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Ryan, R., Gutierrez-Quintana, R., ter Haar, G. and De Decker, S. 'Prevalence of thoracic vertebral malformations in french bulldogs, pugs and english bulldogs with and without associated neurological deficits', *The Veterinary Journal*.

The final version is available online: <http://dx.doi.org/10.1016/j.tvjl.2017.01.018>.

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The full details of the published version of the article are as follows:

TITLE: Prevalence of thoracic vertebral malformations in french bulldogs, pugs and english bulldogs with and without associated neurological deficits

AUTHORS: R. Ryan, R. Gutierrez-Quintana, G. ter Haar, Steven De Decker

JOURNAL: The Veterinary Journal

PUBLISHER: Elsevier

PUBLICATION DATE: 31 January 2017 (online)

DOI: 10.1016/j.tvjl.2017.01.018

1 **Original Article**

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4 **Prevalence of thoracic vertebral malformations in French bulldogs, Pugs and English**
5 **bulldogs with and without associated neurological deficits**

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8 R. Ryan ^a, R. Gutierrez-Quintana ^b, G. ter Haar ^a, Steven De Decker ^{a, *}

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10 ^a *Department of Veterinary Clinical Science and Services, Royal Veterinary College,*
11 *University of London, Hawkshead lane, AL9 7TA North Mymms, Hatfield, England*

12 ^b *School of Veterinary Medicine, College of Medical, Veterinary and Life Sciences,*
13 *University of Glasgow, Bearsden Road, Glasgow, G61 1QH, Scotland*

14

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16

17 * Corresponding author. Tel.: +44 1707 666366.

18 *E-mail address: sdedecker@rvc.ac.uk (S. De Decker)*

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24 **Highlights**

- 25 • Thoracic vertebral malformations are common in neurologically normal French
26 bulldogs, Pugs and English bulldogs.
- 27 • There is an influence of breed on the prevalence of different types of vertebral
28 malformations.
- 29 • Hemivertebrae occur more often in neurologically normal French bulldogs and
30 less often in Pugs.
- 31 • Transitional vertebrae and spina bifida occur more often in neurologically normal
32 Pugs.
- 33 • Hemivertebrae are more likely to be associated with neurological deficits in Pugs
34 than the other breeds.

36 **Abstract**

37 Congenital vertebral malformations are common incidental findings in small breed
38 dogs. This retrospective observational study evaluated the type and prevalence of thoracic
39 vertebral malformations in 171 neurologically normal and 10 neurologically abnormal screw-
40 tailed brachycephalic dogs. Neurologically normal dogs underwent CT for reasons unrelated
41 to spinal disease, while affected dogs underwent MRI. Imaging studies were reviewed and
42 vertebral malformations including hemivertebrae, block vertebrae, transitional vertebrae, and
43 spina bifida were documented.

44
45 The group of clinically normal dogs consisted of 62 French bulldogs, 68 Pugs and 41
46 English bulldogs. The group of affected dogs consisted of one French bulldog and nine Pugs.
47 Overall, 80.7% of neurologically normal animals were affected by at least one vertebral
48 malformation. There was a significant influence of breed, with thoracic vertebral
49 malformations occurring more often in neurologically normal French bulldogs ($P < 0.0001$)
50 and English bulldogs ($P = 0.002$). Compared to other breeds, hemivertebrae occurred more
51 often in neurologically normal French bulldogs (93.5%; $P < 0.0001$ vs. Pugs; $P = 0.004$ vs.
52 English bulldogs) and less often in neurologically normal Pugs (17.6%; $P = 0.004$ vs. English
53 bulldogs). Neurologically normal Pugs were more often diagnosed with transitional vertebrae

54 and spina bifida compared to other breeds ($P < 0.0001$ for both malformations). Of Pugs
55 included in the study, 4.7% were diagnosed with clinically relevant thoracic vertebral
56 malformations. When compared to the general veterinary hospital population, this was
57 significantly more than the other two breeds ($P = 0.006$). This study indicates that thoracic
58 vertebral malformations occur commonly in neurologically normal screw-tailed
59 brachycephalic dogs. While hemivertebrae are often interpreted as incidental diagnostic
60 findings, they appear to be of greater clinical importance in Pugs compared to other screw-
61 tailed brachycephalic breeds.

62

63 *Keywords:* Brachycephalic; Hemivertebra; Kyphosis; Spina bifida; Transitional vertebra

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65 **Introduction**

66 Congenital vertebral malformations are common incidental findings in small
67 brachycephalic dogs. Although the terminology used for these malformations is controversial,
68 they have been classified as defects in segmentation (block vertebrae), defects in formation
69 (wedge or hemivertebrae) and other defects, including transitional vertebrae and spina bifida
70 (Westworth and Sturges, 2010). Hemivertebrae are frequently reported in screw-tailed
71 brachycephalic breeds such as the French bulldog (Moissonnier et al., 2011; Aikawa et al.,
72 2014; Gutierrez-Quintana et al., 2014). Although the exact aetiology is unclear, they are
73 assumed to be hereditary (Schlensker and Distl, 2016). These vertebral malformations are
74 most frequently found in the thoracic vertebral column and can affect single or multiple
75 vertebrae (Faller et al., 2014; Guevar et al., 2014).

76
77 Despite the potential to cause clinical signs of spinal cord dysfunction, vertebral
78 malformations are frequently not associated with disease. Approximately 78% of
79 neurologically normal French bulldogs had radiographic evidence of hemivertebrae, with or
80 without spinal kyphosis (Moissonnier et al., 2011). This indicates that caution should be used
81 when evaluating imaging studies of French bulldogs with suspected spinal disease (Dewey et
82 al, 2016). More specifically, other spinal conditions, including intervertebral disc disease
83 (Aikawa et al., 2014) and spinal arachnoid diverticula (Mauler et al., 2014), should be
84 considered more likely causes of clinical signs in French bulldogs with thoracic vertebral
85 body malformations. Vertebral body malformations can result in alterations of vertebral
86 angulation, such as spinal kyphosis and scoliosis. Although the development of clinical signs
87 in dogs with congenital vertebral body malformations is thought to be multifactorial in
88 aetiology, it has been suggested that spinal kyphosis is a key factor in the development of
89 clinical signs and that kyphosis needs to reach a threshold point before clinical signs are

90 likely to occur (Moissonnier et al., 2014; Guevar et al., 2014). Although this situation is well
91 recognised for French bulldogs (Moissonnier et al., 2011), it is currently unknown if other
92 screw-tailed brachycephalic dogs demonstrate a similar prevalence of clinically irrelevant
93 thoracic vertebral malformations.

94

95 The primary aims of this study were to describe and compare the type and prevalence
96 of thoracic vertebral malformations in French bulldogs, Pugs and English bulldogs with and
97 without associated neurological deficits. It was hypothesised that although thoracic vertebral
98 malformations would occur commonly in each of the three evaluated breeds, they would only
99 rarely result in spinal cord dysfunction. Our secondary hypothesis was that the prevalence of
100 thoracic vertebral malformations would be breed-associated.

101

102 **Materials and methods**

103 The digital medical database of the Small Animal Referral Hospital, Royal Veterinary
104 College, was reviewed between October 2010 and February 2016 to identify two groups of
105 dogs. Group 1 included French bulldogs, Pugs and English bulldogs which underwent
106 thoracic CT under sedation or general anaesthesia for reasons unrelated to spinal disease.
107 Dogs were excluded if medical records or imaging studies were incomplete or unavailable for
108 review, or if the dog demonstrated a gait abnormality. A study was considered incomplete if
109 the complete thoracic vertebral column was not included. Group 2 included French bulldogs,
110 Pugs and English bulldogs with clinically relevant vertebral malformations diagnosed by
111 MRI under general anaesthesia. A vertebral malformation had to be the only identified cause
112 of spinal dysfunction in these dogs.

113

114 Information retrieved from the medical records included signalment, reason for
115 presentation, results of general physical examinations and, if available, neurological
116 examinations. CT was performed with a 16-slice helical CT scanner (PQ 500, GE
117 Healthcare), 2 mm slice thickness and -1 interval between slices After completion of the axial
118 CT study, sagittal, dorsal and 3D reconstructions were made. MRI was performed with a 1.5
119 Tesla magnet and included a minimum of T2 –and T1 – weighted sagittal and transverse
120 images. Slice thickness was 3.5 mm in all planes with an interslice gap of 0.9 mm in the
121 sagittal planes and 1 mm in the transverse planes. Imaging studies were independently
122 evaluated by two observers (RR and RGQ for CT studies; RR and SDD for MRI studies),
123 after which a consensus opinion was reached. The observers were not masked to the breed of
124 dog. For each imaging study, the number of thoracic vertebrae were recorded and each
125 thoracic vertebra was subsequently assessed for the presence of hemivertebra, block vertebra,
126 spina bifida and transitional vertebra (Fig. 1). Hemivertebrae were defined as any defect in
127 vertebral body formation as outlined by Gutierrez-Quintana et al (2014). Block vertebrae
128 were defined as failure of vertebral segmentation with absence of the intervertebral disc space
129 between two adjacent vertebral bodies (Westworth and Sturges, 2010). Spina bifida was
130 defined as incomplete closure of the vertebral arches resulting in a cleft through the dorsal
131 spinous process (Westworth and Sturges, 2010). Transitional vertebrae were defined as
132 thoracic vertebrae at the cervicothoracic or thoracolumbar junctions displaying characteristics
133 of cervical or lumbar vertebrae, respectively, including the absence or hypoplasia of a rib or
134 an abnormal transverse process (Westworth and Sturges, 2010). Standard image archiving
135 and communication system software (Osirix Foundation, V.5.5.2) was used to evaluate all
136 imaging studies.

137

138 *Statistical methods*

139 Data was analysed using commercial software (IBM SPSS Statistics version 22).
140 Association between breed and total number of malformations, and the total number of
141 hemivertebrae and block vertebrae, was evaluated with Kruskal-Wallis tests. Post-hoc
142 analysis with Mann-Whitney tests was used to determine the relative prevalence in each
143 breed. The presence of spina bifida and transitional vertebra and the prevalence of clinically
144 relevant vertebral malformations were determined using Fisher's exact tests; Pugs were
145 compared to the other breeds as a collective. Values of $P < 0.05$ were considered statistically
146 significant for all analyses; the significance level for multiple comparisons was adjusted for
147 using the Bonferroni method.

148

149 **Results**

150 *Dogs without neurological signs*

151 A total of 171 dogs, comprising 62 French bulldogs, 68 Pugs and 41 English bulldogs
152 were included in this group. All underwent CT for a variety of clinical indications, including
153 brachycephalic obstructive airway syndrome ($n=124$), other respiratory disease, neoplastic
154 disease ($n=17$ for both), cardiac disease ($n=5$), gastrointestinal disease, and trauma ($n=4$ for
155 both). CT imaging did not reveal any malformations in 33 dogs (19.3%), while single ($n=36$)
156 or multiple ($n=102$) malformations were observed in 138 dogs (80.7%).

157

158 The group of French bulldogs consisted of 52 males and 10 females between 2 and
159 135 months old (median, 19.5; mean, 29.9 months) and weighing between 6.7 and 16 kg
160 (median, 11.45; mean, 11.4kg). All French bulldogs had 13 thoracic vertebrae. Four (6.5%)
161 French bulldogs had no thoracic vertebral malformations, while 58 (93.5%) had one or more
162 hemivertebrae. A total of 243 hemivertebrae were present; nine (14.5%) dogs had single
163 hemivertebrae and 49 (79.0%) had multiple hemivertebrae. T9 was most often affected ($n=31$)

164 dogs), followed by T10 ($n=30$), T5 and T6 ($n=29$ for both). Block vertebrae were diagnosed
165 in four (6.5%) French bulldogs; T11-T12 ($n=4$) were most often affected, followed by T12-
166 T13 ($n=2$). Fourteen (22.6%) dogs had fused dorsal spinous processes. Transitional vertebrae
167 were present in three French bulldogs (4.8%). In these dogs, T13 had characteristics of a
168 lumbar vertebra. Four (6.5%) dogs had vertebrae with evidence of more than one type of
169 malformation. These consisted of block and hemivertebrae ($n=3$) and spina bifida and
170 hemivertebrae ($n=1$) combinations. No French bulldogs in this population had evidence of
171 spinal bifida alone.

172

173 The group of Pugs consisted of 31 males and 37 females between 4 and 151 months
174 old (median, 31.5; mean, 42.7 months) and weighing between 4 and 14kg (median, 8.15;
175 mean, 8.3kg). Seventeen (25.0%) Pugs had only 12 thoracic vertebrae. Eighteen (26.5%)
176 Pugs had no thoracic vertebral malformations. Twelve (17.6%) Pugs had hemivertebrae. A
177 total of 19 hemivertebrae were present; six (8.8%) Pugs had single hemivertebrae and six had
178 multiple hemivertebrae. The most commonly affected vertebra was T8 ($n=8$), followed by T7
179 ($n=5$) and T9 ($n=3$). No Pugs had block vertebrae or fused spinous processes. Transitional
180 vertebrae were present in 21 Pugs (30.9%). T13 in these dogs had characteristics of a lumbar
181 vertebra. Twenty-six (38.2%) Pugs had spina bifida. This anomaly was exclusively observed
182 at T1.

183

184 The group of English bulldogs consisted of 29 males and 12 females between 10 and
185 132 months old (median, 21.0; mean, 40.9 months) and weighing between 10.3 and 37.1kg
186 (median, 23.25; mean, 23.9kg). All dogs had 13 thoracic vertebrae. Ten (24.4%) English
187 bulldogs had no thoracic vertebral malformations. Thirty (73.2%) dogs had hemivertebrae. A
188 total of 100 hemivertebrae were present; three (7.3%) dogs had single hemivertebrae and 27

189 (65.9%) had multiple hemivertebrae. The most commonly affected vertebra was T9 ($n=18$),
190 followed by T7 ($n=14$) and T8 ($n=13$). Block vertebrae were diagnosed in four (9.8%) dogs,
191 with T11-T12 ($n=3$) being most frequently affected, followed by T12-T13 ($n=1$). Eight
192 (19.5%) English bulldogs had fused dorsal spinous processes. Transitional vertebrae were
193 present in four (9.8%) English bulldogs. In three of these dogs, T13 had characteristics of a
194 lumbar vertebra and in one dog, T1 had characteristics of a cervical vertebra. One (2.4%)
195 English bulldog had evidence of spina bifida at T10. Two English bulldogs had vertebrae
196 with characteristics of more than one type of malformation, which consisted of transitional
197 vertebra and hemivertebra ($n=1$) and a transitional and block vertebra ($n=1$).

198

199 *Dogs with clinically relevant vertebral malformations*

200 During the study period, a total of 105 French bulldogs, 192 Pugs and 120 English
201 bulldogs were presented for a wide variety of clinical indications (neurological and non-
202 neurological) at the Small Animal Referral Hospital, Royal Veterinary College. Of these
203 dogs, one French bulldog (0.95% of all French bulldogs presented) and nine Pugs (4.7% of
204 all pugs presented) were diagnosed with a thoracic vertebral malformation as the cause of
205 their clinical signs. These 10 affected dogs included eight males and two females, aged
206 between 4 and 57 months (median, 8.0 months; mean, 20.0). Duration of clinical signs at
207 presentation varied from 7 days to 2 years (median, 61 days; mean, 173 days) and consisted
208 of ambulatory paraparesis and ataxia of the pelvic limbs in all dogs. Spinal hyperaesthesia
209 could be elicited in two dogs. In each of these dogs, thoracic hemivertebrae were the cause of
210 their clinical signs (Fig. 2). Other abnormalities, not considered to be associated with clinical
211 signs, included spina bifida at the level of T1 in three Pugs and a transitional T13 vertebra in
212 two Pugs.

213

214 *Comparison between breeds*

215 There was a significant influence of breed on the overall prevalence of thoracic
216 vertebral malformations in neurologically normal dogs ($P < 0.0001$). More specifically,
217 neurologically normal French bulldogs had significantly more thoracic vertebral
218 malformations than neurologically normal Pugs ($P < 0.0001$). Neurologically normal English
219 bulldogs had significantly more thoracic vertebral malformations than neurologically normal
220 Pugs ($P = 0.002$). There were no significant differences between the other individual breeds
221 ($P > 0.01$). There was a significant influence of breed on the prevalence of hemivertebrae in
222 neurologically normal dogs ($P < 0.0001$). Hemivertebrae were diagnosed more often in
223 neurologically normal French bulldogs than in neurologically normal Pugs ($P < 0.0001$) and
224 English bulldogs ($P = 0.004$). Hemivertebrae were diagnosed less often in neurologically
225 normal Pugs than in neurologically normal English bulldogs ($P = 0.004$). There was a
226 significant influence of breed on the prevalence of spina bifida and transitional vertebrae in
227 neurologically normal dogs, with Pugs significantly more often affected compared to the
228 other two breeds ($P < 0.0001$ for both malformations). There was no significant influence of
229 breed on the prevalence of block vertebrae in neurologically normal dogs ($P = 0.086$).

230

231 Finally, there was a significant influence of breed on the prevalence of clinically
232 relevant thoracic vertebral malformations, with Pugs being overrepresented compared to the
233 other two breeds ($P = 0.006$).

234

235 **Discussion**

236 This study described and compared thoracic vertebral malformations in French
237 bulldogs, Pugs and English bulldogs with and without associated neurological deficits. The
238 results of this study support previous reports indicating a high prevalence of vertebral

239 malformations in neurologically normal screw-tailed brachycephalic breeds (Moissonnier et
240 al., 2011; Guevar et al., 2014; Gutierrez-Quintana et al., 2014). Overall, 80.7% of
241 neurologically normal animals were affected by at least a single malformation and 59.6% had
242 multiple malformations. This is comparable to previous work reporting a 64.2% prevalence
243 of multiple thoracic malformations in a population of neurologically normal and abnormal
244 brachycephalic screw-tailed dogs (Gutierrez-Quintana et al., 2014). This information has
245 clinical importance for the interpretation of imaging studies in animals with suspected spinal
246 disease and underlines the importance of questioning the clinical relevance of vertebral
247 malformations observed on radiological studies. This is highlighted by the fact that, although
248 thoracic vertebral malformations were commonly encountered on imaging studies, they were
249 only rarely considered the direct cause of clinical signs in the studied breeds. While the
250 prevalence of clinically relevant thoracic vertebral malformations was 4.7% in our hospital
251 population of Pugs overall, the clinical importance of this diagnosis was negligible in the
252 other two breeds.

253

254 Hemivertebrae were the most frequently diagnosed vertebral malformation in French
255 bulldogs and English bulldogs, which is consistent with previous publications (Westworth
256 and Sturges 2010; Moissonnier et al., 2011; Faller et al., 2014; Gutierrez-Quintana et al.,
257 2014). Additionally, the most frequent location for hemivertebrae was in the mid thoracic
258 region (T7-T9; Moissonnier et al., 2011; Faller et al., 2014; Guevar et al., 2014; Gutierrez-
259 Quintana et al., 2014). In our study, French bulldogs were significantly overrepresented for
260 hemivertebrae compared to the other breeds, with over 90% of neurologically normal animals
261 affected and the majority of cases showing multiple hemivertebrae. Interestingly, while
262 hemivertebrae occurred significantly less common in neurologically normal Pugs,
263 hemivertebrae accompanied by neurological deficits was diagnosed significantly more often

264 in this breed compared to the two other breeds. This finding suggests that although
265 hemivertebrae are less common in Pugs, this type of vertebral malformation is more likely to
266 be associated with the development of clinical signs in this breed. It is currently unclear why
267 hemivertebrae are more often associated with clinical signs in Pugs compared to French and
268 English bulldogs. A recent study proposed a classification system of canine hemivertebrae
269 into seven different subtypes (Gutierrez-Quintana et al., 2014). Development of clinical signs
270 in animals with hemivertebrae is thought to have a multifactorial aetiology, with vertebral
271 instability and vertebral canal stenosis considered to be contributing factors (Westworth and
272 Sturges, 2010; Moissonier et al., 2011; Dewey et al., 2016). Hemivertebrae can result in an
273 abnormal angulation of the vertebral column, referred to as kyphosis or scoliosis
274 (Moissonier et al., 2011; Aikawa et al., 2014; Faller et al., 2014; Guevar et al., 2014).
275 Recent studies have suggested that severity of kyphosis is a key factor in the development of
276 clinical signs in dogs with hemivertebrae and that a certain degree of vertebral kyphosis must
277 be exceeded before clinical signs are likely to occur (Moissonier et al., 2011; Guevar et al.,
278 2014). Although beyond the scope of this study, it is possible that Pugs are affected by a
279 different subtype of hemivertebrae which results in more severe kyphosis. Further studies are
280 therefore needed to evaluate the influence of breed on hemivertebra subtype and the influence
281 of hemivertebra subtype on the degree of vertebral kyphosis.

282

283 Pugs were significantly more often diagnosed with thoracolumbar transitional
284 vertebrae compared to French bulldogs and English bulldogs. In almost one third of
285 neurologically normal Pugs, T13 demonstrated characteristics of both thoracic and lumbar
286 vertebrae, including unilateral or bilateral absence of ribs and the presence of rudimentary
287 transverse processes. Although transitional vertebrae are not considered a direct cause of
288 spinal cord dysfunction, these anomalies have been associated with alterations in vertebral

289 biomechanics and can complicate the approach of routine spinal surgeries (Morgan et al.,
290 1968; Flückiger et al., 2006; Westworth and Sturges, 2010).

291

292 Spina bifida refers to failure of closure of one or more vertebral arches over the spinal
293 cord. It is categorised based on the level of neuroectoderm involved, and subtypes include
294 aperta (open), cystica (closed) and occulta (hidden; Song et al., 2016). It has been reported in
295 a variety of breeds and is probably related to genetic and environmental causes (Wilson et al.,
296 1979; Song et al., 2016). Less severe malformations are commonly encountered without
297 clinical signs, with malformations typically found incidentally on diagnostic studies (Song et
298 al., 2016). Our study reported that spina bifida occulta was an infrequent finding in English
299 and French bulldogs as only one case identified among these dogs, which is similar to the
300 findings of previous studies of comparable populations (Gutierrez-Quintana et al., 2014). In
301 contrast, in our study, Pugs were affected by thoracic spina bifida occulta significantly more
302 often than other breeds and this abnormality was found exclusively at T1. It is currently
303 unclear why spina bifida occulta was only present at T1 and what the clinical relevance of
304 this finding may be. This information should however be taken into account when evaluating
305 imaging studies of Pugs with suspected spinal disease.

306

307 Our study was limited by its retrospective design which meant the majority of this
308 population of dogs did not receive a neurological examination. Therefore, we cannot exclude
309 the possibility that a number of animals developed clinical signs relating to vertebral
310 malformations later in life. However, the retrospective study design enabled the inclusion of a
311 large population of dogs. Furthermore, neurologically normal animals underwent CT
312 imaging, while clinically affected dogs underwent MR imaging. Both advanced imaging
313 techniques are associated with specific advantages and disadvantages and differ clearly in

314 their diagnostic potential for imaging the bony vertebral column and the parenchymal spinal
315 cord. Therefore, we decided not to compare imaging findings of affected and unaffected dogs
316 directly. Although not evaluated in this study, further classification of hemivertebra subtype
317 might have determined whether a specific hemivertebra subtype was more prevalent in one of
318 the studied breeds and if there was an association with different degrees in spinal kyphosis. A
319 previous study has demonstrated differences in hemivertebra subtyping when survey
320 radiographs and CT were compared (Brocal et al., 2016). It is possible that differences would
321 also exist when comparing MRI and CT. For this reason, a comparison of hemivertebra
322 subtypes in affected and unaffected dogs was not attempted in our study. It is unclear whether
323 a classification system designed to evaluate specific bony vertebral body abnormalities could
324 be used reliably for MRI.

325

326 **Conclusions**

327 There is a high prevalence of vertebral malformations in neurologically normal
328 ‘screw-tailed’ brachycephalic breeds and there is a significant influence of breed on the
329 prevalence of each type of malformation. Neurologically normal French bulldogs are
330 significantly more often diagnosed with hemivertebrae than neurologically normal Pugs and
331 English bulldogs, while neurologically normal Pugs are diagnosed with hemivertebra
332 significantly less frequently than neurologically normal French and English bulldogs.
333 However, Pugs are diagnosed with hemivertebra as the cause of neurological deficits more
334 often than the other breeds evaluated. Further studies are needed to evaluate the influence of
335 breed on hemivertebra subtype and the influence of hemivertebra subtype on the degree of
336 vertebral kyphosis.

337

338 **Conflict of interest statement**

339 None of the authors has a financial or personal relationship with people or
340 organisations that could inappropriately influence or bias the content of this paper.

341

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399 Fig. 1. Sagittal reconstructed (A and B), transverse and three-dimensional reconstructed CT
400 images illustrating the evaluated thoracic vertebral malformations; hemivertebra (A), block
401 vertebra (B), spina bifida (C) and a thoracolumbar transitional vertebra (D). Vertebral
402 malformations indicated by white arrow.

403

404 Fig. 2. (A) T2-weighted sagittal and (B) T1-weighted sagittal magnetic resonance images of a
405 9-month old Pug with a clinically relevant thoracic vertebral body malformation.
406 Hemivertebra indicated by white arrow.

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