Prognostic value of echocardiographic and Doppler parameters in horses admitted for colic complicated by systemic inflammatory response syndrome

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Abstract

Objective – To assess the prognostic value of echocardiographic parameters of left ventricular (LV) function in horses with systemic inflammatory response syndrome (SIRS).

Design – Prospective observational study.

Setting – Veterinary teaching hospital.

Animals – Forty-one horses admitted for colic with clinical evidence of SIRS.

Interventions – All horses underwent Doppler echocardiographic examination on admission. LV echocardiographic parameters, including pulsed-wave tissue Doppler imaging parameters, were compared between nonsurvivors (n = 29) and horses that survived to discharge (n = 12).

Measurements and Main Results – With comparable heart rate and LV preload estimate, LV stroke volume index, the velocity time integral, deceleration time, ejection time of Doppler aortic flow, and peak early diastolic myocardial velocity were lower in the nonsurviving than in the surviving horses, while pre-ejection period to ejection time ratio (PEP/ET) of Doppler aortic flow and the peak early diastolic filling velocity to peak early diastolic myocardial velocity ratio (E/Em) were higher (P < 0.05). A cut-off value of 0.26 for PEP/ET predicted mortality with 100% sensitivity and 42% specificity (area under the receiver operating characteristic curve: 0.71), whereas a cut-off value of 2.67 for E/Em predicted mortality with 100% sensitivity and 83% specificity (area under the receiver operating characteristic curve: 0.89).

Conclusions – Echocardiography may provide prognostic information in colic horses with clinical evidence of SIRS. Especially, PEP/ET and E/Em could be useful markers of systolic and diastolic dysfunction, respectively, to detect horses with a high risk of death requiring more intensive cardiovascular monitoring as it has been reported in human patients with septic shock.


Keywords: Endotoxemia, equine intensive care, hemodynamics, prognosis

Abbreviations

CI cardiac index

CO cardiac output

2D two-dimensional mode

EDV left ventricular end-diastolic volume

E/Em peak early left ventricular filling velocity to peak early diastolic myocardial velocity ratio

Em peak early diastolic myocardial velocity

EF left ventricular ejection fraction

ET left ventricular ejection time

ETI left ventricular ejection time index corrected for heart rate

HR heart rate

IVRT isovolumetric relaxation time

LV left ventricle

MODS multiple organ dysfunction syndrome

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Introduction

Endotoxia is a common and potentially fatal complication of acute intestinal disease in adult horses. Endotoxic shock, like septic shock, involves hypovolemic, cardiogenic, and distributive components resulting in tissue hypoperfusion with a high risk for multiple organ dysfunction syndrome (MODS) and increased mortality in equine and human ICUs. As a result, most clinical and laboratory parameters currently used as prognostic indicators of colic in adult horses are related to cardiovascular status.

Endotoxins stimulate a systemic inflammatory response syndrome (SIRS), which is associated with severe hemodynamic disturbances via vasoplegia, a drastic reduction in venous return and myocardial depression causing a drop in systemic vascular resistance and cardiac output (CO). Sympathetic stimulation and release of various endogenous substances provide a compensatory vasoconstriction and an increase in heart rate (HR), myocardial contractility and venous return to preserve CO. However, loss of vascular responsiveness to catecholamines and myocardial depression finally contribute to a progressive drop in CO and systemic vascular resistance despite persistent tachycardia.

A shock score (SS) combining several clinical and laboratory parameters on admission enabled prediction of survival in colic horses but could not discriminate, which severely affected horses with shock were likely to respond to treatment. A comparable “Mortality in Emergency Department Sepsis Score,” based on clinical and laboratory variables is currently considered inaccurate in predicting mortality in people with severe sepsis or septic shock, especially in the presence of tissue hypoperfusion associated with a high risk of MODS. Hemo-

dynamic monitoring is required in these patients to predict the response to fluid therapy, the need for vasopressors or inotrope drugs and outcome. A poor prognosis has been associated with persistent tachycardia along with low systemic vascular resistance, a refractory response to catecholamines, and slow lactate clearance or low central venous oxygen saturation postresuscitation. In critically ill horses, lactate clearance after resuscitation recently emerged as a promising marker of systemic hypoperfusion with some prognostic value.

Although frequently observed in people with septic shock, myocardial failure is rarely the cause of death in these patients who succumb mostly from MODS. Nevertheless, serum concentrations of cardiac biomarkers (ie, cardiac troponin T, amino-terminal pro-brain natriuretic peptide), and echocardiographic and Doppler parameters of left ventricular (LV) systolic and diastolic function proved to be strongly associated with outcome in people with septic shock.

A moderate increase in serum cardiac troponin I in the postoperative period has also recently been associated with a poor outcome in 34 horses undergoing emergency laparotomy. A previous echocardiographic study has suggested a systolic dysfunction in colic horses with clinical signs of endotoxic shock. To date however, the prognostic value of echocardiographic parameters is unknown.

The aim of this prospective study was to investigate the value of two-dimensional (2D), Motion-Mode (M-mode), Doppler, and pulsed-wave tissue Doppler imaging (PW-TDI) echocardiography in predicting the outcome of colic in adult horses with clinical signs and laboratory parameters compatible with SIRS.

Material and Methods

Study population

Horses referred between March 2009 and September 2011 for colic with clinical and laboratory parameters compatible with SIRS at admission were included in the study (Table 1). The presence of SIRS was assumed when horses were hyperlactatemic (plasma lactate > 2 mmol/L) or had low systolic arterial pressure (< 90 mm Hg) and fulfilled at least 2 of the following criteria: 1. hypo- or hyperthermia (rectal temperature <36.7°C or > 38.6°C), 2. leukopenia or leukocytosis (WBC < 5,000/μL or >14,500/μL), 3) tachycardia (HR >50/min) or 4) tachypnea (respiratory rate > 25/min). Horses euthanized for financial considerations, immature horses (< 2 years old), or horses that received α2-receptor agonists before echocardiographic examination were excluded from the study.

Clinical data and blood analysis recording

HR, respiratory rate, and rectal temperature were recorded at admission for each horse. Jugular vein EDTA blood samples were obtained and immediately analyzed. Lactate concentration was measured using a portable device and WBC was determined using an automatic analyzer. Systolic arterial pressure was
measured noninvasively by a manometer equipped with an occlusion cuff placed over the coccyeal artery.

**Echocardiography**

Each horse underwent echocardiographic examination on admission. All echocardiographic records were performed and analyzed by the same operator (LB) using a portable echocardiograph equipped with a 1.5–3.6 MHz transducer. It permitted echocardiographic view numeric recording (still frames or cine-loops) with a simultaneous continuous single lead base-apex ECG for delayed analysis. All examinations were recorded digitally and analyzed offline.

Echocardiographic recordings and measurements were obtained conforming to conventional methods. Assessment of aorta and cardiac chamber sizes was performed using standard transthoracic 2D and M-mode echocardiography. The following standard views were recorded for analysis: a right parasternal 2D long-axis 5 chamber view, a TM short-axis view of the LV at the level of the chordae tendinae, and a left parasternal 2D long-axis 4 chamber angled view.

Assessment of LV wall motion was obtained from the standard left parasternal short-axis view of the LV at the level of the chordae tendinae by placing a sample volume of 5.9 mm in the endocardial region of the LV free wall in diastole with the PW-TDI mode. Special attention was paid to keeping the sample volume on the myocardium during all phases of the cardiac cycle. The velocity scale was set to −30 cm to +20 cm/s.

Assessment of aortic flow was performed using standard transthoracic PW-Doppler echocardiography. Transaortic flow velocity profiles were obtained from the standard left parasternal long axis 5-chamber view in PW-Doppler mode by placing the sample volume (4.91 mm) downstream from the aortic valve in the aortic root.

Transmitral flow velocity profiles were recorded by positioning the transducer as ventrally as possible and angling dorsally from the standard left parasternal long axis 4-chamber view in PW-Doppler mode with the sample volume placed between the opened tips of the mitral leaflets in diastole.

Table 1: Comparison of the means ± SD of the parameters used to determine the presence of systemic inflammatory response syndrome in 12 surviving and 29 nonsurviving adult horses admitted for colic. Least-squares-mean test was significant if P < 0.05 (ANOVA model including breed, body weight, age, sex, and outcome).

<table>
<thead>
<tr>
<th></th>
<th>Survivors</th>
<th>Nonsurvivors</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Heart rate (per min)</td>
<td>65.64 ± 18.43</td>
<td>79.71 ± 26.52</td>
<td>0.0818</td>
</tr>
<tr>
<td>Respiratory rate (per min)</td>
<td>27.1 ± 10.6</td>
<td>30.7 ± 14</td>
<td>0.1544</td>
</tr>
<tr>
<td>WBC (count/µL)</td>
<td>12.01 ± 4.69</td>
<td>9.31 ± 5.32</td>
<td>0.26</td>
</tr>
<tr>
<td>Rectal temperature (°C)</td>
<td>37.62 ± 0.85</td>
<td>37.8 ± 0.66</td>
<td>0.6135</td>
</tr>
<tr>
<td>Plasma lactate</td>
<td>3.57 ± 3.67</td>
<td>3.43 ± 2.28</td>
<td>0.9999</td>
</tr>
<tr>
<td>concentration (mmol/L)</td>
<td>94.7 ± 0</td>
<td>122.9 ± 26</td>
<td>–</td>
</tr>
</tbody>
</table>

SAP: systolic arterial pressure (noninvasive); ‘ insufficient data for statistical analysis; only available in 9 horses including only 1 survivor.
In case of impaired relaxation (but also with tachycardia or a falling preload), the relaxation continues until the late diastolic filling phase and the atrial contribution increases in the ventricular filling leading to a prolonged IVRT associated with a low E/A ratio on the mitral flow profile combined with a low Em on PW-TDI. Conversely, a restrictive ventricular filling leads to a normal or shorter IVRT associated to high E/A and E/Em.²⁷

**Statistical analysis**

For statistical analysis, horses were separated into 2 groups according to outcome: nonsurvivor or survived to discharge. Means and SDs were calculated for each group and for each studied parameter. Some parameters were corrected for body weight before statistical analysis (index = variable/body weight). After verifying that breed, sex, and outcome were independent (Pearson chi-square test of independence), a series of ANOVA models were used to test whether echocardiographic and Doppler parameters were statistically different across the 2 groups, after adjusting for the effects of breed, sex, weight, and age. The ANOVA model was as follows:

\[ y_{ijkl} = \mu + C_i + B_j + S_k + b_1 W_{ijkl} + b_2 A_{ijkl} + e_{ijkl}, \]

where \( y \) was the echocardiographic or Doppler parameter on the \( m \)th horse with survival \( C_i \) (\( i = 0, 1 \)), breed \( B_j \) (\( j = 1, 2, \ldots, 5 \)), sex \( S_k \) (\( k = 1, 2, 3 \)), weight (\( W_{ijkl} \)) and age (\( A_{ijkl} \)), \( \mu \) was the overall mean, \( b_1 \) and \( b_2 \) are the linear regression effect of \( W_{ijkl} \) and \( A_{ijkl} \) on \( y_{ijkl} \), and \( e_{ijkl} \) was the residual term, assuming independence between error terms. The null hypothesis of no difference between ordinary least-squares means for the survival was tested.

The ability of each parameter to predict survival was assessed by calculating the area under the receiver operating characteristic (ROC) curve and the Youden index for each significant echocardiographic parameter (logistic procedure) separately. The cut-off value offering the best balance between sensitivity and specificity. A commercial statistical software package was used for all statistical analyses. Statistical significance was set at \( P < 0.05 \).

**Results**

Forty-one horses fulfilled the inclusion criteria during the study period. Each horse had an SS of at least 3 of 4 according to criteria set by Grulke et al.⁶ (ie, HR>80/min, RR>35/min, PCV>55%, systolic arterial pressure <90 mm Hg or plasma lactate >2 mmol/L).⁶ Animals included were 20 geldings, 17 mares, and 4 stallions. Breed distribution was 31 Warmbloods, 1 Arabian, 1 Spanish Breed, 1 Frisian, and 7 riding ponies. Mean ± SD age was 11.1 ± 6.6 years (range: 2–27 years) and mean ± SD body weight was 484.5 ± 78.7 kg (range: 270–620 kg). Twelve horses survived to discharge from the hospital. One horse died spontaneously at induction of anesthesia. Twenty-eight horses were subjected to euthanasia, either early after admission due to a poor surgical prognosis (15 horses), or during the postoperative period (13 horses) because of reocurrence of colic (8 horses) or multiple organ failure (5 horses).

Clinical and laboratory parameters were not significantly different between survivors versus nonsurvivors horses (Table 1). Echocardiographic studies for data collection (3 views recording) did not exceed 15 minutes and were easily integrated into the admission protocol during the time allowed for rectal palpation and abdominal ultrasound. Values for mean ± SD echocardiographic parameters derived from 2D-mode obtained in the 2 groups and results of the ANOVA test are shown in Table 2. Although the influence of body weight was significant for several morphological echocardiographic variables including end-diastolic and end-systolic LV internal diameter, LV end-diastolic volume (EDV) and end-systolic volume (ESV), none of these variables were significantly different between the 2 outcome groups (survivors versus nonsurvivors). The EDV and ESV indexes corrected for body weight (respectively, EDVI and ESVI), LV fractional shortening and LV EF did not differ significantly between survivors and nonsurvivors.

Values for mean ± SD Doppler parameters of systolic function obtained in each group and the results of the ANOVA test are shown in Table 3. CO, cardiac index (CI), as well as maximal velocity (Vmax), pre-ejection-period (PEP), acceleration time and acceleration rate of aortic flow ejection, were not significantly different between the 2 groups. However, the velocity time integral, deceleration time and ET of aortic flow, stroke volume, stroke index (SI), and HR-corrected ET (ETI) were significantly lower, and the PEP to ET ratio (PEP/ET) as well as the mean velocity of circumferential fiber shortening (Vcf) were significantly higher in nonsurvivors compared to survivors.

The area under the ROC curve was 0.73 for SI, 0.76 for ET, 0.74 for ETI, and 0.71 for PEP/ET. A cut-off value of 1.78 mL/kg for SI predicted mortality with 83% sensitivity and 62% specificity (Youden index \( Y = 0.56 \)). A cut-off value of 340 ms for ET predicted mortality with 75% sensitivity and 69% specificity (\( Y = 0.44 \)). A cut-off value of 377 ms for ETI predicted mortality with 75% sensitivity and 69% specificity (\( Y = 0.44 \)). A cut-off value of 0.26 for PEP/ET predicted mortality with 100% sensitivity and 42% specificity (\( Y = 0.41 \); Figure 1).

Values for mean ± SD Doppler parameters and PW-TDI parameters of diastolic function obtained in each
Table 2: Comparison of the means ± SD values of 2D and M-mode echocardiographic parameters in 12 surviving and 29 nonsurviving adult horses admitted for colic complicated by systemic inflammatory response syndrome. Least-squares-mean test was significant if P < 0.05 (ANOVA model including breed, body weight, age, sex, and outcome).

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Survivors n = 12</th>
<th>Nonsurvivors n = 29</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Ao (cm)</td>
<td>7.8 ± 0.82</td>
<td>7.55 ± 0.68</td>
<td>0.0818</td>
</tr>
<tr>
<td>IVSd (cm)</td>
<td>3.61 ± 0.48</td>
<td>3.61 ± 0.57</td>
<td>0.7191</td>
</tr>
<tr>
<td>IVSs (cm)</td>
<td>4.6 ± 0.49</td>
<td>4.52 ± 0.56</td>
<td>0.8162</td>
</tr>
<tr>
<td>LVIDd (cm)</td>
<td>9.26 ± 1.29</td>
<td>8.84 ± 1.55</td>
<td>0.4835</td>
</tr>
<tr>
<td>LVIDs (cm)</td>
<td>5.71 ± 1.24</td>
<td>5.34 ± 1.26</td>
<td>0.2900</td>
</tr>
<tr>
<td>LVPWd (cm)</td>
<td>3.09 ± 0.61</td>
<td>3.08 ± 0.65</td>
<td>0.7371</td>
</tr>
<tr>
<td>LVPWs (cm)</td>
<td>4.24 ± 0.52</td>
<td>4.34 ± 0.57</td>
<td>0.5889</td>
</tr>
<tr>
<td>LA (cm)</td>
<td>10.95 ± 1.27</td>
<td>10.27 ± 1.37</td>
<td>0.1827</td>
</tr>
<tr>
<td>EDV (mL)</td>
<td>489.95 ± 152.72</td>
<td>447.16 ± 167.06</td>
<td>0.5353</td>
</tr>
<tr>
<td>EDVI (mL/kg)</td>
<td>0.991 ± 0.318</td>
<td>0.905 ± 0.287</td>
<td>0.4142</td>
</tr>
<tr>
<td>ESV (mL)</td>
<td>177.35 ± 88.11</td>
<td>148.64 ± 70.43</td>
<td>0.2371</td>
</tr>
<tr>
<td>ESVI (mL/kg)</td>
<td>0.355 ± 0.175</td>
<td>0.3 ± 0.134</td>
<td>0.2154</td>
</tr>
<tr>
<td>%FS (%)</td>
<td>38.4 ± 7.94</td>
<td>39.73 ± 9.9</td>
<td>0.3784</td>
</tr>
<tr>
<td>EF (%)</td>
<td>65.55 ± 9.12</td>
<td>66.81 ± 11.18</td>
<td>0.4399</td>
</tr>
</tbody>
</table>

Ao: aortic diameter; EDV: LV end-diastolic volume; EDVI: LV end-diastolic volume index = EDV/body weight; EF: LV ejection fraction; ESV: left ventricular end-systolic volume; ESVI: ESV index = ESV/body weight; %FS: fractional shortening; IVS: interventricular septum thickness; LA: left atrium; LV: left ventricle; LVPW: LV free wall thickness; LVd: LV internal diameter; n: number of horses; s: end-systolic; d: end-diastolic.

Discussion

Our results suggest that in horses with colic and naturally-occurring SIRS, those patients which succumb or are euthanized show more pronounced LV systolic and diastolic dysfunction compared to those which survive. Certain markers of systolic (SI, ET, ETI, PEP/ET) or diastolic (Em, E/Em) function emerged as valuable predictors of mortality in these cases. As the SS based mainly on HR and blood lactate concentration permits to establish a prognosis in colic horses on admission, echocardiography could be used to predict more accurately the chances of survival of already shocked horses presented with tachycardia and hyperlactatemia.

A previous echocardiographic study in horses admitted for colic signs suggested that systolic dysfunction occurs with endotoxin shock; however, the results were confused by concurrent tachycardia and a drastic drop of EDV, which can both mimic systolic as well as diastolic dysfunction. In the present study, all cases were selected because of the presence of SIRS. Therefore, they all exhibited a high HR and 2D and TM-mode echocardiographic evidence of a drastically low LV preload as indicated by a very low EDV compared to normal values in euvhylated horses or even experimentally hypohydrated horses. Since HR or EDV were not significantly different between surviving and nonsurviving horses in the present study, tachycardia and falling preload were assumed not to have greatly influenced outcome.

Although EDV is recognized as one of the best preload estimates in people with early septic shock, the frequent systolic dysfunction and dynamic changes in LV compliance associated with septic shock preclude the...
Echocardiography and prognosis in colic horses

Figure 1: Receiver operating characteristic curves of PEP/ET and E/Em to predict hospital mortality in 41 horses admitted for colic complicated by systemic inflammatory response syndrome. AUC: area under the receiver operating characteristic curve; E/Em: peak early transmitral flow velocity to peak early diastolic myocardial velocity ratio; PEP/ET: aortic flow pre-ejection period to ejection time ratio.

use of EDV to predict fluid responsiveness in these patients unless LV size is very small (preload dependent patients).37,38

While indicating a favorable prognosis in people with septic shock by predicting a better response to fluid resuscitation,5,11 in the present study a low SI (<1.78 mL/kg) on admission predicted an unfavorable outcome (83% sensitivity, 62% specificity). This discrepancy could be due to impaired myocardial contractility which may worsen the influence of reduced preload in nonsurviving horses, or may be due to the large volumes of fluid required by adult horses compared to people because of body size. Therefore, any preload-dependent systolic impairment is less likely to be corrected in adult horses.7

Systolic time intervals are known to be more reliable indices of systolic function than Vcf or EF.39 However, the changes observed in this study may also be due to tachycardia or falling preload,29,39 as has been described in hemodialyzed human patients.32 Considering the comparable HR and preloading conditions in both groups, the lower ET with a higher PEP/ET ratio in the nonsurvivors may indicate a more compromised systolic function compared to surviving horses.

An increased PEP/ET ratio (>0.26) predicted mortality in colic horses with a high sensitivity (100%) but a low specificity (42%) in the present study. Since PEP and ET change in opposite directions with systolic dysfunction, PEP/ET is known to be more sensitive for predicting systolic dysfunction than either of these parameters alone.20 In an early study in people with sepsis, a high PEP/ET on admission also carried a poor prognosis,40 and this ratio could be used as in human ICUs to identify horses with severe LV systolic dysfunction and a high risk of death.

Concerning diastolic function, impaired relaxation was suspected in both groups, based on the mitral flow profile with a low E/A ratio combined with a prolonged IVRT compared to previously published reference values in adult horses.28,30 This pattern may however simply result from the reduced preload22 or from the tachycardia33 observed in both groups. Indeed, in people with septic shock, reversal of the transmitral flow pattern has been attributed to falling preload and did not carry a poor prognosis unless combined with a poor LV systolic function.22

On the contrary, a low Em (<0.177 m/s) predicted mortality with 100% sensitivity and 73% specificity in the present study, suggesting a more compromised relaxation in nonsurviving horses. Em, which reflects relaxation kinetics,33 is typically reduced in case of impaired LV relaxation associated with septic shock in people,41
and is considered to be preload independent in case of impaired relaxation. However, a mixed pattern combining impaired relaxation with restrictive ventricular filling was suspected based on a higher E/Em ratio in the nonsurviving group with a cut-off value of 2.67 offering the best sensitivity (100%) and specificity (83%) to predict mortality. By correcting the effect of LV relaxation on E velocity, the E/Em ratio is considered one of the best Doppler estimates of LV filling pressure. Despite being observed less frequently than impaired myocar- dial relaxation, this restrictive filling pattern has been described in human septic patients where E/Em recently proved to be an independent predictor of survival, even demonstrating a higher sensitivity (100%) and specificity (83%) than blood cardiac biomarkers or systolic function parameters. The origin of this sepsis-induced diastolic dysfunction is currently incompletely understood but has been associated with histological evidence of inflammatory infiltrates, interstitial edema, apoptosis and necrosis of the myocardium resulting in decreased compliance and subsequent LV pressure elevation.

The main limitation of this study was related to the use of echocardiography itself, which typically provides a noninvasive assessment of LV systolic and diastolic function but is highly dependent on loading conditions of the heart and HR, which are substantially altered by systemic inflammatory response syndrome. Least-squares-mean test was significant if P < 0.05 (ANOVA model including breed, body weight, age, sex, and outcome).

Table 4: Comparison of the means ± SD of echocardiographic parameters of diastolic function derived from pulse wave Doppler-mode and pulse wave tissue Doppler imaging mode in 12 surviving and 29 nonsurviving horses admitted for colic complicated by systemic inflammatory response syndrome. Least-squares-mean test was significant if P < 0.05 (ANOVA model including breed, body weight, age, sex, and outcome).

<table>
<thead>
<tr>
<th>Outcome</th>
<th>Survivors n = 12</th>
<th>Non survivors n = 29</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>E (m/s)</td>
<td>0.555 ± 0.13</td>
<td>0.571 ± 0.193</td>
<td>0.686</td>
</tr>
<tr>
<td>VTIa</td>
<td>9.4 ± 3.25</td>
<td>9.03 ± 4.94</td>
<td>0.9964</td>
</tr>
<tr>
<td>dE (ms)</td>
<td>259.92 ± 51.39</td>
<td>230.42 ± 87.35</td>
<td>0.5355</td>
</tr>
<tr>
<td>DTE (ms)</td>
<td>141.32 ± 48.02</td>
<td>122.02 ± 51.68</td>
<td>0.6982</td>
</tr>
<tr>
<td>Ddv/dtE (m/s²)</td>
<td>4.61 ± 1.94</td>
<td>5.52 ± 2.81</td>
<td>0.4166</td>
</tr>
<tr>
<td>A (m/s)</td>
<td>0.493 ± 0.14</td>
<td>0.579 ± 0.2</td>
<td>0.2397</td>
</tr>
<tr>
<td>VTIA</td>
<td>5.53 ± 1.82</td>
<td>6.46 ± 2.43</td>
<td>0.2464</td>
</tr>
<tr>
<td>dA (ms)</td>
<td>171.58 ± 31.78</td>
<td>174.65 ± 41.19</td>
<td>0.6146</td>
</tr>
<tr>
<td>E/A</td>
<td>1.149 ± 0.416</td>
<td>1.062 ± 0.373</td>
<td>0.4912</td>
</tr>
<tr>
<td>IVRT (ms)</td>
<td>83.61 ± 30.96</td>
<td>69.33 ± 26.68</td>
<td>0.1938</td>
</tr>
<tr>
<td>Em (m/s)</td>
<td>0.234 ± 0.034</td>
<td>0.135 ± 0.057</td>
<td>0.0010</td>
</tr>
<tr>
<td>E/Em</td>
<td>2.08 ± 0.42</td>
<td>5.11 ± 2.49</td>
<td>0.0051</td>
</tr>
</tbody>
</table>

*Data from pulsed-wave tissue Doppler mode were available for only 25 horses (7 surviving and 18 nonsurviving horses); A: peak late LV filling velocity; dA: late LV filling duration; Ddv/dtE: deceleration rate of early LV filling; dE: early LV filling duration; DTE: deceleration time of early LV filling; E: peak early LV filling velocity; E/A: peak early to late LV filling velocity ratio; Em: peak early diastolic myocardial velocity; E/Em: peak early LV filling velocity to peak early diastolic myocardial velocity ratio; IVRT: isovolumic relaxation time; LV: left ventricular; VTI: velocity time integral.

In the absence of a well-established definition in horses, the diagnosis of SIRS in this study was based on clinical and blood parameters mirroring the defining criteria of SIRS in human medicine associated with hyperlactatemia or low systolic arterial pressure as markers of hypoperfusion without attempting to identify endotoxins in the blood. Nevertheless, the detection of lipopolysaccharides in the blood suffered from low sensitivity (only 58.4%) and specificity (87.5%) to detect clinical endotoxemia and added little to the prognosis of equine colic compared to clinical parameters.

The underlying mechanisms of sepsis-induced myocardial failure are not well understood, but several pathways have been hypothesized in human patients. They include excess production of nitric oxide in cardiomyocytes with direct effect on contractility and via a cytopathic hypoxia as a consequence of depressed mitochondrial respiration and impaired oxygen utilization. This hypoxia could be worsened by an impairment of auto-regulating mechanisms of microcirculation, which generates a heterogenous myocardial oxygen supply. Furthermore, endotoxins disrupt calcium transfer at various sites, causing a decrease in calcium sensitivity and an unresponsiveness to β-adrenergic stimulation of the cardiomyocytes. Other factors such as autonomic nervous system dysfunction, myocardial inflammatory infiltration, or a tachycardia-induced cardiomyopathy may similarly contribute to myocardial depression. All or some of these factors could result in a decrease in myocardial contractility and compliance.

Therefore, cardiac performance may be altered by impairment of either diastolic function (ventricular filling defect secondary to combined low venous return and...
impaired relaxation or compliance) or systolic function (lack of contractility). Decreased cardiac performance with low EF and SV could be transiently compensated by sympathetic stimulation and tachycardia to maintain CO but may result in circulatory failure with a drop of CO (e.g., cardiogenic shock). In the present study, CI showed a high interindividual variability in nonsurviving horses, and this could be due to different hemodynamic profiles mirroring those observed in human patients who succumb from septic shock. This way, a cardiogenic shock with low CI was suspected in 3 horses (including the one spontaneous death) although a classic distributive shock characterized by refractory hypotension despite a preserved CI (with late appearance of MODS in some cases) was recognized in most horses subjected to euthanasia.

This myocardial dysfunction (systolic or diastolic) is known to be reversible within 7 to 10 days in human patients who survive septic shock. This phenomenon is sometimes considered a protective mechanism to reduce energy expenditure and oxygen demand during endotoxic shock. This “cardiac hibernation” presumably allows preservation of heart viability and integrity. Although an excellent recovery of cardiac performance has been observed in most surviving patients, some children suffered from long-term cardiac complications such as exercise-induced arrhythmia or decreased LV function after septic shock. The athletic prognosis and any delayed effects of SIRS on cardiac function are currently unknown in horses and warrants further investigation.

Conclusion

In the present study, certain echocardiographic markers of either systolic (PEP/ET > 0.26) or diastolic (E/Em>2.67) dysfunction predicted a poor outcome in colic horses with SIRS, suggesting that echocardiography may provide useful prognostic information in these horses as has been shown in human patients with septic shock. The clinical significance of the more marked myocardial depression occurring with SIRS in nonsurviving horses warrants further investigation. Early use of hemodynamic monitoring during the course of severe sepsis and septic shock with the implementation of early goal directed therapy has significantly lowered mortality in human ICUs. As already used in people, echocardiography could further be used in equine critical care to assess the response to fluid delivery and to guide early selection of appropriate support treatments (ie, fluids, positive inotropes, vasopressors).

Footnotes

4 Accutrend Roche Diagnostics, Mannheim, Germany
5 Medonic, CA 530 Oden, Sweden.

References