The prognostic value of left ventricular systolic function measured by tissue Doppler imaging in septic shock

Li Weng2†, Yong-tai Liu1†, Bin Du2, Jian-fang Zhou2, Xiao-xiao Guo1, Jin-min Peng2, Xiao-yun Hu2, Shu-yang Zhang1, Quan Fang1 and Wen-ling Zhu1

Abstract

Introduction: Left ventricular (LV) dysfunction is common in septic shock. Its association with the clinical outcome is still controversial. Tissue Doppler imaging (TDI) is a useful tool to quantify LV function; however, little knowledge is available about the prognostic value of these TDI variables in septic shock. Therefore, we performed this prospective study to determine the role of TDI variables in septic shock.

Methods: Patients with septic shock in a medical intensive care unit were studied with transthoracic echocardiography with TDI within 24 hours after the onset of septic shock. Baseline clinical, laboratory, and echocardiographic variables were prospectively collected. Independent predictors of 90-day mortality were analyzed with the Cox regression model.

Results: During a 20-month period, 61 patients were enrolled in the study. The 90-day mortality rate was 39%; the mean APACHE IV score was 84 (68 to 97). Compared with survivors, nonsurvivors exhibited significantly higher peak systolic velocity measured at the mitral annulus (Sa) (11.0 (9.1 to 12.5) versus 7.8 (5.5 to 9.0) cm/sec; P < 0.0001), lower PaO2/FiO2 (123 (83 to 187) versus 186 (142 to 269) mm Hg; P = 0.002), higher heart rate (120 (90 to 140) versus 103 (90 to 114) beats/min; P = 0.004), and a higher dose of norepinephrine (0.6 (0.2 to 1.0) versus 0.3 (0.2 to 0.5) μg/kg/min; P = 0.007). In the multivariate analysis, Sa > 9 cm/sec (hazard ratio (HR), 5.559; 95% confidence interval (CI), 2.160 to 14.305; P < 0.0001), dose of norepinephrine (HR, 1.964; 95% CI, 1.338 to 2.883; P = 0.001), and PaO2/FiO2 (HR, 0.992; 95% CI, 0.984 to 0.999; P = 0.031) remain independent predictors of 90-day mortality in septic-shock patients.

Conclusions: Our study demonstrated that LV systolic function as determined by TDI, in particular, Sa, might be associated with mortality in patients with septic shock.

Introduction

Although left ventricular (LV) depression in sepsis was first reported decades ago [1], it has not been well recognized until the recent widespread use of echocardiography in the intensive care unit (ICU) [2]. A variety of echocardiographic parameters have been developed to assess LV function [3]. Among these parameters, ejection fraction (EF) is most commonly used to evaluate LV systolic function, although studies exploring its association with clinical outcome have demonstrated conflicting results in high-risk patients, especially in patients with septic shock [2].

Tissue Doppler imaging (TDI) has been shown to be useful for quantifying global systolic and diastolic LV function [4-6]. The peak systolic velocity measured at the mitral annulus (Sa) reflects the long-axis systolic motion of the ventricle, whereas the early diastolic velocity of the mitral annulus (Ea) reflects the rate of myocardial relaxation. Both Sa and Ea have been demonstrated as useful tools to predict prognosis in a variety of cardiovascular diseases [7]. However, the
prognostic value of the TDI variables in septic shock requires further clarification.

Therefore, we performed a prospective, observational study to evaluate the prognostic significance of TDI variables in septic shock.

Materials and methods

Patients

The study was performed in a nine-bed medical ICU of a university teaching hospital. Between January 2010 and August 2011, all patients admitted for septic shock that developed within 24 hours before ICU admission were prospectively screened for eligibility. Sepsis, severe sepsis, and septic shock were defined according to consensus definition [8] (see Additional file 1 Figure S2), and the differentiation between infectious and noninfectious etiologies was made at the discretion of the ICU consultant. In patients with multiple episodes of septic shock, only the first episode was included in this study.

Exclusion criteria included age younger than 18 years; pregnancy; presence of moderate to severe valvular heart disease; patients or their relatives declined participation; suboptimal echocardiograms; postthoracic operation; documented myocardial infarction at any point in the medical history; and a decision of withdraw or withhold life-sustaining therapy.

Baseline clinical variables during the first 24 hours after admission (day 1) were collected prospectively, including age, gender, comorbidities, hemodynamic parameters, vasopressor or inotropic dose, Acute Physiology and Chronic Health Evaluation (APACHE) IV score [9], and Sequential Organ Failure Assessment (SOFA) score [10].

Echocardiographic examination

Two-dimensional conventional Doppler echocardiography and TDI studies were performed with commercially available equipment (Vivid I; GE Vingmed Ultrasound, Tirat Hacarmel, Israel). All studies were performed and reviewed by cardiologists with advanced training in echocardiography.

The transthoracic echocardiographic examination was performed within 24 hours after the onset of septic shock at the first day of ICU stay (day 1). LV end-diastolic volume (LVEDV), LV end-systolic volume (LVESV), and LV ejection fraction (LVEF) were assessed by using the modified biplane Simpson equation in the apical four- and two-chamber views, according to the American Society of Echocardiography Guidelines [11]. Mitral inflow was assessed with pulsed-wave Doppler echocardiography from the apical four-chamber view. The Doppler beam was aligned parallel to the direction of flow, and a 1- to 2-mm sample volume was placed between the tips of mitral leaflets during diastole [12]. From the mitral inflow profile, the E- and A-wave velocity and the E/A velocity ratio were measured. At least three consecutive beats were measured, and the average value was taken. In patients with tachycardia, the fused EA wave was considered an E wave to calculate the E/Ea.

TDI was performed at the apical four-chamber view for the long-axis motion of the heart [13,14]. Two-dimensional echocardiography with color-TDI imaging was performed. The imaging angle was adjusted to ensure a parallel alignment of the sampling window with the myocardial segment of interest. Gain settings, filters, pulse repetitive frequency, sector size, and depth were adjusted to optimize color saturation. The frame rate was adjusted to > 100. At least three consecutive beats were stored, and the images were digitized and analyzed off-line by EchoPac software (EchoPac 6.3.6; Vingmed-General Electric, Horten, Norway). Pulse-Doppler sample volume was placed at the septal and lateral MV annulus to obtain the average value of systolic (Sa) and early diastolic velocity (Ea) (see Additional file 1 Figure S2).

The intraobserver and interobserver variability in the measurements of Sa were 1.8% and 4.2%, respectively.

Follow-up

Follow-up was performed for 90 days after the onset of septic shock. The primary end point was 90-day all-cause mortality, defined as death within 90 days after onset of septic shock. Death was identified from hospital records or telephone interviews with relatives.

The Institutional Review Board of Peking Union Medical College Hospital approved this study protocol. Written informed consent was obtained from either the patients or their authorized relatives.

Statistical analysis

Deviations from a gaussian distribution were tested by the Kolmogorov-Smirnov test. Continuous variables were presented as median (25th to 75th percentiles). Categoric variables were expressed as percentages of the group from which they were derived. Continuous variables were compared with the use of the Student t test or Mann-Whitney test. Categoric variables were compared with the $\chi^2$ test or Fisher Exact test. Linear regression was used to investigate the correlation between EF and Sa. A receiver-operating characteristic (ROC) curve analysis was performed to determine the cutoff value of Sa for the prediction of 90-day mortality. The optimal cut-off value was defined as the point at which the value of “sensitivity + specificity - 1” was maximum (Youden index [15,16]).

A survival curve was performed by using the Kaplan-Meier method, and mortality rates were compared.
according to the cut-off value of Sa by using the log-rank test.

Cox proportional hazards regression model was used to estimate the risk of death by multivariate analysis (backward stepwise selection method with probability for the removal of 0.10) for the whole population. The multivariate analysis selection criterion from the univariate analysis was P value < 0.05 and absence of collinearity. Collinearity was defined as variance inflation factor (VIF) > 10 by using linear regression analysis.

All analysis was performed by using software (SPSS for Windows 11.5; SPSS, Chicago, IL, USA). Statistical significance was considered at P < 0.05.

Results
Baseline characteristics
During the study period, from January 2010 to August 2011, 132 patients with septic shock were eligible for assessment. Seventy-one were excluded in the final analysis, including consent refusal (n = 24), severe regurgitation (n = 10), postthoracic operation (n = 5), withheld or withdrawn therapy (n = 14), and suboptimal echocardiograms (n = 18). As a result, 61 patients were analyzed. Patients excluded had higher proportion of coronary heart disease. No significant difference was noted in other baseline characteristics between patients excluded and included (see Additional file 1 Table S1).

The 33 (54%) men had a median age of 68 (52 to 77) years, and an APACHE IV score of 84 (68 to 97). Twenty-two patients died during ICU stay, and 24 patients died at 90 days after the onset of septic shock, with a 90-day all-cause mortality of 39%. Five patients died within 48 hours of ICU admission. A total of 36 (59%) patients had documented comorbidities, including coronary heart disease (13%), hypertension (46%), and diabetes (28%). For all patients, no episode of active ischemia was documented in the last 3 months before inclusion. The most common infection was pneumonia (56%). All patients required vasoactive medications to maintain blood pressure, and all were mechanically ventilated because of acute lung injury. Table 1 compared baseline clinical variables on ICU admission (day 1) between survivors and nonsurvivors. Nonsurvivors had significantly lower PaO2/FiO2 (123 (83 to 187) versus 186 (142 to 269) mm Hg; P = 0.002) than did survivors.

Table 2 summarized hemodynamic and echocardiographic parameters on day 1. Heart rate was significantly higher in nonsurvivors (120 (90 to 140) versus 103 (90 to 114) beats/min; P = 0.004). Forty-nine (80%) patients received norepinephrine infusion at a median dose of 0.5 (0.2 to 0.7) μg/kg/min. The dose of norepinephrine was significantly higher in nonsurvivors (0.6 (0.2 to 1.0) versus 0.3 (0.2 to 0.5) μg/kg/min; P = 0.007). Twenty-seven (44%) patients were treated with dopamine, the median dose being 6.0 (5.0 to 10.0) μg/kg/min. During the study periods, one patient was treated with dobutamine (5 μg/kg/min), and another, with epinephrine (0.5 μg/kg/min).

Echocardiographic variables
Five patients were in atrial fibrillation at the time of echocardiography study. Ten of 61 patients had fused E/A wave, six in the group of survivors. Sa was significantly lower in the survivors group than in the nonsurvivors (7.8 (5.5 to 9.0) versus 11.0 (9.1 to 12.5) cm/sec; P < 0.0001), with a mean value of 9.0 (6.6 to 11.0) cm/sec for the whole cohort. Sixteen (27%) patients had an LVEF < 50%. LVEF values for survivors and nonsurvivors were 56% (36% to 65%) and 63% (52% to 66%), respectively. Other parameters, including those evaluating diastolic function, did not show any statistical difference between survivors and nonsurvivors (Table 2). A moderate correlation between LVEF and Sa was identified with linear regression (Figure 1).

The ability of Sa to predict 90-day mortality according to an ROC curve is shown in Figure 2, the area under the curve being 0.83. With a cut-off value of 9 cm/sec, the sensitivity and specificity of Sa to predict 90-day mortality was 75% and 86%, respectively. Patients with a higher Sa value (> 9 cm/sec) had a significantly higher mortality rate (75% versus 17%; P < 0.0001; log-rank = 24.03; P < 0.0001) (Table 2 and Figure 3).

Predictors of 90-day mortality in septic shock patients
In the final multivariate analysis, Sa > 9 cm/sec remained the strongest independent predictor of 90-day mortality in septic shock patients (HR, 5.559; 95% CI, 1.749 to 11.819; Wald, 2.160 to 14.305; Wald, 9.651; P = 0.002) (Table 3).

Discussion
The major finding of our study was that increased Sa is an independent predictor of 90-day mortality in patients with septic shock.

In the landmark study of Parker et al. [1], 10 of the 20 patients with septic shock exhibited global hypokinesia and ventricular dilation during the first 48 hours after admission. Contrary to common sense, the authors found significantly impaired LV systolic function in survivors compared with nonsurvivors. Subsequent studies [17,18] demonstrated similar reversible global hypokinesia by echocardiography. Vieillard-Baron and colleagues...
observed global hypokinesia in 26 of 67 patients. Moreover, LVEF was compromised in survivors during the first 24 hours (49% ± 18% versus 55% ± 15%) [17]. In another study performed over a period of 5 years [18], survivors showed evidence of septic myocardial dysfunction, as suggested by compromised LVEF (43.9% ± 16.4% versus 52.0% ± 14.0%) and higher LVEDV (75.3 ± 20.1 ml/m² versus 64.9 ± 25.0 ml/m²; P < 0.05). However, similar to our study, the difference in LVEF between survivors and nonsurvivors was not significantly different. The linear correlation between Sa and ejection in our study was similar to that of a previous study [5]. Unlike LVEF, Sa was a sensitive marker of LV systolic function in patients with cardiovascular disease, which showed that Sa could predict clinical outcome in a more sensitive manner than could the LVEF [7]. In a study of hypertrophic cardiomyopathy, TDI revealed myocardial contractive abnormalities before any clinical presentations [19].

| Table 1 Baseline characteristics and comparison between survivors and nonsurvivors at the onset of septic shock (day 1) |
|---------------------------------------------------------------|
| Characteristics                                             |
| Survivors (n = 37)                                           |
| Nonsurvivors (n = 24)                                        |
| P value                                                      |
| Age, years                                                  | 68 (49-76) | 74 (61-82) | 0.167 |
| Male, n (%)                                                 | 18 (49)    | 15 (63)    | 0.289 |
| BMI, kg/m²                                                  | 23 (21-25) | 23 (20-26) | 0.732 |
| APACHE IV score                                             | 79 (66-94) | 93 (69-99) | 0.339 |
| APACHE IV predicted mortality, %                            | 28 (17-53) | 48 (35-61) | 0.039 |
| SOFA score                                                  | 10 (8-12)  | 10 (8-12)  | 0.222 |
| PaO₂/FIO₂, mm Hg                                            | 186 (142-269) | 123 (83-187) | 0.002 |
| SOFA cardiovascular score                                   | 4 (4-4)    | 4 (4-4)    | 0.911 |
| Days on vasoactive medications                              | 5 (3-9)    | 7 (4-9)    | 0.340 |
| ICU LOS, days                                               | 12 (8-22)  | 12 (4-20)  | 0.515 |
| Hospital LOS, days                                          | 29 (17-49) | 17 (6-52)  | 0.150 |
| Comorbidities                                               |
| Coronary heart disease, n (%)                                | 6 (16)     | 2 (8)      | 0.373 |
| Hypertension, n (%)                                          | 18 (49)    | 10 (42)    | 0.593 |
| Diabetes, n (%)                                              | 12 (32)    | 5 (21)     | 0.324 |
| Chronic renal failure, n (%)                                 | 6 (16)     | 2 (8)      | 0.373 |
| Primary diagnosis of infection                               |
| Pneumonia, n (%)                                             | 18 (49)    | 16 (67)    | 0.166 |
| Bacteremia, n (%)                                            | 5 (14)     | 2 (8)      | 0.355 |
| Peritonitis, n (%)                                           | 5 (13)     | 1 (4)      | 0.231 |
| Others, n (%)                                                | 9 (24)     | 5 (21)     | 0.751 |
| Laboratory data                                             |
| Lactate, mM                                                  | 1.75 (1.30-2.88) | 2.00 (1.80-3.98) | 0.095 |
| WBC, x10⁹/L                                                  | 11.36 (7.18-19.99) | 12.07 (6.91-18.90) | 0.623 |
| Procalcitonin, ng/ml                                         | 2.01 (0.59-7.40) | 1.50 (0.53-5.71) | 0.435 |
| cTnI, μg/L                                                   | 0.17 (0.06-1.14) | 0.15 (0.04-0.93) | 0.952 |
| CKMB, μg/L                                                   | 1.80 (0.60-3.83) | 1.65 (1.03-4.90) | 0.455 |
| NTproBNP, pg/ml                                              | 4,072.00 (2,006.50-11,885.50) | 3,710.00 (1,361.50-10,618.25) | 0.693 |

APACHE, Acute Physiology And Chronic Health Evaluation; BMI, body mass index; Hospital LOS, hospital length of stay; SOFA, Sequential Organ Failure Assessment.
preserve cardiac myocytes by downregulation of oxygen consumption and energy requirements. It is an adaptive response to maintain myocardial viability for prevention of cell-death pathway activation and to aid the future full recovery. The slightly increased cardiac biomarkers (that is, cTnI) in the study population also support that physical myocardial injury is negligible. Instead, the heart was injured “functionally.” Such a potential beneficial response must be based on an assumption that tissue perfusion might be maintained with the depressed heart. With a close look at our data, lactate, a good marker of tissue perfusion [24], was not elevated, despite myocardial depression. However, serial echocardiograms

### Table 2 Baseline hemodynamic and echocardiographic data of survivors and nonsurvivors at the onset of septic shock (day 1)

|                     | Survivors (n = 37) | Non-survivors (n = 24) | P value |
|---------------------|--------------------|------------------------|---------|
| **Hemodynamic parameters** |                    |                        |         |
| Heart rate, beats/min | 103 (90-114)       | 120 (90-140)           | 0.004   |
| Mean arterial pressure, mm Hg | 76 (74-83)         | 74 (70-82)             | 0.189   |
| Central venous pressure, mm Hg | 16 (12-17)        | 14 (11-19)             | 0.911   |
| Dopamine, n (%) | 17 (46)            | 10 (42)                |         |
| Dose, μg/kg/min | 5.0 (4.0-10.0)     | 6.0 (4.5-10.0)         | 0.836   |
| Norepinephrine, n (%) | 30 (81)           | 19 (79)                |         |
| Dose, μg/kg/min | 0.3 (0.2-0.5)      | 0.6 (0.2-1.0)          | 0.007   |
| Balance on day 0, ml/24 hours | 1,180 (445-2,140) | 1,850 (15-3,004)       | 0.640   |
| **Echocardiographic data** |                    |                        |         |
| **Systolic parameters** |                    |                        |         |
| LVEDV, ml | 72 (54-98)         | 63 (56-78)             | 0.110   |
| LVESV, ml | 30 (20-52)         | 26 (19-34)             | 0.169   |
| LVEF biplane, % | 56 (36-65)        | 63 (52-66)             | 0.111   |
| LVEF biplane < 50%, n (%) | 12 (33)          | 4 (17)                 | 0.234   |
| Sa, cm/sec | 7.8 (5.5-9.0)      | 11.0 (9.1-12.5)        | < 0.0001|
| Sa > 9 cm/s, n (%) | 6 (17)            | 18 (75)                | < 0.0001|
| **Diastolic parameters** |                    |                        |         |
| E/A | 0.9 (0.7-1.4)      | 0.7 (0.6-1.2)          | 0.171   |
| Ea, cm/sec | 8.3 (5.8-10.0)    | 7.0 (6.0-11.0)         | 0.634   |
| Ea < 8 cm/sec, n (%) | 15 (42)           | 12 (52)                | 0.429   |
| E/Ea | 11.1 (8.5-14.6)   | 11.1 (6.6-14.1)        | 0.206   |

E, peak velocity of early diastolic transmitral flow; Ea, early diastolic velocity of the mitral annulus; E/A, the ratio of mitral valve peak E-wave and peak A-wave velocity; EDT, deceleration time of mitral E wave; LVEDV, left ventricular end-diastolic volume; LVEF, left ventricle ejection fraction; LVESV, left ventricular end-systolic volume; Sa, peak systolic velocity measured at mitral annulus.

---

**Figure 1** The line regression between left ventricular ejection fraction (LVEF) and mitral annulus (Sa).

**Figure 2** Receiver-operating characteristic (ROC) curve for predicting 90-day mortality by using the peak systolic velocity measured at the mitral annulus (Sa). Area under the curve is 0.83.

**Figure 3** The 90-day mortality in the study population classified according to the peak systolic velocity measured at mitral annulus (Sa) < 9 cm/sec or Sa > 9 cm/sec.
were performed for only some of our patients; further 
serial study was warranted to support this hypothesis.
Persistent vasoplegia might be another explanation for 
our finding. Although the Sa has advantages over previ-
ously used measures of LV systolic function, such as 
LVEF, it still is load dependent, afterload especially [25,26]. In the study of Robotham et al. [26], the same 
level of LVEF may correspond to very different level of 
intrinsic LV contractility. For instance, an LVEF of 55% 
may correspond to severe impressed intrinsic LV con-
tractility in the presence of decreased vascular tone. It 
would not be surprising to find relatively normal or 
supernormal Sa in nonsurvivors in our study, which 
reflected a hyperkinetic state associated with persistent 
and profound vasoplegia that, in turn, could be a marker 
of sustained cytokine release. This kind of persistent 
vasoplegia was associated with a high mortality rate, 
which is consistent with our findings.

Sturgess et al. [27] also reported the role of TDI to 
assess LV function in septic shock patients. They failed 
to find any difference in LV systolic function between 
survivors and nonsurvivors. This might be explained by 
the small sample size (n = 21) and high prevalence of 
cardiac diseases (43%) in the study population. Myocar-
dial infarction may influence systolic and diastolic TDI 
values, as previously described by Alam et al. [28]. We 
also included patients with coronary heart disease in our 
study, but patients with myocardial infarction were 
excluded. Furthermore, after the exclusion of patients 
with coronary heart disease, the predictive value of Sa 
still remained. Similar to the study of Sturgess [27], 
Landesberg et al. [29] did not find evidence of LV systo-
lic dysfunction in survivors. However, only 62% in the 
study population had septic shock. This, in addition to 
the imbalanced distribution of septic shock between sur-
vivors and nonsurvivors (57% versus 72%; P = 0.012), 
precluded direct comparison between their study result 
and that of our study.

In our cohort of patients, MAP was maintained at a 
higher level than that recommended by the guideline 
[30]. This can be explained by the high prevalence of 
hypertension in the study population. Although no 

evidence suggested the benefit of hyperdynamic support, 
Dünser et al. [31] reported that the time spent below the 
MAP of 75 mm Hg would increase the risk of sub-
sequent renal-replacement therapy. Such findings sug-

gest the importance of addressing ischemic acute renal 
failure in the absence of frank hypotension [32]. Accord-
ingly, we try to individualize the target of blood pressure 
in our patients, based on usual level. The study of Vieil-
lard-Baron et al. [17] suggested that the increased nore-
pinephrine loads necessary to maintain high blood 
pressures were likely to cause LV hypokinesia. However, 
the dose of norepinephrine to maintain blood pressure in 
our groups was lower in survivors (0.3 (0.2 to 0.5) 
versus 0.6 (0.2 to 1.0) μg/kg/min), who had a higher inci-
dence of LV hypokinesia. In the final multivariate 
analysis, even after adjustment for norepinephrine treat-
ment, the prognostic value of Sa still attained statistical 
significance.

Limitations
First, the sample size in our study was relatively small, 
but the robust association between Sa and mortality rate 
suggests that this was not just an accidental finding. 
Possible selection biases might exist because more 
patients were excluded than were studied. However, the 
baseline characteristics, except for coronary heart dis-
ease, did not show significant differences between the 
patients excluded and included. Second, this study is a 
single-center study. Our local management strategy may 
influence the patient’s outcome, which might preclude 
the generalization of the study findings. Third, the 
potential confounding factors for TDI variables were not 
explored in this study. A more-detailed study focusing 
on these confounding factors is highly desirable. Fourth, 
correlation between TDI variables and blood flow-
derived parameters was not performed because pulmon-
ary artery catheters were inserted in only half of the 
study population.

Conclusion
Our study demonstrated that LV systolic function, as 
determined by TDI, in particular, by Sa, might be asso-
ciated with mortality in patients with septic shock. Con-
cerning the limitations as discussed earlier, further 

studies are warranted to confirm our findings.

Key messages
- In patients with septic shock, compared with non-
survivors, survivors exhibited more marked myocar-
dial depression.
- Evaluation of LV function by TDI, in particular, by 
Sa, might be associated with mortality in patients 
with septic shock.
Additional material

Additional file 1: Supplement. Supplement to Methods Results.

Abbreviations
APACHE: Acute Physiology And Chronic Health Evaluation; Ea: peak velocity of early diastolic transmural flow; EF: ejection fraction; ICU: intensive care unit; LV: left ventricular; LVEDV: LV end-diastolic volume; LVESV: LV end-systolic volume; PCWP: pulmonary capillary wedge pressure; ROC: receiver-operating characteristic; SIRS: systemic inflammatory response syndrome; SOFA: Sequential Organ Failure Assessment; TDI: tissue Doppler imaging; Sa: peak systolic velocity measured at the mitral annulus.

Author details
1Department of Cardiology, Peking Union Medical College Hospital, Peking Union Medical College and Chinese Academy of Medical Sciences, 1 Shuaifuyuan, Dongcheng district, Beijing, 100730, China. 2Medical ICU, Peking Union Medical College Hospital, Peking Union Medical College and Chinese Academy of Medical Sciences, 1 Shuaifuyuan, Dongcheng District, Beijing, 100730, China.

Authors’ contributions
YL, LW, BD, JZ, XG, JP, XH, SZ, QF, and WZ participated in the design of the study and performed the statistical analysis. YL, LW, and BD conceived of the study and participated in its design and coordination. All authors read and approved the final manuscript.

Competing interests
The authors declare that they have no competing interests. All authors report no funding for support of this work.

Received: 22 December 2011 Revised: 29 February 2012 Accepted: 3 May 2012 Published: 3 May 2012

References
1. Parker MM, Shelhamer JH, Bacharach SL, Green MV, Natanson C, Frederick TM, Damksie BA, Parrillo JE. Proven but reversible myocardial depression in patients with septic shock. Ann Intern Med 1984, 100:483-490.
2. Hunter JD, Dodd M. Sepsis and the heart. Br J Anaesth 2010, 104:3-11.
3. Dittoe N, Stultz D, Schwartz BP, Hahn HS. Evidence of myocardial hibernation in the septic heart. J Am Coll Cardiol 2002, 39:254-259.
4. Sibbald WJ: How to diagnose diastolic heart failure: a practical approach. Can J Cardiol 2011, 27:51-57.
5. Frasier AG, Brutsaert DL: Systolic and diastolic factors in the heart. J Cardiovasc Magn Reson 2010, 12:85-97.
6. Weng et al. Critical Care Medicine. 2012, 16(3):R71
7. Yu C, Sanderson J, Marwick TH, Oh J. Tissue Doppler imaging: a new prognosticator for cardiovascular diseases. J Am Coll Cardiol 2007, 49:1903-1914.
8. Bone RC, Balk RA, Cerra FB, Dellinger RP, Fein AM, Knusso WA, Schein RM, Sibbald WH. Definitions for sepsis and organ failure and guidelines for the use of innovative therapies in sepsis: The ACCP/SCCM Consensus Conference Committee, American College of Chest Physicians/Society of Critical Care Medicine. Chest 1992, 101:1644-1655.
9. Zimmerman JE, Kramer AA, McNair DS, Malila FM. Acute Physiology and Chronic Health Evaluation (APACHE): IV: hospital mortality assessment for today’s critically ill patients. Crit Care Med 2006, 34:1297-1310.
10. Vincent JL, Moreno R, Takala J, Willatts S, de Mendonça A, Bruining H, Reinhart CK, Suter PM, Thijs LG. The SOFA (Sepsis-related Organ Failure Assessment) score to describe organ dysfunction/failure, on behalf of the Working Group on Sepsis-related Problems of the European Society for Intensive Care Medicine. Intensive Care Med 1992, 22:707-710.
11. Lang RM, Biering M, Devereux RB, Flachskampf FA, Foster E, Pellika T, Picard M, Roman M, Seward J, Shawere Jos, Solomon SD, Spencer RT, Sutton MS, Stewart WJ, Chamber Quantification Writing Group, American Society of Echocardiography’s Guidelines and Standards Committee, European Association of Echocardiography. Recommendations for chamber quantification: a report from the American Society of Echocardiography’s Guidelines and Standards Committee and the Chamber Quantification Writing Group, developed in conjunction with the European Association of Echocardiography, a branch of the European Society of Cardiology. J Am Soc Echocardiogr 2005, 18:1440-1463.
12. Quinones MA, Otto CM, Stoddard M, Waggoner A, Zoghbi WA. Tissue Doppler imaging DQTFotNaSCotASo: Recommendations for quantification of Doppler echocardiography: a report from the Doppler Quantification Task Force of the Nomenclature and Standards Committee of the American Society of Echocardiography. J Am Soc Echocardiogr 2002, 15:167-184.
13. Yu CM, Wang Q, Lau CP, Tse HF, Leung SK, Lee KL, Tsang V, Ayers G. Reversible impairment of left and right ventricular systolic and diastolic function during short-lasting atrial fibrillation in patients with an implantable atrial defibrillator: a tissue Doppler imaging study. Pacing Clin Electrophysiol 2001, 24:979-988.
14. Pai RG, Gill KS. Amplitudes, durations, and timings of apically directed left ventricular myocardial velocities: IL Systolic and diastolic asynchrony in patients with left ventricular hypertrophy. J Am Soc Echocardiogr 1998, 11:112-118.
15. Fluss R, Faraggi D, Reiser B. Estimation of the Youden Index and its associated cutoff point. Biometrics 2005, 47:438-472.
16. Schisterman EF, Perkins NJ, Liu A, Bondell H. Optimal cut-point and its corresponding Youden Index to discriminate individuals using pooled blood samples. Epidemiology 2005, 16:73-81.
17. Veillard-Baron A, Caille V, Charron C, Bellard G, Page B, Jardin F. Actual incidence of global left ventricular hypokinesia in adult septic shock. Crit Care Med 2008, 36:1707-1706.
18. Jardin F, Fourme T, Page B, Loubieres Y, Veillard-Baron A, Beauchet A, Bourdais JP. Persistent preload defect in severe sepsis despite fluid loading: a longitudinal echocardiographic study in patients with septic shock. Chest 1999, 116:1354-1359.
19. Naghue SF, Bachinski LL, Meyer D, Hill R, Zoghbi WA, Tam JW. Quinones MA, Roberts R, Marian AJ. Tissue Doppler imaging consistently detects myocardial abnormalities in patients with hypertrophic cardiomyopathy and provides a novel means for an early diagnosis before and independently of hypertrophy. Circulation 2001, 104:128-130.
20. Chahal NS, Lim TK, Jain P, Chambers JC, Kooner JS, Senior R. Normative reference values for the tissue Doppler imaging parameters of left ventricular function: a population-based study, Eur J Echocardiogr 2010, 11:51-56.
21. Wang M, Yip GW, Wang SY, Zhang Y, Ho PY, Tse MK, Lam PK, Sanderson JE. Peak early diastolic mitral annulus velocity by tissue Doppler imaging adds independent and incremental prognostic value. J Am Coll Cardiol 2003, 41:820-825.
22. Veillard-Baron A, Septic cardiomyopathy. Ann Intensive Care 2011, 1:6.
23. Levy RJ, Piel DA, Acton PD, Zhou R, Ferrari VA, Deutschman CS. Evidence of myocardial hibernation in the septic heart. Crit Care Med 2005, 33:2725-2756.
24. Nguyen HB, Corbett SW, Steele R, Banta J, Clark RT, Hayes SR, Edwards J, Cho TW, Wittlake WA. Implementation of a bundle of quality indicators for the early management of severe sepsis and septic shock is associated with decreased mortality. Crit Care Med 2007, 35:1105-1112.
25. Uemura K, Kawada T, Sunagawa K, Sugimachi M. Peak systolic mitral annulus velocity reflects the status of ventricular-arterial coupling-theoretical and experimental analyses. J Am Soc Echocardiogr 2011, 24:582-591.
26. Robotham JL, Takata M, Berman M, Harasawa Y: Ejection fraction revisited. Anesthesiology 1991, 74:172-183.

27. Sturgess DJ, Marwick TH, Joyce C, Jenkins C, Jones M, Masci P, Stewart D, Venkatesh B: Prediction of hospital outcome in septic shock: a prospective comparison of tissue Doppler and cardiac biomarkers. Crit Care 2010, 14:R64.

28. Alam M, Wardell J, Andersson E, Samad BA, Nordlander R: Effects of first myocardial infarction on left ventricular systolic and diastolic function with the use of mitral annular velocity determined by pulsed wave Doppler tissue imaging. J Am Soc Echocardiogr 2000, 13:343-352.

29. Landesberg G, Glon D, Meroz Y, Georgieva M, Levin PD, Goodman S, Avidan A, Beeri R, Weissman C, Jaffe AS, Sprung CL: Diastolic dysfunction and mortality in severe sepsis and septic shock. Eur Heart J 2011, 33:895-903.

30. Dellinger R, Carlet J, Masur H, Gerlach H, Calandra T, Cohen J, Gaa-Banacloche J, Keh D, Marshall JC, Ramsay G, Zimmerman JL, Levy MM, Surviving Sepsis Campaign Management Guidelines Committee: Surviving Sepsis Campaign guidelines for management of severe sepsis and septic shock. Crit Care Med 2004, 32:858-873.

31. Dünser MW, Takala J, Ulmer H, Mayr VD, Luckner G, Rochberger S, Daudel F, Lepper P, Hasibeder WR, Jakob SM: Arterial blood pressure during early sepsis and outcome. Intensive Care Med 2009, 35:1225-1232.

32. Abuelo JG: Normotensive ischemic acute renal failure. N Engl J Med 2007, 357:797-805.

doi:10.1186/cc11328
Cite this article as: Weng et al.: The prognostic value of left ventricular systolic function measured by tissue Doppler imaging in septic shock. Critical Care 2012, 16:R71.