

Speckle-tracking analysis of left ventricular systolic function in the intensive care unit

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Abstract

Speckle-tracking analysis is a new available tool in order to assess left ventricular function in cardiology. Its novelty relies on the technological ability to track natural acoustic markers (known as speckle) within the myocardium during the cardiac cycle. This technology allows the evaluation of myocardium strain during systole and diastole. To date, global longitudinal strain (GLS) has been extensively studied in cardiology. It is now well established that GLS is more sensitive than left ventricular ejection fraction with 2D echocardiography in detecting systolic function impairment. It is also superior to left ventricular ejection fraction in the prediction of major cardio-vascular events. In the intensive care unit (ICU) setting, data are scarce. In experimental model and human studies in septic shock, speckle-tracking analysis suggests that GSL is impaired along with preserved left ventricular ejection fraction. Recent data also suggest that GLS impairment could predict in-ICU mortality in septic shock. In severe subarachnoid haemorrhage patients, speckle-tracking analysis could be more sensitive in detecting stress cardiomyopathy. However, there are many gaps to fill in the critically ill patient. For instance, the influence of mechanical ventilation on GLS is not fully elucidated, and there are, to date, too few data to exactly assess potential GLS alterations on the patient's outcome. Nonetheless, this new tool provides objective and sensitive data with acceptable intra and inter-observer variability and may be of primary interest in the evaluation of left-ventricular systolic function in the ICU.

Key words: global longitudinal strain, shock, intensive care, mortality, speckle tracking echocardiography

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Left ventricular (LV) contraction during systole is a complex phenomenon. Owing to the myocardial muscular fibres' orientation, it is a combination of longitudinal, circumferential and radial contraction (Fig. 1). A 15% shortening of longitudinal myocardial fibres leads to a 40% radial LV thickening and to a 60% modification of LV ejection fraction (LVEF) in a normal heart [1]. Myocardial contraction in the circumferential-longitudinal plane, results in twist or torsional LV deformation during systole. Altogether, these specific dynamic modifications play a key role for the generation of blood suction in the LV chamber during diastole, as well as ejection fraction during systole. Thus, myocardial deformation is of major functional interest during the cardiac cycle [2].

Several echographic echocardiographic techniques have been developed in order to study myocardial deformations and strain. Although tissue Doppler-derived strain imaging was introduced several years ago, this technology suffers from few drawbacks, such as angle-dependency, noise interference, as well as intraobserver and interobserver variability [4]. Speckle-tracking echocardiography (STE) has more recently emerged as an alternative by studying natural acoustic markers within an ultrasonographic window of the LV. The image algorithm tracks these markers in several blocks of regions of interest of approximately 20 to 40 pixels. Speckles are tracked consecutively using a sum-of-absolute differences algorithm to resolve angle-independent sequences of tissue deformation or motion.

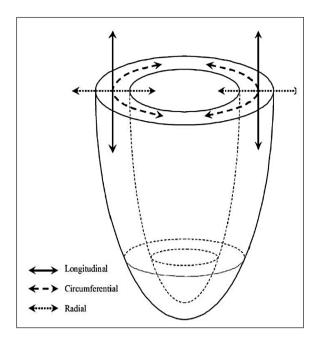


Fig. 1. Example of local heart coordinate system illustrating the 3 orthogonal axes: longitudinal, circumferential, radial [3]

Strain is therefore a dimensionless index and represents a fractional change in one dimension, typically length, and can be mathematically written as: $(L-L_0)/L_0$. In this formula, L is the myocardium length after deformation and L_0 is the original length. By convention, when the length is increasing, strain is expressed positively, while shortening is expressed negatively. When acquired at the apex with echo-

cardiography, normal LV myocardium strain has a negative value during systole and a positive value during diastole, in the longitudinal direction. Other values can be retrieved with speckle-tracking acquisition such as strain rate, which is the rate of deformation. Mathematically, it is expressed as the change in velocity between two points divided by the distance between 2 points. The strain rate has a negative value during systole and a positive value during diastole, when acquired at the apex of a normal LV [3].

STE allows a frame-by-frame tracking of natural acoustic markers within the myocardium with standard echocardiography. Unlike Doppler-Tissue Imaging, STE is not a Doppler-based tracking method and is not angle-dependant. However, similar to Doppler-Tissue Imaging, a high frame rate is necessary to achieve STE (average 80–90 Hz), while high quality echocardiography is required to allow proper tracking of the myocardium during the cardiac cycle [3] (Fig. 2). Eventually, STE provides regional strain data in each LV segments (Fig. 3).

CLINICAL APPLICATIONS OF STE FOR MYOCARDIAL DEFORMATION ASSESSMENT

Longitudinal LV mechanics are predominantly governed by the sub-endocardial area, which is the most vulnerable component of LV mechanics and the most vulnerable area to myocardial disease. The midmyocardial and epicardial function may remain initially unaffected; therefore circumferential strain and twist may remain normal or even supranormal in order to preserve LV systolic performance [2]. The detection of early altered longitudinal

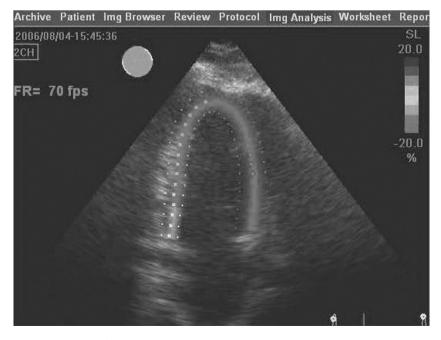


Fig. 2. Example of speckle-tracking analysis of a normal LV

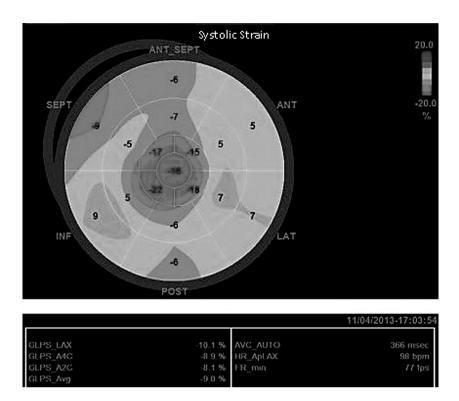


Fig. 3. Example of regional longitudinal strain and longitudinal peak strain. Basal and medio-ventricular strain are severely impaired. Apical segments display a preserved strain in this patient victim of a severe subarachnoid haemorrhage (personal data)

mechanics is, therefore, an early detector of myocardial disease and systolic function impairment. Thus, longitudinal strain has been mostly studied in cardiology in ischemic cardiomyopathy, valvular diseases, chronic hypertension, amyloidosis etc.

During aortic stenosis, LV progressively thickens, due to the increased afterload while LVEF remains preserved. STE showed that although longitudinal strain is impaired during aortic stenosis [5], it improves after aortic valve replacement [6]. In a cohort of 65 patients with aortic stenosis, longitudinal strain impairment was significantly associated with poorer exercise response and more cardiac events during a 12-month follow-up, in spite of a preserved LVEF [7]. During aortic insufficiency, strain rates are decreased and correlated to LV end-systolic and end-diastolic volumes [8]. After myocardial infarction, longitudinal strain is reduced in patients in correlation with the area of infarction and ejection fraction [9–11].

During stress cardiomyopathy, STE has helped in describing an LV failure pattern. Some authors have found the presence of hyperkinesis in the basal segments [12], attenuation of the longitudinal strain in the medio-ventricular segments [13] or dyskinesis in the apex [12]. Strain abnormalities in stress cardiomyopathy do not follow a unique coronary territory, which is helpful in distinguishing them from acute coronary syndromes. During stress cardiomyopathy, strain specifically improves over time [13–15].

In summary, strain provides robust, operator-independent data on LV systolic function, which can be more sensitive in detecting systolic function impairment than classical 2-dimension echocardiography. In a recent meta-analysis including 16 studies and 5,721 patients [16], global longitudinal strain impairment was more robust than LVEF in the prediction of death or major cardio-vascular events in cardiology, underlining the importance that STE will take in the next years.

SPECKLE-TRACKING IN THE INTENSIVE CARE UNIT

Owing to the growing interest of STE in cardiology, its superiority in the prediction of cardio-vascular events compared to standard LVEF, STE has been recently introduced in the Intensive Care Unit (ICU). In a recent experimental study, Hestenes *et al.* [17] studied strain in a porcine model of septic shock. They demonstrated that LV longitudinal strain was impaired along with preserved LVEF and cardiac output but that there was a decrease in the index of regional work. The first ICU-description of STE was in a paediatric population of septic shock [18]. The authors found that LV longitudinal strain and the strain rate were impaired in septic children compared with controls. At the same time, LVEF was preserved in both the septic children and controls. In adult septic shock, several authors found similar data. In a mono-centric study of 106 adult patients with septic shock,

the authors found a severe longitudinal strain impairment along with preserved LVEF [19]. In 60 patients with septic shock, the authors found systolic function impairment in 69% of patients assessed with STE, whereas only 33% patients displayed LVEF alteration [20]. In another cohort of 50 patients with septic shock, the authors also described a preserved LVEF when LV global longitudinal peak strain was altered [21]. These findings were confirmed in 35 patients with septic shock [22]. Longitudinal strain worsens in the first hours following septic shock, when adjusting for the level of vasopressors and LV end-diastolic volume. However, in this study [22], 15 patients with severe sepsis did not display the same longitudinal strain alterations, suggesting a specific signature in septic shock. In another mono-centric study in patients with septic shock [23], the authors found a correlation between strain impairment and the level of hyperlactatemia and low central venous oxygen saturation. LV strain could, therefore, be used in septic shock as a surrogate marker of oxygenation deliverance inadequacy. To the best of our knowledge, one study pointed out that right ventricle strain impairment was associated with long-term mortality in septic shock [20] and a large mono-centric study of 111 patients pointed out that a GLS impairment $\geq -13\%$ was the best marker of in-ICU mortality [24]. Our group published a mono-centric observational study about LV systolic function assessment with STE in patients with severe subarachnoid haemorrhage [25]. In this 46-patient cohort, we also found that while global longitudinal strain was altered, LVEF was preserved. All these data advocate the superiority of strain on classical LVEF in detecting mild systolic impairment.

LV SYSTOLIC FUNCTION ASSESSED WITH STE IN PATIENTS UNDERGOING MECHANICAL VENTILATION

There are currently no data in the literature regarding LV systolic assessment with STE in healthy patients undergoing mechanical ventilation. In a mono-centric study of 20 patients in the general ICU setting [26], the authors found that a PEEP elevation from 5 to 15 cm H₂0 would not modify LV longitudinal strain but would alter left peak-atrial longitudinal strain. It is, therefore, difficult to evaluate the accurate effects of mechanical ventilation on strain parameters. Until now, authors have extrapolated from data in the cardiologic literature in an attempt to set a threshold for normal GLS values in ICU ventilated patients. In a large Norwegian multi-centric nation-wide study [27], the authors studied 1,266 healthy subjects. Although global longitudinal strain increases with age (implying strain impairment), the overall normal global longitudinal strain was -17.4%. In another multi-centric study of 250 volunteers without cardio-vascular diseases, the average global longitudinal strain was –18.6% [28]. However, numerous factors are susceptible to modify a patient's global longitudinal strain. Chronic hypertension can alter longitudinal strain. In a cohort of 56 patients, longitudinal strain was decreased along with increased LV torsion. The authors also found a correlation with serum levels of aminoterminal propeptide of procollagen I/III and tissue inhibitor of matrix metalloproteinase, suggesting a modification of the myocardial collagen turnover and fibrotic process which could alter LV systolic function [29].

WHAT COULD STE ADD IN THE ICU FIELD?

One of the major drawbacks of classical 2D echocardiography parameters the ICU setting is their high inter-observer variability. In a cohort of 50 patients undergoing mechanical ventilation with shock, LVEF evaluation with the Simpson technique displayed the worst intra and inter-observer variability [30]. STE is an operator-independent technique with low inter-observer variability (from 2 [31] to 13% [32]) regarding strain data but higher variability regarding strain rate [28]. In patients with mechanical ventilation, we found an inter-observer variability of 8% [25] while others [20, 24] have also found a good inter-observer agreement with a Bland-Altman analysis. STE could, therefore, provide objective data on systolic function with an acceptable variability.

CONCLUSION

STE is more sensitive in detecting mild systolic function impairment than classical LVEF. It is well-established in cardiology that STE is superior to standard echocardiography in order to predict major cardio-vascular events. However, there are, to date, too few data in the ICU to delineate the potential strain alterations on patient's outcome. However, further studies should definitely provide substantial data regarding cardio-vascular events in specific settings, such as septic shock or neuro-ICU.

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References:

- Covell JW: Tissue structure and ventricular wall mechanics. Circulation 2008; 118: 699–701. doi: 10.1161/CIRCULATIONAHA.108.797399.
- Geyer H, Caracciolo G, Abe H et al.: Assessment of myocardial mechanics using speckle tracking echocardiography: fundamentals and clinical applications. J Am Soc Echocardiogr 2010; 23: 351–369. doi: 10.1016/j. echo.2010.02.015.
- Leung DY, Ng ACT: Emerging clinical role of strain imaging in echocardiography. Heart Lung Circ 2010; 19:161–174. doi: 10.1016/j. hlc.2009.11.006.
- Castro PL, Greenberg NL, Drinko J, Garcia MJ, Thomas JD: Potential pitfalls of strain rate imaging: angle dependency. Biomed Sci Instrum 2000; 36:197–202.
- Dal-Bianco JP, Khandheria BK, Mookadam F, Gentile F, Sengupta PP: Management of Asymptomatic Severe Aortic Stenosis. J Am Coll Cardiol 2008; 52:1279–1292. doi: 10.1016/j.jacc.2008.07.020.

- Becker M, Kramann R, Dohmen G et al.: Impact of left ventricular loading conditions on myocardial deformation parameters: analysis of early and late changes of myocardial deformation parameters after aortic valve replacement. J Am Soc Echocardiogr 2007; 20: 681–689.
- Lafitte S, Perlant M, Reant P et al.: Impact of impaired myocardial deformations on exercise tolerance and prognosis in patients with asymptomatic aortic stenosis. Eur J Echocardiogr 2009; 10: 414–419. doi: 10.1093/ejechocard/jen299.
- Marciniak A, Sutherland GR, Marciniak M et al.: Myocardial deformation abnormalities in patients with aortic regurgitation: a strain rate imaging study. Eur J Echocardiogr 2009; 10: 112–119. doi: 10.1093/ejechocard/ien185.
- Gjesdal O, Hopp E, Vartdal T, Claus P, Bijnens B, Jahangiri M: Global longitudinal strain measured by two-dimensional speckle tracking echocardiography is closely related to myocardial infarct size in chronic ischaemic heart disease. Clin Sci 2007; 113: 287–296.
- Chan J, Hanekom L, Wong C, Leano R, Cho GY, Marwick TH: Differentiation of subendocardial and transmural infarction using two-dimensional strain rate imaging to assess short-axis and long-axis myocardial function. J Am Coll Cardiol 2006: 48: 2026–2033.
- Delgado V, Mollema SA, Ypenburg C et al.: Relation between global left ventricular longitudinal strain assessed with novel automated function imaging and biplane left ventricular ejection fraction in patients with coronary artery disease. J Am Soc Echocardiogr 2008; 21: 1244–1250. doi: 10.1016/j.echo.2008.08.010.
- Baccouche H, Maunz M, Beck T, Fogarassy P, Beyer M: Echocardiographic assessment and monitoring of the clinical course in a patient with Tako-Tsubo cardiomyopathy by a novel 3D-speckle-tracking-strain analysis. Eur J Echocardiogr 2009; 10: 729–731. doi: 10.1093/ejechocard/iep064.
- Heggemann F, Weiss C, Hamm K et al.: Global and regional myocardial function quantification by two-dimensional strain in Takotsubo cardiomyopathy. Eur J Echocardiogr 2009; 10: 760–764. doi: 10.1093/ejechocard/jep062.
- Mansencal N, Abbou N, Pillière R, El Mahmoud R, Farcot JC, Dubourg O: Usefulness of two-dimensional speckle tracking echocardiography for assessment of Tako-Tsubo cardiomyopathy. Am J Cardiol 2012; 103: 1020–1024. doi: 10.1016/i.amicard.2008.12.015.
- Burri MV, Nanda NC, Lloyd SG et al.: Assessment of systolic and diastolic left ventricular and left atrial function using vector velocity imaging in Takotsubo cardiomyopathy. Echocardiography 2008; 25: 1138–1144. doi: 10.1111/j.1540-8175.2008.00819.x.
- Kalam K, Otahal P, Marwick TH: Prognostic implications of global LV dysfunction: a systematic review and meta-analysis of global longitudinal strain and ejection fraction. Heart 2014; 100:1673–1680. doi: 10.1136/heartjnl-2014-305538.
- Hestenes SM, Halvorsen PS, Skulstad H et al.: Advantages of strain echocardiography in assessment of myocardial function in severe sepsis. Crit Care Med 2014; 42: e432–440. doi: 10.1097/CCM.0000000000000310.
- Basu S, Frank LH, Fenton KE, Sable CA, Levy RJ, Berger JT: Two-dimensional speckle tracking imaging detects impaired myocardial performance in children with septic shock, not recognized by conventional echocardiography. Pediatr Crit Care Med 2012; 13: 259–264. doi: 10.1097/PCC.0b013e3182288445.
- Landesberg G, Jaffe AS, Gilon D et al.: Troponin elevation in severe sepsis and septic shock. Crit Care Med 2014; 42: 790–800. doi: 10.1097/CCM.000000000000107.
- Orde SR, Pulido JN, Masaki M et al.: Outcome prediction in sepsis: Speckle tracking echocardiography based assessment of myocardial function. Crit Care 2014; 18: R149. doi: 10.1186/cc13987.
- De Geer L, Engvall J, Oscarsson A: Strain echocardiography in septic shock — a comparison with systolic and diastolic function parameters, cardiac biomarkers and outcome. Crit Care 2015; 19: 122. doi: 10.1186/s13054-015-0857-1.

- Shahul S, Gulati G, Hacker MR et al.: Detection of Myocardial Dysfunction in Septic Shock: A Speckle-Tracking Echocardiography Study. Anesth Anala 2015 (Epub ahead of print).
- Lanspa MJ, Pittman JE, Hirshberg EL et al.: Association of left ventricular longitudinal strain with central venous oxygen saturation and serum lactate in patients with early severe sepsis and septic shock. Crit Care 2015; 19: 493. doi: 10.1186/s13054-015-1014-6.
- Chang W-T, Lee W-H, Lee W-T et al.: Left ventricular global longitudinal strain is independently associated with mortality in septic shock patients. Intensive Care Med 2015; 41: 1791–1799. doi: 10.1007/s00134-015-3970-3.
- Cinotti R, Piriou N, Launey Y et al. on behalf of the ATLANRÉA study group: Speckle tracking analysis allows sensitive detection of stress cardiomyopathy in severe aneurysmal subarachnoid hemorrhage patients. Intensive Care Med 2015. doi: 10.1007/s00134-015-4106-5.
- Naidech AM: Cardiac troponin elevation, cardiovascular morbidity, and outcome after subarachnoid hemorrhage. Circulation 2005; 112: 2851–2856.
- van der Bilt I, Hasan D, van den Brink R et al. and SEASAH (Serial Echocardiography After Subarachnoid Hemorrhage) Investigators: Cardiac dysfunction after aneurysmal subarachnoid hemorrhage: relationship with outcome. Neurology 2014; 82: 351–358. doi: 10.1212/WNL.0000000000000057.
- Franchi F, Faltoni A, Cameli M et al.: Influence of positive end-expiratory pressure on myocardial strain assessed by speckle tracking echocardiography in mechanically ventilated patients. Biomed Res Int 2013; 2013: 918548. doi: 10.1155/2013/918548.
- Dalen H, Thorstensen A, Aase SA et al.: Segmental and global longitudinal strain and strain rate based on echocardiography of 1266 healthy individuals: the HUNT study in Norway. Eur J Echocardiogr 2010; 11: 176–183. doi: 10.1093/ejechocard/jep194.
- Marwick TH: Measurement of strain and strain rate by echocardiography: ready for prime time? J Am Coll Cardiol 2006; 47: 1313– 1327
- Kang S-J, Lim H-S, Choi B-J et al.: Longitudinal strain and torsion assessed by two-dimensional speckle tracking correlate with the serum level of tissue inhibitor of matrix metalloproteinase-1, a marker of myocardial fibrosis, in patients with hypertension. J Am Soc Echocardiogr 2008; 21: 907–911. doi: 10.1016/j.echo.2008.01.015.
- Bergenzaun L, Gudmundsson P, Öhlin H et al.: Assessing left ventricular systolic function in shock: evaluation of echocardiographic parameters in intensive care. Crit Care 2011; 15: R200. doi: 10.1186/cc10368.
- Chow P-C, Liang X-C, Cheung EWY, Lam WWM, and Cheung Y-F: New two-dimensional global longitudinal strain and strain rate imaging for assessment of systemic right ventricular function. Heart 2008; 94: 855–859. doi: 10.1136/hrt.2007.131862.
- Weidemann F, Eyskens B, Jamal F et al.: Quantification of regional left and right ventricular radial and longitudinal function in healthy children using ultrasound-based strain rate and strain imaging. J Am Soc Echocardiogr 2002; 15: 20–28.

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