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▶ To cite this version:

Maxime Di Folco, Patrick Clarysse, Pamela Moceri, Nicolas Duchateau. Learning interactions between cardiac shape and deformation: application to pulmonary hypertension. 10th International Statistical Atlases and Computational Modeling of the Heart (STACOM) Workshop, Held in Conjunction with MICCAI 2019, 2019, Shenzen, China. hal-02309296

HAL Id: hal-02309296 https://hal.science/hal-02309296

Submitted on 9 Oct 2019

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Learning interactions between cardiac shape and deformation: application to pulmonary hypertension

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Abstract. Cardiac shape and deformation are two relevant descriptors for the characterization of cardiovascular diseases. It is also known that strong interactions exist between them depending on the disease. In clinical routine, these high dimensional descriptors are reduced to scalar values (ventricular ejection fraction, volumes, global strains...), leading to a substantial loss of information. Methods exist to better integrate these high-dimensional data by reducing the dimension and mixing heterogeneous descriptors. Nevertheless, they usually do not consider the interactions between the descriptors. In this paper, we propose to apply dimensionality reduction on high dimensional cardiac shape and deformation descriptors and take into account their interactions. We investigated two unsupervised linear approaches, an individual analysis of each feature (Principal Component Analysis), and a joint analysis of both features (Partial Least Squares) and related their output to the main characteristics of the studied pathology. We experimented both methods on right ventricular meshes from a population of 254 cases tracked along the cycle (154 with pulmonary hypertension, 100 controls). Despite similarities in the output space obtained by the two methods, substantial differences are observed in the reconstructed shape and deformation patterns along the principal modes of variation, in particular in regions of interest for the studied disease.

1 Introduction

Clinical routine mainly uses simplified measurements of the cardiac function to characterize cardiovascular diseases, among which scalars such as ejection fraction or volumes, and myocardial strain. These scalars summarize high dimensional information about the cardiac shape and its deformation, and may therefore represent a substantial loss of information for complex pathologies.

Dimensionality Reduction Methods (DRM) have been used in cardiac imaging applications to better consider high-dimensional descriptors encoding shape and deformation, either individually [1–4] or jointly [5]. Very recently, [6] went beyond these techniques and used auto-encoders on dynamic cardiac shapes to compute a latent low-dimensional space optimized for survival prediction. $\mathbf{2}$

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However, cardiac shape and deformation undergo strong interactions depending on the disease. For example, in the case of pulmonary hypertension, a pressure overload forces the Right Ventricle (RV) to adapt by increasing its contractility to maintain a normal RV function. After a certain amount of time, the contractility can no longer increase and a progressive RV shape dilation and motion abnormalities may appear [7].

Methods that mix several high-dimensional heterogeneous descriptors all at once do not take into account their interactions and redundancy, which may limit the analysis. In methods reported in [8-10], an affinity matrix is established as composed of blocks that represent either the affinity between samples according to a single descriptor (diagonal blocks), or the interactions between two descriptors (extra diagonal blocks). Because of the nonlinearity of the methods and the unique output space returned, the reconstruction to the input space is not obvious. Other works proposed to link two learnt representations using additional constraints that express the hypothesized interactions between descriptors [11, 12]. These methods first determine the low-dimensional representations independently and then consider interactions, which can substantially alter the initially learnt representations. In [13], linear techniques consider interactions while learning the low-dimensional representation, demonstrated for linking two different imaging modalities from the same heart. Nonetheless, this application concerned two descriptors that theoretically represent very similar information, while we target the study of shape and deformation descriptors, with known but only partial interactions.

In this paper, we explore the interactions between two partially interacting high-dimensional descriptors (cardiac shape and deformation), with unsupervised linear DRM either applied to descriptors independently or to both descriptors jointly. In the perspective of risk stratification from the low-dimensional output space, we notably evaluate how critical DRM can be for the assessment of the main local characteristics of a given disease: pulmonary hypertension, looked at through 3D RV echocardiographic data. In particular, we consider deformation patterns or shape as whole entities and not at each location independently from the others [14–16].

2 Methods

2.1 Data and pre-processing

We processed a database of right ventricular meshes tracked on 3D echocardiographic images along the cardiac cycle, obtained from the commercial software 4D-RV Function 2.0 (TomTec Imaging Systems GmbH, Unterschleißheim, Germany). The meshes have 822 cells and 1587 points. The database is composed of 254 cases with 100 controls and 154 patients with pulmonary hypertension.

Point-to-point correspondences already exist between the meshes from different subjects, which allows straightforward comparisons within a population. Nonetheless, to reduce the bias due to different spatial heart positions, we realigned the meshes rigidly with Procrustes alignment (limited to global translations and rotations) [17]. This also provides an average shape for the controls population, used as template for display and analysis.

Then, two descriptors representing shape and deformation respectively were computed. The first one, called area strain (Fig.1a), is the relative area change (in %) of each mesh cell between End-Diastole (ED) and End-Systole (ES). It summarizes the deformation of the RV over the systole. The second one characterizes shape at ED, through a distance vector (Fig.1b) computed at each mesh point between a given case and the average shape of the controls, used as reference for normality.

These two features are high-dimensional: 822×1 for area strain (a scalar at each mesh point) and 822×3 for the distance vector (a 3D vector at each mesh point).

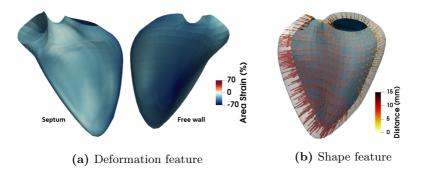


Fig. 1: (a) Area strain pattern at end-systole for a healthy case expressed in % (b) Distance vector (mm) colored by magnitude for the same healthy case. The blue mesh stands for the reference mesh used for the computations.

2.2 Dimensionality reduction

DRM take as input high-dimensional data and return a low-dimensional space encoding the main characteristics of the input space. Here, we focus on the following two linear approaches. Principal Component Analysis (PCA) looks for the main directions of variance in the data. This can be written as the following eigenvalues problem, for a single descriptor \mathbf{x} :

$$\mathbf{C}_{\mathbf{x}}\mathbf{w} = \lambda \mathbf{w},\tag{1}$$

where $\mathbf{C}_{\mathbf{x}}$ is the covariance matrix of the studied descriptor, \mathbf{w} and λ are the eigenvectors and the eigenvalues of $\mathbf{C}_{\mathbf{x}}$, respectively.

The second considered method is the two-block Partial Least Square (PLS) regression [18], which mutually maximizes the covariance between the set of projections for two features, which amounts at solving the following eigenvalues problem:

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$$\mathbf{C}_{\mathbf{x}\mathbf{y}}\mathbf{C}_{\mathbf{y}\mathbf{x}}\mathbf{w}_{\mathbf{x}} = \lambda_{PLS}^{2}\mathbf{w}_{\mathbf{x}},\tag{2a}$$

$$\mathbf{w}_{\mathbf{y}} = \frac{1}{\lambda_{PLS}} \mathbf{w}_{\mathbf{y}},\tag{2b}$$

where $\mathbf{C}_{\mathbf{x}\mathbf{y}}$ and $\mathbf{C}_{\mathbf{y}\mathbf{x}}$ are the cross-covariance matrices of the descriptors \mathbf{x} and \mathbf{y} , and $\mathbf{w}_{\mathbf{x}}$ and λ_{PLS} are the eigenvectors and eigenvalues of the matrix $\mathbf{C}_{\mathbf{x}\mathbf{y}}\mathbf{C}_{\mathbf{y}\mathbf{x}}$. Note that with PLS, we obtain at once two output spaces, one for each descriptor.

Due to the linearity of both methods, the inverse transformation (from the output space to the input space) is known. Therefore, the reconstruction of meaningful points from the output space is straightforward in particular along the first components. Note that for the shape descriptors, the reconstruction from the output space provides a vector field encoding distances from the template shape: a final step is needed for visualization and analysis purposes, which consists in reconstructing the corresponding shape by applying this vector field to the template shape.

3 Experiments and results

The following section reports on the application of the DRM to the entire population (patients and controls). Considering patients and controls at the same time allows to study the continuum that exists from normality to the most severe grades of pulmonary hypertension, in line with what has recently been recommended for cardiac applications [19].

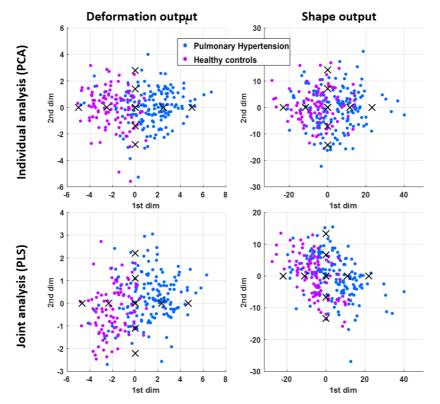
3.1 Output spaces

We first compare the two methodologies (individual analysis of each features using PCA and joint analysis using PLS) based on their output spaces. Each analysis returns two output spaces (one per descriptor, Fig. 2).

A comparison between the obtained components by each strategy reveals that the first and second components are highly correlated (shape: $r^2 = 0.99$ first component, $r^2 = -0.59$ second component; deformation: $r^2 = 0.98$ first component, $r^2 = -0.69$ second component).

Nevertheless, another way of comparing the output spaces reveals differences unseen with the previous assessment. We evaluated the correlation between the components of the two output spaces, for each strategy (Table. 1). With PCA, the first components of a given descriptor are linked with all the other components of the other descriptor. This is different with PLS, where each component of a given descriptor is only linked to the corresponding component of the other descriptor (low extra-diagonal coefficients in the table). This comes from the fact that PLS maximizes the covariance of both spaces at the same time, so the corresponding components are linked together but not with the others. This link is the main difference between the output spaces of both analysis.

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Fig. 2: Output spaces for each descriptor, from the two different strategies. Crosses indicate the samples at $[-2, -1, 0, +1, +2] \sigma$ used to reconstruct the modes of variations examined in Sec. 3.2.

	Deformation					Deformation		
		1st 2nd 3	Brd			1 st	$\mathbf{2nd}$	$\mathbf{3rd}$
	1 st	-0.39 0.19 -0	0.28		1 st	0.51	1e-17	1.5e-16
Shape	$\mathbf{2nd}$	-0.27 0.023 -0	0.13	Shape	2nd	-4.3e-18	0.48	3.8e-16
	$\mathbf{3rd}$	-5e-3 4e-3 -0 .	.065		$\mathbf{3rd}$	1e-17	2.4e-16	0.37
(a) Individual analysis (PCA)				(b) Joint analysis (PLS)				

Table 1: Correlation coefficient (r^2) between the components of the two output spaces.

3.2 Principal modes of variations

The first mode of variations for the shape descriptor quantifies the size of the right ventricle (Fig. 3). The differences plotted on the right side of the figure

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correspond to the point-to-point distance between the PCA and PLS shapes at $+2\sigma$. Samples around $+2\sigma$ (+2 standard deviations) lie in the side of the output space where pulmonary hypertension patients predominate, and have bigger shapes (characteristic RV dilatation in pulmonary hypertension). The second mode of variations encodes the intra-valve distance. The meshes obtained from the two DRM substantially differ close to the apex (near 10mm for a ventricle around 80 mm long). The third mode encodes changes in the curvature of the septum, and corresponds to reported changes in this zone for severe pulmonary hypertension cases ("septal bowing", which means that the septum bows itself and becomes almost straight [20]).

Note that the differences between meshes at -2σ is equal to the differences at $+2\sigma$ due to the linear dimensionality reduction and reconstruction methods, if the differences are unsigned (absolute differences). If signed, differences at -2σ are the exact opposite of differences at $+2\sigma$.

We should also keep in mind that the commercial software that tracks the RV along the cycle uses a surface model adjusted to the image data [21], which may reduce the amount of modes of variation in the mesh data we study.

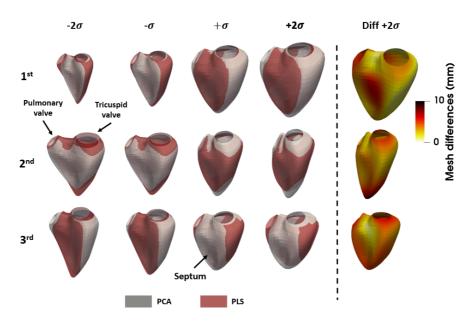


Fig. 3: Comparison of the first three modes of variations for the shape descriptor for both dimensionality reduction strategies.

The first mode of variations for area strain mostly quantifies the global amplitude of deformation (increase of the absolute strain values along the component). Lower strain is observed for samples at $+2\sigma$, which corresponds to the lower deformation observed in pulmonary hypertension cases [14] (Fig. 4). Differences

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between the two DRM (difference between the two area strain pattern at $+2\sigma$) stay under 10% except close to the pulmonary valve (top left when facing the septum). The second mode encodes more local information and in particular the location of the maximal strain on the free wall (moving from left to right) and on the septum (close to the valve, positive to negative strain). Up to 30% differences are observed in some zones of the ventricle, although these do not correspond to characteristics zones reported for this disease.

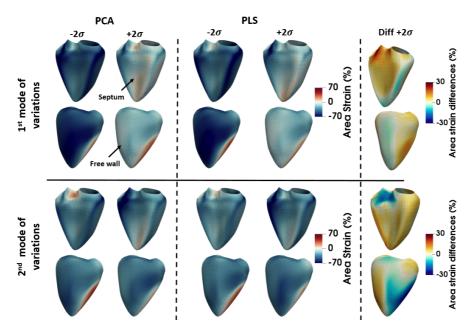


Fig. 4: Comparison of the first three modes of variations for the strain descriptor for both dimensionality reduction strategies. We show both septal and free wall views for better visualization.

4 Conclusion

In this paper, we explored ways to analyse cardiac shape and deformation, which are two high-dimensional descriptors partially interacting. We aimed at obtaining a relevant low-dimensional space that can be used later on for risk stratification. To take into account the interaction between both descriptors, we assessed two unsupervised linear dimensionality reduction strategies that consider the descriptors either independently or jointly. Similarities exist between the outputs of the two strategies (correlated first dimensions, similar patterns encoded along the first dimensions). Nevertheless, some substantial differences have been identified on zones of the right ventricle that convey relevant information for the study of pulmonary hypertension. 8

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Recent work explored supervised techniques to represent shape data against survival in the context of pulmonary hypertension [6]. We preferred unsupervised learning for our analysis, as it allows identifying hidden data structures that can be used for risk stratification (for example, identifying clusters of patients at higher risk, or situating a new patient's risk against known cases) without being conditioned by clinical labels, which may be arguable in some cases [5].

Nonetheless, PLS may be restrictive in the sense that it does not allow controlling the weight given to the interactions between descriptors, which may limit the analysis on partially interacting heterogeneous descriptors such as cardiac shape and deformation. Further work will also extend the analysis to non-linear methods, which may better preserve the structure of the data spaces associated to each descriptor. It may also include recent developments for transporting spatiotemporal shape data among a population through diffeomorphic transformations [22], which explicitly aims at preserving the structure of the space encoding cardiac meshes.

Acknowledgements

The authors acknowledge the partial support from the French ANR (LABEX PRIMES of Université de Lyon [ANR-11-LABX-0063], within the program Investissements d'Avenir [ANR-11-IDEX-0007]), and from the EEA doctoral school.

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