Automatic Localization of the Left Ventricle from Cardiac Cine Magnetic Resonance Imaging: A New Spectrum-Based Computer-Aided Tool

Liang Zhong^{1,2}, Jun-Mei Zhang¹, Xiaodan Zhao¹, Ru San Tan^{1,2}, Min Wan^{1,3}*

1 Bioengineering Department, National Heart Centre Singapore, Singapore, Singapore, 2 Cardiovascular & Metabolic Disorders Program, Duke-NUS Graduate Medical School Singapore, Singapore,

Abstract

Traditionally, cardiac image analysis is done manually. Automatic image processing can help with the repetitive tasks, and also deal with huge amounts of data, a task which would be humanly tedious. This study aims to develop a spectrum-based computer-aided tool to locate the left ventricle using images obtained via cardiac magnetic resonance imaging. Discrete Fourier Transform was conducted pixelwise on the image sequence. Harmonic images of all frequencies were analyzed visually and quantitatively to determine different patterns of the left and right ventricles on spectrum. The first and fifth harmonic images were selected to perform an anisotropic weighted circle Hough detection. This tool was then tested in ten volunteers. Our tool was able to locate the left ventricle in all cases and had a significantly higher cropping ratio of 0.165 than did earlier studies. In conclusion, a new spectrum-based computer aided tool has been proposed and developed for automatic left ventricle localization. The development of this technique, which will enable the automatic location and further segmentation of the left ventricle, will have a significant impact in research and in diagnostic settings. We envisage that this automated method could be used by radiographers and cardiologists to diagnose and assess ventricular function in patients with diverse heart diseases.

Citation: Zhong L, Zhang J-M, Zhao X, Tan RS, Wan M (2014) Automatic Localization of the Left Ventricle from Cardiac Cine Magnetic Resonance Imaging: A New Spectrum-Based Computer-Aided Tool. PLoS ONE 9(4): e92382. doi:10.1371/journal.pone.0092382

Editor: Jeroen Hendrikse, University Medical Center (UMC) Utrecht, Netherlands

Received June 6, 2013; Accepted February 21, 2014; Published April 10, 2014

Copyright: © 2014 Zhong et al. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Funding: This work was supported by the National Research Foundation (NRF2011NRF-POC001-068), the National Medical Research Council (NMRC/EDG/1037/2011), and SERC Biomedical Engineering Programme Grants 1211480003 and 1321480012. The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

Competing Interests: The authors have declared that no competing interests exist.

* E-mail: wanmin1983@gmail.com

Introduction

Cardiovascular Disease (CVD) is currently the leading cause of death, killing 17.3 million people worldwide each year. This figure represents one-third of total death, a proportion that is still increasing. It is estimated that by 2030, CVD will kill 23.6 million people annually [1]. For clinicians, the examination and analysis of medical cardiac images is of great significance for diagnosis and treatment. The reliability of quantitative assessment of cardiac functions such as muscle deformation and ventricular ejection fraction is dependent on the precision and correctness of the heart chamber segmentation [2].

Usually, this segmentation task is performed manually via software assistance, which is quite time-consuming. One trained clinician can delineate one image in half a minute, and it would take him/her more than two hours to finish the whole data set of 11 slices by 25 frames. Furthermore, the results often present a high intra/inter clinician variability [3]. Over the past decade, automation of this tedious yet significant procedure has received a lot of attention from not only medical imaging but also the computer vision community [4–9]. Of the four heart chambers, the imaging of the left ventricle (LV) is of the greatest importance and interest due to its physiological fatality. Of the 70 studies surveyed in Petitjean and Dacher [3], 49 of them address the LV

segmentation problem exclusively, with 18 solving segmentation of both ventricles.

One important stage in the automatic segmentation approach is to locate the left ventricle in the whole image and limit the computation domain within a vicinity of the left ventricle. The localization procedure is sometimes referred to as region of interest (ROI) identification. This procedure not only decreases the computational load, but also provides an initial guess for deformable segmentation approaches. Previously, it was preferred that the ROI identification be done manually by selecting the center of the ROI [10–12] or drawing a bounding contour. To make the whole segmentation procedure automatic, it will be necessary to automate LV localization as well.

The existing approaches for the automatic localization of the LV can be generally categorized into two groups. One group is pattern recognition approaches [13], which require a learning stage. The desired "donut" pattern is decomposed into a multidimensional feature vector, which is further modelled as a Markov process. The system is trained using both positive and negative examples identified manually. The trained system is utilized to classify the image pixels as the LV boundary [14–17]. The whole procedure is a specific application of pattern recognition, the success of which is dependent on a proper training examples and the studied cases.



(a) variance image

(b) \bar{u}_1 image

Figure 1. The imperfect detection. doi:10.1371/journal.pone.0092382.g001

The other group is temporal dimensional approaches. These methods are based on the fact that the heart is the only organ with substantial motion in the field of view (FOV). In an ideal cine sequence, the image is stationery aside from the heart. The variance image is calculated from a 2 D+t original image sequence. The high intensity in the variance image corresponds to the vigorous movement in the original image, which usually happens at the endocardium and epicardium of both ventricles as well as at the septum. A circle Hough transform [18] is used to detect the potential circle approximating the endo/epi-cardium of the LV. Interested readers are referred to previous resources [19-23]. One alternative approach based on Fourier analysis has previously been proposed [24,25].Instead of using the variance image to recognize the motion, the first Harmonic image is used. The circle Hough detection procedure is identical to that of the variance image approaches.

Compared to the pattern recognition methods, the temporal approaches do not require a large training set, which is also a tedious task. They also exploit the information in between the frames. However, both the variance image approach and Fourier analysis approach lack robustness and precision in locating the LV. The LV is not the only moving chamber in the FOV. The variance or the first harmonic image contains much more than the LV boundary and this could hardly be extinguished. The movement of the right ventricle (RV) would unavoidably lead to imprecise or even incorrect circle detection and ROI identification. A more detailed analysis and some examples are provided in Section 2.

In this study, short axis cardiac images at the basal location were prepared for ten normal volunteers. Harmonic images of all orders were examined visually and quantitatively. Two observations were seen regarding the LV and RV in Harmonic images as follows, based on which had more information inferred from Harmonic images of high order that could be utilized in a sophisticated way:

- In all Harmonic images except the zeroth one, the RV region presents higher brightness than the LV region.
- As the Harmonic order increases, the superiority of the RV to the LV increases.

Two Harmonic images were utilized, the first Harmonic image (\bar{u}_1) and a higher-order Harmonic image (the fifth Harmonic image (\bar{u}_5) in our study). A new circle detector was designed based on the anisotropic weighted circle Hough transform. The \bar{u}_1 image with the most substantial LV visibility was an input of the detector with a positive weight, while \bar{u}_5 was the other input with a negative weight to suppress the RV interruption. This dual input Hough detector could largely eliminate the RV interruption, leading to a robust and precise LV localization.

The remainder of this article is organized as follows. Section 2 gives a brief review on earlier methods and shows their relation to Harmonic images. In Section 3, Harmonic images of all orders are



Figure 2. The variance image and the first harmonic image. doi:10.1371/journal.pone.0092382.g002

Volunteer	height	weight	BSA	LV mass	RR	SBP	DBP	LVEDV
	(m)	(kg)	(m^2)	(6)	(ms)	(mmHg)	(mmHg)	(ml)
-	1.71	68.2	1.8	102	745	115	63	130
2	1.8	97.5	2.2	125	1005	123	72	202
3	1.66	78	1.9	115	780	136	79	125
4	1.56	59.2	1.6	78	765	111	60	100
5	1.53	46.5	1.4	66	830	110	67	101
Q	1.47	47.1	1.4	85	880	117	75	125
7	1.62	66	1.7	120	1085	126	78	151
∞	1.47	55	1.5	67	860	92	55	98
6	1.6	49.8	1.5	82	1000	100	63	136
10	1.74	77.7	1.9	105	985	128	77	149
Average	1.616	64.5	1.69	94.5	893.5	115.8	68.9	131.7
Volunteer	LVESV	LVSV	LVEF	LVMassindex	RVEDV	RVESV	RVSV	RVEF
	(ml)	(m)	(%)	(g/m ²)	(m)	(ml)	(ml)	(%)
-	57	73	56	57	169	86	83	49
2	79	123	61	57	237	110	127	54
3	52	73	59	61	161	87	73	46
4	30	70	70	49	105	40	65	62
5	32	69	68	47	105	40	65	62
6	36	89	71	61	155	69	86	55
7	47	104	69	70	177	78	66	56
ω	32	66	67	45	127	58	69	54
0	70	66	48	57	117	61	56	48
10	58	91	61	54	160	72	88	55
Average	49.3	82.4	63	55.75	151.3	70.1	81.1	54.1
doi:10.1371/journal.pc	one.0092382.t001							

Table 1. Statistics on Subjects.



Figure 3. All frames of a cine image. doi:10.1371/journal.pone.0092382.g003

analysed and two facts concerning the LV and RV are presented. An anisotropic weighted circle Hough detector is then proposed for LV localization. Section 4 shows the experimental results of the proposed localization method. Section 5 concludes the article.

Related Work on Harmonic Images of Cine Magnetic Resonance Imaging

Several LV localization methods based on the variance image have been developed [19–23]. A similar approach based on Fourier analysis has been described [24,25]. In this section, we first define the Harmonic images of a cine image sequence. We then provide a brief review of those earlier studies and a reformulation of them into the Harmonic image framework. At the end of the section, we present their limitations and illustrate these using a concrete example.

Discrete Fourier Transform and Harmonic Images

Given a 2 D+t cine Magnetic Resonance (MR) image u with the resolution of $M \times N \times T$, each pixel position over time presents a

discrete time series, (i.e. $u(m,n,t), 0 \le m \le M - 1, 0 \le n \le N - 1, t = 0, 1, \ldots, T-1$). For the rest of the discussion in this section, we omit the spatial coordinates m,n for simplicity, (e.g., $u(t), t=0,1,\ldots,T-1$).

Discrete Fourier transform (DFT) of u(t) is as follows.

$$U(k) = \sum_{t=0}^{T-1} u(t) e^{-j2\pi \frac{k}{T}t}, \ k = 0, \dots, T-1.$$
(1)

Inverse Discrete Fourier transform (IDFT) of U(k) is as follows.

$$u(t) = \frac{1}{T} \sum_{k=0}^{T-1} U(k) e^{j2\pi \frac{k}{T}t}, \ t = 0, \dots, T-1.$$
 (2)

U(k), k = 0, 1, ..., T - 1 presents the decomposition of the time sequence u(t) in term of frequency. We define the *k*-th harmonic series of *u* as IDFT to the *k*-th frequency component separately as follows,



Figure 4. All Harmonic images of the cine image. doi:10.1371/journal.pone.0092382.g004



Figure 5. The variance image and the first harmonic image. doi:10.1371/journal.pone.0092382.g005

$$u_k(t) = \frac{1}{T} U(k) e^{j2\pi \frac{k}{T}t}, t = 0, \dots, T-1, k = 0, \dots, T-1.$$
(3)

The k-th Harmonic image is defined as the L^2 -norm of the k-th harmonic series.

The LV localization for cine images could be dated to early this century. Lin et al. and Jolly [24,25] used the first Harmonic image \bar{u}_1 to detect LV. Others [19–23] provided similar detection methods based on the variance image. We will first recall the

definition of the variance image and then show how it is related to

the Harmonic images. Let u(t) be the same as in Section 2.1. The

Earlier Methods on Harmonic Images

$$\bar{u}_k = \frac{1}{T} \sum_{t=0}^{T-1} \|u_k(t)\|^2.$$
(4)

$$\bar{u} = \frac{1}{T} \sum_{t=0}^{T-1} u(t)$$
⁽⁵⁾

$$Var_{u} = \frac{1}{T} \sum_{t=0}^{T-1} \|u(t) - \bar{u}\|^{2}$$
(6)

The equivalence between the mean and the direct current (dc) image, (i.e., the 0-th harmonic image) is also obvious:

$$u_0(t) = \frac{1}{T} U(0) = \frac{1}{T} \sum_{t=0}^{T-1} u(t) = \bar{u}$$
(7)

The relation between the variance image and other harmonic images could be derived as well.



Figure 6. Predefined RV and LV region as well as masked images. doi:10.1371/journal.pone.0092382.g006





(a) Histogram of masked RV and LV







(b) Normalize Histogram

As the above derivation shows, the variance image is the sum of Harmonic images of all orders except the zeroth order (i.e., the dc image or the mean image).

Most of the earlier methods based on Harmonic images have one disadvantage, the RV interruption. The Harmonic images reflect the organ motion pattern from the spectrum aspect. Motions of both ventricles are present in he Harmonic images. Hence, the precision of LV detection would be undermined by RV interruption. Such a limitation could be observed in most of the earlier studies that we mentioned [19–25]. Figure 1 shows an imperfect detection using \bar{u}_1 and Var_u .

To tackle the RV interruption, some earlier studies made the assumption that the LV is located around the center of the FOV [26,27]. Based on this assumption, a bell shaped or cone shaped filter centered in the FOV is applied to the detection procedure. Figure 2 shows several options regarding the filter shape. However, the aforementioned assumption is strongly relevant to the scanning operator or the participant. The variability of both would cause the assumption to no longer hold. This limitation motivates our study.



Figure 8. Surface plots of all harmonic images. doi:10.1371/journal.pone.0092382.g008



Figure 9. Histograms of all harmonic images. doi:10.1371/journal.pone.0092382.g009

Methods

In this study, we tested algorithm on 10 healthy volunteers. This study was approved by the Singhealth Centralised Institutional Review Board (CIRB No: 2009/705/C) for human research. All enrolled participants gave written informed consent. The MR data are deposited in hospital and are available for research and education purpose. Cardiac relating measurements for each volunteer were give in Table 1.

Two Observations of the LV and RV in Harmonic Images

In this study, we developed a new LV localization method based on Harmonic images and an anisotropic circle Hough transform. First we examined the different patterns of the LV and RV in Harmonic images. An example containing 20 frame cine MR images is illustrated in Figure 3. The Harmonic images from the zeroth to the ninth order were generated as shown in Figure 4. We gave the first Harmonic image a close examination. The colorrendered visualization of \bar{u}_1 and the surface plot are shown in Figure 5(a) and (b), respectively. The LV and RV regions are circled. We could observe that the left "peak" (RV region) was higher than the right "peak" (LV region). The RV region had a larger number of brighter pixels in \bar{u}_1 .

We quantified the observed result beyond the observation and visual analysis. The LV and RV regions were identified manually in \bar{u}_1 , see Figure 6(a). The masked RV and LV images are shown in Figure 6(b) and (c), respectively. The statistics of these two regions were computed. The histogram of each is shown in Figure 7 along with a normalized version. From the histograms, it could be observed that the LV region had fewer pixels with high brightness. We present the first observation:

Observation 1. In the first Harmonic image, the RV region presents higher brightness than the LV region.

This observation explains the limitation of earlier studies. If the circle Hough detection is carried out upon the first Harmonic image or the variance image, the brighter RV region would interrupt the LV detection. This observation also implies that the detected circle tends to shift toward the RV region (Figure 1).

The observation represents the superiority of the RV to the LV in the first Harmonic image. We next inspected the existence of such superiority in other Harmonic images. Surface plots are shown in Figure 8 and the histograms are shown in Figure 9. The superiority of the RV was evaluated in all Harmonic images except



Figure 10. ARLR of different *thresh* parameter. doi:10.1371/journal.pone.0092382.g010



(a) Isotropic (b) Anisotropic

Figure 11. Two circle Hough transform. doi:10.1371/journal.pone.0092382.g011

the zeroth Harmonic image, i.e., \bar{u}_0 . Observation 1 could then be extended as follows:

Observation 1. In all Harmonic image except \bar{u}_0 , the RV region presents higher brightness than the LV region.

It could also be observed from Figure 8 that the LV peaks diminish quickly as the Harmonic order increases. From the fourth harmonic image, the LV could hardly be observed from either the image or the surface plot. Meanwhile, RV peaks could be observed in the first through the last harmonic images. This trend could also be verified by the diminishing white bars in Figure 9.

To quantize this trend, we defined an index of Average RV to LV ratio (ARLR) as follows. With the RV and LV regions, D_{RV} and D_{LV} , as well as a threshold parameter *thresh*, the average of all observable pixels (brighter than *thresh*) within $D_{RV} \cup D_{LV}$ is calculated. Then the pixels that had a higher intensity than this average within the two regions were counted, respectively. The

ratio of the two count numbers was called ARLR. Mathematically, this index is defined as follows:

$$ARLR = \frac{\sum\limits_{x \in D_{RV}} \mathbf{1}_{\{x > \bar{x}\}}}{\sum\limits_{x \in D_{LV}} \mathbf{1}_{\{x > \bar{x}\}}},$$
(8)

where

$$\bar{\mathbf{x}} = \frac{\sum\limits_{x \in D_{RV} \cup D_{LV}} \mathbf{x} \cdot \mathbf{1}_{\{x > thresh\}}}{\sum\limits_{x \in D_{RV} \cup D_{LV}} \mathbf{1}_{\{x > thresh\}}}.$$
(9)



Figure 12. Edge maps e_1 and e_5 from x^{H_1} and x^{H_5} . doi:10.1371/journal.pone.0092382.g012

Table 2. Anisotropic Weighted Circle Hough Transform.

Inputs					
1.	Edge map e_1 from x^{H_1} ;				
2.	Edge map e_5 from x^{H_5} ;				
3.	Gradient map ∇x^{H_1} ;				
4.	Gradient map ∇x^{H_5} ;				
Algorithm					
1	Initialize A to 0				
2	For each (m,n) in e_1				
3	If $e_1(m,n) > thresh$				
4	For (p,q,r) in <i>A</i> , s.t. $ (p-m)^2 + (q-n)^2 - r^2 \le \epsilon$				
5	$A(p,q,r) = A(p,q,r) + \left \left\langle \frac{(p-m,q-n)}{ (p-m,q-n) }, \frac{\nabla_{\mathbf{x}}^{H_1}(m,n)}{ \nabla_{\mathbf{x}}^{H_1}(m,n) } \right\rangle \right $				
6	End For				
7	End If				
8	End If				
9	For each (m,n) in e_5				
10	If $e_5(m,n) > thresh$				
11	For (p,q,r) in <i>A</i> , s.t. $ (p-m)^2 + (q-n)^2 - r^2 \le \epsilon$				
12	$A(p,q,r) = A(p,q,r) - \left \left\langle \frac{(p-m,q-n)}{ (p-m,q-n) }, \frac{\nabla_{\mathbf{X}}^{H_{\mathbf{S}}}(m,n)}{ \nabla_{\mathbf{X}}^{H_{\mathbf{S}}}(m,n) } \right\rangle \right $				
13	End For				
14	End If				
15	End If				
Outputs 1.	Parameter Space A				
Abbreviations:	x ^{H1} , Harmonic image 1;				
	x^{H_5} , Harmonic image 5;				
	A, 3D parameter space;				
	(<i>m</i> , <i>n</i>), spatial coordinates;				
	(p,q,r), parameter coordinates;				
	s.t., such that;				
	thresh, threshold				

doi:10.1371/journal.pone.0092382.t002



(a) A better detection than



(b) The assigned ROI

Spectrum-Based Localization Tool of Left Ventricle

The plot of ratio against harmonic order is shown in Figure 10. A rising trend could be found from the curve for different threshold parameters. From this comes our second observation:

Observation 2. As the Harmonic order increases, the superiority of the RV to the LV increases.

From the visualization and quantitative analysis in the previous subsection, we could draw the conclusion that the RV motion involves more high frequency motion than does the LV motion. Regarding the observation for this special cardiac motion pattern, we could provide the following explanation. Covered by the supporting pericardium, the heart lies close to the middle of the breast-bone. The heart position is oblique. The inferior wall of the LV lies against the diaphragm, while the lateral wall and the anterior wall of the LV lie against the left lung. In contrast, the RV has less support. The superior and anterior position of the RV relative to the LV results in less support from the diaphragm and no support from the lungs. When pumping, the LV moves with the everlasting resistance of the stationary lungs and diaphragm (when the participant holds his/her breath). This causes the selfoscillation frequency of the LV to decrease, thus lowering the high frequency portion. In contrast, the RV motion contains more high frequency motion due to it being relatively free of burden.

Anisotropic Weighted Circle Hough Transform

In this subsection, we first introduce the classic circle Hough transform. We next present two modifications on the algorithm, as well as their advantages.

Classic circle hough transform. Given an edge map derived from an image containing potential circles, the classic circle Hough transform is described as follows. A parameter space is constructed at the same size as the original image. Each pixel in the parameter space has an accumulator, all of which are initialized to zero. Provided an estimate of the radius, R, and a pixel x in the edge map, with an intensity larger than the threshold, one is added to all accumulators that intersect with the circle centered at x with a radius of R. After completing this voting procedure with respect to every point on the edge map, the highest accumulator is considered to be the center of the detected circle with radius R.

When a good estimate of the radius R is not available, an interval of possible radius is provided instead. There are usually two options to adapt the circle Hough transform. 1) Repeat the above voting procedure with respect to every R in the interval. After all loops of voting, the final winning accumulators suggest the circle center with the radius of the mean of the interval. 2) Instead of a 2 D parameter space, the point x votes into a 3 D



(c) The correct detection when LV is not in center

Figure 13. Better detection. doi:10.1371/journal.pone.0092382.g013



Figure 14. Cropped images for all frames of Volunteer 1. doi:10.1371/journal.pone.0092382.g014

parameter space. The extra dimension is for the radius parameter and the size is the length of the radius interval. The final winning accumulator provides not only the center but also the radius of the detected circle.

Anisotropic weighted circle hough transform. In the classic circle Hough transform, the point x votes to all candidates on the circle $B_{x,R}$, equally. This kind of circle Hough transform is isotropic. See Figure 11(a), in which the diameters of the candidate rings stand for the voting weights.

However, the probabilities of the target center falling into the candidates on the circle are not the same. If the gradient map of the original image is available as well, the gradient at x could provide more precise voting if one imagines the gradient vector points to the target center. The voting into the candidates is weighted according to the extent the candidate deviates from the direction the gradient vector points. In our study, we use cosine of the deviation angle to determine the weight. If the candidate is parallel to the gradient vector, the voting weight is maximized; if it is orthogonal to the gradient, the voting weight is zero. See Figure 11(b).

Considering the discussion on the increasing ARLR with the harmonic order and our goal of identifying the LV, we use two harmonic images, x^{H_1} and x^{H_5} to cooperate in the circle detection. Both of these images are passed to the edge detector. Two edge maps e_1 and e_5 are obtained, as are the gradient maps ∇x^{H_1} and ∇x^{H_5} as the by-product. The e_1 and e_5 for the above case are presented in Figure 12. Notice the smaller number of concentric rings conforming to the LV in e_5 . These two edge-gradient pairs are processed to vote into a unique parameter space, only with different weights. Since ARLR is minimized in x^{H_1} , suggesting that the LV has the most influential impact relatively in x^{H_1} , the pair $\{e_1, \nabla x^{H_1}\}$ votes with a positive weight, which is selected as 1 in our study. In the contrast, the pair $\{e_5, \nabla x^{H_5}\}$ votes with a negative weight (-1) due to the fact that ARLR is maximized in x^{H_5} . The candidates from $\{e_5, \nabla x^{H_5}\}$ mostly deviate from the LV center. Negative weights for them would largely suppress the incorrect candidates such as the ones in Figure 1. The whole algorithm is presented in Table 2.

After two voting rounds are completed, the highest accumulator in the 3 D parameter space suggests the LV center as well as the diameter of the LV cavity. We can see a much more precise localization result in Figure 13(a). This could provide a better initialization for edge-based or region-based segmentation approaches. Compared to the approach that counts on the LVcentered assumption in [27], our method is intrinsically rotational and translational invariant. The correct localization result for the cropped cine image, in which LV is far away from the center of FOV, is shown in Figure 13(c).

Results

After the circle detection, a 3 D parameter space A is obtained with $M \times N \times L$ accumulators. The highest accumulator A_{win} corresponds to the detected circle centered at $(A_{win}(1), A_{win}(2))$ with radius $A_{win}(3)$. Such a circle is plotted in Figure 13(a) and (c). In practice, the region of interest is identified as the rectangle that bounds this circle with some extension. Such a extended ROI is also plotted in the same case, shown in Figure 13(b) with the expanding ratio set to 1.5 in this study. The cropped images through the whole cardiac cycle are shown in Figure 14. The cine image contains 20 frames, the end-diastole frame of which is shown in the first sub-figure.

The proposed method was tried in 10 volunteer participants, 5 males and 5 females, mean age 40.5, age range from 24 to 59 years. The cine MR images were acquired on a 1.5 T Siemens scanner. The cine images was ECG gated. The slice thickness is 8 mm. The pixel spacing is from 1.25 mm to 1.87 mm, typically 1.71 mm. The TR/TE/flip angle is between $26.6-68.4/1.13-3.39/30^{\circ}-70^{\circ}$, typically $64.56/1.19/70^{\circ}$. A cine image slice contains typically 22 time frames over a cardiac cycle. The algorithm was implemented by Matlab and applied on the basal image sequence of each volunteer case. The computational time



Figure 15. Cropped images for end-diastole frames of ten volunteers. doi:10.1371/journal.pone.0092382.g015

Volunteers	Gender	Age	Original Image Size	Cropped Image Size	Cropping Ratio
1	Male	29	192*162	79*73	0.18541
2	Male	44	192*162	71*69	0.157504
3	Male	46	192*150	66*65	0.148958
4	Female	43	192*150	64*75	0.166667
5	Female	24	192*150	68*71	0.167639
6	Female	59	192*150	62*63	0.135625
7	Male	50	192*150	72*71	0.1775
8	Female	52	192*144	69*67	0.167209
9	Female	33	192*150	69*75	0.179688
10	Male	25	192*150	70*67	0.162847
Average	N/A	40.5	192*152	approx 69*69	0.165

Table 3. Statistics on Experiment Results.

doi:10.1371/journal.pone.0092382.t003

for each case averaged 3.94 seconds. Generating Harmonic images took 2.987 seconds, while LV detection took 0.956 seconds.

The cropped ROI of all cases contains the LV, which is visually inspected. The size ratio of the cropped ROI to the whole image ranges from 0.136 to 0.185, average is 0.165, which is a large reduction for the computation load. The statistics of experiments with all participants are shown in Table 3 and the cropped images of the end-diastole frame of all participants are shown in Figure 15.

Discussion

A new spectrum-based computer-aided tool has been proposed and developed for automatic LV localization.We found that the RV presented higher brightness than the LV in all harmonic images. The combination of the first and fifth harmonic images selected for an anisotropic weighted circle Hough detector was found to be the most robust for locating the LV. The development of this technique, which enables the automated location and further segmentation of the LV, will have a significant impact in research and in diagnostic settings, since we envisage that the automated method can be used by radiographers and cardiologists who will perform the diagnosis and assessment of ventricular function in patients with diverse heart diseases, like ischemic dilated cardiomyopathy [28], repaired Tetralogy of Fallot [29], heart failure before and after surgery [30], diastolic heart failure [31,32].

Only the basal short-axis images were used for analysis in our experiments. We selected the top basal images (i.e., the first slice below the slice containing LV Outflow Tract). The purpose of the automatic localization method proposed in our paper is to provide an initialization for the segmentation. The propagation of a single image segmentation result to other frames or other slices is quite common in approaches for the whole LV segmentation. Hence choosing the best starting slice and time frame is significant for minimizing the propagation error. The top basal images are chosen for the large dimension of both the LV and RV on such

References

 World Health Organization (2011) Fact sheet no. 317. Cardiovascular diseases.
 Wu KC, Lima JA (2003) Noninvasive imaging of myocardial viability: current techniques and future developments. Circulation Research 93: 1146–58.

 Petitjean C, Dacher JN (2011) A review of segmentation methods in short axis cardiac mr images. Medical Image Analysis 15: 169–184. slices. In contrast, the LV and RV occupy only small portion of the image and contain a small number of pixels. Any noises would affect the detection precision. A greater viability of harmonic images at apical slices could introduce more difficulties for our algorithm, while containing more pathological implication. Hence, a future study interest of ours is the application of this method on images that have more slice locations.

To clarify, we tested our algorithm on 10 healthy volunteers without any cardiovascular pathology. These 10 subjects constitute the study population used to demonstrate the proof-of-concept of our method. It should be noted that this is the first study to develop the described novel method for automated localization with the aim to simplify segmentation of LV cine images, which might represent a step forward in this field. Since the shape and motion pattern of the left ventricle and right ventricle may vary between patients, a validation study to test the approach in a bigger population with and without pathologies would be a logical next step.

In this study, we conducted the experiments in normal participants. The methodology proposed is also from an image analysis aspect. Furthermore, the relationship between harmonic images and the heart atlas concerning anatomy and functions would be more interesting for clinicians. The analysis of the correlation between harmonic image patterns and pathologies would be a powerful diagnosis aid tool, and this is a future research interest of ours.

Acknowledgments

The authors appreciate the medical editing assistance of Jon Kilner, MS, MA (Pittsburgh, Pennsylvania, USA).

Author Contributions

Conceived and designed the experiments: LZ RST. Performed the experiments: MW J-MZ LZ RST. Analyzed the data: MW J-MZ XDZ LZ RST. Contributed reagents/materials/analysis tools: MW J-MZ. Wrote the paper: MW LZ XDZ.

- Xu C, Pham D, Prince J (2000) Image segmentation using deformable models. Handbook of Medical Imaging 2: 129–174.
- Xu C, Prince JL (1998) Snakes, shapes, and gradient vector flow. IEEE Transactions on Image Processing 7: 359–369.

- Gupta A, Von Kurowski L, Singh A, Geiger D, Liang CC, et al. (1993) Cardiac mr image seg-mentation using deformable models. In: Computers in Cardiology 1993, Proceedings. IEEE, 747–750.
- Ranganath S (1995) Contour extraction from cardiac mri studies using snakes. IEEE Transactions on Medical Imaging 14: 328–338.
- Paragios N (2002) A variational approach for the segmentation of the left ventricle in cardiac image analysis. International Journal of Computer Vision 50: 345–362.
- Huang W, Gao Y, Chan KL (2010) A review of region-based image retrieval. Journal of Signal Processing Systems 59: 143–161.
- Nascimento JC, Marques JS (2008) Robust shape tracking with multiple models in ultrasound images. IEEE Transactions on Image Processing 17: 392–406.
- Nascimento J, Sanches J (2008) Ultrasound imaging lv tracking with adaptive window size and automatic hyper-parameter estimation. In: Image Processing, 2008. ICIP 2008. 15th IEEE Inter-national Conference on. IEEE, 553–556.
- Santarelli MF, Positano V, Michelassi C, Lombardi M, Landini L (2003) Automated cardiac mr image segmentation: theory and measurement evaluation. Medical Engineering & Physics 25: 149–159.
- Muller A, Neitmann A, Merkle N, Wohrle J, Hombach V, et al. (2005) Contour detection of short axis slice mr images for contraction irregularity assessment. In: Computers in Cardiology, 2005. IEEE, 21–24.
- 14. Jolly M, Duta N, Funka-Lea G (2001) Segmentation of the left ventricle in cardiac mr images. In: Computer Vision, 2001. ICCV 2001. Proceedings. Eighth IEEE International Conference on. IEEE, volume 1, 501–508.
- Viola PA, Jones MJ (2001). Rapid object detection using a boosted cascade of simple features.
- Pavani SK, Gomez DD, Frangi AF (2010) Haar-like features with optimally weighted rectangles for rapid object detection. Pattern Recognition 43: 160–172.
- Danilouchkine MG, Westenberg JJM, Reiber JHC, Lelieveldt BPF (2005) Accuracy of short-axis cardiac mri automatically derived from scout acquisitions in free-breathing and breath-holding modes. Magnetic Resonance Materials in Physics Biology and Medicine 18: 7–18.
- Ballard DH, Ballard DH (1981) Generalizing the hough transform to detect arbitrary shapes. Pattern Recognition 13: 111–122.
- Cocosco CA, Netsch T, Sngas J, Bystrov D, Niessen WJ, et al. (2004) Automatic cardiac region-of-interest computation in cine 3d structural mri. International Congress Series 1268: 1126–1131.
- Gering D (2003) Automatic segmentation of cardiac mri. Medical Image Computing and Computer-Assisted Intervention-MICCAI 2003 : 524–532.

Spectrum-Based Localization Tool of Left Ventricle

- Pednekar A, Kurkure U, Muthupillai R, Flamm S, Kakadiaris IA (2006) Automated left ventricular segmentation in cardiac mri. IEEE Transactions on Biomedical Engineering 53: 1425–1428.
- Huang J, Huang X, Metaxas D, Axel L (2007) Dynamic texture based heart localization and segmentation in 4-d cardiac images. In: Biomedical Imaging: From Nano to Macro, 2007. ISBI 2007. 4th IEEE International Symposium on. IEEE, 852–855.
- Sorgel W, Vaerman V (1997) Automatic heart localization from a 4d mri data set. In: Proceedings of SPIE. volume 3034, p.333.
- 24. Lin X, Cowan B, Young A (2006) Automated detection of left ventricle in 4d mr images: experience from a large study. Springer, 728–735.
- Jolly MP (2008) Automatic recovery of the left ventricular blood pool in cardiac cine mr images. Medical Image Computing and Computer-Assisted Intervention, 110–118.
- Kaus MR, Berg JV, Weese J, Niessen WJ, Pekar V (2004) Automated segmentation of the left ventricle in cardiac mri. Medical Image Analysis 8: 245– 254.
- Silva I, Almeida AG, Sanches J (2010) Automatic cropping for lv segmentation in cardiac mri. International Journal of Tomography & Simulation 13: 60–71.
- Zhong L, Su Y, Yeo SY, Tan RS, Ghista DN, et al. (2009) Left ventricular regional wall curvedness and wall stress in patients with ischemic dilated cardiomyopathy. American Journal of Physiology-Heart and Circulatory Physiology 296: H573–H584.
- Zhong L, Gobeawan L, Su Y, Tan JL, Ghista D, et al. (2012) Right ventricular regional wall curvedness and area strain in patients with repaired tetralogy of fallot. American Journal of Physiology-Heart and Circulatory Physiology 302: H1306–H1316.
- Zhong L, Su Y, Gobeawan L, Sola S, Tan RS, et al. (2011) Impact of surgical ventricular restoration on ventricular shape, wall stress, and function in heart failure patients. American Journal of Physiology-Heart and Circulatory Physiology 300: H1653–H1660.
- Zhong L, Poh KK, Lee LC, Le TT, Tan RS (2011) Attenuation of stress-based ventricular con-tractility in patients with heart failure and normal ejection fraction. Annals of the Academy of Medicine-Singapore 40: 179.
- Zhong L, Ng KK, Sim LL, Allen JC, Lau YH, et al. (2013) Myocardial contractile dysfunction associated with increased 3-month and 1-year mortality in hospitalized patients with heart failure and preserved ejection fraction. International journal of cardiology 168: 1975–1983.