

# Statistical Shape Analysis for Population Studies via Level-set based Shape Morphing



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#### **Motivation**



NC













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# STG of First Episode Schizophrenics



# Objective

 Detect, Locate and Quantify spatial morphological differences between two shape populations (e.g. healthy controls and patients).

# Shape Representations and Related Methods

- Surface representation: methods such as SPHARM-PDM. Require one-to-one correspondences. [Styner 2004]
- Medial representation: more compact. Similar challenges. [Bouix 2005]
- Feature vectors: Robust but not intuitive.
   [Reuter 2006 (Shape DNA), Niethammer 2007]

# Method Outlines

- Surface representation: Signed Distance Transform
- Shape metric: modified Hausdorff distance -> does not require point-to-point correspondence.
- Shape Alignment: Align shape using 12-affine transformation by minimizing shape distances.
- Shape morphing: construct mean shape via level-set framework by minimizing shape distances.
- Shape statistics: Find statistically significant differences by calculating the signed distances to the mean shape.

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# Shape Representation

A shape  $S_i$  is defined by the image region  $\omega_i \in \Omega$ that corresponds to the structure of interest.  $S_i$  is represented by its signed distance function (SDF) :  $\phi_{S_i} : \Omega \to \mathbb{R}$ 

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We define the distance between  $S_i$  and  $S_j$ by the modified symmetrical Hausdorff distance between their boundaries:

$$\mathbf{dist}(S_i, S_j) = \int_{\partial \omega_i} |\phi_{S_j}| d\mathbf{x} + \int_{\partial \omega_j} |\phi_{S_i}| d\mathbf{x}$$

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$$D_{H}(X,Y) = \max\{\sup_{x \in X} \inf_{y \in Y} \mathbf{d}(x,y), \sup_{y \in Y} \inf_{x \in X} \mathbf{d}(x,y)\}$$

$$D_{mH}(X,Y) = \sum_{x \in X} \inf_{y \in Y} \mathbf{d}(x,y) + \sum_{y \in Y} \inf_{x \in X} \mathbf{d}(x,y)$$

$$\lim_{y \in Y} \inf_{x \in X} \mathbf{d}(x,y) = \sum_{x \in X} \inf_{y \in Y} \mathbf{d}(x,y) + \sum_{y \in Y} \inf_{x \in X} \mathbf{d}(x,y)$$



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-10

-15

-20

-25

-30

-35

$$\mathbf{dist}(S_i, S_j) = \int_{\partial \omega_i} |\phi_{S_j}| d\mathbf{x} + \int_{\partial \omega_j} |\phi_{S_i}| d\mathbf{x}$$





#### Mean Shape

We define the mean  $S^M$  of a shape ensemble  $\{S_1 \dots S_N\}$ as the shape that minimizes the sum of the distances from all the shapes in the set:

$$\hat{S}^M = rg\min_{S^M} \sum_{i=1}^N \mathbf{dist}(S_i \circ \hat{T}_{i,M}, S^M),$$

where  $T_{i,M}$  is the estimated affine (12 parameters) transform that aligns a shape  $S_i$  to the mean shape.

### Alignment of shapes



#### **Alignment of Shapes**



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#### **Smoothed Heaviside function**



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#### Level set Framework

$$\begin{split} & \int_{\partial\omega_i} \to \int_{\Omega} |\nabla H_{\epsilon}(\phi_{S_i})| \\ \mathbf{dist}(S_i, S_j) = \int_{\partial\omega_i} |\phi_{S_j}| d\mathbf{x} + \int_{\partial\omega_j} |\phi_{S_i}| d\mathbf{x} \end{split}$$

 $\mathbf{dist}(S_i, S_j) = \int_{\Omega} [|\nabla H_{\epsilon}(\phi_{S_i})| |\phi_{S_j}| + |\nabla H_{\epsilon}(\phi_{S_j})| |\phi_{S_i}|] d\mathbf{x}$ 



$$H_{\epsilon}(\phi) = \frac{1}{2} \left( 1 + \tanh\left(\frac{\phi}{2\epsilon}\right) \right) = \frac{1}{1 + e^{-\phi/\epsilon}}$$

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# Mean Shape Initialization



#### Mean Shape Morphing



# $D(S^M, \{S_i\}) = \sum_i \int_{\Omega} \left[ |\phi_{S^M}| |\nabla H_{\epsilon}(\phi_{S_i})| + \|\phi_{S_i}| |\nabla H_{\epsilon}(\phi_{S^M})| \right] d\mathbf{x}$

### Mean Shape Morphing



$$D(S^M, \{S_i\}) = \sum_i \int_{\Omega} \left[ |\phi_{S^M}| |\nabla H_{\epsilon}(\phi_{S_i})| + \|\phi_{S_i}| |\nabla H_{\epsilon}(\phi_{S^M})| \right] d\mathbf{x}$$

$$\hat{\phi}_{S^M} = \arg\min_{\phi_{S_M}} D(S^M, \{S_i\})$$

### Mean Shape Morphing

$$\begin{split} & \widehat{\phi}_{SM} = \arg\min_{\phi_{S_M}} D(S^M, \{S_i\}) \end{split}$$

$$\phi_t^M = \sum_{\text{May 8, 2013}} \left[ \text{sign}(\phi_{S^M}) |\nabla H_{\epsilon}(\phi_{S_i})| + \delta_{\epsilon}(\phi_{S^M}) \text{div}\left( \frac{\nabla \phi_{S^M}}{|\nabla \phi_{S^M}|} |\phi_{S_i}| \right) \right]_{31}$$

Population 1: 
$$\phi_{S_1}^{sz}, \dots, \phi_{S_N^{sz}}^{sz}$$
  
Population 2:  $\phi_{S_1}^{nc}, \dots, \phi_{S_N^{nc}}^{nc}$ 

For each voxel on the boundary of the mean shape calculate:

$$\vec{d}^{\rm sz}(\mathbf{x}) = \{ d(\mathbf{x} \in \partial \omega^M, \partial \omega_1^{\rm sz}), \dots, d(\mathbf{x} \in \partial \omega^M, \partial \omega_N^{\rm sz}) \}$$

$$\vec{d}^{\rm nc}(\mathbf{x}) = \{ d(\mathbf{x} \in \partial \omega^M, \partial \omega_1^{\rm nc}), \dots, d(\mathbf{x} \in \partial \omega^M, \partial \omega_N^{\rm nc}) \}$$

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$$= \{ \phi_{S_1}^{\rm nc}(\mathbf{x}), \dots \phi_{S_N^{\rm nc}}^{\rm nc}(\mathbf{x}) \}$$

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For each voxel on the boundary of the mean shape apply the

two-sample t-test on  $\vec{d}^{\rm sz}({\bf x})$  and  $\vec{d}^{\rm nc}({\bf x})$ .

The p-value threshold is determined by using



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# Shape Analysis Algorithm

<u>Input:</u> two populations of shapes (e.g. brain structures). <u>Algorithm:</u>

- 1. Use probability maps of the SDTs to initialize the mean shape
- 2. Iterate
- a. Align the shapes represented by their SDTs by minimizing the modified Hausdorff distances (mHDs).
- b. Morph the mean shape to minimize the mHDs to the shape ensemble.
- 3. Calculate the signed distances between the mean shape surface to each shape in the two populations.
- 4. Use two-sample t-test to find statistically significant differences between the two populations.
  - Use False Discovery Rate (FDR) to correct for multiple comparisons.
- <u>Output:</u> Highlighted regions on the mean shape surface for which statistically significant differences were found.

## Results: Synthetic Amygdala-Hippocampus Complexes

#### **Bump databases**



# Results: Synthetic Amygdala-Hippocampus Complexes

#### **Dimple databases**



# Results: Synthetic Amygdala-Hippocampus Complexes



Radius of deformation

### **Results: Synthetic Striatum**



Gao, Y., Bouix, S., October 2012. Synthesis of realistic subcortical anatomy with known surface deformations. In: MICCAI Workshop on Mesh Processing in Medical Image Analysis.

### **Results: Synthetic Striatum**



# Shape Analysis of First-episode Schizophrenics

- Lower left temporal lobe MRI volumes in patients with first-episode schizophrenia compared with psychotic patients with first-episode affective disorder and normal subjects Y. Hirayasu et. al. Am J Psychiatry 1998
- Planum temporale and heschl's gyrus volume reduction in schizophrenia: A MRI study of first-episode patients Y. Hirayasu *et. al.* Arch Gen Psychiatry 2000
- Hippocampus and superior temporal gyrus volume in first-episode schizophrenia Y. Hirayasu et. al. Arch Gen Psychiatry 2000

# Results: Left STG of First Episode Schizophrenics



Mean left STG

Mean left STG with p-values color map

# Results: Caudate of Schizotypal Personality Disorder (SPD) Subjects





Mean right caudate with p-values color map

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# Summary and Future Work

- Detection, localization and quantification of shape Deformations for population studies.
- One-to-one point correspondences are not require!
- -> Low computational complexity; robustness
- Intuitive output
- Handles shapes with complicated morphology
- Does not require pre-processing such as smoothing.
- Next step:
- Incorporate the concept of boundary/surface uncertainty within the analysis.

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