LV Volume Quantification via Spatiotemporal Analysis of Real-Time 3-D Echocardiography

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Abstract—This paper presents a method of four-dimensional (4-D) (3-D + Time) space–frequency analysis for directional denoising and enhancement of real-time three-dimensional (RT3D) ultrasound and quantitative measures in diagnostic cardiac ultrasound. Expansion of echocardiographic volumes is performed with complex exponential wavelet-like basis functions called brushlets. These functions offer good localization in time and frequency and decompose a signal into distinct patterns of oriented harmonics, which are invariant to intensity and contrast range. Deformable-model segmentation is carried out on denoised data after thresholding of transform coefficients. This process attenuates speckle noise while preserving cardiac structure location. The superiority of 4-D over 3-D analysis for decorrelating additive white noise and multiplicative speckle noise on a 4-D phantom volume expanding in time is demonstrated. Quantitative validation, computed for contours and volumes, is performed on in vitro balloon phantoms. Clinical applications of this spatiotemporal analysis tool are reported for six patient cases providing measures of left ventricular volumes and ejection fraction.

Index Terms—Echocardiography, LV volume, spatiotemporal analysis, speckle denoising.

I. INTRODUCTION

CARDIAC three-dimensional (3-D) imaging was conceived as a method to circumvent the shortcomings of two-dimensional (2-D) echoplanar imaging. Volume quantification and performance assessment of irregularly shaped 3-D cardiac chambers or the description of valve morphology using 2-D images is inherently problematic because of the dynamic nature of the heart. There has been a tremendous effort within the ultrasound community over the last decade dedicated to the development of new cardiac 3-D echo technology. In the first generation of 3-D echo systems (3D FreeScan), a spatial locator was incorporated on the ultrasound beam for the reconstruction of a 3-D representation of the heart. A sound or magnetic signal-emitting device was positioned in a free-hand ultrasound transducer, allowing the position of any particular 2-D plane to be registered by an external locator and used for later reconstruction. Since five to ten planes through the heart were required at a preset point in each cardiac cycle, this system was cumbersome to use and only a static 3-D image could be reconstructed [1], [2]. The second-generation 3-D imaging system, originally developed by Hewlet-Packard, took the form of a transducer that rotated around a fixed axis or moved along a fixed path, obtaining images of the heart at regularly spaced intervals. The 2-D images were later reconstructed into a 3-D volume. By obtaining 2-D images at different time points in the cardiac cycle, these axial slices could be later reconstructed to show the motion of the heart. However, the acquisition remained cumbersome and reconstruction was a time-consuming, off-line process. Nevertheless, these rotational probes have yielded accurate results for mass and volume calculations of cardiac chambers [3], [4] and described well a variety of cardiac lesions [5]. However, because of inherent difficulties these systems have not become clinically applicable. In order to circumvent the problems associated with the previous forms of 3-D imaging, a third-generation of volumetric imaging (Volumetrics Medical Imaging, Durham, NC) was developed [6]–[10] introducing real-time 3-D ultrasound (RT3D). This modality is fundamentally different from the previous generations of 3-D systems as a true volume rather than a summation or interpolation of 2-D planes is acquired. This system uses a 2.5-MHz matrix-array transducer that consists of 2-D phased arrays and offers steering in both the azimuth and elevation of the beam, permitting interrogation of a pyramidal volume through the body. In each of these 2-D arrays, 512 elements connect to the system, 256 of the 512 elements are used for transmission, and 256 are used for reception. Because the RT3D system has 16×1 parallel processing, the volumetric scanner consists of a 4096 channel-receive system (256 elements×16 parallel processing). The RT3D transducer can accomplish a typical cardiac scan at a rate of 20 volumes/s at a depth setting of 15 cm. Therefore, for a typical heart rate of 60 beats/min, 1 s is required to obtain an entire cardiac cycle with 20 volumes captured from systole to diastole. During a clinical examination, the data is displayed as two adjustable orthogonal B-scans, which are perpendicular to the transducer, and two or three C-scans that are parallel to the face of the transducer [7], [11]. The transducer acquisition volume and clinical display interface tool (FreeScan from Echotech 3D Imaging Systems) are presented in Fig. 1. All scan angles are steerable so that various planes within each of the volume sets can be reviewed on screen while the data is recorded on a computer. Clinical studies on cardiac function require an accurate method to extract myocardial boundaries from 3-D echo data sets. Several approaches for epicardial and endocardial border

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Recent publications in the literature on 3-D segmentation of 4-D freehand (first-generation) and rotational (second-generation) ultrasound include the work of Salustri et al. [26], Rohling et al. [27], Treece et al. [28], Ofili et al. [29], Noble et al. [30]. Stetten et al. developed a medial-node model for rapid identification and measurement of objects in RT3D ultrasound data [31].

Our spatiotemporal analysis tool is based on phase information rather than intensity. This strategy is best suited for denoising and cardiac boundary enhancement as first suggested by Noble et al. [32] with directional Gabor filters. In this paper, we use a new set of basis functions called brushlets, introduced by Coifman and Meyer in 1997 [33].

Spatiotemporal analysis of 4-D cardiac volumes with brushlet basis functions provides projected coefficients that are associated with distinct “brush strokes” of a particular size and orientation in four dimensions (3-D volume + Time). Brushlet coefficients may be modified in the transform domain via resetting of high frequency and nonlinear enhancement of low frequency coefficients.

In 1996, McInerney and Terzopoulos published an extensive survey of the use of deformable models in medical image analysis [34]. More recently, several papers on cardiac echocardiography with deformable models have been published. Chalana et al. [32] used active contours to detect both epicardium and endocardium borders on echocardiographic images, Papademetris et al. [35] used integrated deformable models to segment in vitro ultrasound of dog hearts acquired with a rotational-probe; Montagnat et al. [36] used 3-D deformable models with constrained deformations on cylindrical echocardiograms. In this study, segmentation was performed on denoised volumes, after enhancement and denoising in the transform domain. A 2-D deformable model performed segmentation of the endocar- dial borders and LV volumes were then reconstructed for quantitative measures. This segmentation was tested and validated on phantom and clinical RT3D data sets. Quantitative measures of the extracted volumes are reported for both validation studies.

The remainder of this paper is organized as follows. Section II describes the methodology for multidimensional space–frequency analysis with brushlet functions, nonlinear denoising-enhancement schemes and a deformable-model segmentation algorithm. Section III first presents results on denoising performance of the brushlet analysis and then describes quantitative segmentation results obtained from phantom and clinical RT3D ultrasound data sets. Section IV discusses the quality of analysis and segmentation. Section V concludes with an emphasis on future work for validation and possible enhancements of this research.

II. METHODS

A. Multidimensional Space–Frequency Analysis

Speckle noise corrupts ultrasonic data by introducing sharp changes in an image intensity profile, while attenuation alters the intensity of equally significant cardiac structures. These properties introduce inhomogeneity in the spatial domain and suggest that measures based on phase information rather than intensity profiles are more suited for analysis of cardiac volumes. Brushlet functions are a new family of steerable wavelet...
packets based on the expansion of the Fourier transform (FT) of a signal onto windowed complex exponential functions. These functions, first introduced by Coifman and Meyer [33] for compression of highly texturized images, are well localized in both time and frequency. However, we point out that the goals of compression are completely opposite to this application. We modified this original expansion to provide an overcomplete (redundant) representation. Such intentional redundancy has been shown beneficial for denoising applications [37]–[41].

1) Definition of Brushlet Basis Functions in One Dimension: An initial goal of the construction is the desire to build an orthonormal basis of transient functions with good time–frequency localization. For this purpose, windowed complex exponential functions for Lapped orthogonal transforms have been used for many years in the context of sine-cosine transforms [42], [43] and Gabor functions. Following the Balian–Low theorem there is no differentiable and compactly supported window \( g \) such that: \( \forall t \in \mathbb{R} \) and \( \mathcal{C}_0 \), \( \{ g(t - m) e^{j2\pi ft} \}, m \in \mathbb{Z} \) forms an orthonormal basis of \( L^2(\mathbb{R}) \). In order to overcome this limitation while working with complex exponentials, Meyer and Coifman introduced a division of the real axis into subintervals \( [a_n, a_{n+1}] \), along with a new set of analysis basis functions

\[
\begin{align*}
    & u_{jn}(x) = b_n(x - c_n) e^{jn(x)} + \nu(x - a_n) e^{jn(2a_n - x)} - \nu(x - a_{n+1}) e^{jn(2a_{n+1} - x)}, \\
& \quad \text{if } t \leq -1 \\
\end{align*}
\]

The two window functions \( b_n \) and \( \nu \) are defined from the ramp function

\[
r(t) = \begin{cases} 
0, & \text{if } t \leq -1 \\
1, & \text{if } t \geq 1 
\end{cases}
\]

and

\[
r^2(t) + r^2(-t) = 1, \quad \forall t \in \mathbb{R}.
\]

The function \( \nu \) is defined as

\[
\nu(t) = r \left( \frac{t}{\varepsilon} \right) r \left( \frac{-t}{\varepsilon} \right), \quad t \in [-\varepsilon, \varepsilon].
\]

The function \( b \) is defined on \( [-l_n/2 - \varepsilon, l_n/2 + \varepsilon] \) as

\[
b(t) = \begin{cases} 
{r^2 \left( \frac{t + l_n/2}{\varepsilon} \right),} & \text{if } t \in \left[ \frac{-l_n}{2} - \varepsilon, \frac{l_n}{2} + \varepsilon \right] \\
1, & \text{if } t \in \left[ \frac{l_n}{2} - \varepsilon, \frac{l_n}{2} + \varepsilon \right] \\
{r^2 \left( \frac{t - l_n/2}{\varepsilon} \right),} & \text{if } t \in \left[ \frac{-l_n}{2} + \varepsilon, \frac{l_n}{2} - \varepsilon \right]
\end{cases}
\]

where \( l_n = a_{n+1} - a_n \) and \( c_n = l_n/2 \).

The complex-valued exponential \( e_{jn} \) is defined as

\[
e_{jn}(x) = \frac{1}{l_n} \exp \left[ -2i\pi j \left( \frac{x - a_n}{l_n} \right) \right].
\]

The windowing functions \( b_n \) defined on \( [-l_n/2 - \varepsilon, l_n/2 + \varepsilon] \) and \( \nu \) defined on \( [-\varepsilon, \varepsilon] \) are displayed in Fig. 2. The parameter \( \varepsilon \) controls the degree of localization of the brushlet function and its inverse FT in time and frequency. As illustrated in Fig. 3, the smaller the value of \( \varepsilon \), the better the localization in frequency (smaller second peak) but localization in time becomes less (spread of the central peak). This tradeoff in time–frequency resolution is analogous to the Heisenberg uncertainty principle that constrains wavelet packet resolution and all other methods of time–frequency analysis.

Let us call a given one-dimensional (1-D) signal \( f \), and its FT \( \hat{f} \). We can project \( \hat{f} \) on a brushlet basis

\[
\hat{f} = \sum_n \sum_j \hat{f}_{n,j} u_{n,j}
\]

with \( u_{n,j} \) the brushlet analysis functions and \( \hat{f}_{n,j} \) the brushlet coefficients. By doing so, the FT of the signal is divided into subintervals. Each interval, indexed by \( n \), and of size \( l_n \) is projected onto \( u_{n,j} \), with \( n = 0, 1, \ldots, (l_n - 1)/l_n \). By applying an inverse FT, we can then compute a decomposition of \( f \)

\[
f = \sum_n \sum_j \hat{f}_{n,j} u_{n,j}
\]

on the orthonormal basis \( u_{n,j} \), inverse FT of \( u_{n,j} \). The basis functions \( u_{n,j} \) are expressed as

\[
u_{n,j}(x) = \sqrt{l_n} \exp \left[ 2i\pi n x - \frac{l_n}{2} x \right]
\]

\[
\times \left\{ (-1)^j \hat{b}_n(l_n x - j) - 2i \sin(\pi l_n x) \nu(\pi l_n x + j) \right\}.
\]

We observe here the wavelet-like structure of the \( u_{n,j} \) synthesis functions with scaling factor \( l_n \) and translation factor \( j \).

The major difference between the brushlet basis and wavelet packets is the arbitrary tiling of the time–frequency plane and the perfect localization of a single frequency in one coefficient.

The projection of \( \hat{f} \) on the analysis functions \( u_{n,j} \) can be implemented efficiently by a folding technique and fast Fourier transform (FFT). The folding technique, described by Wick-erhauser [44], consists of folding the overlapping parts of the window function \( b_n \) and the bump function \( \nu \) across the end points of the interval, back inside itself.
If we call \( T_{\alpha_0 \alpha_0+1} \) a folding smooth periodic restriction operator and \( T_{\alpha_0 \alpha_0+1}^* \) its adjoint unfolding operator, we have the following properties for the projection of a signal \( \hat{f} \) on the functions \( u_{j,n} \):

\[
\langle \hat{f}, u_{j,n} \rangle = \langle \hat{f}, T_{\alpha_0 \alpha_0+1}^* c_{j,n} \rangle = \langle T_{\alpha_0 \alpha_0+1} \hat{f}, c_{j,n} \rangle.
\]

(10)

This shows how folding and unfolding operators can be computed efficiently to project a signal on the complex exponential functions with an FFT instead of computing the inner product of the original signal with the brushlet basis functions. The reconstruction can be simply computed in the same manner by an inverse FFT and an unfolding operation.

Brushlet functions constitute an orthogonal basis of a domain divided into subintervals. There are as many basis functions as there are subintervals defining brushstrokes associated with the center frequency of each interval. The resolution of each brushstroke is inversely proportional to the size of the interval. This property is illustrated in Fig. 4, where two examples of pairs of analysis and synthesis brushlet functions associated with two intervals of different length but centered at the same frequency are plotted.

In order to control the oscillations of the brushlet synthesis function \( u_{n,j} \), it is desirable to have a positive FT of the windowing function \( b \). This condition is not compatible with the original construction of the \( u_{n,j} \) functions. Meyer and Coifman then introduced two biorthogonal windowed Fourier bases: synthesis function \( u_{n,j} \) and dual analysis functions \( \tilde{u}_{n,j} \). In order to have \( \tilde{b} \), the FT of \( b \), positive, they relaxed conditions on the ramp function defined in (2)–(3) and defined a new ramp function with the following properties:

\[
r(t) = 0, \quad \text{if } t \leq 1
\]

(11)

and

\[
r^2(t) + r^2(-t) > 0, \quad \forall t \in \mathbb{R}.
\]

(12)

It is important to point out that since the second derivative of \( b \) is:

\[
\nu''(t) = -4\pi^2 \int_{-\infty}^{\infty} \xi^2 \hat{b}(\xi)e^{2\pi i \xi t} d\xi,
\]

the constraint \( \tilde{b} \geq 0 \) implies that \( \nu''(0) = -4\pi^2 \int_{-\infty}^{\infty} \xi^2 \hat{b}(\xi)d\xi \). This verifies that \( \nu''(0) < 0 \) and, thus, the bell function \( b \) is not flat around zero. This condition further imposes that \( \varepsilon = L_n/2 \). In our implementation, we followed the sugestion of Coifman and Meyer and defined

\[
\mathcal{U}(t) = \chi_{[-L/4, L/4]}(\tilde{b}) \mathcal{X}(\tilde{b})(-\mathcal{X}(b)) \chi_{[-L/4, L/4]}(\tilde{b})
\]

(13)

with \( \chi_{[-L/4, L/4]} \) the characteristic function defined on the interval \([-L/4, L/4] \). In this case, \( b \) is compactly supported on \([-L, L] \) and we write

\[
\hat{b}(\xi) = \left( \frac{\sin(\pi \xi)}{\pi \xi} \right)^4 \geq 0.
\]

(14)

The synthesis functions \( u_{n,j} \) are defined as stated in (9) with the cubic spline window function \( b \) and associated bump function \( v \). The dual analysis functions \( \tilde{u}_{n,j} \) are defined in (1) by replacing \( b \) and \( v \) by \( \tilde{b} \) and \( \tilde{v} \) defined with the new ramp function

\[
\tilde{r}(t) = \frac{r(t)}{r^2(t) + r^2(-t)}.
\]

(15)

2) Brushlet Basis Functions in \( n \)-D: We can extend brushlet analysis to \( n \)-D via separable tensor products. For a \( n \)-D signal the following steps are involved in the analysis (expansion).

a) Computation of the \( n \)-dimensional (\( n \)-D) FFT of the signal.

b) Tiling of the Fourier domain with arbitrary subvolume sizes along each dimension.

c) Folding along each dimension,

d) \( n \)-D FFT on each subvolume to project a folded signal on complex exponential terms of the brushlet functions.

Synthesis (reconstruction) of the signal includes:

a) \( n \)-D inverse FFT on each subvolume;

b) unfolding along each dimension;

c) reassemble the subvolumes according to the original tiling of the Fourier domain;

d) \( n \)-D inverse FFT of the signal.

3) Tiling of Fourier Domain in Multiscale Analysis: The brushlet basis offers the possibility to decompose and represent \( n \)-D signals in terms of texture patterns with different orientations and resolutions. The original work of Meyer and Coifman focused on image compression and showed that brushlets are a powerful tool for efficiently representing richly textured images [45], [46]. In this paper, we exploit this oriented-frequency representation to accomplish 4-D volume denoising prior to segmentation. The tiling of the Fourier domain determines the center frequencies of the decomposition and the resolution of the analysis. In other words, the tiling of the Fourier domain determines entirely the structure of the transform domain and can be viewed as a hyper-dimension of the analysis. The choice of the tiling determines at which frequencies the original signal is analyzed and at which resolution it is expanded. This additional dimension in the analysis process is illustrated in Fig. 5. In this figure, we illustrate in 3-D tiling in the FT domain with arbitrary sampling along each direction and the corresponding structure of brushlet coefficients in the transform domain. This flexibility in partitioning the transform domain is extremely powerful as it allows us to accommodate and precisely match the sampling rates of each dimension obtained during acquisition (space spacing in \( \bar{x}, \bar{y}, \bar{z} \), and time). By comparison,
4) An Overcomplete Framework: A problem of particular interest for us is the manipulation of an overcomplete decomposition on brushlet basis. Overcomplete multiscale representations are well suited for image analysis and enhancement, because they avoid aliasing effects introduced by critical sampling [47] and yield a translation-invariant representation. This property is crucial for feature analysis such as edge (surface or volume) enhancement. Indeed, without this translation-invariant property, manipulation of the coefficients for feature extraction in the transform domain or thresholding for selective reconstruction could not be accurately carried out. In our case, the aliasing effects arise from the selection of overlapping subintervals of the Fourier plane expanded into a local Fourier basis. To avoid this and at the same time increase the number of coefficients for the same subinterval size along each dimension, we project onto an extended Fourier basis. The overcomplete projection is efficiently implemented by padding the folded signals with zeros along each dimension and computing its FT. Since padding a signal will increase the resolution of the FT, overcomplete projections increase the number of coefficients for the same interval and, therefore, increase resolution in the transform (coefficient) plane. For an overcomplete projection, each subinterval of the Fourier domain is projected onto a brushstroke of dimension equal to the original number of elements. The orientation and size of the original brushstroke are preserved, as the phase of the brushstroke is not modified. However, the number of points defining each brushstroke in the Fourier domain is increased to match the original size of the volumetric data. Inside a subvolume of the FT, the coefficients are stored in the same manner as the data points in the original signal. Therefore, in the overcomplete case, there is a perfect homomorphism between the location of data points in the original set and the position of the coefficients in each projected subvolume. This bijection (1:1 mapping) introduces redundancy in the transform domain, which results in a dramatic improvement in feature representation in each dimension, as observed in Fig. 6.

We point out that extension of brushlet analysis in four dimensions and overcompleteness of the expansion are two innovations from the initial 2-D decimated implementation. The computational cost added is not prohibitive since the 2-D FT can be efficiently implemented with fast algorithms similar to the 2-D FFT.

B. Nonlinear Modification of Brushlet Coefficients in the Transform Domain

In order to extract features of interest and eliminate speckle noise components, we applied nonlinear thresholding on coefficients in the transform domain, in a similar fashion as previous works in cardiac denoising [48]–[50]. Recall that overcompleteness of the analysis guarantees that no distortion is introduced and a direct correspondence of feature location in both the spatial and transform domains exists. Because of the high frequency nature of the speckle noise and the absence of correlation in space and time we chose to decompose each dimension into four equal length intervals to divide the frequency spectrum of the signal into low frequencies (two inner intervals centered around the zero frequency) and high frequencies (two outer intervals). For an n-D signal we apply a 4\(^n\) tiling of the Fourier domain (four subintervals/dimension) and extract only the 2\(^n\) inner subvolumes centered around zero frequency. These subvolumes encode only the low frequency components of the signal along 45° diagonal directions in n-D. Since we wanted to detect rather isotropic endocardial and epicardial borders, we did not privilege any specific direction with tiling of the Fourier domain. Denoising in each subquadrant was performed via hard thresholding of the coefficients at 25% of maxima in the spatial domain and soft-thresholding at 15% in the time direction. These levels were selected via experimentation with a judicious trade off between attenuation of noise components and enhancement of directional features. Reconstruction was then performed with these thresholded coefficients and segmentation, described next, was then applied on the resulting reconstructed volumes.

C. Deformable-Model Segmentation

Segmentation of anatomical (and functional) structures remains in practice the most time consuming part of modeling a dynamic process. Automatic segmentation of anatomical structures in medical images remains an open problem. In clinical practice, manual segmentation is often considered the most reliable technique (i.e., the gold standard). Efforts by researchers in this field have shown the advantage of deformable models and templates coupled with object parameterization, such as those reported by McInerney and Terzopoulos in a recent survey [51].
Along these lines, techniques based on the use of deformable models for reconstructing 3-D surface shape and motion of the left ventricle from computed tomography and magnetic resonance imaging data have been developed [51], [52]. Our approach more closely follows the work of Metaxas [53] and uses a balloon deformable model first introduced by L. Cohen [54].

The class of deformable models originated with the introduction of the “snake” by Kass, Witkin, and Terzopoulos [55] and has been widely used to detect smooth contours in 2-D images. Deformable models were then applied for 21/2-D and 3-D object extraction from 2-D slice images [56].

A “snake” can be viewed as a curve that deforms under the influence of internal and external forces. Internal forces depend on the snake’s intrinsic properties such as its elasticity and curvature. External forces are derived from image properties (intensity gradient and edge maps).

Let a curve \( v \) be defined on \( \Omega = [0, 1] \) through the following mapping:

\[
\Omega \rightarrow \mathbb{R}^2 \\
v \mapsto v(s) = (x(s), y(s))
\]  

(16)

We define a snake as the space of admissible deformations that minimize the functional \( E \)

\[
E(v) = \int_{\Omega} \alpha ||\v' ||^2 + \beta ||\v'' ||^2 + P_{\text{ext}}(\nu(s)) \, ds
\]  

(17)

with \( \alpha \) and \( \beta \) weighting parameters that control respectively the elasticity and rigidity of the snake and \( P_{\text{ext}} \) the potential of external forces, derived from the image edges. Let \( v \) be a local minimum of \( E \), the associated Euler–Lagrange equation verified for \( v \) is

\[
\begin{cases}
- \alpha \nu'' + (\beta \nu')'' + \nabla P_{\text{ext}}(v) = 0 \\
\text{for some } \nu(0), \nu'(0), \nu(1), \nu'(1).
\end{cases}
\]  

(18)

The first two terms represent the internal force that impose regularity to the curve, and the last term represents the potential of the external force that attracts the curve to features of interest. There are several approaches for defining this potential. In order to attract the snake to minima of the potential force, a simple model is to define an edge map

\[
P(v) = -||\nabla I(v)||^2
\]  

(19)

on the image \( I \), or

\[
P(v) = -||\nabla(G_\sigma * I(v))||^2
\]  

(20)

with \( G_\sigma \) a Gaussian kernel of standard deviation \( \sigma \). Even though the convolution of the image with a Gaussian kernel blurs the edges it has the effect of increasing the capture range of an active contour so that the initialization does not need to be close to the correct position to converge. Cohen and Cohen [57] modified the traditional gradient-based potential force to produce more stable results. They proposed to define an external force as

\[
F = k_2 \bar{n}(s) - k \frac{\nabla P(\nu(s))}{||\nabla P||}
\]  

(21)

where \( \bar{n}(s) \) is the unit vector normal to the snake curve at point \( \nu(s) \). The first term represents an internal pressure force that inflates the snake as a "balloon." This prevents the curve from shrinking or being trapped by spurious isolated edges and makes the minimization process less sensitive to the initialization. The second term normalizes the potential force to simulate a local time step. This applies the same speed to every point in the snake to overcome issues related to the selection of a common time step for every snake point. Coefficients \( k \) and \( k_2 \) are chosen of the same subpixel order with \( k > k_2 \) to let the snake stop at edge locations. In our work, in order to keep implementation simple, the "balloon" snake was programmed using a finite difference approximation scheme as in [58] and [59] with similar parameter values as suggested in [59]. However, rigidity was increased to diminish the time step of the dynamic equations and ensure the smoothness of a contour in situations where part of the LV wall was missing (as was often the case in our data). The edge map was defined as the gradient of the image blurred with 2-D Gaussian filter of standard deviation \( \sigma = 2 \). Three-hundred iterations were run for each short axis slice in the transform domain.

III. RESULTS

A. Comparison of Denoising Performance in Three and Four Dimensions

To quantitatively evaluate the performance of dynamic 4-D analysis in decorrelating noise components in low spatial frequencies, we first experimented with mathematical phantoms corrupted with white or speckle noise. The phantom consisted of an ovoid volume growing in time that schematically mimicked the aspect of the left ventricle with an inner gray cavity surrounded by a thick white wall on a black background. The size of a single volume matrix was \( 64 \times 64 \times 64 \) and there were 16 volume “snap shots” growing in time. The volume size increased by 70% over 16 time frames, similar to the average ejection fraction in normal patients. We corrupted the volumes with 1) additive uniform white noise; 2) multiplicative speckle noise derived from uniform distribution; and 3) multiplicative speckle noise derived from Rayleigh distribution. All noisy signals were constructed with a SNR equal to \(-15 \) dB. Corrupting noise is commonly modeled as Gaussian white noise. Ultrasound noise has been extensively studied in the literature. Because the ultrasound signal acquired by a transducer is not the raw wave signal itself but the envelope of the complex radio frequency signal, it can be shown that the initial Gaussian random noise is transformed into a signal dependent noise that follows a Rayleigh distribution. We have tested this hypothesis of Rayleigh distributed noise in data acquired with RT3D transducer and we found that it was indeed verified with good agreement. Brushlet analysis was performed with four subintervals in each direction. Denoising was performed via resetting of the higher frequency components and hard-thresholding of the lower frequency coefficients at 15% of coefficient maxima in each 3-D or 4-D subvolumes. In order to compare denoising performance to methods of reference we also performed denoising with Wiener filtering. Wiener filtering is optimal for denoising of additive white noise in the mean square error sense. We further adapted the filter to use variance...
estimated from a Rayleigh noise model as presented in [61] to improve the performance on noisy data corrupted with speckle noise. A single slice from the original volumes, noisy volumes and denoised volumes processed with the four different methods is displayed in Fig. 7 for white noise, Fig. 8 for speckle noise derived from uniform distribution and Fig. 9 for speckle noise derived from Rayleigh distribution. Detailed results for the three different types of noise are summarized in Table I and commented below.

- In the case of additive white noise, the SNR of the noisy volume was $-15$ dB. In Fig. 7, denoised results are displayed after brushlet analysis in 3-D and 4-D, and Wiener filtering with white and Rayleigh noise models. Visually, 4-D brushlet analysis clearly performed better at denoising the data. Quantitatively, we measured a SNR improvement of 51% with Wiener filtering using white noise model and 43% with 4-D brushlet analysis. Quantitative SNR improvement with 3-D brushlet analysis and Wiener filtering for speckle noise was negligible.

- In the case of multiplicative speckle noise derived from a uniform distribution, the SNR of the noisy volume was again $-15$ dB. In Fig. 8, denoised results are displayed after brushlet analysis in three and four dimensions, and Wiener filtering with white and Rayleigh noise models. Visually, 4-D brushlet analysis clearly performed better at denoising the data. Quantitatively, significant SNR improvement of 55% was achieved by 4-D brushlet analysis while the other methods only achieved negligible improvements.

- In the case of multiplicative speckle noise derived from a Rayleigh distribution, the SNR of the noisy volume was again $-15$ dB. In Fig. 9, denoised results are displayed after brushlet analysis in three and four dimensions, and Wiener filtering with white and Rayleigh noise models. Visually, 4-D brushlet analysis clearly performed better at denoising the data. Quantitatively, significant SNR improvement of 55% was achieved by 4-D brushlet analysis while the other methods only achieved negligible improvements.

B. Segmentation for Volume Extraction

Segmentation was executed on two types of data: in vitro RT3D contrast echocardiogram phantoms, and in vivo clinical RT3D volume data sets.

1) In Vitro Contrast Echocardiogram Phantoms: In vitro contrast echocardiogram phantoms consisted of balloons filled with human albumin. A layer of bubbles was artificially created on the surface of the internal balloon wall to mimic the appearance of the white myocardium and increase the contrast at the wall interface. Ultrasonic RT3D data was acquired with a setup identical to clinical conditions. The phantom database used in this study consisted of two balloons of 35.0 ml and 65.0 ml. Each RT3D volume had a size of $64 \times 64 \times 258$ voxels. Since the volume spanned by the transducer beam is conic, voxel position is determined in cylindrical coordinates. Each sample slice in the $z$ direction (depth) was separated by 0.308 mm. For each slice, $64 \times 64$ points were recorded spanning a total
azimuth and elevation angles of 64° with 1° increment. With a total depth of acquisition of 7.92 cm, the optimal resolution in the short axis direction (perpendicular to the transducer) was 1.736 mm/pixel.

Brushlet analysis was performed in three dimensions with a tiling of the Fourier domain partitioned by $4 \times 4 \times 4$ cubes as described previously in Section II. Denoising was performed in space via hard-thresholding of the lower-frequency coefficients as described in the previous paragraph. Segmentation was carried out using deformable-model on each slice. The snake was initialized with a four-pixel-radius circle inside the cavity to segment. The center of the circle was identified by a circular Hough transform applied on the edges extracted with a Prewitt filter for every ten slices of the volume. For each inspected slice, if more than one circle was detected, no center point was selected and the next slice inspected. In this fashion, for every group of ten slices we either selected one center point or none. We then computed parameters of the best linear fit in the least square sense over the entire set of center points detected. We, thus, obtained two linear models $x = a_2 + b$ and $y = a_2 + b$ to determine the $x$ and $y$ coordinates of the center points for each slice $z$. The circular Hough transform has been applied previously to RT3D data by Stetten et al. [62], [63] for volume measurements on balloon phantoms. Because of the high incidence of spurious edges in ultrasound data, a simple Hough transform on single slices might detect several different circles. Missing edges on the other hand can lead the Hough transform to shift the position of the center of the circular cavity. The linear best fit on the series of center points was observed to be well suited to handle these errors and provided a more robust initialization than a single slice Hough transform.

Parameters values for the active contour were set to: $\alpha = 0.2$, $\beta = 1$, $\gamma = 2$, $K = 3$, $k_3 = 0.2$, and $\sigma = 2$. An expert cardiologist performed manual segmentation of the inner balloon wall on the original volumes. Table II reports volume measures for manually traced balloon and deformable model segmentation. Examples of segmentation of the inside cavity of the two balloons are displayed in Fig. 10 for two slices.

In both cases, there is a consistent underestimation of volume estimation when using the deformable model while manual tracing estimations were within 5% of accuracy. This difference can be explained by the fact that the expert cardiologist knew that the real balloon wall borders were actually located inside the thin layer of bubbles that was inserted to increase the contrast. From its construction and the constraints applied to it, the active contour is forbidden to go inside the white-appearance bubble layer, so that the final contours always underestimated the real balloon cavity. Testing on the balloon data sets allowed us to tune the deformable-model parameters on denoised data leading to a stable behavior in situation where part of the wall borders is missing. This situation is illustrated in Fig. 10.

2) In Vivo Clinical RT3D Volume Data Sets: A database of six patients co-screened with cardiac functional MRI was used for validation of LV volume segmentation and measure. Clinical data sets consisted of either: 21 time frames of size $(64 \times 64 \times 373)$ voxels (two cases), 26 time frames of size $(64 \times 64 \times 438)$ voxels (one case), or 21 time frames containing $(64 \times 64 \times 438)$ voxels (three cases).

Brushlet analysis was performed in three dimensions for spatial denoising followed by 1-D analysis along time for enhance-
ment of cardiac structures. Tiling of the Fourier domain was set to four subintervals in each direction. Denoising in space was performed via hard thresholding of the lower frequency components at 25% of coefficient maxima. Enhancement was then performed in time via soft-thresholding at 15% of coefficient maxima. Threshold values in space and time were empirically set via testing on clinical data sets to achieve a good tradeoff between removal of noise artifacts and enhancement of cardiac structures.

Rescaling of the volumes from spherical coordinates to Cartesian coordinates via linear interpolation was performed prior to segmentation. Rescaling produces isotropic volumes with identical resolution in each of the three dimensions. The best spatial resolution in the acquired volumes is along the long-axis with pixel size of 0.308 mm with over 400 slices. Because of the computational time needed for the segmentation process, whether manually or with the deformable model, we decided to downsample the rescaled volumes. We initially downsampled by a factor of four but manual tracing ended up too far off from the correct values. We finally opted for a downsampling by a factor of two, reaching an acceptable compromise between accuracy of tracing and processing time. On average, 100 slices were segmented for each volume with a voxel resolution of $2 \times 0.308^3 = 0.23 \mu l$.

End-diastole and end-systole time frames were identified and manually segmented by an expert cardiologist. Manual tracing was performed on the unprocessed data while deformable-model segmentation was performed on the denoised data. The deformable model was initialized with a five-pixel-radius circle inside the cavity to segment. An operator manually selected the centers of the circle for every ten slices since the circular Hough transform applied to the balloon data sets failed to produce accurate center estimations in this case. Center locations for in-between slices were determined via linear best fit. By identifying the best linear fit to the center points detected inside the LV cavity we defined an approximation of the LV medial axis. Parameters values for the active contour were set to: $\alpha = 0.07$, $\beta = 2$, $\gamma = 1$, $K = 2$, $k_2 = 0.09$, and $\sigma = 2$. Examples of LV endocardium segmentation via manual tracing and deformable model are displayed in Fig. 11 for one short-axis and one long-axis view. After segmentation, the LV cavity was reconstructed to compute end-diastolic volume (EDV) and end-systolic volume (ESV). An example of LV volume reconstruction is also provided in Fig. 11.

Ejection-fraction (EF) was then computed as

$$ EF = \left( \frac{EDV - ESV}{EDV} \right) \times 100, \quad (22) $$

The six patients were co-screened with cardiac functional MRI, which is considered as the method of reference in this study. An expert cardiologist performed manual tracing on the MRI data. Quantitative measures and errors for the six clinical cases are reported in Tables III–V.

Absolute errors of quantitative measures were computed for volumes and ejection-fraction. We evaluated mean-error values and standard deviation over the six cases for the three quantitative measures. These results are reported in Table IV. Maximum and minimum error values are provided to better assess the range of variability achieved by the different segmentation methods. We also report in Table V percentage mean error for volumes and ejection-fraction measures over the six cases. We observed
with these results a significant improvement of quantitative accuracy with the deformable model when comparing to manual tracing. EDV accuracy was improved by 50.1%, ESV accuracy was improved by 60.6%, and ejection fraction measure accuracy improved by 61.4%.

IV. DISCUSSION

This paper presented a method for feature extraction from RT3D ultrasound that combines 4-D directional denoising and model-based segmentation.

Denoising of ultrasound data was performed via space–frequency brushlet analysis of the 4-D Fourier domain and thresholding of the coefficients to remove speckle noise and enhance coherent anatomical structures in space and time. Brushlet analysis identifies efficient tilings of the Fourier domain, along each dimension of a signal. Through characterization of coefficients within sets of redundant articulated (orientation rich) bases we can separate signal and noise components. The performance of brushlet analysis to decorrelate signal from white additive and multiplicative speckle noise components was demonstrated on a 4-D mathematical phantom. These examples also showed the superiority of 4-D brushlet denoising over 3-D for 4-D data sets. We have previously shown that we can characterize and isolate features of interest in LV volumes by selection of specific brushlet coefficients [64]. Since decomposition on a brushlet basis can efficiently isolate directional features at specific frequencies, preprocessing of RT3D volumes via thresholding of lower frequency brushlet coefficient can assist segmentation by removing noise components and enhancing anatomical features. The RT3D clinical data sets suffer from very low spatial resolution due to subsampling by the transducer during acquisition of the echo, the high level of speckle noise and the motion of the cardiac muscle. These three factors diminish the resolution of the acquisition and create artifacts that can either remove parts of the myocardial wall or introduce “myocardium-like” bright signals inside the LV cavity or outside the ventricle. Since the temporal resolution is much better compared to the three spatial dimensions, the inclusion of time in the analysis of the data was a tremendous advantage. It helped to remove artifacts not persistent in adjacent time frames and added myocardial components (fill in holes) when the cardiac surface was present in adjacent time frames. We qualitatively compared denoising performance on clinical data using 3-D or 4-D brushlet analysis and Wiener filtering adapted to speckle noise. Results for the different denoising schemes are illustrated in Fig. 12 for four long-axis and four short-axis slices extracted from one clinical data set. It is interesting to note that Wiener filtering produced a high quality denoising in long-axis views but introduced significant blurring that corrupted the data when displayed in short-axis views. Blurring of the data, which is a common artifact with denoising filters, prevents segmentation from realizing accurate volume quantification. Such blurring was not observed with brushlet expansions. When comparing 3-D and 4-D brushlet denoising we observed that the introduction of the temporal dimension improved the contrast of the denoised data with enhanced myocardial wall and a more homogeneous inside LV cavity. We observed that segmentation performance was enhanced by such contrast improvement as the deformable model was not trapped in local minima inside the cavity, moved faster toward the myocardial border, and localized border edges more accurately. The main power of brushlet analysis is its flexibility in decomposing heterogeneous signals and its ability to accommodate and precisely match nonuniform sampling rates of each dimension obtained during RT3D acquisition (sample spacing in z, y, z and time). The tiling of the Fourier domain determined the orientation and the resolution of the brush strokes on which the signal was projected. In different terms, the tiling selected the textural patterns used for the analysis of the original signal. From this point of view, tiling can be considered as an added dimension of the analysis. As described in Section II, brushlet analysis was performed for a fixed tiling of the FT that divided each dimension into four intervals. This choice allowed us to extract and denoise only the lower frequency components without selecting any specific direction. This approach was originally motivated by the isotropic nature of the LV myocardium shape in space and identification of low-frequency movements in time (i.e., slowly moving structures). The next step is to investigate mathematical and empirical methods for identifying the most “efficient” brush strokes sizes and orientations for decomposing and reconstruction. From an empirical point of view, the shape of the left ventricle is highly isotropic in short axis planes (circular wall) while rather elongated in long-axis views. This observation suggests that a selective tiling for horizontal features might be more appropriate in the z dimension. A mathematical framework for identifying an optimal basis was introduced by Coifman and Wickerhauser [65] for the wavelet packets and cosine transforms. By measuring a predefined cost function, such as entropy, the algorithm selects an optimal tiling of the Fourier domain where the energy of the signal is “best” decomposed. Meyer and Coifman [33] applied this algorithm on brushlet functions for compression of highly textured images. In the current application, optimal speckle noise removal is desired and a cost function adapted to denoising performance should be designed to carry out an optimal-basis search. We presented some preliminary work in that direction [61] with qualitative evaluation of denoising quality as a function of the tiling. However, optimization of the tiling needs to be studied further and would enable us to systematically adjust specific brushlet coefficients for noise reduction and selective reconstruction of more salient cardiac temporal features. Computation time for the 4-D brushlet analysis applied to clinical data sets will also need to be optimized for future real-time applications. Currently, the whole process of analysis and segmentation takes less than one hour, running on a Pentium II 400 MHz under Matlab.
Regarding volume quantification using RT3D, we showed that the deformable model could accurately segment balloon phantom data with consistency in observed behavior. Quantitative measures obtained from manual tracing were biased as detailed in Section III such that validation of volume measurements could not be obtained on this data.

Testing of volume quantification using a database of six patients showed that simple 2-D deformable-model segmentation applied on denoised data improved by over 50% the accuracy of volume and ejection fraction measures when compared to manual tracing by an expert cardiologist. The gold standard used in this clinical study was functional cardiac MRI. Takuma et al. [66] reported an inter-observer variability of 8.3% and intra-observer variability of 3.7% for LV volume estimation with parallel-plane-disks summation on RT3D echocardiograms. These values set the variability limits for acceptability of automatic volume extraction. Any segmentation method should achieve this range of accuracy in order to be used in clinical settings. As reported in Table V, mean percentage errors for volumes and ejection fraction measures are higher than these values for manual tracing while they fall below 8.7% with deformable-model-based segmentation.

These results are quite encouraging and suggest that a deformable model applied to denoised RT3D ultrasound could be used in clinical practice for rapid quantification of cardiac function. To fully validate our methods we need to carry out further testing on a larger database of patients to evaluate the clinical significance of these results. We are currently working on the implementation of a deformable model in three dimensions following the work of Jones and Metaxas [67]. Three-dimensional deformable surfaces applied to medical image segmentation have drawn a lot of interest in recent years. A good survey of the different approaches and implementations developed for these surfaces is provided in [68]. We believe that extending the segmentation algorithm to true three dimensions will have a great impact on the quality and the robustness of the extracted volumes. When using 2-D deformable models there are cases where the model is stopped by false edges inside the cavity and does not fully expand to reach the actual myocardium wall border. This can be avoided with 3-D deformable models whose deformations are controlled in 3-D space, ensuring smoothness of the contours in every dimension.

After building a 3-D deformable model, we plan to further adapt the method for segmentation of the epicardium borders. A long-term goal of this research project is to extract the myocardium wall borders and analyze wall deformation in time as studied during stress-echo examinations. As true volumetric data is acquired in time, usual post processing tasks such as interpolation for volume reconstruction and registration of volumes from different times is not required, leaving segmentation
as the final step to realize a patient-specific dynamic model of the left ventricle. This property makes RT3D ultrasound a very strong candidate for becoming a powerful echocardiographic screening modality that would enable fast and accurate computation of dynamic computer heart models for standard screening, patient monitoring and stress-echo testing.

V. CONCLUSION

This paper presented a spatiotemporal analysis method for feature extraction using a third-generation RT3D ultrasound system. The challenge of developing a segmentation tool for quantification of cardiac function from RT3D ultrasound lies in the novelty of the data itself and the wealth of dynamic information that can be extracted despite low spatial resolution and high levels of speckle noise. Overcomplete wavelet-like brushlet functions were used to analyze the multidimensional FT of echocardiographic data and perform denoising and enhancement on low-frequency directional components. Speckle noise was decorrelated well from the original signal and anatomical structures were enhanced at the same time via nonlinear thresholding of analysis coefficients. The deformable-model based segmentation method was performed on the denoised data in two dimensions.

We illustrated the superiority of 4-D (3-D + Time) over 3-D analysis in decorrelating white and speckle noise from 4-D phantom and clinical data. We have shown that we can extract LV endocardial borders using 2-D deformable models and quantify volumes of interest with a higher degree of accuracy than those achieved with manual tracing. Future work will extend the segmentation process itself to higher dimensions to integrate the continuity of the echo data in space and time. Finally, we plan to investigate mathematical and empirical methods for identifying the most “efficient” brushlet analysis tiling for denoising and enhancement.

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REFERENCES


