This is the peer-reviewed, manuscript version of the following article:


The final version is available online via [http://dx.doi.org/10.1016/j.jvc.2015.09.008](http://dx.doi.org/10.1016/j.jvc.2015.09.008).

© 2015. This manuscript version is made available under the CC-BY-NC-ND 4.0 license [http://creativecommons.org/licenses/by-nc-nd/4.0/](http://creativecommons.org/licenses/by-nc-nd/4.0/).

The full details of the published version of the article are as follows:

**TITLE:** Risk factors associated with sudden death vs. congestive heart failure or arterial thromboembolism in cats with hypertrophic cardiomyopathy

**AUTHORS:** Payne, J. R., Borgeat, K., Brodbelt, D. C., Connolly, D. J. and Luis Fuentes, V.

**JOURNAL TITLE:** Journal of Veterinary Cardiology

**VOLUME/EDITION:** 17 Supplement 1

**PUBLISHER:** Elsevier

**PUBLICATION DATE:** December 2015

**DOI:** 10.1016/j.jvc.2015.09.008
Factors associated with sudden death versus congestive heart failure or arterial thromboembolism in cats with hypertrophic cardiomyopathy

Jessie Rose Payne BVetMed PhD*
Kieran Borgeat BSc (Hons) BVSc MVetMed†
David Charles Brodbelt MA VetMB PhD*
David J Connolly BSc BVetMed PhD*
Virginia Luis Fuentes MA VetMB PhD*

*Clinical Science and Services, Royal Veterinary College, Hawkshead Lane, North Mymms, Hatfield, Hertfordshire, AL9 7TA, United Kingdom
†Highcroft Veterinary Referrals, 615 Wells Road, Whitchurch, Bristol, BS14 9BE, United Kingdom

Corresponding author: Virginia Luis Fuentes. Clinical Science and Services, Royal Veterinary College, Hawkshead Lane, North Mymms, Hatfield, Hertfordshire, AL9 7TA, United Kingdom.
vluisfuentes@rvc.ac.uk

Keywords
- HCM
- sudden death

- CHF

- ATE

Abbreviations:

ATE – Arterial Thromboembolism

CHF – Congestive Heart Failure

E/A – ratio of mitral inflow peak early filling (E) to late filling (A) velocities

FS% – Left ventricular fractional shortening

HCM – Hypertrophic cardiomyopathy

LA – Left atrial

LA:Ao – short axis ratio of diastolic left atrial diameter to aortic root diameter

LAD – the diameter of the left atrium measured parallel with the mitral annulus in the last frame before mitral valve opening

LA-EF% – Left atrial emptying fraction

LA-FS% – Left atrial fractional shortening

LV – Left ventricle

LVWd – end-diastolic left ventricular septal or free wall thickness

MR – Mitral regurgitation

QMHA – Queen Mother Hospital for Animals

SAM – Systolic Anterior Motion of the mitral valve

SEC – Spontaneous echo-contrast

Conflict of Interest
The authors have no conflict of interest to declare with regard to this study.

This study was presented at the ECVIM-CA, Seville, September 2011

Acknowledgments: The authors gratefully acknowledge the Everts Luff feline endowment and IDEXX Laboratories for financial support for Dr Payne’s PhD studies. The authors would like to thank the owners, referring vets and QMHA clinicians involved in the care of these cats.
Abstract

Objectives

To evaluate differences in risk factors associated with death due to congestive heart failure (CHF), arterial thromboembolism (ATE) or sudden death in cats with hypertrophic cardiomyopathy (HCM).

Animals

255 cats with HCM referred to a veterinary teaching hospital.

Methods

Retrospective study. Cats with HCM were identified that had either died within 2 years of diagnosis or were known to be alive 2 years after initial examination. Signalment, physical exam, electrocardiographic and echocardiographic data were analyzed separately for association with death due to congestive heart failure (CHF) vs. aortic thromboembolism (ATE) vs. sudden death. Results:

Results

Within 2 years of follow-up, 23/255 (9.0%) cats had died with ATE, 44/255 (17.3%) cats had died with CHF and 12/255 (4.7%) cats had experienced a sudden death, with 141/255 (55.3%) cats still alive at the end of 2 years. Presence of CHF at presentation and reduced left ventricular fractional shortening (FS%) were independently associated with a CHF death within 2 years of diagnosis. Presence of ATE and reduced left atrial fractional shortening (LA-FS%) were independently associated with dying with ATE within 2 years. No multivariable models were generated for risks of dying a sudden death owing to the low event rate, but syncope at presentation and arrhythmias on auscultation were associated with sudden death on univariable analysis.
Conclusions

Asymptomatic cats have a reduced risk of all three types of death. Reduced FS% and a history of CHF independently predict CHF death, and reduced LAFS% and history of ATE independently predict ATE death. Sudden death is less commonly reported but is associated with syncope.
Hypertrophic cardiomyopathy (HCM) is defined as a hypertrophied, non-dilated left ventricle (LV) in the absence of other systemic or cardiac disease capable of producing a similar degree of hypertrophy.\(^1\) It is the most common human familial heart disease\(^2\)-\(^3\) and the most commonly diagnosed feline myocardial disease.\(^4\) HCM is a heterogeneous disease, both in terms of presentation and outcome. Affected cats may be asymptomatic, or may show signs associated with congestive heart failure (CHF), arterial thromboembolism (ATE), syncope or sudden death. Some cats will die of their cardiac disease but others may have long survival times and die of non-cardiac causes.\(^5\)-\(^10\) Previous studies have reported clinical features at presentation in cats that have subsequently died with cardiomyopathy and CHF, ATE or sudden death. These features include presence of clinical signs, left atrial (LA) enlargement, the absence of systolic anterior motion of the mitral valve (SAM), increasing age and breed.\(^6\)-\(^8,11\)-\(^13\) We have previously reported reduced LA fractional shortening, left ventricular (LV) fractional shortening and extreme LV hypertrophy as independent predictors of cardiac mortality in 282 cats with HCM.

Studies evaluating the risk factors for specific types of cardiac death (such as sudden death vs. CHF death or ATE death) are lacking in cats. An association between LA enlargement and ATE was noted in one study, and a breed predisposition noted for sudden death in another study, but in general our understanding is poor of the risk factors predicting death due to CHF instead of ATE or sudden death.

We hypothesized that the risk factors in cats with HCM would be different for death due to CHF compared with thromboembolic death or sudden cardiac death. The aim of this study was to conduct a sub-analysis of a previously reported population of cats with HCM to investigate risk factors for different mortality end-points (CHF death, ATE death and sudden cardiac death).
Animals, Materials and Methods

These data were originally collected to investigate overall survival for this population and as such, the methods used have been described previously. In brief, cases of feline HCM diagnosed by a board-certified cardiologist or a cardiology resident supervised by a board-certified cardiologist at the Royal Veterinary College, Queen Mother Hospital for Animals (QMHA) on the basis of 2D and/or M-mode echocardiography\(^a\) between June 2004 and August 2009 were included in this study. Cats were excluded from the study if they had a concurrent diagnosis of hyperthyroidism or hypertension, defined as either systolic blood pressure $\geq 180$ mmHg; systolic blood pressure $\geq 160$ mmHg with retinal changes suggestive of hypertension; medically controlled hypertension; or where renal disease was present and blood pressure had not been determined. Cats were also excluded if they were still alive with less than 2 years of follow-up.

The medical records for each cat were reviewed for signalment data, physical exam findings, systolic arterial blood pressure assessment, serum total thyroxine concentrations, ECG findings, clinical status and echocardiographic findings. Clinical status at presentation was recorded and grouped into asymptomatic, syncope, exertional dyspnea, CHF and ATE. Congestive heart failure was defined as present based on previous radiographic evidence of pulmonary edema, ultrasonographic evidence of pleural or pericardial effusion, or severe tachypnea responsive to furosemide. Cats presented to our center without congestive signs were designated as having CHF if there was prior evidence of CHF at either our hospital or the referring practice. Aortic thromboembolism was defined as either sudden onset painful pulseless limb paresis; central nervous system signs associated with magnetic resonance imaging findings of a well-demarcated brain lesion, hyperintense on a T2-weighted image; or a history of

\(^a\) Vivid 7, GE Medical Systems Ltd, Hatfield, Hertfordshire, UK
acute onset abdominal pain, high creatinine kinase levels, and echocardiographic evidence of SEC in the LA for mesenteric thromboembolism. For the original study, all echocardiographic examinations were reviewed and remeasured by one board-certified cardiologist (VLF) or a cardiology resident (KB). Details of echocardiographic measurements are listed in Table 1. Left ventricular hypertrophy was defined as LV septal or free wall end-diastolic thickness >= 6 mm in 2D or M-mode measurements.

M-mode images were used to obtain LV fractional shortening (FS%) values whilst 2D images were used to record maximal LV end-diastolic wall thickness from either the septum or free wall (LVWd). The presence of any left ventricular regional wall hypokinesis, subjectively assessed based on 2D images, was noted. Presence of systolic anterior motion of the mitral valve (SAM) or mitral regurgitation (MR) was recorded.

Mortality was determined by reviewing QMHA medical records and contacting referring veterinarians. Date of death, whether the cat died naturally or due to euthanasia, and whether death was related to cardiac disease (sudden death, CHF, ATE) or non-cardiac causes was recorded. Where insufficient evidence was available, owners were contacted and asked to complete a questionnaire. Sudden death was defined as being found dead without obvious cause at home or as a witnessed event where the cat had been apparently well in the preceding 24 hours, and was assumed to be cardiac. CHF death was defined as dying with dyspnea, crackles, cyanosis, fluid pouring out of mouth and/or euthanasia due to becoming refractory to CHF medication. ATE death was defined as death or euthanasia following a new episode of ATE or worsening of a current ATE. If a cat was alive 2 years after diagnosis it was classified as alive, irrespective of whether or not further long term follow up was available.
Statistical analysis was performed using commercially available software\(^b\) and values are reported as mean ± standard deviation (normally distributed data) and median (interquartile range (IQR)) for non-normally distributed data. Cats reaching a specific end point (sudden cardiac death, CHF death or ATE death), were compared to a composite of cats that had either died of the other two end points or a non-cardiac death within 2 years of diagnosis or were still alive 2 years after diagnosis (irrespective of their ultimate outcome). Univariable analyses on continuous data were performed using t-tests or Mann Whitney U test as appropriate and for proportions using Chi-squared or Fisher’s exact as appropriate. Multivariable analysis was performed using binary logistic regression for the outcomes that had at least 20 cats. Overall model fit was assessed using the Hosmer-Lemeshow test, positive and negative predictive values from classification tables and assessment of DFBeta and deviance residuals. A value of p<0.05 was considered statistically significant.

\(^b\) GraphPad Prism 5, GraphPad Software, 2007 and PASW Statistics 20, 2011
Results

Between June 2004 and August 2009, 282 cats were diagnosed with HCM. Of those, 255 had either died within 2 years of diagnosis or were known to be still alive 2 years after diagnosis. The other 27 cats were known to be still alive at the last point of contact but had less than 2 years of follow up and were therefore excluded. Most cats were male (75.3%) and neutered (94.5%), and 81.2% were nonpedigree. Mean body weight was 4.69 ± 1.09 kg. The study population were not different in terms of sex (p=1.000), neutering status (p=0.375), breed (p=0.442) or weight (p=0.477) to the 27 excluded cats. The study population was significantly older (7.0 [3.1-10.0] years) than the excluded 27 cats (3.6 [2.2-7.5] years, p=0.016). On presentation, the majority (139, 54.5%) of cats were asymptomatic, but 84 (32.9%) had CHF, 16 (6.3%) had ATE, 13 (5.1%) were presented for syncope and 3 (1.2%) had exertional dyspnea (Table 2). Of the 16 cats presenting with ATE, four also had CHF but were classed as ATE. Four of the cats presenting with ATE also had a history of ATE. A minority of cats designated as having CHF (10/84 with CHF and 2/4 with ATE and CHF) had a history of CHF but were not showing overt signs of congestive failure at presentation on the visit used in this study. At the initial visit, antithrombotic treatment was started in 44 (17.3%) cats. Antithrombotic treatment included aspirin (n=40), clopidogrel (n=2), or both aspirin and clopidogrel (n=2). Of those started on antithrombotic treatment, 15 (34.1%) subsequently presented with ATE, and 25 (56.8%) presented with CHF. Within 2 years of diagnosis, 79/255 cats had died of their heart disease (31%). Death was associated with CHF in 44 of the 255 (17.3%) cats, ATE in 23 (9.0%) cats, and sudden death in 12 (4.7%) cats. Sixteen (6.3%) cats did not survive to discharge; eight (50%) cats were euthanized because of noncardiac disease, five (31.3%) were euthanized because of CHF, two (12.5%) died suddenly and one (6.3%) was euthanized because of ATE. Of the eight cats euthanized in hospital because of noncardiac disease, seven (87.5%) were classified as asymptomatic and one cat (12.5%) developed CHF during investigations for neoplastic disease and was
euthanized because of the neoplastic disease but was classified as having CHF. One of the cats that died suddenly was witnessed to develop ventricular fibrillation and resuscitation was unsuccessful. Of the 44 deaths due to CHF, 37 (84.1%) were euthanized and seven (15.9%) died spontaneously. Of the 23 deaths due to ATE, nine (39.1%) also had CHF at the time of death, though they were classed as ATE deaths. Seventeen (73.9%) of the cats that died because of ATE were euthanized and six (26.1%) died spontaneously. Other than the two sudden deaths that occurred during the original visit, all sudden deaths occurred at home. For all the cats that died at home, the primary veterinarian was unaware of their death and information was gained from the owners. Death was due to noncardiac disease in 35 (13.7%) cats, and 141 (55.3%) cats were still alive 2 years after diagnosis.

**CHF death**

Cats that died because of CHF were compared to a composite of cats that died suddenly, died because of ATE, died a noncardiac death or were still alive 2 years after diagnosis. On univariable analysis, the hazard of a CHF death (Fig. 1) increased with increasing age, presence of an arrhythmia, presenting with CHF, decreasing left ventricular fractional shortening (FS%), increasing LA size (LA:Ao and LAD), decreasing left atrial emptying fraction (LA-EF%) and left atrial fractional shortening (LA-FS%), increasing maximal 2D LVWd, presence of regional wall hypokinesis, presence of SEC/thrombus, and having a restrictive filling pattern. The hazard of a CHF death was decreased in cats with a murmur, cats that were asymptomatic at presentation, and cats with SAM.

**ATE death**

Cats that died because of ATE were compared to a composite of cats that died suddenly, died because of CHF, died a noncardiac death or were still alive 2 years after diagnosis. The hazard of an ATE death (Fig. 2) was increased in male cats, cats with a gallop sound, presenting with an ATE, increasing LA size
(LA:Ao and LAD), decreasing LA-EF% and LA-FS %, increasing maximal 2D LVWd, presence of SEC/thrombus and cats with a restrictive filling pattern. The hazard of an ATE death was decreased in cats that were asymptomatic at presentation.

**Sudden death**

Cats that died suddenly were compared to a composite of cats that died because of an ATE, died because of CHF, died a noncardiac death or were still alive 2 years after diagnosis. The hazard of sudden death (Fig. 3) was increased in cats with an arrhythmia, those with syncope, decreasing FS %, increasing LA size (LA:Ao and LAD), decreasing LA-EF%, presence of regional wall hypokinesis and presence of SEC/thrombus. The hazard of sudden death was decreased in cats that were asymptomatic at presentation. Separate multivariable models were generated for predicting either a CHF death (Table 3) or an ATE death (Table 4) within 2 years of diagnosis. Presence of CHF at diagnosis and decreasing LV FS% predicted a CHF death while presence of an ATE at presentation and decreasing LA-FS% predicted an ATE death. Owing to low numbers of sudden deaths, a multivariable model could not be generated. Model fit for the CHF model (Hosmer-Lemeshow \( \chi^2 = 0.329 \), positive predictive value \( \% = 82.4\% \), negative predictive value \( \% = 90.1\% \), correctly classified \( \% = 89.5\% \) ) and for the ATE model (Hosmer-Lemeshow \( \chi^2 = 0.895 \), positive predictive value \( \% = 88.9\% \), negative predictive value \( \% = 93.7\% \), correctly classified \( \% = 93.5\% \) ) were both considered good. Analysis of the residuals for each model showed good model fit.
Numerous studies have reported factors associated with a cardiac death in cats with HCM, including our previous report of the population included in this study, but the risk factors have not been reported separately for CHF vs. ATE or sudden death. If risk factors vary for different forms of cardiac death, this could have important implications for design of therapeutic trials and the prevention of these complications of HCM. Although no treatment is known to prevent the onset of CHF in cats with HCM, antithrombotic treatment is commonly recommended for cats believed to be at risk of ATE, even though risk factors for ATE have largely been presumed based on clinical experience and anecdotal evidence. Our previous analysis of the cohort of cats with HCM reported here yielded multiple echocardiographic risk factors for cardiac death, but no attempt was made to link these risk factors with specific types of mortality end-point. We reanalyzed the data from the previous study to evaluate the specific risk factors for different cardiac mortality end-points: death due to CHF; death due to ATE; and death due to sudden death. We found that initial clinical presentation remained an important predictor for all three types of cardiac death, with asymptomatic cats at lower risk of all forms of cardiac mortality. Multivariable analysis showed CHF on presentation and LV systolic dysfunction to be independent predictors of CHF death, and ATE on presentation and reduced LA-FS% as independent predictors of ATE death. Univariable analysis demonstrated an increased risk for sudden death in cats with a history of syncope, or arrhythmias. In the current population, cardiac mortality was most commonly associated with CHF (55.7%), then ATE (29.1%) and least commonly sudden death (15.2%). In the 2002 study by Rush et al., ATE was the most common cause of cardiac death, followed by CHF and then sudden death. Where type of cardiac death has been reported in survival studies in cats with HCM, sudden death is usually the least common type of death, but data on prevalence of sudden cardiac death in cats with HCM are generally scarce. The proportion of sudden deaths reported by owners to
their own veterinarian is not known. In our cohort, none of the primary veterinarians of the cats that experienced sudden death at home were aware that the cat had died, supporting the notion of under-reporting. One cat in this study developed ventricular fibrillation, but the cause of sudden death in the other cats was not known. Alternative possible causes include bradyarrhythmias, coronary or central nervous systemic thromboembolic events or a thrombus within the LV outflow tract. In people, sudden cardiac deaths appear to be more common in young adults, with CHF related deaths more common in middle-aged people, and embolic related deaths more likely to occur in the elderly. In the study by Trehiou-Sechi et al., Maine Coons that died suddenly were younger than other breeds with sudden death. In our population of cats, CHF deaths within 2 years of diagnosis occurred in older cats than in those who did not experience a CHF death, but age was not associated with either ATE deaths or sudden deaths.

Factors previously associated with ATE in the literature include LA enlargement, increased LV diameter, and SEC. All the LA variables (LA:Ao, LAD, LA-FS%, LA-EF%) were also associated with ATE risk in our study, as well as SEC or presence of a thrombus, increasing LV wall thickness, presence of a gallop, and restrictive diastolic filling, but not LV systolic dysfunction or regional wall motion abnormalities. The multivariable model showed that a history of ATE and reduced LA-FS% were independent predictors. The initial clinical presentation of the cat was an important predictor of specific mortality endpoints. Cats presenting with CHF were at increased risk of dying with CHF but not of ATE, and cats presenting with ATE were at increased risk of dying with ATE but not of CHF. Cats with syncope were more likely to experience sudden death, although this was only explored at the univariable level because of a low event rate. Syncope is an important risk factor for sudden death in people and our data suggest this may also be true in cats. Left atrial variables were predictive of CHF deaths and ATE deaths. Cats with CHF have been reported to have worse LA function than healthy controls. Left atrial enlargement has
consistently been one of the most important risk factors reported for cardiac mortality in previous
studies and reduced LA-FS% was an independent predictor of cardiac mortality in the previous overall
survival analysis of the cats in the present study.11 It is therefore likely that a large, poorly contractile
left atrium increases the risk of death due to both CHF and ATE. In human cardiovascular disease, the
presence of SEC has been associated with embolic events and it has been assumed that this is also the
case in cats with HCM.16 In these results, however, SEC appears to predict any type of cardiac end-point
rather than being specific for ATE. Given the previously reported association with SEC and left auricular
appendage velocities, it is likely that SEC is a surrogate marker for LA dysfunction and as such can
predict a CHF or ATE death. At the univariable level LA:Ao, LAD and LA-EF% were predictive of sudden
death. However, it is more difficult to assess the predictive value for sudden death because of low
numbers.

Cats with LV systolic dysfunction (FS% <30%) or regional wall hypokinesis were at increased risk of a CHF
death or sudden death at the univariable level, with FS% <30% being an independent predictor of a CHF
death. Human HCM patients that have an end-stage phenotype (LV systolic dysfunction, defined as a
left ventricular ejection fraction <50%) are more likely to progress to advanced CHF than those without
systolic dysfunction20, 24, 25 and FS% <30% been reported to be associated with increased risk of
cardiovascular mortality in people.46 The end-stage phenotype of HCM has been suggested as a
predictor of sudden death in people.24, 25 LV systolic function appears to have similar predictive value in
cats as in people, although further work investigating the role in sudden death is required.

In our original survival analysis of this population, LV hypertrophy was an independent predictor of
cardiac mortality and this sub-analysis suggests the increased risk is primarily associated with death due
to CHF or ATE, which contrasts with findings in people where extreme hypertrophy is associated with an increased risk of sudden death.

Auscultation findings showed some predictive value at the univariable level, with the presence of a murmur significantly associated with freedom from a CHF death within 2 years of diagnosis. The presence of a murmur is often due to the presence of SAM causing dynamic left ventricular outflow tract obstruction (LVOTO), and SAM was also associated with freedom from a CHF death. A number of feline studies have reported a similar association between a favorable outcome and a murmur or dynamic LVOTO. It could be speculated that cats with SAM are detected earlier in their disease course than cats without SAM because diagnostic investigations are undertaken following detection of a murmur rather than demonstration of clinical signs, which might explain the apparent survival benefit of both a murmur and SAM in cats in the short term. There are no studies in cats to evaluate the long term effect of dynamic LVOTO on cardiac mortality, so it is not clear if cats genuinely differ from people with HCM, in whom the presence of LVOTO is associated with progression to severe heart failure. Neither the presence of a murmur nor the presence of SAM was associated with a risk of an ATE death or sudden death. Gallop sounds were associated with an ATE death but not with a CHF death. Arrhythmias (both on auscultation and on electrocardiography) were associated with an increased risk of a CHF death or a sudden death, but it was not possible to analyze individual arrhythmia types because of low numbers.

This is a retrospective study conducted in a referral environment and as such, there are limitations. Blood pressure was not recorded in every cat, so that it is possible that hypertensive cats were included, although efforts were made to exclude cats without a blood pressure measurement if they had factors predisposing to hypertension, such as renal disease. Serum thyroxine concentrations were not routinely measured unless clinically indicated and so it is possible that some hyperthyroid cats may have been
included. Each cat only had 1 echocardiographic examination measured and used for the study. As HCM is a disease that progresses at a variable rate, some cats may have died a cardiac death within two years of diagnosis without any of the risk factors being obvious at the point of examination. Therefore the absence of these risk factors does not exclude the potential for a cat dying within two years of diagnosis. At the time of admission some cats were showing signs of CHF and so did not receive a full echocardiographic examination until stabilized. Some echocardiographic measurements (eg left atrial size or function) are likely to have changed during stabilization. Equally, some cats were already receiving treatment at admission, which may have affected their echocardiographic measurements. Not all echocardiographic measurements were available for every cat, either due to clinician preference, patient stability or temperament. Owner descriptions and clinical records of referring practices were used to determine the outcome for the cats. It is possible that some deaths were missed, particularly as 27 cats could not be included due to lack of follow up data. Reliance was placed on owner description for deciding what type of death the cat had experienced. There were only low numbers of sudden deaths and ATE deaths within 2 years of diagnosis, limiting the ability to generate meaningful multivariable models. Veterinary survival studies are always confounded by variability in selection of timing for euthanasia versus survival attributable to spontaneous death, and this is also true of our study.

In conclusion, the most important predictor of the type of cardiac death appears to be symptomatic status at first presentation, with those presenting with either CHF or an ATE most likely to die of the same complication within 2 years, and those presenting with syncope are at higher risk of sudden death. The variables LA:Ao, LAD and LA-EF% were all associated with an increased risk for all three types of cardiac death. Left atrial fractional shortening was associated with an increased risk for CHF and ATE deaths.
327 References


Figure 1 – Risk of death due to congestive heart failure (CHF); univariable hazard ratios of physical exam and echocardiographic findings at presentation between cats that did and did not suffer a CHF death within 2 years of diagnosis. CHF – congestive heart failure, ATE – arterial thromboembolism, FS% - Left ventricular fractional shortening, LA:Ao - short axis ratio of diastolic left atrial diameter to aortic root diameter, LAD – the diameter of the left atrium measured parallel with the mitral annulus in the last frame before mitral valve opening, LA-EF% – left atrial emptying fraction, LA-FS% – left atrial fractional shortening, Max 2D LVWd – maximum end-diastolic left ventricular septal or free wall thickness measured on a 2D image, SEC – spontaneous echo contrast, MR – mitral regurgitation, SAM – systolic anterior motion of the mitral valve, \( a \) – for a unit change of 1 year, \( b \) – for a unit change of 1 beat per minute, \( c \) – for a unit change of 1 breath per minute, \( d \) – for a unit change of 10%, \( e \) – for a unit change of 1, \( f \) – for a unit change of 1 mm
Figure 2 - Risk of death due to arterial thromboembolism (ATE); univariable hazard ratios of physical exam and echocardiographic findings at presentation between cats that did and did not suffer an ATE death within 2 years of diagnosis. CHF – congestive heart failure, ATE – arterial thromboembolism, FS% - Left ventricular fractional shortening, LA:Ao - short axis ratio of diastolic left atrial diameter to aortic root diameter, LAD – the diameter of the left atrium measured parallel with the mitral annulus in the last frame before mitral valve opening, LA-EF% – left atrial emptying fraction, LA-FS% – left atrial fractional shortening, Max 2D LVWd – maximum end-diastolic left ventricular septal or free wall thickness measured on a 2D image, SEC – spontaneous echo contrast, MR – mitral regurgitation, SAM – systolic anterior motion of the mitral valve, a – for a unit change of 1 year, b – for a unit change of 1 beat per minute, c – for a unit change of 1 breath per minute, d – for a unit change of 10%, e – for a unit change of 1, f – for a unit change of 1 mm
Figure 3 - Risk of death due to sudden death; univariable hazard ratios of physical exam and echocardiographic findings at presentation between cats that did and did not suffer a sudden death within 2 years of diagnosis. Nb No pedigree cat died suddenly so it was not possible to generate hazard ratios, a chi-squared was non-significant (p=0.130). No cat that presented with ATE died suddenly so it was not possible to generate hazard ratios, a chi-squared was non-significant (p=1.000).

CHF – congestive heart failure, ATE – arterial thromboembolism, FS% – Left ventricular fractional shortening, LA:Ao - short axis ratio of diastolic left atrial diameter to aortic root diameter, LAD – the diameter of the left atrium measured parallel with the mitral annulus in the last frame before mitral valve opening, LA-EF% – left atrial emptying fraction, LA-FS% – left atrial fractional shortening, Max 2D LVWd – maximum end-diastolic left ventricular septal or free wall thickness measured on a 2D image, SEC – spontaneous echo contrast, MR – mitral regurgitation, SAM – systolic anterior motion of the mitral valve, a – for a unit change of 1 year, b – for a unit change of 1 beat per minute, c – for a unit change of 1 breath per minute, d – for a unit change of 10%, e – for a unit change of 1, f – for a unit change of 1 mm.
<table>
<thead>
<tr>
<th></th>
<th>OR (95% CI)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>CHF at first presentation</td>
<td>6.318 (1.699 – 23.496)</td>
<td>0.006</td>
</tr>
<tr>
<td>LA-FS%</td>
<td>0.892 (0.799 – 0.994)</td>
<td>0.039</td>
</tr>
<tr>
<td>FS%</td>
<td>0.965 (0.917 – 0.997)</td>
<td>0.036</td>
</tr>
<tr>
<td>Respiratory rate</td>
<td>1.023 (1.001 – 1.045)</td>
<td>0.045</td>
</tr>
</tbody>
</table>

Table 1 - Binary logistic regression model, predicting CHF death within two years of diagnosis, (25 events of 172 cases)

<table>
<thead>
<tr>
<th></th>
<th>OR (95% CI)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>ATE at first presentation</td>
<td>28.58 (6.05 – 135.03)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>LA-FS%</td>
<td>0.84 (0.77 – 0.92)</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>

Table 2 – Binary logistic regression model, predicting ATE death within two years of diagnosis, (20 events of 200 cases)