

1 **Survey-based pilot study into the chosen therapy and prophylaxis used by UK primary**
2 **care veterinary surgeons against canine angiostrongylosis**

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4 Adam Shuttleworth^a, Mark Dunning^a, Ian Wright^b, Hany M. Elsheikha^{a,*}

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6 ^a *School of Veterinary Medicine and Science, University of Nottingham, Sutton Bonington*
7 *Campus, Leicestershire LE12 5RD, UK*

8

9 ^b *Mount Veterinary Practice, Fleetwood FY7 6QX, UK*

10

11 *Corresponding author. Tel: +44 1159516445; fax: +44 1159516440;

12 *E-mail address: hany.elsheikha@nottingham.ac.uk (H.M. Elsheikha)*

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22 **ABSTRACT**

23 Canine Angiostrongylosis (CA), a gastropod-borne parasitic infection caused by the
24 metastrongyloid nematode *Angiostrongylus vasorum*, is an important cause of significant
25 morbidity to domestic dogs across the UK as well as in other European countries. This study
26 aimed to ascertain the frequency at which particular drugs were used by primary care
27 practitioners in the UK for therapy against and prophylaxis for CA. Primary care veterinary
28 clinicians were surveyed using an online questionnaire and face-to-face or telephone
29 interviews. Eighty-six veterinary surgeons responded. The majority of practices (n = 52)
30 included lungworm in their standard anthelmintic protocols; moxidectin was the most
31 common drug used for prophylaxis (n = 71). Fenbendazole was the most frequently selected
32 drug, by 45% of vets, for treatment of confirmed cases of CA despite it being unlicensed for
33 this purpose in the UK and the absence of a clear treatment protocol. The results of this pilot
34 study provide an initial insight into the approach taken by primary care practitioners in their
35 approach to CA. This provides an important starting point for future studies investigating the
36 decision-making for CA amongst UK veterinary surgeons, particularly to clarify whether in a
37 larger cohort an unlicensed drug remains the treatment of choice. The absence of a clear
38 protocol for fenbendazole means that treatment of dogs affected by CA may be suboptimal,
39 increasing the risk of morbidity and mortality.

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42 **Keywords:**

43 *Angiostrongylus vasorum*; Angiostrongylosis; Dogs; Moxidectin; Milbemycin oxime;
44 Fenbendazole

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47 **List of abbreviations:**

48 Bronchoalveolar lavage (BAL), canine angiostrongylosis (CA), enzyme-linked
49 immunosorbent assay (ELISA), polymerase chain reaction (PCR), royal college of veterinary
50 surgeons (RCVS), small animal practice (SAP)

51 **1. Introduction**

52 Canine angiostrongylosis (CA) is a parasitic disease caused by the cardiorespiratory
53 nematode *Angiostrongylus vasorum* (Superfamily: Metastrongyloidea). Dogs become
54 infected after ingesting the third larval stage (L3), usually within an intermediate gastropod
55 host or a paratenic host, such as the common frog (*Rana temporaria*) (Bolt et al., 1993) and
56 chicken (*Gallus gallus domesticus*) (Mozzer and Lima, 2015). In addition, experimental
57 infection of *Biomphalaria glabrata* demonstrated that infected snails shed L3 into the
58 environment, which could create a potential free-living reservoir of infection (Barçante et al.,
59 2003). Rising numbers of confirmed cases of CA have thrown *A. vasorum* into the spotlight
60 globally, driving research into its epidemiology, diagnosis and risk factors (Bwangamoi,
61 1972; Helm et al., 2010; Kistler et al., 2014; Penagos-Tabares et al., 2018). It is well-
62 established that *A. vasorum* has spread from its original hot spot in South West (Simpson and
63 Neal, 1982) and South East England (Chapman et al., 2004) and is now found throughout the
64 UK. Cases have been reported as far north as Scotland (Helm et al., 2009); this expanding
65 parasite's territory has probably expanded faster than our knowledge of its pathogenicity and
66 epidemiology. The full spectrum of the factors related to increasing prevalence of *A. vasorum*
67 in dogs in Great Britain remains unexplained. However, a study conducted in the fox
68 populations showed that the increased prevalence and geographic spread of *A. vasorum* in
69 this reservoir host may have contributed to the increased risk of infection in dogs (Taylor et
70 al., 2015).

71 CA presents with variable clinical signs and in some cases the infection may be entirely
72 sub-clinical. The most common presenting signs relate to the respiratory tract and include
73 dyspnoea and coughing. Other clinical signs include pulmonary hypertension, primary and
74 secondary disorders of haemostasis (with associated haemorrhage) and neurological signs,
75 including seizures. This is a disease with a myriad of possible presentations and therefore

76 remains a diagnostic challenge for veterinarians (Chapman et al., 2004; Koch and Willesen,
77 2009). The lack of a clear pathognomonic profile and the non-specific clinical signs may
78 delay anthelmintic treatment while another suspected cause is investigated, resulting in more
79 severe pathology. This is particularly the case for dogs presenting with coagulopathies or
80 neurological signs in the absence of obvious respiratory signs. A major concern with CA is
81 that it may remain largely asymptomatic and can manifest as sudden death (Bourque et al.,
82 2002; Brennan et al., 2004).

83 In the UK, a number of macrocyclic lactone-based products are licensed to treat
84 angiostrongylosis, containing moxidectin in a topical form (along with imidacloprid) or
85 milbemycin oxime in tablet form. The treatment efficacy for a single dose of the formulation
86 of imidacloprid/moxidectin was found to be 85.2%. There was no significant difference
87 between this and the 91.3% efficacy of fenbendazole administered daily for 20 days. The
88 former of these treatments being licensed for use against AV in dogs, whereas the latter is not.
89 Of those dogs still shedding larvae which received a further dose of imidacloprid/moxidectin,
90 all were then found to be Baermann negative (Willesen et al., 2007).

91 Besides being licensed for treatment of adult worms imidacloprid/moxidectin is licensed
92 for monthly use for the prevention of angiostrongylosis and prevention of patent infection,
93 with 100% efficacy against L4 larvae and immature adults (L5) of *A. vasorum* (Schnyder M.
94 et al., 2009). In regards to milbemycin oxime, commonly available in combination with
95 praziquantel, a two-dose protocol achieved a clinical improvement, but without clearance of
96 larval shedding and treatment was therefore extended to a protocol of using 4 weekly
97 treatments to reduce the level of infection, achieving negative faecal Baermann results in 14
98 out of 16 dogs (Conboy, 2004). Monthly use of milbemycin oxime is licensed for prevention
99 of angiostrongylosis by reduction in the level of infection by immature adult (L5) and adult
100 parasite stages, with a recent study showing a worm count reduction efficacy of 94.9%

101 (Lebon et al., 2016). Whilst there are additional publications testifying the efficacy of
102 fenbendazole in naturally occurring CA infections (Brennan et al., 2004; Chapman et al.,
103 2004; Manning, 2007), in the absence of licensing, optimal dose, frequency and duration
104 remain unclear. The risks of using unlicensed products are highlighted by levamisole, which
105 was historically advocated for the treatment of CA due to its potency and rapid onset of
106 action. However despite efficacy against CA, this is no longer recommended due to the
107 occurrence of significant side-effects including anaphylaxis in levamisole-treated patients,
108 thought to result at least in part from the rapid increase in circulating worm antigen (Søland
109 and Bolt, 1996).

110 The efficacies of many of these drugs have been studied experimentally and in a number
111 of clinical studies, and they have been licensed as a result. It is at present however unknown
112 which products primary care veterinary surgeons in the UK are choosing to treat CA. It is
113 suspected anecdotally that fenbendazole, whilst unlicensed remains the first-choice
114 anthelmintic treatment for most primary care vets. In addition to this, it also remains
115 unknown as to their choice of prophylaxis. A review of the therapeutic choices made by
116 veterinary surgeons around CA treatment and prophylaxis across the UK is therefore
117 important. In addition, the reasons behind these choices and the ‘perceived’ efficacy of this
118 choice made by primary care clinicians are also of value. Whilst the latter point is difficult to
119 prove retrospectively, the vet’s opinion is important as this will influence their future
120 therapeutic choices. Perhaps a more important rationale for this study in the UK is the
121 importance of prescribing veterinary medication according to the cascade. Therefore, the
122 aims of this pilot survey were to identify which anthelmintic drugs were being used for
123 therapy and prophylaxis of CA by UK primary care veterinary surgeons. An additional aim
124 was to review the reported clinical signs in the confirmed cases of CA along with the method
125 by which the disease was confirmed. Finally, the clinical outcome of cases diagnosed with

126 CA and the perspective of the veterinary surgeons on the efficacy of drugs used in the
127 treatment was also examined.

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129 **2. Material and methods**

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131 *2.1. Ethics statement*

132 The Institutional ethical review committee approved this study. Responses to the survey were
133 entirely voluntary and all data obtained was anonymised and stored securely.

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135 *2.2. Data collection*

136 Data were collected for this study using a combination of an online questionnaire ([Table 1](#)),
137 and telephone or face-to-face interviews, all targeting small animal veterinary surgeons in
138 primary care practice. The online survey contained 4 sections, beginning with a demographic
139 question asking for the number of years the clinician had been working in small animal
140 practice (SAP). This was followed by questions on the drugs used to prevent CA (clinicians
141 could indicate more than one), and whether the practice's standard worming prophylaxis was
142 licensed for lungworm prevention. The final section was focused on treatment of CA, asking
143 the clinician for the drug they used and their efficacy rating for this drug. For ease of
144 comparison, drugs were listed according to their classification and not trade name for ease of
145 understanding. Clinicians were allowed to indicate more than one drug. The survey provided
146 an optional 'other' field which contained drugs less frequently mentioned (ivermectin and
147 selamectin). The online questionnaire was piloted by using other primary care veterinary
148 surgeons and changes were implemented where required. The questionnaire was available for
149 completion by veterinary surgeons during the pilot project period: between 11/10/2016 and
150 28/10/2016 via a specific URL provided to 287 primary care practices throughout England.

151 Additional interviews were conducted to supplement the online questionnaire, using the same
152 questions and additionally requesting the standard worming protocol of the practice, then
153 determining if the product was licensed in the UK to prevent CA. Eleven interviews were
154 conducted in-person and 28 (of 59 practices contacted) consented to a telephone interview.
155 Direct interviews were conducted across Greater Manchester, Lancashire, Leicestershire,
156 Nottinghamshire and Norfolk. This produced a distribution of 326 practices (287 via email,
157 28 via telephone and 11 via direct interview).

158

159 *2.3. Data analysis*

160 Responses to the online survey and interview locations were plotted on a map of England in
161 order to assess geographical coverage. Data from all response types was collected,
162 categorized and analysed using Excel 2016 (Microsoft Corporation©) and SPSS Version 23
163 2005 (IBM®). Maps were created using the mapping feature of Excel 2016 (Microsoft
164 Corporation©).

165

166 **3. Results**

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168 *3.1. Participants and distribution*

169 Eighty-six responses were obtained from the 326 practices contacted representing a 26%
170 response rate. The online questionnaire provided 47 responses (representing 14% of total
171 distribution), 28 were from the telephone interviews (8.6% of total) and 11 from face-to-face
172 interviews (3.4% of total). In order to assess how demographics of the participants influenced
173 their responses, the number of years' post-qualification that the veterinary surgeons had been
174 in SAP was collated into a histogram (Fig. 1). The spread of all responses by county is shown
175 in Fig. 2A. Greater London provided the greatest number of responses (13), followed by

176 Bristol (5) and then Cornwall, Norfolk, Nottinghamshire and Warwickshire (all of which
177 provided 4). An additional map was created, displaying the number of confirmed cases of CA
178 varying by county (Fig. 2B). Of the 86 responding practices, 19 (22%) stated that one or
179 more confirmed cases of CA had been examined at that practice within the previous year.
180 When the respondents were asked for the exact number of cases examined, the total came to
181 41 across all practices. The maximum number of cases examined at any one practice was 10
182 with a median of 2 cases per practice (IQR 1-2).

183

184 *3.2. Frequency of clinical signs*

185 A variety of clinical signs were reported by the study participants, those associated with the
186 haematologic, respiratory and neurological systems were the most commonly reported (Table
187 2). One case was excluded from this analysis due to the fact CA was only discovered at
188 referral and the primary care clinician completing the survey was unaware of the presenting
189 clinical signs. Three respondents (handling 14 serologically confirmed cases of CA between
190 them) observed subclinical cases and another three (6 confirmed cases in total) witnessed the
191 sudden death of infected dogs. Signs relating to the gastrointestinal system were noted
192 infrequently (one case reporting vomiting and one diarrhoea).

193

194 *3.3 Diagnostic testing*

195 Clinicians were asked to select the method(s) they had used to confirm a suspected case of
196 CA from the options of: coproscopic examination (sub-options: Baermann, faecal smear),
197 cytological diagnosis by bronchoalveolar lavage (BAL), in-house serological assay (Angio
198 Detect™), external laboratory ELISA and other (Fig. 3). Two clinicians reported that they
199 were unaware of the diagnostic test used for diagnosis. Over 50% of the veterinary surgeons

200 reported using the Angio Detect™ (IDEXX Laboratories, Wetherby, West Yorkshire, UK) to
201 confirm their diagnosis of CA whilst none sent serum samples to an external laboratory for
202 ELISA diagnosis. Of the 13 clinicians who used the Angio Detect™ test, three did so in
203 conjunction with another testing modality. One respondent used a combination of BAL and
204 PCR.

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206 *3.4. Treatment of CA*

207 The treatment choices for confirmed cases of CA by those clinicians surveyed are presented
208 in **Fig. 4A**. There were 51 responses to this question, 17 of which related to treatment given
209 in a confirmed case with 34 relating to which drug the clinician would select, were they to
210 encounter a case of CA. In the 17 confirmed cases, 45.5% of clinicians selected
211 fenbendazole, which was the most popular drug used. Moxidectin and milbemycin oxime
212 were each selected by 27.3% of responding clinicians. No other drugs were recommended as
213 treatment for CA. In contrast, when asked which drug the clinician would use in a
214 hypothetical case of CA, the most frequent response was moxidectin (41%), followed by
215 fenbendazole (28%) and milbemycin oxime (23%).

216 There were a variety of treatment protocols for CA reported by the responding clinicians.
217 The most consistent was monthly moxidectin, with 19/22 (86%) of respondents advising this
218 was given to treat CA. Four respondents using milbemycin oxime to treat cases of CA
219 advised that this should be given weekly for 4 weeks, one respondent prescribed this weekly
220 for 2 weeks and one recommended a single dose to be sufficient for treatment. The remaining
221 9 respondents selecting milbemycin oxime to treat CA did not specify the frequency or
222 duration of treatment. The dosing schedule reported by 43% of the respondents for
223 fenbendazole was once daily. However, there was considerable discrepancy in duration of
224 therapy. One clinician reported 3 days, one recommended 5 days, three recommended 7 days,

225 two recommended 10 days and two recommended 14 days. Unfortunately, 12 clinicians using
226 fenbendazole therapy failed to provide a therapeutic frequency or duration. We asked the 19
227 clinicians reporting confirmed cases of CA to describe response to therapy. Interestingly,
228 only 4 clinicians reported that they re-tested dogs after therapeutic intervention,
229 corresponding to 6 of the 41 (15%) infected dogs. Of these six re-tested dogs, all produced
230 negative results using the in-house serological assay (Angio Detect™).

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232 *3.5. Prevention of CA*

233 When asked the clinicians if their practice's standard prophylactic de-worming protocol
234 included lungworm prevention, 52 agreed and 28 disagreed. Six respondents reported using
235 more than one standard protocol, depending on the dog's 'risk of infection'. Surveyed
236 clinicians provided information on which drugs were most frequently used as prophylaxis
237 against CA. [Fig. 4B](#) displays the frequency of drugs being selected as prevention for CA.
238 Those respondents that included lungworm prevention in their standard protocol were then
239 asked which product was chosen to achieve this (more than one product could be selected).
240 Products containing moxidectin and milbemycin oxime were the most commonly prescribed
241 preventatives, with 71 clinicians selecting moxidectin and 56 selecting milbemycin oxime.
242 One respondent selected fenbendazole as their prophylactic treatment of choice and two
243 selected selamectin.

244

245 **4. Discussion**

246 This pilot study was conducted to begin to understand more about how anthelmintics are used
247 for treatment of and prophylaxis against CA by primary care clinicians in the UK. The survey
248 responses provided a reasonable spread across the UK and there was a reasonable response
249 rate of 26%. Despite the reasonable response rate, the absolute number of responses was low

250 which represents a limitation. This is likely to reflect the short period of data collection,
251 which was necessitated due to the timescale over which this pilot study was conducted. There
252 was a predominance of responses from those vets with fewer years in small animal practice.
253 It is difficult to know whether this reflects that newer graduates are more likely to respond to
254 questionnaires perhaps because they did not anticipate compensation or monetary incentives
255 for the time spent completing the survey or because newer and updated knowledge about
256 diseases and treatments have become available in recent years. Also, young veterinarians,
257 aged between 26-35, represent a third of veterinary surgeons in the UK according to the 2014
258 RCVS survey (Buzzeo et al., 2014); thus it is feasible to expect younger vets to represent a
259 large proportion of the respondents. The highest correlation between the number of responses
260 and the frequency of detection of clinical cases has been detected in Greater London, which is
261 in agreement with others (Kirk et al., 2014). This correlation might be due to previous
262 knowledge and experience in CA therapeutics in clinics in this geographic area, which had
263 prolonged history of receiving cases of CA than those in the low endemic areas. However,
264 the small sample size limits the generalization of our finding and does not allow a robust
265 correlation between the participant responses to the questionnaire and the spatial distribution
266 and intensity of clinical *A. vasorum* cases seen in dogs in England to be made. How this
267 correlation may differ with age of the clinicians remains to be determined.

268 Under the veterinary prescribing cascade in the UK, clinicians must first use a drug that
269 is licensed to treat the condition in question (in this case CA) in the patient's species (in this
270 case dogs) before moving onto alternatives in justifiable circumstances (VMD, 2015).
271 Therefore, a clinician's first choice of treatment for CA should theoretically contain either
272 moxidectin or milbemycin oxime. The results obtained here suggest that the majority of
273 clinicians are using fenbendazole to treat CA, which is off license and does not follow the
274 cascade. The reasons behind this were not interrogated in this pilot study, but a subsequent

275 study is planned to understand more about what drives decision-making for treatment of CA.
276 It is possible that the first-line use of fenbendazole is as a result of its long-standing presence
277 in the market place, suggesting veterinary surgeons feel more familiar with the outcome for
278 CA cases. That said, the majority of our responses came from relatively recent graduates and
279 not from clinicians that had been in the profession for a considerable time period. Moxidectin
280 became available in April 2003, shortly followed by milbemycin oxime tablets, compared to
281 fenbendazole which was first produced in January 1993 (VMD, 2017). This clearly warrants
282 further investigation to provide important insight into understanding decision-making around
283 this condition. This is important, as at least from this pilot study, the licensed products are
284 less frequently used than the unlicensed one, despite the advice of the RCVS and the VMD.

285 In trying to seek some consensus as to the treatment strategies employed by the cohort of
286 clinicians surveyed, it seems that selected fenbendazole is being used for differing periods of
287 time. Periods ranged from 3 to 14 days. The literature would suggest that fenbendazole may
288 be effective in treating CA with courses lasting between 5 and 21 days (Chapman et al., 2004;
289 Willesen et al., 2007). There was no association between those vets using fenbendazole for a
290 shorter period and ineffective therapy or relapse. Further research is necessary to clarify the
291 most appropriate duration of therapy required to obtain maximum efficacy of fenbendazole.

292 Whilst all drugs, fenbendazole, moxidectin and milbemycin oxime (Conboy, 2004;
293 Willesen et al., 2007), have been demonstrated to be efficacious, Moxidectin was found to be
294 the most frequently prescribed drug in hypothetical cases of CA. The reasoning behind this is
295 currently unclear but may be because it only requires a single application or that it doesn't
296 rely on having to tablet the dog. Milbemycin oxime was the least frequently prescribed drug
297 (excluding ivermectin and selamectin, categorised into the field entitled "other") in a
298 hypothetical case. This could be due to a perceived low efficacy, the fact that it comes in
299 tablet form, the need for multiple doses or another reason. Another study into the treatment

300 efficacy of milbemyacin oxime to ascertain the minimum number of weeks treatment needed,
301 would benefit clinicians.

302 The low number of treated cases that are re-tested at a later date could be influenced by a
303 host of factors, such as the cost of such testing, perceived hassle or possible fatality. Whilst
304 there are many reasons clinicians may avoid re-testing patients, it must be remembered that
305 without this information it cannot be concluded as to whether the apparent improvement in
306 clinical signs was associated with true resolution of infection. It is important to reiterate that
307 the eradication of the infection means that there is no residual worm burden that could lead to
308 recrudescence. Treatment of CA does not result in lasting immunity and as such treated dogs
309 (even those that re-test negative) should continue to receive a monthly anthelmintic for
310 preventative purposes (Böhm et al., 2014; Lebon et al., 2016; Schnyder et al., 2009).

311 This study found that, moxidectin and milbemyacin oxime were the most frequently
312 prescribed preventative drugs, reflecting their general popularity in primary practice.
313 Alternatively, this may represent an attempt to include lungworm prevention in the standard
314 de-worming protocol of a practice, without the need to use an additional drug. In this small
315 survey, moxidectin was more frequently prescribed than milbemyacin oxime. Whether this is
316 owing to its higher efficacy or simply its existing popularity in veterinary practice is unclear.
317 The majority of clinicians responded yes when asked if their practice's standard de-worming
318 protocol included lungworm prophylaxis. Whilst important that 65% of the surveyed
319 practices are including prophylaxis for CA, this means that 35% of practices do not. This may
320 relate to a perceived lower prevalence in their area, however it is now widely accepted that *A.*
321 *vasorum* is spreading throughout the UK implying that no area can be guaranteed as
322 lungworm-free (Kirk et al., 2014). Some clinicians may wish to adopt a risk-based approach
323 to CA prevention in areas where the parasite is not endemic. Assessing the relative risk is one
324 way that this could be carried out. The risk factors include: age (dogs less than 18 months old

325 were most likely to test positive), no CA prophylaxis for the past 3 months and possibly
326 season (dogs were more likely to test positive in winter and spring) (Morgan et al., 2010). It
327 must be mentioned here that the online respondents were directly asked if their standard de-
328 worming protocol included lungworm prevention and not to provide details of this protocol.
329 This creates the possibility that these clinicians may in fact not be using effective CA
330 prophylaxis however given the nature of the study, the protocol could not be determined from
331 the responses.

332 A secondary aim of this survey was to look at the range of clinical signs associated with
333 CA, encountered in primary care practice. Morgan et al. (2010) found the 3 most common
334 clinical signs to be coughing, dyspnoea and lethargy respectively. This pilot study identified
335 coughing, haemorrhage and depression as the most frequently reported signs. These are
336 therefore very similar in as much as the interpretation of lethargy and depression could be
337 considered variations on the same presenting sign associated with the neurological system.
338 An important highlight from this study is that CA may be an incidental finding and therefore
339 particular presenting clinical signs may not be associated with the infection per se, but the
340 result of an unconnected comorbidity. Although respiratory signs and coagulopathy are well-
341 accepted presenting signs association with CA, vomiting/diarrhoea have been less frequently
342 reported to be significant findings in previous studies (Chapman et al., 2004; Koch and
343 Willesen, 2009; van Doorn et al., 2009). The single episodes of vomiting and diarrhoea
344 occurring in this pilot study are unlikely to be directly related to CA. As expected, the
345 preferred diagnostic approach to confirm a suspected case of CA was the Angio Detect™
346 test. Whilst the specific reasons for this were not surveyed, it is perhaps due to its simplicity,
347 reliability and rapidity of results (Liu et al., 2017). Despite a specificity of 100%, it must be
348 remembered that with a sensitivity of 84.6% (lower than the commercially available CA
349 ELISA 94.9%) it is less suited as a screening test (Schnyder et al., 2014). Therefore given the

350 potential for a false negative result with Angio Detect™, it is important that additional
351 diagnostics are employed to effectively rule out CA when it is a major differential.

352

353 **5. Conclusion**

354 Responding clinicians indicated that they would be likely to use moxidectin to treat a
355 hypothetical case of CA, however contrary to this, the most commonly prescribed drug in
356 confirmed cases of CA was actually fenbendazole. It is therefore unclear as to what
357 influences UK veterinary surgeon's choice of drug for CA, particularly when fenbendazole is
358 unlicensed. From this small pilot study and in line with the literature on CA regarding the
359 optimal duration and dose of fenbendazole, there appeared to be no consensus amongst the
360 vets as to the optimal duration and dose of therapy for fenbendazole. This study highlights the
361 need for larger and more detailed studies in to the choices that veterinary surgeons in the UK
362 are making in relation to CA. Once these have been completed the reasons for particular
363 therapeutic choices can be better understood.

364

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374

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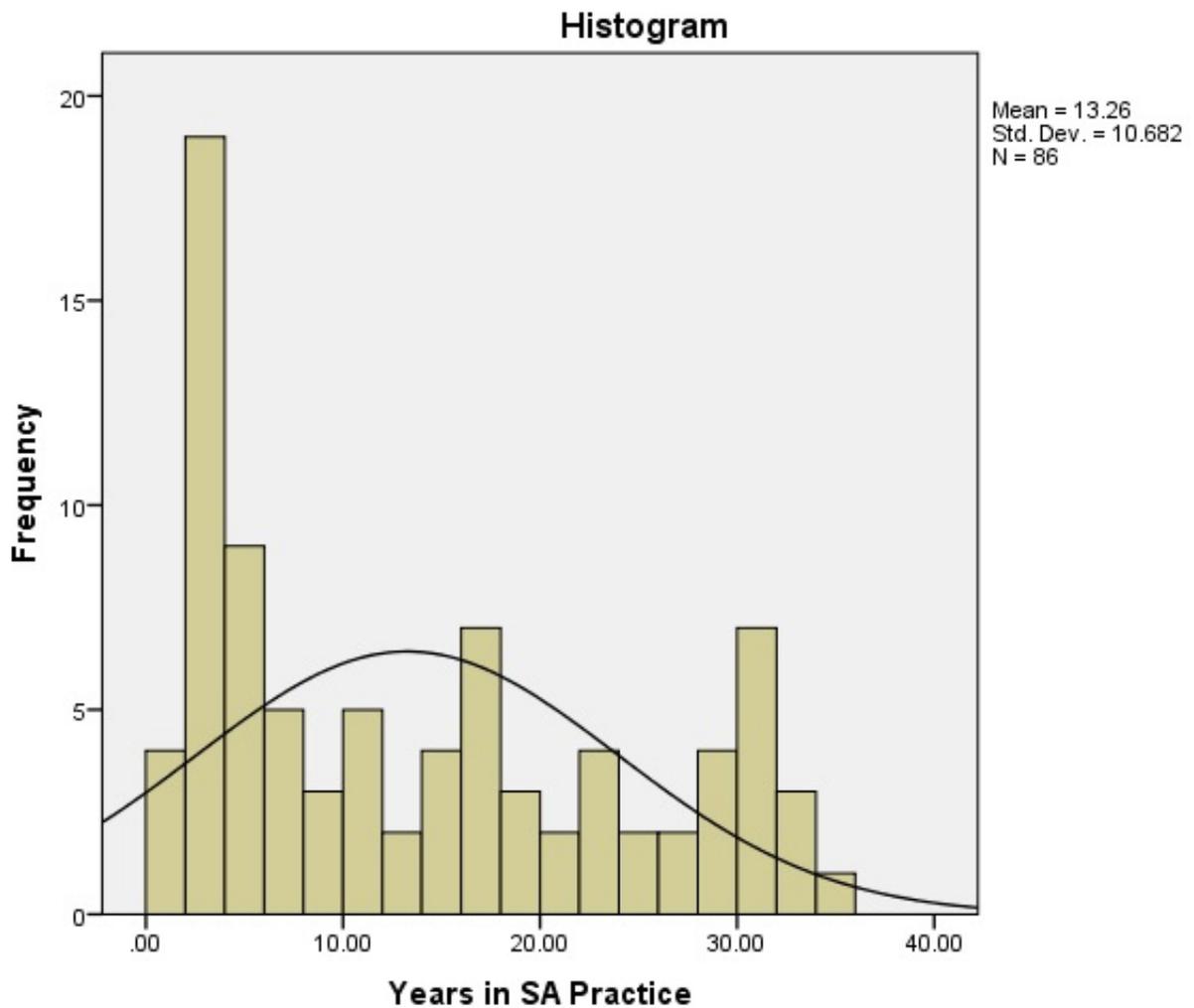
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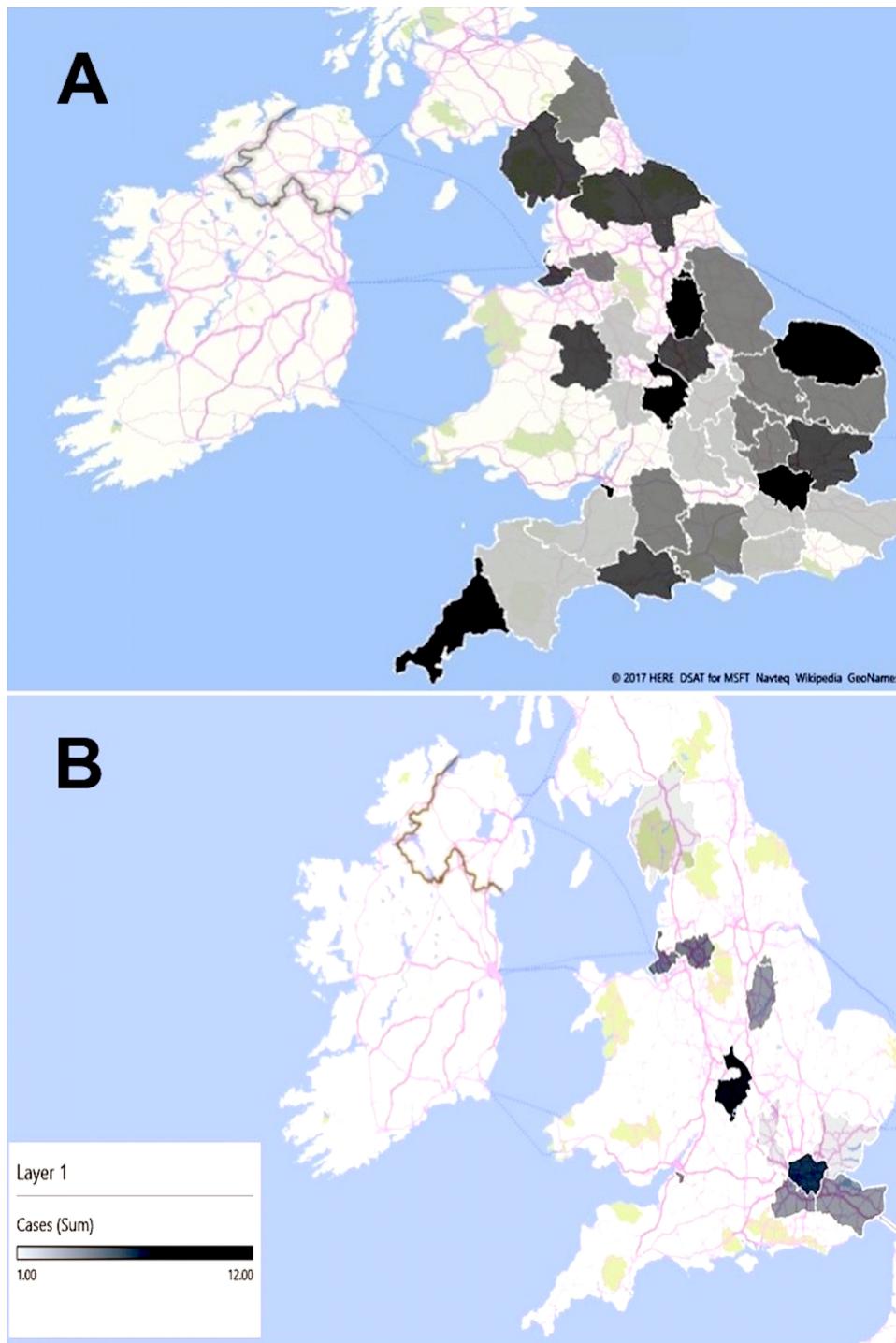
461 **Figure legends:**
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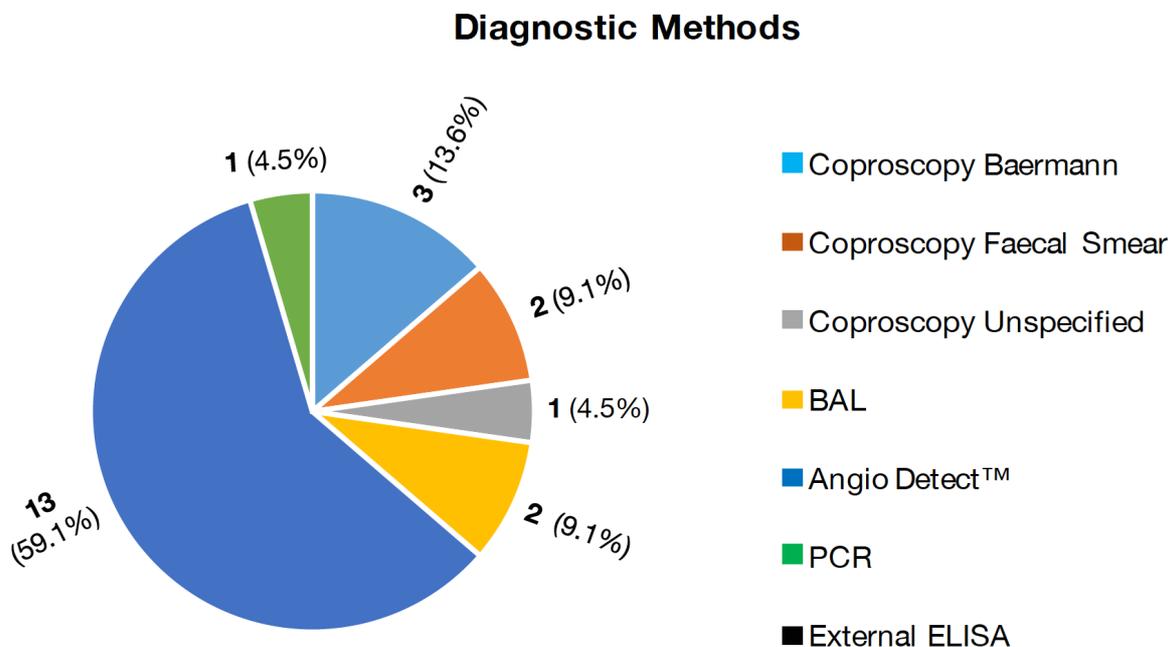
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465 **Fig. 1.** A histogram displaying the frequency of responses from different brackets of years in
466 SAP. The slight left skew in the curve indicates that more survey responses were obtained
467 from clinicians who had spent less time working in SAP, whereas those with more years in
468 SAP were less represented.

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501 **Fig. 2.** Mapping the distribution of the survey respondents and clinical cases of CA in
502 *England.* (A) Geographical distribution of responses to the survey by county. A relatively
503 even coverage of England was achieved with Greater London providing the highest number
504 of total responses. Shading intensity reflects the number of responses – darker reflects more
505 responses. (B) Geographical distribution of CA cases reported by the responding clinicians.
506 Confirmed cases were reported in Bristol (3), Buckinghamshire (1), Cumbria (1), Essex (1),
507 Greater London (9), Greater Manchester (4), Kent (2), Merseyside (4), Nottinghamshire (3),
508 Surrey (2) and Warwickshire (12).



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511 **Fig. 3.** The method(s) used by primary care clinicians to diagnose CA. The numbers at the
512 outer edge of the chart represent the percentage of clinicians using that diagnostic test.
513 Coproscopic examination consisted of: Baermann test, faecal smear or unspecified, and
514 clinicians could select more than one. The most frequently used test was Angio Detect™
515 while none of the surveyed clinicians used the external ELISA.

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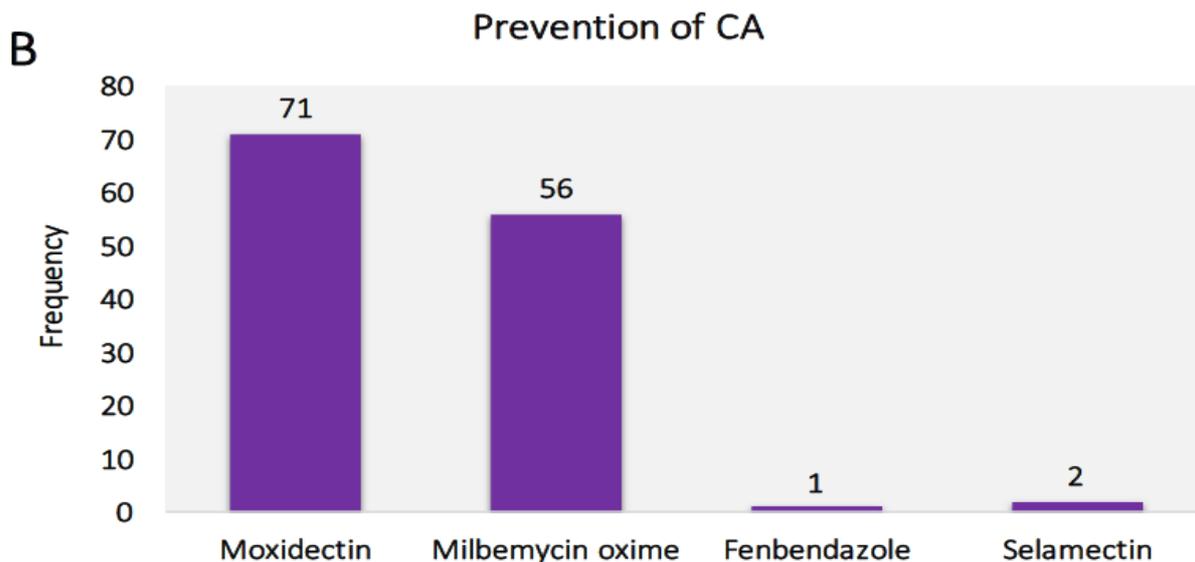
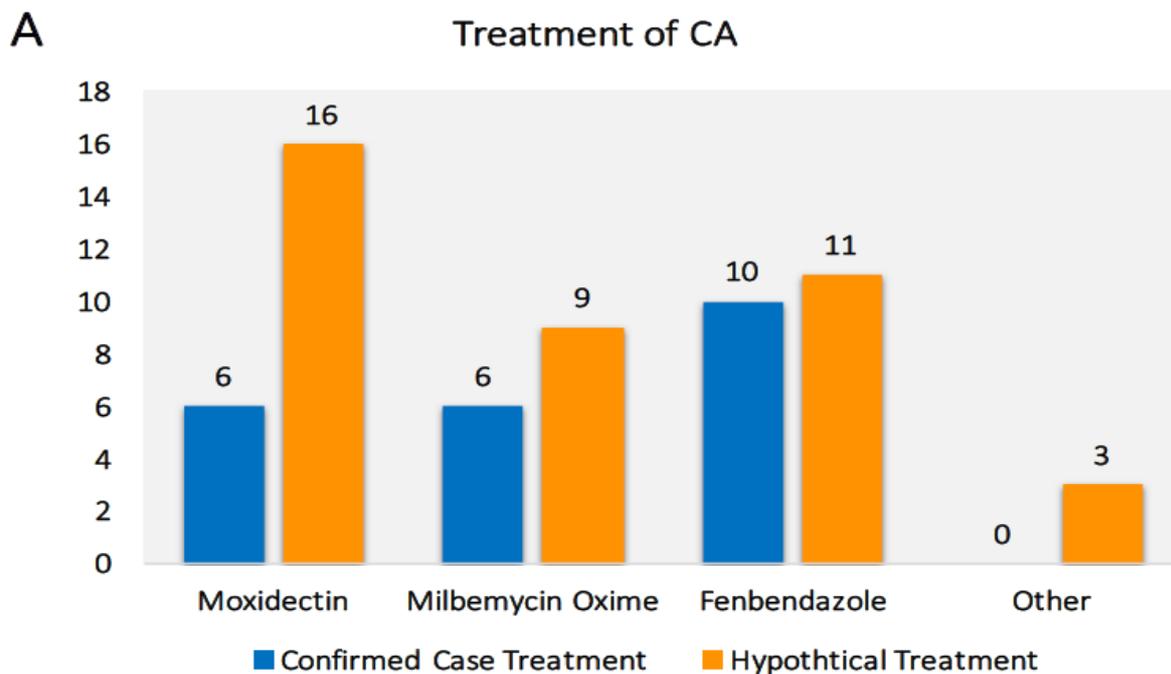
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532 **Fig. 4.** (A) The percentage of clinicians selecting the drugs to treat both confirmed and
 533 hypothetical cases of CA. Although the most frequently selected drug in a hypothetical case
 534 was moxidectin, fenbendazole was the most commonly used drug in confirmed cases. Other
 535 responses include selamectin (selected by 2 clinicians) and ivermectin (selected by 1
 536 clinician). (B) The most frequently used drugs for the prevention of CA. Moxidectin was the
 537 most commonly advised preventative drug, with milbemycin oxime second. Two clinicians
 538 recommended selamectin and only one advised fenbendazole.

539 **Table 1**

540 Questions completed by clinicians who took part in the online survey.

Category	Question
Demographic	<ul style="list-style-type: none"> – Please provide your practice’s post code – How many years have you worked in SAP?
Prophylaxis	<ul style="list-style-type: none"> – Which de-worming product(s) do you recommend for the prevention of <i>A. vasorum</i> infection and at what dose/frequency? – To your knowledge, how many times has a dog become infected with <i>A. vasorum</i> whilst using this prevention protocol? – Does your practice’s standard de-worming protocol include lungworm prevention?
Cases	<ul style="list-style-type: none"> – Have you seen a confirmed case of angiostrongylosis at your practice within the last year (if yes, please indicate how many cases)? – Which of the following symptoms were exhibited by the patient(s)? * – By what method was the diagnosis of angiostrongylosis confirmed? *
Treatment	<ul style="list-style-type: none"> – Which de-worming product(s) have you used in the past to treat a confirmed case of angiostrongylosis and at what dose/frequency? – How would you rate the efficacy of this treatment? – If the treatment was successful, after what period of time was the patient considered to be cured of infection?

- How many of the treated dogs were re-tested at a later date?
 - Of these re-tested dogs, how many displayed a positive result?
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542 *For these questions, clinicians could select multiple options from lists on the screen, or add
543 their own choice.

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570 **Table 2**

571 The frequency of clinical signs relating to CA. Clinical signs reported by responding
 572 clinicians associated with confirmed cases of CA categorised into those presenting with
 573 single or multiple clinical signs. Coughing, haemorrhage and depression were the most
 574 commonly reported signs.

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Clinical presentation	Clinical signs (number of confirmed cases)
Individual signs	Coughing (2)
	Dyspnoea (1)
	Haemorrhage (1)
	Total number of cases = 4
Combined signs	Coughing, exercise intolerance, haemorrhage (3)
	Coughing, haemorrhage, sudden death (3)
	Dyspnoea, coughing, exercise intolerance (2)
	Dyspnoea, coughing, depression, haemorrhage, sudden death (2)
	Depression, haemorrhage (2)
	Coughing, depression, haemorrhage (2)
	Depression, haemorrhage, vomiting, neurological deficits (2)
	Coughing, haemorrhage (1)
	Dyspnoea, coughing, depression (1)
	Depression, exercise intolerance, diarrhoea (1)
	Dyspnoea, haemorrhage, sudden death (1)
	Dyspnoea, coughing, depression, exercise intolerance (1)
	Coughing, asymptomatic (10)
	Haemorrhage, asymptomatic (2)
Total number of cases = 33	
Asymptomatic	No clinical signs (2)
	Total number of cases = 2
Unspecified*	Unknown (2)
	Total number of cases = 2

576 *One respondent (2 confirmed cases) failed to specify which clinical signs had been
 577 displayed as CA was discovered at referral.

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