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### Shape Analysis of Segmentation Variability

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#### Abstract

Patient-specific cardiac simulation rely on accurate geometric models extracted from medical images. Segmentation of cardiac images is a key, yet possibly error-prone part of patient-specific simulations, e.g., heart propagation models, ECG forward simulation, and ECG Imaging. In this study, we performed shape analysis on multiple segmentations of the same patient to quantify variability. We found that segmentation shape varied most in the basal region of the ventricles and the right ventricular outflow tract in all three structures, which could have significant impact on pipelines that depend on geometric models. The statistical shape-model generated using ShapeWorks provides a pathway to subsequently quantify the impact of the segmentation variability on modeling pipelines with uncertainty quantification.

#### 1. Introduction

Patient-specific cardiac simulation is becoming increasingly relevant as a research and clinical tool for predicting arrhythmias and guiding treatments. These simulation methods, such as heart propagation models [1], ECG forward simulation, and Electrocardiographic Imaging (ECGI)[2, 3], rely on interpreting clinical data, especially medical images, into computer models. Although many of these pipelines are continually improving and emerging into clinical use, variability from the interpreting clinical data remains largely unquantified for many approaches.

Many assumptions and estimations comprise the interpretation of clinical data into a tractable cardiac models, leading to multiple possible sources of uncertainty. One source of uncertainty, often overlooked, is the segmentation of the geometric model, particularly the heart, because it usually requires manual input and user judgements. We have previously shown that segmentations of the same patient geometry can vary widely, especially the cardiac surface [4] and likely affects the ECGI solutions [5]. However, we still do not have a sufficiently quantifiable description of the variability of segmentation shape and its resulting affect on cardiac simulations.

In this study, we quantified segmentation shape variability and generated a statistical shape-model to facilitate uncertainty quantification of cardiac modeling pipelines. We used the collaborative framework of the Consortium for ECG Imaging (CEI) to generate multiple segmentations of the same patient to provide sufficient data for computing shape variability of cardiac segmentations, and used a correspondence-based shape analysis to calculate shape statistics and a parameterized shape space. We found the areas of segmentation variability agreed, generally, with our previous analysis of ECGI variability [5], and that the shape space provided a statistical shape-model for use in uncertainty quantification.

#### 2. Methods

To quantify variability of cardiac segmentation, we used statistical shape analysis on multiple ventricular segmentations. A single patient CT scan was segmented by nine researchers within the CEI and shape analysis was applied using ShapeWorks [6] (https://www.sci.utah.edu/software/shapeworks.html). Shape-Works performs shape analysis by finding correspondence points across all input geometries using a particle system optimizer. The optimization attempts to maximize distance between points while minimize the number modes of variation across the cohort of geometries. The correspondence points, shown in Figure 1, facilitate regional comparison of the geometries and subsequent shape analysis. The original ventricular segmentations did not produce stable correspondence solutions in ShapeWorks, likely due to the concave shape of the ventricles, so each was split into three separate surfaces: epicardium, left ventricular (LV) endocardium, and right ventricular (RV) endocardium. This

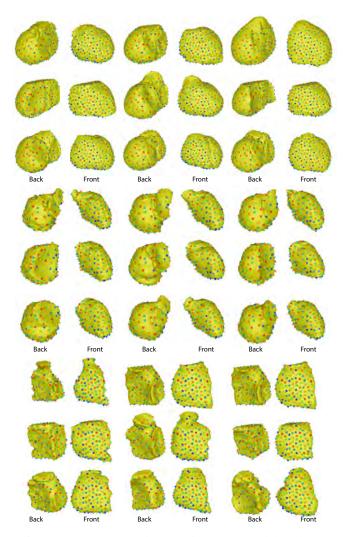


Figure 1. Correspondence points generated by Shape-Works for all segmentations.

produced three cohorts with correspondence points for shape analysis.

The shape analysis performed in this study included euclidean distance statistics and a principle component analysis (PCA) to determine modes of shape variation. For each of the three set of surfaces, a mean geometry was created from the computed correspondences. The standard deviation of the euclidean distance from each correspondence point to the mean geometry was calculated to create a map of local shape variability. PCA was also performed on the correspondence point sets, identifying seven or eight orthogonal axes, or modes, of variation for each of the 3 surface sets. We limited analysis to the first four modes of variation, which encompassed of 89.3 %, 87.2 %, and 89.2 % of the total variability for the epicardium, LV endocardium, and RV endocardium. For the four major modes of variation, we computed the standard deviation of the

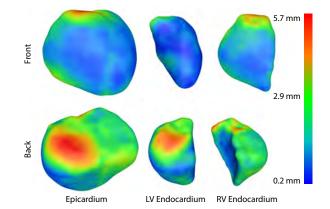


Figure 2. Local cohort variability computed by the standard deviation of the euclidian distance of each point set to the mean point set.

distribution and generated geometries at intervals along the principle axes. The axes of variation and the distribution along each axis constitute the statistical shape-model.

The patient data used in this study were collected by Sapp et al. [7] and is available for open use on the EDGAR database (http://edgar.sci.utah.edu) [8] a shared resource of the CEI.

#### 3. **Results**

Figure 1 demonstrates the results of the particle system optimization, i.e., the computed correspondence points. Qualitatively, the points are generally evenly spaced and cover the macro structures of each of the surfaces. There are regions, such as the RV apex and RV outflow tract (RVOT) which have relatively sparser sampling than the rest of the surface in some submissions. However, the coverage of the correspondences generated was sufficient to allow for suitable shape analysis.

Shape analysis of the segmentations showed regions of high variability across the three surfaces. Figure 2 shows high variability in the euclidean distances of the point sets in regions along the base and the RVOT. Other areas of variability include the posterior wall and the RV apex. Figure 4 shows high qualitative agreement between the local variability along the primary mode of variation and the euclidean distance variation (Figure 2), albeit with a higher amplitude. Furthermore, Figure 4 shows that other modes of variation contain the most variability in the same locations as the primary mode, yet the relative variability among those regions shift. For instance, while the RVOT demonstrates high variability in all modes, the area of highest standard deviation can occur at different locations along the annulus of the pulmonary valve: the whole annulus in axis 0, the center and the endocardium in axis

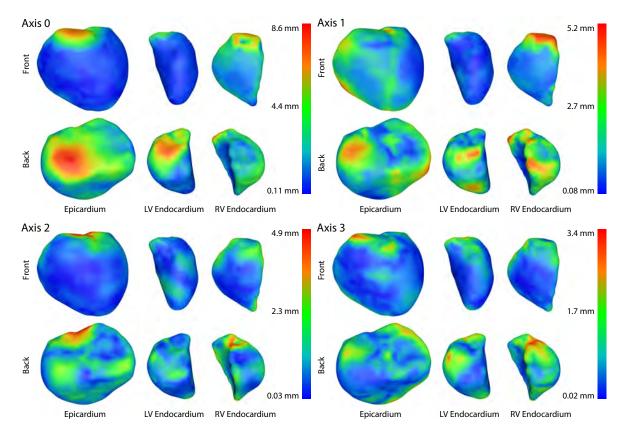


Figure 3. Local standard deviation along the 4 major modes of variation.

1, on the base in axes 2 and 3, and on the anterior annulus in axis 3. Figure 4 shows the geometric changes that occur along each of the principle axes. The standard deviations in shape space along each of the axis of variation are 63.8, 42.7, 29.2, and 20.8 mm for the epicardium; 35.6, 23.0, 15.1, and 13.5 mm for the LV endocardium; 42.1, 31.2, 22.5, and 17.2 for the RV endocardium.

#### 4. Discussion and Conclusions

The results in this study indicate substantial variability in regions of the cardiac segmentation that may affect the results of cardiac simulations. Additionally, the work in study involves generating a parameterized shape-model that can be used in cardiac modeling pipelines to compute uncertainty.

The high variability regions shown in near the base and RVOT (Figures 2 & 3) matches results found in our previous work that analyzed segmentation based on closest distance to an aggregate segmentation generated with STA-PLE [5]. These regions aligned with high variability of the ECGI solution, suggesting a link between simulation variability and segmented shape variability. However, our current shape analysis also identified regions of variability on the posterior wall and near the apex, areas which previously showed less variability in the ECGI solution [5]. However, more more analysis is needed to explore the link between the high cardiac segmentation variability simulation results.

The shape analysis performed with ShapeWorks includes generating a shape space from the PCA which acts as a statistical shape-model. Any geometry within the shape space can be recreated by a linear combination of the modes of variation. Additionally, the projection of the cohort geometries onto each of the axes can be described with a scalar distribution. The mean geometries, axes of variation, each defined as a set of 3D vectors for each point, and the distributions on each axis can be used as the inputs of uncertainty tools such as polynomial chaos [9] to provide a estimation variability.

While a relatively small part of many cardiac modeling pipelines, segmentation of the heart is an uncertain aspect of simulation. The work in this study will lead to better quantification of these sensitivities by combining shape-models with uncertainty tools like UncertainSCI (https://github.com/SCIInstitute/UncertainSCI).

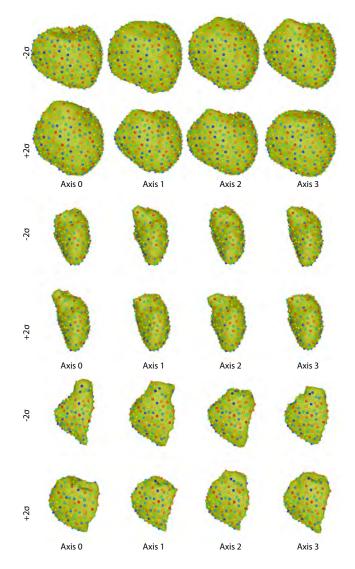


Figure 4. Shape changes along the four major modes of variation at two standard deviations ( $\sigma$ ).

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