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TITLE: Gastric dilation-volvulus in dogs attending UK emergency-care veterinary practices: prevalence, risk factors and survival

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- 50 0.001). Odds increased as dogs aged up to 12 years and neutered male dogs had 1.3
- 51 (95% CI 1.0-1.8, P = 0.041) times the odds compared with entire females. Of
- 52 presumptive gastric dilation-volvulus cases that presented alive, 49.7% survived to
- 53 discharge but 79.3% of surgical cases survived to discharge.
- 54 **Clinical importance:** Approximately 80% of surgically managed cases survived to
- 55 discharge. Certain large breeds were highly predisposed.
- 56

# 57 Keywords

- 58 VetCompass, gastric dilation-volvulus, GDV, emergency, bloat
- 59

### 60 Abbreviations

- 61 CI confidence interval
- 62 EPR electronic practice record
- 63 GDV gastric dilation-volvulus
- 64 IQR interquartile range
- 65 OR odds ratio
- 66 RVC Royal Veterinary College
- 67 SD standard deviation

68

## 69 Introduction

- 70 Gastric dilation-volvulus (GDV) is generally an acute and life-threatening condition with
- severe multi-systemic effects in dogs (Brockman 2012, Sharp and Rozanski 2014). The
- 72 precise pathophysiology of GDV is unclear but the presentation is characterised by rapid

73 accumulation of various combinations of gas, fluid and ingesta in the stomach, with 74 increased intra-gastric pressure (Hendriks and others 2011). Dilation of the stomach and 75 rotational twist along its horizontal axis compress the major abdominal blood vessels and 76 impair venous return to the heart, leading to impaired cardiac output and shock 77 (Glickman and others 1994). If left untreated, severe systemic hypoperfusion can ensue 78 via multiple mechanisms (hypovolaemic, distributive and obstructive shock) along with 79 additional respiratory compromise and decreased tidal volume that results from 80 restriction of diaphragmatic excursions by the enlarged stomach (Monnet 2003). 81 Diagnosis is commonly based on a combination of characteristic historical and physical 82 examination findings (notably abdominal distension and pain, tachycardia, poor 83 peripheral pulses and unproductive retching), combined with evidence of gastric 84 malpositioning and enlargement apparent from diagnostic imaging (Tivers and Brockman 85 2009).

A typical combination of significant pain and a high mortality rate make GDV both a clinical and welfare concern in affected dogs. Among UK pedigree dogs participating in a survey-based retrospective study, GDV was reported to cause 2.5% of all deaths, with a median age at death of 7.9 years (Evans 2010). A US study of large and giant dog breeds reported that 16% of deaths in these breeds were from GDV (Glickman and others 2000) and that 28.6% of GDV cases died directly from the disorder (Glickman 2000).

GDV is a complex disorder with multiple interacting inherited and environmental factors reported to affect the probability of a GDV event (Bell 2014). Reported prevalence values for GDV in dogs varies widely across differing breeds and populations but it is generally recognised as a disorder that mainly affects large and giant breeds (Glickman and others 1994). Purebred dogs are reported to be predisposed to GDV compared with crossbred dogs, with reported odds ratios (OR) ranging from 1.8 (95% confidence interval [CI] 1.1-2.9) (Bellumori and others 2013) to 2.5 (95% CI 2.1-3.0) (Glickman and others 100 1994). Larger body size (Glickman and others 1994, Glickman and others 2000) and 101 deep-chested conformations (Bell 2014) have been reported as risk factors for GDV. 102 Many breeds are reported as predisposed to GDV, and these include great Dane, 103 bloodhound, German shepherd, standard poodle, grand bleu de Gascogne, German 104 pointer, akita, Irish setter, Weimaraner and Neapolitan mastiff (Bell 2014, Brockman 105 1995, Evans 2010, Glickman 2000). Advancing age has also been reported to 106 substantially increase risk of GDV (Elwood 1998, Glickman 2000, Theyse and others 107 1998) but evidence to support a sex predisposition to GDV has been more equivocal 108 (Eggertsdóttir and Moe 1995, Glickman and others 1997, Glickman and others 1994, 109 Glickman and others 2000).

110 Research using primary-care veterinary clinical records has been recommended as a 111 means to generate reliable and generalisable information on the occurrence and risk 112 factors for disorders affecting the wider animal population (O'Neill and others 2014a). 113 This study aimed to analyse a database of merged emergency-care practice electronic 114 patient records (EPRs) to estimate the prevalence of presumptive GDV diagnoses among 115 an emergency-care caseload of dogs in the UK and to evaluate demographic risk factors 116 for the occurrence of GDV. The study additionally aimed to report on clinical 117 management and survival among this presumptive GDV caseload. To unpick the 118 interacting effects of breed and body size, it was hypothesised that, within breeds, 119 animals with bodyweight above their breed mean have an increased odds ratio of 120 presumptive GDV diagnosis compared with those at or below the breed mean.

121

#### 122 Materials and methods

123 The current study was part of the VetCompass<sup>™</sup> Programme of research at the Royal 124 Veterinary College (RVC) (VetCompass 2017) and included all dogs attending Vets Now 125 practices with at least one electronic patient record (EPR) recorded within the

126 VetCompass<sup>™</sup> database from September 1<sup>st</sup>, 2012 to February 28<sup>th</sup>, 2014 (Vets Now 127 2015). These dates were selected because they covered the span of available clinical 128 records at the time of the study. Vets Now clinics use a standard practice management 129 system called Helix and Vets Now team members are required to record presenting signs 130 and encouraged to record diagnoses using the Venom Coding standardised terminology 131 (The VeNom Coding Group 2017). A clinical query was used to extract EPR data from the 132 Helix system that were then uploaded to a secure structured query language database 133 (O'Neill and others 2014b). Data available for the current study included demographic 134 (breed, date of birth, sex, neuter status, insurance status and bodyweight) and clinical 135 (clinical notes, treatment, presenting signs and diagnosis terms with relevant dates) 136 information.

137 A cross-sectional analysis was used to estimate prevalence and evaluate associations 138 between risk factors and GDV diagnosis. Based on the main study hypothesis, sample 139 size calculation estimated that a cross-sectional study would require 5,148 dogs of 140 below-average weight and 5,148 dogs of above-average weight to identify within-breed 141 bodyweight as a risk factor with an odds ratio  $\geq 2.0$  (unexposed:exposed ratio 1:1, 95%) 142 confidence level, 80% power and 0.5% of the unexposed animals with GDV) (Epi Info 7 143 CDC 2015). Ethics approval was granted by the RVC Ethics and Welfare Committee 144 (reference number S25/2014).

145 In this study, the case definition for diagnosis of GDV required that the dog was 146 presented either dead or alive at a participating clinic and that a final diagnosis of GDV 147 (or synonyms covering either torsion or volvulus) was recorded in the EPR based on first 148 opinion emergency-care diagnostic criteria. Because not all cases underwent definitive 149 diagnosis by radiography, surgery, or both, the diagnosis must be considered to be 150 presumptive in some cases. However, dogs that were specified as having gastric dilation 151 without volvulus did not meet the case definition for this study and were excluded. On 152 the other hand, animals that were already dead at the time of first presentation and that 153 met the case definition were included because of the risk that overall prevalence and risk 154 factor values might otherwise be biased by preferentially removing categories of dog 155 with more acute presentations and that tended to die more rapidly.

156 The case-finding process involved an initial EPR screening of all study dogs to identify 157 potential GDV candidate cases by multiple searches of the clinical free-text (*gdv*, *volvul*, 158 torsion, gastric dilat, bloat, twisted stom, gastropexy) and the VeNom term fields 159 (Gastric (stomach) dilation, Gastric (stomach) torsion - chronic and Gastric dilation-160 volvulus syndrome (GDV)). Candidate GDV cases were randomly sorted using the 'RAND' 161 function in Microsoft Excel (Microsoft Office Excel 2007, Microsoft Corp.) to avoid 162 temporal bias during review and the full clinical records were read manually by one of 163 the authors (JC) before deciding on case inclusion and extracting clinical information 164 following a standardised process. For included cases, any previous episodes of GDV were 165 recorded and the date of diagnosis of the current presumptive GDV event were 166 described. Additionally, the presenting status (ambulatory or collapsed), blood lactate 167 concentration at presentation, surgical management (including whether a gastropexy or 168 splenectomy was performed) and clinical outcome (dead on arrival, non-survival, 169 survived) were reported. For the analysis, all dogs not meeting our GDV case definition 170 were included as non-cases for presumptive GDV.

171 Following data checking and cleaning in Excel using internal data validity evaluations 172 (Microsoft Office Excel 2007, Microsoft Corp.), statistical analyses were conducted using 173 Stata Version 11.2 (Stata Corporation). Prevalence values with 95% CI were reported 174 overall and for individual common breeds. The 95% CI estimates were derived from standard errors based on approximation to the normal distribution (Kirkwood and Sterne 175 176 2003). Descriptive statistics characterised purebred status, breed, sex/neuter, age, 177 actual bodyweight and bodyweight relative to breed mean for the GDV case and non-case 178 dogs separately. The breed variable included individual breeds with 10 or more 179 presumptive GDV cases recorded, a group containing the remaining purebred dogs

180 ('other purebred') and a group containing the crossbreds. Actual bodyweight (kg) 181 described the maximum recorded bodyweight for dogs of any age. Six actual-bodyweight 182 categories were generated: (0.0-9.9, 10.0-19.9, 20.0-29.9, 30.0-39.9, ≥ 40.0 and no 183 bodyweight recorded). The mean adult bodyweight was calculated for individual breeds 184 using bodyweight data from dogs aged over 18 months and these values were used to 185 categorise individual dogs aged over 18 months as being either above or at/below their 186 breed mean value ('bodyweight relative to breed mean'). Age (years) described the age 187 at diagnosis for case animals and the age at the mid-point between the first and final EPR 188 for the non-case animals. Age was categorised into six groups (< 3.0, 3.0-5.9, 6.0-8.9, 189 9.0-11.9,  $\geq$  12.0 and not available). Analysis of age and bodyweight as categorical 190 variables was planned during the study design because associations between these 191 variables and clinical outcomes are rarely linear and therefore analysis as continuous 192 variables is statistically inappropriate (Kirkwood and Sterne 2003, O'Neill and others 193 2016, Taylor-Brown and others 2015). The additional categories that described missing 194 data for the bodyweight and age variables were included in the analyses in order to 195 better understand the impact that missing data may have had on the results for these 196 variables. Blood lactate concentrations (mmol/L) were categorised into 5 groups (< 2, 2 197 to < 4, 4 to < 6, 6 to < 8 and  $\geq$  8). These cut-offs were selected to be consistent with 198 previously published results (Green and others 2011). Animals discharged alive from Vets 199 Now clinical care were defined as having survived while those that died or were 200 euthanased whilst still under the care of Vets Now were classified as non-surviving. 201 Categorical variables were compared between groups using the chi-square test or Fishers 202 exact test and quantitative variables were compared using the unpaired t-test or 203 Wilcoxon rank sum test as appropriate (Kirkwood and Sterne 2003). A Bonferroni 204 adjusted P-value of 0.007 was accepted for statistical significance to account for the 205 effects of multiple testing (Aickin and Gensler 1996).

206 Binary logistic regression modelling was used to evaluate univariable risk factor 207 associations with presumptive GDV occurrence. Purebred status (highly correlated with 208 breed) and actual bodyweight (bodyweight was considered to be a defining characteristic 209 of individual breeds) were excluded from multivariable modelling. Other factors with 210 liberal associations in the univariable modelling (P < 0.2) were taken forward for 211 multivariable evaluation. Model development used manual backwards stepwise 212 elimination. Clinic attended was entered as a random effect to quantify the diagnostic 213 variation for presumptive GDV between clinics (Bolker and others 2009). Such variation 214 could result, for example, from regional geographic or breed associations with 215 occurrence across the UK. Pair-wise interaction effects were evaluated for the final model 216 variables (Dohoo and others 2009). The Hosmer-Lemeshow test statistic and the area 217 under the ROC curve were used to evaluate model fit (non-random effect model) (Dohoo 218 and others 2009). Statistical significance was set at P < 0.05 for the logistic regression 219 modelling.

220

#### 221 **Results**

### 222 Descriptive results: demography and clinical outcomes

223 Data completeness overall differed widely between the variables: sex 82.5%, neuter 224 82.5%, insurance 100.0%, date of birth 84.7%, actual bodyweight 33.4% and breed 225 88.9%. The study population comprised 77,088 dogs attending 50 Vets Now clinics 226 across the UK. There were 492 dogs that met our presumptive GDV diagnosis case 227 definition, giving a prevalence of 0.64% (95% CI 0.58% - 0.70%) of the overall canine 228 emergency-care caseload. The prevalence of presumptive GDV varied widely across the 229 breeds. Breeds with the highest prevalence included great Dane (14.0% prevalence, 95% 230 CI 9.8-19.1), akita (9.2%, 95% CI 4.5-16.2), dogue de Bordeaux (7.2%, 95% CI 4.6-231 10.7), Irish setter (7.1%, 95% CI 3.7-12.1) and Weimaraner (7.1%, 95% CI 5.0-9.8). 232 The prevalence among crossbreds was 0.1% (0.1-0.2) (Table 1).

233 Of the presumptive GDV dogs with information available, 431 (96.2 %) were purebred, 234 167 (40.2%) were female, 85 (17.3%) were insured and 168 (40.5%) were neutered 235 (Table 2). The median actual bodyweight was 38.8 kg (IQR: 30.0 – 48.5, range: 3.7 – 236 87.0) and the median age was 8.0 years (IQR: 6.0 – 10.0, range: 0.3 – 20.0) (Figure 237 1). The most common 11 breeds accounted for 264 (53.7%) of the case dogs. Eleven 238 breeds had 10 or more affected individuals recorded: akita, boxer, dogue de Bordeaux, 239 great Dane, basset hound, Dobermann pinscher, Labrador retriever, Rhodesian 240 ridgeback, Irish setter, German shepherd dog and Weimaraner. Of these, the highest 241 breed prevalence was recorded for the great Dane (14.0%, 95% CI 9.8-19.1) and akita 242 (9.2%, 95% CI 4.5-16.2) (Table 2). Data completeness for the presumptive GDV cases 243 was: sex 84.3%, neuter 84.3%, insurance 100.0%, date of birth 90.7%, actual

245

244

bodyweight 37.8% and breed 91.1%.

246 Of the non-case dogs with information available, 55,676 (81.8%) were purebred, 30,046 247 (47.5%) were female, 11,854 (15.5%) were insured and 22,690 (35.9%) were neutered. 248 The median bodyweight of non-case dogs was 14.7 kg (IQR: 7.8-25.7, range: 0.2 – 249 100.0) and the median age was 5.0 years (IQR: 1.7 – 10.0, range: 0.0 – 25.0) (Table 250 2). The most common breed types in the overall study were Labrador retriever (9.8%), 251 Jack Russell terrier (5.5%), Staffordshire bull terrier (4.3%) and cocker spaniel (4.1%). 252 Data completeness for the non-cases was: sex 82.5%, neuter 82.5%, insurance 100.0%, 253 date of birth 84.6%, actual bodyweight 33.4% and breed 88.9%.

Nine of the 492 presumptive GDV cases overall (1.8%) presented as 'dead on arrival'. Of the dogs that were alive at presentation, 243/483 (50.3%) did not survive to discharge, with 215 (88.5%) of these deaths involving euthanasia. Of the 198 dogs that were alive at presentation and did not undergo surgery, 184 (92.9%) did not survive the emergency-care period, with the remaining 14 (7.1%) alive at discharge to their 259 primary-care practices or to their owners' care after which no further information was 260 available. Of these 184 deaths, 178 (96.7%) were due to euthanasia and 6 (3.3%) were 261 unassisted. Of the 285 dogs that underwent surgery, 226 (79.3%) survived and 59 262 (20.7%) did not survive the emergency-care period. Of these 59 deaths, 37 (62.7%) 263 were due to euthanasia and 22 (37.3%) were unassisted (Figure 2). Overall, 169/215 264 (78.6%) presumptive GDV cases that were euthanased during the study period had 265 information specified in the free-text notes describing the main reasons that the owner 266 elected for euthanasia. Of these, 118 (69.8%) euthanasia decisions were taken to avoid 267 further animal suffering and 51 (30.2%) were taken following financial concerns related 268 to ongoing treatment.

269 Of dogs that were alive at presentation, 309 of 388 (79.6 %) presumptive GDV cases 270 with information available were ambulatory. Cases that were ambulatory at presentation 271 were significantly more likely to survive to discharge than non-ambulatory cases 272 (survival: 158 (51.1%) ambulatory versus 17 (21.0%) non-ambulatory, P < 0.001). 273 Ambulatory cases were also significantly more likely to have surgery than non-274 ambulatory cases (surgery: 189 (61.2%) ambulatory versus 27 (33.3%) non-275 ambulatory, P < 0.001). Of 481 presumptive GDV cases alive at presentation with 276 information available, 189 (39.3%) did not receive any diagnostic imaging, 267 (55.5%) 277 received radiology alone, 5 (1.0%) received ultrasonography alone and 20 (4.1%)278 received both radiology and ultrasonography. Of the dogs presented alive but that did 279 not receive surgery, 81 (41.1%) received diagnostic imaging. Dogs that received some 280 form of diagnostic imaging were more likely to have surgery than dogs that did not 281 receive diagnostic imaging (221 (72.3%) versus 73 (38.6%), P < 0.001). Blood lactate 282 concentrations were recorded for 181 (36.8%) cases and showed a median of 3.5 (IQR 283 2.3-6.3, range 0.6-18.3) mmol/L. Blood lactate concentrations at presentation 284 (categorised data) were not associated with the likelihood of surgery (P = 0.227). Dogs

with blood lactate concentrations < 4 mmol/L had an increased probability of survival to discharge both overall (P < 0.001) and among the surgical cases (P < 0.001) (Table 3).

287 Of the presumptive GDV cases that were alive at presentation with information available, 288 182/398 (45.7%) had an orogastric tube inserted and 285/483 (59.0%) were surgically 289 managed. Dogs that underwent surgery were younger than dogs that did not undergo 290 surgery (mean (standard deviation (SD)) 7.0 (3.1) years versus 9.2 (3.3) years, P <291 0.001) but bodyweight was not associated with surgery (surgical cases: mean (SD) 40.3 292 (15.2) kg versus non-surgical cases: 40.9 (15.6) kg, P = 0.845). Insured cases were 293 more likely to receive surgery than uninsured cases (67/85 insured [78.8%] versus 294 218/406 uninsured [53.7%] respectively, P < 0.001). Of surgical cases with information 295 available that survived to discharge, 17.5% (25 of 143 dogs) had a recorded gastrotomy 296 procedure, 97.0% (191 of 197 dogs) underwent a recorded gastropexy procedure and 297 14.9% (20 of 134 dogs) underwent a recorded splenectomy procedure. Splenectomised 298 and non-splenectomised surgical cases did not differ in their proportional survival: 20 299 (71.4%) versus 114 (73.1%), P < 0.001. Information on the duration of general 300 anaesthesia was available for 242 dogs. The median (IQR, range) duration of general 301 anaesthesia was 90.0 (60.0-120.0, 0.0-300.0) minutes (Table 4). The differing counts of 302 cases with procedural information available is explained by differing clarity of the notes 303 recorded in the EPRs about the procedures performed.

304

#### **305** Risk factors for diagnosis of presumptive GDV

306 Univariable logistic regression modelling identified six variables with liberally significant 307 (P < 0.20) association with presumptive GDV diagnosis: purebred status, breed, actual 308 bodyweight, bodyweight relative to breed mean, age and sex/neuter status. Although not 309 included in multivariable modelling as explained above, the univariable results indicated 310 that purebred dogs had 5.6 (95% CI 3.5-9.2, P < 0.001) times the odds of presumptive GDV compared with crossbred dogs and that increasing actual bodyweight was strongly associated with increased odds of the diagnosis, with dogs weighing  $\geq$  40 kg showing 148.7 (95% CI 54.5-406.0, P < 0.001) times the odds compared with dogs weighing < 10.0 kg. No association was identified between insurance status and diagnosis of presumptive GDV (P = 0.272).

316 The final multivariable model comprised three risk factors: breed, age and sex/neuter 317 status but did not identify bodyweight relative to breed mean as a significant risk factor. 318 The final model showed acceptable model-fit (Hosmer-Lemeshow test statistic: P = 319 0.680) and good discrimination (area under the ROC curve: 0.843). The final model was 320 not improved by inclusion of the clinic attended as a random effect (P = 0.095) and no 321 biologically significant interactions were identified. After accounting for the effects of the 322 other variables evaluated in the multivariable model, the individual breeds with the 323 highest odds ratios for presumptive GDV diagnosis compared with crossbred dogs were 324 the great Dane (OR: 114.3, 95% CI 55.1-237.1, P < 0.001), akita (OR: 84.4, 95% CI 325 33.6-211.9, P < 0.001) and dogue de Bordeaux (OR: 82.9, 95% CI 39.0-176.3, P < 326 0.001). The odds of diagnosis increased as dogs aged up to 12 years and then 327 decreased. Compared with dogs aged < 3 years, dogs aged from 6 to < 9 years had 9.5 328 (95% CI 6.1-14.8, P < 0.001) times, and dogs aged from 9 to < 12 years 10.0 (95% CI 329 6.4-15.6, P < 0.001) times, the odds of presumptive GDV diagnosis. Neutered male dogs 330 had 1.3 (95% CI 1.0-1.8, P = 0.041) times the odds of diagnosis compared with entire 331 females (Table 5).

332

#### 333 Discussion

To our knowledge, this is the most comprehensive epidemiologic study of presumptive GDV diagnoses relative to all veterinary emergency cases that has been published to date. This study highlights the relevance of presumptive GDV to the canine emergency 337 caseload, with a prevalence of 0.64% and an overall survival of under 50% of all cases. 338 Of dogs that underwent surgery, approximately 80% survived to discharge from 339 emergency care. Risk factors for diagnosis identified in this study included breed, age > 340 3 years and sex. The great Dane, akita and dogue de Bordeaux breeds had the highest 341 odds of presumptive GDV. Ambulatory status and blood lactate concentrations at initial 342 presentation appeared to be useful survival indicators but their true association with 343 survival may be confounded by their influence on euthanasia decision-making by 344 veterinarians and owners (i.e. these factors may have been used in making the decision 345 for euthanasia).

The diagnostic processes used in the current study were typical of first opinion emergency-care clinicians and may differ from strict diagnostic criteria that might include right lateral radiography of the cranial abdomen, post-mortem or laparotomy to confirm the GDV diagnosis (Glickman and others 1998, Tivers and Brockman 2009, Zacher and others 2010). Because not all of the cases included in the current study met this full diagnostic definition before being assigned a GDV diagnosis, the cases included in the study have been labelled as presumptive GDV cases.

The results of the study did not support the hypothesis that, within breeds, animals with bodyweight above their breed mean have an increased odds ratio of presumptive GDV compared with those at or below the breed mean. Although some association was indicated in the univariable analysis, this effect was no longer significant after accounting for other confounding differences (*e.g.* sex) using multivariable modelling.

358 The current study reported a presumptive GDV prevalence of 0.64% of the overall

359 canine emergency-care caseload, which is in broad agreement with a study of UK

360 pedigree dogs (n = 36,006) that reported a GDV prevalence of 0.7% (Evans 2010). A US

361 study of referral cases (n = 27,254) reported a lower prevalence of GDV of 0.2%

362 (Bellumori and others 2013). Another US study reported that 2.4% of large-breed dogs

and 2.7% of giant-breed dogs had at least one GDV episode annually (Glickman and
others 2000). However, it is difficult to compare prevalence results between studies that
have widely differing populations at risk.

366 First-opinion veterinary clinical records can be a valuable resource for reliable and 367 generalisable health information on the wider animal population (O'Neill and others 368 2014a). Dogs affected by GDV require prompt management to prevent poor clinical and 369 welfare outcomes and are therefore commonly presented as emergency-care cases to 370 out-of-hours clinics (Brockman 2012). Consequently, analysis of combined clinical 371 records from a large number of emergency-care clinics in the UK has potential to provide 372 valuable insights into the epidemiology of GDV that are difficult to gain from other data 373 resources. Emergency-care clinical records benefit from reduced recall and 374 misclassification bias because of contemporaneous recording of health information by 375 qualified professionals at the point of clinical care when and can be geographically 376 representative when the contributing clinics span the entirety of the UK (Bateson 2010, 377 McGreevy 2007, O'Neill and others 2014a). Electronic patient records also enable 378 researchers to identify and include all diagnosed cases, regardless of their level of clinical 379 work-up and management, and to explore the free-text clinical notes to answer 380 contextual questions concerning aspects of welfare, presentation, diagnosis and 381 management.

382 Purebred status was not evaluated in the multivariable modelling because it was highly 383 correlated with breed. However, based on the univariable analysis in the current study, 384 purebred dogs had over five times the odds of presumptive GDV compared with 385 crossbreds. This finding is consistent with US studies emanating from referral clinics that 386 reported odds ratios for purebred predisposition of 1.8 (Bellumori and others 2013), 2.5 387 (Glickman and others 1994) and 4.8 (de Battisti and others 2012). A purebred 388 predisposition for GDV may indicate a genuine inheritable predisposition but, alternatively, could reflect differing body-size distributions between purebred and 389

390 crossbred dogs. Although the median bodyweight of purebred and crossbred dogs is quite 391 similar, purebred dogs tend to have a greater proportion of dogs with extreme large and 392 small body size (O'Neill and others 2013) and GDV has been reported to be strongly 393 associated with large or giant breed body types (Evans 2010).

394 After accounting for the other factors analysed, the breeds with the highest risk of 395 presumptive GDV compared with crossbred dogs in the current study included great 396 Dane (OR 114.3), akita (OR 84.4), dogue de Bordeaux (OR 82.9), Irish setter (OR 67.4) 397 and Weimaraner (OR 50.8). A previous UK study additionally identified high GDV 398 prevalence in the grand bleu de Gascogne, bloodhound and otterhound (Evans 2010) 399 while US studies also reported high odds for Saint Bernard (Bellumori and others 2013, 400 Brockman 1995, Glickman and others 1994). Many of these predisposed breeds are 401 considered as 'deep-chested', with this description written into their breed standards 402 (The Kennel Club 2014). This suggests an association between this conformation and 403 GDV, thereby offering opportunities to reduce GDV hazard via prophylactic gastropexy 404 (Ward and others 2003) or breed selection against extreme deep-chested conformation 405 (Bell 2014).

406 Large or giant breeds feature heavily among the predisposed breeds for GDV (Bell 407 2014). The current study did not include actual bodyweight in the multivariable analysis 408 because the breed variable already somewhat accounted for bodyweight. However, the 409 univariable results did support an association between increasing bodyweight and 410 increased odds of presumptive GDV, with dogs weighing 30.0-39.9 kg having 38.5 times 411 the odds compared with dogs weighting < 10.0 kg. Similar associations between 412 increasing bodyweight and risk of GDV have been reported in US studies of referred GDV 413 cases (Glickman 2000, Glickman and others 1994). However, it is worth noting for the 414 current study, that bodyweight data were available for less than 40% of the study dogs 415 and that these results may be confounded by breed and sex effects. Consequently, 416 exploration in future studies that control for these factors is warranted.

417 Older dogs were identified at higher risk of presumptive GDV in the current study, with 418 dogs aged between 9 and 12 years showing 10 times the odds of dogs aged under 3 419 years. A previous study reported significantly increasing odds of GDV in Irish setters as 420 dogs aged (Elwood 1998). The risk of GDV was reported to increase by approximately 421 20% with each year of age in large and giant breeds in the US (Glickman and others 422 2000) and the odds of GDV rose by 1.9 times for each year as great Danes in the 423 Netherlands aged from 1 to 10 years old (Theyse and others 1998). Progressive 424 stretching of the hepatogastric ligament with increasing age has been suggested as a 425 pathogenic pathway for this increase GDV risk with aging (Hall and others 1995).

The current study did identify some evidence of a male predisposition, consistent with the results of a small US study (Glickman and others 1997). However, other previous studies have failed to identify a sex predisposition (Glickman and others 1994, Glickman and others 2000). Sex associations may be confounded by bodyweight, neutering and other factors that complicate the interpretation of true sex effects.

431 The current study identified that 50.3% of the emergency-care presumptive GDV 432 caseload that presented alive did not survive to discharge, with 88.5% of these deaths 433 involving euthanasia. This figure is very similar to the 86% of dog deaths that involve 434 euthanasia under general primary veterinary care in the UK (O'Neill and others 2013). 435 However, 79.3% of dogs that underwent surgery survived to discharge from emergency 436 care. Although the relatively high survival rate for surgical cases may reflect effective 437 case selection for surgical intervention, it is possible that many animals that did not 438 receive surgery may also have survived if this option had been elected. There was a 439 trend for breeds with a higher prevalence of presumptive GDV to be more likely to 440 undergo surgery and therefore to have better overall survival (Table 2). This may reflect 441 the probability that the owners of predisposed breeds are more aware of the disease in 442 general and therefore better prepared to make the decision for surgical treatment during 443 the emergency-care consultation at a time of extreme emotional distress.

444 It is difficult to compare the proportions of dogs undergoing surgery and their survival 445 rates against previous publications because of differing study case definitions, case 446 populations, methodologies and data sources between the published works. A study of 447 working farm dogs in New Zealand reported 65% survival overall but the study included 448 dogs affected with gastric dilation either with or without volvulus and included only dogs 449 that received either radiography, surgery or post-mortem affected with GDV (Hendriks 450 and others 2011). US studies of surgically-managed cases in referral clinic settings have 451 reported survival rates of 90% (Mackenzie and others 2010), 88% (Green and others 452 2011), 85% (Brockman 1995), 84% (Beck and others 2006) and 82% (Brourman and 453 others 1996). The lower survival reported in the current study may reflect inclusion of 454 more acute and severe cases in the first-opinion emergency-care setting and differing 455 case management compared with the referral situation and suggests that the results of 456 the current study may be more applicable to the wider canine population. In addition, 457 many of the deaths in the current study involved euthanasia, and over 30% of these 458 deaths were related to financial concerns, which may have biased the survival rates 459 downwards.

460 Taken at face value, the current findings might suggest that assessing ambulation and 461 blood lactate concentrations at initial clinical presentation may be useful survival 462 indicators for presumptive GDV. In the current study, 51.1% of dogs that were 463 ambulatory at presentation survived compared with 21.0% survival in alive but non-464 ambulatory dogs. Lower initial lactate concentrations were also associated with higher 465 survival rates in the current study (Table 3). These findings are consistent with other 466 reports that suggest the prognostic value of blood lactate estimation (Beer and others 467 2012). Among surgical GDV cases in the US, the median initial plasma lactate 468 concentration in dogs that survived was 3.4 mmol/L compared with 6.80 mmol/L for 469 non-survivors (Green and others 2011). A US study of a general emergency caseload 470 reported that 4.9% of patients with blood lactate concentrations up to 2.5 mmol/L died 471 compared with 28.4% of patients with lactate values at or above 4.0 mmol/L (Shapiro 472 and others 2005). Among systemically ill dogs in Canada, dogs with lactate 473 concentrations above 2.3 mmol/L had 16 times more risk of dying than dogs with lactate 474 concentrations at or below 2.3 mmol/L (Stevenson and others 2007). However, it is 475 likely that associations between ambulation and blood lactate concentrations with 476 subsequent survival are highly complex and so it may be unsafe to draw definitive 477 conclusions from the current study results because of possible reverse-causality effects. 478 Non-ambulatory dogs or those with high blood lactate concentrations may have been 479 considered a priori to have a poorer prognosis by the owner or the attending veterinary 480 surgeon and thus have been more likely to be euthanased. . It should also be noted that 481 data availability for ambulatory status (79.2%) and plasma lactate concentration 482 (36.8%) were not complete and therefore some biases relating to missing data may 483 have affected the results. Survival analyses where clinicians were blinded to the values 484 of potential prognostic indicators are needed to more definitively clarify the predictability 485 of ambulation and blood lactate concentration as useful clinical indicators for survival, 486 especially when they may be applied to individual animals.

487 Gastropexy, where the stomach is securely adhered to the abdominal wall, is clinically 488 indicated in all dogs that undergo surgical correction of GDV to prevent recurrence (Allen 489 and Paul 2014). GDV cases that receive gastropexy may have less than 5% recurrence 490 and a median survival time of 547 days whereas those that do not receive gastropexy 491 may have up to 80% recurrence and a median survival time of 188 days (Allen and Paul 492 2014, Glickman and others 1998). In this current study 3% of the surgical cases did not 493 have a gastropexy procedure recorded in their clinical notes and so it appears that 494 gastropexy is widely accepted as a clinical standard in the UK.

495

The current study had some limitations. Because of the urgent nature of emergency-carepresentations and because these data were not recorded primarily for research

498 purposes, many of the available data fields in the clinical records were incomplete or 499 missing and it cannot be assumed that these data were missing at random. 500 Consequently, 'missingness' may have introduced some bias into the final results and 501 categories that described missing data were included in the analyses to try to quantify 502 such biases. The EPRs of emergency-care patients described mainly the current 503 presentation and often provided very limited information on prior history. Dogs were 504 generally lost to longer-term follow-up after the immediate emergency-care treatment 505 period. Serial blood lactate concentrations values were not available for this study. This 506 study reports the prevalence of presumptive GDV within the first-opinion emergency-507 care population and the management and outcomes of these cases may not be fully 508 representative of the wider dog population (Bartlett and others 2010). However, these 509 differences should have less impact on the generalisability of results from the risk factors 510 which are more dependent on basic physiology and therefore should be more constant 511 across all dogs in the UK (Elwood 2007).

512

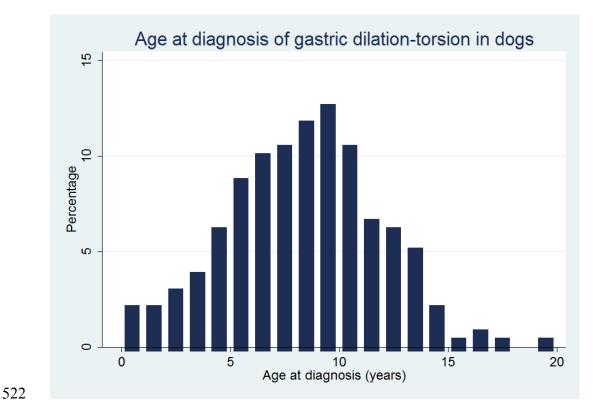
### 513 Conclusion

These results provide a baseline against which future studies of GDV in the UK primarycare population can be judged and provide information that may help to inform both veterinary surgeons and dog owners about GDV risk and prognosis.

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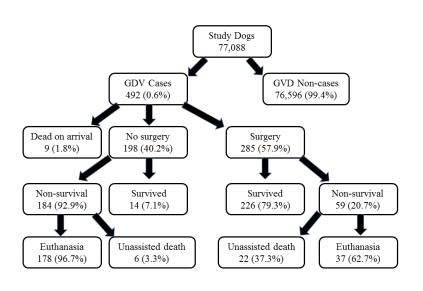
# 521 Figures



523 Figure 1: Age at diagnosis of presumed gastric dilation-volvulus in 466 dogs attending

524 first opinion emergency-care veterinary practices in the UK

525



- 527 Figure 2: Flowchart showing some outcomes for presumed gastric dilation-volvulus in
- 528 dogs attending first opinion emergency-care veterinary practices in the UK

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530

# 531 **Tables**

Breed type	Total no. dogs	No. cases	Prev alen ce (%)	95% CI	No. (%) cases receivin g surgery	No. (%) surgical cases died or were euthanased	No. (%) cases died or were euthanased overall
Crossbreed	10,713	12	0.1	0.1-0.2	4 (33.3)	1 (25.0)	9 (75.0)
Great Dane	236	33	14.0	9.8- 19.1	24 (72.7)	3 (12.5)	12 (36.4)
Akita	109	10	9.2	4.5- 16.2	4 (40.0)	0 (0.0)	6 (60.0)
Dogue de Bordeaux	318	23	7.2	4.6- 10.7	9 (39.1)	2 (22.2)	16 (69.6)
Irish Setter	169	12	7.1	3.7- 12.1	10 (83.3)	1 (10.0)	3 (25.0)
Weimaraner	480	34	7.1	5.0-9.8	21 (61.8)	6 (28.6)	17 (50.0)
Rhodesian Ridgeback	194	10	5.2	2.5-9.3	8 (80.0)	3 (37.5)	5 (50.0)
Basset Hound	241	11	4.6	2.3-8.0	7 (63.6)	1 (14.3)	5 (45.5)
German Shepherd Dog	1,910	74	3.9	3.1-4.8	41 (55.4)	10 (24.4)	42 (56.8)
Dobermann Pinscher	417	12	2.9	1.5-5.0	5 (45.5)	1 (20.0)	8 (66.7)
Boxer	1,308	23	1.8	1.1-2.6	8 (34.8)	3 (37.5)	17 (73.9)
Labrador Retriever	6,707	22	0.3	0.2-0.5	10 (45.5)	2 (20.0)	13 (59.1)
Other purebreds	54,264	216	0.4	0.3-0.5	134 (62.0)	26 (19.4)	100 (46.3)

532 Table 1: Prevalence with 95% confidence intervals (95% CI) for dog breed types

533 commonly diagnosed with presumed gastric dilation-volvulus at first opinion emergency-

534 care veterinary practices in the UK

535

Variable	Category	Case No. (%)	Non-case No. (%)	Odds ratio	95% CI*	P-Value
Purebred status	Crossbred	17 (3.4)	12,377 (18.2)	Base		
	Purebred	431 (96.2)	55,676 (81.8)	5.6	3.5-9.2	< 0.001
Breed	Crossbreed	12 (2.4)	10,701 (14.0)	Base		
	Great Dane	33 (6.7)	203 (0.3)	145.0	73.8- 284.7	< 0.001
	Akita	10 (2.0)	99 (0.1)	90.1	38.0- 213.3	< 0.001
	Dogue de Bordeaux	23 (4.7)	295 (0.4)	69.5	34.3- 141.1	< 0.001
	Irish Setter	12 (2.4)	157 (0.2)	68.2	30.2- 154.1	< 0.001
	Weimaraner	34 (6.9)	446 (0.6)	68.0	35.0- 132.2	< 0.001
	Rhodesian Ridgeback	10 (2.03)	184 (0.2)	48.5	20.7- 113.6	< 0.001
	Basset Hound	11 (2.2)	230 (0.3)	42.6	18.6- 97.7	< 0.001
	German Shepherd Dog	74 (15.0)	1,836 (2.4)	35.9	19.5- 66.3	< 0.001
	Dobermann Pinscher	12 (2.4)	405 (0.5)	26.4	11.8- 59.2	< 0.001
	Boxer	23 (4.7)	1,285 (1.7)	16.0	7.9- 32.2	< 0.001
	Other breed types	216 (43.9)	54,048 (70.6)	3.6	2.0-6.4	< 0.001
	Labrador Retriever	22 (4.47)	6,707 (8.8)	2.9	1.4-5.9	< 0.001
Actual bodyweight (kg)	< 10.0	4 (0.8)	9,021 (11.8)	Base		
	10.0-19.9	5 (1.0)	6,856 (9.0)	1.7	0.4-6.1	0.458
	20.0-20.9	36 (7.3)	5,169 (6.8)	15.7	5.6- 44.2	< 0.001
	30.0-30.9	55 (11.2)	3,220 (4.2)	38.5	13.9- 106.4	< 0.001
	≥ 40.0	86 (17.5)	1,304 (1.7)	148.8	54.5- 406.0	< 0.001
	No recorded bodyweight	306 (62.2)	51,026 (66.6)	13.5	5.0- 36.3	< 0.001

Adult (>18 months)		70 (14 2)	0.000 (11.7)	D		
bodyweight relative to breed mean	Lower/Equal	70 (14.2)	8,989 (11.7)	Base		
	Higher	85 (17.3)	7,644 (10.0)	1.4	1.0-2.0	0.028
_	Not available	337 (68.5)	59,963 (78.3)	0.7	0.6-0.9	0.013
Age category (years)	< 3.0	31 (6.3)	22,030 (28.8)	Base		
	3.0 - 5.9	85 (17.3)	12,593 (16.4)	4.8	3.2-7.2	< 0.001
	6.0 - 8.9	147 (29.9)	10,558 (13.8)	9.9	6.7- 14.6	< 0.001
	9.0 - 11.9	137 (27.9)	8,898 (11.6)	10.9	7.4- 16.2	< 0.001
	≥ 12.0	66 (13.4)	10,741 (14.0)	4.4	2.8-6.7	< 0.001
	No age data available	26 (5.3)	11,776 (15.4)	1.6	0.9-2.6	0.091
Sex/neuter	Female entire	100 (24.1)	18,609 (29.4)	Base		
	Female neutered	67 (16.1)	11,437 (18.1)	1.1	0.8-1.5	0.586
	Male entire	147 (35.4)	21,917 (34.7)	1.2	1.0-1.6	0.088
	Male neutered	101 (24.3)	11,253 (17.8)	1.7	1.3-2.2	< 0.001
Insurance	Non-insured	407 (82.7)	64,742 (84.5)	Base		
	Insured	85 (17.3)	11,854 (15.5)	1.1	0.9-1.4	0.272

Table 2: Descriptive and univariable logistic regression results for risk factors associated 

with a diagnosis of presumed gastric dilation-volvulus in dogs attending first opinion

emergency-care veterinary practices in the UK.

Blood lactate concentration at presentation (mmol/L)	Overall: No. (%)	Not received surgery: No. (%)	Received surgery: No. (%)	Overall - survived to discharge: No. (%)	Surgical cases - survived to discharge: No. (%)
< 2	32 (17.7)	2 (6.2)	30 (93.8)	30 (93.8)	29 (96.7)
2  to < 4	67 (37.0)	7 (10.4)	60 (89.6)	56 (83.6)	52 (86.7)
4  to < 6	29 (16.0)	4 (13.8)	25 (86.2)	16 (55.2)	16 (64.0)
6 to < 8	22 (12.2)	5 (22.7)	17 (77.3)	13 (59.1)	12 (70.6)
$\geq 8$	31 (17.1)	7 (22.6)	24 (77.4)	10 (32.3)	10 (41.7)

	Overall	181 (100.0)	25 (13.8)	156 (86.2)	125 (69.1)	119 (76.3)			
544	Table 3: Initial blood lactate concentration (mmol/L) and associations with surgery and								
545	survival to discharge in dogs alive at presentation that were diagnosed with presumptive								
546	gastric dilation-volvulus attending first opinion emergency-care veterinary practices in								
547	the UK.								
548									
549									

Duration of general anaesthesia (minutes)	No. (%) not receiving surgery	No. (%) receiving surgery	No. (%) died or were euthanased	No. (%) survived to discharge
< 50	8 (42.1)	11 (57.9)	13 (68.4)	6 (31.6)
50 - <100	3 (2.9)	100 (97.1)	25 (24.3)	78 (75.7)
100 - <150	0 (0.0)	77 (100.0)	15 (19.5)	62 (80.5)
≥150	0 (0.0)	43 (100.0)	8 (18.6)	35 (81.4)
Overall	11 (4.5)	231 (95.5)	61 (25.2)	181 (74.8)

552 Table 4: Duration of general anaesthesia (minutes) and associations with surgery and

553 survival to discharge in dogs diagnosed with presumptive gastric dilation-volvulus

attending first opinion emergency-care veterinary practices in the UK.

Variable	Category	Odds ratio	95% CI*	P- Value
Breed	Crossbreed	Base		
	Great Dane	114.3	55.1-237.1	< 0.001
	Akita	84.4	33.6-211.9	< 0.001
	Dogue de Bordeaux	82.9	39.0-176.3	< 0.001
	Irish Setter	67.4	28.9-157.2	< 0.001
	Weimaraner	50.8	25.2-102.7	< 0.001
	Basset Hound	39.5	16.8-92.7	< 0.001
	Rhodesian Ridgeback	31.0	12.2-78.6	< 0.001
	German Shepherd Dog	27.5	14.4-52.4	< 0.001

	Dobermann Pinscher	20.8	8.7-49.5	< 0.001
	Boxer	13.4	6.5-27.9	< 0.001
	Labrador Retriever	2.4	1.2-5.1	0.019
	Other purebreds	3.2	1.7-5.8	< 0.001
Age category (years)	< 3.0	Base		
	3.0 - 5.9	5.2	3.3-8.3	< 0.001
	6.0 - 8.9	9.5	6.1-14.8	< 0.001
	9.0 - 11.9	10.0	6.4-15.6	< 0.001
	$\geq$ 12.0	5.6	3.4-9.2	< 0.001
	No age data available	1.6	0.9-3.0	0.111
Sex/neuter	Female entire	Base		
	Female neutered	0.8	0.6-1.1	0.271
	Male entire	1.2	0.9-1.6	0.168
	Male neutered	1.3	1.0-1.8	0.041

558 Table 5: Final multivariable logistic regression model for risk factors associated with a

559 diagnosis of presumptive gastric dilation-volvulus in dogs attending first opinion

560 emergency-care veterinary practices in the UK.

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