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Heart rate turbulence after ventricular premature beats in healthy Doberman pinschers and those with dilated cardiomyopathy

J.D. Harris, BVSc ^{a,*}, C.J.L. Little, BVM&S PhD ^b, J.M. Dennis, BSc MSc ^c, M.W. Patteson, MA VetMB PhD ^a

^a HeartVets, The Animal Hospital, Stinchcombe, Dursley, Gloucestershire, GL11 6AJ, UK ^b Barton Veterinary Hospital, 34 New Dover Road, Canterbury, Kent, CT1 3BH, UK ^c Health Statistics Group, Institute of Health Research, University of Exeter Medical School, Exeter, EX1 2LU, UK

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| KEYWORDS | Abstract Objectives: To describe the measurement of heart rate turbulence |
|-------------------------|---|
| Heart rate variability; | (HRT) after ventricular premature beats and compare HRT in healthy Doberman |
| Canine; | pinschers and those with dilated cardiomyopathy (DCM), with and without conges- |
| Congestive heart | tive heart failure (CHF). |
| failure; | Animals: Sixty-five client-owned Dobermans: 20 healthy (NORMAL), 31 with precli- |
| Autonomic nervous | nical DCM and 14 with DCM and CHF (DCM $+$ CHF). |
| system; | Methods: A retrospective study of data retrieved from clinical records and ambula- |
| Holter monitoring | tory ECG (Holter) archives, including data collected previously for a large-scale pro- |
| | spective study of Dobermans with preclinical DCM. Holter data were reanalysed |
| | quantitatively, including conventional time-domain heart rate variability and the |
| | HRT parameters turbulence onset and turbulence slope. <i>Results:</i> Heart rate turbulence could be measured in 58/65 dogs. Six Holter record- |
| | ings had inadequate ventricular premature contractions (VPCs) and one exhibited |
| | VPCs too similar to sinus morphology. Heart rate turbulence parameter, turbulence |
| | onset, was significantly reduced in DCM dogs, whereas conventional heart rate |
| | variability measures were not. Heart rate variability and HRT markers were reduced |
| | in DCM $+$ CHF dogs as expected. |
| | Conclusions: Heart rate turbulence can be measured from the majority of good |
| | quality standard canine 24-hour Holter recordings with $>$ 5 VPCs. Turbulence onset |

* Corresponding author.

E-mail address: jo@heartvets.co.uk (J.D. Harris).

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is significantly reduced in Dobermans with preclinical DCM which indicates vagal withdrawal early in the course of disease. Heart rate turbulence is a powerful prognostic indicator in human cardiac disease which can be measured from standard 24-hour ambulatory ECG (Holter) recordings using appropriate computer software. Further studies are warranted to assess whether HRT may be of prognostic value in dogs with preclinical DCM and in other canine cardiac disease. © 2017 Elsevier B.V. All rights reserved.

| A 1. 1. | |
|------------|---------------|
| Δnn | reviations |
| NDD | i c v lacions |
| | |

| ANS BRS | autonomic nervous system baroreflex sensitivity |
|------------|--|
| CHF | congestive heart failure |
| CIPA | Cardiac Index of Parasympathetic |
| | Activity |
| DCM | dilated cardiomyopathy |
| ECG | electrocardiogram |
| HRT | heart rate turbulence |
| HRV | heart rate variability |
| LV | left ventricle |
| LVIDd | left ventricular internal dimension at |
| | end-diastole |
| LVIDs | left ventricular internal dimension at |
| | end-systole |
| RMSSD | square root of the mean of the sum |
| | of the squares of differences |
| | between adjacent NN intervals |
| SDANN | standard deviation of the averages of |
| | NN intervals in all 5 min segments of |
| | the entire recording |
| SDNN | standard deviation of all intervals |
| | between normal beats (NN intervals) |
| SMOD | Simpson's method of discs |
| TI | triangular index |
| TO | turbulence onset |
| TS | turbulence slope |
| VPC | ventricular premature complex |
| VVTI | vasovagal tonus index |
| | |

Introduction

The autonomic nervous system (ANS) and its influence on cardiovascular health has been extensively studied in recent years. This has led to a shift in the understanding of the pathophysiology of the heart failure syndrome, particularly the neurohumoral response which becomes abnormally activated in heart disease. It has become widely accepted that there is a relationship between ANS dysfunction and cardiovascular disease [1], which in humans manifests as a progressive withdrawal of parasympathetic tone and stimulation of sympathetic activity. This has provoked significant interest in quantitative estimates of autonomic tone to provide risk stratification for stroke and cardiac mortality in humans including sudden cardiac death.

Various markers of ANS function have been proposed, the simplest of which are derived from changes in heart rate in response to various stimuli. In resting humans and dogs, the heart rate is predominantly under parasympathetic control, mediated through reflex baroreceptor stimulation. Sympathetic influence affects the heart more slowly, both by direct effect on heart rate and by changes in peripheral vascular resistance [2]. Heart rate alone indicates the net effect of the sympathetic and parasympathetic nervous systems at any one time but does not disclose the two components of the ANS individually.

Heart rate variability (HRV) is the collective term for changes in instantaneous heart rate, measured by beat-to-beat (RR interval) changes on an electrocardiogram (ECG) [3]. Different HRV parameters reflect either predominantly sympathetic or parasympathetic activity and hence can more accurately guantify the interaction between the two systems than measurement of heart rate alone. Statistical calculations on direct measurements of intervals between normal beats (NN intervals) or differences between NN intervals are known as time-domain measures of HRV. In humans, HRV can predict severity of myocardial failure [4] as well as risk of cardiac mortality in both ischaemic and non-ischaemic cardiomyopathy [5,6]. Attempts have been made to demonstrate a similar relationship in dogs. Whilst this has improved the understanding of autonomic dysfunction in canine cardiac disease, these studies have thus far failed to identify conventional HRV techniques as clinically useful tools before the onset of congestive heart failure (CHF) [7-9]. This has been attributed to the predominance of vagal withdrawal in dogs rather than the accompanying sympathetic activation observed in humans [10,11]. It has been concluded that better non-

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