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Original Article**Patterns of antimicrobial agent prescription in a sentinel population of canine and feline veterinary practices in the United Kingdom**

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3 Highlights

- 4 • Antimicrobial agent prescription was monitored in a large UK population of cats and dogs over a
5 2 year period (2014-2016).
- 6 • Systemic antimicrobial agents were prescribed more frequently to cats; topical prescription was
7 more frequent in dogs.
- 8 • A temporal reduction (2014-2016) in antimicrobial agent prescription was observed in both cats
9 and dogs in this population.
- 10 • Premises which prescribed antimicrobial agents commonly to cats generally also prescribed
11 commonly to dogs.
- 12 • The most frequently prescribed antibiotics were ceftiofur in cats and clavulanic acid potentiated
13 amoxicillin in dogs.

14 Abstract

15 Antimicrobial resistance is an increasingly important global health threat and the use of
16 antimicrobial agents is a key risk factor in its development. This study describes antimicrobial
17 agent prescription (AAP) patterns over a 2 year period using electronic health records (EHRs)
18 from booked consultations in a network of 457 sentinel veterinary premises in the United
19 Kingdom. A semi-automated classification methodology was used to map practitioner defined
20 product codes in 918,333 EHRs from 413,870 dogs and 352,730 EHRs from 200,541 cats,
21 including 289,789 AAPs. AAP as a proportion of total booked consultations was more frequent
22 in dogs (18.8%, 95% confidence interval, CI, 18.2-19.4) than cats (17.5%, 95% CI 16.9-18.1).
23 Prescription of topical antimicrobial agents was more frequent in dogs (7.4%, 95% CI 7.2-7.7)
24 than cats (3.2%, 95% CI 3.1-3.3), whilst prescription of systemic antimicrobial agents was more
25 frequent in cats (14.8%, 95% CI 14.2-15.4) than dogs (12.2%, 95% CI 11.7-12.7). A decreasing
26 temporal pattern was identified for prescription of systemic antimicrobial agents in dogs and
27 cats. Premises which prescribed antimicrobial agents frequently for dogs also prescribed
28 frequently for cats. AAP was most frequent during pruritus consultations in dogs and trauma
29 consultations in cats. Clavulanic acid potentiated amoxicillin was the most frequently
30 prescribed antimicrobial agent in dogs (28.6% of prescriptions, 95% CI 27.4-29.8), whereas
31 ceftiofur, a third generation cephalosporin, was the most frequently prescribed antimicrobial

32 agent in cats (36.2%, 95% CI 33.9-38.5). This study demonstrated patterns in AAP over time
33 and for different conditions in a population of companion animals in the United Kingdom.

34 *Keywords:* Canine; Feline; Antimicrobial resistance; Antibiotic prescribing practices; Surveillance

35 Introduction

36 Antimicrobial resistance (AMR) is widely recognised as an increasingly important
37 global health threat.^{1,2,3,4} Evidence of transmission of bacterial resistance amongst human
38 beings, livestock (Cuny et al., 2015) and companion animals¹ (Zhang, 2016) demonstrates the
39 necessity of a ‘one health’ approach to preserve treatment efficacy.² Although use of
40 antimicrobial agents selects for and promotes transfer of resistance (Rantala et al., 2004;
41 Magalhaes et al., 2010; Cantón and Bryan, 2012), data on antimicrobial agent prescription
42 (AAP) to date are limited in animals.

43 Antimicrobial agents are frequently prescribed in dogs and cats (Mateus et al., 2011;
44 Radford et al., 2011; Buckland et al. 2016), and there is evidence of development of resistance
45 in response to treatment¹ (Trott et al., 2004), and transmission of antimicrobial resistant isolates
46 between human beings and pets (Johnson et al., 2008a, b; Zhang et al., 2016). Specific guidance
47 for practice level prescription policies have been published^{5,6} (Beco et al., 2013a, b); however,
48 there is a need to understand how these are being applied in practice.

49 Data on human AAP in the United Kingdom (UK) are freely available, in part because
50 of a national health system.⁷ For animals, the Veterinary Medicines Directorate (VMD) is
51 constructing a central body collating data on AAP for the UK; however data currently available
52 cannot identify antimicrobial agents administered under the cascade prescribing system, which
53 species they have been prescribed to, practice level prescription variability or why the

¹ See:

http://www.ema.europa.eu/docs/en_GB/document_library/Scientific_guideline/2015/01/WC500181642.pdf
(accessed 15 July 2016).

² See: <https://www.gov.uk/government/publications/uk-one-health-report-antibiotics-use-in-humans-and-animals>
(accessed 15 July 2016).

³ See: <http://amr-review.org/home> (accessed 15 July 2016).

⁴ See: http://apps.who.int/iris/bitstream/10665/112642/1/9789241564748_eng.pdf (accessed 15 July 2016).

⁵ See: <http://www.bsava.com/Resources/PROTECT.aspx> (accessed 4 October 2016).

⁶ See: <http://www.fecava.org/content/guidelines-policies> (accessed 15 July 2016).

⁷ See: <http://fingertips.phe.org.uk> (accessed 15 July 2016).

54 antimicrobial agents were prescribed.⁸ Advances in veterinary health informatics provides
55 opportunities to fill this gap, particularly for companion animals where Electronic Health
56 Records (EHR) are most developed and accessible (O'Neill et al., 2014a).

57 Early studies of companion animal AAP in the UK were limited in size, but have
58 consistently pointed to frequent use of β -lactams (Mateus et al., 2011; Radford et al., 2011).
59 More recently, using a much larger data set, 25% of dogs and 21% of cats seen at veterinary
60 practices received at least one AAP over a 2 year period (2012-2014), the most frequent being
61 penicillins and cephalosporins (Buckland et al., 2016). Whilst such 'big data' studies have
62 started to report on AAP, this study aims to describe a near real-time, on-going, AAP
63 surveillance system from a diverse range of veterinary premises ($n = 457$) that also consider
64 AAP in a broad range of practitioner defined clinical presentations.

65 **Materials and methods**

66 *Data collection*

67 The Small Animal Veterinary Surveillance Network (SAVSNET) collected EHRs in
68 near real-time from booked consultations in volunteer UK veterinary practices (1 April 2014-
69 31 March 2016). A full description of the data collection protocol has been described by
70 Sánchez-Vizcaíno et al. (2015). A practice ($n = 216$) was defined as a single business, whereas
71 premise(s) ($n = 457$) included all branches that form a practice (see Appendix: Supplementary
72 Figure 1). Before submitting each consultation to SAVSNET, the practitioner selected one of
73 10 main presenting complaints (MPCs), consisting of a pre-determined list grouped into
74 healthy, unhealthy and post-operative categories (see Appendix: Supplementary Table 1). The
75 EHR further included product codes as text strings defined by individual practices.

76 *Antimicrobial agent identification*

⁸ See: <https://www.gov.uk/government/publications/veterinary-antimicrobial-resistance-and-sales-surveillance-2014> (accessed 15 July 2016).

77 The product codes of the EHR were utilised to identify AAP. A set of 52,267 codes
78 (extracted 26 August 2015) were manually categorised. Pharmaceutical products were defined
79 with reference to the VMD's Product Information Database for veterinary authorised products,
80 and the electronic Medicines Compendium (Datapharm Communications) for human
81 authorised products. An identifying string was ascribed to each antimicrobial agent product and
82 was used to identify the product code. This process was reiterated until all pharmaceutical and
83 non-pharmaceutical product codes were classified to further validate antimicrobial agent
84 identification. When applied to the complete list of 95,709 codes (extracted 31 March 2016),
85 416 antimicrobial agent identifying strings were utilised.

86 Where possible, product codes for antimicrobial agents were further characterised to
87 specific species authorisation and administration by systemic (oral or injectable) or topical
88 (topical, aural or ocular) routes. Whilst not all products were authorised for human use at the
89 time of the study, we considered all fluoroquinolones, macrolides and third generation
90 cephalosporins as highest priority critically important antimicrobial agents (HPCIA), as defined
91 by the World Health Organization (WHO).⁹

92 *Statistical analysis*

93 Consultation and prescription-level proportions and confidence intervals were
94 calculated to adjust for clustering (bootstrap method, $n = 5000$ samples) within premises and at
95 animal level within practices.¹⁰ Pearson correlations (t test to reject null hypothesis) were
96 performed to explore prescription frequency for dog and cat total, systemic and topical AAP as
97 a proportion of total submitted consultations for each premises. Paired t tests with Bonferroni
98 corrections were used for a matched pairs premises level sample to investigate total, systemic
99 and topical AAP as a proportion of total submitted consultations for each MPC.

⁹ See: <http://www.who.int/foodsafety/publications/antimicrobials-fourth/en/> (accessed 13 February 2017).

¹⁰ See: <http://cran.r-project.org/package=aod> (accessed 11 October 2016).

100 A mixed effects binomial regression model, incorporating practice and premise as
101 random effects, was utilised to examine quarterly variation in total, systemic and topical canine
102 and feline AAP as a proportion of total consultations. The variable time was categorised as an
103 ordinal variable into quarters of the year (Q1, Q2, Q3 and Q4) and included as a fixed effect.
104 Quarter was codified using two contrasting coding systems: (1) an orthogonal polynomial
105 method¹¹ to analyse for overall trend (see Appendix: Supplementary Table 2); and (2) a
106 backward differencing method¹² to investigate quarter-by-quarter variation in a backward
107 pairwise manner (e.g. Q1 2016 compared with Q4 2015). A further model was fitted for canine
108 and feline HPCIA prescription as a proportion of total AAP. A likelihood ratio test (LRT)
109 indicated that including practice and premise as random effects in all models provided the best
110 fit. Statistical significance was defined as $P < 0.05$ and all analyses were carried out using R
111 (version 3.2.3).¹³

112 **Results**

113 A total of 918,333 canine EHRs (from 413,870 dogs) and 352,730 feline EHRs (from
114 200,541 cats) were obtained from 216 veterinary practices (457 premises) from 1 April 2014 to
115 31 March 2016.

116 *Consultation and animal level*

117 The percentage of consultations where at least one antimicrobial agent was prescribed
118 (AAPC) was significantly greater for dogs (18.8%, 95% confidence interval, CI, 18.2-19.4)
119 than cats (17.5%, 95% CI 16.9-18.1). Systemic AAPC was significantly less frequent in dogs
120 (12.2%, 95% CI 11.7-12.7) than cats (14.8%, 95% CI 14.2-15.4), representing 64.9% (95% CI
121 63.8-66.0) and 84.5% (95% CI 83.9-85.2) of total canine and feline AAPC, respectively (paired
122 t test; $P < 0.001$). Topical AAPC was significantly more frequent in dogs (7.4% of

¹¹ See: http://www.ats.ucla.edu/stat/r/library/contrast_coding.htm#ORTHOGONAL (accessed 11 October 2016).

¹² See: http://www.ats.ucla.edu/stat/r/library/contrast_coding.htm#backward (accessed 11 October 2016).

¹³ See: <http://www.R-project.org/> (accessed 23 November 2016).

123 consultations, 95% CI 7.2-7.7) than cats (3.2%, 95% CI 3.1-3.3), representing 39.6% (95% CI
124 38.5-40.6) and 18.3% (95% CI 17.7-19.0) of AAPC, respectively ($P < 0.001$). Dogs and cats
125 were co-prescribed systemic and topical antimicrobial agents in 0.87% (95% CI 0.84-0.94) and
126 0.59% (95% CI 0.54-0.64) of total consultations, respectively. Significant positive correlations
127 were found between dogs and cats at premise level for total (0.62, 95% CI 0.56-0.67, $P < 0.001$),
128 systemic (0.61, 95% CI 0.54-0.66, $P < 0.001$) and topical (0.21, 95% CI 0.12-0.30, $P < 0.001$)
129 AAPC (Fig. 1).

130 Fig. 2 shows AAPC categorised by quarter. A significant negative linear trend was
131 observed for canine total and systemic AAPC, and feline total, systemic and topical AAPC (P
132 < 0.001 ; see Appendix: Supplementary Table 3). A significant negative trend by quarter was
133 observed for canine topical AAPC ($P < 0.001$). Results of quarter-by-quarter comparison
134 models can be found in Supplementary Table 4 (see Appendix).

135 Over the 2 year period, at the animal level, 28.4% (95% CI 27.2-29.7) of dogs were
136 prescribed an antimicrobial agent, compared with 23.3% (95% CI 22.3-24.4) of cats. When
137 route of administration was considered, 19.6% (95% CI 18.4-20.7) of dogs and 20.0% (18.9-
138 21.0) of cats were prescribed a systemic antimicrobial agent, and 12.9% (95% CI 12.3-13.5) of
139 dogs and 5.0% (95% CI 4.7-5.2) of cats were prescribed a topical antimicrobial agent.

140 Total AAPC was 35.5% (95% CI 34.5-36.5) of unhealthy dogs, 35.1% (95% CI 34.1-
141 36.1) of unhealthy cats, 7.4% (95% CI 6.7-8.0) of healthy dogs and 5.5% (95% CI 4.9-6.2) of
142 healthy cats. Systemic AAPC was more frequent in unhealthy cats (30.5%, 95% CI 29.5-31.5)
143 than unhealthy dogs (24.1%, 95% CI 23.1-25.0). The MPCs with the highest frequencies of
144 AAPC were pruritus in dogs (51.0%, 95% CI 49.8-52.2) and trauma in cats (53.5%, 95% CI
145 52.1-54.8). Antimicrobial agents were prescribed in a significantly greater proportion of dogs
146 than cats for gastroenteric ($P < 0.001$), pruritus ($P < 0.001$), kidney disease ($P < 0.001$), other
147 unwell ($P = 0.012$), vaccination ($P < 0.001$), other healthy ($P = 0.001$) and post-operative ($P =$

148 0.003) consultations. Cats were prescribed antimicrobial agents significantly more frequently
149 than dogs for respiratory ($P < 0.001$) and trauma ($P < 0.001$) consultations. Full results are
150 presented in Tables 1 and 2.

151 *Level of antimicrobial agent prescription*

152 A total of 218,700 canine and 71,089 feline AAPs were made from 215 practices (455
153 premises) in the UK.

154 *Authorisation* - For systemic AAP, 90.0% (95% CI 88.5-91.4) of canine and 92.9%
155 (95% CI 91.7-94.1) of feline AAPs were species authorised, with 0.6% (95% CI 0.2-0.9) and
156 5.2% (95% CI 4.0-6.5) authorised in other veterinary species; of these, 8.2% (95% CI 7.0-9.4)
157 and 1.7% (95% CI 1.4-2.1) were human authorised, 0.9% (95% CI 0.4-1.3) and 0.05% (95%
158 CI 0.03-0.07) were dual generic and 0.4% (95% CI 0.1-0.6) and 0.04% (95% CI 0.00-0.09)
159 were expired or of unknown authorisation, respectively. Metronidazole was the most frequently
160 prescribed human authorised systemic antimicrobial agent in dogs (96.7% of human authorised
161 systemic AAP, 95% CI 95.3-98.1) and cats (94.2%, 95% CI 92.1-96.3).

162 *Class of antimicrobial agent* - Clavulanic acid potentiated amoxicillin was the most
163 frequently prescribed antimicrobial agent in dogs (28.6% of total AAP, 95% CI 27.4-29.8) and
164 cefovecin was the most frequently prescribed antimicrobial agent in cats (36.2%, 95% CI 33.9-
165 38.5) (Tables 3, 4 and 5). Fusidic acid was the most frequently prescribed topical antimicrobial
166 agent in dogs (44.3% of topical AAP, 95% CI 43.1-45.4) and cats (55.1%, 95% CI 53.6-56.6).

167 *Highest priority critically important antimicrobial agents* - Canine and feline HPCIA
168 prescriptions were 5.4% (95% CI 4.6-6.1) and 39.2% (95% CI 36.8-41.7) of total AAPs
169 respectively. On consideration of temporal trend, for canine HPCIA prescription, a significant
170 positive cubic trend was noted ($P < 0.001$). Similarly, in cats, a significant positive linear trend
171 was found ($P < 0.001$) (see Appendix: Supplementary Tables 3 and 4). The most frequently

172 prescribed HPClAs in dogs were fluoroquinolones and in cats was cefovecin, a third generation
173 cephalosporin (Fig. 3).

174 *Main presenting complaint* - Total canine and feline AAPs summarised by MPCs are
175 shown in Supplementary Tables 5 and 6 (see Appendix). Clavulanic acid potentiated
176 amoxicillin was the most commonly prescribed antimicrobial agent in dogs for respiratory
177 conditions, trauma, tumours and kidney disease, as well as other unwell, post-operative and
178 other healthy MPCs. In cats, cefovecin was the most commonly prescribed antimicrobial agent
179 for respiratory conditions, pruritus, trauma, tumours and kidney disease, as well as other unwell,
180 post-operative and other healthy MPCs.

181 **Discussion**

182 In this study, EHRs were used to describe AAP in a large population of companion
183 animal veterinary premises. Quantitative differences in AAP were found between dogs and cats,
184 and according to MPC. AAPC decreased significantly over the course of the study in this
185 population of animals.

186 Broadly similar levels of total AAP were found in dogs and cats. However, when route
187 of administration was considered, dogs were significantly more likely to be prescribed topical
188 antimicrobial agents than cats, whereas cats were significantly more likely to be prescribed
189 systemic antimicrobial agents than dogs. Such differences may reflect an increased prevalence
190 of pruritus (and other dermatological diseases) in dogs compared to cats (Sánchez-Vizcaíno et
191 al., 2016). They may also reflect the challenge of giving oral and topical medication to cats
192 when compared to injectable antimicrobial agents (Burke et al., 2016).

193 Using data derived from EHRs, it was not possible to determine whether individual
194 prescriptions were appropriate, nor whether the overall frequency of AAP in this population
195 was appropriate. However, there was a significant reduction in canine and feline AAP within
196 this population over the 2 years of the study. Whether this reflects the success of awareness

197 campaigns is not known.^{14,15} It is possible that changes in AAP might reflect changes in other
198 aspects of veterinary activity, such as vaccination. Furthermore, previous human AAP
199 surveillance has noted short-term temporal variability that is not necessarily reflective of longer
200 term patterns.¹⁶ As a consequence, there is a need to for ongoing monitoring of AAP.

201 Buckland et al. (2016) found that 25.2% of dogs and 20.6% of cats in the UK received
202 systemic antimicrobial agents from 2012 to 2014. Whilst our results (2014-2016) were lower
203 for dogs (19.6%), they were similar for cats (20.0%). In a smaller study conducted in the UK
204 in 2010 (Radford et al., 2011), the proportion of consultations involving unhealthy animals
205 where systemic antimicrobial agents were prescribed was 35.1% for dogs and 48.5% for cats.
206 In our study, these values were lower (unhealthy dogs 24.1%, unhealthy cats 30.5%). It is
207 unclear whether differences between these studies reflect a reduction in frequency of
208 prescription of systemic antimicrobial agents, or are related to population differences or
209 methods used to identify AAP.

210 Considerable variation in AAPs according to premise was identified in our study, as
211 well as in the previous study by Radford et al. (2011). Premises that prescribed antimicrobial
212 agents more frequently to dogs also tended to prescribe more frequently to cats. Such a
213 correlation may be explained by geographical variation in risk (perceived or actual), either for
214 AMR or for bacterial infections capable of infecting both species. Other complex factors,
215 extending beyond the risk of antimicrobial agent responsive disease, can influence AAP
216 decisions, such as clinical experience, perceived owner and/or pet compliance and practice
217 policy (Hughes et al., 2012; Mateus et al., 2014).

218 It is not surprising that certain MPCs were more commonly associated with AAP,
219 suggesting that practitioners believe that the risk of infection responsive to antimicrobial agents

¹⁴ See: <http://www.fecava.org/content/guidelines-policies> (accessed 15 July 2016).

¹⁵ See: <http://www.bsava.com/Resources/PROTECT.aspx> (accessed 4 October 2016).

¹⁶ See: <http://ecdc.europa.eu/en/publications/Publications/Antimicrobial-consumption-europe-esac-net-2012.pdf>
(accessed 26 January 2017).

220 is higher in certain MPCs. Pruritus in dogs is frequently associated with bacterial pyoderma
221 (Summers et al., 2014) and was associated with the most frequent use of topical antimicrobial
222 agents in our study. However, acute respiratory disease in cats is generally considered to have
223 a viral origin, although primary bacterial disease has been described and secondary bacterial
224 infections can increase the severity of disease (Jacobs et al., 1993). Prescription of antimicrobial
225 agents in feline trauma may reflect a high frequency of cat bite abscesses associated with this
226 MPC (Radford et al., 2011; O'Neill et al., 2014b).

227 In dogs, clavulanic acid potentiated amoxicillin was the most frequently prescribed
228 antimicrobial agent, as found in previous studies (Mateus et al., 2011; Radford et al., 2011;
229 Buckland et al., 2016). In our study and that of Buckland et al. (2016), cefovecin was the most
230 frequently prescribed antimicrobial agent in cats, in contrast to previous studies, where
231 amoxicillin and clavulanic acid potentiated amoxicillin were more frequently prescribed
232 (Mateus et al., 2011; Radford et al., 2011). This suggests that there has been a recent shift in
233 choice of antimicrobial agents for cats. Prescription of cefovecin was common for MPCs
234 associated with authorised indications for use, such as pruritus and kidney disease¹⁷ (Burke et
235 al., 2016). However, cefovecin was also prescribed frequently in MPCs, such as respiratory and
236 gastroenteric disease in cats, where there was no apparent indication for prescription by the
237 datasheet¹ or practice prescribing policy.^{18,19} It is also possible that relying on MPCs as declared
238 by veterinary practitioners might fail to include other clinical conditions found during the same
239 consultation. Collection and analysis of clinical free text presents an opportunity to characterise
240 each consultation based on clinical signs and duration, which would provide further information
241 to support the rationale for any given prescription (Burke et al., 2016).

¹⁷ See: http://www.ema.europa.eu/docs/en_GB/document_library/EPAR_-_Product_Information/veterinary/000098/WC500062067.pdf (accessed 12 December 2016).

¹⁸ See: <http://www.fecava.org/content/guidelines-policies> (accessed 15 July 2016).

¹⁹ See: <http://www.bsava.com/Resources/PROTECT.aspx> (accessed 4 October 2016).

242 Although ceftiofur is not authorised for human use, it is a third generation
243 cephalosporin and is classified as an HPCIA.^{20,21} Relevant product information sheets state that
244 ceftiofur should be reserved for clinical conditions which have responded poorly, or are
245 expected to respond poorly, to other classes of antimicrobial agents.²² In our study, it was not
246 possible to determine to what extent the use of ceftiofur is in compliance with these
247 recommendations. A recent study showed that veterinary surgeons prescribing ceftiofur rarely
248 justified its use within the clinical narrative (Burke et al., 2016). Relative ease of administration
249 and duration of action, together aiding compliance, may be important motivating factors for the
250 use of ceftiofur in veterinary practice. We noted considerable variation in prescription of
251 ceftiofur between premises, suggesting that there are differences in cat populations,
252 presentations or justification for veterinary prescription. We further observed a slight increase
253 in overall HPCIA prescription in dogs and cats throughout the study, and that many of the most
254 commonly prescribed antimicrobial agents in both species are considered to be critically
255 important.²³

256 Whilst such large volumes of data provide new insights into AAP, the nature of these
257 data have their own inherent limitations. Quantification of AAP relies on practitioners charging
258 for antimicrobial agents through their practice management software, which means that any
259 antimicrobial agents not charged for will be missed. The SAVSNET population of practices is
260 recruited on the basis of convenience and so cannot necessarily be considered to be
261 representative of the wider UK population. In order to fully place findings in context, there is a
262 need for in depth analysis of the animal populations monitored. The use of the MPC function

²⁰ See: <http://www.who.int/foodsafety/publications/antimicrobials-fourth/en/> (accessed 13 February 2017).

²¹ See: <http://www.noah.co.uk/wp-content/uploads/2016/12/NOAH-briefing-on-CIAs-07122016.pdf> (accessed 14 February 2017).

²² See: http://www.ema.europa.eu/docs/en_GB/document_library/EPAR_-_Product_Information/veterinary/000098/WC500062067.pdf (accessed 12 December 2016).

²³ See: <http://www.who.int/foodsafety/publications/antimicrobials-fourth/en/> (accessed 13 February 2017).

263 allows all consultations to be coded in real time; variations in individual interpretation of the
264 MPC case definition are possible.

265 **Conclusions**

266 AAP frequency decreased from 2014 to 2016 in this population of dogs and cats in the
267 UK. Additionally, some MPCs were more likely to be associated with AAP than others, both
268 within and between the two species. There is considerable variability in AAP amongst different
269 premises and there is a need to understand factors that influence AAP at the individual animal,
270 owner and premise level, particularly for HPCIAAs. To aid responsible use, SAVSNET provides
271 a mechanism for participating practices to benchmark their prescription against anonymised
272 peers via an online portal. This and other studies are now providing the valuable tools and data
273 that the profession needs to ensure antimicrobial agents are used responsibly.

274 **Conflict of interest statement**

275 None of the authors of this paper have a financial or personal relationship with other
276 people or organisations that could inappropriately influence or bias the content of this paper.

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286 **Appendix: Supplementary material**

287 Supplementary data associated with this article can be found, in the online version, at
288 doi: ...

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359 **Table 1**

360 Canine antimicrobial agent prescription percentage (total, systemic and topical) by practitioner badged main
 361 presenting complaint calculated from total number of consultations for each category in a network of United
 362 Kingdom small animal veterinary premises.

Main presenting complaint	Dog						
	Number (%) of EHRs ^a	Total %	95% CI ^b	Systemic %	95% CI ^b	Topical %	95% CI ^b
Pruritus	62,655 (6.8)	51.0	49.8-52.2	25.5	24.2-26.9	30.0	29.0-31.0
Respiratory	14,359 (1.6)	42.2	40.5-44.0	40.4	38.7-42.2	2.7	2.2-3.2
Gastroenteric	38,954 (4.2)	39.4	37.0-41.7	38.2	35.8-40.6	1.7	1.2-2.2
Trauma	58,033 (6.3)	26.7	25.5-27.9	21.3	20.3-22.4	6.2	5.8-6.6
Kidney disease	2607 (0.28)	29.1	26.6-31.7	26.8	24.3-29.3	3.0	2.2-3.7
Tumour	20,938 (2.3)	22.0	21.1-23.0	17.5	16.7-18.3	5.4	5.0-5.8
Other unwell	156,197 (17.0)	32.8	31.8-33.8	20.3	19.5-21.2	13.9	13.4-14.5
Post-operative	98,753 (10.8)	13.0	12.2-13.8	9.9	9.3-10.5	3.5	3.1-3.8
Vaccination	277,246 (30.2)	4.3	3.9-4.7	1.4	1.1-1.7	3.0	2.8-3.2
Other healthy	188,582 (20.6)	11.8	10.7-13.0	7.0	6.1-7.8	5.3	4.8-5.9

363
 364 ^a Number (%) of electronic health records (EHRs). Relative occurrence of badged consultations as a frequency
 365 and as a percentage of total consultations.

366 **Table 2**

367 Feline antimicrobial agent prescription percentage (total, systemic and topical) by practitioner badged main
 368 presenting complaint calculated from total number of consultations for each category in a network of United
 369 Kingdom small animal veterinary premises.

Main presenting complaint	Cat						
	Number (%) of EHRs ^a	Total %	95% CI ^b	Systemic %	95% CI ^b	Topical %	95% CI ^b
Pruritus	13,749 (3.9)	33.5	31.9-35.2	24.9	23.3-26.6	10.3	9.5-11.1
Respiratory	7681 (2.2)	52.0	49.8-54.3	59.9	47.6-52.2	5.3	4.6-5.9
Gastroenteric	11,206 (3.2)	29.8	27.4-31.8	28.9	26.7-31.1	1.0	0.7-1.4
Trauma	22,796 (6.5)	53.5	52.1-54.8	50.1	48.8-51.4	4.3	4.0-4.7
Kidney disease	4009 (1.1)	19.6	17.9-21.3	18.9	17.2-20.6	0.7	0.5-1.0
Tumour	5330 (1.5)	21.3	19.8-22.7	19.8	18.3-21.3	1.7	1.4-2.0
Other unwell	72,189 (20.5)	30.5	29.5-31.6	24.9	23.9-26.0	6.5	6.3-6.8
Post-operative	32,136 (9.1)	11.1	10.0-11.9	9.6	8.7-10.6	1.7	1.4-2.0
Vaccination	115,394 (32.6)	2.5	2.2-2.8	1.4	1.2-1.6	1.2	1.1-1.3
Other healthy	68,236 (19.4)	10.5	9.1-11.9	8.4	7.1-9.6	2.4	2.1-2.7

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 371 ^a Number (%) of electronic health records (EHRs). Relative occurrence of badged consultations as a frequency
 372 and as a percentage of total consultations.

373 ^b 95% Confidence interval.

374 **Table 3**

375 Percentage breakdown of canine antimicrobial agent prescriptions by antimicrobial agent class prescribed for
 376 total, systemic and topical prescriptions from a network of United Kingdom small animal veterinary premises.

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Antimicrobial agent class	Total		Systemic		Topical	
	%	95% CI ^a	%	95% CI ^a	%	95% CI ^a
Aminoglycoside	12.0	11.4-12.6	0.1	0.0-0.2	29.1	28.0-30.2
Amphenicol	1.9	1.6-2.1	0.0	< 0.00	4.5	3.9-5.2
Other antimicrobial agent ^b	7.2	6.6-7.8	0.0	< 0.00	17.4	16.1-18.8
β-lactam	43.6	42.3-44.8	73.8	72.2-75.4	0.1	0.0-0.2
Fluoroquinolone	4.4	3.6-5.1	4.1	3.1-5.2	4.6	4.0-5.2
Fusidic acid	18.2	17.4-19.0	0.0	< 0.00	44.3	43.1-45.4
Lincosamide	4.7	4.2-5.2	7.9	7.0-8.8	0.0	< 0.00
Macrolide	0.2	0.0-0.3	0.3	0.0-0.6	0.0	< 0.00
Nitroimidazole	4.7	4.0-5.4	8.0	6.7-9.2	0.0	< 0.00
Nitroimidazole-macrolide	0.8	0.5-1.0	1.3	0.8-1.7	0.0	< 0.00
Rifamycin	0.0	< 0.00	0.0	< 0.00	0.0	< 0.00
Sulphonamide	1.5	1.1-1.9	2.5	1.9-3.2	0.0	< 0.00
Tetracycline	1.2	1.0-1.3	2.0	1.7-2.2	0.0	0.00-0.01

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394 ^a 95% Confidence interval.395 ^b Consists of polymyxin b sulphate; mupirocin; novobiocin; thymol and bronopol.

396 **Table 4**
 397 Percentage breakdown of feline antimicrobial agent prescriptions by antimicrobial agent class prescribed for
 398 total, systemic and topical prescriptions from a network of United Kingdom small animal veterinary premises.
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Class of antimicrobial agent	Total		Systemic		Topical	
	%	95% CI ^a	%	95% CI ^a	%	95% CI ^a
Aminoglycoside	4.5	4.2-4.8	0.2	0.1-0.3	22.1	20.7-23.6
Amphenicol	1.3	1.1-1.5	0.0	< 0.00	6.5	5.6-7.4
Other antimicrobial agent ^b	2.7	2.4-2.9	0.0	< 0.00	13.5	12.4-14.6
β-lactam	70.8	69.3-72.3	87.9	86.1-89.7	0.3	0.0-0.6
Fluoroquinolone	3.0	1.7-4.3	3.1	1.6-4.7	2.5	2.0-3.0
Fusidic acid	10.8	10.2-11.3	0.0	< 0.00	55.1	53.6-56.6
Lincosamide	4.1	3.5-4.7	5.2	4.4-5.9	0.0	< 0.00
Macrolide	0.05	0.01-0.09	0.07	0.01-0.12	0.0	< 0.00
Nitroimidazole	1.3	1.1-1.6	1.6	1.3-2.0	0.0	< 0.00
Nitroimidazole-macrolide	0.4	0.2-0.5	0.5	0.3-0.7	0.0	< 0.00
Rifamycin	0.0	< 0.00	0.0 ^c	< 0.00	0.0	< 0.00
Sulphonamide	0.05	0.03-0.07	0.06	0.03-0.09	0.0	< 0.00
Tetracycline	1.1	1.0-1.3	1.4	1.2-1.6	0.0	< 0.00

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418 ^a 95% Confidence interval.419 ^b Polymyxin b sulphate, mupirocin, novobiocin, thymol and bronopol.420 ^c One recorded prescription of rifampicin for systemic administration (authorised for oral administration).

421 **Table 5**

422 Percentage breakdown of β -lactam antimicrobial agent prescription by species and β -lactam sub-categories as a
 423 percentage of total and systemic antimicrobial agent prescriptions from a network of small animal veterinary
 424 premises in the United Kingdom.

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Class of antimicrobial agent	Total prescription				Systemic prescription			
	Dog		Cat		Dog		Cat	
	%	95% CI ^a	%	CI ^a	%	CI ^a	%	CI ^a
Amoxicillin	5.3	4.1-6.5	12.5	10.0-15.0	9.0	7.1-10.9	15.3	12.2-18.3
Other β -lactams ^b	0.4	0.0-0.8	0.07	0.01-0.13	0.5	0.0-1.3	0.02	0.00-0.05
First generation cephalosporin	8.4	7.8-9.0	0.4	0.3-0.5	14.2	13.2-15.3	0.5	0.4-0.6
Second generation cephalosporin	0.04	0.01-0.07	0.01	0.00-0.02	0.07	0.02-0.12	0.02	0.00-0.03
Third generation cephalosporin	0.9	0.7-1.0	36.2	33.9-38.5	1.5	1.3-1.8	45.1	42.1-48.2
Clavulanic acid potentiated amoxicillin	28.6	27.4-29.8	21.6	19.6-23.6	48.5	46.0-50.9	26.9	24.5-29.3
Penicillin	0.03	0.01-0.05	0.03	0.01-0.05	0.04	0.01-0.07	0.04	0.01-0.06
Total	43.6		70.8		73.8		87.9	

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427 ^a 95% confidence interval.428 ^b Ampicillin and cloxacillin.

Figure legends

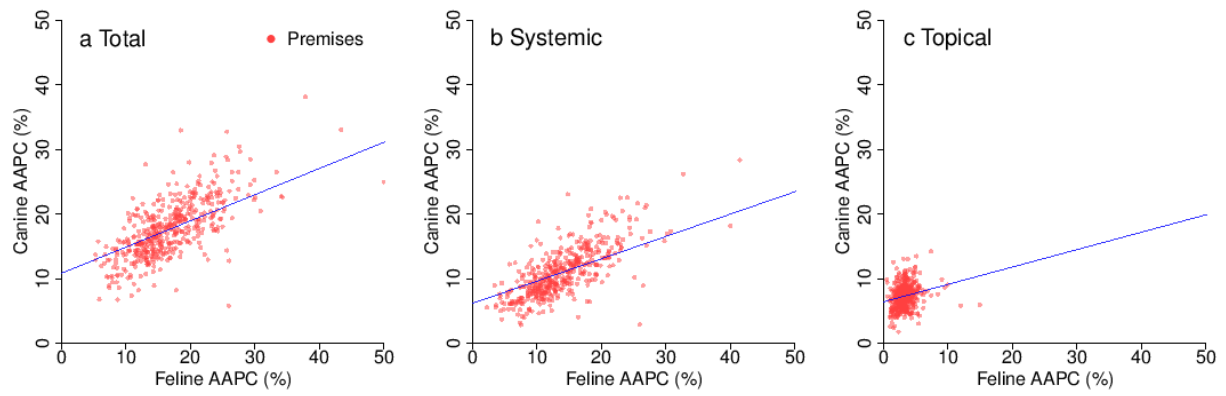
Fig. 1. Comparison of canine and feline antimicrobial agent prescription as a percentage of total consultations (AAPC) by premises ($n = 457$) split by (a) total, (b) systemic and (c) topical antimicrobial agent prescription.

Fig. 2. Comparison of (a) canine ($n = 918,333$ electronic health records) and (b) feline ($n = 352,730$) total, systemic and topical antimicrobial agent prescription as a percentage of total consultations (95% confidence interval) by quarter (Q2 2014-Q1 2016).

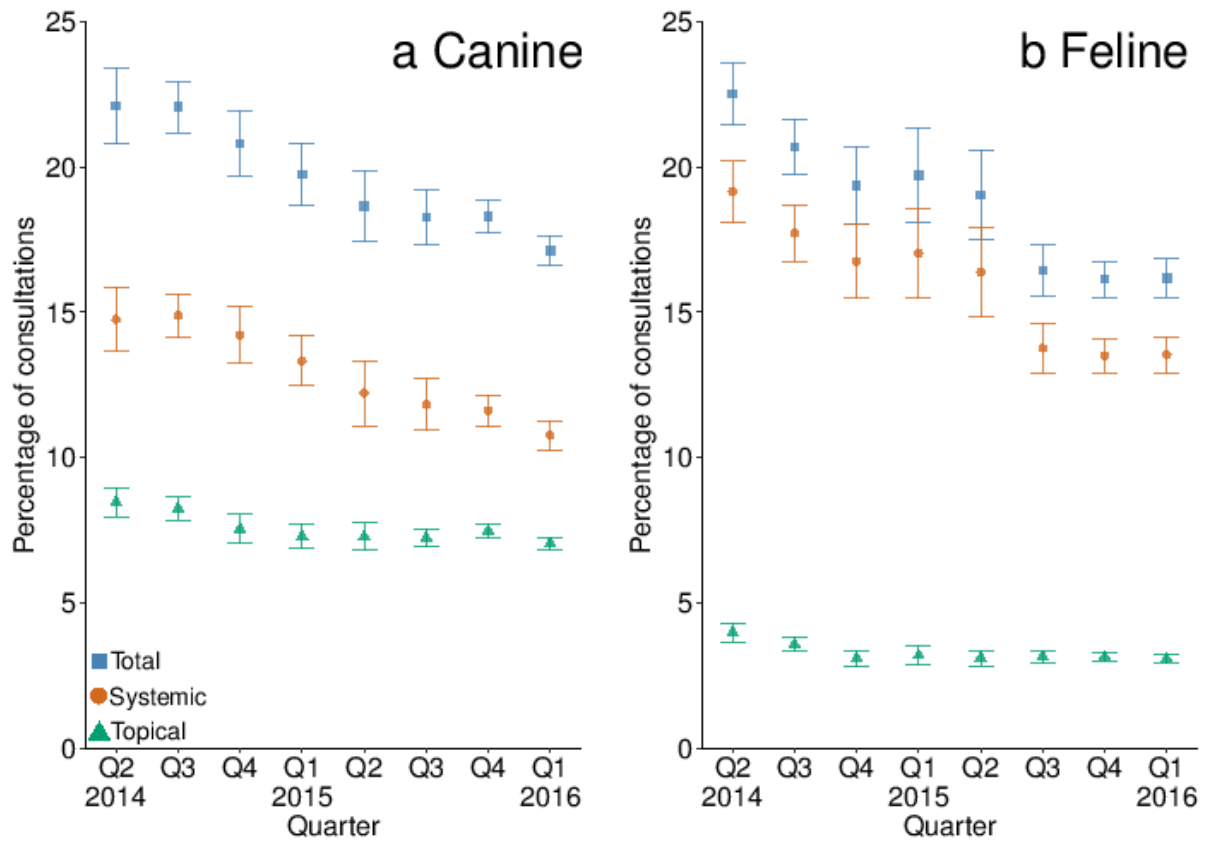
Fig. 3. Comparison of (a) canine and (b) feline highest priority ‘critically important antimicrobial agent’ (HPCIA) prescription as a percentage of total antimicrobial agent prescriptions (95% confidence interval) by quarter (Q2 2014-Q1 2016).

^b 95% Confidence interval.

Fig-1



Figr-2



Figr-3

