

1 **Original Article**

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4 **Difficulties in administration of oral medication formulations to pet cats; an e-survey of**  
5 **cat owners**

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**20 Abstract**

21           The purpose here was to determine the problems cat owners encounter in medicating  
22 their cats with orally administered drugs at home. The study was carried out as an open e-  
23 questionnaire survey addressed to cat owners in which we focused on the oral administration  
24 route. A total of 46 completed questionnaires were included in the survey. In the study, 46  
25 cats received 67 orally administered drugs. Approximately half of the drugs were registered  
26 for use in cats by the European Medicines Agency (54%), and there were also off-label drugs  
27 registered for human (36%) and canine medication (7.4%) and an *ex tempore* drug (3.0%).  
28 The owners were unable to give the doses as prescribed for their cats for one fourth of the  
29 medications (16/67). Drugs that were registered for feline medication were significantly more  
30 palatable than drugs registered for other species (odds ratio (OR) 4.9), and liquid formulations  
31 were significantly more palatable than solid formulations (OR 4.8). However, most of the  
32 owners (22/38) preferred a solid dosage form, while few (4/38) chose a liquid formulation.  
33 The results indicate that there is still a need for more palatable and easily administered oral  
34 drugs for cats.

35

36 *Keywords:* Compliance; Dosage form; Feline; Medicine; Palatability

## 37 **Introduction**

38           Cats are popular as companion animals and are generally well cared for throughout  
39 their lifespan. Owners are willing to provide their cats with good health care and medication  
40 for illnesses on veterinarians' recommendations (Jevring, 2005). However, owners often  
41 express difficulty in medicating their pets.

42

43           Medication compliance in companion animals may be compared with that in  
44 paediatrics, since owners and parents are responsible for medicating the patient (Grave and  
45 Tanem, 1999). All medications face similar challenges, such as the patient often being  
46 reluctant to take the drug, due to unpleasant taste and a high degree of noncompliance  
47 (Matsui, 1997; Haynes et. al., 2002). In veterinary medication, owners may prefer oral solid  
48 dosage forms, such as tablets, for long-term administration because they are familiar with  
49 how to administer these dosage forms to the pet (Khor et al., 2012). A choice of palatable  
50 formulations may increase pet acceptance (Thombre, 2004). Nevertheless, the degree of  
51 compliance varies widely. In compliance studies in dogs, 27-84% of the medications were  
52 given as prescribed (Bomzon, 1978; Grave and Tanem, 1999; Adams et al., 2005). In many  
53 cases the reason for non-compliance was that the owner was unable to follow the dosing  
54 regimen, rather than the dog's resistance to medication. The owners' medication experiences,  
55 as well as animal habituation in medication, also affect compliance. Compliance is clearly a  
56 multifaceted issue, in which the owners' abilities to follow the medication instructions play an  
57 important role (AAHA, 2003). Thus, pet compliance can be evaluated indirectly through  
58 owner consumption success or as the free-choice acceptance of the drug by the pet.

59

60           Few results are available on medication compliance in cats. Cats are more difficult to  
61 medicate than dogs, due to their discriminating nature (Thombre, 2004). They can be less

62 accustomed to being restrained than dogs and may display fear or resistance when medicated.  
63 The free-choice acceptability in cats is typically less than 50% for conventionally flavoured  
64 tablets (Ahmed and Kasraian, 2002). Various methods for administering oral solid dosage  
65 forms to cats have been used, such as forcing the animal to swallow a tablet ('dry  
66 swallowing') by placing it in the back of the oral cavity with the fingers or by a specific  
67 device, hiding the tablet in a highly palatable food or treat, and liquefying or crushing the  
68 tablet (Thombre, 2004; Bennett et al., 2010). Consumption success has rarely been evaluated  
69 in cats. For bitter-tasting drugs, owner-estimated consumption success rates have been as high  
70 as 90% for conventional tablets (dry swallowing) and 93% for extemporaneously prepared  
71 flavoured suspensions (Khor et al., 2012). In another study, dissolving oral film strips were  
72 easier to administer than gelatin capsules (Traas et al., 2010). Dosage form palatability has  
73 been assessed in studies in association with drug efficacy evaluations (Ahmed and Kasraian,  
74 2002; Gunew et al., 2008; Giraudel et al., 2010). In general, flavoured formulations (liquids  
75 or solids) are more palatable than conventional tablets, but comparison between studies is  
76 difficult, because various criteria for acceptance have been used. The criteria for determining  
77 palatability have been clarified quite recently by European Medicines Agency (EMA, 2014).  
78

79         The aim here was to determine the difficulties cat owners encounter in administering  
80 oral medication to their pets. Information is needed for developing palatable formulations for  
81 cats, as well as the methodology for assessment of acceptance. Detailed knowledge also  
82 supports veterinarians in counselling owners. More specifically, the purpose was to evaluate  
83 the palatability of dosage forms (expressed as free-choice acceptance), ease of administration  
84 of the dosage forms and ease in following the medication schedules (owner compliance). The  
85 hypothesis was that a marked number of owners experience difficulty with one or more of  
86 these aspects in medicating their cats at home.

## 87 **Materials and methods**

### 88 *Study outline*

89           The study was carried out as an open e-questionnaire survey addressed to cat owners  
90 responsible for medicating their cats at home. The platform chosen was a secured online tool  
91 provided by the University of Helsinki (e-form). The term compliance was adopted, as  
92 described by Cramer et al. (2008), in which the medication compliance (synonym adherence)  
93 refers to the degree or extent of conformity to the recommendations for daily treatment by the  
94 provider with respect to the timing, dosage and frequency. The theoretical framework of the  
95 “five interacting dimensions of adherence” (Sabaté, 2003) was implemented in exploring the  
96 study aims, and the e-questionnaire was adopted within the context of veterinary medication.

97

98           The questionnaire was developed in collaboration with veterinarians and pharmacists  
99 (seven specialists, University of Helsinki). In developing the questionnaire, we focused on the  
100 five dimensions of adherence and especially on the therapy-related factors, such as the  
101 medical regimen, ease in following the regimen and ease in administration of the dosage form  
102 to the cat. Pet compliance was also evaluated as free-choice acceptance of the drug by the cat.  
103 Originally, the oral, topical, eye and ear administration routes were included, but in this study  
104 we focused on orally administered dosage forms, since it is a common route of  
105 administration. The drugs were categorized according to their target species registration;  
106 feline, canine or other (human medication or *ex tempore* drugs).

107

108           The Viikki Campus Research Ethics Committee (University of Helsinki) approved the  
109 study protocol (Statement of Approval 14.1.2010). In a pilot study, preliminary suggestions  
110 by 14 cat owners resulted in rewording of one background question and of the instructions for  
111 answering the questions. The Checklist for Reporting Results of Internet E-Surveys

112 (CHERRIES) (Eysenbach, 2004) was followed in reporting the results of the e-questionnaire  
113 survey.

114

#### 115 *Recruitment of cat owners*

116 Owners who had medicated their cats were recruited from the Veterinary Teaching  
117 Hospital (University of Helsinki), four randomly chosen municipal clinics and four private  
118 veterinary clinics. One private cat clinic and the Veterinary Teaching Hospital were chosen  
119 without randomizing, due to the large numbers of their feline patients. The inclusion criteria  
120 for the cat owners was medication experience during the previous 3 months at home. The  
121 recruitment time span was 3 months.

122

123 The clinics were informed of the study via telephone calls, cover letters and e-mails.  
124 In all, 840 invitation letters were sent to the clinics that were willing to participate. The  
125 invitation letters included information on the study for the cat owners to support their decision  
126 to participate, and an Internet link to the questionnaire. The letters were distributed among the  
127 personnel of the clinics. Additionally, notices were distributed at the University Pharmacy  
128 (Viikki) and at one boarding cattery. Cat owners were also recruited from four Internet cat-  
129 themed discussion forums. One notice was posted at the University Library (Viikki).  
130 Participation in the study was voluntary, and no material incentives were provided to the  
131 participants.

132

#### 133 *Questionnaire*

134 Demographic data on the cats and their owners were collected (Table 1). The  
135 questions concerning the dosage form and administration of the drug are described in Table 2.  
136 A total of 46 completed questionnaires on 46 cats were included in the study. Each cat

137 received one or more drugs; in all, 67 orally administered drugs were included (Table 3).  
138 Three completed questionnaires on orally administered drugs were excluded, one because the  
139 owner lived abroad, another because the respondent was an employee of a rescue shelter and  
140 therefore not a regular cat owner and a third because the owner did not define the drug.  
141 Another 16 completed questionnaires were excluded, because the reported administration  
142 route was not oral. The clinics were asked to return the unused invitation letters; 447/850  
143 (53%) letters were returned.

144

#### 145 *Statistical methods*

146 Descriptive statistics on the number of observations and percentage frequencies were  
147 presented. Statistical analyses were performed using SAS System for Windows, version 9.3  
148 (SAS Institute Inc., Cary, NC, USA). The outcome variables concerning the owner's ability to  
149 give all the doses to the cat, free-choice acceptance of the drug, salivation, gagging or  
150 vomiting, following the medication schedule and ease in following the schedule were  
151 measured categorically, as were the explanatory variables (drug, dosage form, number of  
152 daily administrations, duration of medication, adverse effects). Frequency tables were  
153 constructed between the categories of the explanatory variables on all outcome variables. A  
154 cumulative logistic regression model was fitted to outcome variables that were evaluated  
155 through an agree-neutral-disagree-scale. The probabilities were modelled for responses  
156 having lower values (i.e. more agreement). Due to small cell frequencies, the original five-  
157 step Likert-scale (Strongly agree/Agree, Neither agree nor disagree, Disagree/ Strongly  
158 disagree) questions were transformed into three-class categorical variables, the middle class  
159 representing neutral, the lower and upper classes representing agreement and disagreement,  
160 respectively. The variable 'Number of doses administered daily' was transformed into a  
161 dichotomous variable (1 or less, more than 1). For the binary response 'Ability to give all

162 doses', a logistic regression model was fitted to the data, and the probability was modelled for  
163 the response value being 'yes'. All explanatory variables were analysed separately. In all  
164 models the explanatory variable in question was included in the model as the sole fixed factor.  
165 The effect of the cat was included in the model as a random effect. For all analyses, the odds  
166 ratios (ORs) and their 95% confidence intervals (CIs) were constructed to describe the group  
167 differences.

168

## 169 **Results**

### 170 *Dosage form and administration schedule*

171       Approximately half of the drugs were registered for use in cats by the EMA (Table 3).  
172 Most of these were solid dosage forms: tablets or capsules. Off-label drugs were registered for  
173 human (36%) or dog medication (7.4%) and one was an *ex tempore* tablet. All drugs were  
174 immediate-release formulations. In nearly half of the cases, the drug was used as long-term  
175 medication for chronic illnesses (Table 4). Most of the oral drugs were administered once or  
176 twice per day. However, the total administration frequency could have been higher if the cat  
177 had concurrent medications (orally administered or other routes of administration). This was  
178 the case for one third of the cats (n = 16). The maximum dosing requirement was five to six  
179 times per day.

180

181       When asked "*Did the cat accept the drug willingly?*", the owners agreed in 35% of  
182 the cases if the drug was registered for feline medication (Table 5). The free-choice  
183 acceptance was nearly five-fold higher (OR 4.9, P < 0.05) for drugs registered for cats than  
184 those registered for other species. A statistically significant relationship was also evident for  
185 the dosage form, with solutions and suspensions being more acceptable than tablets or  
186 capsules (OR 4.8). For the other questions, no statistically significant effects were observed



187 (Table 6). Most of the owners agreed that precisely following the medication schedule was  
188 easy.

189

#### 190 *Dosage form and administration practice*

191 In only two cases did the cat willingly accept the drug by consuming it ‘as a treat’  
192 (Table 7). Most of the drugs were introduced orally by dry swallowing (36/67). In addition,  
193 various methods for facilitating drug administration were described, such as mixing the drug  
194 in cat’s food or covered with a palatable treat. The cat either consumed the modified dosage  
195 form as such or else the form was introduced orally. It was also common practice for owners  
196 to first crush the tablet or dissolve it in water, after which the drug was mixed in the food or  
197 given by a syringe. If the package included an administration device, the owners found that  
198 using it was easy, and the guidance of the device was deemed adequate and clear (Table 8).

199

#### 200 *Difficulties encountered in drug treatment*

201 No medications in which all the doses were missed were reported. However, some  
202 doses were missed in one fourth (n = 16) of the cases (Table 6). At most, four doses were  
203 missed for one prescribed drug (n = 8), and most of these cases were associated with short-  
204 term medication (n = 6). In general, the missed doses occurred at the beginning (n = 6) or in  
205 the middle (n = 2) of the course of medication. In response to the open question “*Why could*  
206 *you not give all the doses to the cat?*” (n = 15), most of the answers (n = 11) were associated  
207 with adverse effects or the individual behaviour of the cat; the cat spat the drug out (n = 5),  
208 vomited (n = 2), salivated strongly (n = 2) or resisted the medication (n = 2). For some  
209 owners, scanty medication experience made it difficult to administer the drug in the beginning  
210 (n = 2). Answers to the question “*Why was it difficult to follow the medication schedule?*” (n  
211 = 15) included adverse effects (n = 2) or some other feline response (n = 2), such as the drug

212 given with the food was not accepted, due to lack of appetite. One third of the responses were  
213 related to the owners' working hours (n = 5) or, in the case of chronic illness, it was not  
214 always possible for the owner to commit to long-term medication (n = 2). One owner stopped  
215 the medication, because the symptoms disappeared, but then started it again after consulting  
216 the veterinarian.

217

218 For the question, "*What was the major problem related to the drug itself or*  
219 *administering the drug to the cat?*", 41 out of 46 responses were obtained. One fourth of the  
220 answers concerned the bad taste of the drug (n = 11). Other characteristics of the dosage form  
221 were also considered problematic; these included taking the right dose by splitting the tablet  
222 (n = 7), the tablet size being too large (n = 4) or the syringe for a liquid drug being a bit  
223 difficult to use (n = 1). Again, the individual behaviour of the cat and adverse effects were  
224 mentioned; the cat did not willingly take the drug, and the owner had to force it (n = 8) or the  
225 drug caused increased salivation or other side effects (n = 4). However, approximately half of  
226 the owners agreed with the question "*Did the administration of the drug become easier over*  
227 *time?*" (Table 8).

228

### 229 *Dosage form and administration preferences*

230 An open question "*What kind of dosage form would you prefer in medicating your*  
231 *cat?*" received 38 answers (38/46). A solid dosage form was the most preferred formulation;  
232 58% of the owners (n = 22) chose a tablet or capsule, while some also mentioned that the  
233 tablet should be small (n = 8), palatable (n = 2) or tasteless (n = 1). Four owners did not  
234 define the dosage form, only that the drug should be palatable or tasteless. Some (n = 3) chose  
235 either a solid or liquid formulation, if it were palatable enough, or a liquid or semisolid  
236 dosage form (n = 4). Few preferred injections (n = 2) or a pour-on formulation (n = 1).

237

238           The question “*What would you consider to be the easiest method for administering the*  
239 *drug to your cat?*” received 43 responses (43/46). Few (n = 3) answered that their cat  
240 consumed the drug willingly without any modification to the dosage form or administration  
241 aid. One third of the owners mentioned a solid dosage form by dry swallowing (n = 16). Some  
242 considered that the drug would be easiest to administer orally, but only after coating it with  
243 some palatable viscous material, such as butter (n = 4). In two cases, the owner specified that  
244 two persons would be needed to administer the drug orally. Some owners (n = 7) named  
245 liquids as the easiest dosage form; the drug should be either in a liquid form or the owner  
246 would dissolve a solid drug and then administer it orally with a syringe. For others, the easiest  
247 way was to give the drug hidden in the cat’s food or in a treat (n = 8). One owner mentioned  
248 that the cat should be hungry before medication, and one stated that the cat would lick a liquid  
249 drug from its fur. One considered injections as the easiest means of medication.

250

#### 251 *Owner counselling supported drug treatment*

252           The owners considered themselves well informed about the medication (88%) (Table  
253 8). In an open question, however, owners reported that they would have liked to have more  
254 information on administration methods (n=2) or how to take the right dose from the package  
255 (n=1). One owner had hoped for more information on the active pharmaceutical ingredient  
256 (drug-drug interactions, adverse effects and if the drug was sufficiently efficacious) and the  
257 presence of generic products on the market. For the question “*Did you read the drug*  
258 *description before administering the drug?*”, the owners agreed in 85% of the cases.

259

## 260 **Discussion**

261           The hypothesis was that a marked number of owners experience difficulty in

262 medicating their cats at home. The owners were unable to give the doses as prescribed for one  
263 fourth of the medications (24%), and therefore the owner compliance and owner consumption  
264 success can be considered to be 76%. The data indicate that off-label use of drugs is still  
265 common in the feline setting, since almost half of the drugs used were off-label. Pet  
266 compliance expressed as free-choice acceptance was higher for feline formulations than for  
267 off-label drugs, yet only in 35 % of these feline formulations did the owner agree that the cat  
268 took the drug willingly. Solutions and suspensions were significantly more acceptable than  
269 solid dosage forms. It is noteworthy that these were practically all (excluding one off-label  
270 drug) feline medications in which the palatability issues were considered in formulation  
271 development by the pharmaceutical company. In other studies, oral suspensions registered for  
272 feline medication were also well accepted by cats, either given alone or mixed in food (Litster  
273 et al., 2007; Gunew et al., 2008). Product development for feline medication is apparently  
274 progressing in the right direction, although there is further need for drugs registered  
275 specifically for cats, based on the amount of off-label usage.

276

277       Most consumption failures were related to the individual behaviour of the cat (the cat  
278 resisted the medication or spat the drug out) or adverse effects (such as strong salivation).  
279 Some failures may have been caused by the bad taste of the drug, which the owners presented  
280 as an important reason. Interestingly, the owners preferred tablets as a dosage form, although  
281 there was significantly better pet compliance with the liquids. In most cases, tablets were  
282 administered orally 'dry swallowing'. Such preference may be explained by the owner being  
283 familiar with the administration technique (Khor et al., 2012). The 'dry swallowing' method  
284 applies when medicating cats with a compliant nature, but is problematic in cases in which  
285 the cat shows fear or resistance. In forcing, the cat may become even more unwilling to take  
286 the drug, which may negatively affect both the human animal relationship and owner

287 compliance. The safety of the person administering the medicine should also be considered  
288 (Bennett et al., 2010). An alternative approach to support pet compliance could be training the  
289 pet to be more favourable to handling and drug administration already as a kitten. In  
290 medicating cats with chronic illnesses, maintaining compliance is crucial (Jevring, 2005). In  
291 our data, nearly half of the medications were long-term.

292

293 Administration practice varied markedly and owners described several methods for  
294 avoiding consumption failure. The tablets may have been easier to handle than liquid dosage  
295 forms, because they (if small enough) could be given inside a treat or with other palatable  
296 material. Nevertheless, it is known that solid dosage forms given by dry swallowing can  
297 become trapped in the oesophagus and cause esophagitis or even stricture formation (German  
298 et al., 2005; Beatty et al., 2006). Therefore, several authors have recommended that solid oral  
299 dosage forms should be given with a water bolus or a small amount of food to facilitate  
300 oesophageal clearance, or otherwise with an administration aid such as a pill delivery treat or  
301 flavoured liquid (Graham et al, 2000; Westfall et al., 2001; Bennett et al., 2010). Furthermore,  
302 owners facilitated administration by giving the drug mixed in the cat's food. The food  
303 approach may, however, be problematic, because the food effects on bioavailability of the  
304 drug substance or the dosage form performance may not be easily managed (Ahmed and  
305 Kasraian, 2002).

306

307 Our results suggest that the free-choice acceptance and ease of dosage form  
308 administration are still problematic in feline medication. This often seems to be related to the  
309 use of off-label drugs. Thus, there is need for pharmaceutical development of solid dosage  
310 forms for cats in particular. Species-specific considerations are needed, including tablet size  
311 and formulations that enhance free-choice acceptance and ease in swallowing. Dosage form

312 characteristics, including small tablet size, palatability or tastelessness were preferred by cat  
313 owners in the present study.

314

315         One limitation of the study was the relatively small sample size. Information on a total  
316 of 46 cats and 67 orally administered drugs was provided in the study. The low survey  
317 response rate made the study subject to non-response bias, and the results may have led to  
318 overestimation of the owners' compliance and abilities to medicate the cat, using oral  
319 medication formulations. The more compliant owners were more likely to have been  
320 respondents. As an open e-questionnaire, the survey relied on a convenience sample, which  
321 may have led to undercoverage of noncompliant owners or owners who have limited access to  
322 the Internet. The retrospective study setting increased the risk for pet owner recall bias for  
323 self-reported data.

324

### 325 **Conclusion**

326         Our results confirm that there is need for developing new palatable and easily  
327 administered drugs for feline medication. Medications registered for feline administration  
328 were more palatable than off-label drugs, even though the owners reported that their cats took  
329 the drugs willingly in only 35 % of these cases. To improve the willingness of cats to ingest  
330 the drugs, as well as owner compliance, the individual nature of the cat should be taken into  
331 account, and not only effective, but also easily administrable dosage forms should be  
332 developed for cats.

333

334

335 **References**

- 336 AAHA (American Animal Hospital Association) (2003) AAHA study finds millions of pets  
337 aren't getting maximum health care. *Journal of the American Veterinary Medical Association*  
338 222, 1488-1488
- 339 Adams, V.J., Campbell, J.R., Waldner, C.L., Dowling, P.M., Shmon, C.L. (2005) Evaluation  
340 of client compliance with short-term administration of antimicrobials to dogs. *Journal of the*  
341 *American Veterinary Medical Association* 226, 567-574
- 342 Ahmed, I., Kasraian, K. (2002) Pharmaceutical challenges in veterinary product development  
343 *Advanced Drug Delivery Reviews* 54, 871
- 344 Beatty, J.A., Swift, N., Foster, D.J., Barris, V.R. (2006) Suspected clindamycin-associated  
345 oesophageal injury in cats: five cases. *Journal of Feline Medicine and Surgery* 8, 412-419
- 346 Bennett, A.D., MacPhail, C.M., Gibbons, D.S., Lappin, M.R. (2010) A comparative study  
347 evaluating the esophageal transit time of eight healthy cats when pill with the FlavoRx pill  
348 glide versus pill delivery treats. *Journal of Feline Medicine and Surgery* 12, 286-290
- 349 Bomzon, L. (1978) Short-term antimicrobial therapy - A pilot compliance study using  
350 ampicillin in dogs. *Journal of Small Animal Practice* 19, 697-700
- 351 Cramer, J.A., Roy, A., Burrell, A., Fairchild, C.J., Fuldeore, M.J., Ollendorf, D.A., Wong,  
352 P.K. (2008) Medication Compliance and Persistence: Terminology and Definitions. *Value in*  
353 *Health* 11, 44-47
- 354 EMA (European Medicines Agency) (2014) Guideline on the demonstration of palatability of  
355 veterinary medicinal products. EMA/CVMP/EWP/206024/2011  
356 [http://www.ema.europa.eu/docs/en\\_GB/document\\_library/Scientific\\_guideline/2014/07/WC5](http://www.ema.europa.eu/docs/en_GB/document_library/Scientific_guideline/2014/07/WC500170030.pdf)  
357 [00170030.pdf](http://www.ema.europa.eu/docs/en_GB/document_library/Scientific_guideline/2014/07/WC500170030.pdf) (Accessed 27<sup>th</sup> June 2016)
- 358 Eysenbach, G. (2004) Improving the Quality of Web Surveys: The Checklist for Reporting  
359 Results of Internet E-Surveys (CHERRIES). *Journal of Medicinal Internet Research* 6(3), e34
- 360 German, A.J., Cannon, M.J., Dye, C., Booth, M.J., Pearson, G.R., Reay, C.A., Gruffyd-Jones,  
361 T.J. (2005) Oesophageal strictures in cats associated with doxycycline therapy. *Journal of*  
362 *Feline Medicine and Surgery* 7, 33-41
- 363 Giraudel, J.M., Gruet, P., Alexander, D.G., Seewald, W., King, J.N. (2010) Evaluation of  
364 orally administered robenacoxib versus ketoprofen for treatment of acute pain and  
365 inflammation associated with musculoskeletal disorders in cats. *American Journal of*  
366 *Veterinary Research* 71, 710-719
- 367 Graham, J.P., Lipman, A.H., Newell, S.M., Roberts, G.D. (2000) Esophageal transit of  
368 capsules in clinically normal cats. *American Journal of Veterinary Research* 61, 655-657
- 369 Grave, K., Tanem, H. (1999) Compliance with short-term oral antibacterial drug treatment in  
370 dogs. *Journal of Small Animal Practice* 40, 158-162
- 371 Gunew, M.N., Menrath, V.H., Marshall, R.D. (2008) Long-term safety, efficacy and  
372 palatability of oral meloxicam at 0.01-0.03 mg/kg for treatment of osteoarthritic pain in cats.

- 373 *Journal of Feline Medicine and Surgery* 10, 235-241
- 374 Haynes, R.B., McDonald, H.P., Garg, A.X. (2002) Helping patients follow prescribed  
375 treatment: clinical applications. *Journal of the American Medical Association* 288, 2880-2883
- 376 Jevring, C. (2005) Compliance in veterinary practice. *European Journal of Companion  
377 Animal Practice* 15, 205-209
- 378 Khor, K.H., Campbell, F., Rathbone, M.J., Greer, R.M., Mills, P.C. (2012) Acceptability and  
379 compliance of atenolol tablet, compounded paste and compounded suspension prescribed to  
380 healthy cats. *Journal of Feline Medicine and Surgery* 14, 99-106
- 381 Litster, A., Moss, S., Honnery, M., Rees, B., Edingloh, M., Trott, D. (2007) Clinical efficacy  
382 and palatability of pradofloxacin 2.5% oral suspension for the treatment of bacterial lower  
383 urinary tract infections in cats. *Journal of Veterinary Internal Medicine* 21, 990-995
- 384 Matsui, D.M.S. (1997) Drug compliance in pediatrics. *Pediatric Clinics of North America* 44,  
385 1-14
- 386 Sabaté, E. Editor (2003) Adherence to long term therapies: Evidence for action. Section II:  
387 Improving adherence rates: Guidance for countries, Chapter V, 27-30. World Health  
388 Organization. [www.who.int/chp/knowledge/publications/adherence\\_full\\_report.pdf](http://www.who.int/chp/knowledge/publications/adherence_full_report.pdf) (Accessed  
389 30<sup>th</sup> June 2016)
- 390 Thombre, A.G. (2004) Oral delivery of medications to companion animals: Palatability  
391 considerations. *Advanced Drug Delivery Review* 56, 1399-1413
- 392 Traas, A.M., Fleck, T., Ellings, A., Mahabir, S., Stuebner, K., Brown, D.C., Durso, D.,  
393 DiGregorio, M., Bode, L., Kievit, K.I., McCall, R. (2010) Ease of oral administration and  
394 owner-perceived acceptability of triglyceride oil, dissolving thin film strip, and gelatin  
395 capsule formulations to healthy cats. *American Journal of Veterinary Research* 71, 610-614
- 396 Westfall, D.S., Twedt, D.C., Steyn, P.F., Oberhauser, E.B., VanCleave, J.W. (2001)  
397 Evaluation of esophageal transit of tablets and capsules in 30 cats. *Journal of Veterinary  
398 Internal Medicine* 15, 467-470



## TABLES

Table 1. Demographic data on cats and their owners (a total of 67 oral medications in 46 cats).

Pedigree cats	25 (54%)
Domestic cats <sup>a</sup>	21 (46%)
Age of the cat (years)	0.25 to 17 years (average 7.1 years, SD $\pm$ 5.1)
Castrated males	23 (50%)
Spayed females	13 (28%)
Intact males	6 (13%)
Intact females	4 (9%)
Number of cats in household	median 2.5 (average 3.6, SD $\pm$ 2.8, range 1 to 12 cats)
Persons responsible for medicating the cat <sup>b</sup>	
One	28 (61%)
Two	17 (37%)
Three	1 (2%)

All owners reported previous experience in medicating cats (the level of experience was not rated)

<sup>a</sup> Short hair or long hair domestic cat

<sup>b</sup> One of which was the owner

Table 2. Questions concerning the dosage form and administration of medication to the cat. The type of question is indicated by the letters a, b, c and d.

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Specify the dosage form administered to the cat \*  
 How many doses were administered daily? a  
 How many drugs were administered concurrently? a  
 If the cat had concurrent medications, what was the total daily administration frequency? a  
 What was the duration of drug treatment? b

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How was the drug given to the cat? b \*\*  
 Did you use some kind of administration aid in giving the drug to the cat? b \*\*\*  
 Did the cat accept the drug willingly? c  
 Did the salivation of the cat increase while giving the drug? c  
 Did the cat begin to gag or vomit during ingestion of the drug? c

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Did the drug cause any adverse effects? Which adverse effects? b \*\*\*\*

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Was it easy to follow the medication schedule? c  
 Did you follow the medication schedule precisely? c  
 Why was it difficult to follow the medication schedule? d

Was it easy to provide a single dose from the package? c  
 Was the guidance of the administration device provided in the package adequate and clear? c  
 Was the use of the administration device easy? c  
 Did the administration of the drug become easier over time? c

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Were you able to give all the doses? c  
 How many doses were missed? d  
 When were the doses missed? d  
 Why could you not give all the doses to the cat? d

What was the major problem related to the drug itself or administering the drug to the cat? d  
 What kind of dosage form would you prefer in medicating your cat? d  
 What would you consider to be the easiest method for administering the drug to your cat? d

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Did you read the drug description before administering the drug? c  
 Did you receive enough information about the correct use of the drug? c  
 What kind of information would you have liked to have more of? d

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a Numeric field

b Multiple choice question

c Likert scale (Strongly agree/Agree, Neither agree nor disagree, Disagree/Strongly disagree)

d Open question

\* The drug was picked from a list of names and images of 50 commonly used drugs registered for feline medication in the European Medicines Agency (EMA). If the drug was not on the list, it was reported in open question.

\*\* Orally (“dry swallowing”), from a cat’s food bowl mixed with food, inside a treat, crushed, dissolved, other (please describe how), cat refuses to take the medication

\*\*\* Inside a treat, with palatable viscous paste, pill gun, dispensed in a gelatin capsule, other administration aid (please describe which?), no administration aid was used

\*\*\*\* Diarrhoea, constipation, nausea, dizziness and/or fatigue, skin symptoms other side effects, (please describe which)

Adoptive questioning was used; where appropriate the question was conditionally displayed, based on response to the previous question. Review of the answers by the respondents (back button) and completeness check by the system for mandatory items (highlighted) were enabled before the questionnaire was submitted.

Table 3. Frequency of different types of oral dosage forms (a total of 67 medications in 46 cats).

	Tablet/ capsule	Solution/ suspension	Total
	n	n	n (%)
Registered for cats	25 <sup>a</sup>	11 <sup>b</sup>	36 (54)
Registered for dogs	5 <sup>c</sup>	0	5 (7)
Drugs for human medication (off-label drugs)	23 <sup>d</sup>	1 <sup>e</sup>	24 (36)
<i>Ex tempore</i> drugs	2 <sup>f</sup>	0	2 (3)
Total n (%)	55 (82%)	12 (18%)	67 (100)

<sup>a</sup> Axilur 250 mg or 500 mg tabl., Clavubactin 50/12,5 mg or 250/62,5 mg tabl., Drontal 230/20 mg tabl., Fortekor 2,5 mg or 5 mg tabl., Medrol vet 4 mg tabl., Perlutex vet 5 mg tabl., Synulox 40 mg, 200 mg or 400 mg tabl., Xeden 15 mg or 50 mg tabl.

<sup>b</sup> Amovet 50 mg/mL susp., Metacam 0,5 mg/mL susp., Flubenol 44 mg/mL paste, Mirrix 11,5% paste

<sup>c</sup> Atopica 25 mg tabl., Barbivet 30 mg tabl., Cerenia 16 mg tabl., Furovet 20 mg tabl., Metacam 1 mg tabl.,

<sup>d</sup> Anafranil 10 mg tabl., Atarax 25 mg tabl., Clinaxin 75 mg caps., Dexametason 1,5 mg tabl., Dilzem 30 mg tabl., Disperin 50 mg tabl., Doximycin 100 mg tabl., Norvasc 5 mg tabl., Pepcid 10 mg tabl., Prednisolon 5 mg tabl., Tenobloc 25 mg tabl., Trikozol 200 mg tabl., Tyrazol 5 mg tabl.

<sup>e</sup> Primperan 1mg/mL sol.

<sup>f</sup> Tylosin tabl. (University Pharmacy)

Table 4. Administration frequency of oral dosage forms and duration of the treatment (a total of 67 medications in 46 cats). Number of concurrently administered medications (orally administered and other routes of administration, if any) and subsequent administration frequency for the cat (n = 46).

	One n (%)	Two n (%)	Three or more n (%)	Less than one n (%)
Administration times per day for one medication	30 (45)	33 (49)	1 (1.5)	3 (4.5)
Medications for one cat	30 (65)	13 (28)	3 (7)	0
Total administration times per day for one cat	17 (37)	17 (37)	10 (22) <sup>a</sup>	2 (4)
Duration of drug treatment				
Short term (10 days or less)				
Long term (more than 10 days)	38 (57)			
	29 (43)			

<sup>a</sup> Four, three, one and two cats were medicated three, four, five and six times per a day, respectively

Table 5. Drug and dosage form as explanations for the question “Did the cat accept the drug willingly?” (a total of 67 medications in 46 cats).

		NA	Agree	Neither agree nor disagree	Disagree
		n	n (%)	n (%)	n (%)
Drug <sup>a</sup>	Registered for cats	2	12 (35.3)	4 (11.8)	18 (52.9)
	Registered for dogs	0	1 (20.0)	0	4 (80.0)
	Other	2	2 (8.3)	2 (8.3)	20 (83.3)
	Total	4	15 (23.8)	6 (9.5)	42 (66.7)
Drug formulation <sup>b</sup>	Tablet/capsule	4	9 (17.6)	5 (9.8)	37 (72.5)
	Solution/suspension	0	6 (50.0)	1 (8.3)	5 (41.7)
	Total	4	15 (23.8)	6 (9.5)	42 (66.7)

NA = Not answered

<sup>a</sup> Statistically significant ( $P < 0.05$ ) when drug registered for cats was compared with other drugs, odds ratio OR (95% confidence interval CI) 4.947 (1.086 – 22.525).

<sup>b</sup> Statistically significant ( $P < 0.05$ ) when solution/suspension was compared with tablet/capsule, OR (95% CI) 4.776 (1.072 – 21.274).

Table 6. Other questions concerning the dosage form and administration schedule (a total of 67 medications in 46 cats).

	NA	Agree	Neither agree nor disagree	Disagree
	n	n (%)	n (%)	n (%)
Was the owner able to give all the doses?	0	51 (76.1)	0	16 (23.9)
Did the salivation of the cat increase while giving the drug?	4	22 (34.9)	7 (11.1)	34 (54.0)
Did the cat begin to gag or vomit during ingestion of the drug?	7	12 (20.0)	3 (5.0)	45 (75)
Was it easy to follow the medication schedule?	0	58 (86.6)	2 (3.0)	7 (10.4)
Did the owner follow the medication schedule precisely?	0	53 (79.1)	7 (10.4.)	7 (10.4)

NA = Not answered. No statistically significant effects were found when the drug, dosage form, daily administration times, duration of the medication and adverse effects were tested as explanations for the question.

Table 7. Questions concerning administration practice (a total of 67 medications in 46 cats).

	Dry swallowing	Mixed with food in a bowl	Inside a treat <sup>b</sup>	Crushed or dissolved <sup>c</sup>	Other <sup>d</sup>
How was the drug given to the cat? <sup>a</sup>	36	8	11	13	15
Did you use some kind of administration aid in giving the drug to the cat?	No aid		Inside a treat (9) Palatable viscous paste (2)		Pill gun (1) Syringe (9) Gelatin capsule (2) Spoon (1)

<sup>a</sup> Multiple responses possible, <sup>b</sup> inside a treat or the tablet was covered with Easypill® or butter, <sup>c</sup> crushed and then mixed with food (n = 5), dissolved in water and then mixed with food or given with syringe (n = 8), <sup>d</sup> other: some kind of administration aid was used (n = 13), in two cases the drug was consumed as a treat. No answers were obtained for the category “cat refuses to take the medication”.

Table 8. Other questions concerning administration practice and administration device (a total of 67 medications in 46 cats).

	NA	Agree	Neither agree nor disagree	Disagree
	n	n (%)	n (%)	n (%)
Was it easy to provide a correct single dose from the package?	0	49 (73.1)	1 (1.5)	17 (25.4)
Was the guidance of the administration device adequate and clear? <sup>a</sup>	0	39 (58.2)	24 (35.8)	4 (6.0)
Was the use of the administration device easy? <sup>a</sup>	0	32 (47.8)	31 (46.2)	4 (6.0)
Did the administration of the drug become easier over time?	0	35 (52.2)	12 (17.9)	20 (29.9)
Did you read the drug description before administering the drug?	0	57 (85.0)	2 (3.0)	8 (12.0)
Did you receive enough information about the correct use of the drug?	0	59 (88.0)	4 (6.0)	4 (6.0)

NA = Not answered

<sup>a</sup> Administration device provided in the drug package was related to solution/suspension formulations (n = 12)