

1   **An endoscopic method for semi-quantitatively measuring internal pyloric diameter in  
2   healthy cats: a prospective study of 24 cases.**

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25     **Abstract**

26     The objectives of this study were to describe an endoscopic technique for semi-quantitative  
27     measurement of the internal pyloric diameter and apply this method to determine its typical size  
28     in a population of healthy cats.

29     Twenty-four healthy adult cats, privately owned or originating from catteries, were prospectively  
30     recruited. Endoscopies were performed by the same investigator and cats with moderate to  
31     marked macroscopic inflammation were excluded. The internal pyloric diameter was measured  
32     with bespoke interchangeable biocompatible ‘olives’ (ranging from 4 to 12 mm in diameter) that  
33     could be attached to a guidewire. Attempts were made to pass the olives through the pylorus, in  
34     decreasing order of size, and the internal pyloric diameter was assumed to be equivalent to the  
35     size of the first olive that could successfully be passed.

36     The median duration of the endoscopic procedure was less than 5 (interquartile range 2.7-5.4)  
37     minutes and all cats recovered quickly from the procedure without any complications. The  
38     median internal pyloric diameter in this population was 9 (interquartile range 9-10) mm, with  
39     most (23/24) cats having an internal pyloric diameter within  $\pm 1$  mm of this measurement. There  
40     was no apparent effect of age, sex, breed or weight on the pyloric size.

41     This study is the first to describe a quick and safe method for semi-quantitatively assessing the  
42     internal pyloric diameter in healthy adult cats. A prospective study is now warranted in order to  
43     determine the impact of gastrointestinal disease on pyloric diameter, for example cats with  
44     possible pyloric stenosis.

45

46     **Keywords:** pyloric stenosis, gastric emptying, endoscopy, feline, pylorus, vomiting

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48

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60

61 **Introduction**

62 The pylorus plays an important role in digestion, acting as a valve to regulate the flow of  
63 partially-digested food (Simpson, 2013). Gastric outlet obstruction is a syndrome where there is  
64 an inability of gastric contents (food or water) to exit the stomach, usually arising from a  
65 mechanical blockage at or near the pylorus (Hall, 2013). Stenosis of the pyloric canal is one of  
66 the most commonly reported causes of gastric outlet obstruction in dogs and humans, but is  
67 poorly-documented in cats.

68

69 Histologically, the layers of the gastric wall are organized as in all parts of the digestive tract with  
70 a mucosa, submucosa, muscularis and serosa (Simpson, 2013); however, in the distal part of the  
71 stomach, the inner circular layer of the muscularis is ring-shaped at the pylorus and thickens to  
72 form the sphincter pylori muscle (Couturier et al., 2012). The evaluation of the internal pyloric

73 diameter in live animals is difficult, and is mainly subjectively evaluated by whether or not it can  
74 be successfully intubated by gastroscope (Tams, 2011). The normal ultrasonographic appearance  
75 of the pylorus has also been described, with the annular thickening of the muscularis layer  
76 appearing as a hypoechoic triangle in longitudinal section (Couturier et al., 2012).  
77 Ultrasonographic evaluation of the pylorus enables evaluation of a thickening of individual layers  
78 of the gastric wall, but the evaluation of the internal pyloric diameter is not possible and, to the  
79 authors' knowledge, a reference range has never been established for healthy cats.

80

81 Currently, the diagnosis of pyloric stenosis (PS) is based on clinical signs of pyloric outflow  
82 obstruction, mainly chronic food vomiting delayed after eating (a few hours) which may even be  
83 projectile (Hall, 2013); and on imaging signs of delayed gastric emptying and abnormal pyloric  
84 conformation, detected by ultrasonography and characterized by severe fluid distension of the  
85 stomach and circumferentially thickening of the pylorus, respectively (Syrkle et al., 2013). In a  
86 retrospective study, we recently described 34 cats with chronic alimentary vomiting and in which  
87 the pylorus was not passed with an 8.6 mm endoscope (Freiche et al., 2016). The pylorus  
88 appeared to be abnormally small on direct examination and a presumptive diagnosis of acquired  
89 PS was made. Although, a reliable method of diagnosis is needed to establish a definitive  
90 diagnosis of PS in cats. The first aim of this study was to describe an endoscopic procedure  
91 allowing the measurement of the internal pyloric diameter in cats. Having done this, a second  
92 aim was to create a reference range for this measurement in healthy cats.

93

94 **Materials and methods**

95 **Study population**

96 All procedures were approved by the Ethics Committee of the National Veterinary School of  
97 Alfort (agreement 2015-10-27) and informed owner consent was obtained. Healthy cats were  
98 prospectively recruited from November 2015 to July 2016 at the teaching hospital of Ecole  
99 Nationale Vétérinaire d'Alfort. Study cats were owned by the staff or the students from the  
100 veterinary school or originated from a colony of reproductive laboratory cats and a colony of  
101 specific-pathogen-free (SPF) laboratory cats. Cats were eligible for inclusion if they were  
102 between 1 and 8 years old and if their body weight was between 2.5 kg and 5.5 kg. Further, there  
103 could be no history of chronic vomiting, no signs of vomiting in the last four weeks, and had not  
104 received any medication in the previous 3 months, except for deworming and insecticides. The  
105 signalment of each cat was also reviewed (for age, breed, diet and deworming history), a physical  
106 examination was performed, and a blood biochemistry panel was performed; cats were not  
107 included if any abnormalities were identified on these tests or if it was decided that anaesthesia  
108 was contraindicated. Finally, cats were also excluded from the final study population if there was  
109 macroscopic evidence of possible gastric lesions during endoscopy (see below).

110

### 111 **Anaesthesia and endoscopic evaluation**

112 Cats were fasted for 18 hours prior to the procedure, to ensure that the stomach was empty.  
113 Water intake was possible until 12 hours before the procedure. General anaesthesia was induced  
114 with propofol (4 mg/kg, IV), cats were then intubated and isoflurane gas anaesthesia was used  
115 (1.6 L/min). Cats were positioned in left lateral recumbency, as previously recommended (Tams,  
116 2011), and underwent a gastroscopy with an 8.6 mm diameter insertion tube (Video endoscope  
117 GIF 160, Olympus, France) performed by the same clinician (VF). The internal macroscopic  
118 appearance of the oesophagus and stomach was evaluated according to recommendations from  
119 the World Small Animal Veterinary Association (WSAVA) Gastrointestinal Standardization

120 Group (Washabau et al., 2010). Even without any clinical signs, cats were immediately excluded  
121 from the study, and their endoscopic procedure terminated, if macroscopic gastric hyperaemia  
122 was scored as 2/3 or 3/3 or if others macroscopic gastric lesions were observed, according to the  
123 WSAVA endoscopic examination report form (<http://www.wsava.org/guidelines/gastrointestinal-guidelines>). No other procedures were undertaken during the procedure including collection of  
124 mucosal biopsy samples. One cat underwent surgery for bone fracture repair following the  
125 endoscopic procedure. All other cats were anaesthetised for the sole purpose of the planned  
126 endoscopic procedure.  
127

128

### 129 **Measurement of internal pyloric diameter**

130 To measure the internal pyloric diameter, interchangeable biocompatible resin ‘olives’ of  
131 diameter ranging from 4 to 12 mm were designed and manufactured for this study (Figure 1).  
132 Olives could be screwed onto a specific endoscopic guidewire that had been passed through the  
133 working channel of the video endoscope (Figure 2). To determine the internal pyloric diameter,  
134 attempts were made to pass the olives through the pylorus, in decreasing order of size (i.e. largest  
135 to smallest). To do this, the gastroscope and olive were carefully passed through the oesophagus  
136 and stomach to the pylorus (Figure 3). The olive was then placed in front of the pylorus and a  
137 gentle pressure was applied to pass the olive through (Figure 4). If it was not possible to pass the  
138 olive through the pylorus, the endoscope and the olive were gently removed, and the procedure  
139 repeated sequentially with smaller olives (i.e. 1 mm smaller each time), until one of the olives  
140 could be passed. Once this was done, the olive was then carefully withdrawn, and the cat  
141 allowed to recover from the procedure. Therefore, the internal pyloric diameter was assumed to  
142 be equivalent to the diameter of the first olive that could be passed into the duodenum.

143

144 **Data collection and statistical analysis**

145 The parameters recorded for the study included the diameter of the first olive that could be  
146 passed, the number of manipulations, the duration of the endoscopic procedure and macroscopic  
147 gastric inflammation according to WSAVA criteria, as mentioned above. Basic descriptive  
148 statistical analyses were performed using Microsoft Excel. Medians and Interquartile ranges  
149 (IQR; first and third quartiles) were provided for all quantitative variables. Because of the small  
150 number of cats, two distinct groups were considered in order to test the influence of  
151 epidemiological data on the internal pyloric diameter. One group “small pylorus” was defined by  
152 the smallest pylorus diameter measured (8 or 9 mm), while the other group “large pylorus” was  
153 defined by the largest pylorus diameter measured (10 or 11 mm). Age, sex, weight and breed  
154 were compared between the small and large pylorus groups by univariate logistic regression  
155 analysis, using a software program (XLSTAT 2017 software, Addinsoft, Paris, France). Odds  
156 ratio and 95% confidence intervals were calculated for variables. Statistical significance was set  
157 as  $P<0.05$  using 2-sided analyses.

158

159 **Results**

160 **Study population**

161 Twenty-seven clinically healthy cats were initially included in the study, but three cats were  
162 excluded after the endoscopic procedure due to macroscopic gastric hyperaemia graded at 2/3  
163 according to the WSAVA criteria (Washabau et al., 2010), leaving 24 cats whose internal pyloric  
164 diameter was measured and used in data analysis. The age of three cats was not known, although  
165 they were determined to be young adult cats based on clinical examination. For the remaining  
166 cats, median (IQR) age was 25 (16-66) months. Median (IQR) weight in the 24 cats included  
167 was 3.8 (3.2-4.5) kg (8.4 [7.1-9.9] lb). There were 11 males (1 neutered) and 13 females (3

168 neutered), with breeds represented including Domestic Shorthair (15/24), Siamese cross (8/24)  
169 and Bengal (1/24). All eight mixed Siamese cats were part of a reproductive laboratory cattery  
170 and all were entire males, whilst all entire females were SPF cats experimentally infected with  
171 toxoplasmosis a few months previously (as part of a separate trial), but free from any clinical  
172 signs at the time the current study was conducted. Three cats had a history of bone fracture, 1 cat  
173 had a history of previous calicivirus infection; 3 cats presented some moderate lesions of  
174 gingivitis and stomatitis (all from the catteries). Only three cats had recently been dewormed; the  
175 others lived strictly indoors. All were eating a veterinary dry diet.

176

### 177 **Endoscopic findings**

178 The oesophagus and the stomach were evaluated in each cat for macroscopic lesions. Five cats  
179 had mild (1/3) gastric hyperaemia without any other significant lesion. No mucosal abnormality  
180 was observed in the other 19 cats. The median (IQR) duration of the endoscopic procedure was  
181 4.9 (2.7-5.4) minutes and the median (IQR) number of manipulations was 4 (3-4). No adverse  
182 event occurred during the procedure for any cat. All cats recovered from anaesthesia within  
183 minutes following the end of the procedure without any complication.

184

### 185 **Internal pyloric diameter measurement**

186 The internal pyloric diameter was 8 mm in three cats, 9 mm in twelve cats, 10 mm in eight cats  
187 and 11 mm in one cat. The median (IQR) internal pyloric diameter was 9 (9-10) mm, with 23 of  
188 24 cats (96%) having an internal pyloric diameter of  $9 \pm 1$  mm. Fifteen cats were included in the  
189 group of “small pylorus”, as described above, with an internal pyloric diameter of 8 or 9 mm; and  
190 nine cats were included in the group of “large pylorus” with an internal pyloric diameter of 10 or

191 11 mm. Univariate logistic regression did not show significantly increased odds of having a  
192 smaller pylorus with all variables tested (Table).

193

194

195 **Discussion**

196 In this population of 24 healthy cats, the internal pyloric diameter was easily measured by  
197 endoscopy by an experienced endoscopic manipulator. The technique developed for the study  
198 was safe and quick, with results suggesting a median internal pyloric diameter of 9 mm, with  
199 most cats being within  $\pm$  1 mm of this measurement. With further validation, this approach could  
200 be used in the future as a means of identifying and classifying PS.

201

202 Congenital PS is a rare condition in cats with only a few case reports and case series published  
203 (Pearson et al., 1974; Syrcle et al., 2013; Twaddle, 1970, 1971). Of the 19 cases described in the  
204 veterinary literature, sixteen were young Siamese cats and a predisposition of this breed has been  
205 assumed. Furthermore, Twaddle described two cases of feline congenital PS in litter sisters born  
206 to a dam who had herself a PS diagnosed and an inherited pattern was suspected (Twaddle,  
207 1971). In one case report, histological and immunohistochemical findings were consistent with a  
208 diagnosis of hypertrophy of the tunica muscularis (Syrcle et al., 2013).

209 In a retrospective study performed by our team, 34 cats underwent upper gastrointestinal  
210 endoscopy as part of chronic alimentary vomiting investigation (Freiche et al., 2016). In these  
211 cases, a presumptive diagnosis of benign acquired PS was made on the basis that the pylorus was  
212 abnormally small on direct examination and could not be intubated with an 8.6 mm gastroscope,  
213 which should be possible in healthy cats when performed by an experienced clinician (Tams,

214 2011). However, only a presumptive diagnosis was established because, for now, no  
215 complementary exam allowing a definitive diagnosis of PS has been validated.

216

217 In fact, the main thing that this study highlighted is the fact that the exact size of the feline  
218 internal pyloric diameter was not known. As a result, we thought that the first step to document  
219 benign acquired PS would be to evaluate the internal pyloric diameter in healthy cats. To achieve  
220 this, we developed a new endoscopic technique using bespoke measurement devices with an  
221 olive shape. The shape of the devices was deliberately chosen to have rounded edges so as to  
222 minimize the risk of trauma during insertion, whilst the rest of the device was of constant  
223 diameter in order to avoid any risk of bougienage during the passage of the olive through the  
224 pylorus. The procedure was easily performed by an experienced clinician in endoscopy and was  
225 of short duration with a median duration time less than 5 minutes. Soft pressure was applied and  
226 the operator waited for 15 seconds to minimize the possibility that pyloric tone (rather than the  
227 internal pyloric diameter) was causing any resistance to passage of the olive. Therefore, while  
228 pyloric tone might occasionally have been responsible, we believe that its effect was minimized.  
229 No adverse event occurred during any procedure and all cats recovered well in the minutes  
230 following the end of the manipulations. The anaesthetic protocol used in this study was chosen in  
231 order to avoid the use of any drug known to affect pyloric tone or function (Smith et al., 2004).

232

233 Cats included in the study were determined to be healthy based on history, clinical examination  
234 and biochemistry. Most of them had an internal pyloric diameter of  $9 \pm 1$  mm, consistent with the  
235 suggestion that the pylorus is supposed to be easily passable in cats with a flexible gastroscope of  
236 8.6 mm (Tams, 2011). However, 3/24 cats in our study had an internal pyloric diameter of 8 mm  
237 despite no clinical signs of pyloric outflow obstruction. Rather than suggesting subclinical PS, it

238 is possible that, for some cats, a narrower pyloric diameter is a normal finding. Further work  
239 involving more cats would be required to confirm normal pyloric diameter. A second  
240 observation of note was that moderate to severe macroscopic gastric hyperaemia was evident in 3  
241 cats (which were excluded), even though they were apparently healthy with no evidence of  
242 clinical signs including vomiting. Given that the internal pyloric diameter was not measured, it is  
243 unclear as to whether or not the pyloric size was different. Further, whilst these cats might have  
244 had a subclinical gastropathy, no biopsies were taken to confirm this hypothesis.

245

246 Siamese cats are reportedly predisposed both to congenital and acquired PS and, as a result, we  
247 expected that the Siamese-cross cats in our study would have a smaller internal pyloric diameter  
248 (Freiche et al., 2016; Pearson et al., 1974; Twaddle, 1970, 1971). However, there was no  
249 significant difference in breed between cats with different pyloric sizes and, in fact, most  
250 Siamese-cross cats had an internal pyloric diameter of 10 mm. This might suggest that the  
251 predisposition of this breed to PS is not due to the fact that the pyloric diameter is inherently  
252 smaller in this breed. That said, the cats included here were mixed Siamese and so the findings  
253 might have been different if a purebred population of Siamese cats had been studied. Therefore,  
254 a further study is required to determine the normal internal pyloric diameter of cats of different  
255 breeds.

256

257 In humans, infantile hypertrophic PS is a well-known condition of newborns and infants and is  
258 the most common condition requiring surgery in infants (Zhu et al., 2017). It is characterized by  
259 acquired narrowing of the pylorus secondary to hypertrophy of the pyloric musculature (Peters et  
260 al., 2014). Its aetiology is largely unknown but a genetic background is suspected and  
261 environmental factors as well as perinatal factors are also likely to play an important role (Peters

262 et al., 2014; Zhu et al., 2017). In contrast, the adult form of idiopathic hypertrophic PS is a rare  
263 disorder in humans and only over 200 cases have been described in the medical literature  
264 (Gurvits et al., 2013). In contrast, two forms of benign PS have been described in dogs (Hall,  
265 2013). The congenital form is the least common, most notably in brachycephalic breeds and there  
266 are similarities with infantile hypertrophic PS, with selective hypertrophy of the muscularis of the  
267 pylorus (Hall, 2013). The acquired form of the disease is more common with the stenosis being  
268 due to mucosal hypertrophy in addition to a thickened muscular layer (Hall, 2013). Male dogs  
269 and those of smaller breeds (Lhasa Apso, Shih Tzu, Pekingese, Maltese) are overrepresented  
270 (Bellenger et al., 1990). In cats, ultrasonographic and endoscopic findings suggest that the  
271 acquired form of PS predominantly affects the mucosa (unpublished data). However, full-  
272 thickness biopsies have never been performed in cases of acquired PS in cats to confirm these  
273 findings.

274

275 In children, ultrasonography is the modality of choice for the diagnosis of hypertrophic PS and  
276 the main diagnostic criterion is measurement of the thickness of the muscular layer with the use  
277 of an abnormal cut off value of 3 mm in thickness (Costa Dias et al., 2012; Peters et al., 2014).  
278 Other criteria include abnormal elongation of the canal (greater than 12 or 15 mm in length) and a  
279 markedly distended stomach with active peristalsis. However, these criteria are difficult to use in  
280 cats because pyloric wall thickness and thickness of the pyloric muscularis vary widely between  
281 studies and because several other diseases can affect the pyloric antrum in adult cats, such as  
282 neoplasia, benign polyps, ulcers and eosinophilic sclerosing fibroplasia (Couturier et al., 2012;  
283 Goggin et al., 2000). In fact, ultrasonography is useful to identify a thickening of one layer or a  
284 thickened gastric wall but the internal pyloric diameter cannot be evaluated.

285

286 Until now, the presumptive diagnosis of PS in cats was based on the association of clinical  
287 findings, radiographic and/or ultrasonographic findings and endoscopic appearance with  
288 exclusion of other causes by histologic analysis of multiple biopsies (Syrkle et al., 2013).  
289 Knowing the size of the internal pyloric diameter in healthy cat will now allow a more accurate  
290 endoscopic evaluation of cats with a suspicion of PS. For comparison, in humans, the size of the  
291 internal pyloric diameter has been estimated between 1.2 and 1.5 cm in adult patients, as  
292 measured using the index finger during gastric surgery (Maylard, 1904). The size of the internal  
293 pyloric diameter has not yet been determined precisely in dogs.

294

295 The current study has several limitations that should be considered. First, the repeatability and  
296 reproducibility of this technique was not assessed for ethical reasons, for example to avoid  
297 prolonged and repeated anaesthesia. However, the procedure was easy to perform and most  
298 clinicians experienced in endoscopy should be able to perform the same procedure. Second, only  
299 a small number of measurement devices were available, meaning that there was a limitation in  
300 the sizes that could be assessed, and measurements could only be made to the millimetre.  
301 However, whilst more devices could have been created (for example differing in size by 0.2-0.5  
302 mm), this would have increased the number of manipulations and the duration of the procedure  
303 overall. A third study limitation was the fact that mucosal biopsies were not collected during the  
304 procedure, meaning that the presence of gastric disease (even subclinical) could not be  
305 determined. In fact, gross findings frequently do not correlate with histopathological results in  
306 the digestive tract (Hall and Day, 2017). That being said, all cats were healthy without any signs  
307 of gastrointestinal disease. However, we thought that adding biopsies at the design of this study  
308 would have make the study more invasive, even if the risk of complication would have still been  
309 low. A fourth study limitation was that ten cats were SPF cats experimentally infected with

310 toxoplasmosis in a previous study and the effect that this might have had on the measurements  
311 taken is not known. However, all of these cats were clinically healthy and did not excrete oocysts  
312 at the time of the study. A fifth limitation was that the number of cats included was small and the  
313 absence of statistical significant differences between groups could be due to the fact that the  
314 study was underpowered. Finally, since there were many Siamese cats and intact males and  
315 females, the cats studied might not be totally representative of cats in the general pet population.  
316 Further studies should be considered involving more cats, with a wider range of ages and breeds.  
317 Cats with gastrointestinal signs, including those with signs of delayed gastric emptying, should  
318 also prospectively be evaluated to determine how the internal pyloric diameter changes with  
319 disease. Finally, it would be useful to assess the performance of this technique in young kittens  
320 to determine if it might be useful for the diagnosis of congenital PS, and also adapted to dogs.

321

322 **Conclusion**

323 Pyloric diameter can be easily measured in cats by endoscopy using specially-designed olive-  
324 shaped measurement devices. This endoscopic technique was safe and of short duration. In this  
325 study, based on 24 healthy cats, the median diameter of the pylorus was measured at 9 mm with  
326 96 % of cats having an internal pyloric diameter within 1 mm of this, using propofol and  
327 isoflurane for anaesthesia. A prospective study is now warranted in cats with chronic delayed  
328 food vomiting and with a high suspicion of benign PS to compare their internal pyloric diameter  
329 to the group of cats described in the present study.

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