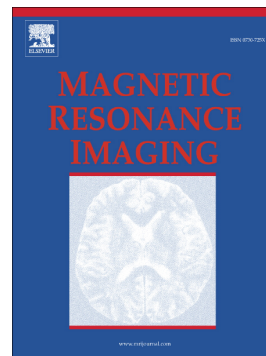


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## **Fully Automated Contour Detection of the Ascending Aorta in Cardiac 2D Phase-Contrast MRI**

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**Abstract**

**Purpose:** In this study we proposed a fully automated method for localizing and segmenting the ascending aortic lumen with phase-contrast magnetic resonance imaging (PC-MRI).

**Material and Methods:** Twenty-five phase-contrast series were randomly selected out of a large population dataset of patients whose cardiac MRI examination, performed from September 2008 to October 2013, was unremarkable. The local Ethics Committee approved this retrospective study. The ascending aorta was automatically identified on each phase of the cardiac cycle using a priori knowledge of aortic geometry. The frame that maximized the area, eccentricity, and solidity parameters was chosen for unsupervised initialization. Aortic segmentation was performed on each frame using active contouring without edges techniques. The entire algorithm was developed using Matlab R2016b. To validate the proposed method, the manual segmentation performed by a highly experienced operator was used. Dice similarity coefficient, Bland-Altman analysis, and Pearson's correlation coefficient were used as performance metrics.

**Results:** Comparing automated and manual segmentation of the aortic lumen on 714 images, Bland-Altman analysis showed a bias of  $-6.68 \text{ mm}^2$ , a coefficient of repeatability of  $91.22 \text{ mm}^2$ , a mean area measurement of  $581.40 \text{ mm}^2$ , and a reproducibility of 85%. Automated and manual segmentation were highly correlated ( $R = 0.98$ ). The Dice similarity coefficient versus the manual reference standard was  $94.6 \pm 2.1\%$  (mean  $\pm$  standard deviation).

**Conclusion:** A fully automated and robust method for identification and segmentation of ascending aorta on PC-MRI was developed. Its application on patients with a variety of pathologic conditions is advisable.

**Keywords:** Aorta, Magnetic Resonance Imaging, Computer-Assisted Image Processing

## 1. Introduction

Monitoring cardiovascular disease progression from subclinical changes to advanced stages is crucial for the definition of a target population to apply preventive strategies and perform longitudinal studies [1,2]. Structural and functional changes in the aortic wall can induce arterial stiffness, which is correlated with hypertension, associated with heart failure, and is a major predictor of stroke and myocardial infarction [3–5]. Aortic stiffness can be non-invasively measured through different methodologies. An average estimate of aortic stiffness can be obtained measuring pulse wave velocity by applanation tonometry [6]. On the other hand, local markers of aortic elasticity in the different aortic segments can be estimated using magnetic resonance imaging (MRI) [7,8]. In particular, phase-contrast MRI (PC-MRI) sequences application in cardiovascular imaging allows to obtain quantitative information on blood flow. These sequences can be easily performed during normal respiration or during breath-hold and MRI data acquisition can be synchronized with the cardiac cycle using both prospective and retrospective gating techniques [9]. Due to their high spatial and temporal resolution, these sequences are currently used to evaluate aortic elasticity in clinical practice [8].

The estimation of local parameters of aortic elasticity strictly rely on the accuracy and reproducibility of its lumen segmentation. In the last years, several attempts were performed to overcome the limitations of manual tracing, which is time-demanding and operator-dependent [2,10]. In particular, the high temporal resolution that can be achieved with PC-MRI sequence make the manual delineation of the aortic lumen time-consuming, unsuitable for daily clinical workflow and prone to measurement variability [2]. Moreover, the increasing number of data that can be generated with advanced PC sequences can increase the time needed for manual segmentation up to approximately 30 minutes for a real time PC image series of 148 frames [11]. To overcome the limitation of manual aortic lumen segmentation, in the last years several attempts were made to reach the highest possible level of automation.

Automated aortic lumen segmentation on cine cardiac imaging requires the detection of the aorta within the image and its segmentation throughout the whole cardiac cycle. Goel et al. proposed a possible solution to the localization of ascending and descending aorta on 2D PC-MRI images in 2014. However, once detected, vessel lumen boundaries were approximated to a perfect circle, so that local distensibility parameters could not be calculated [12]. In the last years, other research groups proposed semi-automated methods for aortic lumen segmentation still requiring an operator to perform a manual initialization of the segmentation process [8,11].

To the best of our knowledge, no methods were published that combine fully automated detection and segmentation of the ascending aorta on PC-MRI images. Thus, the aim of our study was to propose a fully automated method for localizing and segmenting the ascending aortic lumen on 2D PC-MRI images.

## **2. Materials and methods**

### *2.1. Dataset*

The local Ethics Committee approved this retrospective study and informed consent by patients was obtained. Twenty-five cardiac MR studies in 25 different subjects were randomly selected from a database of 192 patients, whose cardiac MR examination was unremarkable and negative with regard to the clinical question, being part of a large study dataset composed of 1027 consecutive cardiac 2D PC-MRI images performed at our institution from September 2008 to June 2014. Our sample was composed of 13 men and 12 women, whose age (mean  $\pm$  standard deviation) was  $37 \pm 23$  years (range 7-77 years). The selected MR studies were performed from September 2008 to October 2013.

Images were acquired using a 1.5-T unit (Magnetom Sonata Maetsro Class, Siemens, Erlangen, Germany). Breath-hold retrospectively electrocardiographically-gated two-dimensional phase-contrast gradient recalled echo sequences were acquired on a transverse plane about 3 cm above the aortic anulus, with a through-plane velocity encoding gradient ranging from 150 to 350 cm/s. In 23

out of 25 selected datasets, sequence parameters were set as follows: repetition time 41 ms; echo time 3.2 ms; flip angle  $30^\circ$ ; parallel imaging with acceleration factor 2, retrospective electrocardiographic gating with 30 phases per cycle; in-plane spatial resolution of  $1.3672 \times 1.3672$  mm<sup>2</sup>. The remaining two image series had the following acquisition parameters: repetition time 55 ms, echo time 3.2 ms, flip angle  $30^\circ$ ; in-plane spatial resolution of  $1.5625 \times 1.5625$  mm<sup>2</sup>; retrospective electrocardiographic gating with 11 phases per cycle in one series and 13 phases per cycle in the other one.

## 2.2. Ascending Aorta Detection

The automated detection of ascending aorta on 2D PC-MRI magnitude images required the aortic contour to be automatically selected and distinguished by the algorithm from other closed-line vessel-like structures. To this aim, the segmentation method started with the application of an edge detection filter to delineate the anatomical structures included in each frame. To facilitate this process, a sharpening preparatory step was performed in order to improve the contrast along image edges. The unsharp masking technique was used to this aim setting the standard deviation of the Gaussian lowpass filter at 2 pixels. After this preprocessing step, edge detection was performed using the Canny method [13]. Once the edge-image was created, the closed edges were automatically filled using morphological operators. This step allowed the detection of the vessels included in each image frame. Thereafter, a binary image containing all vessels was created. Within this image, for each vessel lumen were calculated: the area (A); the distance of its barycenter from the image center (D); the solidity (S), which is the ratio between object area and the area of smallest convex polygon that can contain it; the eccentricity (E), a scalar that indicates how much a conic section deviates from being circular ( $E = 0$  for a perfect circle). Finally, the ascending aorta was automatically detected in each frame using a priori knowledge of aortic morphology. During this process, for the three larger vessels depicted in each frame of the cardiac cycle, the “ascending aortic score” ( $AA_S$ ) was calculated as follow:

$$AA_S = \frac{A \cdot S}{D^2 \cdot E^3}$$

The vessels that maximized this score and had an eccentricity value lower than 0.55 was labeled as aorta in each frame of the cardiac cycle.

Due to the flow-induced variation of phase-contrast intensity during the cardiac cycle [8], it was not possible to detect the ascending aorta in all the frames of the sequence. For this reason, among vessels labeled as aorta in each frame of the cardiac cycle the one with the highest  $AA_S$  was chosen as “best frame”, which was used as initialization mask for the segmentation process.

### 2.3. Ascending Aorta Segmentation

The aortic lumen was segmented through the whole cardiac cycle using the active contour without edges technique, which is suited to detect objects whose boundaries are not necessarily defined by the image gradient, such as PC-MRI images [14]. The segmentation started with the “best frame” using the binary image with the previously localized aortic lumen as initialization mask. After that, the segmentation continued in a two steps process where the aortic lumen was segmented from the best frame to the first frame (best-to-first branch) and from the best frame to the last frame of the cardiac cycle (best-to-last branch). In each frame, the initialization mask was set as the binary image obtained segmenting the  $t+1$  or  $t-1$  frame depending on the algorithm branch.

The active contour technique has two main regularization factors, namely the smoothness of the boundaries ( $b_S$ ) and the tendency of the contour to grow or shrink ( $b_G$ ). In order to make the algorithm able to segment ascending aortae with different stiffness, these parameters were pre-set to standard values ( $b_S = 2.0$ ;  $b_G = -0.3$ ) and then iteratively optimized in each frame. The optimization process stopping criteria were: i) an aortic eccentricity lower than 0.60; and ii) an area difference between the initialization mask (namely the previous or subsequent segmented frame) and the segmented area lower than 7% of the initialization mask. This parameter was empirically set to this value based on the maximum area difference recorded in the original dataset of 1,027 2D PC-MRIs,

which comprises both healthy and pathological subjects. In that dataset the maximum area difference between the systolic and diastolic phase was equal to 176% in image series of 30 frames. For this reason, we set the upper limits of area increment to 200% on the whole cardiac cycle that, assuming a gradual area increment during the cardiac cycle, corresponds to an area increment of 6.6% in each frame (successively rounded to 7%).

At the end of this process, the aortic lumen was segmented in each frame of the cardiac cycle. The whole algorithm was developed using Matlab (MathWorks, Natick, MA, USA), version R2016b, on an Intel i7 processor 2.5 GHz with 8 GB of RAM and an NVIDIA GeForce 940MX graphic card. Its flowchart is depicted in Figure 1.

#### *2.4. Validation*

To validate the proposed segmentation method, manual tracing of the aortic lumen performed by an expert operator was used as a reference standard. This operator was a radiology resident who had previously manually segmented more than 1,300 2D PC-MRI series of the ascending aorta in a large retrospective study (10). In order to evaluate the intra-operator agreement of repeated measurements, the manual segmentation was performed twice on 10 randomly selected subjects (300 images) with at least 3 months of interval between the two repetitions. Moreover, to evaluate algorithm repeatability the operators run it two times on the whole dataset. Bland-Altman analysis and the one-way random model inter-class correlation coefficient (ICC) were used to assess operator and algorithm repeatability [15]. Elapsed time was also recorded for both manual and automated segmentations, which were performed on the same computer used to develop the proposed algorithm.

Bland-Altman analysis was also used to evaluate the agreement between the manual and automated segmentation [15]. In addition, linear regression and Dice similarity coefficient [16] were used as performance metrics. In Figure 2 an example of the comparison between manual and automated segmentation is shown.



### 3. Results

The proposed segmentation method of the ascending aorta on 2D PC-MRI images was fully automated and provides full repeatability, with an ICC value between repeated measurement equal to 1.00 (95% confidence interval [CI]: 1.00 -1.00). The Bland-Altman analysis on repeated measurements performed with the proposed algorithm shows a bias of 0.00 mm<sup>2</sup> (95% CI: 0.00 – 0.00), a coefficient of repeatability (CoR) of 0.00 mm<sup>2</sup> over a mean measurements of 622.63 mm<sup>2</sup>, resulting in a reproducibility of 100% (Figure 3a). The mean segmentation time for a 2D PC-MRI image series composed of 30 frames ( $\pm$  standard deviation) was equal to 19.4  $\pm$  11.7 sec in the case of automatic segmentation.

On the other hand, intra-operator analysis of the manual segmentation process showed an ICC value on single measurements equal to 0.989 (95% CI: 0.986 – 0.991). Intra-operator Bland-Altman analysis showed: a mean area measurement equal to 612.45 mm<sup>2</sup>, a bias equal to -10.90 mm<sup>2</sup> (1.7% of the average measurement), a 95% CI equal to -80.85 – 59.05 mm<sup>2</sup>, a CoR equal to 71.38 mm<sup>2</sup>, with a resulting reproducibility of 88% (Figure 3b). The mean manual segmentation time was 877.0  $\pm$  43.5 sec.

The algorithm was able to segment all frames of the analyzed dataset. Comparing the automated and the manual segmentation approaches, the linear regression showed a high correlation value ( $R = 0.98$ , automated = 0.96 manual + 27 mm<sup>2</sup>). The Bland-Altman analysis showed a mean area measurement equal to 581.4 mm<sup>2</sup>, a bias equal to -6.68 mm<sup>2</sup> (1.2% of the average measurement), with a 95% CI equal to -96.07 – 82.72 mm<sup>2</sup>, a CoR equal to 91.22 mm<sup>2</sup> and a resulting reproducibility equal to 85% (Figure 4). The Dice similarity coefficient resulted to be 94.6  $\pm$  2.1%

### 4. Discussion

In this study, we developed a robust, accurate, and fully automated method for localization and segmentation of the ascending aorta on two-dimensional PC-MRI magnitude images.

The proposed algorithm is fully repeatable, therefore minimizes the operator dependency of the manual segmentation and all the limitations that stem from it, as can be seen in the Bland-Altman plot of repeated manual measurements. Bland-Altman analysis demonstrated that the measurement variability on repeated measurements is null in case of automated segmentation, making it suitable for use in longitudinal assessment.

The algorithm was validated by comparison with manual tracing performed by a highly experienced operator on 25 image series, corresponding to 714 images. The validation showed a high correlation between the automated and manual tracing. However, considering that manual tracing suffers from the operator-dependency, we also used the Bland-Altman analysis, more suitable when two methods are compared without preliminary preference for one of them [15]. Finally, the Dice coefficient demonstrated a high correspondence between automatically and manually traced contours, comparable to those obtained by others using semi-automated methods [8].

During the last years, several semi-automated methods were developed using different segmentation approaches, which still need to be manually initialized, thus compromising their reproducibility [8,11,17–19]. In particular, Herment et al. developed a robust approach that requires the operator to select a squared region of interest around the aorta [8]. On the other hand, Goel et al, to overcome the initialization problem, proposed an automated tool for both ascending and descending aortic lumen localization on 2D PC-MRI images. Although they developed a robust localization algorithm, after being localized, the aortic lumen was just approximated as a perfect circle. For this reason, their solution did not replace previous approaches that semi-automatically segment the aorta because an accurate segmentation and the adherence to the actual anatomical structure is fundamental to evaluate local aortic elasticity parameters [12].

Our results in term of Dice similarity coefficient (mean 94.6%) are very close to those obtained from Herment et al. (mean 94.5%). However, in addition to the good results of this study, we minimized the operator dependency. Moreover, the algorithm is designed to converge to a unique

solution, with null algorithm outcome variability; this makes it suitable for follow-up assessments of aortic morphology and elasticity.

The algorithm was tested on a series of randomly selected subjects with negative cardiac PC-MRI examination. One might argue that applying the algorithm only on healthy aortae is a relevant limitation. Nevertheless, the wide age range of the included subjects (7-77 years) made it possible to assess aortic elasticity at different ages, given the well-known reduction of aortic distensibility even with normal ageing. [1]. Moreover, due to the design of the proposed algorithm, an enlarged aortic lumen of an elderly man would have been easier to segment than a healthy child's aorta, which is more elastic. This apparent paradox is due to the reduced distensibility of an enlarged aorta. Once the algorithm detected and segmented the aorta in the "best frame", it easily tracked its contours through the whole cardiac cycle. Moreover, the iterative calculation of the active contouring regularization factors should make the algorithm able to segment different aortic morphologies; some examples of segmentation outcome are depicted in Figure 5.

This study has limitations. Firstly, manual tracing was used as reference standard, even if it is affected by intra- as well as inter-operator variability with a intra-operator reproducibility equal to 88%. However, manual contouring is still one of the most practiced segmentation methods in cardiovascular MR imaging, when it comes to precisely delineate structures. Secondly, the methods proposed here have been validated only for the ascending aorta. In order to extend the algorithm application to different aortic segments or to other vessels, the localization parameters would need to be adapted to the anatomical structure of interest.

In this study, the elastic deformation of the aortic wall was approximated as a linear area increment. In particular, we set the maximum area increment or decrement for each phase of the cardiac cycle equal to 7% based on the maximum area increment observed in the above mentioned population database of 1,027 subject.

This assumption represents a simplification of the complex deformation process and could make algorithm performance sample dependent. Nevertheless, the wide age range of this sample and the full spectrum of physiological aortic deformations (i.e. low and high ascending aortic strain) on which this algorithm has been tested make this experience-based choice mostly able to suite the everyday clinical practice. However, the validation of the proposed algorithm on subjects belonging to a larger and different sample will prove the strength of this assumption.

In this study, we used only the magnitude images to segment the ascending aortic lumen in order to take advantage of the morphological information contained therein, which is crucial for ascending aorta detection and segmentation initialization. Nevertheless, further performance improvement could be obtained by combining both magnitude and phase image information to improve aortic boundary detection during the active contour segmentation step.

Finally, the computational time of the algorithm is still relatively long (with a mean of about 20 seconds), albeit low if compared to the time needed to perform the manual segmentation (approximately equal to 15 minutes).

## **5. Conclusions**

In conclusion, this study proposes a fully automated detection and segmentation method for the ascending aorta on 2D cardiac PC-MRI images. Results show high performances in terms of accuracy and reproducibility. This method may accelerates the evaluation of ascending aortic morphology and elasticity minimizing operator dependency during lumen segmentation. Its validation requires the application on larger series of patients with a variety of pathologic conditions.

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**Conflicts of interest**

None.

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**Figure 1.** Flowchart of the proposed automated segmentation method. The flowchart shows the main steps of the proposed algorithm: edge detection, vessel detection, ascending aorta detection and its segmentation through the entire cardiac cycle.

**Figure 2.** Phase-contrast MR images of the ascending aorta on in a 44-year old female subject. Of the 30 frames acquired, only 10 are shown, one frame every 3 frames. Comparison between automated segmentation (green contour) and manual segmentation (red contour). The near-complete overlapping of the two segmented areas can be appreciated. Area values obtained with each method for each phase are reported in the table.  $\Delta$  = the absolute difference between manually and automatically obtained areas.

**Figure 3.** Bland-Altman plot resulting from the comparison between repeated area measurements performed automatically (A) and manually (B) on phase-contrast MR images in a subset of 10 subjects.

**Figure 4.** Bland-Altman plot resulting from the comparison between automatically ( $A_A$ ) and manually ( $A_M$ ) segmented luminal areas of the ascending aorta on phase-contrast MR images in the entire analyzed dataset.

**Figure 5.** Examples of segmentation outcomes (right) and their corresponding original images (left) in case of different aortic lumen morphology.

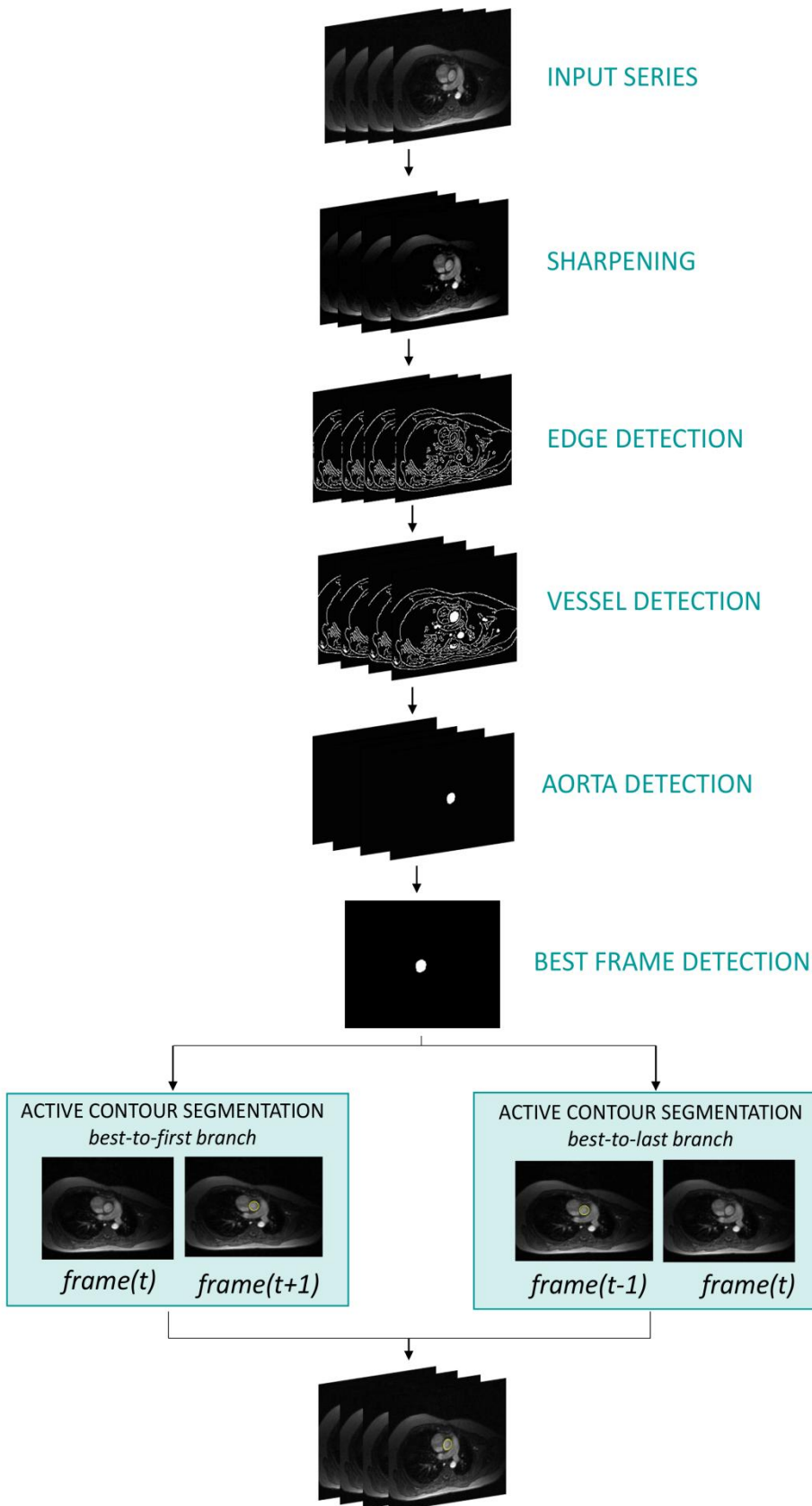


Fig. 1

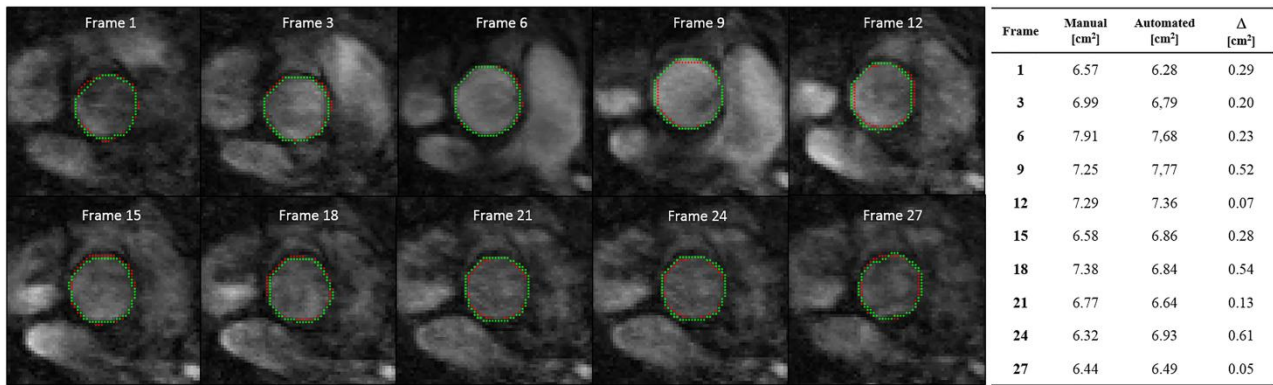


Fig. 2

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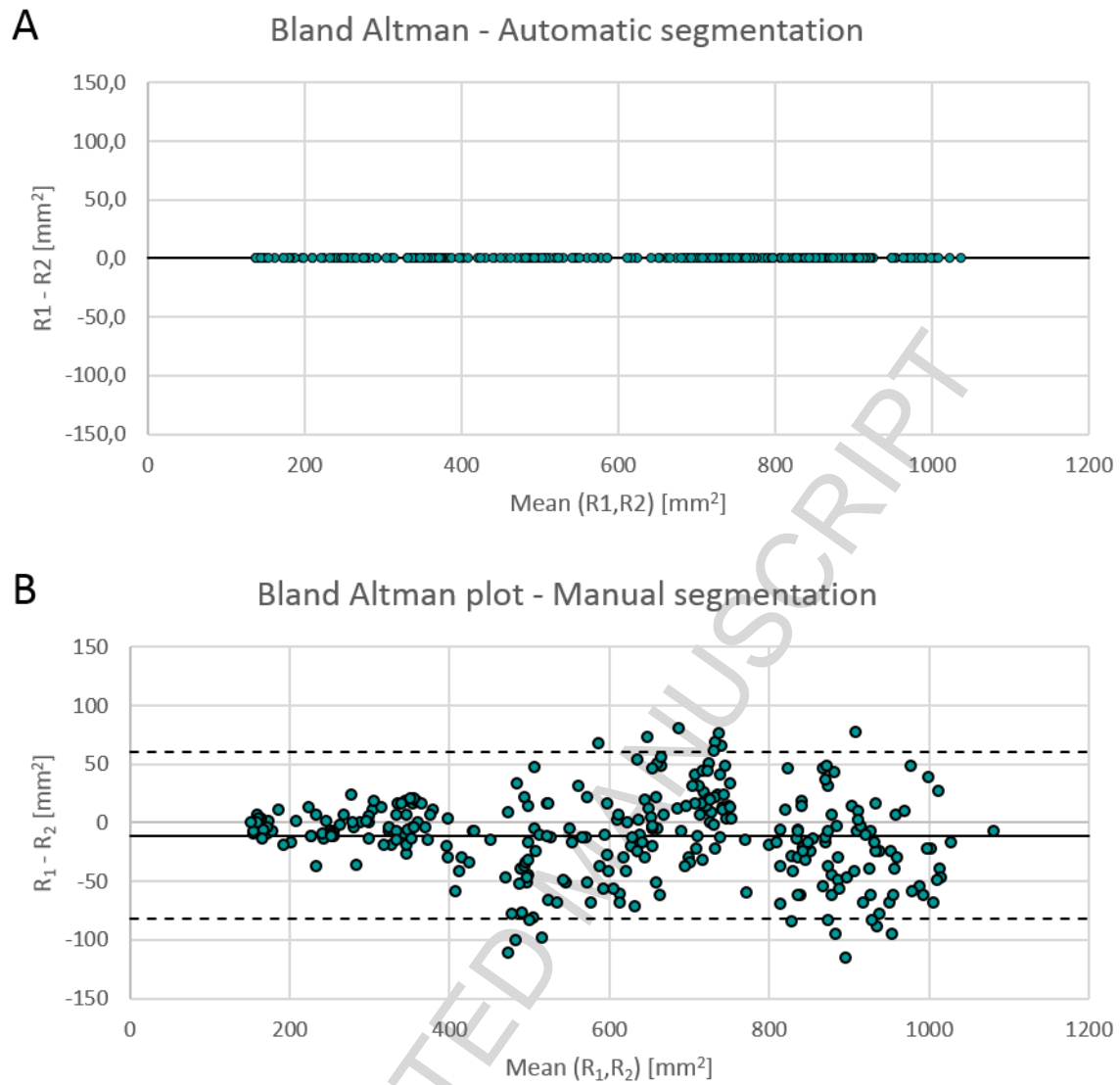


Fig. 3

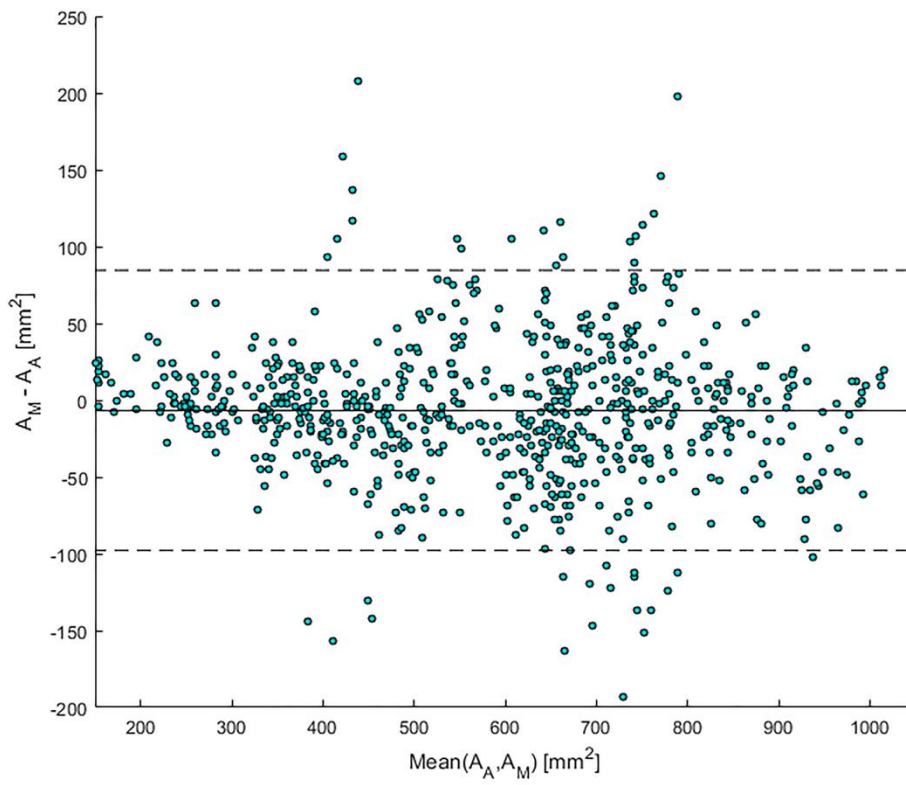


Fig. 4

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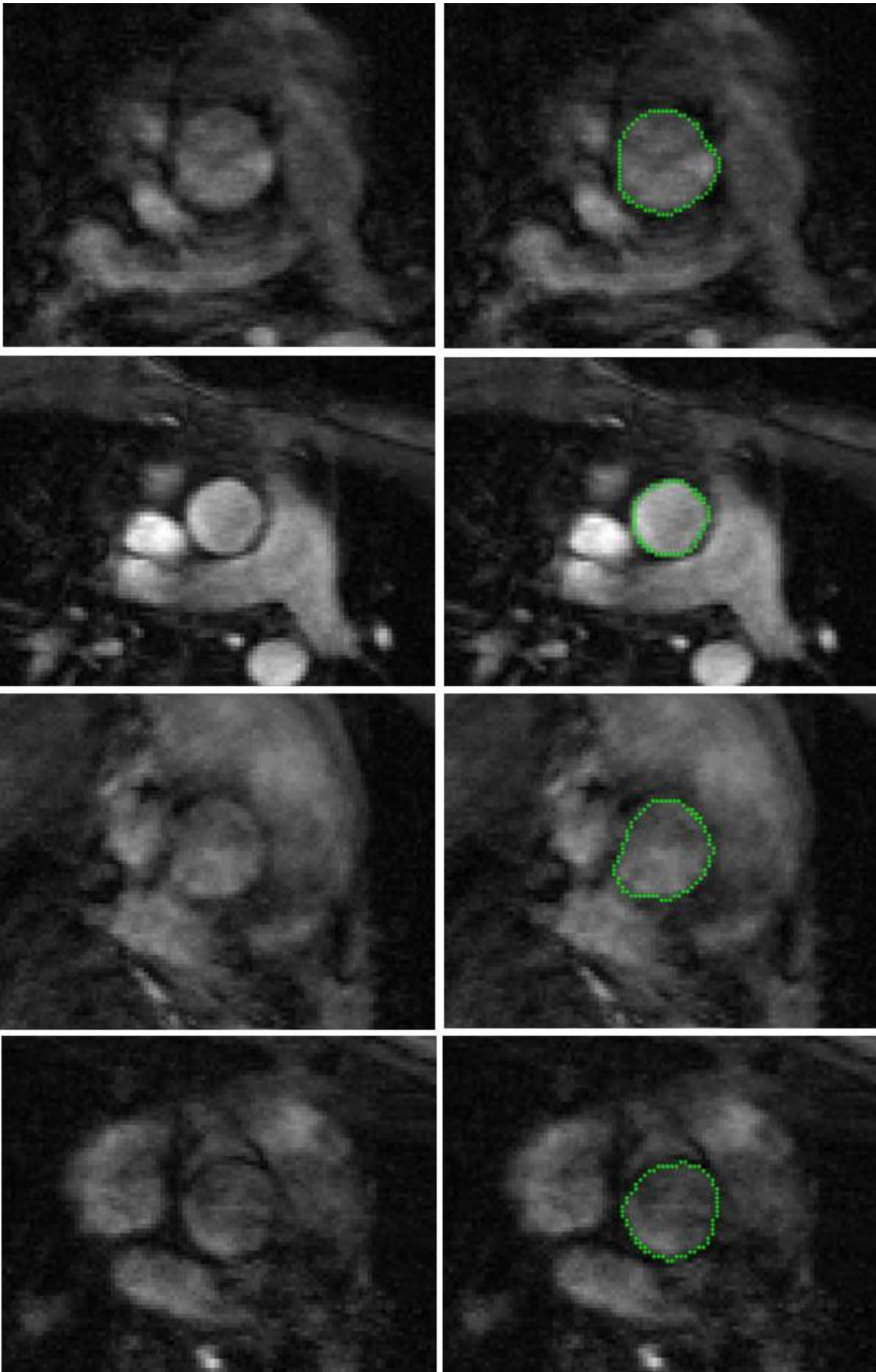


Fig. 5

## Highlights

- 1) Fully automated and robust detection and segmentation of the ascending aorta lumen
- 2) Fully reproducible segmentation tool suitable for longitudinal examinations
- 3) High correlation between automatically and manually segmented areas ( $R=0.98$ )

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