

1 Brachycephalic obstructive airway surgery outcome assessment using the 6-minute walk test:

2 a pilot study

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8 Brachycephalic obstructive airway syndrome (BOAS) is used to describe a combination of
9 upper airway problems in brachycephalic dog breeds (such as English bulldogs, French
10 bulldogs and pugs) that lead to partial obstruction of the upper respiratory tract (Lecoindre &
11 Richard 2004, Packer 2013). Many affected dogs require surgical intervention such as
12 rhinoplasty, soft palate resection and laryngeal sacculotomy (Meola, 2013). The outcome of
13 BOAS surgery has been subjectively evaluated with 88 to 94% of dogs showing improvement
14 after surgery according to owner questionnaires (Poncet et al. 2006, Torrez & Hunt 2006,
15 Riecks et al. 2007, Pohl et al. 2016). Whole-body barometric plethysmography can objectively
16 evaluate respiratory function (Bernaerts et al. 2010, Liu et al. 2015) and has shown a significant
17 improvement in respiratory function after surgery but, in contrast, a simple objective
18 assessment of respiratory function has not been reported.

19

20 The 6-minute walk test (6-MWT) is a safe and simple test that is commonly used to assess
21 functional exercise capacity in humans with impaired cardiorespiratory function (Olsson et al.
22 2005, Bellet et al. 2012). The 6-MWT has been assessed in dogs with idiopathic pulmonary
23 fibrosis (Lilja-Maula et al. 2014), induced obesity (Manens et al. 2014) and induced congestive
24 heart failure (Boddy et al. 2004).

25

26 The aim of this pilot study was to assess the practicality and feasibility of the 6-MWT in
27 brachycephalic dogs as a test for assessment of functional exercise capacity after airway
28 surgery.

29

30 **Materials and methods**

31 *Recruitment of dogs*

32 Privately owned pugs, French bulldogs (FB) and English bulldogs (EB) clinically affected by
33 BOAS were prospectively recruited between August 2015 and February 2016 at the Queen
34 Mother Hospital for Animals (Royal Veterinary College, London). Dogs were excluded if
35 they were oxygen-dependent and deemed unable to be without supplemental oxygen to allow
36 the 6-MWT to be performed, or had severe orthopaedic or neurological disease that impaired
37 walking. None of the recruited dogs had detectable cardiac disease at the time of the study.
38 Body condition score was recorded based on a 1 to 9 scale on which 4 to 5 is considered
39 ideal. Approval from the Queen Mother Hospital for Animals Ethics and Welfare committee
40 had been granted (URN 2015 1337).

41

42 *6-MWT and measured values*

43 Dogs were walked along a 75-m corridor at a quiet time, for 6 uninterrupted minutes. Dogs
44 were walked on a leash with a harness and were allowed to walk at their own pace and to stop
45 if needed. The 6-MWT was performed by two of the investigators (EV, LR). The distance
46 walked (forward motion only) within the 6 minutes was recorded in metres.

47

48 Heart rate, respiratory rate and oxygen saturation (SpO₂) by pulse oximetry (Viamed Ltd,
49 VM-2500-S) were recorded before walking, immediately after walking and 2 and 5 minutes
50 after the exercise test. Rectal temperature was recorded before and immediately after the
51 walk. SpO₂ was recorded on the ear, lip or prepuce.

52

53 *Data collection*

54 The 6-MWT was performed for all dogs before BOAS surgery (and before any
55 sedation/anaesthesia) after being hospitalised for a few hours as an acclimatisation period,
56 approximately 24 hours after surgery, and again at least 6 weeks after surgery. Preoperative

57 evaluation included clinical scoring according to the Poncet clinical scoring system (Poncet et
58 al. 2005); this clinical scoring is based on the frequency of snoring, inspiratory effort,
59 exercise intolerance and syncope. All dogs were anaesthetised using the same protocol under
60 supervision of a ECVAA board-certified veterinary anaesthetist. Oropharyngeal examination
61 included description of stenotic nares, presence of elongation of the soft palate and degree of
62 laryngeal collapse. A subjective evaluation of degree of stenosis of the nares was made as
63 follows: mild stenosis of the nares (narrowing of the nostril by <25% compared to what is
64 considered a “normal adequate” opening of the nostril for the breed); moderate stenosis of the
65 nares (narrowing of the nostril of between 25 and 50%) and severe stenosis of the nares
66 (narrowing of the nostril of >50%). Laryngeal sacculle eversion was seen in all animals and
67 considered to be part of a grade 1, 2 or 3 laryngeal collapse. The physical examination was
68 performed on all the included cases by the same ECVS board-certified surgeon. After
69 oropharyngeal examination, CT of the head, neck and thorax was performed. Tracheal
70 hypoplasia was not evaluated.

71

72 BOAS surgery consisted of partial staphylectomy, using a modified technique described by
73 Bright & Wheaton (1983). Briefly, the palate was sharply cut with Metzenbaum scissors in an
74 arch shape, taking most tissue from the medial part of the soft palate, approximately to the
75 level of the cranial third of the tonsils. The oropharyngeal and nasopharyngeal mucosa were
76 apposed using simple interrupted absorbable sutures (Bright & Wheaton 1983, Riecks et al.
77 2007) (polyglactin 910, Vicryl Rapide™, Ethicon Inc.). A modified horizontal wedge
78 resection rhinoplasty was also performed (Schmiedt & Creevy 2011) and closed with simple
79 interrupted absorbable sutures (Monocryl®, polyglecaprone 25, Ethicon Inc.). Laryngeal
80 saccullectomy was not performed in any patient. All the surgeries were performed by the
81 same ECVS board-certified surgeon or ECVS resident under direct supervision.

82

83 *Statistical analysis*

84 Statistical analysis was performed using commercially available software (SPSS version 22).

85 Mean \pm sd values are given for age and body weight of all dogs and for distance walked, heart
86 rate (HR), respiration rate (RR), temperature and SpO₂ for each time point. Repeated

87 measures analysis of variance (ANOVA) were used to compare the distance walked, HR, RR,

88 temperature and SpO₂ between the different time points and between the different tests

89 (preoperatively, 24 hours postoperatively and 6 weeks postoperatively). P values <0.05 were

90 considered significant.

91

92 One-way ANOVA were used to compare the distance walked between different grades for

93 stenotic nares, elongation of soft palate, nasopharyngeal turbinate protrusion, laryngeal

94 collapse and Poncet clinical score. P values <0.05 were considered significant.

95

96 **Results**

97 *Patients and preoperative assessment results*

98 A total of 24 dogs were recruited to this pilot study: 10 FB, nine EB and five Pugs. Seventeen

99 dogs were male (12 entire, five neutered) and seven were female (five entire, two neutered).

100 The mean (\pm sd) age was 21.7 (\pm 16.6) months (range 4.6 to 79 months). The mean (\pm sd) weight

101 was 15 (\pm 7.7) kg (7.2 to 34.6 kg) and none of the dogs were considered clinically obese.

102 According to the Poncet clinical scoring system 83% of the dogs were grade 3, 11% of dogs

103 were grade 2 and 6% were grade 1. Nose and oropharyngeal examination findings are presented

104 in Table 1. CT findings included nasopharyngeal turbinate protrusion of grade 1 in seven dogs,

105 grade 2 in six dogs, grade 3 in four dogs and grade 4 in three dogs (Vilaplana Grosso et al.

106 2015). All dogs performed the preoperative and 24 hours postoperative 6-MWT. Four dogs did

107 not perform the 6-week postoperative 6-MWT because of failure to attend. These four dogs
108 were excluded from the repeated measures ANOVA analysis.

109

110 *Results of 6-MWT*

111 The results for HR, RR, SpO₂, rectal temperature and distance walked are presented in Table
112 2. Fig. 1 also shows the distance walked by each dog at the different 6-MWT. There was a
113 significant change in distance walked by each dog at 6 weeks postoperatively compared to
114 preoperatively, with an average increase of $13.7 \pm 28.8\%$ (range -31.3 to 66.6%). At 24 hours
115 postoperatively there was a significant decrease in distance walked, HR and temperature. There
116 was no significant difference in distance walked for the different grades of clinical scoring or
117 anatomical abnormalities.

118

119 **Discussion**

120 In this pilot study, the 6-MWT was feasible and relatively easy to perform on brachycephalic
121 dogs. Most dogs demonstrated an improvement in distance walked on the >6 weeks 6-MWT
122 consistent with improved cardiopulmonary function after airway surgery.

123

124 The clinical signs were evaluated using the Poncet respiratory clinical scoring system (Poncet
125 et al. 2005): 83% of our dogs fell in the grade 3 for respiratory signs, in line with the previous
126 reports (Poncet et al. 2006, Torrez & Hunt 2006, Pohl et al. 2016). No significant difference in
127 distance walked was found for the different clinical scores perhaps as type II error because of
128 the small sample size. Postoperative Poncet clinical scoring was not performed in this study,
129 which prevented comparison with the postoperative 6-MWT.

130

131 Although assessment of outcome after BOAS surgery currently mainly relies on the perception
132 of the owner (Poncet et al. 2006, Torrez & Hunt 2006, Riecks et al. 2007, Pohl et al. 2016),
133 they may not recognise respiratory compromise in their dogs (Packer 2013, Liu et al. 2015)
134 and there may well be individual variability in owner assessment. The 6-MWT would allow
135 assessment of large numbers of dogs and could allow for evaluation of the influence of type of
136 surgery or clinical grade on the outcome after BOAS surgery.

137

138 Our study showed an average increase of 13.7% in the distance walked during the 6-MWT at
139 6 weeks after BOAS surgery, indicating an improved exercise tolerance. However, not all
140 individual dogs demonstrated an improvement. Four dogs, all EBs, demonstrated decreased
141 exercise tolerance. Four dogs, all FBs, walked 35 to 66% further after surgery compared to
142 before. This could possibly be because FB are more afflicted with anatomical abnormalities
143 that are amenable to surgery, but larger numbers of dogs would need to be examined to
144 investigate this in more detail. Tracheal hypoplasia and thickness of the soft palate (as opposed
145 to length) were not evaluated and could also have influenced our results.

146

147 Our results also show a decrease in resting respiratory rate and an increase in SpO₂ following
148 surgery. SpO₂ measurement can be insensitive and variable when using pulse oximetry, which
149 could explain the wide range of measures obtained, and the changes might not be clinically
150 important. Arterial oxygen saturation measurement is more reliable but is not easily accessible
151 for all patients. Arterial blood sampling was not deemed clinically necessary for these patients.
152 The decrease in resting respiratory rate at 6 weeks after surgery might reflect an improvement
153 in respiratory function although the change observed is small and might not be clinically
154 important.

155

156 Interestingly, the 6-MWT results obtained at 24 hours after surgery were generally worse than
157 those just before surgery. We consider that this might be a consequence of the recent general
158 anaesthesia and concurrent opioid analgesia, and/or surgery-related swelling.

159

160 In our results, the distance walked (mean 446 m before surgery) is still far lower than that
161 published for other breeds; Manens et al. (2014), Boddy et al. (2004) and Swimmer & Rozanski
162 (2011) report mean distances walked of 589, 573 and 522.7 m in healthy beagles, research
163 hounds and various breeds, respectively. This difference likely highlights the brachycephalic-
164 related exercise intolerance, although direct comparison between studies is difficult and other
165 factors such as stride length or leg height could also account for the differences. In these reports,
166 control dogs walked 9 to 33% more in the 6-MWT than affected dogs (namely obese dogs,
167 dogs in congestive heart failure and dogs with pulmonary disease). The 6-MWT could be used
168 to explain and demonstrate the severity of BOAS to pet owners and as a screening tool for
169 breeding to detect the most clinically affected dogs and thus be used to decrease the severity
170 of respiratory compromise that greatly impacts the quality of life of these popular pets (Packer
171 2013, Liu et al. 2015).

172

173 A limitation of this study was the small number of dogs included, and the fact that four dogs
174 did not attend their postoperative 6-MWT. This could have influenced results (as the owners
175 might have declined re-examination because patients were either doing very well or owners
176 may instead have been dissatisfied with the surgery), and decreased the power of the study. A
177 larger number of dogs would be needed to assess breed-related, gender-related or body
178 condition score-related changes in the 6-MWT and its correlation with anatomical
179 abnormalities and clinical grade.

180

181 Although each patient acted as its own control and the test was designed to be repeatable and
182 met the recommended guidelines (Enright 2003), the breed, temperament, body condition
183 score, concurrent undetected conditions or overall fitness level of each dog could influence
184 results of the 6-MWT. Patients might refuse to walk, some might perform better with repeated
185 testing (although in our study the 6-MWT was performed 6 weeks apart), and some might have
186 occult orthopaedic or cardiac disease affecting their exercise capacity. Daily intra-patient
187 variability has not been assessed for the 6-MWT.

188

189 **Acknowledgements**

190 The authors would like to acknowledge Yu-Mei Ruby Chang for her help with the statistical
191 analysis.

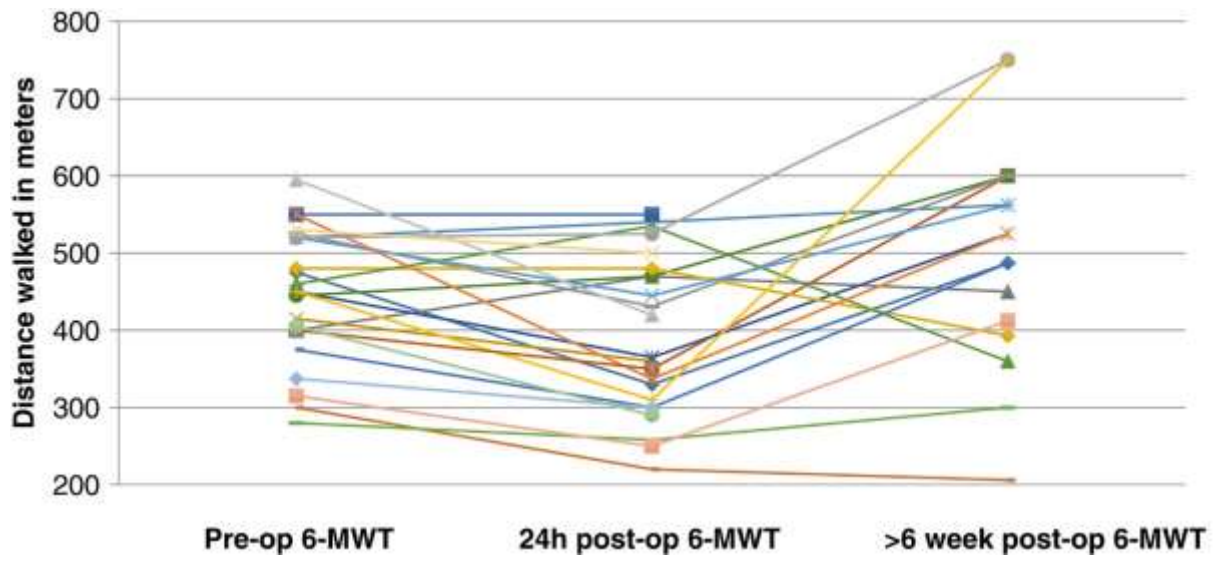
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- 250

251 **Figure 1.** Distance in metres walked by each dog for the 6-minute walk test at the different
252 time points



253
254
255

256 **Table 1.** Nose and oropharyngeal examination findings including degree of stenotic nares,
 257 elongation of the soft palate, laryngeal collapse and nasopharyngeal turbinate protrusion
 258

	Not recorded	Mild	Moderate	Marked	
Stenotic nares	3	3/21 (14.3%)	7/21 (33.3%)	11/21 (52.4%)	
Elongation of the soft palate	4	0	7/20 (35%)	13/20 (65%)	
	Not recorded	Grade I	Grade II	Grade III	
Laryngeal collapse	3	3/21 (14.3%)	16/21 (76.2%)	2/21 (9.5%)	
	Not recorded	Grade I	Grade II	Grade III	Grade IV
Nasopharyngeal turbinates	3	7/21 (33.3%)	6/21 (28.6%)	4/21 (19%)	3/21 (14.3%)

259

260

261 **Table 2.** Means (\pm sd) for heart rate, respiratory rate, SpO₂ and rectal temperature for all times
 262 points for each 6-MWT. Means (\pm sd) and range for distance walked for each 6-MWT

		Preoperative	24-hour postoperative	>6 weeks postoperative
Heart rate (mean beats per minute)	Pre	107 (\pm 22)	93 (\pm 18)	100 (\pm 19)
	Immediately post	126 (\pm 25)	111 (\pm 25)	115 (\pm 26)
	2 min post	117 (\pm 21)	103 (\pm 24)	111 (\pm 30)
	5 min post	109 (\pm 21)	96 (\pm 20)	99 (\pm 14)
Respiratory rate (mean breaths per minute)	Pre	48 (\pm 27) *	52 (\pm 46)	35 (\pm 15)*
	Immediately post	82 (\pm 48)	70 (\pm 39)	103 (\pm 56)
	2 min post	80 (\pm 53)	72 (\pm 51)	102 (\pm 54)
	5 min post	68 (\pm 45)	70 (\pm 46)	82 (\pm 43)
SpO ₂ (mean %)	Pre	94 (\pm 3) *	94 (\pm 5)	96 (\pm 2)*
	Immediately post	93 (\pm 6) *	95 (\pm 4)	95 (\pm 3)*
	2 min post	93 (\pm 5) *	95 (\pm 3)	97 (\pm 1)*
	5 min post	95 (\pm 3) *	96 (\pm 3)	97 (\pm 2)*
Temperature (mean degrees Celsius)	Pre	38.2 (\pm 0.5)	37.7 (\pm 0.5)	38.4 (\pm 0.5)
	Post	38.6 (\pm 0.5)	38.1 (\pm 0.7)	39.0 (\pm 0.5)
Distance walked in metres		446 (\pm 85) * (280 to 595)	391 (\pm 101) (220 to 550)	504 (\pm 144)* (206 to 750)

263 * Statistical significance between preoperative and >6 weeks postoperative