

1 **CANINE BREEDS PREDISPOSED TO DEVELOP DISKOSPONDYLITIS:**
2 **A RETROSPECTIVE STUDY OF 181 CASES (2009-2018)**

3
4 **PREDISPOSIÇÃO RACIAL CANINA PARA O DESENVOLVIMENTO DE**
5 **DISCOESPONDILITE: ESTUDO RETROSPECTIVO DE 181 CASOS**
6 **(2009-2018)**

7
8 **RESUMO**

9 Para determinar a prevalência e predisposição racial da discoespondilite (DS) em cães
10 realizou-se uma pesquisa numa população de 5497 animais submetidos a exame de
11 tomografia computadorizada ou radiografia digital da coluna entre 2009 e 2018. Variáveis
12 como raça, sexo, idade, segmento vertebral e total de vértebras acometidas foram coletadas e
13 submetidas aos testes de prevalência, Qui-quadrado e odds ratio. Foram identificados 181 cães
14 com DS, prevalência de 3,4%. Destes, 65% eram machos, probabilidade 1,6x maior que
15 fêmeas (CI 1.17-2.17). Cães maiores que 10 anos tem probabilidade 1,5x maior (CI 1.10-
16 2.05), enquanto em cães entre 2-5 anos a probabilidade diminui 51% (CI 0.34-0.77).
17 Observou-se o predomínio de cães de grande porte (>30 kg; 45%), com 3,8x mais chances de
18 DS (CI 2.56-5.33); seguido de 28% de cães de pequeno porte, ainda que demonstrada uma
19 probabilidade 34% menor (CI 0.24-0.47). O labrador apresenta 3,7x mais chances que todas
20 as raças estudadas (CI 2.56-5.33) e o buldogue francês, entre as raças de pequeno porte, 2,8x
21 mais susceptibilidade (CI 1.51-5.06). Conclui-se que fatores como idade avançada, grande
22 porte e, especialmente labradores, apresentam maior probabilidade a serem portadores de DS.
23 O buldogue francês deve ser mais estudado quanto a sua discrepância em comparação a raças
24 de mesmo porte.

25 **Palavras-chave:** Buldogue Francês. Cão. Coluna Vertebral. Labrador Retriever. Mielopatias.

26
27 **ABSTRACT**

28 A study to determine the prevalence and predisposition of dog breeds to develop
29 diskospondylitis (DS) was carried out on a population of 5,497 animals submitted to
30 computed tomography or digital radiography of the spine between 2009 and 2018. Variables
31 such as breed, gender, age, vertebral segment and total number of vertebrae affected were
32 collected and submitted to the prevalence tests, chi-square and *odds ratio*. A total of 181 dogs
33 presented DS, a prevalence of 3.4%. Of these, 65% were males with a probability 1.6x greater

34 than females (CI 1.17-2.17). Dogs more than 10 years old have a 1.5x higher probability (CI
35 1.10-2.05), while those between 2-5 years the probability decreases 51% (CI 0.34-0.77).
36 Large dogs (>30 kg; 45%) showed a 3.8x greater chance to develop DS (CI 2.56-5.33) than
37 small dogs (<15 kg; 28%), although the small dogs showed a 34% lower probability (CI 0.24-
38 0.47). The Labrador Retriever breed was 3.7x more likely to develop DS than all the other
39 breeds studied (CI 2.56-5.33) and the French Bulldog, among the small breeds, was 2.8x more
40 susceptible (CI 1.51-5.06). In conclusion older dogs, large dogs, especially Labrador
41 Retrievers, are more likely to develop DS. The French bulldog should be studied further.

42 **Keywords:** Dog. French Bulldog. Labrador Retriever. Myelopathies. Spine.

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INTRODUCTION

45 Diskospondylitis (DS) is a primary infection of the cartilaginous vertebral endplates
46 with secondary involvement of the intervertebral disk that leads to neurologic disorders
47 (THOMAS, 2000). Conventionally, the diagnosis of diskospondylitis has relied on the
48 presence of characteristic radiographic findings such as the loss of the vertebral end plate
49 margin definitions, narrowing or collapse of the intervertebral disk (IVD) space and various
50 degrees and combinations of endplate lysis, sclerosis, and bony proliferation (GENDRON et
51 al., 2012; HARRIS et al., 2013; RUOFF et al., 2017).

52 Spinal pain is the most common finding; however, clinical signs can be non-specific
53 such as weight loss, depression, fever, and anorexia (HARRIS et al., 2013). The dogs may be
54 neurologically normal or with various degrees of neurologic impairment (BURKERT et al.,
55 2005). Several pathological processes can account for the neurologic deficits in this disease
56 and include extrusion of intervertebral disk material secondary to collapse of an affected disk
57 space, osseous or soft tissue proliferation within the vertebral canal in response to chronic
58 inflammation, vertebral subluxation or pathological fracture secondary to marked bone lysis,
59 and secondary meningitis or myelitis (GENDRON et al., 2000; RUOFF et al., 2017; DAVIS
60 et al., 2000).

61 The main differential diagnoses for diskospondylitis are neoplasia and degenerative
62 intervertebral disk disease (IVDD). Vertebral tumors may have features similar to
63 diskospondylitis, although the location of the lesions usually involves a single vertebra and is
64 centered on the vertebral body rather than the endplates, also the intervertebral disk is not
65 affected. Bone-marrow tumors tend to be multifocal and similarly the endplates and the

66 intervertebral disks are not affected. Finally, in IVDD the endplates may appear sclerotic, but
67 do not show any erosion or destruction of the cortical bone (THOMAS, 2000; DA COSTA &
68 MOORE, 2010; GENDRON et al., 2012).

69 Acute cases may be a diagnostic challenge since clinical signs can be non-specific and
70 radiographic findings appear belatedly. Thus, it is important to "keep in mind" the possibilities
71 of DS according to the breed of the dog, its gender, age and medullary neurological
72 dysfunctions. The aim of this study was to describe the signalment of dogs with DS, compare
73 them with a population of dogs submitted to spinal image exams and determine which specific
74 characteristics increase the odds to develop DS.

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MATERIAL & METHODS

77 All the computerized tomography (CT) scans or digital radiography (RX) exams of the
78 electronic medical record system acquired between April 2009 and June 2018 at a Veterinary
79 Reference Center (**HIDDEN FOR SUBMISSION**) were searched for "diskospondylitis". The
80 radiographs and CT scans found were reviewed and only the exams that described features
81 such as loss of definition of end-plate margins, narrowing of the IVD space, lytic bone
82 changes of vertebrae adjacent to the IVD space and sclerosis at the margins of bone lysis were
83 included (DS group). The other canine patients submitted to spine TC or RX exams during the
84 same interval were used as controls (C group). Signalment (sex, age, breed), IVD space and
85 number of vertebrae affected were recorded. The dogs in both groups were categorized
86 according to the standard breed size (small, <15 kg; medium, 15-30 kg; large, >30 kg or
87 mixed-breed) and age (puppy, <2 years old; young, 2-5 years old; adult, 5-10 years old;
88 elderly, >10 years old). The owners of dogs with DS were contacted by phone to obtain the
89 outcome of the disease.

90 Statistical analyses were performed with statistical software packages. The results were
91 described as percentages (frequency). Contingency tables were generated for the categorical
92 variables (sex, breed and age). The twenty most prevalent breeds were compared to the
93 control group and to the same breed size group (small, medium or large breed groups).
94 Distribution of factors were compared between DS and C groups via the X^2 test. Odds ratio
95 (OD) and 95% confidence interval (CI) were determinate for each comparison. Data for age,
96 anatomical site and owner's perception of the outcome was evaluated using Kruskal-Wallis
97 nonparametric analysis of variance. Factors were considered significant when the P value was
98 ≤ 0.05 and the exceeded 1.0.

RESULTS

100 One hundred eighty-one cases with a diagnostic for diskospondylitis (127 CT and 54
101 RX) met the inclusion criteria (DS group) and 5,316 dogs (3,001 TC and 2,315 RX exams)
102 were used as control cases (C group). The prevalence of the disorder was 3.4%. In the DS
103 group 65% (117) were male and 35% (64) were female dogs. Male dogs has 1.6 times more
104 chance to had DS than females (OR 1.6; CI 1.17-2.17) (Table 1).

105 The mean age of the dogs was similar in the DS and C groups (8 ± 4 years, $p=0.898$).
106 Animals more than 10 years old had 1.5 times more chance to develop DS (OR 1.5; CI 1.10-
107 2.05) while dogs between 2 to 5 years old had 51% less probability (OR 0.51; CI 0.34-0.77)
108 (Table 1). The average number of vertebrae affected was 3 varying from 2 to 12. The most
109 prevalent vertebral segments affected were the thoracolumbar and lumbosacral, with 28% (51)
110 each (Table 2).

111 In terms of breeds, 44.7% (81) of the DS cases were large breed dogs, 27.6% (50) were
112 small breeds, 13.3% (24) were medium breeds and 14.4% (26) were mixed-breeds (Table 1).
113 Large breeds had a 3.8 times greater chance to develop DS (OR 3.8; CI 2.80-5.12); however,
114 for small breeds, the chances decreased 34% (OR 0.34; CI 0.24-0.47). The most frequent large
115 breed in the DS group was the Labrador Retriever (22% - $n=40$) which presented significantly
116 increased chances (OR 3.7; CI 2.56-5.33) compared to all the other breeds (Table 3 and 4). In
117 relation to small breeds, the French Bulldog (FB) was the most frequent (32% - $n=16$). When
118 compared to other small breeds, FB presented a significant increase in the possibility to
119 develop this disorder (OR 2.7; CI 1.51-5.06). Also, the age of French Bulldogs affected was
120 significantly lower compared with all groups studied (Table 5).

121 Only sixty-two owners were found and forty (64%) reported that their dogs had no
122 outcome or remained with sequelae. In ten cases, the diskospondylitis was the direct cause of
123 death or euthanasia.

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DISCUSSION

126 This retrospective study confirms the high prevalence of large breeds developing DS,
127 highlighting the significant risk for Labrador Retrievers in relation to other large breeds. In
128 addition, the results highlight the increased chances of French Bulldogs to develop DS, which,
129 to the authors knowledge, has not been reported previously.

130 The results showed that DS had a prevalence of 3.4% in dogs submitted to spinal image
131 exams. In order to reduce the limitation of non-access to any clinical record, this study only

132 included dogs that had undergone CT or RX exams of the vertebral column. These exams are
133 important tools to diagnose spinal-medullary disorders. Another retrospective study used all
134 canines that had been admitted to the veterinary clinic as the control group during the study
135 period (BURKERT et al., 2005) which would certainly justify the lower prevalence observed,
136 compared with the present study.

137 Some epidemiological aspects previously described were endorsed: (1) male dogs are
138 more affected (HUROV et al., 1978; DAVIS et al., 2000; BURKERT et al., 2005; CANAL et
139 al., 2016); (2) Large breeds are more susceptible (HUROV et al., 1978; DAVIS et al., 2002;
140 HARRIS et al., 2013; CANAL et al., 2016); (3) The average age of dogs affected is five years
141 old (HARRIS et al., 2013, CANAL et al., 2016), although older dogs (>10 years) are more
142 prevalent (DAVIS et al., 2000; BURKERT et al., 2005).

143 Interestingly, dogs between two to five years old have a 51% less chance of a spinal
144 disorder being DS. However, various reports have highlighted the occurrence of DS in
145 puppies (ADAMO & CHERUBINI, 2001; FINNEN et al., 2012; KIRBERGER, 2016),
146 especially in large breeds, while prospective/retrospective studies describe a high prevalence
147 in elderly dogs (BURKERT et al., 2005). In humans, the presence of any endplate lesion has
148 statistically and significantly been associated with the elderly (WANG et al. 2012,
149 AGUILAR-COMPANY et al., 2018), but has not been reported in children as observed in
150 young dogs. In general, most studies suggest that DS is frequently associated to canine urinary
151 tract and prostatic infections (BURKERT et al., 2005), common diseases of male and elderly
152 dogs, but fungus (SCHULTZ et al., 2008), previous surgeries (THOMAS, 2000; CANAL et
153 al., 2016; SHWARTZ et al., 2016) and previous infections from foci elsewhere in the body
154 (ADAMO & CHERUBINI, 2001; KIRBERGER, 2016) have also been associated to DS. In
155 humans, the risk factor includes advanced age, diabetes mellitus, hypothyroidism,
156 genitourinary tract infections, respiratory tract infections, rheumatoid arthritis,
157 immunosuppression and surgery (FRIEDMAN et al., 2002; AGUILAR-COMPANY et al.,
158 2018).

159 Although any region of the vertebral column can be affected, in our study 28.1% were
160 observed along the thoracic spine (T1-T13) and the same high prevalence was observed at the
161 lumbosacral IV space (L7-S1). Davis et al. (2000), Kurbert et al. (2005) and Harris et al.
162 (2013) observed similar findings that demonstrate the significance of DS as a differential
163 diagnosis for pain and neurological dysfunctions in these regions. The high prevalence of the
164 LS space may be explained by Carrera et al. (2011) and Gendron et al. (2012) who
165 demonstrated using Magnetic Resonance Imaging that the lumbosacral joint was a

166 predilection site for endplate lesions and the most common presumptive endplate lesion is DS.
167 The vertebral endplate is responsible for transferring stress between disc and vertebral body,
168 and its concavity is important in dispersing compression stress (HE et al., 2012). Humans with
169 osteoporosis, sclerosis, vascular compromise and necrotic areas developed in the vertebral
170 endplates are known to result in microfractures and later in infection – but such microfractures
171 in vertebral endplates as a precursor to vertebral infection have not been detected in dogs
172 (BURKERT et al., 2005). Burkert et al. (2005) hypothesizes that intermittent venous
173 occlusion or stasis of blood flow at the lumbosacral junction during locomotion may lead to
174 focal endplate necrosis and an episode of bacteremia could then lead to focal colonization.

175 The Labrador Retriever is frequently described as one of the most affected breeds in
176 diskospondylitis clinical studies (DAVIS et al., 2000; HARRIS et al., 2013; BURKERT et al.,
177 2005). Also, in this study the prevalence of DS in Labrador Retrievers was 26% and the
178 chances were four times greater compared to all the other breeds studied here, which
179 emphasizes the potential risk for this breed. Labradors are a very popular household pet dog
180 breed and one of the most common breeds used worldwide as working dogs. Recently studies
181 demonstrated that this breed also has a genetic predisposition to similar diseases to DS such as
182 lumbosacral stenosis (MUKHERJEE et al., 2017) and osteoarthritis (ANDERSON et al.,
183 2018). Other hypothesis that may explain the high prevalence includes the genetic
184 predisposition to obesity (MANKOWSKA et al., 2017). Similarly to humans dog obesity can
185 predispose or exacerbate several clinical conditions such as osteoarthritis, respiratory airway
186 distress, renal diseases, diabetes mellitus and metabolic derangements (IMPELLIZERI et al.,
187 2000; TVARIJONAVICIUTE et al., 2013). Notably both over-nutrition and obesity have been
188 associated with impaired immunity and chronic low-grade inflammation in humans and
189 mouse models (NIEMAN et al., 1999; BERG & SCHERER, 2005). In fact, Anderson et al.
190 (2018) reports that breeds at or above mean breed bodyweight were 2.3 times more likely to
191 have osteoarthritis than dogs below average weight.

192 French Bulldogs, a chondrodystrophic, brachycephalic and screw-tailed breed has a
193 high risk of congenital vertebral anomalies (BERTRAN et al., 2018). Theses vertebral
194 malformations can occur in isolation or be multiple and are frequently associated with
195 vertebral malalignment and angulations (BAILEY e MORGAN, 1992; WESTWORTH &
196 STURGES, 2010). Furthermore, it is unclear why most dogs with vertebral malformations are
197 often asymptomatic, while some dogs develop clinical signs (MOISSONNIER et al., 2011;
198 GUEVAR et al., 2014; RYAN et al., 2017; BERTRAM et al., 2018). Since this breed has

199 become increasingly popular, recent epidemiological studies have demonstrated that FBs are
200 prone to several neurological disorders: Mayousse et al. (2017) reported that 18.7% of FBs
201 have neurological disorders whereas 64.7% were myelopathies; O'Neil et al. (2018) reported
202 that the most common cause of death in primary care were brain disorders (11.9%), spinal
203 cord disorders (9.5%) and vertebral spinal disorders (6%). In fact, the most common disorder
204 is IVDD (AIKAWA et al., 2014), but the risk of developing DS reported in our study may
205 provide new insights to clinical practice.

206 Although only 34% (62) of the owners of the DS group were found, 64% (40) of them
207 reported that their animal did not recover from the disease. This represents at least 22% of the
208 DS cases. The prognosis for dogs with DS is considered to be good; however, long-term
209 treatment with antimicrobials and confinement to a cage to rest is necessary, which is more
210 difficult with large breeds (BURKERT et al., 2005). Moreover, elderly dogs may be
211 debilitated by age or have some concurrent disease. Consequently, the morbidity and
212 mortality among cases may appear to be high. Diskospondylitis may be associated with severe
213 morbidity and neurological impairment if the definitive diagnosis is delayed as a result of
214 doubtful or normal radiographic findings in acute cases. Advanced imaging such as computed
215 tomography and magnetic resonance have shown greater sensitivity and precocity diagnosis
216 for DS (GENDRON et al., 2012; HARRIS et al., 2013; RUOFF et al., 2018), and is therefore
217 a differential in the acute phase of the disorder.

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CONCLUSION

220 Large breed dogs are more susceptible to develop DS and the Labrador Retriever
221 breed has a higher risk than all large breeds studied here. Although small breeds have less
222 chance to develop DS, the French Bulldogs require further investigations - as diskospondylitis
223 can be considered an important differential diagnosis for this breed.

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TABLES

Table 1. Results of univariate analysis to identify factors associated with prevalence of diskospondylitis (DS) in 5,497 dogs submitted to vertebral computerized tomography (n=3,128) or digital radiography (n=2,369) from 2009 to 2018.

| Characteristics | | | with DS %(n) | without DS %(n) | OD | 95% CI | P value* |
|---------------------------|----------------|-----------------|-------------------------|----------------------------|-----------|-------------------|---------------------|
| All animals | | | 3.4% (181) | 96.6% (5316) | - | - | - |
| Breeds | | | | | | | |
| <i>small (<15 kg)</i> | | | 27.6% (50) | 53% (2815) | 0.34 | 0.24- 0.47 | <0.001 |
| <i>medium (15-30 kg)</i> | | | 13.3% (24) | 16.2% (863) | 0.79 | 0.51- 1.22 | 0.334 |
| <i>large (>30 kg)</i> | | | 44.7% (81) | 17.6% (937) | 3.78 | 2.80- 5.12 | <0.001 |
| <i>Mixed</i> | | | 14.4% (26) | 13.2% (704) | 1.10 | 0.72- 1.68 | 0.744 |
| Age | | | | | | | |
| <i>puppy (<2ys)</i> | | | 11.0% (20) | 7.2% (382) | 1.60 | 1.00- 2.58 | 0.069 |
| <i>young (2-5ys)</i> | | | 15.5% (28) | 26.4% (1403) | 0.51 | 0.34- 0.77 | 0.001 |
| <i>adult (5-10ys)</i> | | | 38.1% (69) | 39.7% (2110) | 0.936 | 0.69- 1.27 | 0.728 |
| <i>elderly (>10ys)</i> | 35.3% (64) | 26.7% (1418) | 1.504 | 1.10-2.05 | | 0.012 | |
| Sex | | | | | | | |
| <i>Male</i> | 64.6% (117) | 53.4% (2840) | 1.60 | 1.17-2.17 | | 0.004 | |
| <i>Female</i> | 35.4% (64) | 46.6% (2477) | 0.63 | 0.46-0.85 | | 0.004 | |

OD – odds ratio; CI – confidence interval
 * X^2 test

342 **Table 2. BREED** distribution for cases with diskospondylitis (DS) and controls.

| Breeds | with DS %(n) | without DS %(n) | OD | 95% CI | P value* |
|-----------------------|-------------------------|----------------------------|-----------|---------------|---------------------|
| All dogs | n=181 | n=5316 | - | - | - |
| Labrador Retriever | 22.1% (40) | 7.1% (379) | 3.70 | 2.56-5.33 | <0.001 |
| Mixed-breed | 14.4% (26) | 13.4% (713) | 1.05 | 0.69-1.61 | 0.894 |
| French Bulldog | 8.8% (16) | 7.7% (409) | 1.16 | 0.69-1.96 | 0.670 |
| Yorkshire | 5.5% (10) | 6.1% (325) | 0.90 | 0.47-1.72 | 0.867 |
| Golden Retriever | 5% (9) | 2.9% (154) | 1.75 | 0.88-3.49 | 0.163 |
| Poodle | 5% (9) | 11.1% (591) | 0.42 | 0.21-0.82 | 0.013 |
| Rottweiler | 4.4% (8) | 2.6% (139) | 1.72 | 0.83-3.57 | 0.213 |
| Shepherd (all breeds) | 3.9% (7) | 2.9% (155) | 1.34 | 0.62-2.90 | 0.602 |
| Boxer | 2.8% (5) | 2.1% (113) | 1.31 | 0.53-3.24 | 0.749 |
| Dachshund | 2.8% (4) | 10.2% (541) | 0.20 | 0.07-0.54 | <0.001 |
| English Bulldog | 2.2% (4) | 0.8% (45) | 2.65 | 0.94-7.44 | 0.129 |
| Schnauzer | 2.2% (4) | 1.9% (99) | 1.19 | 0.43-3.27 | 0.952 |
| Pit bull | 1.6% (3) | 0.9% (46) | 1.93 | 0.59-6.27 | 0.476 |
| Pug | 1.6% (3) | 1.9% (102) | 0.86 | 0.27-2.74 | 0.981 |
| Maltese | 1.1% (2) | 4.5% (239) | 0.24 | 0.06-0.96 | 0.045 |
| Beagle | 0.5% (1) | 2.3% (123) | 0.23 | 0.03-1.69 | 0.189 |

343 OD – odds ratio; CI – confidence interval

344 * X^2 test

345

346 **Table 3. SMALL BREED** distribution of cases with diskospondylitis (DS) and controls.

| Breeds | with DS %(n) | without DS %(n) | OD | 95% CI | P value* |
|-----------------------|-------------------------|----------------------------|-----------|---------------|---------------------|
| All small breeds | n=50 | n=2815 | - | - | - |
| <i>French Bulldog</i> | 32% (16) | 14.5% (409) | 2.77 | 1.51-5.06 | 0.001 |
| <i>Yorkshire</i> | 20% (10) | 11.5% (325) | 1.91 | 0.95-3.87 | 0.105 |
| <i>Poodle</i> | 18% (9) | 21.0% (591) | 0.83 | 0.40-1.71 | 0.733 |

347 OD – odds ratio; CI – confidence interval

348 * X^2 test

349

350 **Table 4. LARGE BREED** distribution of cases with diskospondylitis (DS) and controls.

| Breeds | with DS %(n) | without DS %(n) | OD | 95% CI | P value* |
|---------------------------|-------------------------|----------------------------|-----------|---------------|---------------------|
| All large breeds | n=81 | n=937 | - | - | - |
| <i>Labrador Retriever</i> | 49.4% (40) | 40.4% (379) | 1.44 | 0.91-2.27 | 0.147 |
| <i>Golden Retriever</i> | 11.1% (9) | 16.4% (154) | 0.64 | 0.31-1.30 | 0.273 |
| <i>Rottweiler</i> | 9.9% (8) | 14.9% (139) | 0.63 | 0.30-1.33 | 0.292 |

351 OD – odds ratio; CI – confidence interval

352 * X^2 test

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357 **Table 5.** Diskospondylitis **SITE** distribution.

| disk spaces affected | (n=181) | LR (n=40)* | FB (n=16)** |
|--------------------------------|------------|------------|-------------|
| <i>cervical (C1 to C7)</i> | 10.4% (19) | 7.5% (3) | (0) |
| <i>cervicothoracic (C7-T1)</i> | 7.7% (14) | 17.5% (7) | (0) |
| <i>thoracic (T1 to T13)</i> | 28.1% (51) | 22.5% (9) | 25% (4) |
| <i>thoracolumbar (T13-L1)</i> | 1.6% (03) | 2.5% (1) | 6.2% (1) |
| <i>lumbar (L1 to L7)</i> | 12.7% (23) | 5.0% (2) | 6.2% (1) |
| <i>lumbosacral (L7-S1)</i> | 28.1% (51) | 40.0% (16) | 50% (8) |
| <i>Multifocal</i> | 11.4% (20) | 5.0% (2) | 12.5% (2) |

358 * Labrador Retriever

359 ** French Bulldog

360