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1	Evaluation of Magnetic Resonance Imaging Guidelines for Differentiation
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4	
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34 Abstract

35 Four MRI variables have recently been suggested to be independently associated with 36 a diagnosis of thoracolumbar intervertebral disk extrusion or protrusion. Midline 37 intervertebral disk herniation, and partial intervertebral disk degeneration were 38 associated with intervertebral disk protrusion, while presence of a single intervertebral 39 disk herniation and disk material dispersed beyond the boundaries of the 40 intervertebral disk space were associated with intervertebral disk extrusion. The aim 41 of this retrospective, cross sectional study was to evaluate if using these MRI 42 variables improves differentiation between thoracolumbar intervertebral disk 43 extrusions and protrusions. Eighty large breed dogs with surgically confirmed 44 thoracolumbar intervertebral disk extrusions or protrusions were included. 45 Randomized MRI studies were presented on two occasions to six blinded observers, 46 which were divided into three experience categories. During the first assessment, 47 observers made a presumptive diagnosis of thoracolumbar intervertebral disk 48 extrusion or protrusion without guidelines. During the second assessment they were 49 asked to make a presumptive diagnosis with the aid of guidelines. Agreement was 50 evaluated by Kappa-statistics. Diagnostic accuracy significantly improved from 51 70.8% to 79.6% and inter-observer agreement for making a diagnosis of intervertebral 52 disk extrusion or intervertebral disk protrusion improved from fair ($\kappa = 0.27$) to 53 moderate ($\kappa = 0.41$) after using the proposed guidelines. Diagnostic accuracy was 54 significantly influenced by degree of observer experience. Intra-observer agreement 55 for the assessed variables ranged from fair to excellent and inter-observer agreement 56 ranged from fair to moderate. The results of this study suggest that the proposed 57 imaging guidelines can aid in differentiating thoracolumbar intervertebral disk 58 extrusions from protrusions.

59

60 Two types of degenerative thoracolumbar intervertebral disk disease have been 61 recognized; intervertebral disk extrusion or Hansen Type-I, and intervertebral disk protrusion or Hansen Type-II intervertebral disk disease.¹⁻³ Both types of 62 63 intervertebral disk disease are associated with different pathological and clinical characteristics.^{4,5} Thoracolumbar intervertebral disk extrusion is characterized by a 64 65 sudden herniation of degenerated and calcified nucleus pulposus through a fully ruptured anulus fibrosus into the vertebral canal.^{1,5,6} Affected dogs can be young and 66 67 present often with an acute onset of clinical signs, which can vary from spinal hyperesthesia to paraplegia with loss of nociception.^{4,5} Intervertebral disk protrusion 68 69 is characterized by slowly progressive and focal extension of the anulus fibrosus and dorsal longitudinal ligament into the vertebral canal.^{1,5-8} Affected animals are 70 typically older, present with a more insidious clinical history and milder clinical 71 signs, such as ambulatory paresis and ataxia of the pelvic limbs.^{4,5,9,10} Apart from the 72 73 above mentioned differences in clinical presentation, thoracolumbar intervertebral 74 disk extrusion and protrusion are also associated with different suggested surgical 75 techniques ¹¹⁻¹⁵, postoperative complications, and possibly also a different prognosis.⁴ 76 Although outcomes after medical and surgical treatment have been extensively 77 reported for dogs with extrusions, this information is only scarce for dogs with 78 thoracolumbar intervertebral disk protrusions. It seems therefore important to 79 accurately differentiate extrusions from protrusions before treatment options and 80 associated outcomes are discussed with owners of affected dogs. Although magnetic 81 resonance imaging (MRI) is considered the imaging modality of choice to diagnose intervertebral disk disease in dogs ¹⁶, it is currently unknown how well this technique 82 83 can be used to differentiate between both types of intervertebral disk herniation. A recent study ⁹ identified four MRI variables, which were suggested to be 84

85 independently associated with a diagnosis of thoracolumbar intervertebral disk 86 extrusion or protrusion in large breed dogs (Figure 1): (1) midline instead of 87 lateralized intervertebral disk herniation, and (2) partial instead of complete 88 intervertebral disk degeneration were associated with a diagnosis of intervertebral 89 disk protrusion, while (3) the presence of a single instead of multiple intervertebral 90 disk herniations and (4) dispersed intervertebral disk material beyond the borders of 91 the intervertebral disk space were associated with a diagnosis of intervertebral disk 92 extrusion.⁹ Before these MRI variables can be suggested as diagnostic guidelines, it is 93 however important to evaluate if applying them would be clinically useful and 94 reliable. In other words, it should be assessed how well thoracolumbar intervertebral 95 disk extrusion can be differentiated from intervertebral disk protrusion without using 96 these guidelines, if using these guidelines improves indeed accuracy and reliability to 97 differentiate both types of intervertebral disk herniation, and if assessment of the 98 proposed MRI variables is associated with acceptable intra -and inter-observer 99 agreements. The aim of this retrospective, cross sectional study was therefore to 100 evaluate if implementation of the above mentioned MRI variables would facilitate 101 obtaining an accurate and reliable diagnosis of thoracolumbar intervertebral disk 102 extrusion or protrusion. An additional aim was to evaluate how levels of experience 103 would influence MRI assessments with and without the proposed imaging guidelines. 104 It was hypothesized that application of the proposed MRI guidelines would improve 105 diagnostic accuracy and reliability of diagnosing thoracolumbar intervertebral disk 106 extrusions and protrusions, that the proposed variables could be assessed with 107 acceptable intra and interobserver agreements, and that observers with limited MRI 108 experience would benefit the strongest from using the proposed MRI guidelines.

109

110 Methods

111 Included dogs were presented to the Royal Veterinary College (RVC), 112 University of London, between July 2002 and January 2014 for further assessment of 113 a suspected spinal condition. The studied animals were also included in a previous 114 study, evaluating MRI characteristics of 95 large breed dogs with surgically 115 confirmed thoracolumbar intervertebral disk extrusion (n=52) or intervertebral disk 116 protrusion (n=43).⁹ Of these 95 animals, 40 dogs with thoracolumbar intervertebral 117 disk extrusion and 40 dogs with intervertebral disk protrusion were randomly selected 118 using a random number generator. This selection aimed to obtain an equal number of 119 dogs in each group. To assess intraobserver agreement, the MRI studies of 20 dogs of 120 each group were randomly selected and duplicated. A total of 80 dogs and 120 MRI 121 studies were therefore included (Figure 2). All dogs included in this study received 122 general physical and complete neurological examinations. Dogs were included if (1) they were large breed dogs, defined as a body weight exceeding 20kg^4 , (2) underwent 123 124 an MRI study of the thoracolumbar or lumbar vertebral column, (3) following a 125 diagnosis of intervertebral disk herniation underwent spinal surgery consisting of a 126 hemilaminectomy or hemilaminectomy combined with a partial discectomy and (4) 127 the type of intervertebral disk herniation (extrusion or protrusion) was clearly noted in 128 the surgical reports. Dogs were excluded if the medical records or imaging studies 129 were incomplete, if they were not available in a digital format, or if the type of 130 intervertebral disk herniation (extrusion or protrusion) was not clearly noted in the 131 surgical reports, if more than one type of intervertebral disk herniation (both extrusion 132 and protrusion present), or acute herniations of flaps of anulus were observed during 133 surgery. Information retrieved from the medical records included signalment, 134 duration, type, and severity of clinical signs, general physical and neurological

examinations findings and type of surgery with surgical confirmation of the type of
intervertebral disk herniation (extrusion or protrusion). Severity of neurological
deficits was graded by the modified Frankel score, which was defined as paraplegia
with no deep nociception (grade 0), paraplegia with no superficial nociception (grade
paraplegia with nociception (grade 2), non-ambulatory paraparesis (grade 3),
ambulatory paraparesis and ataxia (grade 4), spinal hyperesthesia only (grade 5), or
no dysfunction.¹⁷

142

143 All dogs underwent MRI under general anesthesia. MRI was performed with a 144 1.5T magnet (Intera, Philips Medical Systems, Eindhoven, the Netherlands). The 145 selection of variables was based on the results of a previous study evaluating MRI 146 characteristics of thoracolumbar intervertebral disk disease in large breed dogs and 147 were suggested to be independently associated with a diagnosis of intervertebral disk extrusion or intervertebral disk protrusion.⁹ Assessed variables included (1) 148 149 lateralization of herniated disk material, (2) degree of intervertebral disk 150 degeneration, (3) the presence of multiple intervertebral disk herniations, and (4) 151 location of herniated disk material relative to the intervertebral disk space (Figure 1). 152 Lateralization of herniated intervertebral disk material was assessed on transverse 153 images and was described as being exclusively in the midline ventral to the spinal 154 cord or lateralized. Assessment of intervertebral disk degeneration was based on 155 nucleus pulposus signal intensity on sagittal T2-weighted images. A non-degenerate 156 intervertebral disk (grade 0) had a homogenous hyperintense signal, a partially 157 degenerate intervertebral disk (grade 1) had heterogeneous loss of hyperintense 158 signal, and a completely degenerate intervertebral disk (grade 2) had complete loss of hyperintense signal.¹⁸⁻²⁰ Presence of a single or multiple intervertebral disk 159

160 herniations was evaluated on sagittal and transverse T2-weighted images. Location of 161 herniated disk material relative to the affected intervertebral disk space was assessed 162 on T1-and T2-weighted sagittal images and was described as dispersed or confined 163 to the intervertebral disk space. Dispersed intervertebral disk material was defined as 164 intervertebral disk material beyond the borders of the affected intervertebral disk space and associated vertebral endplates.²¹ Disk material confined to the intervertebral 165 166 disk space was defined as herniated disk material not exceeding the limits of the intervertebral disk space or associated vertebral endplates.²¹ 167

168 The 120 MRI studies were presented twice in a randomized order to six 169 blinded observers (Figure 2). The sequence of MRI studies and the identity of 170 duplicate studies differed among observers. The observers were not informed about 171 the clinical history, type of clinical signs, the number of dogs with thoracolumbar 172 intervertebral disk extrusions or protrusions, were not informed about the inclusion of duplicate MRI studies, were not aware of the results of our previous study⁹, and were 173 174 also not involved in assessment of MRI studies for our previous study⁹. Standard image archiving and communication system software (Osirix Foundation, V.5.5.2 175 176 Geneva, Switzerland) was used to view the imaging studies. During the first round of 177 assessments, the observers were asked for each MRI study to make a presumptive 178 diagnosis of thoracolumbar intervertebral disk extrusion or protrusion without any 179 guidelines. They were however informed about the location of the affected and 180 surgically confirmed intervertebral disk space. After returning their answers, they 181 received again 120 MRI studies (different sequence and different identity of 182 duplicates) with at least a 1-month interval between the two rounds of assessments. 183 During the second round of assessments the observers were again asked for each MRI 184 study to make a presumptive diagnosis of intervertebral disk extrusion or protrusion.

185 This time they were however provided with the proposed MRI guidelines. 186 Additionally, they were asked to record the presence or absence of each of the 187 proposed MRI variables (Figure 2). They were again informed about the location of 188 the affected intervertebral disk space. To evaluate the role of experience, the six 189 observers were divided into three groups of different experience levels; expert, 190 moderate, and no experience. Two board-certified neurologists represented the group 191 of expert observers (PJK and EB). Two final year residents in neurology represented 192 the group of observers with moderate experience (BP and JF). The group of observers 193 without experience was represented by two veterinary surgeons completing a small 194 animal rotating internship (DN and GN). Because of their lack of experience and 195 unfamiliarity with the imaging software, the two observers without experience 196 received a training session by the first author of this study. During this session, they 197 learnt how to use the imaging viewing software, were instructed about the 198 pathophysiological differences between intervertebral disk extrusions and protrusions, 199 and were handed a recent review manuscript discussing intervertebral disk disease in 200 chondrodystrophic and non-chondrodystrophic dog breeds.⁵ After the first round of 201 assessments they received illustrated guidelines with examples of the proposed MRI 202 variables. While these training sessions were deemed necessary to facilitate study 203 enrollment of observers without experience, it was considered unnecessary for the 204 more experienced observers. 205 Wilcoxon signed ranks tests were used to detect the influence of diagnostic

205 whicoxon signed ranks tests were used to detect the influence of diagnostic 206 guidelines on diagnostic accuracy and agreement. Chi-squared tests were used to 207 detect differences in accuracy between the three experience groups, and the effect of 208 intervertebral disk herniation type (intervertebral disk extrusion vs. protrusion) on 209 diagnostic accuracy with and without use of guidelines. These statistics were carried

210	out in IBM SPSS Statistics v21 (IBM SPSS Statistics, New York) by one of the
211	authors (RMAP). Agreement statistics were calculated using Minitab v17 (Minitab
212	version 17, Pennsylvania). Fleiss' kappa (κ) for more than two observers was
213	calculated ²² for diagnosis and each assessed MRI variable. The strength of agreement
214	was interpreted on the basis of the κ values suggested by Altman $^{23},$ as adapted from
215	the method of Landis and Koch 24 : κ -values of $0.81 - 1.00$ indicated very good
216	agreement; 0.61 – 0.80, good agreement; 0.41 – 0.60, moderate agreement; 0.21 –
217	0.40, fair agreement; and 0.20 or lower, poor agreement. As a guide, the minimum
218	threshold for κ is often arbitrarily set at $\kappa\!\leq\!0.4,$ below which variables are considered
219	unreliable. ²⁵ Calculation of interobserver agreement and accuracy was based on the
220	assessment of the 80 original MRI studies. Calculation of intraobserver agreement
221	was based on assessment of 40 duplicate MRI studies (Figure 2). Results were
222	considered significant if $P < 0.05$.

223

210

224 **Results**

225 A total of 80 large breed dogs with surgically confirmed thoracolumbar 226 intervertebral disk extrusion (n=40) or protrusion (40) were included in this study. 227 The group of dogs with intervertebral disk extrusion included German Shepherd Dogs 228 (n=10), Staffordshire Bull Terriers (five), Clumber Spaniels (four), Labrador 229 Retrievers (four), Basset Hounds (three), Rottweilers (three); there were six breeds 230 represented by one dog each and five crossbreeds. This group included 21 males and 231 19 females aged between three and 12 years (mean, 7.2 years). Duration of clinical 232 signs ranged from 12 hours to six months (median, three days) and affected dogs 233 presented with neurological grades 0 (n=three dogs), 1 (one), 2 (10), 3 (nine), and 4 234 (17). Affected intervertebral disk spaces in order of occurrence were T12-T13, L1-L2

235 (both n=nine), T13-L1 (seven), L2-L3 (six), L3-L4 (five), T11-T12 (two), T3-T4 and

236 L4-L5 (both one). The group of dogs with thoracolumbar intervertebral disk

237 protrusion included German Shepherd Dogs (n=20), English Staffordshire Bull

238 terriers (eight), Basset Hounds (two), Labrador Retrievers (two); there were four

breeds represented by one dog each and four crossbreeds. This group included 32

240 males and eight females aged between four and 12.2 years (mean, 8.6 years). Duration

of clinical signs ranged from 12 hours to two years (median, 29 days) and affected

242 dogs presented with neurological grades 2 (n=one dog), 3 (seven), and 4 (32).

243 Affected intervertebral disk spaces in order of occurrence were T13-L1 (n=17), T12-

244 T13, L1-L2 (both nine) and L2-L3 (five).

245 MRI studies were performed with dogs in dorsal recumbency, using a 246 dedicated spinal coil. Imaging studies included a minimum of T2-weighted (repetition 247 time (ms) (TR)/ echo time (ms) (TE); 3000/120) and T1-weighted (TR/TE, 400/8) 248 sagittal and transverse images. Slice thickness for sagittal and transverse images were 249 respectively 1.75 and 2.5mm with an interslice gap of 0.3mm in both planes. The 250 transverse images were aligned perpendicular to the spinal cord. Diagnostic accuracy 251 was calculated using the assessments of the 80 original MRI studies (Table 1). 252 Overall diagnostic accuracy without using the proposed guidelines for the six 253 observers combined was 70.8% (340/480) and ranged from 45% (36/80) to 88.8% 254 (71/80) for individual observers (Table 1). There was a significant influence of 255 observer experience with observers without experience being least accurate (58%) and observers with moderate experience being most accurate (78%) ($X^2 = 18.78$, P <256 257 0.001). There was no significant difference between the number of accurately 258 diagnosed thoracolumbar intervertebral disk extrusions or protrusions before 259 guidelines were included (74% vs. 75%, respectively, P > 0.05). Overall diagnostic

260 accuracy for the six observers combined improved significantly (P = 0.028) from 261 70.8% to 79.6% (382/480) after using the proposed diagnostic guidelines and ranged 262 from 52.5% (42/80) to 92.5% (74/80) after using the MRI guidelines. Additionally, all 263 six observers diagnosed a higher number of thoracolumbar intervertebral disk 264 extrusions or protrusions correctly after using the guidelines (mean improvement; 13.4%). There was again a significant influence of observer experience ($X^2 = 31.72$, P 265 266 < 0.001) with observers without experience being least accurate (65%) and observers 267 with expert experience being most accurate (88%). After using the guidelines 268 thoracolumbar intervertebral disk protrusions (84%) were significantly more often (P = 0.024) correctly diagnosed than intervertebral disk extrusions (75%) ($X^2 = 5.13$, P =269 270 0.02)

271

272 Intraobserver agreement was calculated using the assessments of the 40 273 duplicate MRI studies (Table 2). Intraobserver agreement for making a diagnosis of 274 thoracolumbar intervertebral disk extrusion or protrusion ranged from 0.4 to 0.95 275 before, and from 0.51 to 0.85 after applying the proposed guidelines. This difference 276 was not significant and there was no significant influence of observer experience. 277 Although a high degree of variability was seen among individual observers, most 278 variables were associated with moderate or good intraobserver agreement (Table 2). 279 Assessing if multiple intervertebral disk herniations were present was associated with 280 the highest intraobserver agreement ($\kappa = 0.43 - 0.79$) and assessing if an 281 intervertebral disk herniation was midline or lateralized was associated with the 282 lowest intraobserver agreement ($\kappa = 0.33 - 0.75$). There was no significant influence 283 of level of experience on intraobserver agreement of the assessed MRI variables. 284

285	Interobserver agreement was calculated using the assessments of the 80
286	original MRI studies. Overall interobserver agreement for making a diagnosis of
287	thoracolumbar intervertebral disk extrusion or protrusion significantly improved ($P <$
288	0.001) from fair (κ =0.27) to moderate (κ = 0.41) after using the proposed guidelines.
289	Assessing if an intervertebral disk herniation was midline or lateralized and if and
290	intervertebral disk herniation was confined to or exceeded the borders of the affected
291	intervertebral disk space were associated with moderate interobserver agreement
292	(both $\kappa = 0.43$). Assessing the presence of partial intervertebral disk degeneration ($\kappa =$
293	0.35), complete disk degeneration ($\kappa = 0.40$), and multiple intervertebral disk
294	herniations ($\kappa = 0.35$) were associated with fair interobserver agreement.
295	

296 **Discussion**

297 This study evaluated the accuracy, reliability and reproducibility of MRI guidelines to 298 improve the differentiation of thoracolumbar intervertebral disk extrusion and disk 299 protrusion in large breed dogs. The results of this study suggest that applying the 300 proposed guidelines improves the accuracy and inter-observer agreement of 301 diagnosing thoracolumbar intervertebral disk extrusion and intervertebral disk 302 protrusion. The assessed MRI variables were associated with fair to excellent 303 intraobserver and fair to moderate interobserver agreements. The results of this study 304 indicate further that accuracy of assessing spinal MRI studies is influenced by the 305 observer's degree of training and experience. 306 307 Differentiating thoracolumbar intervertebral disk extrusions from protrusions is

308 clinically important. Although several studies have characterized the clinical

309 presentation and outcomes after medical or surgical treatment for dogs with

thoracolumbar intervertebral disk extrusion ²⁶⁻³⁰, this information is less well 310 311 characterized for dogs with intervertebral disk protrusion. While surgical treatment of 312 thoracolumbar intervertebral disk extrusion typically consists of a hemilaminectomy with fenestration of the affected intervertebral disk $^{28-30}$, it is suggested that a lateral 313 314 corpectomy or vertebral stabilization should be considered for dogs with intervertebral disk protrusion.¹¹⁻¹⁵ While the former can be considered a basic spinal 315 316 surgical technique, the latter are probably more technically demanding. Furthermore, 317 little is known about results of medical management in dogs with thoracolumbar intervertebral disk protrsuins⁴, it has been suggested that dogs with thoracolumbar 318 319 intervertebral disk extrusion have a better prognosis after surgery than dogs with intervertebral disk protrusion ⁴, and that dogs with thoracolumbar intervertebral disk 320 protrusion are at increased risk of early postoperative neurological deterioration.^{4,31} 321 322 This illustrates that reaching an accurate and reliable imaging diagnosis of 323 thoracolumbar intervertebral disk extrusion or protrusion is not only important for 324 surgical planning, but also for managing the expectations of owners and clinical staff. 325 The overall accuracy to differentiate thoracolumbar intervertebral disk extrusion from 326 intervertebral disk protrusion improved significantly from 70.8 to 79.6%, and the 327 interobserver agreement for making a diagnosis improved significantly from fair to 328 moderate after applying the proposed MRI guidelines. Although these findings 329 indicate clinical usefulness, a proportion of cases were still diagnosed incorrectly after 330 using the proposed guidelines. This suggests that MRI in isolation cannot be completely relied on to differentiate between thoracolumbar extrusions and 331 332 protrusions and confirms that imaging findings should always be interpreted in light of a thorough clinical history and results of a complete clinical examination.³² In 333 334 combination to the evaluated MRI variables, a clinical variable has been suggested to

be independently associated with a diagnosis of thoracolumbar intervertebral disk
extrusion or protrusion. Increased duration of clinical signs, with a threshold value of
21 days, was significantly associated with a diagnosis of intervertebral disk
protrusion.⁹ The observers in this study were however unaware of the clinical history
and clinical signs of included dogs. It is therefore possible that better diagnostic
accuracy and reliability would have been reached if this information had been
available.

342

343 Intra -and interobserver agreement for the assessed MRI variables was evaluated with 344 kappa statistics. This is a useful and widespread statistical technique to evaluate reliability and reproducibility.^{25,33} Obtained kappa coefficients are commonly 345 translated into poor, fair, moderate, good or very good agreement.^{23,24} Although this 346 347 can provide useful information, results of kappa statistics do not directly determine if a certain variable can be assessed with acceptable reliability and reproducibility.^{25,33} 348 349 Although such a decision is dependent on the specific variable and clinical context, 350 studies in musculoskeletal research have suggested that kappa coefficients lower than 0.4 should be considered clinically unacceptable.^{25,33} When applying this threshold 351 352 value to the study presented here, intraobserver agreement for the assessed MRI 353 variables could be considered clinically acceptable. Only one observer did not reach 354 intraobserver agreement coefficients ≥ 0.4 for all assessed MRI variables. 355 Interobserver agreement for the assessed variables ranged from fair to moderate with 356 3 of the 5 assessed variables reaching the arbitrary threshold value (≥ 0.4) for 357 acceptable agreement. Presence of multiple intervertebral disk herniations and degree 358 of intervertebral disk degeneration were associated with only fair interobserver 359 agreement. The lower interobserver agreement for the presence of multiple

360 intervertebral disk herniations is in agreement with a previous study evaluating the 361 number of spinal cord compressions in dogs with disk-associated cervical spondylomyelopathy.³⁴ In the current study, assessment of multiple intervertebral disk 362 363 herniations was among the variables with the highest intraobserver agreement, 364 indicating consistent rating within an individual observer. Different degrees of intervertebral disk herniation can occur in clinically normal dogs ^{20,35} and it is not 365 366 always possible to clearly differentiate clinically relevant from irrelevant intervertebral disk herniations.³² It is therefore possible that different observers will 367 368 have different, but individually consistent, criteria for evaluating if an intervertebral 369 disk herniation is present or not. This variable is therefore likely associated with 370 inherent subjectivity and it should be considered to develop a more objectively 371 definition of intervertebral disk herniation. Although a grading system based on accepted terminology ^{19,20,36} with easily recognizable characteristics was used, 372 373 evaluation of degree of intervertebral disk degeneration was also associated with 374 lower values for interobserver agreement. This finding is however in contrast with several human and veterinary studies.^{19,32,37-40} A previous veterinary study 375 376 demonstrated good agreement between experienced observers for rating intervertebral disk degeneration using the same criteria as this study.³² Although other reasons 377 378 cannot be excluded, it is possible that the inclusion of observers with varying degrees 379 of training and experience, including observers without experience, contributed to the lower value for interobserver agreement in this study.^{39,41} 380 381 382 The influence of observer experience was further highlighted by the fact that 'low

383 observer experience' negatively influenced the diagnostic accuracy of the assessed

384 MRI studies. Even after using the diagnostic guidelines, the group of observers

385 without experience had significantly lower accuracy for diagnosing thoracolumbar 386 intervertebral disk extrusion and intervertebral disk protrusion. This finding is in agreement with several human studies ^{39,42,43} and indicates that assessment and 387 388 clinical interpretation of MRI studies should be preserved for people with sufficient 389 experience and training. It is therefore possible that the overall results of this study are 390 influenced by the inclusion of observers with different experience levels and it can be 391 debated if our results therefore appropriately reflect referral practice. For the purpose 392 of this study, observers with different experience categories were included to evaluate 393 if differentiation between thoracolumbar intervertebral disk extrusion and disk 394 protrusion would be more problematic for observers with little experience, if using the 395 proposed guidelines would be easy and objective enough to be used by novices, and if 396 using these imaging guidelines could compensate for lack of training and experience. 397

398 A potential limitation of this study was the limited number of observers in each 399 'experience category'. Although care was taken to select observers of similar 400 experience levels. In addition, there was no standardization across observers in terms 401 of how the assessments were performed (i.e. all in one sitting or a few at a time) or 402 the monitor on which they were viewed. Another limitation was that all patients had 403 surgically confirmed intervertebral disk extrusion or protrusion. The combination of 404 imaging and clinical findings was therefore severe enough to warrant surgical 405 intervention. These factors could have influenced the assessment of the observers; 406 however surgical confirmation was necessary to provide a "golden standard" 407 diagnosis with which to assess diagnostic accuracy. It should further be emphasized 408 that MRI studies were evaluated by general clinicians or veterinary surgeons trained 409 in veterinary neurology and neurosurgery. It can however not be excluded that

410 interpretation of MRI studies by observers with a different training background, such 411 as veterinary radiologists, would have influenced our results. Veterinary neurologists 412 were however preferred in this study, because in our institution, these clinicians 413 perform MRI interpretation and spinal surgery on a daily basis and were therefore 414 considered to have most experience in taking clinical decisions after assessing spinal 415 MRI studies. Although only limited data is available on direct comparisons between 416 veterinary neurosurgeons and radiologists for assessing spinal MRI studies ^{34,44,45}, 417 human studies indicate acceptable agreement between specialities.⁴⁶⁻⁴⁹ Although 418 comparing the accuracy and agreement between veterinary neurologists and 419 radiologists was considered beyond the aims of this study, further studies are 420 necessary to not only evaluate the role of experience, but also the type of training on 421 the reliability and accuracy of evaluating spinal MRI studies 422 423 In summary, the results of this study suggest that differentiation between 424 thoracolumbar intervertebral disk extrusion and protrusion becomes more accurate 425 and reliable after applying the proposed MRI guidelines, that the proposed MRI 426 variables can be assessed with, in general, clinically acceptable agreement, and that 427 diagnostic guidelines cannot replace thorough clinical training and experience. 428 Further studies are necessary to evaluate the influence of observer experience and 429 availability of clinical history and clinical signs on the evaluation of spinal MRI 430 studies. 431 432 433 Acknowledgments: None

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- 590 Legends
- **Figure 1**. Proposed MRI guidelines to differentiate between thoracolumbar
- 592 intervertebral disk extrusion (IVDE) and intervertebral disk protrusion (IVDP) in
- 593 large breed dogs.

MRI variable	Intervertebral disk Extrusion	Intervertebral disk Protrusion
Lateralized intervertebral disk herniation	More likely	Less likely
Partial instead of complete intervertebral disk degeneration	Less likely	More likely
Multiple intervertebral disk herniations	Less likely	More likely
Herniated disk material confined to intervertebral disk space	Less likely	More likely

- 597 Figure 2. Flow-chart illustrating different MRI assessments in order to calculate the
- 598 diagnostic accuracy, inter-observer agreement and intra-observer agreement for
- 599 differentiating between thoracolumbar intervertebral disk extrusions and protrusions
- 600 with and without the use of the proposed imaging guidelines.



- **Table 1.** Number of correctly diagnosed thoracolumbar intervertebral disk extrusionsor protrusions before and after using the proposed MR imaging guidelines.Calculations were based on 80 original MRI studies

Observer	Degree of experience	Correct diagnoses without guidelines (out of 80)	Correct diagnoses with guidelines (out of 80)
Observer 1	Expert	69 (86.2%)	73 (91.2%)
Observer 2	Moderate	53 (66.2%)	63 (78.8%)
Observer 3	None	36 (45%)	42 (52.5%)
Observer 4	Expert	54 (67.5%)	68 (85%)
Observer 5	Moderate	71 (88.8%)	74 (92.5%)
Observer 6	None	57 (71.2%)	62 (77.5%)
Overall	NA	340 (70.8%)	382 (79.6%)

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607 Table 2. Intraobserver agreement κ-values for making a diagnosis of thoracolumbar
 608 intervertebral disk extrusion or protrusion and for assessment of each of the proposed
 609 MRI variables. Calculations were based on 40 duplicate MRI studies

Variable	Observer 1- Expert	Observer 2- Moderate experience	Observer 3- No experience	Observer 4- Expert	Observer 5- Moderate experience	Observer 6 – No experience
Diagnosis without guidelines	0.75	0.81	0.42	0.80	0.95	0.40
Diagnosis with guidelines	0.75	0.85	0.51	0.59	0.70	0.75
IVD herniation midline?	0.75	0.75	0.33	0.69	0.55	0.45
Multiple IVD herniations present?	0.79	0.73	0.46	0.43	0.85	0.75
IVD herniation confined to IVDS?	0.95	0.89	-0.14	0.84	0.69	0.56
Partial IVD degeneration present?	0.68	0.85	0.55	0.65	0.49	0.47
Complete IVD degeneration present?	0.68	0.90	0.44	0.62	0.49	0.75

612 IVD = intervertebral disk, IVDS = intervertebral disk space on and for assessment of

613 each of the proposed MRI variables. Calculations were based on 80 original MRI

614 studies

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