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AUTHORS: Frances E Taylor-Brown, Steven De Decker

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1	Acute non-compressive nucleus pulposus extrusion in cats: clinical features, diagnostic
2	imaging findings, treatment and outcome
3	Taylor-Brown, F.E., De Decker, S.
4	Clinical Science and Services, The Royal Veterinary College, University of London, Hatfield,
5	UK
6	
7	Corresponding author: Frances Taylor-Brown BSc (Hons), BVetMed (Hons), MRCVS
8	Email: <u>ftaylor@rvc.ac.uk</u>
9	Address: Clinical Science and Services, The Royal Veterinary College, University of London,
10	Hawkshead Lane, North Mymms, Hatfield, Hertfordshire, AL9 7TA, UK
11	Tel: +44(0)1707 666366
12	Fax: +44 (0)1707 649384
13	
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15	Keywords:
16	Cat; disc; extrusion; non-compressive; nucleus pulposus; outcome
17	Abstract
18	Objectives: The aim of the study was to describe the clinical features, diagnostic imaging
19	findings, treatment and outcome in cats diagnosed with presumptive acute non-compressive
20	nucleus pulposus extrusion.

Methods: Medical records and imaging studies of cats diagnosed with presumptive acute noncompressive nucleus pulposus extrusion were retrospectively reviewed. Long-term follow-up information was acquired from patient records and from either owners or referring veterinary surgeons via a telephone questionnaire.

Results: Eleven cats met the inclusion criteria. All cats had a peracute onset of clinical signs, 25 with eight cats experiencing witnessed (n = 6) or suspected (n = 2) external trauma. 26 27 Neurological examination findings ranged from ambulatory paresis to plegia with loss of deep nociception. Neuroanatomical localisation included C1-C5 (n = 1), T3-L3 (n = 7) and L4-S3 28 29 (n = 3) spinal cord segments. Ten cats were discharged with a median hospitalisation time of 10 days (range 3 days to 26 days). One cat was euthanised during hospitalisation due to 30 complications unrelated to neurological disease. Cats that presented with paraplegia regained 31 32 voluntary movement within a median of 4 days (range 2 to 7 days). For those cats that presented non-ambulatory, all cats regained an ambulatory status with the median time to ambulation of 33 17 days (range 6 to 21 days). Five cats had absent voluntary urination at presentation; this 34 resolved in all but one cat that had long-term urinary incontinence. Overall the outcome for 35 cats diagnosed with acute non-compressive nucleus pulposus extrusion was good with almost 36 37 90% returning to ambulation with urinary and faecal continence.

38 *Conclusions and relevance*: The majority of cats diagnosed with acute non-compressive 39 nucleus pulposus extrusion had characteristic clinical presentations and good outcomes. Acute 40 non-compressive nucleus pulposus extrusion should be considered as a differential diagnosis 41 for cats presenting with peracute onset of spinal cord dysfunction, particularly if there is a 42 clinical history or evidence of trauma.

44 Introduction

Acute non-compressive nucleus pulposus extrusion (ANNPE), previously referred to as a traumatic intervertebral disc extrusion (IVDE), high-velocity/low-volume IVDE and type III intervertebral disc extrusion occurs when a healthy and hydrated IVDE is exposed to sudden and excessive force and is typically seen following vigorous exercise or trauma¹⁻⁴. This type of intervertebral disc extrusion results in spinal cord contusion with minimal or no spinal cord compression¹⁻³.

ANNPE has been frequently reported in dogs¹⁻³, however there are only single case reports describing ANNPE in cats^{4,5}. Dogs with ANNPE typically present with a peracute onset of spinal cord dysfunction that is non-progressive after 24 hours ^{1,2}. Clinical signs are often strongly lateralised and mild to moderate spinal hyperaesthesia may be seen in approximately half of affected cases^{2,6}.

Definitive diagnosis of ANNPE can only be confirmed by histopathology¹. However, magnetic resonance imaging (MRI) can be used to make a presumptive diagnosis with with specific characteristics identified to reach a presumptive ante-mortem diagnosis of ANNPE.^{2,3}.

Typical treatment involves physiotherapy and supportive care with the use of analgesics as
 required⁶. The outcome is considered good in dogs with only a minority failing to regain normal
 neurological function².

Despite this disorder being well characterised in dogs, little is known about the clinical presentation, imaging findings and outcome in cats. The aims of this study were therefore to describe the clinical features, diagnostic imaging findings, treatment and outcome in a larger number of cats diagnosed with presumptive ANNPE. We hypothesised that cats diagnosed with presumptive ANNPE would have a characteristic presentation and a good long-term outcome.

68 Material and Methods

69 *Ethics Statement*

70 Ethics approval was granted by the Royal Veterinary College (RVC) Ethics and Welfare
71 Committee (reference number 2015 1324).

72 Criteria for inclusion

Medical records of cats that had presumptively been diagnosed with ANNPE at the RVC 73 between 2008 and 2014 were reviewed. In order to be included, cats needed to have had an 74 75 MRI of the affected spinal cord segments within 48 h of the onset of clinical signs, MRI findings consistent with the diagnosis of presumptive ANNPE and have follow-up information 76 for a minimum of 3 months. Recorded information included immediate history preceding onset 77 78 of clinical signs, treatment prior to referral, signalment, general physical examination findings, neurological examination findings, duration of time from detecting neurological signs to MRI, 79 80 treatment administered following diagnosis, duration of hospitalisation and presence of complications. In relevant cases the time to recover nociception, voluntary motor activity and 81 82 unassisted ambulation was also recorded.

83 *Diagnostic imaging*

MRI was performed using a 1.5 Tesla scanner (Intera, Philips Medical Systems) and included 84 85 a minimum of T2- and T1-weighted sagittal and transverse images. All imaging studies were reviewed for diagnostic accuracy by a board certified neurologist (SDD) blinded to the clinical 86 signs and neuroanatomical localisation, and only those cases with imaging features consistent 87 with presumptive ANNPE diagnosis were included in the study. MRI findings compatible with 88 ANNPE included (1) a reduction in volume of the T2- weighted hyperintensity of the nucleus 89 pulposus signal, (2) a focal T2-weighted hyperintensity within the spinal cord overlying an 90 intervertebral disc space, (3) mild narrowing of the intervertebral disc space, and (4) extraneous 91

92 material or signal change within the vertebral canal with absent or minimal spinal cord
93 compression^{2,3,5} (Figure 1 a, b).

94 Assessment of outcome

Short-term outcome was defined as the period between the onset of clinical signs up to 6 95 weeks following presumptive diagnosis of ANNPE, and information was retrieved from 96 medical records. Long-term outcome was defined as a minimum follow-up period of 3 97 months⁷. This information was initially obtained via telephone interview with the referring 98 veterinary surgeons. For cats that were deceased, date and cause of death as well as the last 99 documented neurologic status were recorded. Conforming to local ethics and welfare 100 committee guidelines, only owners of cats that were still alive at the time of data collection 101 were subsequently contacted. Owners were mailed a letter with study details and a 102 103 standardized questionnaire that had been reviewed and approved by a local ethics and welfare committee. Telephone interviews were conducted using the questionnaire, which included 104 questions covering specific aspects of the disease, such as amount of activity, lameness, 105 paresis and incontinence, type of medical and supportive treatment received, response to 106 treatment and quality of life (supplementary material). A successful outcome was defined as 107 resolution or improvement of clinical signs with the cat being able to ambulate independently 108 with control of urination and defaecation, while an unsuccessful outcome was defined as a cat 109 that required support to ambulate or had persistent urinary or faecal incontinence. 110

111

112 **Results**

Of 14 potential cats identified, 11 were included in the study (Table 1). The cats had a median
age of 7 years (range 2 years 9 months to 13 years) at presentation. Eight of the cats were male

neutered and three were female neutered. Breeds comprised the domestic shorthair (n = 6),
domestic longhair (n = 3), Egyptian Mau (n = 1) and British Shorthair (n = 1).

117 *Historical findings*

All cats had an acute or peracute onset of clinical signs. The median time to presentation was 14 h (range 2–48 h) following the onset of neurological signs. Prior to presentation six of the cats had been involved in a witnessed traumatic event (road traffic accident [n = 3] or fall from a height [n = 3]). The remaining five cats were found either in the home or nearby the house and the onset of clinical signs was not witnessed.

123 Clinical findings

The majority of cats (n = 10) had clinical signs referable to the paraparesis or paraplegia (Table 1). Neuroanatomical localisation included the C1–C5 (n = 1), T3–L3 (n = 7) and L4–S3 (n = 3) spinal cord segments. The clinical signs were non-progressive in all cats following presentation. Five of the cats had signs consistent with external trauma, including head trauma, pulmonary contusions and scuffed nails.

129 MRI findings

MRI revealed ANNPE located at C3–C4 (n = 1), T12–T13 (n = 1), T13–L1 (n = 1), L1–L2 130 = 1), L3–L4 (n = 3), L4–L5 (n = 1) and L5–L6 intervertebral disc spaces (n = 3). One cat had 131 a dorsal spinous process fracture of the L7 vertebra, which was not associated with the 132 neuroanatomical localisation nor the anatomical localisation of the ANNPE and was therefore 133 considered incidental. There was evidence of ill-defined T2-weighted hyperintensity within the 134 epaxial musculature compared with surrounding muscle suggestive of contusion, haemorrhage 135 or oedema in five cats (Figure 1c). Of these five cats, two cats had no history or examination 136 findings consistent with trauma, while the other three cats were involved in a witnessed trauma. 137

138 *Treatment and short-term outcome*

All cats received physiotherapy performed by a veterinary physiotherapist and/or qualified veterinary nurse consisting of massage, passive range of motion exercises, assisted standing and exercises to develop strength and coordination, as appropriate and tolerated by each cat. Five cats that demonstrated signs of spinal hyperaesthesia received analgesic medication that included opioids (ie, methadone and buprenorphine; n = 3), non-steroidal anti-inflammatory drugs (n = 1) and gabapentin (n = 1).

The median time for cats with absent deep nociception (n = 3) to regain sensation was 2 days (range 1–3 days). Of the cats that presented with paraplegia, including those with absent deep nociception (n = 5) the median time for them to regain voluntary movement (non-ambulatory) was 4 days (range 2–7 days). For those cats that presented non-ambulatory (including paraplegic cats; n = 9) the median time to ambulation was 17 days (range 6–21 days).

Five cats required bladder management during hospitalisation, including indwelling catheter placement (n = 1), intermittent catheterisation (n = 1) and manual bladder expression (n = 3). Two cats received a sympatholytic medication (prazosin) to aid in bladder management. Three cats were discharged with improved motor function but continued to require manual bladder expression.

155 The cats had a median hospitalisation time of 10 days (range 3–26 days). Four of the 10 cats 156 that survived to discharge were ambulatory at that time. One cat did not survive until discharge, 157 and was euthanased owing to respiratory deterioration as a result of pulmonary contusions.

Short-term outcome (4–6 weeks following diagnosis of presumptive ANNPE) in six cats revealed all cats were ambulatory and had improved neurological function compared with the time of discharge; however, none of the cats were considered to be neurologically normal.

161 *Long-term outcome*

Long-term outcome in eight cats (four cats were also included in the assessment of short-term 162 outcome) was obtained from the referring veterinary surgeons (n = 2) or veterinary surgeons 163 and owners (n = 6). The median duration of time between the onset of clinical signs and 164 assessment of outcome was 44 months (range 4–68 months). None of the cats displayed signs 165 of further improvement 6 months after reaching a presumptive diagnosis of ANNPE. Although 166 all cats were ambulatory and did not demonstrate any signs of spinal hyperaesthesia, none were 167 reported to have become neurologically normal. Owners or veterinary surgeons assessed all 168 cats to have regained a good quality of life; however, quality of life was considered decreased 169 170 compared with before the onset of clinical signs in all of the cats with 3/8 cats now indoor-only cats. 171

One cat (cat 2) had ongoing urinary incontinence requiring twice daily manual bladderexpression, and the same cat had intermittent faecal incontinence (Table 1).

Overall, 7/8 cats (88%) were considered to have a successful long-term outcome, and one cat
was considered to have an unsuccessful outcome.

176 Discussion

177 The differential diagnosis for cats presenting with an acute or peracute onset of paresis or plegia includes aortic thrombo-embolism, ischemic myelopathy, fibrocartilaginous embolism, 178 intervertebral disk disease, and vertebral fractures and luxations^{8,9}. It has previously been 179 reported that trauma accounts for 14% of cases of feline spinal cord injury,¹⁰ and the occurrence 180 of a vertebral fracture or luxation is generally considered the most important differential 181 diagnosis for cats presenting with a peracute onset of spinal cord dysfunction after a witnessed 182 or suspected traumatic event. Of the cats included in this study nearly three-quarters of the cats 183 had experienced a witnessed traumatic event or there was evidence of trauma based on their 184 185 clinical exam or imaging findings. This highlights the need to include ANNPE as a possible

differential diagnosis for any cat presenting with an acute or peracute onset of spinal corddysfunction, particularly if there is any history or evidence of trauma.

When considering the location of the ANNPE the most frequent sites were the L3-L4 and L5-6 intervertebral discs. There was also one patient with a cervical ANNPE. This is consistent with the previous case reports that describe a lumbar and cervical ANNPE^{4,5}. Whilst this contrasts to the findings in dogs, which predominantly have T12-T13 and T13/L1 ANNPE², it is more consistent with data looking at the location of IVDE, with previous studies suggesting that the mid to caudal lumbar region is more commonly affected in cats¹¹⁻¹³.

When considering the outcome for patients diagnosed with ANNPE it is overall very good with 194 almost 90% of the cats being ambulatory with full urinary and faecal continence. None of the 195 cats were described as returning to 'normal' following the onset of clinical signs, and this is 196 197 consistent with one of the previous case reports that suggested there was ataxia present six months following diagnosis⁴. However, 50% of cats had returned to former behaviours 198 including outside activity and climbing on to furniture. It is currently unclear for those cats that 199 were no longer allowed outside, if this reflected a concern on part of the owners or an actual 200 inability to perform activities as before the onset of clinical signs. From the results of this 201 study, it is difficult to draw any conclusions on potential prognostic indicators for cats with a 202 presumptive ANNPE. 203

The incidence of ANNPE in cats is not known, although it appears to be infrequent, however it is possible that this reflects a decreased awareness of the condition and therefore an underdiagnosis. The treatment involved in caring for cats following the diagnosis of ANNPE is primarily supportive, involving the use of analgesics as appropriate, bladder management where required and intensive physiotherapy. The cost of treatment compared to cats diagnosed with vertebral fracture/luxation or IVDD is often reduced owing to the fact that there is no need for surgery. In addition it is often possible for caregivers to be trained to provide physiotherapy and bladder management at home. This combined with the evidence presented in this study that suggests cats with ANNPE appear to have a favourable prognosis highlights the need for ANNPE to be considered as an important differential diagnosis in cats with a peracute onset of spinal cord dysfunction.

215 Conclusions

This study is obviously limited by its retrospective nature and the small number of included 216 cases. However, the majority of cats diagnosed with presumptive ANNPE presented with 217 paraparesis or paraplegia and had neuroanatomical localisation of T3-L3 and L4-S3 spinal 218 cord segments. Nearly 75% of the cats were involved in a witnessed trauma or had evidence of 219 trauma based on clinical examination or imaging findings. The majority of cats diagnosed with 220 221 ANNPE had good outcomes. ANNPE should be considered as a differential diagnosis for cats presenting with peracute onset of spinal cord dysfunction, particularly if there is a clinical 222 history or evidence of trauma. 223

224

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228

229 Conflict of Interest

230 The authors do not have any potential conflicts of interest to declare.

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263 **Table 1**

Cat	Age	Sex	Breed	Neurological Examination Findings	Deep nociception present	Neuro- localisation	Site of ANNPE	Time to Ambulation (days)	Duration of hospitalisation (days)	Follow-up time and outcome
1	12 y 5 mo	MN	DSH	Paraplegic	Yes	T3-L3	L3-L4	15	11	4 mo Now indoor cat
2	2 y 9 mo	MN	Egyptian Mau	Non- ambulatory paraparetic; right pelvic limb plegic	Absent right pelvic limb	L4-S3	L5-L6	9	23	11 mo Urinary incontinence and occasional faecal incontinence
3	10 y 6 mo	FN	DLH	Non- ambulatory paraparetic	Yes	T3-L3	L3-L4	n/a	n/a	Euthanised due to pulmonary contusions
4	6 у	MN	DSH	Non- ambulatory paraparetic	Yes	T3-L3	T12-T13	16	20	20 mo Now indoor cat
5	4 y	MN	BSH	Non- ambulatory tetraparetic	Yes	C1-C5	C3-C4	18	9	LTF
6	4 y	FN	DSH	Ambulatory paraparetic	Yes	T3-L3	L4-L5	n/a	3	41 mo Only allowed outside in daylight hours
7	13 Y	MN	DLH	Non- ambulatory paraparetic; right pelvic limb plegic	Absent right pelvic limb	T3-L3	L3-L4	21	7	46 mo Returned to previous lifestyle; now deceased due to

Signalment, clinical presentation and outcome of 11 cats diagnosed with presumptive ANNPE

										unrelated
										causes
8	5 y	FN	DSH	Ambulatory paraparetic	Yes	L4-S3	L5-L6	n/a	3	67 mo Now indoor cat
9	8 y	MN	DSH	Non- ambulatory paraparetic	Yes	L4-S3	L5-L6	6	7	68m Returned to previous lifestyle;
10	7 y	MN	DSH	Paraplegic	Yes	T3-L3	L1-L2	18	13	LTF
11	8 y 6 mo	MN	DLH	Paraplegic	Absent bilaterally in pelvic limbs	T3-L3	T13-L1	21	26	46 mo Returned to previous lifestyle; now deceased due to unrelated causes

265 ANNPE = acute non-compressive nucleus pulposus extrusion; y = years; mo = months; MN =

266 male neutered; FN = female neutered; DSH = domestic shorthair; DLH = domestic longhair;

267 BSH = British Shorthair; LTF = lost to follow up

269 Figure 1. (a) Sagittal T2-weighted and (b) transverse T2-weighted images at the level of the L5–L6 intervertebral disc space, and (c) L4 vertebral body of an Egyptian Mau aged 2 years 270 and 9 months (cat 2). (a) A focal intraparenchymal hyperintensity is present at the level of the 271 L5–L6 intervertebral disc space (long arrow). Although the nucleus pulposus has a reduced 272 volume compared with the adjacent discs, it has remained a homogeneous hyperintense signal. 273 274 (b) A small amount of extraneous material present in the epidural space (arrow). (a,c) A poorly demarcated hyperintensity within the epaxial musculature at the level of the L4 vertebral body, 275 suggestive of epaxial muscle contusion, oedema or haemorrhage, was considered indicative for 276 external trauma (short arrow [a] and arrow [c]) 277

