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Original Contribution

SHORT-LAG SPATIAL COHERENCE IMAGING OF CARDIAC ULTRASOUND DATA: INITIAL CLINICAL RESULTS

Muyinatu A. Lediju Bell,*[†] Robi Goswami,[‡] Joseph A. Kisslo,[‡] Jeremy J. Dahl,* and Gregg E. Trahey*[§]

*Department of Biomedical Engineering, Duke University, Durham, North Carolina, USA; †Department of Computer Science, Johns Hopkins University, Baltimore, Maryland, USA; †Division of Cardiology, Department of Medicine, Duke University Medical Center, Durham, North Carolona, USA; and §Department of Radiology, Duke University Medical Center, Durham, North Carolina, USA

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Abstract—Short-lag spatial coherence (SLSC) imaging is a novel beamforming technique that reduces acoustic clutter in ultrasound images. A clinical study was conducted to investigate clutter reduction and endocardial border detection in cardiac SLSC images. Individual channel echo data were acquired from the left ventricle of 14 volunteers, after informed consent and institutional review board approval. Paired B-mode and SLSC images were created from these data. Contrast, contrast-to-noise, and signal-to-noise ratios were measured in paired images, and these metrics were improved with SLSC imaging in most cases. Three cardiology fellows rated the visibility of endocardial segments in randomly ordered B-mode and SLSC cine loops. SLSC imaging offered 22%-33% improvement (p < 0.05) in endocardial border visibility when B-mode image quality was poor (i.e., 80% or more of the endocardial segments could not be visualized by the three reviewers). The percentage of volunteers with poor-quality images was decreased from 21% to 7% with the SLSC beamformer. Results suggest that SLSC imaging has the potential to improve clinical cardiac assessments that are challenged by clutter. (E-mail: mledijubell@jhu.edu) © 2013 World Federation for Ultrasound in Medicine & Biology.

Key Words: Echocardiography, Clutter reduction, Endocardial border detection, Coherence-based beamforming, B-mode, M-mode, Contrast, Signal-to-noise ratio.

INTRODUCTION

Clinical cardiac ultrasound images are challenged by several noise artifacts, including speckle, acoustic shadowing, echo dropout and clutter. Though regarded as a noise artifact, speckle is beneficial for tracking myocardial motion in strain imaging and other motion-dependent applications. In addition, acoustic shadowing or echo dropout may be reduced by adjusting transducer placement, limiting ultrasound blockage by the ribs or enhancing the ultrasound angle of incidence with tissue. Clutter, on the other hand, is a persistent noise artifact that is usually difficult to remove or correct. Unlike speckle, clutter has no known benefits in echocardiography.

Clutter appears as a diffuse haze that reduces image contrast (Lediju et al. 2008) and obscures visualization of endocardial borders, tumors, vegetations and other cardiac abnormalities (Mele et al. 2006; Patel et al. 1980). It presents a major challenge in strain imaging and automated or manual border detection (Bezante et al. 2005; Skolnick et al. 1999; Teske et al. 2007; Vandenberg et al. 1992). Sources of cardiac clutter include wavefront distortions encountered as sound travels transthoracically, sound reverberations in multiple tissue layers and reflections from off-axis structures such as the ribs, lungs, chordae tendineae, valves and myocardial walls (Cloutier et al. 2003; Hinkelman et al. 1997; Lediju et al. 2009a; Yeh 1977).

Harmonic imaging is a common clutter reduction method, whereby images contain frequencies that are integer multiples of the frequencies of transmitted pulses. These harmonic frequencies are generated as a pulse travels through tissue. One reason for the reduced clutter content in harmonic images is that harmonic frequencies are not fully developed near the transducer surface, where sound reverberations typically occur (Muir and Carstensen 1980; Shapiro et al. 1998; Starritt et al. 1986; Tranquart et al. 1999). In several studies, harmonic imaging lowered the percentage of patients with

Address correspondence to: Muyinatu A. Lediju Bell, 128 Hackerman Hall, 3400 N. Charles St., Baltimore, MD 21218, USA. E-mail: mledijubell@jhu.edu

suboptimal images from 45%-51% to 11%-24% (Caiati et al. 1999; Chirillo et al. 2005). This indicates that harmonic imaging improves image quality in a subset of patients, yet it does not always sufficiently reduce clutter.

Another approach to clutter reduction is to separate tissue signals from clutter noise using motion filters or principal component analysis-based filters (Gallippi and Trahey 2002; Lediju et al. 2009b; Mauldin et al. 2011; Zwirn and Akselrod 2006). These filters are effective at reducing stationary or slowly moving clutter. In echocardiography, this type of clutter is often due to acoustic reverberations within the chest wall or reflections from stationary and slowly moving extracardiac structures such as the rib cage and lungs. One challenge with these filters is their limited ability to remove the highervelocity clutter that is due to reflections from intracardiac structures like the chordae tendineae, valves and myocardial walls. In addition, these filters require information from multiple frames of image data to differentiate myocardial motion from clutter noise.

In transesophageal echocardiography (TEE), images are acquired by inserting an ultrasound transducer in the esophagus, rather than using the standard transthoracic window (Ragland and Tak 2006; Ward et al. 2004). Despite the improvements in image quality, TEE poses a discomfort to patients and is not recommended for routine clinical use (Ragland and Tak 2006; Hanrath 2001). Another alternative is contrast echocardiography, which uses contrast agents to enhance endocardial border visualization (Zwirn et al. 2009). However, the injection of contrast agents presents an additional expense to patients and necessitates a sterile environment for intravenous access (Al-Mansour et al. 2000; Moir and Marwick 2004; Ophir and Parker 1989).

Short-lag spatial coherence (SLSC) imaging (Lediju et al. 2011), a novel beamforming approach that uses the

spatial coherence of back-scattered ultrasound echoes, overcomes many of the challenges associated with existing clutter-reduction or border-enhancement approaches. The spatial coherence of echoes from myocardium exhibits characteristics different from that of clutter and blood (Bamber et al. 2000). This difference is most noticeable over small spatial differences; thus, SLSC imaging is implemented by computing the spatial coherence of echoes received by closely spaced transducer elements. In this article, we describe initial results of the SLSC beamforming method applied to *in vivo* cardiac data.

METHODS

Study population

The study population consisted of 14 volunteers (9 men, 5 women; age, mean \pm standard deviation [SD]: 55 ± 17 y, range: 33-85 y). Six of the volunteers were employees of Duke University and eight were patients scheduled for an echocardiogram at the Duke University Medical Center. Two of the patients required the use of contrast agents for endocardial visualization during a standard exam, one patient suffered from pulmonary hypertension and two patients had scar tissue in the chest wall, one from a heart transplant and one from breast surgery. This study was approved by the Duke University institutional review board (Protocol ID: Pro00030455), and informed consent was obtained from all volunteers.

Data acquisition

Each volunteer was placed in the left lateral decubitus position to acquire mid-level short-axis views and apical four-chamber views of the left ventricle (LV). These views are illustrated in Figure 1 relative to a schematic of the heart. A Verasonics ultrasound scanner (Redmond, WA, USA) and a 64-element ATL P4–2 phased array transducer were used to acquire the individual

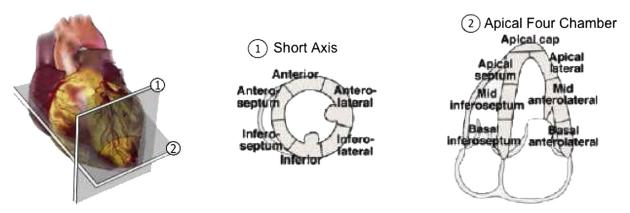


Fig. 1. Schematic of two standard ultrasound views of the heart and associated segments of the left ventricle walls. There are six segements in the mid-level short axis view and seven segments in the apical four-chamber view. Reprinted from Lang et al. (2005) with permission from Elsevier.

channel radiofrequency echoes needed to form 35 frames of image data. Data were acquired at a rate of approximately 7–12 frames/s.

The transmit beam was electronically steered and focused at 50 different lateral locations, such that the sector width of each frame was 45°, and the lateral beam spacing was 0.9° per line. No apodization was applied to the transmit beam. The minimum depth required to display the entire LV in each volunteer was selected from pre-set values of 8, 10, 12, 14, 16 and 18 cm. The transmit focus ranged from 4 to 10 cm, depending on the location of the heart. In the short-axis views, the focus was consistent with the location of the LV, and in the apical views, it was consistent with at least two of the seven endocardial segments. The axial sampling frequency was 30 MHz, unless otherwise stated. The transducer transmit frequency was 2 MHz.

Gain settings were standardized for all volunteers. The programmed pulse sequences were below the Food and Drug Administration limits for spatial-peak pulse-average intensity and mechanical index (Phillips and Harris 2008). The size of one data set (*i.e.*, the individual channel data required to make a cine loop of 35 image frames) was approximately 6–7 gigabytes.

B-mode and SLSC image formation

The ultrasound echo data received by the 64 individual transducer elements were processed offline to create matched (or paired) B-mode and SLSC images and cine loops. B-mode images were created by applying a delay-and-sum beamformer with dynamic receive focusing and no aperture growth. The data were then envelope detected, normalized to the maximum value, scan converted and log compressed using settings that were determined to be optimal by the cardiology fellow who performed the acquisitions.

Short-lag spatial coherence images were formed by first computing a normalized spatial coherence function at one depth (*i.e.*, one axial position) of the delayed back-scattered echoes that correspond to one lateral location in an image, as described by Fedewa et al. (2003):

$$\widehat{R}(m) = \frac{1}{N - m} \sum_{i=1}^{N - m} \frac{\sum_{n=n_1}^{n_2} s_i(n) s_{i+m}(n)}{\sqrt{\sum_{n=n_1}^{n_2} s_i^2(n) \sum_{n=n_1}^{n_2} s_{i+m}^2(n)}}$$
(1)

where $\widehat{R}(m)$ is the normalized spatial correlation measured across a receive aperture, m is the element separation distance (or element lag), N refers to the total number of receive elements (i.e., N=64) and $s_i(n)$ is the time-delayed signal received by the *i*th element at depth, or time, n, expressed in number of samples. The correlation kernel size $(n_2 - n_1)$ used to calculate the spatial coherence function was approximately one wave-

length. The resulting coherence function was then summed over the short-lag region, as described by the equation

$$R_{\rm sl} = \sum_{m=1}^{M} \widehat{R}(m),\tag{2}$$

where $R_{\rm sl}$ is the short-lag spatial coherence and M is equal to 6 lags, unless otherwise stated. Equations (1) and (2) were repeated for all possible axial and lateral positions to create the different pixels in a SLSC image. The SLSC image was then scan converted and normalized to its maximum value. All SLSC images were displayed on a linear scale that ranged from 0 to 0.95 (*i.e.*, 0%–95% of its maximum value). No compression was applied to display SLSC images.

Endocardial visibility and scoring system

Cine loops of the short-axis views were grouped separately from those of the apical four-chamber views. Each group was randomized and reviewed independently by three cardiology fellows, who each had at least 3 years of experience. One reviewer was the cardiology fellow who acquired the images (reviewer 1), and the other two reviewers were blinded to the study (reviewers 2 and 3). Cine loops were observed in the cardiology reading room at Duke University Hospital, with the same equipment and lighting conditions used to diagnose clinical conditions.

The observers were given the segment models shown in Figure 1, which complied with the American Society of Echocardiography standards for short-axis and apical four-chamber views (Lang et al. 2005). There are six segments in the short-axis view and seven segments in the apical four-chamber view. Qualitative assessment of endocardial visualization of each segment of the LV during systole and diastole was performed with a visual scoring system ranging from 1 to 3, where 1 = endocardial border clearly visible; 2 = endocardial border visible, but not clearly (*i.e.*, poorly visible); and 3 = no endocardial border visible.

In the short-axis and apical four-chamber views, results were separated according to B-mode image quality, based on the scores of the three reviewers. Good quality was defined as unanimous visibility (score = 1 or 2) of \geq 80% of the endocardial segments in systole and diastole (three volunteers). Poor quality was defined as no visibility (score = 3) of \geq 80% of the endocardial segments in systole or diastole (three volunteers). The remaining volunteers were grouped as having medium-quality B-mode images (eight volunteers). The number of segments in each "score" category for each "image-quality" category was recorded for each reviewer. The same criteria were used to group SLSC images according

to image quality for comparison with separations based on B-mode image quality.

Performance metrics

For each volunteer, one B-mode image with relatively well-defined endocardial borders (*i.e.*, relative to the quality of all B-mode images in a cine loop) was selected from the short-axis views of the LV. The selected B-mode image and its matched SLSC image were used to characterize performance. Performance was evaluated by measuring contrast (*C*), contrast-to-noise ratio (CNR) and signal-to-noise ratio (SNR) in the same locations in the matched B-mode and SLSC images, using the equations

$$C = 20 \log \left(\frac{S_{\rm e}}{S_{\rm v}} \right) \tag{3}$$

$$CNR = \frac{|S_{v} - S_{e}|}{\sqrt{\sigma_{v}^{2} + \sigma_{e}^{2}}}$$
(4)

$$SNR = \frac{S_e}{\sigma_e}, \tag{5}$$

where $S_{\rm v}$ and $S_{\rm e}$ are the mean signals in the ventricle and endocardium, respectively, and $s_{\rm v}$ and $s_{\rm e}$ are the standard deviations of signals in the ventricle and endocardium, respectively. All calculations were performed on uncompressed data. The signals used to measure B-mode performance metrics were taken from scan-converted, envelope-detected data, whereas the signals used to measure SLSC performance metrics were taken from scan-converted SLSC data. In addition, contrast, CNR and SNR were measured as a function of the short-lag value (i.e., M in eqn [2]) for each volunteer. All performance metrics were calculated independently of the enodcardial visibility scores assigned by each reviewer. Performance metric results were then categorized by image quality after the two independent analyses were completed.

Statistical analysis

B-Mode and SLSC performance metrics (*i.e.*, contrast, CNR and SNR) were compared with a paired t-test to determine statistical significance. Cohen's d was calculated to determine the effect size and clinical significance of the performance metrics. Visibility scores were reported as the median and interquartile range of the percentage of total segments evaluated by each reviewer in each category. The numbers of segments not visible in B-mode and SLSC images (score = 3) for each image-quality category were compared with a Wilcoxon signed rank test for paired data. Differences between B-mode and SLSC images were considered statistically significant for p values <0.05 and clinically significant for Cohen's $d \ge 0.8$.

Data from the apical four-chamber view were also summarized as the mean visibility score of each segment in B-mode and SLSC images. Differences between the means were compared with a paired t-test, and statistical significance was maintained at p < 0.05.

To determine the level of inter-observer agreement among the visibility scores, Fleiss' kappa (κ) statistic was calculated (Cardillo 2007). Separate statistics were calculated for the short-axis and apical four-chamber views. κ values <0.00 were interpreted as poor agreement; values between 0 and 0.20 were interpreted as slight agreement; and values between 0.21 and 0.4 were interpreted as fair agreement, in accordance with Landis and Koch (1977).

Software

All image processing and data analyses were performed with MATLAB (The Mathworks, Natick, MA, USA) software and several MATLAB Executable (MEX) files that enabled the interface of C++ subroutines. The time to calculate one SLSC image was approximately 2 s with a 3.5-GHz processor, when the lag variables m and M were limited to 6, whereas the time to calculate one B-mode image with the same processor was approximately 20 ms.

RESULTS

Short-axis views

Matched B-mode and SLSC images of the LV of one volunteer are displayed in Figure 2(a and b, respectively). The LV and adjacent right ventricular and atrial cavities in Figure 2(b) contain less clutter than the respective locations in the matched B-mode image. Observation of the corresponding cine loops revealed a reduction of both stationary and nonstationary clutter for all frames of acquired data, particularly in the near-field region. The axial sampling frequency was decreased to 10 MHz to increase the frame rate of this acquisition.

Short-lag spatial coherence and traditional M-mode images were created from channel data by forming an image of the same lateral position as a function of time. The comparative M-modes (Fig. 2c) reveal that the SLSC image clarifies the inferior endocardial border (*solid arrow*), while the pericardium and anterior endocardial border are well visualized (*dashed* and *dotted arrows*, respectively).

The SLSC image was used to manually trace the endocardial border in the LV (Fig. 2b). Contrast, CNR and SNR, calculated from signals in the regions of interest (ROIs) indicated in Figure 2(b), were 6.6 dB, 1.1 and 2.3, respectively, in the B-mode image and 9.1 dB, 1.1 and 2.0, respectively, in the SLSC image. Thus, there is a 2.4-dB improvement in contrast in the SLSC

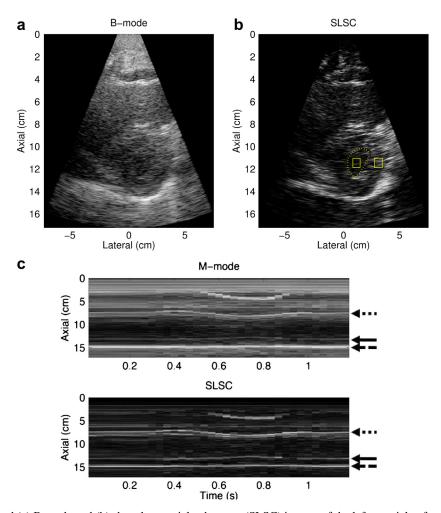


Fig. 2. Matched (a) B-mode and (b) short-lag spatial coherence (SLSC) images of the left ventricle of volunteer 2. The endocardial border was manually outlined (dotted line) with the aid of the associated cine loop, and the outlined boxes were used to calculate contrast, contrast-to-noise ratio and signal-to-noise ratio. (c) Corresponding M-mode and SLSC images as a function of time. The arrows indicate the locations of the anterior endocardial border (*dotted arrow*), inferior endocardial border (*solid arrow*) and pericardium (*dashed arrow*). The axial sampling frequency was 10 MHz, and SLSC images were created with M=10. B-mode and M-mode images are shown with 50-dB dynamic range, whereas SLSC images are shown on a linear scale ranging from 0%-95% of the maximum value.

image, while the CNR and SNR are similar in the B-mode and SLSC images. ROIs in similar locations were used to calculate performance metrics for all of the volunteers. In this example, SLSC images were created with M=10 to improve the resolution of the endocardial border.

Matched B-mode and SLSC images of the LV and mitral valves in another volunteer are displayed in Figure 3(a and b, respectively). The B-mode image is an example of a good-quality image, where more than 80% of the endocardial border is visualized. Clutter is reduced and contrast is improved by approximately 4 dB in the SLSC image. CNR is improved by 0.5 and SNR is increased by 0.4 in the SLSC image. The SLSC image also shows more well-defined borders.

An example of a poorer-quality B-mode image appears in Figure 4(a), where less than 80% of the endo-

cardial border is clearly visualized. The borders are clearer in the matched SLSC image in Figure 4(b). The contrast between the ventricle and the endocardium is improved by 11 dB in the SLSC image. CNR and SNR are improved by 1.5 and 1.6, respectively, in the SLSC image.

Performance metrics

The comparative performance of B-mode and SLSC images from all volunteers was measured using contrast, CNR and SNR metrics. These performance metrics were measured in matched B-mode and SLSC images from each volunteer. Matched values, categorized by the B-mode image quality determined from the endocardial visibility scores (*i.e.*, good, medium or poor), are compared in Figure 5. B-mode performance is displayed

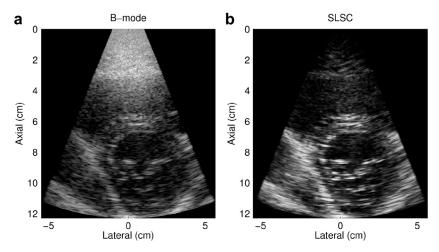


Fig. 3. Matched (a) B-mode and (b) short-lag spatial coherence (SLSC) images of the left ventricle of volunteer 5. The B-mode image is an example of a good-quality image, where more than 80% of the endocardial border is visualized. The corresponding SLSC image shows reduced clutter and more well-defined borders. The B-mode image is shown with 50-dB dynamic range, whereas the SLSC image is shown on a linear scale ranging from 0%–95% of the maximum value.

on the abscissa axis, and SLSC performance is displayed on the ordinate axis. Values above the dashed line indicate improvement with SLSC imaging. Note that contrast, CNR and SNR are mostly improved in SLSC images for each of the three image-quality categories.

The means of measured contrast, CNR and SNR values are 17 dB, 1.6 and 2.2, respectively, in the SLSC images and 8 dB, 0.9 and 1.8, respectively, in the B-mode images. Thus, in the SLSC images, the average contrast is improved by 9 dB (p = 0.007, Cohen's d = 1.0); the average CNR is improved by 0.7 (p = 0.00001, Cohen's d = 1.5); and the average SNR is improved by 0.4 (p = 0.02, Cohen's d = 0.9). The ranges of contrast,

CNR and SNR improvement with SLSC imaging were -2 to 33 dB, -0.1 to 1.5 and -0.4 to 1.6, respectively.

The performance metrics were measured as a function of the short-lag value M in SLSC images from the 14 volunteers. Figure 6 illustrates that as M is increased, the mean contrast increases for M < 6, then remains constant for $M \ge 6$, while the mean SNR decreases and the mean CNR decreases for M > 2. The mean and standard deviation of B-mode measurements are provided as a reference. A value of M = 6 was chosen to display images throughout this article (unless otherwise stated), to compare performance metrics in Figure 5 and to make the cine loops that were reviewed, because it is

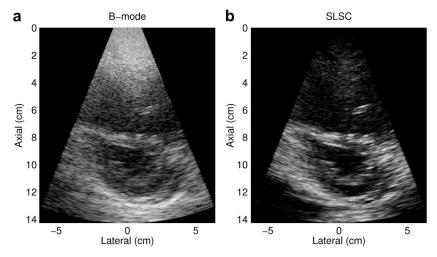


Fig. 4. Matched (a) B-mode and (b) short-lag spatial coherence (SLSC) images of the left ventricle of volunteer 7. The B-mode image is an example of a poorer-quality image, where less than 80% of the endocardial border is clearly visualized. The endocardial border is better defined in the SLSC image. The B-mode image is shown with 50-dB dynamic range, whereas the SLSC image is shown on a linear scale ranging from 0%–95% of the maximum value.

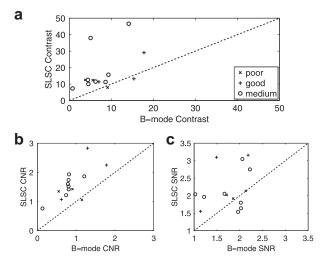


Fig. 5. Scatterplots of (a) contrast, (b) contrast-to-noise ratio (CNR) and (c) signal-to-noise ratio (SNR) measured in B-mode and short-lag spatial coherence (SLSC) images from the 14 volunteers. The data points are categorized by the B-mode image quality determined from the endocardial visibility scores (*i.e.*, good, medium or poor). Data points above the dashed line indicate better contrast, CNR or SNR in the SLSC image compared with the matched B-mode image. The SLSC values were calculated with M=6.

the lowest short-lag value with the best average contrast. Higher values of M have poorer CNRs and SNRs. Examples of three SLSC images formed with different values of M and the matched B-mode image are given in Figure 7. Note the changes in contrast, CNR, and SNR as M is varied. Lateral resolution is also improved as M increases.

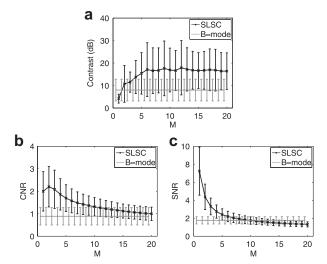


Fig. 6. Mean (a) contrast, (b) contrast-to-noise ratio (CNR) and (c) signal-to-noise ratio (SNR) measured in short-lag spatial coherence (SLSC) images from the 14 volunteers, as a function of the short-lag value *M*. B-mode measurements are provided as a reference. Error bars indicate \pm one standard deviation from the mean.

Observer reviews for the short-axis views

The scores assigned to each segment in the short axis view are summarized in Figure 8. Improvement in segment visibility between B-mode and SLSC results is indicated by an increase in the height of the bars labeled with scores of 1 (clearly visible) and 2 (poorly visible), for each image-quality category in systole or diastole. Improvement is also indicated by a decrease in the height of the bars labeled with a score of 3 (segments not visible). The greatest improvement with SLSC imaging was observed in poor-quality B-mode images, where the percentage of segments not visualized with B-mode imaging decreased by 33% and 22% in systole and diastole, respectively (p < 0.05). In good- and mediumquality images, there were no statistically significant improvements (i.e., no statistically significant differences in the height of the bars labeled 1, 2 or 3).

To compare the percentage of volunteers with poorquality B-mode images with the percentage of those with poor-quality SLSC images, the volunteers were separated according to SLSC image quality, using the same criteria described for separations based on B-mode image quality. The number of volunteers with poor-, medium- or goodquality SLSC images was 1, 10 and 3, respectively. Thus, in data from the short-axis views of the LV, SLSC imaging lowered the percentage of volunteers with poor-quality images from 21% to 7%, increased the percentage of volunteers with medium-quality images from 57% to 71% and did not change the percentage of volunteers with good-quality images (Fig. Supplemental Video 1 shows a cine loop of matched B-mode and SLSC images that were categorized as good-quality. Supplemental Video 2 shows poor-quality B-mode images and matched SLSC images that were categorized as medium quality.

Apical four chamber views

Matched B-mode and SLSC images of an apical four-chamber view from one volunteer (Fig. 10a and 10b, respectively) reveal a portion of chordae tendineae in the LV. The clutter in the near field obscures the apical endocardium segments. The three reviewers rated the apical cap as not visible (score = 3) in the B-mode image. Reviewers 1 and 2 observed better visualization of this segment (score = 2) in the SLSC image, whereas reviewer 3 recorded no difference.

The three reviewers also noted that the three lateral segments (apical lateral, mid anterolateral, and basal anterolateral) were not visualized in the B-mode image. In the SLSC images, these three segments maintained no visibility by Reviewer 1 (score = 3) and were poorly visualized by Reviewer 2 (score = 2). Reviewer 3 recorded poor visibility (score = 2) of the apical lateral segment in the SLSC image during systole, but the other

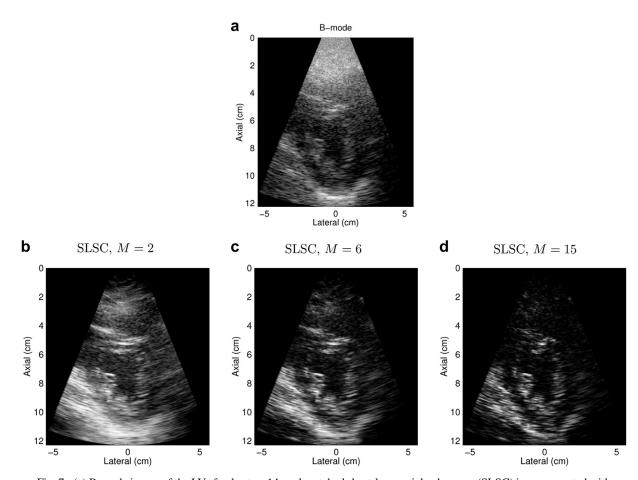


Fig. 7. (a) B-mode image of the LV of volunteer 14, and matched short-lag spatial coherence (SLSC) images created with (a) M = 2, (b) M = 6 and (c) M = 15. Note the changes in contrast, contrast-to-noise ratio (CNR), signal-to-noise ratio (SNR) and resolution as *M* is varied. The B-mode image is shown with 50-dB dynamic range, whereas the SLSC images are shown on a linear scale ranging from 0%–95% of the maximum value.

two lateral segments were rated as not visible by this reviewer.

Matched B-mode and SLSC images of an apical four-chamber view from another volunteer (Fig. 11a and 11b, respectively) include internal structures in the LV. Visibility of the apical segments is obscured by the presence of clutter in the near field. Reviewers 1 and 3 rated the apical cap, apical septum and midinferoseptum as not visible with B-mode imaging during diastole (score = 3). With SLSC imaging, these segments maintained no visibility by reviewer 3 (score = 3) and were poorly visualized by reviewer 1 (score = 2). However, reviewer 2 rated these three segments with the same visibility in B-mode and SLSC images (2, 2 and 1, respectively).

It is interesting to note that the three reviewers visualized (score = 1 or 2) the basal lateral segment in the B-mode and SLSC cine loops represented in Figure 11, but when the images are viewed side-by-side, the segment appears to be missing in the SLSC image. Two of the

reviewers rated this segment with equal visibility in the B-mode and SLSC images, whereas one reviewer recorded that this segment was better visualized in the SLSC image. Supplemental Video 3 contains cine loops of this data acquisition.

The scores assigned to each segment in the apical four-chamber view are summarized in Figure 12. The greatest improvement with SLSC imaging was observed in poor-quality B-mode images during systole, where the percentage of segments not visualized with B-mode imaging decreased by 19%; however, the p-value of 0.06 indicates that this result is not statistically significant. In good- and medium-quality images, there were no statistically significant differences in endocardial border visualization between B-mode and SLSC images (p >> 0.05).

The mean visibility scores (1 = clearly seen, 2 = poorly seen, 3 = not visible) for the apical four-chamber views (Fig. 13) ranged from 2.1 to 2.6, indicating that a majority of these apical views contained segments that

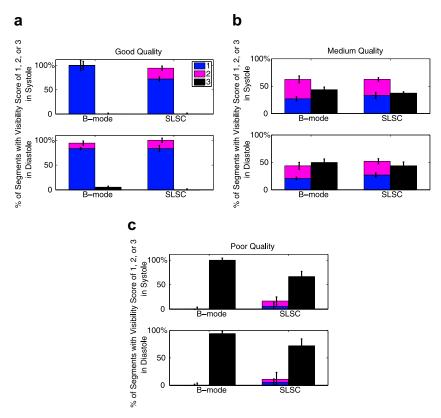


Fig. 8. Visibility of the segments in the short-axis view of the left ventricle. The images were separated by B-mode image quality (good, medium, poor). The number of segments with each visibility score (1 = clearly seen, 2 = poorly seen, 3 = not visible) is expressed as a percentage of the total number of segments in each image-quality category. The height of the bars represent the median of the three independent observers, and the error bars indicate the interquartile range for each score category in each image-quality category in sytole and diastole. The *p*-values were (a) 1 in systole and diastole in good-quality images; (b) 0.45 and 0.64 in systole and diastole, respectively, in medium-quality images; and (c) 0.047 and 0.0078 in systole and diastole, respectively, in poor-quality images. The κ statistics indicate fair agreement among the reviewers ($\kappa = 0.30$).

were not visible or poorly visible. Differences between B-mode and SLSC mean visibility scores were not statistically significant.

DISCUSSION

Improvements with and limitations of SLSC imaging

Examples of patients that are difficult to image with conventional B-mode beamforming methods (Figs. 2 and 4) are contrasted with examples from a less challenging patient (Fig. 3). Clutter is noticeably reduced in the matched SLSC images. Clutter reduction was observed in a majority of SLSC images from the 14 volunteers, as evidenced by the improved contrast values in Figure 5(a). These results suggest that SLSC imaging has the potential to reduce clutter when conventional beamforming methods produce good-, medium- or poor-quality B-mode images. Although clutter was reduced and contrast was improved in medium- and good-quality B-mode images, no statistically significant

differences in segment visibility were observed in these cases.

Statistically significant improvements in segment visibility were observed in the short-axis views of patients with poor-quality B-mode images (Fig. 8c), as SLSC imaging decreased the percentage of myocardial segments not visualized with B-mode imaging by 22%–33%. These results indicate that SLSC imaging improves endocardial border visibility in poor-quality B-mode images without degrading that of medium- and good-quality images. The percentage of volunteers with poor-quality images decreased from 21% with B-mode imaging to 7% with SLSC imaging (Fig. 9).

When low-amplitude signals are adjacent to structures with higher amplitudes, the coherence of the lower-amplitude signal is decreased relative to that of the higher-amplitude signal, thus providing a source of contrast in SLSC images. This occurs because the off-axis reflections from the higher-amplitude structures add incoherently with the on-axis reflections from the

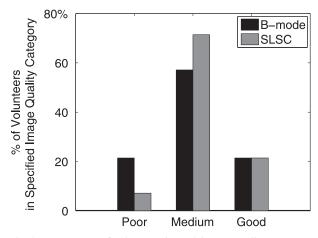


Fig. 9. Percentage of volunteers in each image quality category, as defined by the number of visible segments in B-mode or short-lag spatial coherence (SLSC) images of the left ventricle short-axis view. SLSC imaging decreased the percentage of volunteers with poor-quality images.

lower-amplitude structures (Lediju et al. 2011). With respect to implications for endocardial border detection, this effect can be beneficial in the presence of the mitral valve (*e.g.*, Fig. 3) and detrimental in the presence of a bright pericardium (*e.g.*, Fig. 11).

In SLSC images, there is a known trade-off among contrast, CNR, SNR and lateral resolution that varies with the selected short-lag value, M (Dahl et al. 2011; Lediju et al. 2011). Although resolution is degraded at lower values of M, the spatial correlation between structures in SLSC and B-mode images is expected to be similar if the value of M is optimized, as illustrated in

Figure 7 and observed in previous theoretical, simulated and experimental results (Lediju et al. 2011; Lediju Bell 2011). Note that selecting the highest short-lag value offers the best resolution, but degrades CNR and SNR, as illustrated in Figure 6. The variation in these measurements as a function of M is likely due to the increased spatial frequency content with the inclusion of more lags (Lediju et al. 2011).

Study limitations

B-Mode and SLSC endocardial segment scores were based on visual semi-quantitative estimates, introducing a degree of reviewer subjectivity. Yet, statistically significant improvements with SLSC imaging of poor-quality B-mode data were achieved with this subjective analysis. In addition, objective measurements of contrast, CNR and SNR were performed on the raw data. Additional contrast enhancement and improvements in endocardial border detection may have been achieved with harmonic data (Becher et al. 1998; Dahl et al. 2012; Kasprzak et al. 1999).

Contrast, CNR and SNR measurements were potentially biased toward B-mode images, particularly for the volunteers with poor-quality B-mode images, because the image selection criteria for calculating these performance metrics was based on B-mode images. An alternative would have been to select good SLSC images and make comparisons to matched B-mode images created from the same channel data. This choice would likely improve the contrast and CNR measurements of SLSC images. However, to prevent bias toward SLSC imaging, selections were based on B-mode images. In addition, the

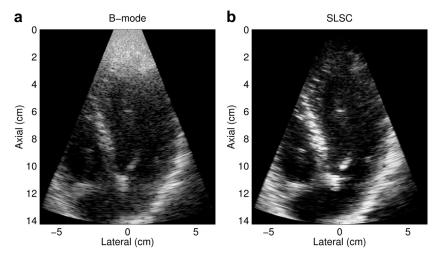


Fig. 10. Apical four-chamber view of the left ventricle of volunteer 3 in matched (a) B-mode and (b) short-lag spatial coherence (SLSC) images. The endocardial borders are more clearly defined, particularly in the near field. The transmit focus was 8.3 cm. The B-mode image is shown with 50-dB dynamic range, whereas the SLSC image is shown on a linear scale ranging from 0%–95% of the maximum value.

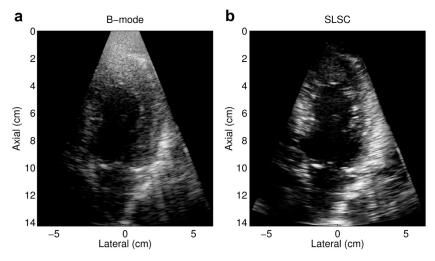


Fig. 11. Apical four-chamber view of the left ventricle of volunteer 2 in matched (a) B-mode and (b) short-lag spatial coherence (SLSC) images. There is reduced clutter in the near field of the SLSC image, and the basal lateral segment appears to be missing in the SLSC image compared with the B-mode image (see Supplemental Video 3 for cine loops of this acquisition). The transmit focus was 8.3 cm. The B-mode image is shown with 50-dB dynamic range, whereas the SLSC image is shown on a linear scale ranging from 0%–95% of the maximum value.

size of the ROIs used to calculate these measurements was limited by the size of the left ventricle. If larger ROIs could be used, the measurements would be independent of additional image statistics (e.g., number of speckles in each ROI and the spacing between them), and the results would be more accurate. Despite the ROI size limitations and the potential biases toward B-mode imaging, the SLSC measurements outperformed B-mode measurements in a majority of cases.

Although segment visibility, contrast, CNR and SNR were quantitatively assessed in B-mode and SLSC images, it is difficult to quantify resolution. SLSC resolution is dependent on an array of factors, including target size and location, short-lag value (*M*) and magnitude of clutter noise present (Lediju Bell et al. 2011).

Prospective studies with larger numbers of patients are required to fully assess the clinical role of SLSC imaging. For example, the high mean visibility scores in Figure 13 indicate that there were few images with visible segments in the apical four-chamber views. This is likely due to the presence of clutter in Bmode images. In SLSC images, the poor visibility is due to a complicated combination of the presence of bright extracardiac scatterers, the absence of intracardiac structures in the image plane and the location of the focus. The poor visibility in SLSC images is also potentially due to the low SNR of the channel signals (Dahl et al. 2011). Nonetheless, the examples presented in Figures 10 and 11 illustrate improved endocardial visualization in the apical segments in the presence of high-amplitude clutter.

Note that the apical four-chamber systolic results in Figure 12(c) are statistically significant at a significance level of 10% (p < 0.1), whereas conclusions about other results remain the same at this higher significance level. It is promising that statistical significance is achieved at the 10% significance level with the limited number of visible segments and the small sample size of three volunteers in the poor-quality image category.

Although one frame of data was acquired and processed in approximately 2 s, real-time SLSC imaging requires faster processing times, which could potentially be achieved with optimized correlation algorithms, advanced processors and graphics processing unit (GPU)-based computing. With access to faster processing methods, the sonographer or cardiologist performing a scan would have the ability to adjust the short-lag value (*M*) and optimize SLSC images in real time.

Clinical implications

Visualization of endocardial borders is essential to determining LV function. In several studies, 50%–64% of patients had inadequate visualization of the heart and the LV endocardial border, resulting in inconclusive diagnoses of abnormal cardiac conditions, such as congestive heart failure, endocarditis, mitral regurgitation, thrombi and aortic dissection (Heidenreich et al. 1999; Hwang et al. 1993; Pavlides et al. 1990).

Short-lag spatial coherence imaging reduces clutter and significantly improves endocardial visualization of the LV short-axis view in patients with poor-quality Bmode images. Thus, there is potential for SLSC imaging

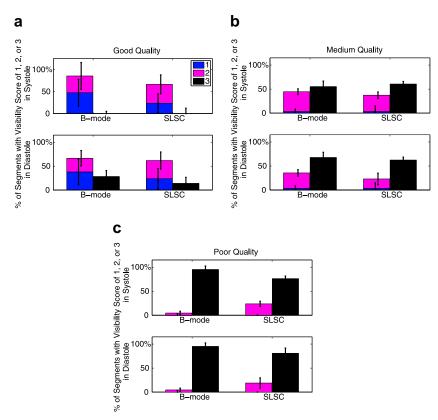


Fig. 12. Visibility of the segments in the apical four-chamber view of the LV. The images were separated by B-mode image quality (good, medium, poor). The number of segments with each visibility score (1 = clearly seen, 2 = poorly seen, 3 = not visible) is expressed as a percentage of the total number of segments in each image-quality category. The height of the bars represents the median of the three independent observers, and the error bars indicate the interquartile range for each score category in each image-quality category in sytole and diastole. The *p*-values were (a) 0.50 and 1 in systole and diastole, respectively, in good-quality images; (b) 0.55 and 0.35 in systole and diastole, respectively, in medium-quality images; and (c) 0.06 and 0.25 in systole and diastole, respectively, in poor-quality images. The κ statistics indicate fair agreement among the reviewers ($\kappa = 0.21$).

to improve qualitative clinical assessments of LV function. There may also be potential to improve cardiac measurements that rely on endocardial border detection, such as volume, mass and ejection fraction. In addition, the reduced clutter in the near-field region of the apical four-chamber views suggests that SLSC might be a preferred method for imaging apical cardiac masses like tumors and thrombi.

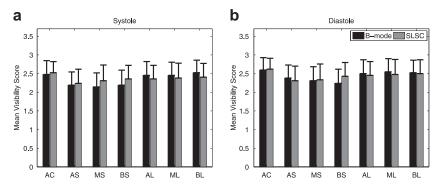


Fig. 13. Mean visibility score (1 = clearly seen, 2 = poorly seen, 3 = not visible) of each segment in B-mode and short-lag spatial coherence (SLSC) images of the apical four-chamber view in (a) systole and (b) diastole. AC = apical cap, AS = apical septum, MS = mid-inferoseptum, BS = basal inferoseptum, AL = apical lateral, ML = mid anterolateral, and BL = basal anterolateral. The *p*-values exceed 0.05 for all segments in systole and diastole, indicating no statistically significant differences.

CONCLUSIONS

Short-lag spatial coherence imaging improves endocardial border detection when B-mode images contain a majority of poorly visualized endocardial segments due to the presence of acoustic clutter. Quantitative metrics revealed better contrast, CNR and SNR with SLSC imaging in most patients. Three cardiology fellows reviewed and scored the images independently, and their scores revealed that SLSC imaging lowered the percentage of volunteers with poor-quality images from 21% to 7%. This work indicates the potential of SLSC imaging to reduce cardiac clutter, clarify endocardial borders and thereby enhance subjective analyses of left ventricular function. There is also potential to improve visualization of cardiac abnormalities and calculations of border-dependent cardiac performance metrics.

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SUPPLEMENTARY DATA

Supplementary data related to this article can be found online at http://dx.doi.org/10.1016/j.ultrasmedbio.2013.03.029.

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