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1 **Clinicopathological Findings, Treatment and Outcome in 60 cats with Feline**
2 **Gastrointestinal Eosinophilic Sclerosing Fibroplasia**

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15
16 **Background** – Feline gastrointestinal eosinophilic sclerosing fibroplasia (FGESF) presents as
17 mass(es) associated with the gastrointestinal tract, mesentery, and abdominal lymph nodes.

18 **Hypothesis/Objectives** – To report the clinicopathological findings, treatment, and outcome of cats
19 with FGESF.

20 **Animals** – Sixty client-owned cats diagnosed with FGESF.

21 **Methods** – Retrospective review of medical records of cats with histopathologically confirmed
22 FGESF.

23 **Results** – The median age was 5.4 years (interquartile range [IQR] 3.3-8.9.); 30% were Domestic
24 Shorthairs and 12% were Domestic Longhair cats, with the most prevalent pedigree breeds being
25 Ragdolls (25%), Exotic Shorthair (10%) and Persian (8%) cats. The median duration of clinical
26 signs was 90 days (IQR 17.5-247.0); the most common clinical signs were weight loss (60%),
27 hyporexia/anorexia (55%), chronic vomiting (37%), lethargy (35%) and chronic diarrhea (27%).
28 Masses were located in the small intestine (32%), stomach (27%), ileocolic junction (15%), colon
29 (10%), lymph node (8%) and mesentery (8%) and 15% of cats had more than one mass.

30 Eosinophilia was present in 50% and hypoalbuminemia in 28% of cats. The mass was removed
31 surgically in 37% of cases. Most cats (98%) were treated with corticosteroids and 1 cat with
32 antibiotics alone. The survival was not statistically different between cats treated with surgical
33 resection and cats treated with medical therapy alone, 88% of the cats still alive at the time of
34 writing.

35

36 Conclusions and clinical importance – FGESF is an important differential diagnosis for
37 abdominal masses in cats, and has a much better prognosis than previously reported.

38

39 **Keywords:** eosinophilia, gastrointestinal mass, mesenteric mass, Ragdoll

40

41 **Introduction**

42 Feline gastrointestinal eosinophilic sclerosing fibroplasia (FGESF) is a recently described disease
43 in cats that presents as eosinophilic mass(es) that are associated with the gastrointestinal tract and
44 associated abdominal lymph nodes, most commonly near the pylorus or ileocolic junction (1, 2).
45 There have also been two case reports where FGESF was localized to the mesentery or
46 retroperitoneum, and even presented as a cavitated mass (3, 4). A recent case report also described
47 the same type of pathology outside of the abdominal cavity; a feline eosinophilic sclerosing
48 lymphadenitis in medial retropharyngeal lymph node that was associated with *Pseudomonas*
49 *aeruginosa* infection (5). Feline gastrointestinal eosinophilic sclerosing fibroplasia is likely
50 underdiagnosed because these mass lesions can be misinterpreted as lymphoma, granuloma,
51 fibrosarcoma, adenocarcinoma, and mast cell tumor, and the histopathological diagnosis can be
52 challenging (2, 6-8). That said, a recent study suggested that immunohistochemical staining (*e.g.*,
53 for transforming growth factor β 1) can aid diagnosis (9). This disease is most commonly seen in
54 middle aged and male cats of all breeds, with Ragdolls being overrepresented in previous studies,
55 and Maine Coons, Persians, Exotic Shorthairs, Bengal and Scottish fold cats also being reported
56 (1, 2, 10-12). The pathogenesis of FGESF is still poorly understood; however, an aberrant response
57 to antigens from bacteria or parasites has been considered, and a case of FGESF associated with
58 fungal organisms (*Phycomycetes*) has also been reported (1, 13, 14). With some breeds, such as
59 Ragdolls, being overrepresented, a genetic predisposition could be considered.

60

61
62 The most common presenting signs of cats with FGESF are chronic vomiting and/or diarrhea,
63 followed by weight loss, lethargy; less commonly an acute onset of vomiting and/or diarrhea has
64 been reported (1, 2). A palpable intestinal mass has been reported in 85-100% of cases, with
65 abdominal pain and pyrexia being less common (1, 2). Surgical removal of the mass has been
66 performed in most cases; however, several studies have also reported medical management with
67 corticosteroids, cyclosporine and/or mycophenolate (2, 15-17). A mass in a second location has
68 been reported to develop in some of the cases, after surgical removal of the initial one (18, 19).

69
70 The prognosis varies between studies; however, no large studies on prognosis or response to
71 treatment have been reported (2). The objective of this study was to retrospectively evaluate a
72 large number of cats with FGESF, including their presentation, diagnosis, treatment and outcome.

73 74 **Materials and methods**

75 *Case recruitment*

76 This is a retrospective, multi-centric study, of cases of FGESF which have been collected by
77 several veterinary hospitals around the world (USA, United Kingdom and Japan) by contacting
78 veterinarians that have previously seen cases of FGESF between 2010 and 2022. Inclusion criteria
79 was confirmation of an FGESF diagnosis by histopathology after surgical removal or biopsy of
80 the mass; histopathology was performed by different pathologist from referral hospitals or referral
81 laboratories. Patient signalment, clinical signs, physical findings, clinicopathological results,
82 surgical reports and medical management were tabulated in an Excel (Microsoft, Redmond, WA,
83 USA) spreadsheet. Ethical approval was gained from University of Edinburgh (VERC Reference:
84 17.22).

85 86 *Statistical analysis*

87 Survival times were measured from the date of presentation until the date of death or last follow-
88 up. Kaplan-Meier analysis and log rank tests were used for survival analysis in order to evaluate
89 the association of survival time with treatment (GraphPad Prism 9, GraphPad Software, Boston,
90 MA, USA). Results were considered significant if $P < 0.05$.

91

92 **Results**

93 *Presentation and clinical signs*

94 A total of 60 cats met the inclusion criteria for the study. The median age was 5.4 years
95 (interquartile range [IQR] 3.3-8.9.). Of the 60 cats, 18 (30%) were Domestic Shorthair, 7 (12%)
96 Domestic Longhair cats, and 35 (58%) were pedigree breeds: Ragdolls 15/60 (25%), Exotic
97 Shorthair 6/60 (10%), Persian 5/60 (8%), Maine Coon 3/60 (5%), Sacred Birman 2/60 (3%),
98 American Shorthair 1/60 (2%), Bengal (1/60 (2%), Bobtail 1/60 (2%) and British Shorthair 1/60
99 (2%). Of the 60 cats, 34 (57%) of the cats were neutered males, 25 (42%) were spayed females
100 and 1 (2%) entire female.

101 The median duration of clinical signs was 90 days (IQR 17.5-247.0) with most cats showing
102 median of 3 (IQR 2-4) clinical signs. The most common clinical signs are reported in Table 1.

103 On physical examination, the most common abnormality was a palpable abdominal mass in 35
104 (58%) of the 60 cats, followed by pyrexia, in 9/60 (15%), dehydration, 7/60 (12%) and abdominal
105 pain in 4/60 (7%). In 16 (26%) of the 60 cats, the body condition score was reported as less than
106 ideal (<4/9).

Clinical sign	Number of cats (%)
Weight loss	36/60 (60)
Hyporexia/anorexia	33/60 (55)
Chronic (>2 weeks) vomiting	22/60 (37)
Lethargy	21/60 (35)
Chronic diarrhea	16/60 (27)
Acute (<2 weeks) vomiting	8/60 (13)
Acute diarrhea	6/60 (10)
Constipation	6/60 (10)
Tenesmus	5/60 (8)
Polyphagia	4/60 (7)

Hematochezia	4/60 (7)
Decreased grooming	3/60 (5)
Melena	1/60 (2)
Excessive grooming	1/60 (2)

107 **Table 1:** Presenting clinical signs of cats with feline gastrointestinal eosinophilic sclerosing
108 fibroplasia.

109

110 ***Clinopathological findings***

111 Complete blood cell count findings were available for 57 (95%) of the 60 cats. The most common
112 abnormalities were eosinophilia which was present in 30 (52%) of the 57 cases - the reference
113 intervals varied between the clinics and the eosinophilia was mostly moderate to severe with the
114 median percentage above RI 243.3 (IQR 188.7-465.2); however not all medical records contained
115 actual eosinophil numbers (some records only mentioned eosinophilia being present). Second most
116 common hematological abnormality was anemia in 16 (28%). Less common findings included
117 neutrophilia 10/57(18%), monocytosis 7/57 (12%), lymphocytosis 5/57 (9%), basophilia 3/57
118 (5%) and neutropenia 1/57 (2%). Serum biochemistry findings were available for 58 (97%) of the
119 60 cats. The most common abnormality was hypoalbuminemia which was seen in 16/58 (28%) of
120 cases. The reference intervals varied between the clinics and the hypoalbuminemia was mostly
121 mild with the median percentage below RI 91.3 (IQR 83.7-96.2). The second most common
122 abnormality was hyperglobulinemia seen in 8/58 (14%), followed by hypocholesterolemia in 6/58
123 (10%) and total hypocalcemia 6/58 (10%); with 4/6 (67%) hypocalcemic cats having normal
124 albumin levels. Serum cobalamin was measured in 11 (18%) of the 60 cats and was normal in all
125 of them, although it was at the low end of the reference interval in 1 cat (278; reference interval
126 [RI] 214-1106ng/L). Folate measurement was available in 8/60 cats and was abnormally high in
127 5/8 cats.

128 ***Diagnostic imaging***

129 Abdominal imaging was performed in all cats; however, ultrasound images were only available
130 for 30 (50%) of the 60 cats; the others had abdominal radiographs performed and/or ultrasound

131 reports was part of the medical records, but ultrasound images were not available for review. In
132 25/30 (83%) of the cats the mass originated from the stomach or intestines. Of the other 5 cases, 3
133 (10%) affected abdominal lymph nodes and 2 (7%) involved the mesentery (7%). The majority of
134 gastrointestinal masses were associated with loss of the intestinal layering (Figure 1), symmetrical
135 or asymmetrical circumferential thickening, eccentric growth and a heterogeneously mixed wall
136 echogenicity which had hyperechoic areas and possible ulceration. In 6 (20%) of the 30 cases,
137 these masses were reported to be associated with altered rather than lost layering. Hyperechoic
138 areas were noted in 84% of the gastric or intestinal cases, and 80% of all cases, while thickening
139 of the muscularis layer in the small intestine was seen in 33% of them. Peritoneal changes were
140 reported in 22/30 (73%) of the cats, of which 19 (86%) had hyperechoic peritoneum and 8 (36%)
141 had a peritoneal effusion (the amount of effusion was not always reported in the medical records).
142 None of the lesions showed ultrasonographic findings compatible with gastrointestinal perforation.
143 Enlarged lymph nodes were present in 27 (90%) of the 30 cases where ultrasound images were
144 available; the most commonly affected lymph nodes were ileocolic in 9/27 (33%), followed by
145 pancreaticoduodenal 8/27 (30%), and mesenteric, 8/27 (30%). For the rest of the cases (30 cats),
146 abdominal ultrasound images were not available to assess the lymph nodes further.

147 ***Location of the mass***

148 The most common location of the masses (Figure 2) was small intestine in 19 (32%) of the 60
149 cases, including the proximal duodenum 15/60 (25%; Figure 3), jejunum 2/60 (3%), ileum 2/60
150 (3%), and the stomach in 16/60 (27%), followed by the ileocolic junction (9/60; (15%), colon 6/60
151 (10%), lymph node 5/60 (8%) and mesentery in 5/60 (8%; Figure 4). Most of the cats, 51/60 (85%)
152 had only 1 mass; however, in 9/60 (15%) of cats a mass was present in more than 1 location. The
153 additional masses most commonly involved the mesentery and surrounding lymph nodes in 4/9
154 (44%); in 3 cats the additional masses affected the stomach and proximal duodenum, in 1 cat the
155 mesentery and jejunum, and in 1 cat the proximal duodenum and jejunum. Of note, 1 cat had an
156 eosinophilic skin mass at the same time as FGESF. One cat had a mass in the ileocolic junction
157 removed, then presented 7 months later with a mesenteric mass. Another cat had a mass removed
158 from its colon, then presented 2 years later with a pyloric mass, which was also removed, the re-
159 presented 3 years after that with another pyloric mass. In both of these cases, the cats were not
160 started on corticosteroid therapy until after surgical resection of the second or third mass,
161 respectively and all of the masses in both cats were consistent with FGESF on histopathology.

162 **Cytology**

163 Cytology of fine-needle aspirates (FNA) of the mass was performed in 22 (37%) of the 60 cats and
164 showed eosinophilic inflammation in 10/22 (45%) of cases. In other cases, the cytology was either
165 non diagnostic, or showed necrosis or mixed inflammation. Cytology on FNA of abdominal lymph
166 nodes was performed 22/60 (37%) of cats but was mostly non-diagnostic or showed reactive lymph
167 nodes; eosinophils were only reported in 8/22 (36%) of cases.

168 **Surgery/biopsy**

169 The mass was removed surgically in 22 (37%) of the 60 cats; with complete microscopic excision
170 achieved in 18/22 (82%) of the cats. In the remaining 29 (76%) of the 38 cases had a surgical
171 biopsy performed, while the diagnosis was achieved on endoscopic biopsies in the other 9 (24%)
172 cats. The cats that had endoscopic biopsies, the mass was located in proximal duodenum in 5/9
173 (56%) or stomach in 4/9 (44%) of the cases. Surgical complications were reported in 5/22 (22%)
174 of cats, with 3/5 of the cats developing anemia and requiring transfusion (14% of the cats that had
175 surgery to remove or biopsy a FGESF mass); all three of these cats were anemic on presentation
176 with HTC on presentation being 18, 20 and 26% respectively. One cat developed septic peritonitis
177 requiring a second surgery, 1 cat became anorexic, 1 cat developed chyloabdomen, which has
178 resolved with treatment, and 1 cat developed persistent fecal incontinent following surgical
179 resection of a colonic mass.

180 **Histopathology and culture**

181 In all 60 cases the mass was confirmed as FGESF on histopathology (Figure 5 and 6). Of the 22
182 (37%) out of 60 cases in which the mass was surgically removed, the lesion was completely
183 excised in 13 (59%) cats. In 19/60 (32%) of cats, intralesional bacteria were present on
184 histopathology, and fungal organisms were detected in 1 cat (by positive periodic acid-Schiff
185 (PAS) staining). Fluorescence *in situ* hybridization (FISH) was performed in 3 cats and showed
186 *Eubacteria* in 1 cat and *Eubacteria*, *Campylobacter jejuni*, *Salmonella* species and *Escherichia*
187 *coli* in the second cat and no invasive bacteria in the third cat. Bacterial culture was performed in
188 18/60 (30%) of cases; 4/18 (22%) were negative, while in the others the most common bacteria
189 were *E. coli* (6/12;50%), *Staphylococcus* species (6/12;50%), *Enterococcus* species (4/12;33%)
190 and *Streptococcus* species (1/12;8%) and *Bacteroides fragilis* (1/12;8%). It was not always clear
191 if effusion or mass or swab of the tissue was cultured in some cats. The biopsy that was positive

192 on PAS staining cultured *Candida albicans* as well as *Enterococcus* species and *E. coli*. In 34
193 (57%) of the 60 cases, additional organs were also biopsied; these included lymph nodes (28/34;
194 82%), stomach (5/34; 15%), liver (4/34;12%), duodenum (2/34;6%), jejunum (1/34;3%) and
195 omentum (1/34;3%). In 12 (42%) of the 28 cases where lymph node histopathology was performed
196 cases, results were consistent with FGESF, as was the omentum in the one case where this site was
197 biopsied.

198 ***Treatment***

199 Most cats 59/60 (98%) were treated with corticosteroids, although 1 cat was treated with antibiotics
200 alone. All except for 1 of the cats that were started on corticosteroids were started on prednisolone
201 once daily with a median dose of 1.5 mg/kg/day (IQR 1.0-2.0). The median time of cats to be
202 started on corticosteroids was 23 (IQR 10.5-49.0) days after first being presented. In 49 cats, the
203 prednisolone dose was changed on with a median of 32 (IQR 16.0-60.0) days after starting
204 corticosteroid therapy, and in 13 cats, the prednisolone was discontinued; however, in 11 of these
205 13 cats, prednisolone had to be restarted at a median time of 114 (IQR 36.0-366.0) days following
206 discontinuation due to recurrence of clinical signs. The median time to the lowest dose of
207 prednisolone was 369 (IQR 195.0-841.0) days, with a median lowest maintenance dose of 0.65
208 (IQR 0.40-0.90) mg/kg/day needed to control the clinical signs. The most common complications
209 from corticosteroid treatment were hypertriglyceridemia in 5 cats. Hypertriglyceridemia was
210 treated with a low-fat diet and fish oil in 1 cat, fish oil alone in 1 cat, with low fat diet, fish oil and
211 fenofibrate in 1 cat, and no treatment in the remaining 2 cats), and the development of diabetes
212 mellitus in 3 cats (2 of which went into diabetic remission on insulin therapy – one of these cats
213 still remain on low dose (0.4mg/kg/day of prednisolone). Secondary immunosuppressive agents
214 (cyclosporin or chlorambucil) were prescribed in 14 (23%) of the 60 cats and were discontinued
215 in 8 (57%) of those cats after a media of 80 (IQR 64.5-236.3) days.

216 Antibiotics were prescribed in 43 (72%) of the 60 cats, with the most common antibiotics being
217 penicillins (16/43;37%), fluoroquinolones (9/43; 26%), metronidazole (6/43;14%), cephalosporins
218 (5/43;12%) and clindamycin (2/43;4%). The average length of treatment with antibiotics was 34
219 days (range 7-204).

220 Hydrolyzed or selected protein diets were advised in 37% (22/60) of cats; 41% (9/22) of the owners
221 reported an improvement of clinical signs on hydrolyzed or selected protein diet, although 3 cats
222 would not eat the diet.

223 ***Survival***

224 As 53 (88%) of the 60 cats still being alive at the time of writing this publication, the median
225 survival time cannot be estimated. Of the 7 cats that died or were euthanized, 4 cats due to poorly
226 controlled FGESF, 2 cats of pancreatic neoplasia and 1 cat died of causes unknown. There was no
227 statistical difference between the survival of cats that had a surgical resection of the mass and cats
228 where the mass was biopsied only ($p=0.16$; Figure 7). There was no statistical difference between
229 the survival of cats that had a complete resection with clear margins confirmed by histopathology
230 and cats with incomplete resection of the mass and cats where the mass was biopsied only ($p=0.67$).
231 There was also no statistical difference between the survival of cats that were treated with
232 corticosteroids only vs secondary immunosuppressive agents ($p=0.41$), nor between cats that were
233 treated with antibiotics and cats that were not treated with antibiotics ($p=0.71$).

234

235 **Discussion**

236 This paper presents the largest study of FGESF to date; cases were collected internationally over
237 a 15-year period. Previous studies had reported the FGESF masses to be associated most
238 commonly with the stomach (often near the pylorus) or the intestines, and may also affect the
239 abdominal lymph nodes (1, 2), the mesentery, and retroperitoneum (3, 4). This is similar to the
240 current study, where the most common location of the FGESF masses were the small intestine,
241 stomach, ileocolic junction or colon, while in 16% of the cats the mass was associated with lymph
242 nodes or mesentery. In 15% of the cats, FGESF masses were present in more than 1 location; in
243 12 of the cases the local lymph nodes were also affected, and in 1 cat the omentum was involved,
244 showing that this disease can affect a number of locations in each cat. Of note, 1 cat also had an
245 eosinophilic skin mass at the same time as FGESF - this has not been previously reported in cats
246 with FGESF; however, subcutaneous masses with eosinophilic infiltration have been reported (20),
247 and a recent case report also described possible FGESF-like pathology outside the abdominal
248 cavity, in medial retropharyngeal lymph node (5).

249

250 When looking at signalment, the median age of cats with FGESF in this study was 5.4 years (range
251 1.3-14.5), which is similar to the previously reported median age of 7 years, with a range of 2-11
252 years (2). Previously, male cats were reported to be more affected by FGESF; however, this was
253 not seen in this study (2). More than half (58%) of the cats in this study were pedigree cats, with
254 Ragdolls comprising a quarter of the study population; this is similar to another study that
255 previously reported Ragdolls to be overrepresented (2). Other breeds commonly seen in the current
256 study were Exotic Shorthair (10%) and Persian (8%) cats. It is unclear why pedigree cats appear
257 to be predisposed to develop FGESF, notably Ragdolls (25%) and Persian/Exotic cats (18%), and
258 further studies, including genetic analysis, are needed to see if these breeds have a genetic
259 predisposition to develop eosinophilic inflammation as a response to enteric antigens which is the
260 likely cause of FGESF. It is important to note that these are also breeds predisposed to feline
261 infectious peritonitis (FIP) (21); FGESF and non-effusive FIP are both differential diagnoses of
262 note for cats presenting with abdominal masses.

263
264 In the current study, the median duration of clinical signs was 90 days with most cats showing
265 median of 3 clinical signs, with the most common being weight loss (60%), hyporexia/anorexia
266 (55%), chronic vomiting (37%), lethargy (35%) and chronic diarrhea (27%), which is very similar
267 to a previously reported study; however, they also reported excessive grooming in 50% of cases
268 which was only seen in 2% of the cats in the current study (2). Palpation of an abdominal mass
269 was present in 85-100% of the cats reported previously (1, 2); however, this was less common in
270 the current study where a mass was only palpable in 58% of the cats. The prevalence of pyrexia
271 was similar to other studies, 15% vs 18% (2).

272
273 The most common bloodwork abnormality was peripheral eosinophilia, which was present in 50%
274 of the cats in the current study, which is similar to previous studies (1, 2). Anemia was present in
275 almost third of the cats in the current study, but was not reported previously. In serum
276 biochemistry, hypoalbuminemia and hyperglobulinemia were the most common abnormalities,
277 occurring in 27% and 14% of cats, respectively, which is less common than the previously reported
278 45% and 67%, respectively (2).

279

280 Large studies evaluating abdominal ultrasonography findings of cats with FGESF are lacking to
281 date; however, a study did report 5 cats that had solitary mass with mural thickening and loss of
282 layering in the stomach, duodenum, jejunum and colon (18). In the current study, abdominal
283 ultrasound images were available for review in 50% of the cats, with most cases (83%) showing
284 that the majority of the masses originated from the stomach or intestines. These masses were
285 associated with loss of the intestinal layering and circumferential thickening in most cases,
286 although in 20% there was alteration of the layering rather than loss of it. Enlarged local lymph
287 nodes were present in 90% of the cases, and peritoneal changes in 73%, of which 36% had a
288 peritoneal effusion; however, none of the lesions showed ultrasonographic findings compatible
289 with gastrointestinal perforation.

290
291 Intralesional bacteria were identified in 56% of the cases overall (all of the ileocecolic junction
292 and colon lesions) in one study (1) and in 69% of cats in another study using either culture or
293 conventional light microscopy, special stains and FISH (2). In the current study, only 32% of the
294 cases had bacteria present on histopathology, and fungal organisms were detected in 1 cat;
295 however, as one limitation of this retrospective study, infectious organisms might have been
296 missed in some cats as FISH was only performed in 3 cats and bacterial culture was performed in
297 only 30% of the cats in this study. Even though bacteria are commonly associated with FGESF,
298 fungal organisms have only been reported once previously, in a case report of FGESF associated
299 with phycomycetes (13).

300
301 The prognosis for cats with FGESF has been reported as variable in previous publications, varying
302 from guarded, to cats living for several years (1, 2, 18). Linton *et al.* reported that most cats
303 surviving the perioperative period remained well for several years (2). In the current study, the
304 median survival time could not be estimated as 88% of the cats still alive at the time of writing this
305 publication. This shows the importance of the correct diagnosis for cats with FGESF, as many of
306 these masses can be misdiagnosed as neoplasia, which usually carries a poor prognosis.

307 It has been reported that cats being treated with surgery alone had a significantly shorter survival
308 time than those cats treated with surgery and corticosteroids (1). Improved survival time was
309 reported when prednisolone was included in the therapeutic regimen in another study, regardless

310 of whether or not they also had surgery (2). In the current study, 98% of the cats were started on
311 corticosteroid therapy, so it is not possible to assess the survival time of the cats with surgery
312 alone; however, there was no statistical difference between the survival of the cats that had their
313 masses surgically resected and cats where their mass were only biopsied including cats with
314 complete resection with clear margins confirmed by histopathology. In the future, a prospective
315 randomized study would be needed to determine the most appropriate therapy for cats with
316 FGESF.

317 Corticosteroids appear to be important in the treatment of cats with FGESF. Re-occurrence of
318 masses has been previously reported when surgery was not followed by corticosteroids (18, 19).
319 In the current study, 1 cat had a mass resected from the ileocolic junction but was not started on
320 corticosteroid therapy, and re-presented 7 months later with a mesenteric mass. Another cat in this
321 study was diagnosed with FGESF in the colon, which was resected, then with a pyloric mass 2
322 years later, which was also resected, and another pyloric mass 3 years after that; while all of these
323 FGESF masses were surgically resected, corticosteroid therapy was not started until after the
324 resection of the third mass. There was no recurrence of abdominal masses in either of these cats
325 after starting corticosteroid therapy for over 1.5 years. The indication to follow surgery with
326 corticosteroid therapy is further supported by 13 cats where prednisolone was discontinued, 85% of
327 these had to have prednisolone restarted a median of 114 days after discontinuation because of
328 recurrence of clinical signs. The median time to the lowest dose of prednisolone was 369 days,
329 with a median lowest maintenance dose of 0.65 mg/kg/day to control the clinical signs; however,
330 as this is a retrospective study, some patients were lost to follow up and it is therefore unclear if
331 prednisolone was tapered further in these cats.

332 The use of secondary immunosuppressive agents and/or antibiotics in cats with FGESF has been
333 reported previously (1, 2). In the current study there was no statistical difference between the
334 survival of cats that were treated with corticosteroids only vs including secondary
335 immunosuppressive agents, regardless of whether or not antibiotics were given. Larger prospective
336 studies are needed to evaluate the effectiveness of secondary immunosuppressive agents and/or
337 antibiotics in the treatment of FGESF.

338 Hydrolyzed or selected protein diets were tried in 37% of the cats in the current study, with 41%
339 of the owners reporting an improvement of clinical signs on these diets. This suggests diet

340 modification as a possible treatment of cats with FGESF. However, all of these cats were already
341 being treated with corticosteroids and further studies are needed to evaluate the use of hydrolyzed
342 diets in the cats with FGESF.

343

344 **Conclusion**

345 Feline gastrointestinal eosinophilic sclerosing fibroplasia is an important differential diagnosis for
346 abdominal masses in cats, and it has a much better prognosis than previously reported, regardless
347 of whether the mass is removed surgically or is treated with medical management alone.

348

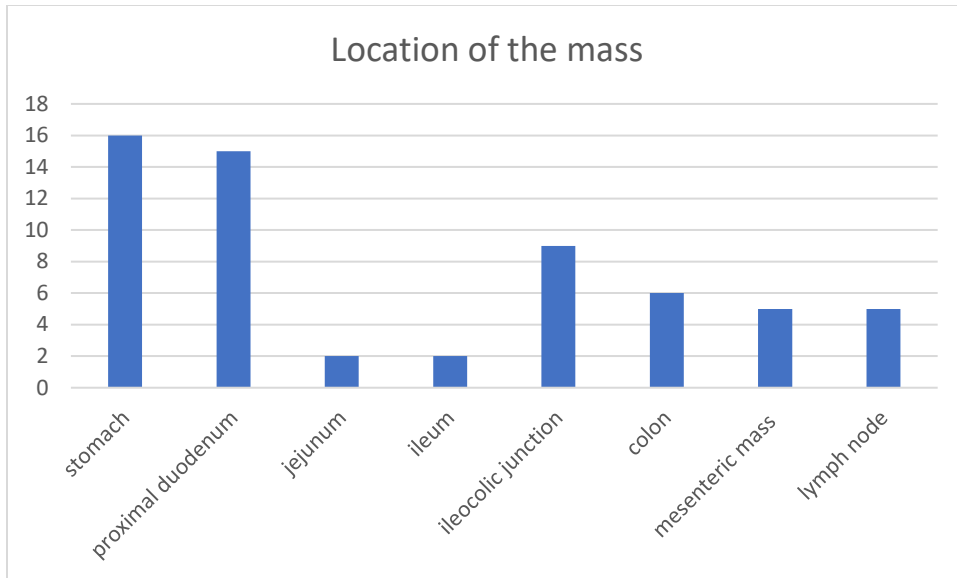


349

350 **Figure 1:** Ultrasonographic image of a mass at the level of the ileocolic junction (white arrow).
351 The mass shows loss of layering, circumferential thickening and eccentric growth. The wall is
352 heterogeneously mixed in echogenicity due to hyperechoic areas. The surrounding peritoneum is
353 hyperechoic (asterisk).

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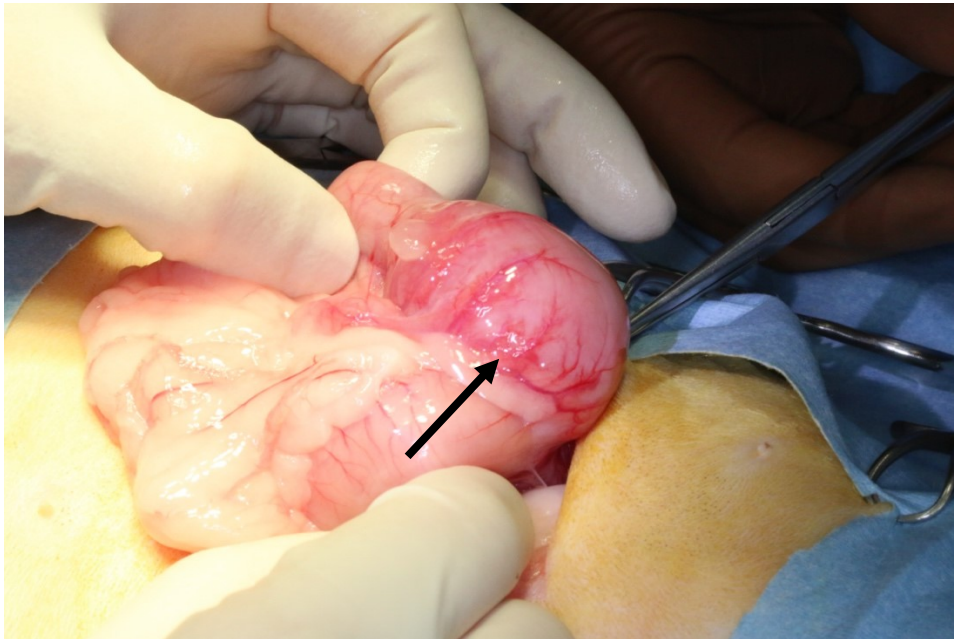


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357

358 **Figure 2:** Common locations of the masses found in cats with feline gastrointestinal sclerosing
 359 fibroplasia in this study. Y-axis represents the number of masses in the location.

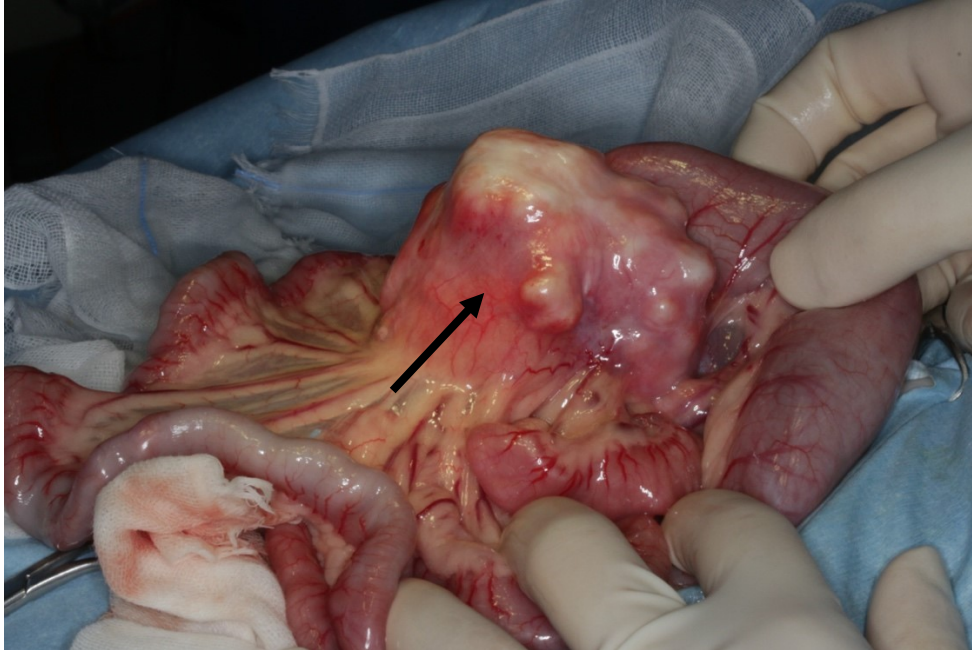
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362 **Figure 3:** Intraoperative photograph of a proximal duodenal mass (arrow) in a cat with feline
 363 gastrointestinal sclerosing fibroplasia. Photo credit: Dr. Atsushi Fujita from Japan Small Animal
 364 Medical Center.

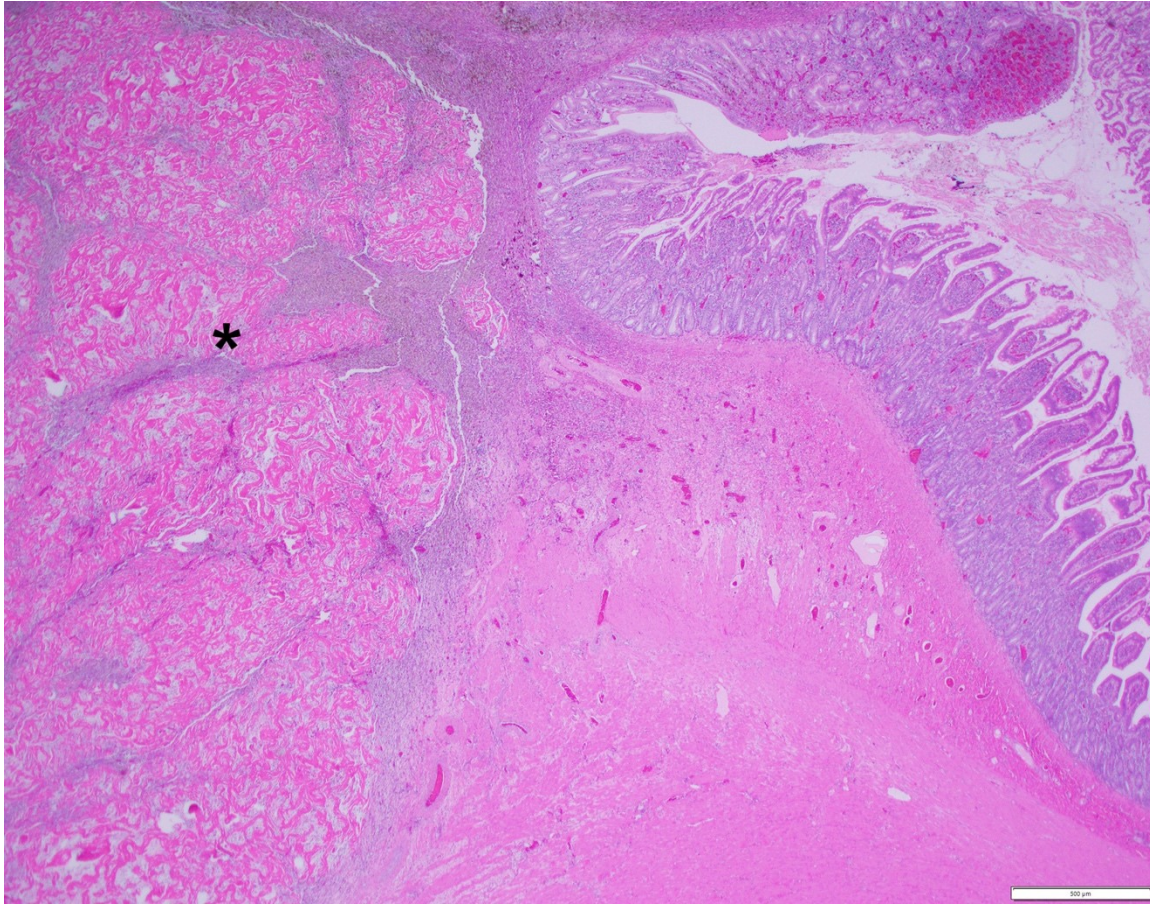
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367 **Figure 4:** Intraoperative photograph of a mesenteric mass (arrow) in a cat with feline
368 gastrointestinal sclerosing fibroplasia. Photo credit: Dr. Atsushi Fujita from Japan Small Animal
369 Medical Center.

370

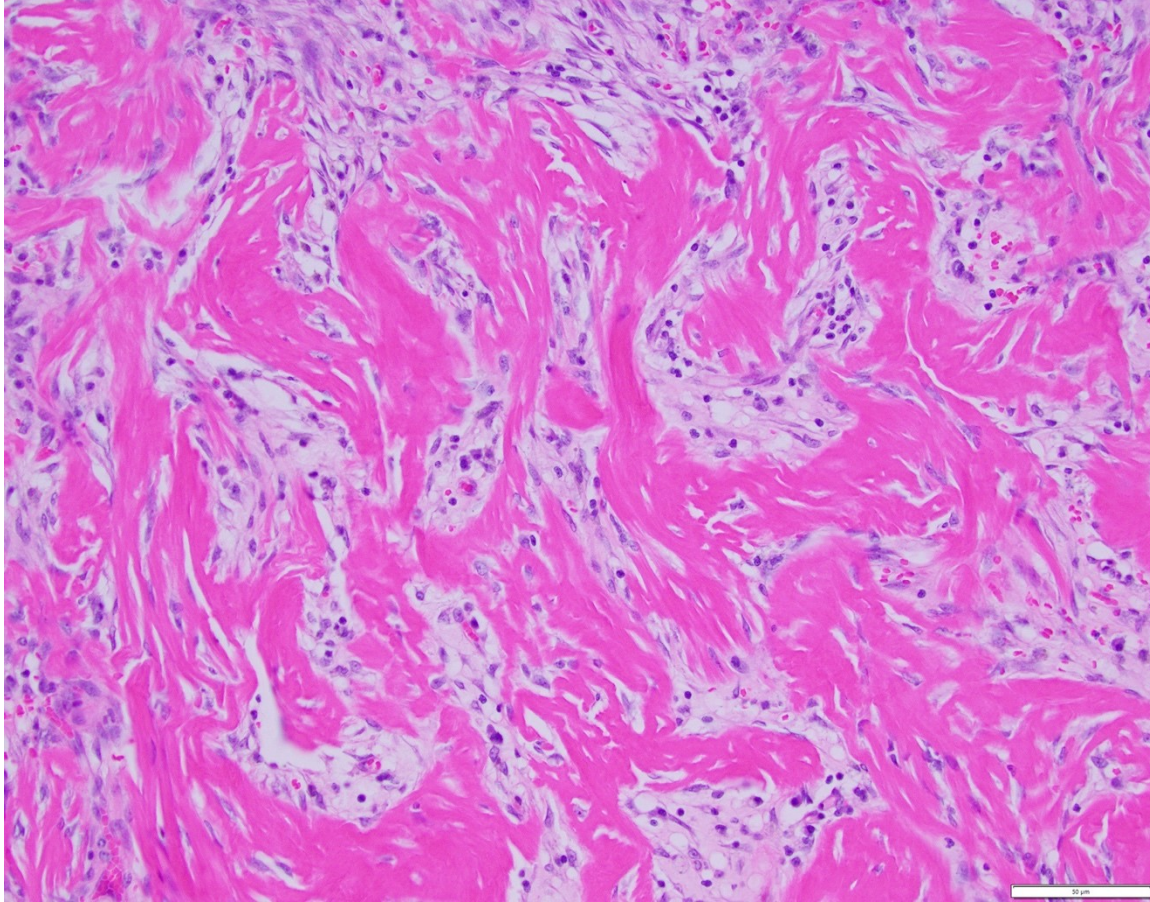


371

372 **Figure 5:** Histological findings of the duodenal mass – the muscularis and submucosa are
373 expanded by a discrete, sparsely cellular mass (*). Hematoxylin & eosin, 20X magnification.

374 Scale bar = 500 μm. Photo credit: Dr. Allison Watson from Colorado State University.

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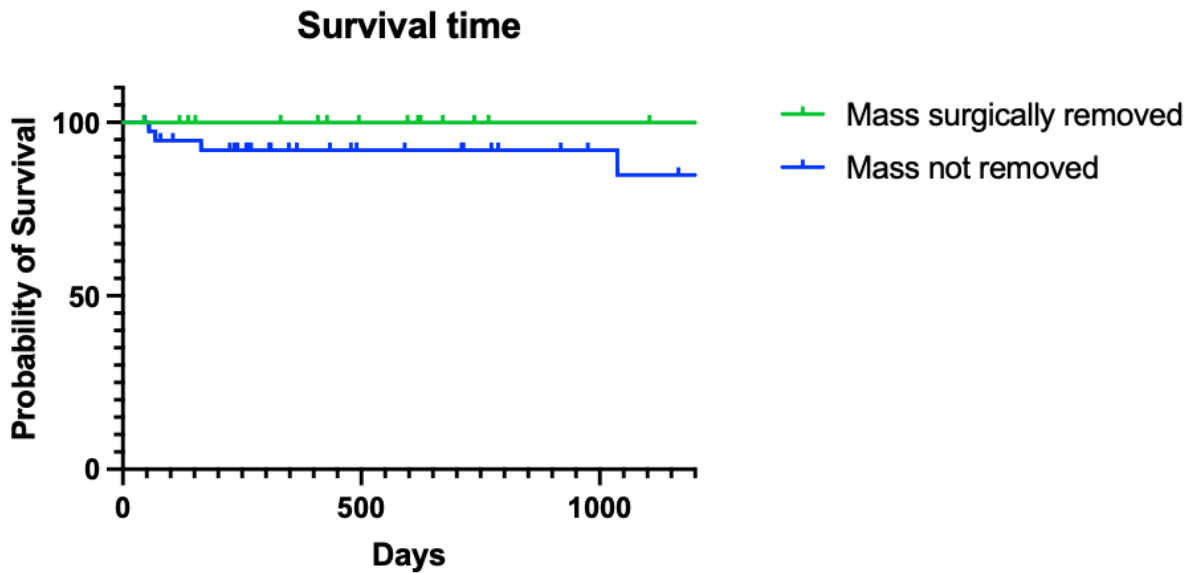


376

377 **Figure 6:** Histological findings of the duodenal mass - the mass is composed of anastomosing
378 trabeculae of sclerotic collagen separated by fibroblasts, macrophages, and small numbers of
379 eosinophils and mast cells. Hematoxylin & eosin, 200X magnification. Scale bar = 50 μ m. Photo
380 credit: Dr. Allison Watson from Colorado State University.

381

382



383
 384 **Figure 7:** Kaplan-Meier plot of survival of cats that had a surgical resection of the mass and cats
 385 where the mass was biopsied only ($p=0.19$). Tick marks represent censored cats.
 386

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392 **OFF-LABEL ANTIMICROBIAL DECLARATION:** Authors declare no off-label use of
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394 **INSTITUTIONAL ANIMAL CARE AND USE COMMITTEE (IACUC) OR OTHER**
 395 **APPROVAL DECLARATION:** Approved by the University of Edinburgh Ethics Review
 396 Committee (VERC Reference: 17.22).

397 **HUMAN ETHICS APPROVAL DECLARATION:** Authors declare human ethics approval
 398 was not needed for this study.

399

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