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AUTHORS: Lamb, C. R. and Jones, I. D.

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1 Abstract

Objectives. To estimate the prevalence of subclinical abnormalities reported in thoracic CT scans of
 cats and to test associations between respiratory signs and CT signs.

Methods. Retrospective review of signalment, indications, respiratory signs and reported CT findings
in a series of cats having thoracic CT scans. Associations between patient variables, respiratory signs
and CT signs were tested using multivariable regression methods.

7 Results. Records of 352 consecutive cats were reviewed. Abnormalities affecting thoracic structures

8 were reported in CT scans of 138/179 (77%) cats without respiratory signs. The most prevalent CT

9 findings in cats without respiratory signs were pulmonary collapse (41%), signs of bronchial disease

10 (24%) and space-occupying lesions (21%). Dyspnoea, cough and tachypnoea were associated with

11 space-occupying lesions. Dyspnoea was also associated with pulmonary consolidation and

12 atelectasis. Increasing body weight was associated with pulmonary atelectasis and increasing age

13 was associated with signs of bronchial disease.

14 Clinical significance. Abnormalities are reported frequently in thoracic CT scans of cats without

15 respiratory signs. The most prevalent abnormality, pulmonary atelectasis, is probably a temporary

16 effect of sedation or anaesthesia for CT. A high prevalence of subclinical abnormalities and limited

17 correlations between clinical signs and CT findings will complicate diagnosis.

18 Introduction

19 Computed x-ray tomography (CT) is used increasingly to investigate cats with thoracic conditions 20 (Henninger 2003; Cohn et al. 2004; Oliveira et al. 2011; Aarsvold et al. 2015). Being a cross-sectional 21 imaging modality, CT eliminates the problem of superimposition that affects radiography and 22 therefore enables clearer depiction of internal anatomy and morphological abnormalities that alter 23 anatomy. As a result CT has a higher sensitivity for pulmonary disease than radiography (Nemanic et 24 al. 2006). In general, higher sensitivity is advantageous, but one of the drawbacks of using more 25 sensitive diagnostic tests is that more of the abnormalities detected may represent clinically silent 26 conditions and, potentially, incidental findings (Aspinall et al. 2013). For example, previous CT 27 studies have documented the tendency for lung collapse in anaesthetised cats (Henao-Guerrero et 28 al. 2012). It is the authors' impression that other abnormalities, such as signs of bronchial disease, 29 are observed frequently in cats having thoracic CT for reasons unrelated to respiratory signs. 30 Clinicians interpreting diagnostic images must always consider the possible relationship between 31 imaging signs and clinical signs in order to assess the likely meaning of the images (Wood, 1999). For 32 example, it is generally accepted that coughing in cats is more associated with bronchial lesions 33 whereas dyspnoea is more associated with cardiac disease or space-occupying lesions, such as a 34 pulmonary mass or pleural fluid (Moise et al. 1989; Corcoran et al. 1995; Dye et al. 1996; Sauve et al. 35 2005; Swift et al. 2009; Foster & Martin 2011; Johnson & Vernau 2011; Sigrist et al. 2011); however, 36 such associations are not pathognomonic, so it can be difficult to judge if the imaging signs in an 37 individual patient fully explain their clinical signs. Particularly in patients with non-specific or vague 38 clinical signs, it can be difficult to decide if an imaging finding represents a clinical or subclinical 39 condition, and whether to treat it or employ further diagnostic tests.

The aims of the present study were to estimate the prevalence of subclinical abnormalities reported
in thoracic CT scans of cats and to test the associations between clinical signs and CT signs.

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43 Methods

44 For this retrospective cross-sectional study, electronic medical records of cats referred to the Queen 45 Mother Hospital for Animals (QMHA) between September 2013 and September 2015 were 46 reviewed. The only inclusion criterion was CT imaging of the thorax. The signalment, body weight, 47 body condition score, indication for CT and respiratory signs (if any) were extracted from 48 contemporaneous clinical notes by one investigator (IDJ). Indications for CT were categorised as 49 follows: suspected respiratory condition, metastasis check, trauma, comprehensive medical work-50 up, other. Respiratory signs were categorised as follows: none, cough, upper respiratory signs 51 (stertor, change of voice, nasal discharge, nasal obstruction, dyspnoea), dyspnoea not associated 52 with upper airway signs, tachypnoea, hyperpnoea, other. 53 CT scans were acquired using a multi-detector row CT scanner (MX 8000 IDT, Philips Medical 54 Systems, Cleveland, USA). CT images were obtained using helical acquisition, 120kVp, 200mAs, 16 x 55 1.5mm collimation, 0.688 pitch, 2mm slice thickness, 150-300mm field of view, 512x512 matrix (pixel size 0.3-0.6mm). All cats had separate CT series to examine the lung and the thoracic soft 56 57 tissues, respectively. Images of the lung were reconstructed using a high frequency (lung enhanced) 58 algorithm whereas images of the thoracic soft tissues were reconstructed using a sharp 59 reconstruction algorithm. When indicated, post-contrast CT images of the thoracic soft tissues were 60 obtained 60s after the start of rapid intravenous injection of 2ml/kg bolus of iohexol (Omnipaque 61 300, GE Healthcare, Oslo, Norway). Use of sedation or anaesthesia for CT was recorded. 62 Abnormalities in CT scans, as reported individually at the time of clinical work-up by the six Board-63 certified radiologists employed at the QMHA in this period, were extracted from imaging reports by 64 another investigator (CRL) without reference to the clinical information. Experience of small animal 65 CT in this group of radiologists ranged from 5 to 26 years. CT signs were categorised as follows: space-occupying lesions (pulmonary mass, other intrathoracic mass, pleural fluid, pneumothorax, 66 67 pneumomediastinum), pulmonary nodules (solid, subsolid, cavitary, bulla/cyst), consolidation

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68 (sublobar, lobar, bronchocentric, multifocal), atelectasis/collapse (sublobar, lobar, bronchocentric, 69 multifocal), ground-glass opacities (focal, multifocal, diffuse), air-trapping (focal/lobar, multifocal, 70 diffuse, interstitial emphysema), bronchial lesions (wall thickening, bronchiectasis, mucus plugging, 71 "tree-in-bud," foreign body, broncholith), other (honeycombing, septal lines, parenchymal bands, 72 adenopathy, vascular congestion) and thoracic wall lesions (rib fractures, barrel-chested 73 conformation, pectus excavatum, peritoneopericardial diaphragmatic hernia, other). The "tree-in-74 bud" pattern applies to centrilobular branching structures that resemble a budding tree. 75 Terminology followed recommendations for thoracic imaging by the Fleischner Society (Hansell et al. 76 2008). 77 Calculation based on the need to estimate a prevalence of 30% to within +/-5% indicated a sample 78 size equal to 322 cats (Eng 2003). Associations between respiratory signs and reported CT signs were 79 tested using multinomial logistic regression with respiratory signs as the dependent variable. 80 Associations between age, body weight, body condition score, use of sedation or anaesthesia and 81 atelectasis were tested using binary logistic regression. The association between age and bronchial 82 lesions was also tested using binary logistic regression. Statistical testing was done using SPSS 83 Statistics, version 22 (IBM Corporation, Chicago, IL). Results with p<0.05 were considered significant. 84 85 Results 86 Records of 352 consecutive cats were reviewed. No cats were excluded. There were 195 (55%) male

cats (186 neutered) and 157 (45%) females (150 neutered). The most prevalent breeds were

88 Domestic Short Hair (n=213, 61%), Domestic Long Hair (34, 10%), British Short Hair (14, 4%), Siamese

89 (14, 4%) and Maine Coon (12, 3%). There were 21 other feline breeds with less than 10 individuals.

90 Age was known in 350 cats. Median (range) age of cats was 9.5y (4m – 18y). Body weight was

91 recorded in 311 cats and body condition score was recorded in 202 cats. Median body weight was

92 4.4kg (range 1.2 - 13.1kg) and median body condition score was 4 (range 1 - 9).

The principal indications for thoracic CT were respiratory signs (n= 144, 41%), comprehensive
medical work-up (107, 30%), search for metastasis (64, 18%) and trauma (37, 11%). Respiratory signs
were present in 173 (49%) cats, including upper respiratory signs in 78 (22%) cats, dyspnoea not
associated with upper airway signs in 42 (12%) cats, combinations of two or more other categories
of respiratory signs in 20 (6%) cats, tachypnoea in 17 (5%) cats and cough in 16 (4%) cats. There were
no cats in which hyperpnoea was recorded as the sole respiratory sign.

99 Two hundred and thirty-seven (67%) cats were anaesthetised for CT and the remainder were 100 sedated. Cats were positioned in sternal recumbency for CT, except for four cats with suspected 101 vertebral fractures, which were placed in lateral recumbency. Abnormalities affecting thoracic 102 structures were reported in CT scans of 138 (77%) cats without respiratory signs, 65 (83%) cats with 103 upper respiratory signs and 91 (96%) cats with other respiratory signs (p=0.0004). CT scans were 104 interpreted as normal in 58 (16%) cats.

105 Abnormalities reported in CT scans of the 179 cats without respiratory signs are summarised in table 106 1. The most prevalent abnormalities in these cats were pulmonary atelectasis (reported in 41%) and 107 signs of bronchial disease (24%). No abnormalities were reported in 23% cats without respiratory 108 signs. Of the 57 cats without respiratory signs in which the indication for CT was to search for 109 metastasis, 50 (88%) had a cellular diagnosis of neoplasia and, of these, 15 (30%) had pulmonary 110 nodules and 9 (18%) had pleural fluid reported on CT. Of the 28 cats without respiratory signs in 111 which the indication for CT was trauma, 9 (32%) had space-occupying lesions, including 112 pneumomediastinum in 5 (18%) and pneumothorax in 4 (14%). The 92 cats without respiratory signs 113 in which the indication for CT was the need for a comprehensive medical work-up included a wide 114 range of conditions. The most prevalent of these were 21 cats subsequently shown to have 115 neoplasia. Of these, 5/21 (24%) had space-occupying lesions (3 pulmonary mass, 1 mediastinal cyst, 116 1 pleural effusion) and 4/21 (19%) had pulmonary nodules on CT. The second most prevalent group 117 were 15 cats with uncontrolled diabetes mellitus (including 11 with hypersomatotropism). Of these,

only one diabetic cat had a CT abnormality related to its condition, which was pleural effusionsecondary to metastatic adrenal carcinoma.

120 The frequencies of reported CT abnormalities CT in cats with different respiratory signs are 121 summarised in table 2 and the results of statistical testing are summarised in table 3. Air-trapping 122 and thoracic wall lesions were excluded from the statistical analysis because of the small number of 123 affected individuals. The strongest association found between respiratory signs and CT abnormalities 124 was between dyspnoea and space-occupying lesion (odds ratio 8.7, 95% confidence interval 3.6-125 21.1) (p=0.00002). Cough and tachypnoea were also significantly associated with a CT report of a 126 space-occupying lesion. Space-occupying lesions reported in 99 cats in this series were pulmonary 127 mass in 25%, other thoracic mass in 27%, pleural fluid in 53%, pneumothorax in 18% and 128 pneumomediastinum in 11% cats. Masses and pleural fluid occurred together in 17% cats with 129 space-occupying lesions. Dyspnoea was also significantly associated with a CT report of pulmonary 130 consolidation or atelectasis. Upper respiratory tract signs were significantly associated with a CT 131 report of bronchial lesions and inversely associated with space-occupying lesions or ground-glass 132 opacities. 133 Overall, pulmonary atelectasis/collapse was the most frequent reported CT abnormality, affecting 134 159 (45%) cats in this series. The most frequently collapsed lobes, in descending order were the right 135 middle (collapsed in 53% affected cats), left cranial (45%), right cranial (39%), left caudal (16%), 136 accessory (11%) and right caudal (9%). All lobes had some degree of atelectasis in 8% affected cats. 137 The predominant distribution of atelectasis was sub-lobar, which was reported in 71% lobes affected 138 by atelectasis. Other reported distributions of pulmonary atelectasis were lobar (19%),

139 bronchocentric (5%) and multifocal (5%) (Figure 1). There was an association between body weight

and CT signs of pulmonary atelectasis (odds ratio 1.2, 95% confidence interval 1.0-1.5) (p=0.02).

141 Neither body condition score (p=0.8) nor use of anaesthesia (p=0.08) were significantly associated

142 with pulmonary atelectasis.

Bronchial abnormalities were reported in CT scans of 103 (29%) cats in this series. Reported
bronchial abnormalities were thickened bronchial walls in 91%, mucus plugging in 14%, tree in bud
in 5%, bronchiectasis in 3% and broncholithiasis in 1% instances (Figure 2). Signs of thickened
bronchial walls and mucus plugging occurred together in 12% cats with bronchial lesions. There was
an association between age and CT signs of bronchial disease (odds ratio 1.12, 95% confidence
interval 1.07-1.19) (p=0.0003).

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150 Discussion

These results support the conclusion that the prevalence of abnormalities reported in thoracic CT scans is high (77%) in cats having CT for reasons other than to investigate respiratory signs. A high prevalence of potentially subclinical abnormalities is problematical because it will complicate diagnosis and tend to undermine the correlations between clinical signs and CT findings that are normally used as a basis for interpretation of findings.

156 Although the high prevalence of lung atelectasis observed in the present study is probably a 157 temporary effect of the routine use of sedation or anaesthesia for CT, signs of bronchial disease and 158 space-occupying lesions were also reported frequently in cats without respiratory signs. Previous studies have found that cats with infectious pneumonia may lack respiratory signs (Macdonald et al. 159 160 2003) and that histological lesions affecting the lung of cats with diabetes mellitus may be 161 unassociated with respiratory signs (Mexas et al. 2006). These findings may correspond with the 162 widely-held belief that the cat – like the sheep – is a species able to hide signs of disease, with the 163 result that by the time an owner recognises their pet is unwell, its condition may be advanced; 164 however, the present study provides no information about the possibility that certain subclinical 165 lesions may eventually become serious. Only a cohort study with long term follow-up of patients 166 could determine the extent to which that occurs. Alternatively, many of the abnormalities reported

168 signs. If so, labelling affected cats carries the potential for overdiagnosis (Moynihan et al. 2012). 169 In this study, respiratory signs were significantly associated with abnormal findings on CT but there 170 were few significant associations between specific respiratory signs and CT-detected abnormalities. 171 Dyspnoea, cough and tachypnoea were associated with space-occupying lesions. Dyspnoea was also 172 associated with pulmonary consolidation and atelectasis. A previous study found that cardiac 173 disease is the most frequent cause of dyspnoea in cats (Swift et al. 2009). This finding is applicable to 174 a referral population of cats, but the present study underrepresented this group of patients because 175 relatively few cats with cardiac disease are considered candidates for CT. Cough was a relatively 176 infrequent clinical sign in this study and was not significantly associated with bronchial abnormalities 177 on CT. An association between expiratory dyspnoea and lower airway disease in cats has been 178 reported (Sigrist et al. 2011), but a similar association was not evident in the present study. 179 Bronchial abnormalities, mainly thickened bronchial walls and signs of mucus plugging, were 180 observed in many cats with no respiratory signs or signs referable to the upper respiratory tract, 181 hence the clinical significance of these findings is uncertain. One of the limitations of the present 182 study is that the severity of radiographic signs cannot be assessed on the basis of CT reports, hence 183 it is unclear if – for example –signs of bronchial disease were more marked in cats with cough than 184 cats without respiratory signs.

in the thoracic CT scans in this series may represent static conditions that will never cause clinical

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185 The tendency for lung atelectasis in anaesthetised cats has been reported previously (Henao-186 Guerrero et al. 2012). In dogs, use of high inspired pO2 during anaesthesia has been associated with 187 a higher prevalence of lung atelectasis than moderate inspired pO2 (Staffieri et al. 2007). The same 188 effect may apply to cats, although this was not evident in the present study because cats having CT 189 under general anaesthesia were no more likely to have lung atelectasis than those having CT under 190 sedation. Avoiding lung atelectasis may be expected to aid interpretation of CT images and to 191 increase sensitivity for lesions that could be obscured by lung atelectasis. A reduced tendency for 192 lung atelectasis may be an advantage of conscious CT for cats (Oliveira et al. 2011). Hence, CT

193 scanning of conscious cats or cats under anaesthesia with positive pressure ventilation may be 194 preferred; however, a recent study of anaesthetised dogs and cats that had brief hyperventilation 195 and CT scan during the subsequent expiratory pause (which is the technique usually employed in the 196 authors' hospital), found that ability to detect pulmonary nodules or bulla was not significantly 197 different than when images were acquired with the lungs inflated by positive pressure ventilation 198 (Simone et al. 2016). Cats in ventral or dorsal recumbency are probably affected less by lung 199 atelectasis than those in lateral recumbency, although this cannot be tested in the present study 200 because at the authors' hospital only cats presented after trauma that are considered at risk of being 201 repositioned have CT scans in lateral recumbency (Figure 1D).

202 In this study, lung atelectasis was associated with increasing body weight, likely reflecting the 203 tendency for reduced lung volume in cats with accumulation of intrathoracic fat. Body condition 204 score is used widely as a method of estimating obesity (Laflamme 1997), and would be expected to 205 be correlated with likelihood of lung atelectasis under anaesthesia, possibly better than body 206 weight, but that was not observed. The lack of recorded body condition score in a large proportion 207 of cats will have reduced statistical power and may at least partly explain this negative result. 208 The most frequently collapsed lung lobes in cats in the present study were the right middle and the 209 left and right cranial lobes. The right middle lobe is most prone to collapse in the cat (and dog) 210 (Robinson 1982) and is frequently found to be collapsed in cats with bronchial disease because of 211 mucus or exudate obstructing the airways (Moise et al. 1989). The association found between CT 212 signs of bronchial disease and patient age may reflect progressive disease that is initially feline 213 asthma. Asthma is thought to the result of an allergic response triggered by inhaled allergens that 214 leads to bronchial smooth muscle constriction and airway narrowing (Trzil & Reinero 2014). Many

cats with asthma have minimal radiographic signs other than pulmonary overinflation (Gadbois et al.

2009); however, chronic antigenic stimulation in affected cats leads to permanent bronchial luminal

217 narrowing and wall thickening because of smooth muscle hyperplasia, goblet cell and submucosal
218 gland hypertrophy and hyperplasia, epithelial erosion and eosinophilic infiltration (Padrid et al. 1995;

219 Reinero et al. 2004) and these pathological changes are more likely to become visible

radiographically.

221 It should be emphasised that this study was not designed to estimate the accuracy of CT for thoracic 222 lesions in cats or to estimate the reliability of observers. Our principal aim was to estimate the 223 prevalence of subclinical abnormalities reported in thoracic CT scans of cats and this is probably best 224 achieved by extracting data from imaging reports produced in a consecutive series of patients by 225 radiologists working according to their normal routine in a clinical environment. As a result, the 226 authors' data include variability occurring as a result of patients (all were included regardless of 227 indication or diagnosis) and radiologists (including hawks and doves, working with clinical histories of 228 varying quality). Other methods, such as limiting the study to cats in which there is a certain final 229 diagnosis, or employing a panel of observers to reach a consensus about the CT findings without 230 knowledge of the clinical history, would produce results less representative of clinical practice.

Table 1. Abnormalities reported in CT scans of 179 cats without respiratory signs

	CT abnormalities									
	None	Space- occupying lesion	Pulmonary nodules	Consolidation	Atelectasis	Ground glass opacity	Air- trapping	Bronchial lesions	Other	Thoracic wall lesion
Indication for CT										
Suspected respiratory condition ^a n=1	0	1 (100%)	0	0	1 (100%)	0	0	0	1 (100%)	0
Metastasis check n=57	11 (19%)	10 (18%)	17 (30%)	4 (7%)	21 (37%)	7 (12%)	0	12 (21%)	7 (12%)	2 (4%)
Trauma n=28	6 (21%)	9 (32%)	1 (4%)	0	16 (57%)	0	0	3 (11%)	3 (11%)	2 (7%)
Comprehensive medical work-up n=92	24 (26%)	18 (20%)	11 (12%)	9 (10%)	35 (38%)	8 (9%)	0	27 (29%)	10 (11%)	1 (1%)
Other ^b n=1	0	0	0	0	1 (100%)	0	0	1 (100%)	0	0
Totals	41 (23%)	38 (21%)	29 (16%)	13 (7%)	74 (41%)	15 (8%)	0	43 (24%)	21 (12%)	5 (3%)

^a Reduced lung sounds on auscultation, subsequently shown to be secondary to pleural effusion

^b Pre-operative assessment of suspected vascular ring anomaly

Table 2. Associations between respiratory signs and abnormalities reported in CT scans of 352 cats

	CT abnormalities									
	None	Space- occupying	Pulmonary nodules	Consolidation	Atelectasis	Ground glass	Air- trapping	Bronchial lesions	Other	Thoracic wall lesion
Respiratory signs		lesion				opacity				
None n=179	41 (23%)	38 (21%)	29 (16%)	13 (7%)	74 (41%)	15 (8%)	0	43 (24%)	21 (12%)	5 (3%)
Cough n=16	0	8 (5%)	2 (13%)	1 6(%)	10 (62%)	2 (13%)	1 (6%)	7 (44%)	0	0
Upper respiratory n=78	13 (17%)	8 (10%)	12 (15%)	5 (6%)	34 (44%)	2 (3%)	0	33 (42%)	4 (5%)	1 (1%)
Dyspnoea n=42	1 (2%)	30 (71%)	3 (7%)	8 (19%)	28 (67%)	3 (7%)	2 (5%)	9 (21%)	9 (21%)	6 (14%)
Tachypnoea n=17	3 (18%)	7 (53%)	4 (24%)	3 (18%)	4 (24%)	0	0	3 (18%)	4 (24%)	0
Combination of respiratory signs n=20	0	6 (30%)	4 (20%)	4 (20%)	9 (45%)	2 (10%)	0	8 (40%)	2 (10%)	0
Totals	58 (17%)	99 (28%)	54 (15%)	34 (10%)	159 (45%)	24 (7%)	3 (1%)	103 (29%)	40 (11%)	12 (3%)

Table 3. Significant associations between respiratory signs and abnormalities reported in CT scans of 352 cats

	CT abnormalities									
Respiratory signs	Space- occupying lesion	Pulmonary nodules	Consolidation	Atelectasis	Ground glass opacity	Bronchial lesions	Other			
Cough	4.1 (1.3-13.3) p=0.017	NS	NS	NS	NS	NS	NS			
Upper respiratory	0.4 (0.17-0.98) p=0.04	NS	NS	NS	0.2 (0.04-0.8) p=0.03	2.0 (1.1-3.8) p=0.04	NS			
Dyspnoea	8.7 (3.6-21.1) p=0.00002	NS	5.0 (1.5-17.0) p=0.009	3.5 (1.4-8.9) p=0.009	NS	NS	NS			
Tachypnoea	3.8 (1.1-13.3) p=0.03	NS	NS	NS	NS	NS	NS			
Combination of respiratory signs	NS	NS	NS	NS	NS	NS	NS			

Values are odds ratio (95% confidence interval)

NS, non-significant, p>0.05

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Legends



Figure 1. Examples of CT images reported as showing signs of pulmonary atelectasis. All are transverse images displayed with window 1500HU and centre -500HU. A) Sub-lobar atelectasis affecting the tip of the right cranial lobe (arrowheads) in a cat with chronic cough. There is mediastinal shift to the right. Other images showed signs of bronchial disease. B) Bronchocentric atelectasis affecting the left cranial lobe (arrowhead) in a cat with metastatic adenocarcinoma and no history of respiratory signs. C) Lobar atelectasis affecting the right middle lobe (between arrowheads) in a cat with chronic cough. The heart (H) is displaced to the right. Other images showed signs of bronchial disease. D) Unilateral left lung atelectasis (*) in an anaesthetised cat that had CT scan in left lateral recumbency because of trauma and suspected spinal fracture. The heart (H) is displaced to the left.



Figure 2. Examples of CT images reported as showing signs of bronchial disease. All are transverse images displayed with window 1500HU and centre -500HU. A) Thickened bronchial walls (arrowheads) in a cat with a suspected vertebral neoplasm and no history of respiratory signs. There is also ground-glass opacity (*) affecting the right cranial lobe. B) Mucus plugging of the right cranial lobar bronchus (between arrowheads) in a cat with chronic cough and tachypnoea. C) Mucus plugging of dilated bronchi (arrowheads) at the periphery of the left caudal lobe in a cat with multiple cutaneous mast cell tumours and no history of respiratory signs. The diameters of each of these bronchi grossly exceed the diameter of the bronchus in the right caudal lobe (arrow) compatible with bronchiectasis. D) Tree-in-bud sign (arrowhead) in the left caudal lobe of a cat with chronic cough.