnostic imager who was blinded to the clinical history/previous radiographs of each kitten. Images were evaluated using dedicated DICOM viewer software (Osirix, Geneva, Switzerland, version 5.8.5 – 64bit) on a computer workstation (Apple Mac Pro, Apple, USA) with a calibrated LCD flat screen monitor (Apple Cinemax Display, 30 inch, Apple, USA). During the course of image evaluation, multiplanar reconstructions and variable windowing were used according to the preference of the diagnostic imager.

107

108 All CT studies were assessed for diagnostic quality and for concurrent musculoskeletal 109 abnormalities. CT determined VI and FSI were recorded for each kitten and compared 110 to the radiographically determined VI/FSI using either recent (within 48 hours of CT) 111 radiographs or CT "scout" images (planning radiographs) if judged to be of sufficient 112 quality. Anticipated low case numbers precluded meaningful statistical analysis so 113 scatter plots were used to illustrate any relationships between the measured 114 radiographic and CT VI and FSI values. Correlation coefficients were calculated with an 115 "r value" >0.5 taken as positively correlated and <0.5 as a negative correlation. Bias was 116 assessed using Bland and Altman plots.

117

Additional CT analysis including measurement of lung volume and assessment of the nature of the sternal deformity. Lung volume was calculated by drawing an ROI around the surface of each lung and then using the ROI volume calculator tool of the imaging software. Results (cm³/kg) were compared against a control population of adult cats who had undergone thoracic CT for non-respiratory disease. The nature of the dorsal deviation of the caudal sternebrae (midline or lateralised) was also recorded as was the location/number of the deviated sternebrae and their proximity to the visible overlying major cardiovascular structures. The presence/absence of lateralisation of the defect was compared to the CSS.

127

Two kittens had an additional dynamic CT performed to assess the degree of movement of the sternum during the respiratory cycle. A cine protocol was used to take sequential transverse sections at the estimated point of maximum sternal deformity with images acquired every 0.5 seconds for 20 seconds. The distance between the sternum and vertebral body was measured at the points of greatest inspiration and expiration for the two dynamic CT studies and the percentage change calculated.

134

135

136 Surgical Methods:

137 Cases that had a severe (<6) VI and clinical signs attributed to PE (CSS 1-5) were 138 considered surgical candidates. The surgical technique was based on that published 139 elsewhere.^{11,12} In brief, the kittens were placed in dorsal recumbency and a ventral 140 sternal cast (Dynacast Prelude, BSN Medical) was made conforming to the anticipated 141 postoperative position of the sternum which was facilitated by applying moderate 142 laterolateral compression to the thorax during the casting process. 3.5M polypropylene 143 (Prolene, Ethicon) circumsternal sutures were then placed as single interrupted sutures 144 starting cranially and progressing caudally. The ends of the sutures were left long and 145 passed through the cast before tying under tension whilst simultaneously applying

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moderate laterolateral thoracic compression. Cases that had no detectable safe corridor
for suture passage underwent a minimal dissection to the caudal sternebrae which were
then directly retracted ventrally allowing the circumsternal sutures to be passed. This
wound was closed using subcutaneous 1.5M or 2M glycomer 631 (Biosyn, Covidien)
before routine cast placement. All casts were covered by chest bandages and
postoperative thoracic radiographs were taken.

152

Kittens who underwent surgery were discharged the next day on 5 day courses of 0.05mg/kg meloxicam sid (Metacam, Boehringer Ingelheim) and 20mg/kg potentiated amoxicillin bid (Synulox Palatable Drops, Zoetis). The casts were maintained for 4 weeks at which point they were removed under anaesthesia using an identical anaesthetic protocol as above. Thoracic radiographs were taken to assess the degree of PE correction and postoperative VI's were calculated by the primary author.

159

Medium term follow up was obtained by email/telephone contact with the owners/referring veterinarians and the kittens were allocated new clinical severity scores. Pre and postoperative CSS were then compared.

163

164 **Results**

165

Ten kittens met the inclusion criteria during the study period. Breed distribution was: Bengal (5), Domestic Short Hair (4) and Maine Coon (1). There were four female and six male kittens. Median weight (kg) at time of surgery was 1.17 (range 0.65-2.0). Clinical signs reported included: palpable abnormality (10), tachypnoea (7), exercise intolerance (4), at least one previous episode of antimicrobial responsive dyspnoea (3), stunting/poor growth (3), dehydration (3), chronic dyspnoea (2), constipation (2) and
PU/PD (1). Allocated CSS were: 1(4 kittens), 2(3 kittens), 4(2 kittens) and 5(1 kitten).

174 All CT scans were deemed to be of diagnostic quality. Two kittens (G, H) did not have 175 radiographs or CT "scout" images of sufficient quality for VI and FSI to be accurately 176 measured and so these cases were omitted from the analysis. Radiographic and CT 177 measurements/indices are given in tables 3 and 4. CT and radiographically determined 178 indices (VI, FSI) were compared and the results shown in figures 1 and 2. CT 179 consistently gave a lower value for the VI with a mean difference of 0.53. FSI calculated 180 from CT images tended to be higher than the value calculated from radiographic 181 measurements

182

The distance between the dorsal most point of the deviated sternum and the overlying vertebra ("c") was determined at maximum inspiration and expiration for each of 3 respiratory cycles for two kittens for which a cine scan was performed. "c" changed by an average of 3.16% in the first kitten and 0.68% in the second.

187

188 Lung volume/kg body weight did not appear to be significantly different from that of 189 the control population (mean PE: $37.1 \text{ cm}^3/\text{kg}$, mean control: $45.3 \text{ cm}^3/\text{kg}$) although 190 low case numbers precluded statistical analysis. CT review showed that the dorsal 191 sternal deviation was lateralised in 5/10 kittens and this was dextral in 4/5 cases. The 192 kitten with sinistral deviation was diagnosed with complete situs inversus. No other 193 musculoskeletal deformities were detected. Kittens which had a lateralised sternal 194 deformity tended to have a lower CSS than those that did not (1,1,1,1,2, versus 195 2,2,4,4,5).

The sternebrae closest to the overlying major cardiovascular structures were 5-7 (4 kittens), 5-8 (2 kittens), 6-8 (2 kittens), 6-7 (2 kittens), 4-6 (1 kittens) (see figure 3). The dorsal aspect of the sternal deformity was judged to be in contact with a major cardiovascular structure in 9/10 cases. The tenth case was judged to have a safe corridor for needle passage of 2mm.

202

203 Short term/medium term follow up:

Cases A, B, C, F, G, H and I underwent surgery. Case D was PU/PD, anorexic and significantly dehydrated at the time of presentation and was euthanased at the owners request. A post-mortem CT examination was performed in this case with the owner's consent. Cases E and J were judged to have PE of medium severity based on VI. Case E had a CSS of 1 and surgery was not recommended. Case J presented with a CSS of 4 and surgery was recommended but was declined by the owner.

210 There were no intraoperative complications in any of the cases that underwent surgery.

Case A developed furosemide responsive dyspnoea 1 week following surgery that required hospitalisation and oxygen supplementation. Dyspnoea partially resolved but required continued medication. Case A developed cardiorespiratory arrest under anaesthesia for cast removal and did not respond to resuscitation attempts. VI was improved (higher) in all cases that underwent surgical correction (table 5).

216 No moist dermatitis or pyoderma secondary to cast placement occurred in any of the

- 217 **cases**. Owners do not report any clinical symptoms of PE in all surviving kittens.
- Follow up data and CSS are listed in table 5.
- 219

220 **Discussion**:

Ten kittens were recruited into this study, five of which were Bengals. Although
adequate population data is not available for statistical analysis, this tends to support
Bengal kittens being at increased risk of PE⁸ although bias in the referred population is
possible.

226

227 The clinical signs reported by the owners and/or breeders were variable. Although 228 some interobserver variation is probable, all clinical histories were taken by the same 229 veterinary surgeon and the severity of historical signs reported seemed to correlate 230 with clinical signs at presentation. Tachypnoea was the most common clinical sign 231 reported after "palpable deformity" which is a diagnostic feature of PE. Tachypnoea 232 was presumed to be due to decreased lung volume and impaired alveolar exchange and 233 forms part of a continuum of signs progressing to exercise intolerance and respiratory 234 distress presumably with increased severity of deformity. This spectrum of signs 235 formed the basis for the CSS which was used in this study. Surprisingly, however, the 236 lung volume (as a proportion of body weight) of affected kittens was not significantly 237 different from a control population of unaffected cats. Larger case numbers and ideally 238 an age-matched control population would allow statistical analysis and verification of 239 these preliminary findings. Results could be further confounded by a compensatory 240 increased depth of respiration seen in affected kittens. More information could be 241 obtained by lung plethysmography but this is not widely available for veterinary 242 patients particularly those with small tidal volumes.

243

Three of the ten kittens had at least one previous episode of antimicrobial –responsivedyspnoea. In each case, dyspnoea responded rapidly to the administration of

potentiated amoxicillin. The exact site and cause of the presumed infection is not known
but this could be caused by ventilation impairment, failure to clear alveolar secretions
or other functional abnormalities.

249

Although only assessed in two of the 10 cases, the sternum did not move significantly during the respiratory cycle with "c" values varying by only 3.16% and 0.68%. This suggests that radiographically determined VI should be relatively constant independent of the respiratory phase at which they were taken.

254

255 There was reasonable correlation between radiographically determined and CT-256 determined VI and FSI, CT consistently gave a lower value for the VI with a mean 257 difference of 0.53 and FSI calculated from CT images tended to be higher than the value 258 calculated from radiographic measurements. The mean difference in FSI was 0.83 but 259 there was a significant association between the variation and the value of the FSI. These 260 findings suggest that separate reference intervals for normality and severity of VI and 261 FSI are appropriate depending on the imaging modality used but that an appropriate 262 reference interval for CT-calculated FSI can be inferred from the published radiographic 263 FSI values.

264

No significant concurrent musculoskeletal deformities were identified in any of the affected kittens. In each case the sternum started to deform in the caudal half and the deformity was dorsal but also lateral in 5/10 cases with 4/5 being deviated to the right hemithorax. The case with sinistral deviation of the xiphoid was the cat with situs inversus. The cause of this lateralisation away from the heart is uncertain but possible

- explanations would include displacement of the caudally deviated sternebrae by theheart and/or traction from the diaphragmatic crura during development.
- 272

273 Cardiac perforation is a recognised complication of placing a ventral corrective splint.¹² 274 The deformed sternum was judged to be in contact with either the heart or the caudal 275 vena cava in 9/10 cases with the 10th case having a safe corridor for suture placement of 276 only 2mm. Having demonstrated how close the dorsal sternebrae were to the heart in 277 all our surgical cases, we modified our surgical technique accordingly. No intraoperative 278 complications were encountered and no postoperative complications relating to this 279 approach were seen.

280

281 VI and FSI are useful for initial screening of cases into an anatomic severity category -282 mild, moderate or severe but they do not appear to be useful for determining whether 283 cases with moderate to severe anatomic defects are likely to have severe clinical signs. 284 This would imply that there may be other factors beyond simple musculoskeletal 285 deformity which are contributing to the severity of the clinical signs observed. Cats with 286 lateral deviation of the xiphoid seemed to be associated with a lower clinical score than 287 those kittens with a midline deviation. It has been suggested that clinical symptoms of 288 PE in people may be partially due to a direct compressive or restrictive effects of the 289 displaced sternebrae on the heart itself.¹³ Athletic performance is compromised by the 290 inability of the heart to increase diastolic volume to meet increased oxygen demands 291 and direct compression of the right side of the heart is considered to be an indication 292 for surgery in people. 5,13 It is possible that clinical signs seemed to be more severe in 293 cats with a midline pectus deformity due to a similar mechanism whereas the kittens 294 with a lateralised defect had more space available for the heart to increase diastolic volume as required. Cardiovascular compromise may therefore be a more significant
driver of clinical signs associated with PE in kittens than altered pulmonary function.
This could be why case J, that had a midline sternal deviation, had severe clinical signs
despite a "moderate" VI (7) and it may be that a different threshold (higher VI) should
be used when deciding if kittens with midline defects should undergo surgery.

300

301 One case (A) developed significant postoperative furosemide-responsive dyspnoea and 302 then died at the time of cast removal. The apparent initial response to diuretics is 303 suggestive of pulmonary oedema that could be caused either by pulmonary re-304 expansion, concurrent cardiovascular disease or pulmonary hypertension.¹⁴ Although 305 no significant concurrent cardiovascular disease was detected on the initial CT scan, this 306 modality is not as sensitive as echocardiography when assessing cardiac function. The 307 cause of death at the time of cast removal remains unknown as no post-mortem analysis 308 was permitted.

309

310 All cases that survived showed full resolution of clinical symptoms with no exercise 311 intolerance or episodes of dyspnoea reported at the time of follow up. Whilst we used a 312 combination of VI and CSS to determine which cases should benefit from surgery, there 313 was no control population for which treatment was intentionally withheld in order to 314 demonstrate a difference in postoperative outcome as this would have been unethical. 315 We suspect that many cases of severe PE are euthanased due to perceived poor 316 prognosis and financial concerns about treatment costs. This, and the rarity of the 317 condition resulted in only low numbers of cases being recruited despite internet based 318 advertising for case enrolment. We are therefore limited to making broad 319 recommendations about patient selection and treatment efficiency. In our study,

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patients were selected for surgery based on VI and clinical signs. All cases had a sternal cast maintained for 4 weeks and all cases that survived are currently asymptomatic at a mean follow up of 15 months. One of the risks of uncorrected PE is the development of pulmonary hypertension and right side heart failure which the authors have observed in multiple cases < 12mths old. It is possible that some of our cases could develop respiratory symptoms at a later stage and we intend to publish longer term follow up (5 year) data when available.

327

328 In summary, conventional radiography yields reasonable approximations of CT-329 determined VI and FSI. CT was useful in determining the presence/absence of safe 330 corridors for circumsternal suture placement leading to a minor modification of the 331 surgical approach employed for ventral cast placement. CT also allowed detection of 332 cats with midline sternal deviation which may be at risk of developing more severe 333 clinical signs due to diastolic restriction despite relatively mild skeletal deformity. Ten 334 to 15 week old kittens with severe deformity as judged by VI and with compatible 335 clinical signs can be treated by placement of a sternal splint for a 4 week period which 336 can be associated with an excellent medium term outcome.

337

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341

- 342 **Conflict of Interest**
- 343 The authors declare that there is no conflict of interest.

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346

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Table 1: Vertebral (VI) and Frontosagittal (FIS) Indices for Assessment of Pectus Excavatum (P.E.)

	FSI	VI
Normal	0.7-1.3 (1.00)	12.6-18.8 (15.0)
Mild P.E.	2.0	>9.0
Moderate P.E.	2.0-3.0	6.0-9.0
Severe P.E.	>3.0	<6.0

Definitions:

FSI: Ratio of the thoracic width at T10 as measured on a dorsoventral radiograph and the distance from the centre of the ventral surface of T10 or vertebra overlying the deformity and the nearest point on the sternum

VI: Ratio of the distance from the centre of the dorsal surface of the vertebral body overlying the deformity to the near point of the sternum and the dorsoventral diameter of the centrum of the same vertebra

Table 2: Clinical Severity Score

No clinical symptoms recorded	0
Elevated respiratory rate (>30 breaths/min)	1
Elevated respiratory rate, exercise intolerance noted	2
Elevated respiratory rate, intermittent (<50% time) dyspnoea	3
Prolonged periods (>50%) of dyspnoea	4
Prolonged periods of dyspnoea with evidence of significant extra-thoracic	5
disease	

Patient ID	Weight (kg)	VI	FSI	Resting Resp (breaths/min)	Age (weeks)	Clinical score
А	0.85	4	4.5	40	15	2
В	0.9	2.2	9.7	40	12	2
С	2	4.3	4.8	40	13	4
D	0.65	4	4.3	60	10	5
Е	1.23	6.3	3.3	40	12	1
F	1.62	5.4	2.4	40	13	1
G	1.4	N/A	N/A	30	13	1
Н	1	N/A	N/A	48	12	2
Ι	1.3	5.8	3.3	36	12	1
J	1.1	7	2.5	30	12	4

Table 3: Clinical data and Radiographically Determined Indices at Presentation

Patient ID	VI	FSI	Lung volume cm³/Kg	Clinical score
А	3.7	5.1	45.71	2
В	1.6	19.8	62.96	2
С	2.9	5.5	43.16	4
D	3.3	5.6	26.02	5
Е	3.4	3.9	28.40	1
F	4.8	2.6	26.87	1
G	4.3	4	27.95	1
Н	3.1	5.9	30.35	2
Ι	4.6	3.2	33.72	1
J	6.2	2.5	46.01	4

Table 4: Calculated indices from CT measurements

O P -P

		Initial	Postop	Defect			
Case ID	Surgery?	VI	VI	Lateralised?	Initial CSS	Folow up CSS	FUP (mths)
А	Y - died	4	N/A	Ν	2	N/A	N/A
В	Y	2.2	6.3	Ν	2	0	18
С	Y	4.3	8.3	Ν	4	0	10
D	N - Euth	4	N/A	Ν	5	N/A	N/A
E	Ν	6.3	N/A	Y-R	1	0	24
F	Y	5.4	6.4	Y-R	1	0	24
G	Y	4.3	7.7	Y-R	1	0	18
Н	Y	3.2	5.5	Y-R	2	0	18
1	Y	5.8	7.9	Y-L	1	0	4
J	Ν	7	N/A	Ν	4	0	4

Table 5: Follow up data. Euth = euthanased; VI = vertebral index; N = No; Y-R = Yes to right; Y-L = Yes to left; CSS = clinical severity score; FUP (mths) = Follow up period (months). N/A = not applicable

NB CT data used for initial VI for cases G, H as radiographic data not available.

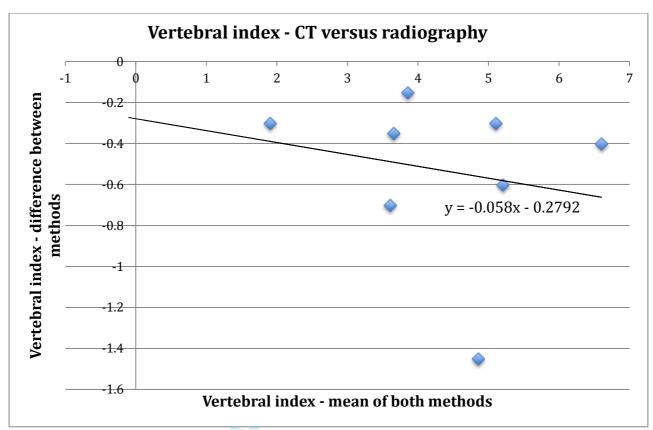


Figure 1 – Bland and Altman plot of the data obtained from 8 paired measurements of the vertebral index using the computed tomography or radiographic image. Correlation R = 0.20 (P = 0.64); slope = -0.058 (P = 0.64); intercept = -0.28 (P = 0.62)

P.C.

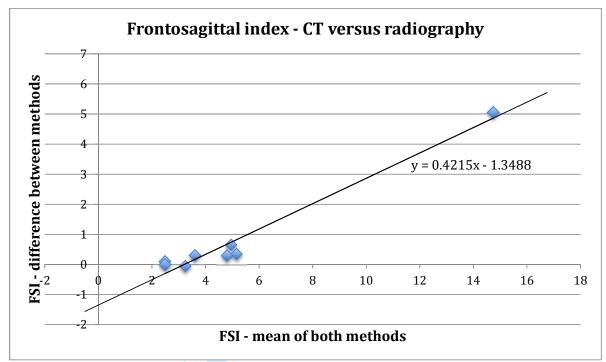


Figure 2 – Bland and Altman plot of the data obtained from 8 paired measurements of the frontosagittal index using the computed tomography or radiographic image. Correlation R = 0.983 (P = <0.01); slope = -0.422 (P = <0.01); intercept = -1.35 (P = <0.01)

Figure 3– Sternebrae closest to overlying cardiovascular structures in each of the ten kittens

S4	S5	S6	S7	S8

1	Pectus Excavatum: Computed Tomography and Medium Term Surgical Outcome
2	in a Prospective Cohort of 10 Kittens
3	
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15	
16	
17	Abstract
18	Objectives
19	To report the use of computed tomography (CT) in conjunction with clinical signs to
20	assess severity of pectus excavatum (PE) in kittens and to guide surgical decision
21	making. To report medium term outcome in a prospective cohort of kittens undergoing
22	surgical correction.
23	

24 Methods

25 Prospective study of ten, 10-15 week old kittens diagnosed with moderate/severe26 pectus excavatum

27

29	CT provides additional information useful for selecting patients for surgical correction
30	and for planning that surgery. Traditional radiographic indices (vertebral,
31	frontosagittal) provide reasonable approximations of the CT determined dimensions
32	but these seem to correlate poorly with the severity of clinical signs. Kittens commonly
33	have lateralised deformities which are associated with less severe clinical symptoms,
34	whilst those with midline deformities are associated with more severe clinical signs.
35	6/7 kittens with severe PE which had a ventral splint applied for 4 weeks had excellent
36	medium term outcomes.

37

38	Clinical S	ignificance

39

40 Restriction of diastolic filling by midline sternal deviation may be an important cause of

41 exercise intolerance in cats with pectus excavatum. CT can be used to assess affected

42 kittens and to plan surgery when indicated.

43

44 Keywords: Feline, Pectus Excavatum, Computed Tomography, Soft Tissue, Thoracic

46 Introduction

47

48 Pectus Excavatum (PE) is an uncommon, congenital thoracic wall deformity that has 49 been previously documented in a variety of species including man, dogs and cats.¹⁻⁴ PE 50 is characterised by a palpable dorsal deviation of the caudal sternebrae resulting in a 51 loss of thoracic volume and potential respiratory compromise.

52

53 In man, PE is the most commonly observed thoracic wall abnormality occurring 54 between 1:400 and 1:1000 live births and is commonly associated with connective 55 tissue disorders such as Marfan and Ehlers Danlos syndromes.^{6,7} Although familial 56 tendencies have been demonstrated, it may well be a phenotypic response to a variety 57 of underlying conditions and its aetiology is incompletely understood.⁷ The incidence of 58 PE in kittens is unknown although the defect seems to be more commonly seen in 59 Bengal cats than domestic short hair (DSH) cats which is suggestive of there being a 60 familial component to its expression.⁸ The presence of PE is also positively correlated 61 with flat-chested kitten syndrome in Burmese cats.⁹

62

In cats, the severity of the deformity is traditionally graded using the vertebral (VI) and
frontosagittal (FSI) indices as measured from orthogonal view thoracic radiographs
(table 1).¹ In man, however, computed tomography (CT) is commonly employed to
assess both the severity of the deformity and to assist with preoperative surgical
planning.^{5,10}

68

69 The authors had noted an apparent discrepancy between the severity of clinical70 symptoms and radiographically determined vertebral and frontosagittal indices. This

study describes the use of CT in assessing severity of PE in cats with the hypotheses that standard radiography accurately approximates CT determined indices but that CT will provide additional information to explain the discrepancy between the radiographic and clinical severity of PE. The authors also provide short and medium term follow up for a cohort of 10 kittens who underwent CT +/- surgical correction.

76

77 Materials and Methods:

78

79 Kittens seen by the primary author in the period 2012-2014 that were between ten and 80 fifteen weeks of age diagnosed with moderate/severe PE (using published VI/FSI 81 ranges) were eligible for inclusion in this study. Full patient data (age breed, weight, 82 history, results of clinical examination) were recorded and cases were allocated a 83 Clinical Severity Score (CSS) (table 2). Kittens who had not had thoracic radiographs 84 taken by the referring veterinarian were radiographed by the primary author during 85 the initial patient assessments. All radiographs were reviewed and VI/FSI calculated by 86 the primary author.

87

88 After a full clinical examination, kittens were premedicated with a standard protocol of 89 0.01mg/kg acepromazine (ACP, Novartis) and 0.02mg/kg buprenorphine (Vetergesic, 90 Alstoe) and preoxygenated in an oxygen cage before induction with propofol (PropoFlo, 91 Abbott Animal Health). Anaesthesia was maintained with isoflurane (IsoFlo, Abbott 92 Animal Health) and supportive intravenous fluid therapy (Hartmann's solution) was 93 administered at 10ml/kg/hr. CT images were acquired using a 2 slice GE Lightspeed 94 scanner and standard helical thoracic protocol with 2.5mm slice thickness with 1.25mm 95 slice interval with the kittens placed in dorsal recumbency. Post contrast series were

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acquired using 2ml/kg ioversol 64% (Optiray 300, Covidien) given as an intravenous
bolus immediately prior to scanning. The kitten's lungs were not hyperinflated prior to
scanning and no attempt was made to induce a respiratory pause.

99

All CT scans were assessed at a later date by a board certified diagnostic imager who was blinded to the clinical history/previous radiographs of each kitten. Images were evaluated using dedicated DICOM viewer software (Osirix, Geneva, Switzerland, version 5.8.5 – 64bit) on a computer workstation (Apple Mac Pro, Apple, USA) with a calibrated LCD flat screen monitor (Apple Cinemax Display, 30 inch, Apple, USA). During the course of image evaluation, multiplanar reconstructions and variable windowing were used according to the preference of the diagnostic imager.

107

108 All CT studies were assessed for diagnostic quality and for concurrent musculoskeletal 109 abnormalities. CT determined VI and FSI were recorded for each kitten and compared 110 to the radiographically determined VI/FSI using either recent (within 48 hours of CT) 111 radiographs or CT "scout" images (planning radiographs) if judged to be of sufficient 112 quality. Anticipated low case numbers precluded meaningful statistical analysis so 113 scatter plots were used to illustrate any relationships between the measured 114 radiographic and CT VI and FSI values. Correlation coefficients were calculated with an 115 "r value" >0.5 taken as positively correlated and <0.5 as a negative correlation. Bias was 116 assessed using Bland and Altman plots.

117

Additional CT analysis including measurement of lung volume and assessment of the nature of the sternal deformity. Lung volume was calculated by drawing an ROI around the surface of each lung and then using the ROI volume calculator tool of the imaging software. Results (cm³/kg) were compared against a control population of adult cats who had undergone thoracic CT for non-respiratory disease. The nature of the dorsal deviation of the caudal sternebrae (midline or lateralised) was also recorded as was the location/number of the deviated sternebrae and their proximity to the visible overlying major cardiovascular structures. The presence/absence of lateralisation of the defect was compared to the CSS.

127

Two kittens had an additional dynamic CT performed to assess the degree of movement of the sternum during the respiratory cycle. A cine protocol was used to take sequential transverse sections at the estimated point of maximum sternal deformity with images acquired every 0.5 seconds for 20 seconds. The distance between the sternum and vertebral body was measured at the points of greatest inspiration and expiration for the two dynamic CT studies and the percentage change calculated.

134

135

136 Surgical Methods:

137 Cases that had a severe (<6) VI and clinical signs attributed to PE (CSS 1-5) were 138 considered surgical candidates. The surgical technique was based on that published 139 elsewhere.^{11,12} In brief, the kittens were placed in dorsal recumbency and a ventral 140 sternal cast (Dynacast Prelude, BSN Medical) was made conforming to the anticipated 141 postoperative position of the sternum which was facilitated by applying moderate 142 laterolateral compression to the thorax during the casting process. 3.5M polypropylene 143 (Prolene, Ethicon) circumsternal sutures were then placed as single interrupted sutures 144 starting cranially and progressing caudally. The ends of the sutures were left long and 145 passed through the cast before tying under tension whilst simultaneously applying moderate laterolateral thoracic compression. Cases that had no detectable safe corridor
for suture passage underwent a minimal dissection to the caudal sternebrae which were
then directly retracted ventrally allowing the circumsternal sutures to be passed. This
wound was closed using subcutaneous 1.5M or 2M glycomer 631 (Biosyn, Covidien)
before routine cast placement. All casts were covered by chest bandages and
postoperative thoracic radiographs were taken.

152

153 Kittens who underwent surgery were discharged the next day on 5 day courses of 154 0.05mg/kg meloxicam sid (Metacam, Boehringer Ingelheim) and 20mg/kg potentiated 155 amoxicillin bid (Synulox Palatable Drops, Zoetis). The casts were maintained for 4 156 weeks at which point they were removed under anaesthesia using an identical 157 anaesthetic protocol as above. Thoracic radiographs were taken to assess the degree of 158 PE correction and postoperative VI's were calculated by the primary author.

159

Medium term follow up was obtained by email/telephone contact with the owners/referring veterinarians and the kittens were allocated new clinical severity scores. Pre and postoperative CSS were then compared.

163

164 **Results**

165

Ten kittens met the inclusion criteria during the study period. Breed distribution was: Bengal (5), Domestic Short Hair (4) and Maine Coon (1). There were four female and six male kittens. Median weight (kg) at time of surgery was 1.17 (range 0.65-2.0). Clinical signs reported included: palpable abnormality (10), tachypnoea (7), exercise intolerance (4), at least one previous episode of antimicrobial responsive dyspnoea (3), stunting/poor growth (3), dehydration (3), chronic dyspnoea (2), constipation (2) and
PU/PD (1). Allocated CSS were: 1(4 kittens), 2(3 kittens), 4(2 kittens) and 5(1 kitten).

173

174 All CT scans were deemed to be of diagnostic quality. Two kittens (G, H) did not have 175 radiographs or CT "scout" images of sufficient quality for VI and FSI to be accurately 176 measured and so these cases were omitted from the analysis. Radiographic and CT 177 measurements/indices are given in tables 3 and 4. CT and radiographically determined 178 indices (VI, FSI) were compared and the results shown in figures 1 and 2. CT 179 consistently gave a lower value for the VI with a mean difference of 0.53. FSI calculated 180 from CT images tended to be higher than the value calculated from radiographic 181 measurements

182

The distance between the dorsal most point of the deviated sternum and the overlying vertebra ("c") was determined at maximum inspiration and expiration for each of 3 respiratory cycles for two kittens for which a cine scan was performed. "c" changed by an average of 3.16% in the first kitten and 0.68% in the second.

187

188 Lung volume/kg body weight did not appear to be significantly different from that of 189 the control population (mean PE: $37.1 \text{ cm}^3/\text{kg}$, mean control: $45.3 \text{ cm}^3/\text{kg}$) although 190 low case numbers precluded statistical analysis. CT review showed that the dorsal 191 sternal deviation was lateralised in 5/10 kittens and this was dextral in 4/5 cases. The 192 kitten with sinistral deviation was diagnosed with complete situs inversus. No other 193 musculoskeletal deformities were detected. Kittens which had a lateralised sternal 194 deformity tended to have a lower CSS than those that did not (1,1,1,1,2), versus 195 2,2,4,4,5).

The sternebrae closest to the overlying major cardiovascular structures were 5-7 (4 kittens), 5-8 (2 kittens), 6-8 (2 kittens), 6-7 (2 kittens), 4-6 (1 kittens) (see figure 3).
The dorsal aspect of the sternal deformity was judged to be in contact with a major cardiovascular structure in 9/10 cases. The tenth case was judged to have a safe corridor for needle passage of 2mm.

202

203 Short term/medium term follow up:

Cases A, B, C, F, G, H and I underwent surgery. Case D was PU/PD, anorexic and significantly dehydrated at the time of presentation and was euthanased at the owners request. A post-mortem CT examination was performed in this case with the owner's consent. Cases E and J were judged to have PE of medium severity based on VI. Case E had a CSS of 1 and surgery was not recommended. Case J presented with a CSS of 4 and surgery was recommended but was declined by the owner.

210 There were no intraoperative complications in any of the cases that underwent surgery.

Case A developed furosemide responsive dyspnoea 1 week following surgery that required hospitalisation and oxygen supplementation. Dyspnoea partially resolved but required continued medication. Case A developed cardiorespiratory arrest under anaesthesia for cast removal and did not respond to resuscitation attempts. VI was improved (higher) in all cases that underwent surgical correction (table 5).

216 No moist dermatitis or pyoderma secondary to cast placement occurred in any of the

217 cases. Owners do not report any clinical symptoms of PE in all surviving kittens.

Follow up data and CSS are listed in table 5.

219

220 **Discussion**:

Ten kittens were recruited into this study, five of which were Bengals. Although adequate population data is not available for statistical analysis, this tends to support Bengal kittens being at increased risk of PE⁸ although bias in the referred population is possible.

226

227 The clinical signs reported by the owners and/or breeders were variable. Although 228 some interobserver variation is probable, all clinical histories were taken by the same 229 veterinary surgeon and the severity of historical signs reported seemed to correlate 230 with clinical signs at presentation. Tachypnoea was the most common clinical sign 231 reported after "palpable deformity" which is a diagnostic feature of PE. Tachypnoea 232 was presumed to be due to decreased lung volume and impaired alveolar exchange and 233 forms part of a continuum of signs progressing to exercise intolerance and respiratory 234 distress presumably with increased severity of deformity. This spectrum of signs 235 formed the basis for the CSS which was used in this study. Surprisingly, however, the 236 lung volume (as a proportion of body weight) of affected kittens was not significantly 237 different from a control population of unaffected cats. Larger case numbers and ideally 238 an age-matched control population would allow statistical analysis and verification of 239 these preliminary findings. Results could be further confounded by a compensatory 240 increased depth of respiration seen in affected kittens. More information could be 241 obtained by lung plethysmography but this is not widely available for veterinary 242 patients particularly those with small tidal volumes.

243

Three of the ten kittens had at least one previous episode of antimicrobial –responsivedyspnoea. In each case, dyspnoea responded rapidly to the administration of

potentiated amoxicillin. The exact site and cause of the presumed infection is not known
but this could be caused by ventilation impairment, failure to clear alveolar secretions
or other functional abnormalities.

249

Although only assessed in two of the 10 cases, the sternum did not move significantly during the respiratory cycle with "c" values varying by only 3.16% and 0.68%. This suggests that radiographically determined VI should be relatively constant independent of the respiratory phase at which they were taken.

254

255 There was reasonable correlation between radiographically determined and CT-256 determined VI and FSI, CT consistently gave a lower value for the VI with a mean 257 difference of 0.53 and FSI calculated from CT images tended to be higher than the value 258 calculated from radiographic measurements. The mean difference in FSI was 0.83 but 259 there was a significant association between the variation and the value of the FSI. These 260 findings suggest that separate reference intervals for normality and severity of VI and 261 FSI are appropriate depending on the imaging modality used but that an appropriate 262 reference interval for CT-calculated FSI can be inferred from the published radiographic 263 FSI values.

264

No significant concurrent musculoskeletal deformities were identified in any of the affected kittens. In each case the sternum started to deform in the caudal half and the deformity was dorsal but also lateral in 5/10 cases with 4/5 being deviated to the right hemithorax. The case with sinistral deviation of the xiphoid was the cat with situs inversus. The cause of this lateralisation away from the heart is uncertain but possible explanations would include displacement of the caudally deviated sternebrae by theheart and/or traction from the diaphragmatic crura during development.

272

Cardiac perforation is a recognised complication of placing a ventral corrective splint.¹² The deformed sternum was judged to be in contact with either the heart or the caudal vena cava in 9/10 cases with the 10th case having a safe corridor for suture placement of only 2mm. Having demonstrated how close the dorsal sternebrae were to the heart in all our surgical cases, we modified our surgical technique accordingly. No intraoperative complications were encountered and no postoperative complications relating to this approach were seen.

280

281 VI and FSI are useful for initial screening of cases into an anatomic severity category -282 mild, moderate or severe but they do not appear to be useful for determining whether 283 cases with moderate to severe anatomic defects are likely to have severe clinical signs. 284 This would imply that there may be other factors beyond simple musculoskeletal 285 deformity which are contributing to the severity of the clinical signs observed. Cats with 286 lateral deviation of the xiphoid seemed to be associated with a lower clinical score than 287 those kittens with a midline deviation. It has been suggested that clinical symptoms of 288 PE in people may be partially due to a direct compressive or restrictive effects of the 289 displaced sternebrae on the heart itself.¹³ Athletic performance is compromised by the 290 inability of the heart to increase diastolic volume to meet increased oxygen demands 291 and direct compression of the right side of the heart is considered to be an indication 292 for surgery in people. 5,13 It is possible that clinical signs seemed to be more severe in 293 cats with a midline pectus deformity due to a similar mechanism whereas the kittens 294 with a lateralised defect had more space available for the heart to increase diastolic

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volume as required. Cardiovascular compromise may therefore be a more significant
driver of clinical signs associated with PE in kittens than altered pulmonary function.
This could be why case J, that had a midline sternal deviation, had severe clinical signs
despite a "moderate" VI (7) and it may be that a different threshold (higher VI) should
be used when deciding if kittens with midline defects should undergo surgery.

300

301 One case (A) developed significant postoperative furosemide-responsive dyspnoea and 302 then died at the time of cast removal. The apparent initial response to diuretics is 303 suggestive of pulmonary oedema that could be caused either by pulmonary re-304 expansion, concurrent cardiovascular disease or pulmonary hypertension.¹⁴ Although 305 no significant concurrent cardiovascular disease was detected on the initial CT scan, this 306 modality is not as sensitive as echocardiography when assessing cardiac function. The 307 cause of death at the time of cast removal remains unknown as no post-mortem analysis 308 was permitted.

309

310 All cases that survived showed full resolution of clinical symptoms with no exercise 311 intolerance or episodes of dyspnoea reported at the time of follow up. Whilst we used a 312 combination of VI and CSS to determine which cases should benefit from surgery, there 313 was no control population for which treatment was intentionally withheld in order to 314 demonstrate a difference in postoperative outcome as this would have been unethical. 315 We suspect that many cases of severe PE are euthanased due to perceived poor 316 prognosis and financial concerns about treatment costs. This, and the rarity of the 317 condition resulted in only low numbers of cases being recruited despite internet based 318 advertising for case enrolment. We are therefore limited to making broad 319 recommendations about patient selection and treatment efficiency. In our study, patients were selected for surgery based on VI and clinical signs. All cases had a sternal cast maintained for 4 weeks and all cases that survived are currently asymptomatic at a mean follow up of 15 months. One of the risks of uncorrected PE is the development of pulmonary hypertension and right side heart failure which the authors have observed in multiple cases < 12mths old. It is possible that some of our cases could develop respiratory symptoms at a later stage and we intend to publish longer term follow up (5 year) data when available.

327

328 In summary, conventional radiography yields reasonable approximations of CT-329 determined VI and FSI. CT was useful in determining the presence/absence of safe 330 corridors for circumsternal suture placement leading to a minor modification of the 331 surgical approach employed for ventral cast placement. CT also allowed detection of 332 cats with midline sternal deviation which may be at risk of developing more severe 333 clinical signs due to diastolic restriction despite relatively mild skeletal deformity. Ten 334 to 15 week old kittens with severe deformity as judged by VI and with compatible 335 clinical signs can be treated by placement of a sternal splint for a 4 week period which 336 can be associated with an excellent medium term outcome.

337

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341

- 342 **Conflict of Interest**
- 343 The authors declare that there is no conflict of interest.

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