

*The Waisman Laboratory
for Brain Imaging and Behavior*

Fourier Spectral Method for Shape Asymmetry Analysis

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Acknowledgments

Kim M. Dalton, Richard J. Davidson

Waisman laboratory for brain imaging and behavior

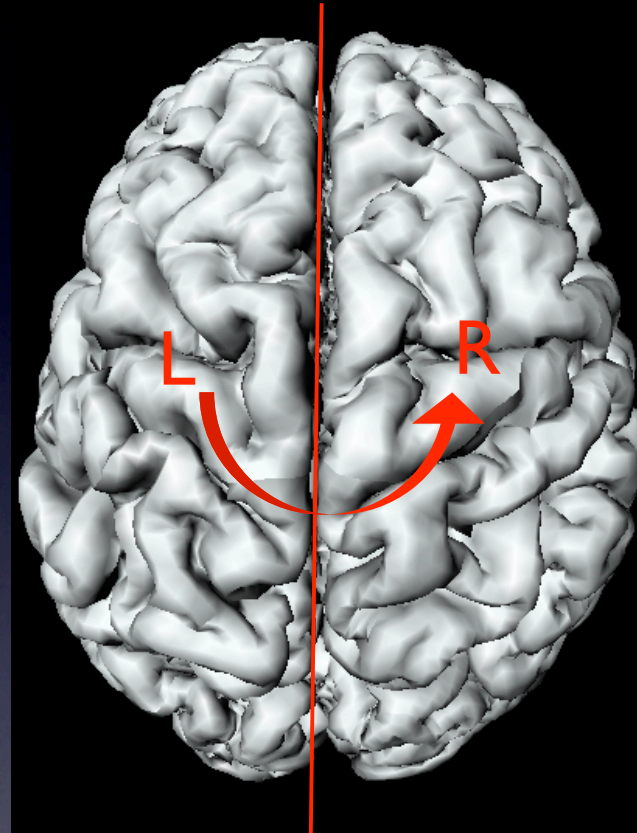
Shubing Wang, Dongjun Chung, Houri Vorperian

Vocal Tract Development Laboratory

Abstracts

Although shape asymmetry has been investigated in many branches of science, there is a lack of unified methodological framework for quantifying local shape asymmetry. Previous literature mainly deals with quantifying a global amount of shape asymmetry. A more interesting question would be to ask if we could spatially localize the source of asymmetry. In brain imaging, this question has been successfully addressed by using the deformable template approach of Grenander and Miller. By registering the original and its mirror reflected 3D magnetic resonance image (MRI), one can establish the correspondence across brain hemispheres and, in turn, able to construct the localize asymmetry index of type $(L-R)/(L+R)$. The additional computational burden of establishing deformation across hemispheres and possible mismatching of sulcal pattern across subjects are two major shortcomings of this widely used approach. In this talk, we present a different framework for shape asymmetry analysis that basically combines the deformable template idea and Brechbuler's 3D Fourier descriptor. Surface shape registration, surface data smoothing and surface parameterizations are all tackled in a unified framework. This is a joint work with Kim Dalton and Richard Davidson of the Waisman Laboratory for Brain Imaging and Behavior. An application of the same technique to longitudinal mandible shape modeling (in collaboration with Houri Vorperian of the Vocal Tract Development Laboratory) on 300 subjects will be also briefly discussed.

Brain hemisphere asymmetry



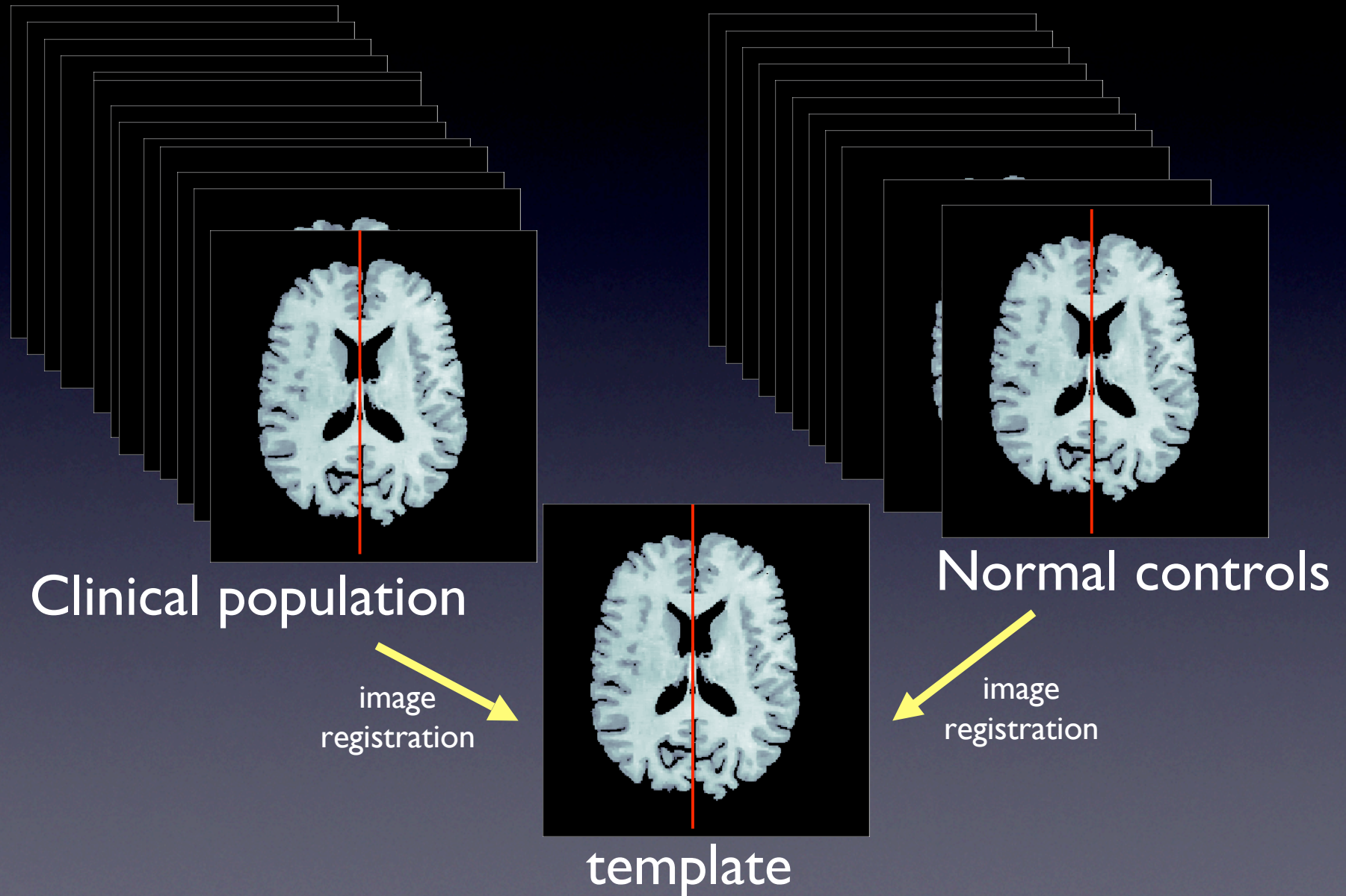
Localized
asymmetry
index
 $(L-R)/(L+R)$

Motivation: quantify abnormal brain structural asymmetry across hemispheres in a group of autistic subjects

Previous 3D approach

1. Image registration across subjects via a template
2. Image registration across hemispheres by registering the original MRI and its mirror reflection.
3. Construct asymmetry index at each voxel.
4. Feed the index into a statistical model.

Two population asymmetry analysis framework



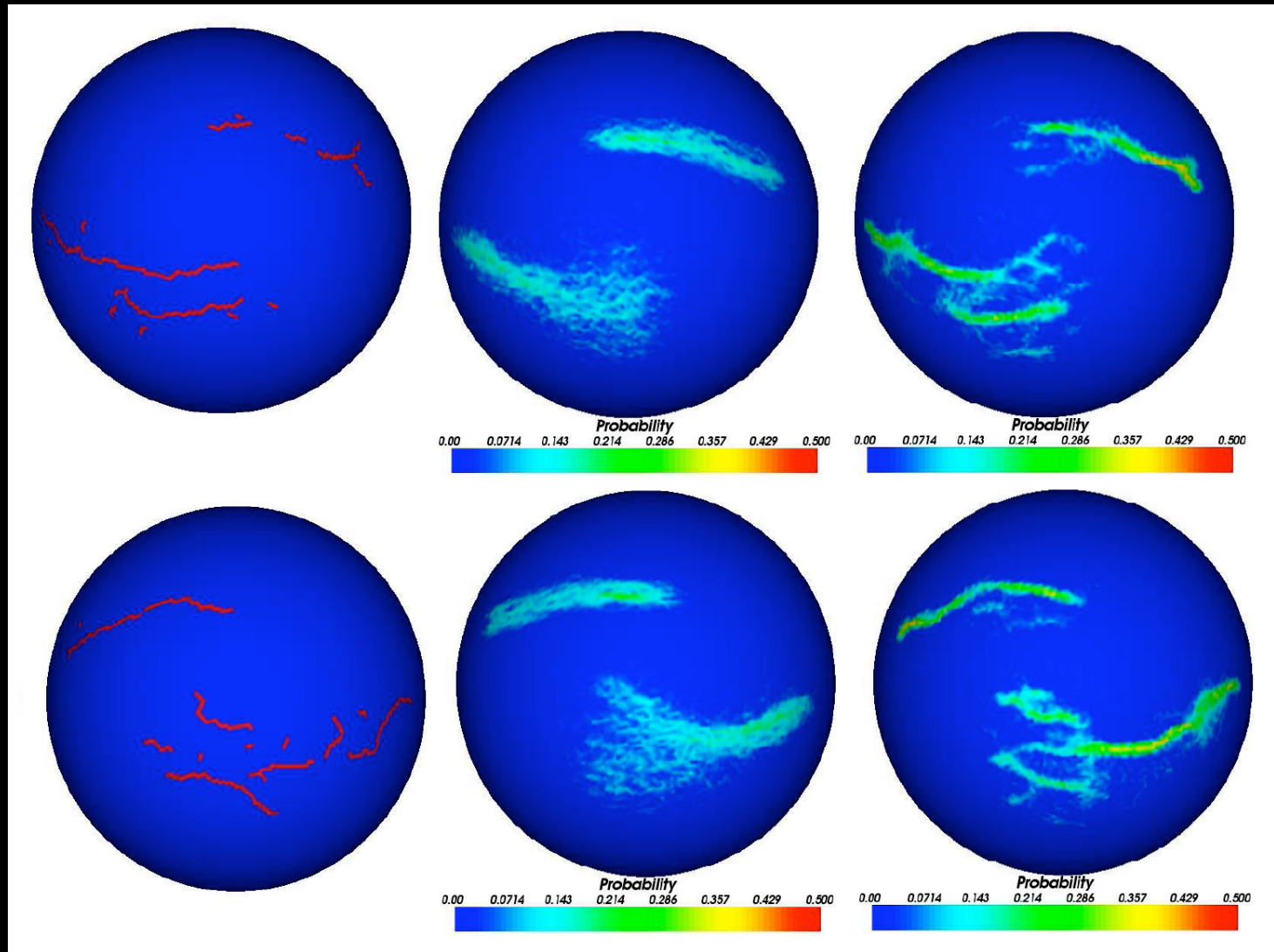
Three issues with this well established 3D approach

1. 3D image registration can easily misalign sulcal pattern.
2. Mirror reflection and doing image registration is an additional computational burden.
3. The 3D approach does not work for 2D cortical surface data. **New 2D framework is needed.**

Comparison of surface registration on 149 subjects

Left
central &
temporal
sulci

Right
central &
temporal
sulci

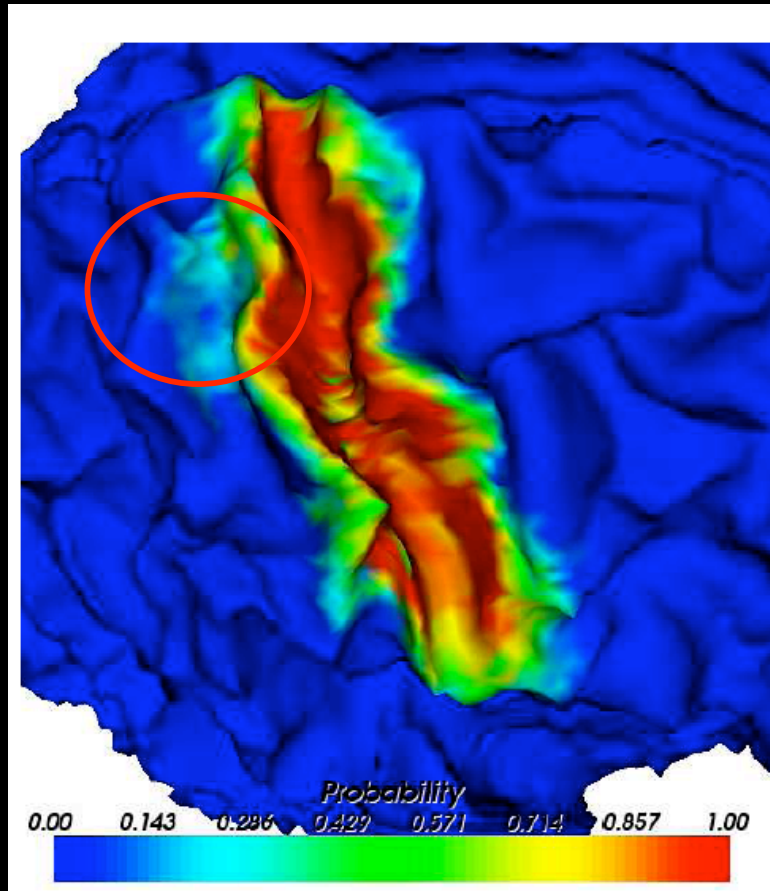


NeuroImage (2003)

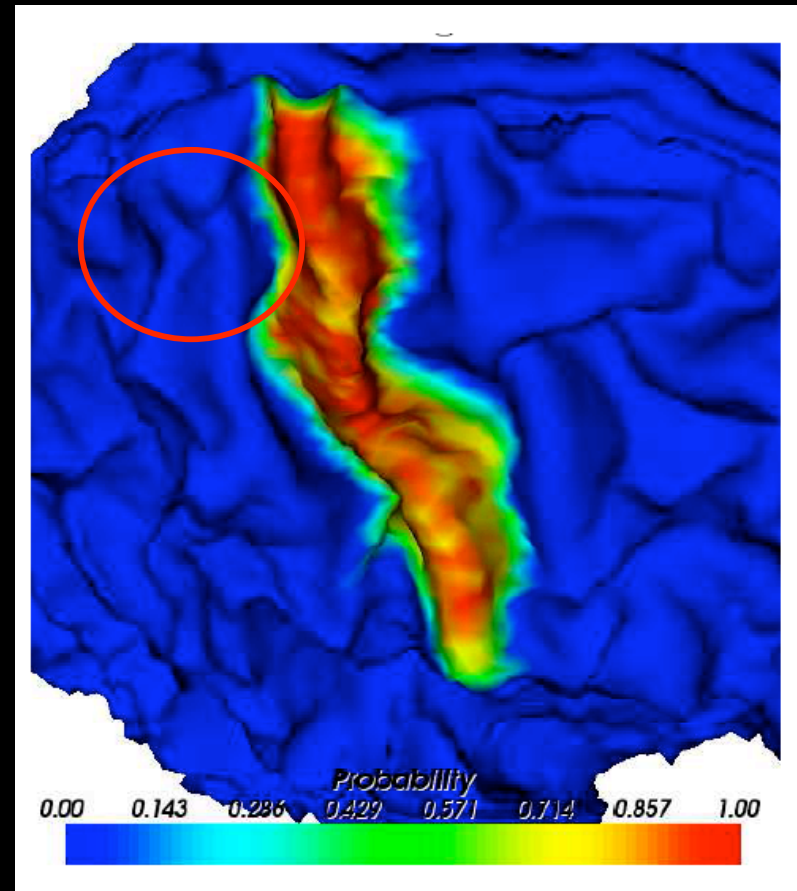
3D registration

2D registration

Probability of matching in the right central sulcus



3D volume registration



2D surface registration

Literature vs. new framework

Surface data
smoothing

diffusion smoothing
(NeuroImage, 2003)
heat kernel
smoothing
(NeuroImage, 2005)

Surface
parameterization

SPHARM
Guido Gerig
Martin Styner
Li Shen

Surface
registration

PDE
Paul Thompson
Michael Miller

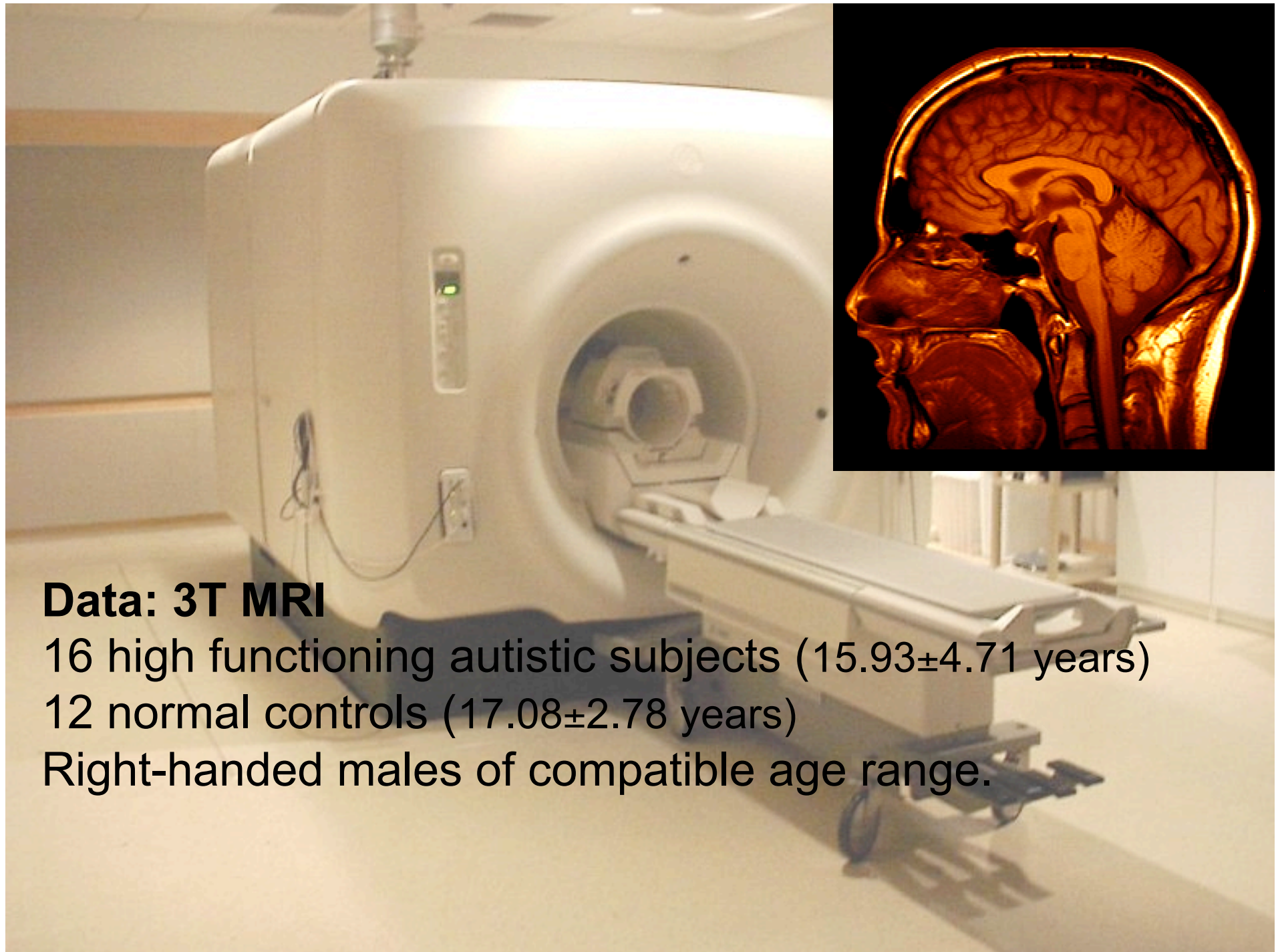
Multiple
comparison
correction

Random field
theory
Keith Worsley
Jonathan Taylor

New unified approach:
Weighted spherical harmonic representation
(TMI, 2007)

Outline of talk

1. Introduction to cortical surface data
2. Weighted Fourier series representation
3. Surface registration
4. Surface asymmetry index
5. Statistical analysis
6. Future research direction



Data: 3T MRI

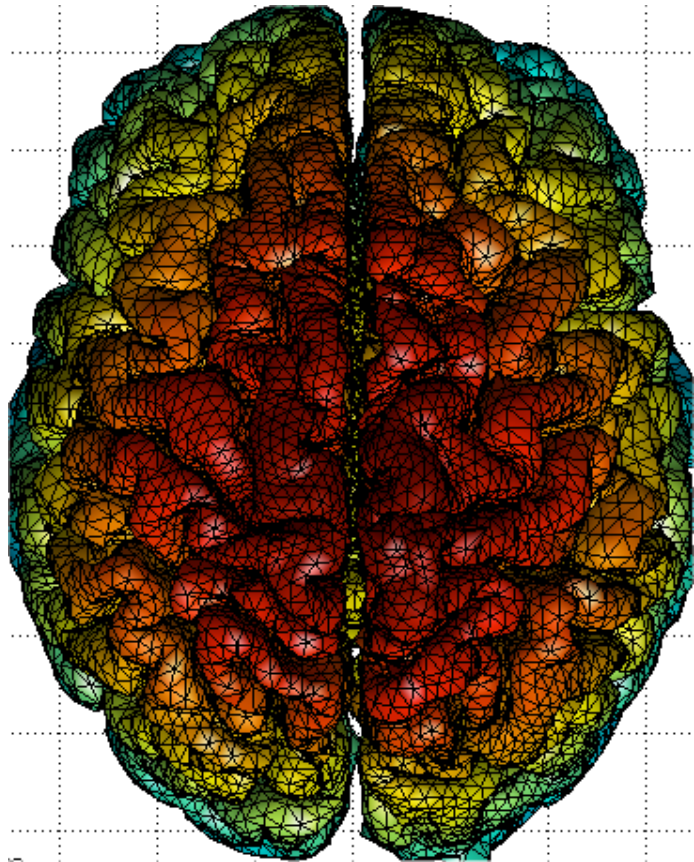
16 high functioning autistic subjects (15.93 ± 4.71 years)

12 normal controls (17.08 ± 2.78 years)

Right-handed males of compatible age range.

Polygonal mesh

Mesh resolution 3mm



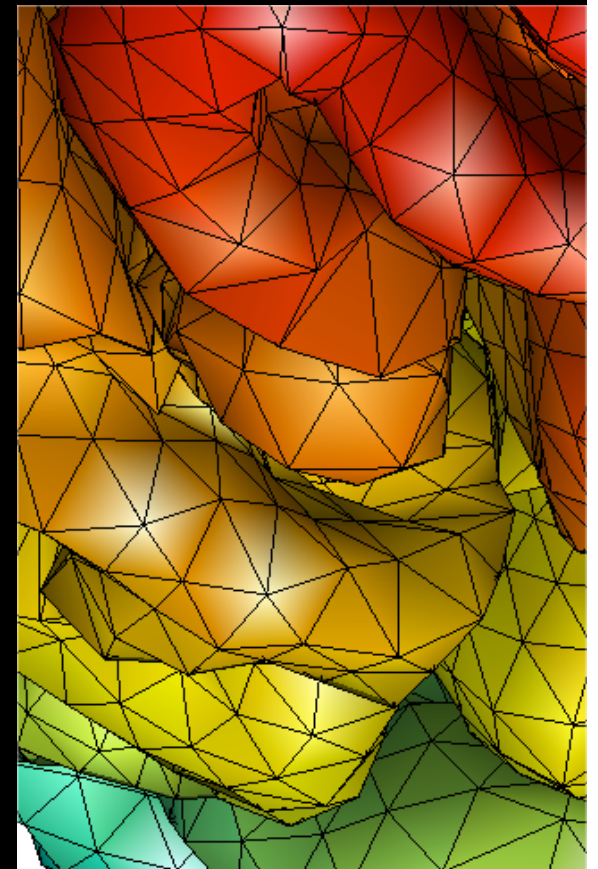
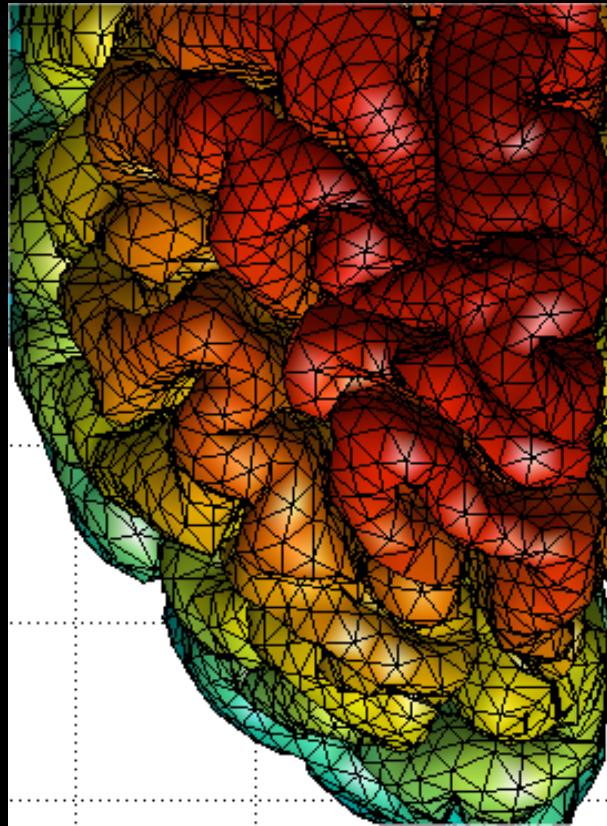
82,190 triangles

40,962 vertices

Our method

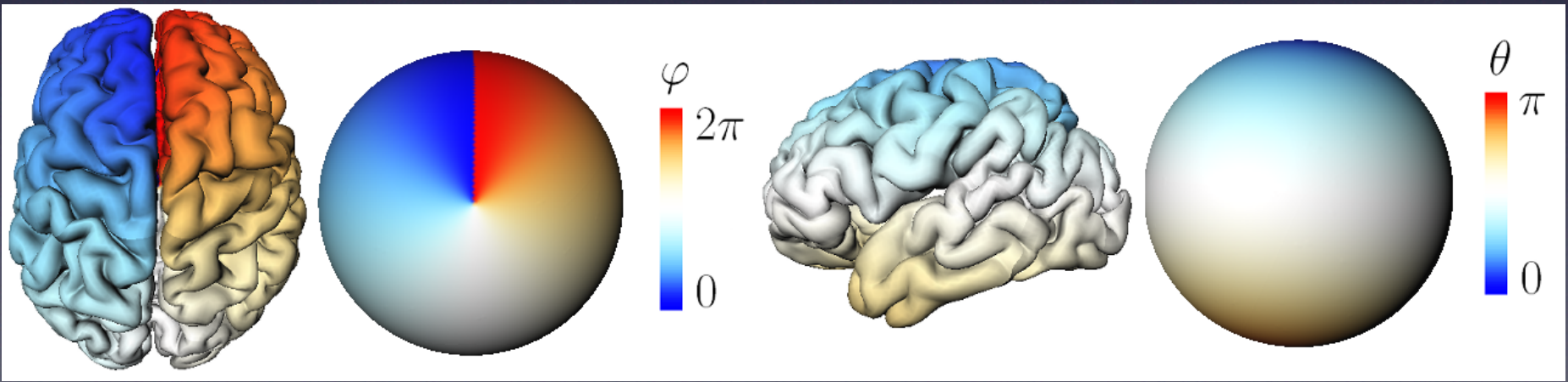


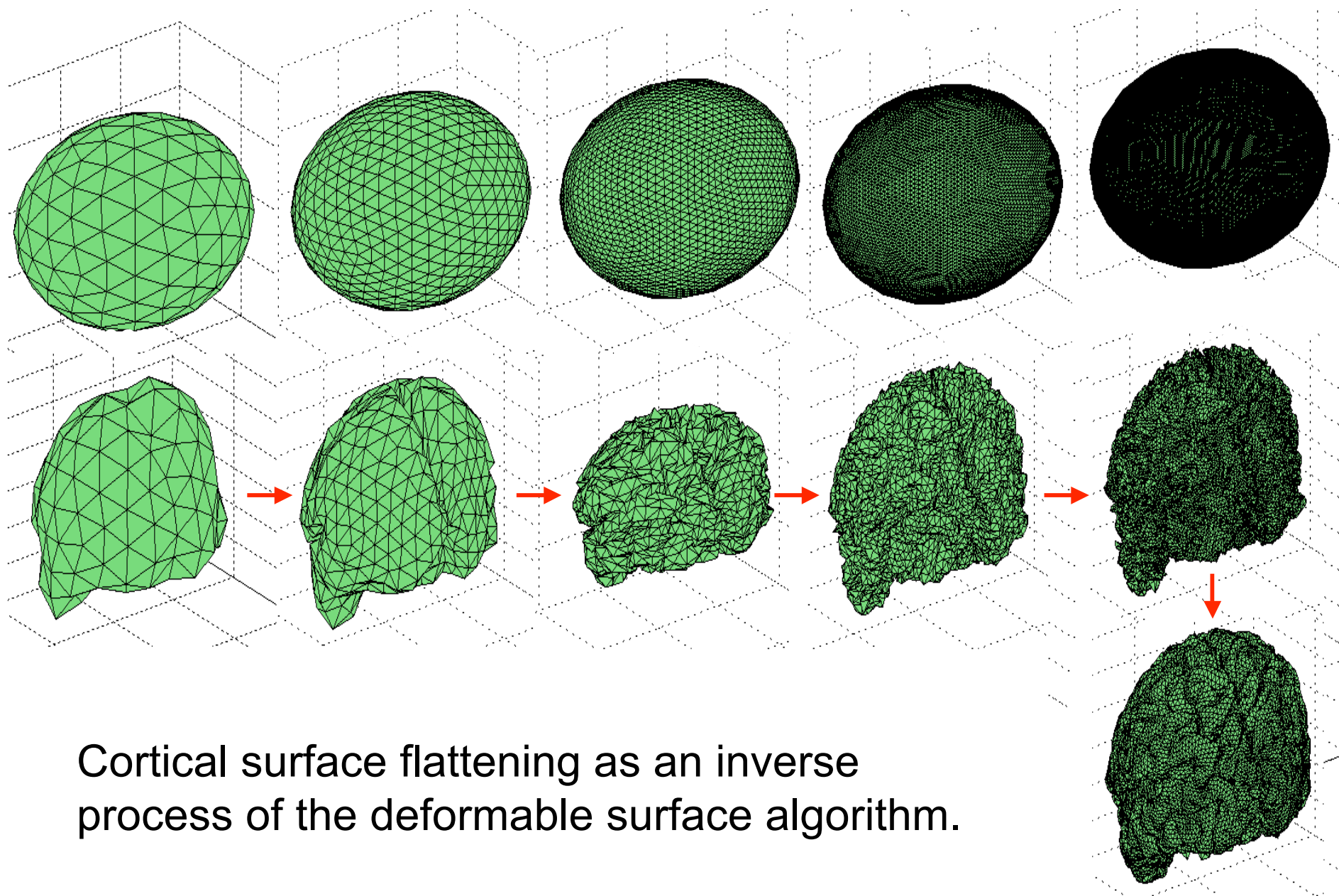
20,000 parameters per surface



Cortical surface flattening

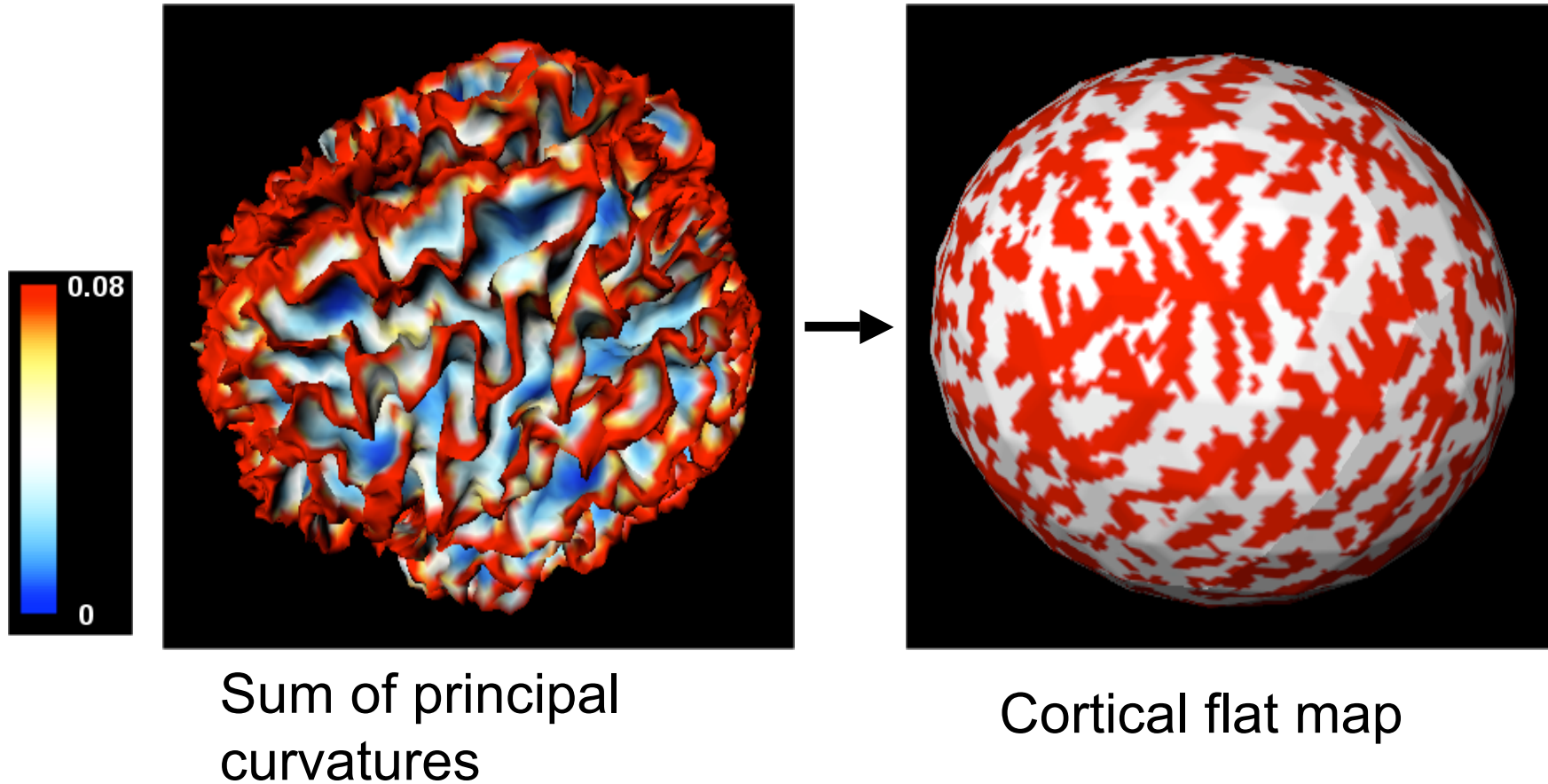
- Deformable surface algorithm (McDonalds et al., 2001) is used to segment surfaces and obtain the mapping from a unit sphere to a cortical surface.
- Functional measurement defined on cortical surface will be pulled back onto the unit sphere.





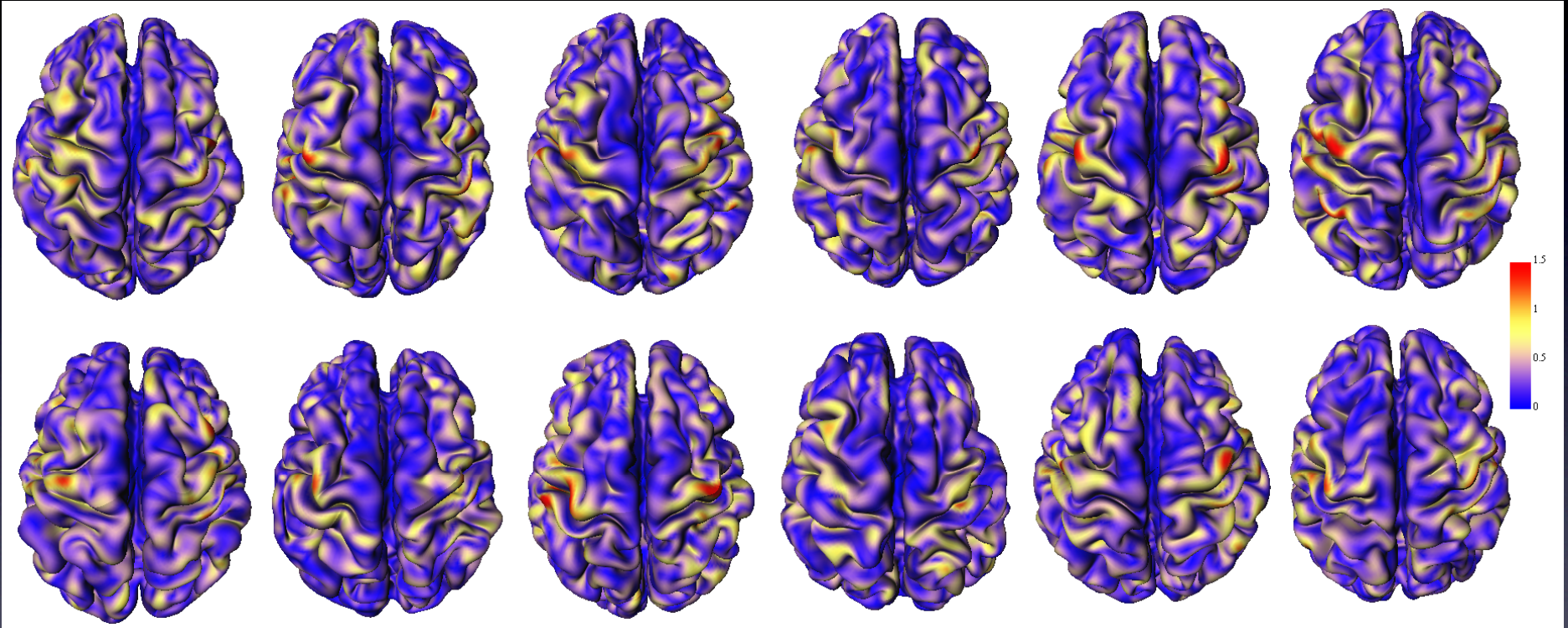
Cortical surface flattening as an inverse process of the deformable surface algorithm.

Example of functional measurement pulled back onto unit sphere



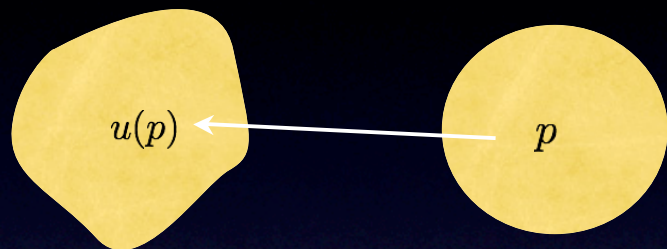
Note: metric distortion might influence the final statistical analysis.

There is a way to address area distortion
Local area element can be obtained by analytically
differentiating VFS and computing metric tensors.



This measures amount of area distortion associated with cortical flattening. It can be used as a nuisance covariate in a statistical analysis.

Cortical manifold and function measurement defined on the manifold



manifold \mathcal{M}

parameter space \mathcal{N}

Anatomical manifold $\mathcal{M} \in \mathbb{R}^d$

Parameter space $\mathcal{N} \in \mathbb{R}^m$

Parameterization $u : \mathcal{N} \rightarrow \mathcal{M}$

Hilbert space $L^2(\mathcal{N})$ with inner product

$$\langle g_1, g_2 \rangle = \int_{\mathcal{N}} g_1(p) g_2(p) \mu(p)$$

Self-adjoint operator \mathcal{L}

$$\langle \mathcal{L} g_1, g_2 \rangle = \langle g_1, \mathcal{L} g_2 \rangle$$



Basis function

$$\mathcal{L}\psi_j = \lambda_j\psi_j$$

Weighted Fourier Series (WFS) representation

cortical thickness
(function)
+ surface coordinates
(surface)

$t = \text{scale, bandwidth}$

PDE

$$\partial_t g + \mathcal{L}g = 0, g(p, t = 0) = f(p)$$

solution

$$g(p, t) = \sum_{j=0}^{\infty} e^{-\lambda_j t} \langle f, \psi_j \rangle \psi_j(p)$$

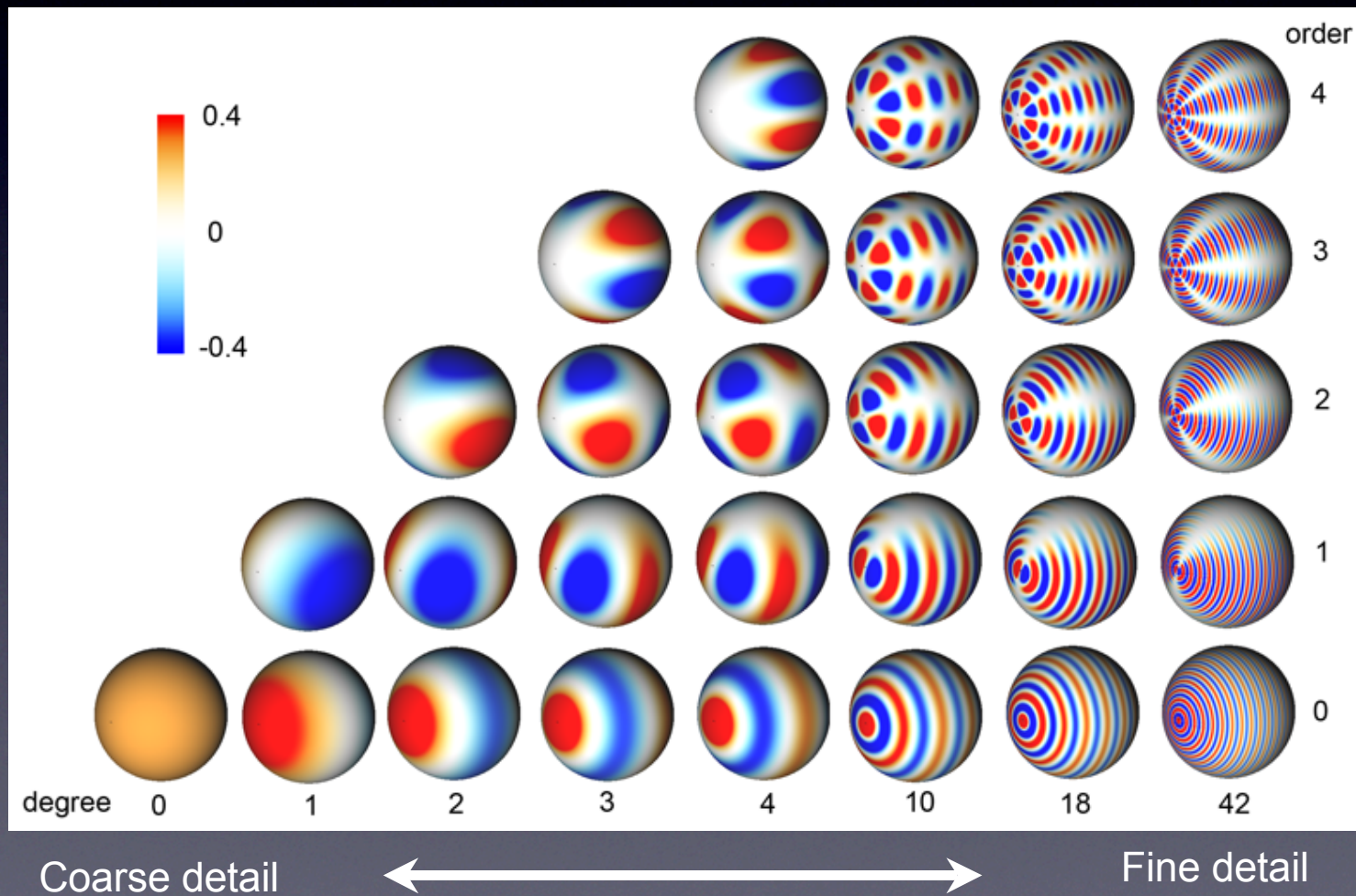
$$= \int_{\mathcal{N}} K_t(p, q) f(q) d\mu(q)$$

Spectral
representation

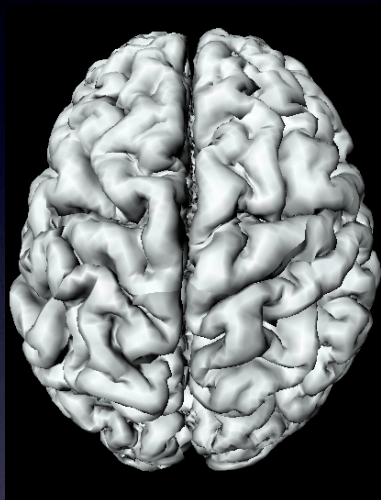
Kernel smoothing

Spherical harmonic of degree l and order m

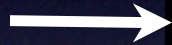
$$Y_{lm} = \begin{cases} c_{lm} P_l^{|m|}(\cos \theta) \sin(|m|\varphi), & -l \leq m \leq -1, \\ \frac{c_{lm}}{\sqrt{2}} P_l^0(\cos \theta), & m = 0, \\ c_{lm} P_l^{|m|}(\cos \theta) \cos(|m|\varphi), & 1 \leq m \leq l, \end{cases}$$



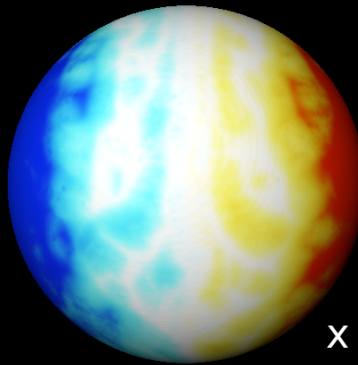
WFS = parameterization + smoothing



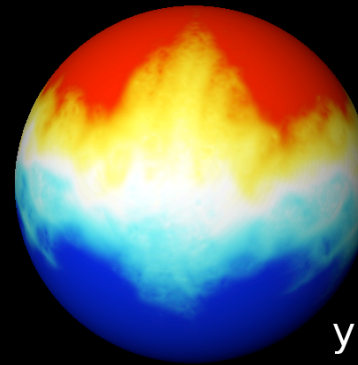
Original
cortical
surface



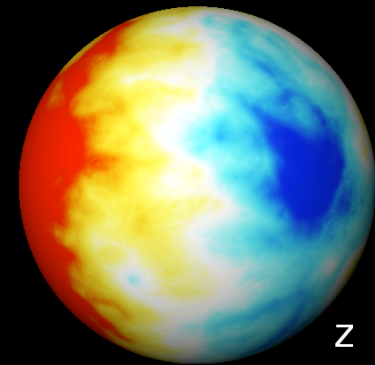
Coordinate functions



x

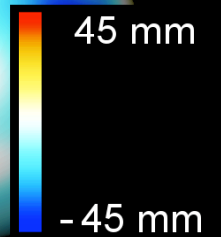
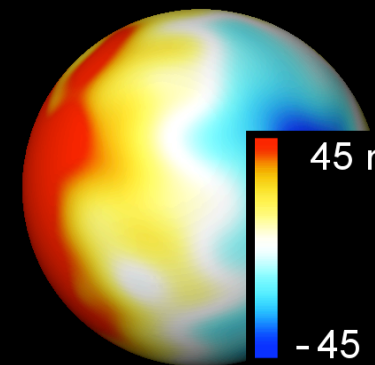
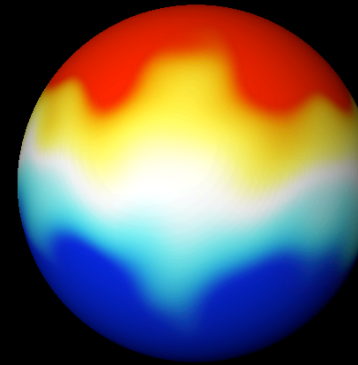
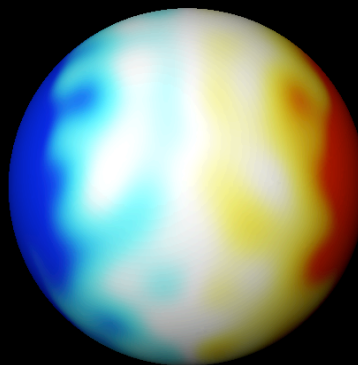


y

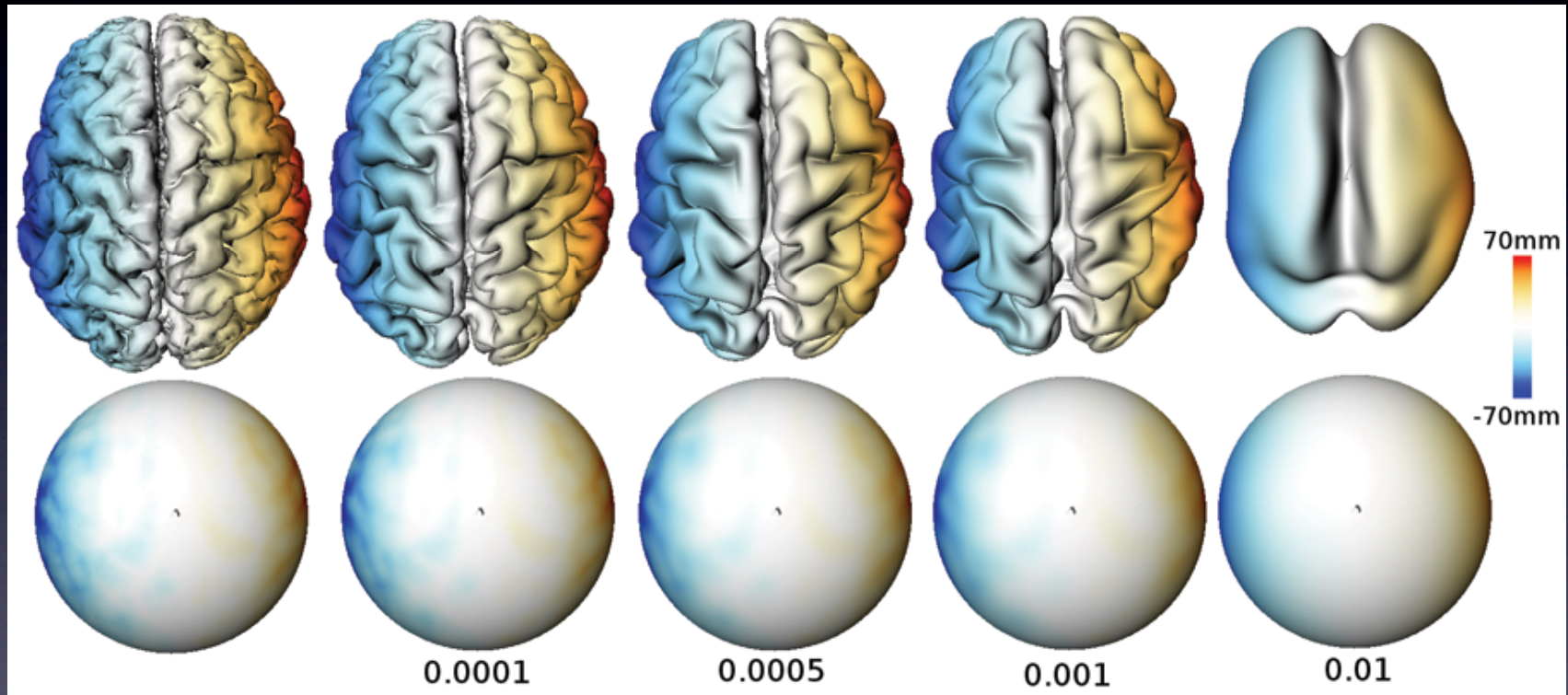


z

Weighted-SPHARM



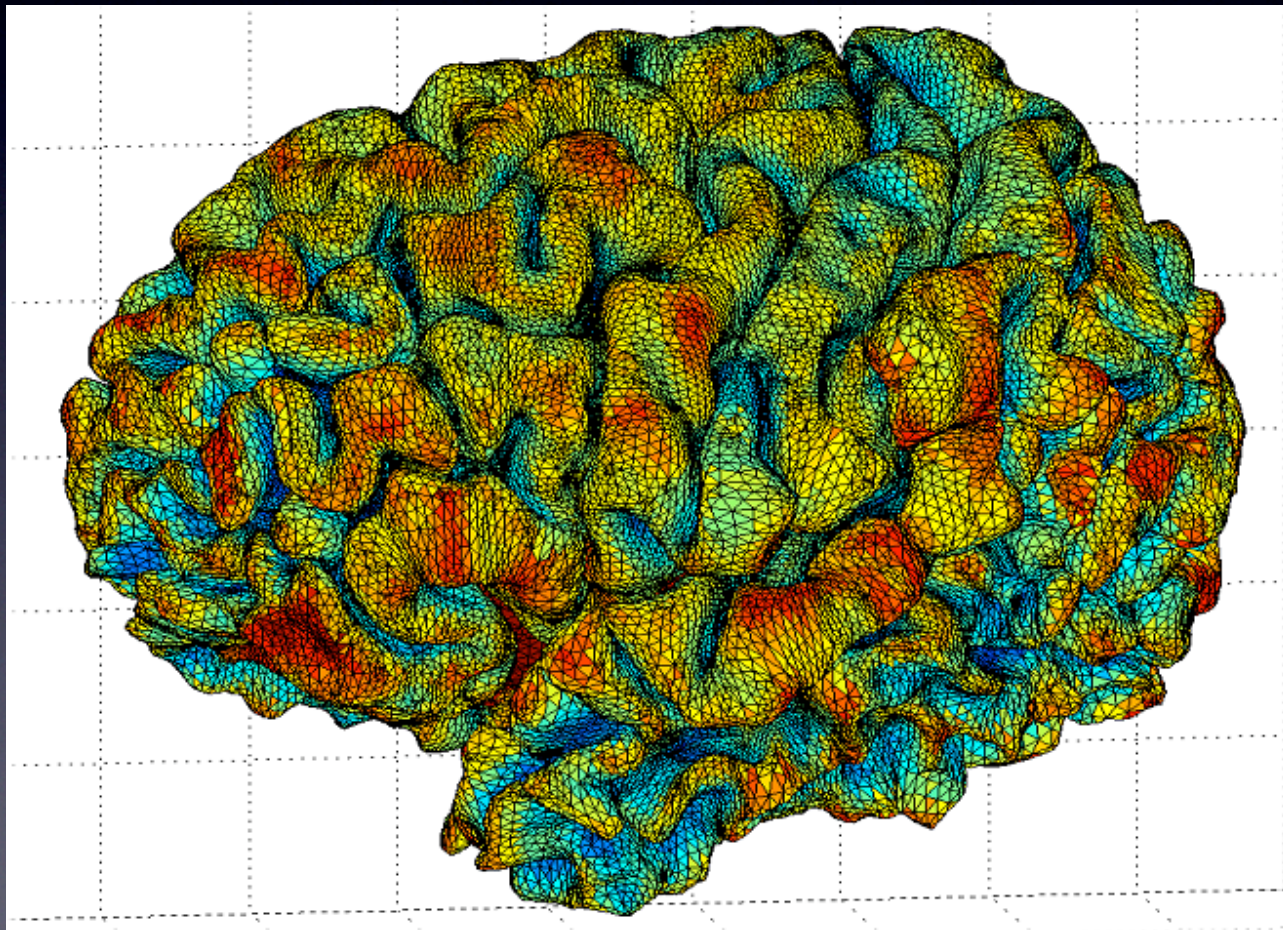
WFS = multiscale representation



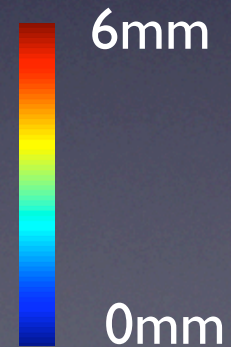
Color scale= x-coordinate

WFS can be applied to functional data like cortical thickness.

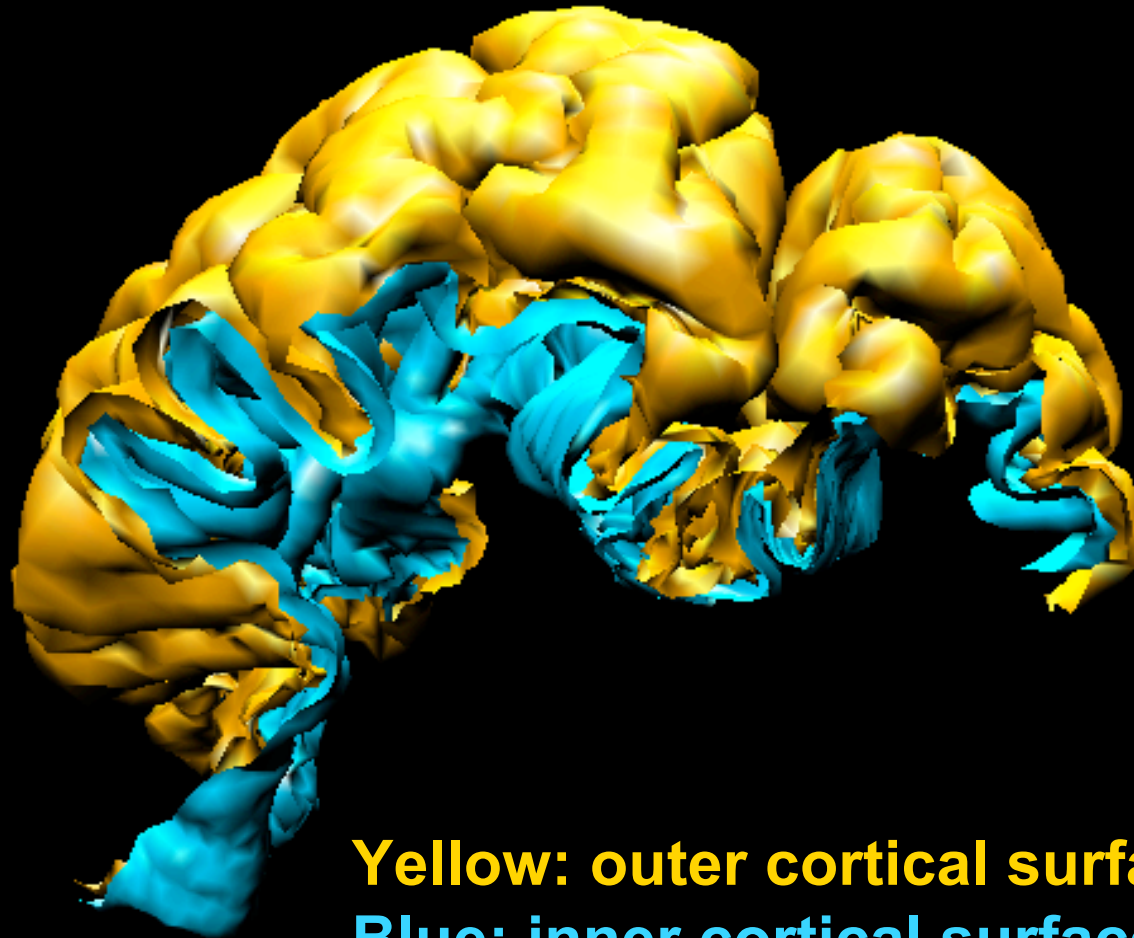
Cortical thickness = most widely used cortical structural measure



Cortical thickness



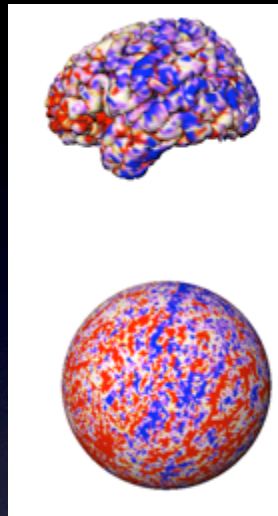
What is cortical thickness ?



Yellow: outer cortical surface
Blue: inner cortical surface

- distance between surfaces
- measures amount of gray matter bounded by these two surfaces

WFS of cortical thickness



Cortical thickness

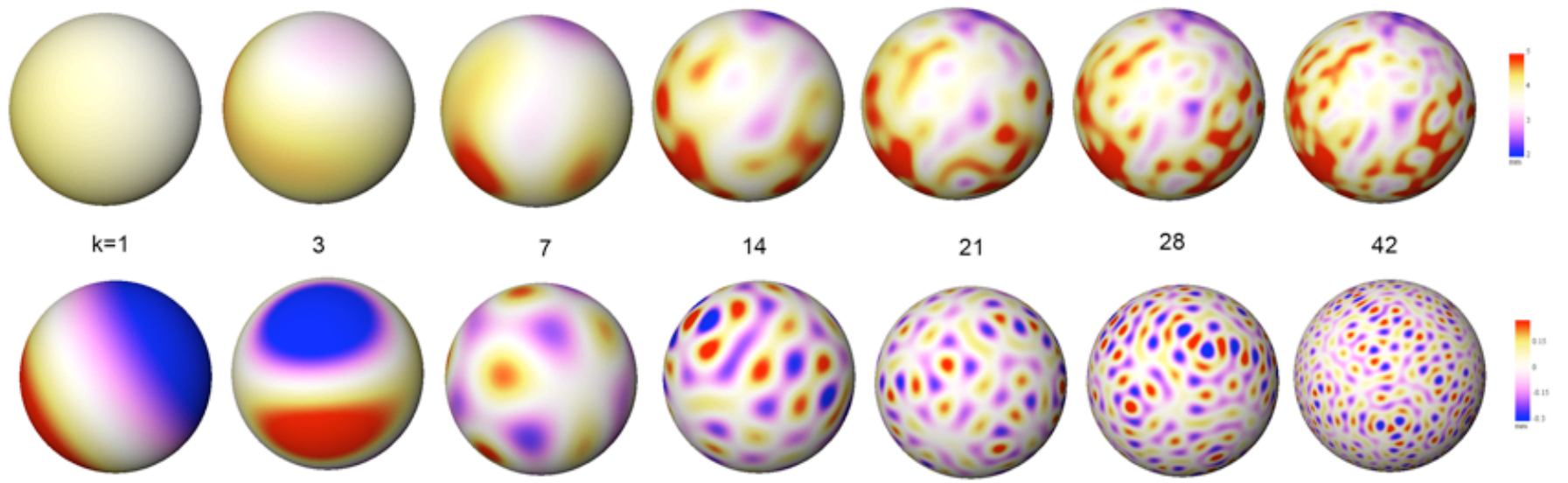


Pull back



1st row:
$$\sum_{l=0}^k \sum_{m=-l}^l e^{-l(l+1)t} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi)$$

2nd row:
$$\sum_{m=-k}^k e^{-l(l+1)t} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi)$$



Iterative residual fitting (IRF) algorithm (TMI, 2007)

Can estimate more than 20,000 coefficients per surface

Joint work with Li Shen

Step 1. measurements $f(p_1), \dots, f(p_n)$

Step 2. Set initial degree=0 $k = 0$

Step 3. Solve $f(p_i) = \sum_{m=-k}^k \beta_{km} Y_{km}(p_i)$ Project data into a finite subspace

Iterate Step 3.5. $f \leftarrow f - \hat{f}$ Once low frequency parts are estimated, we throw them away

Step 4. Set degree $k \leftarrow k + 1$

MATLAB code available at <http://www.stat.wisc.edu/~mchung/>

Similar method in literature: Matching pursuit (MP) method (Mallat and Zhang, 1993, IEEE Trans. Signal Processing)

MR is identical to IRF in principle except the methods for estimating Fourier coefficients are different.

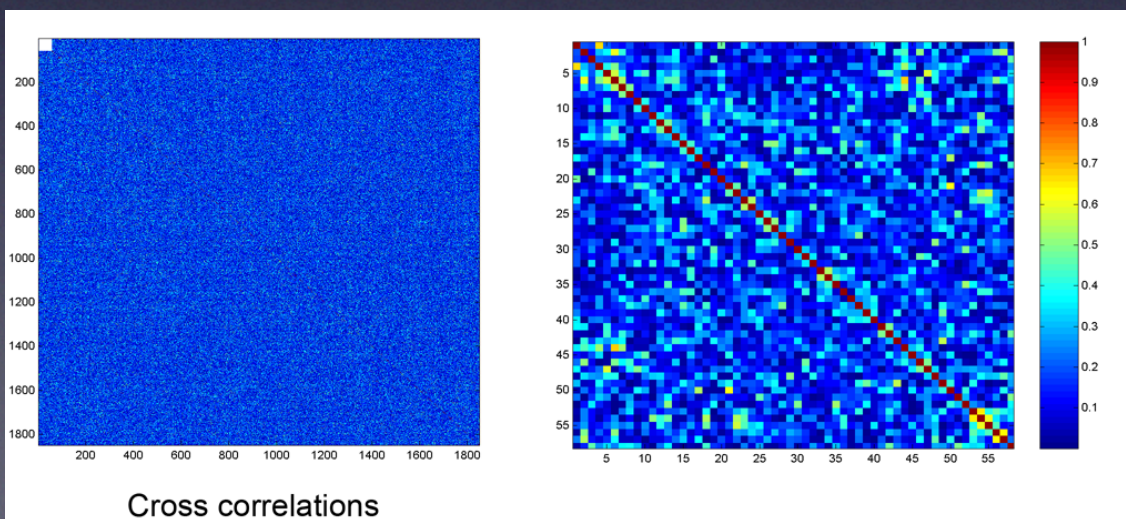
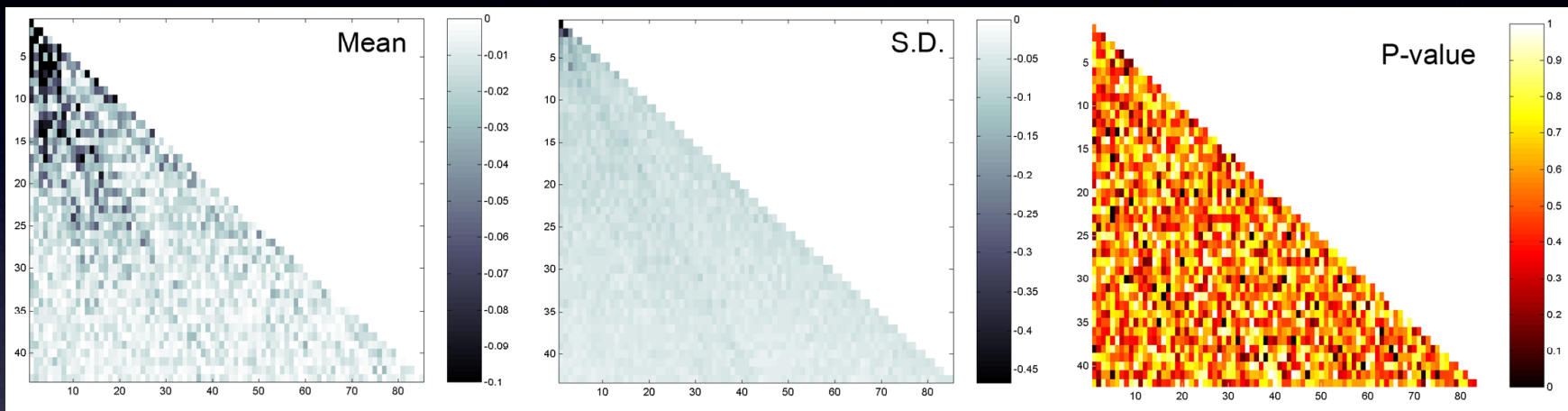
IRF was developed to solve a large linear system (with orthonormality constraint) iteratively.

MP was developed as a way to compactly decompose time frequency signal into a linear combination of basis in a dictionary.

Statistical Model on WFS: Karhunen-Loeve expansion

$$\sum_{l=0}^k \sum_{m=-l}^l e^{-l(l+1)t} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi)$$

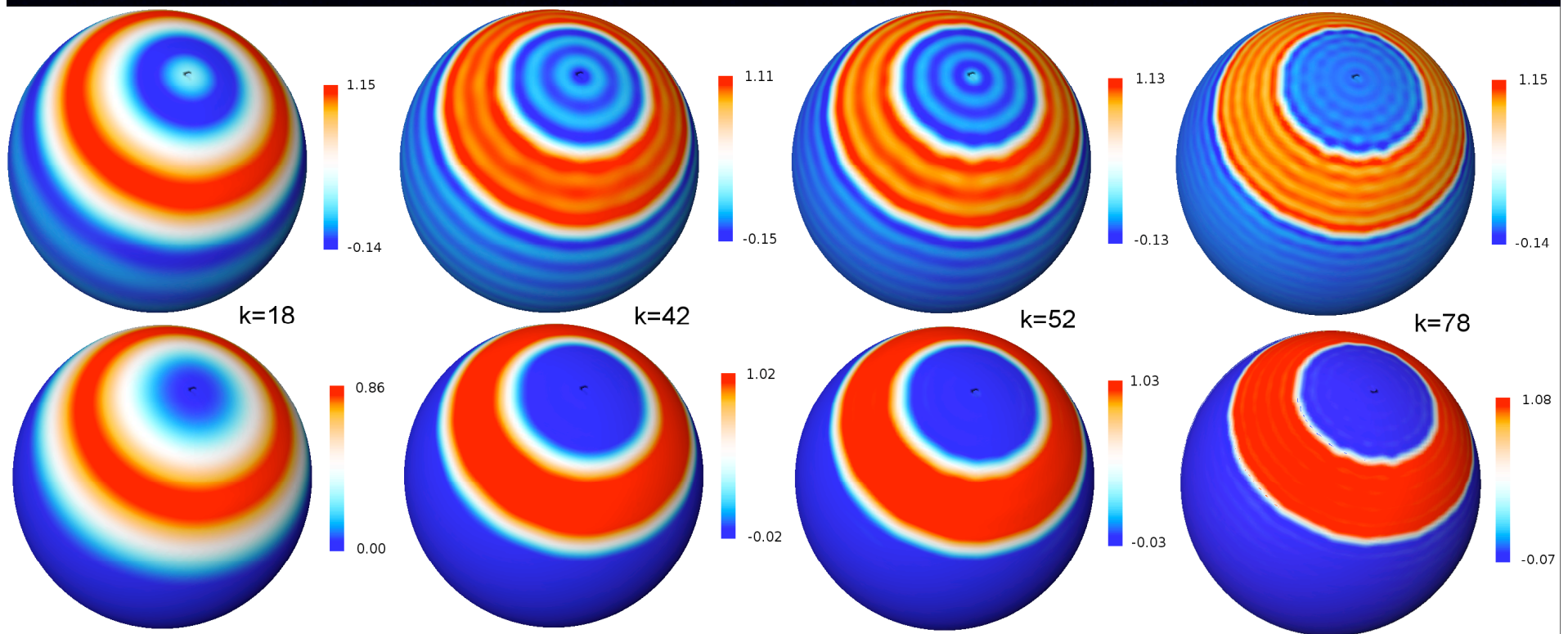
← Uncorrelated normal



Why WFS ?

Reduction of Gibbs phenomenon (ringing artifacts)

Functional data defined on sphere

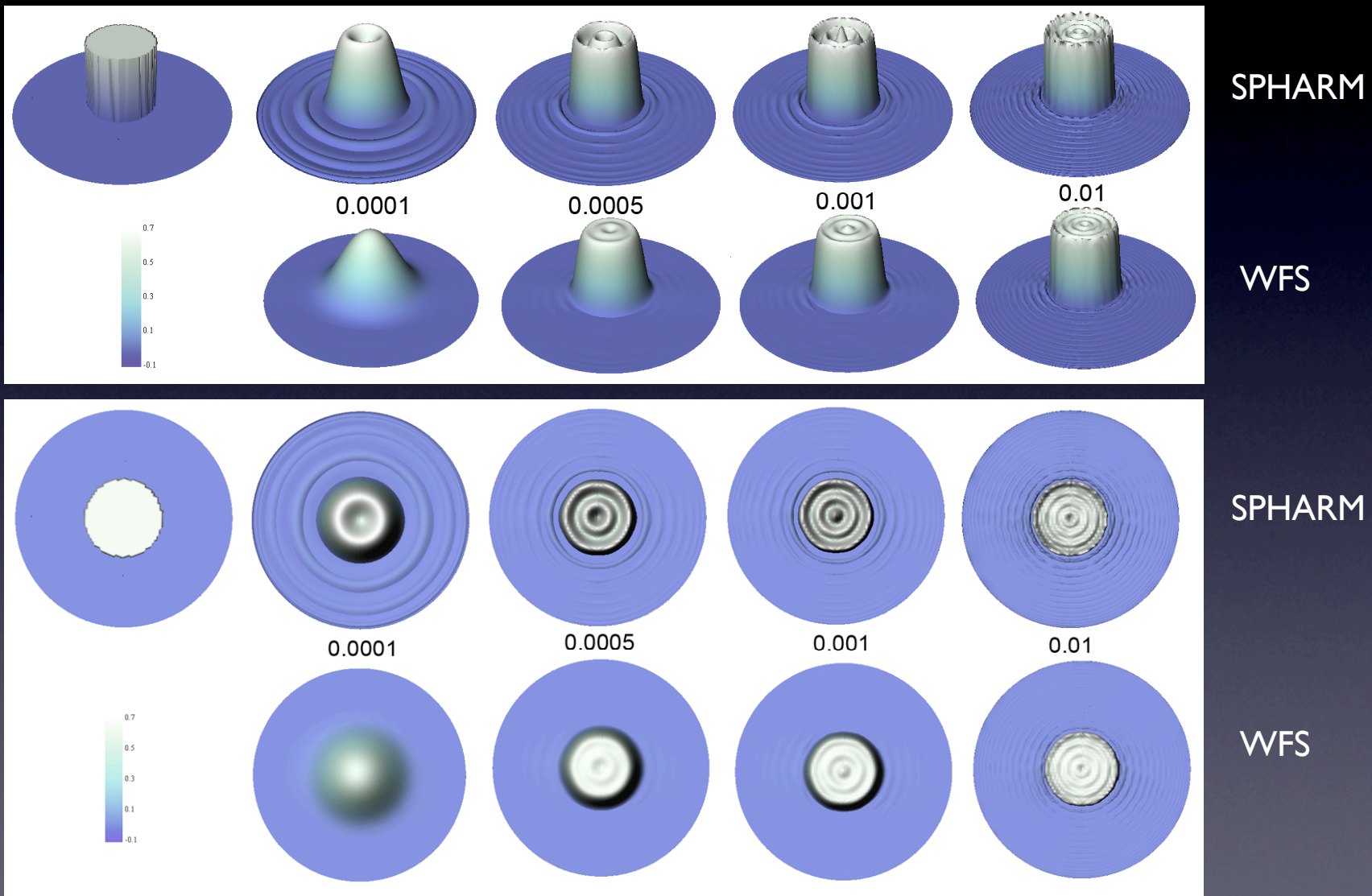


Top: Fourier series expansion (SPHARM)

Bottom: WFS

Why WFS ? Gibbs phenomenon

Anatomical data



Surface registration via WFS

Given two l -th degree WFS surfaces v_{i1}, v_{i2}
find the displacement d_i that minimizes the
discrepancy between two surfaces:

$$v_{i2} - v_{i1} = \arg \min_{d_i \in \mathcal{H}_l} \int_{\mathcal{M}} [v_{i1} + d_i(v_{i1}) - v_{i2}]^2 d\mu(p).$$

\mathcal{H}_l : subspace spanned by up to l -th degree spherical harmonics

$v_{i1} + d_i(v_{i1})$: deformation of coordinates v_{i1}

Consequence: For fixed (θ, φ) ,

$v_{i1}(\theta, \varphi)$ corresponds to $v_{i2}(\theta, \varphi)$.

Example of surface registration



subject 1

$$\alpha = 0$$

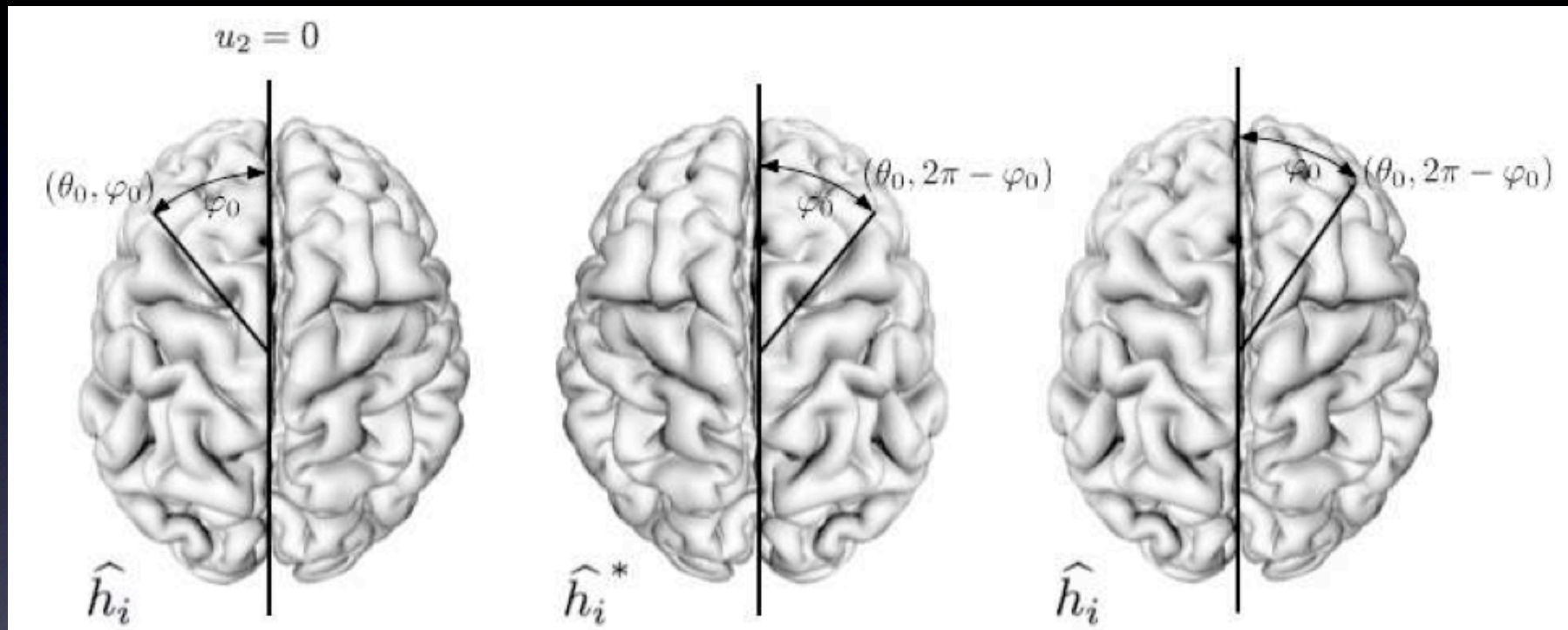
$$v_{i1} + \alpha d_i(v_{i1})$$

subject 2

$$\alpha = 1$$

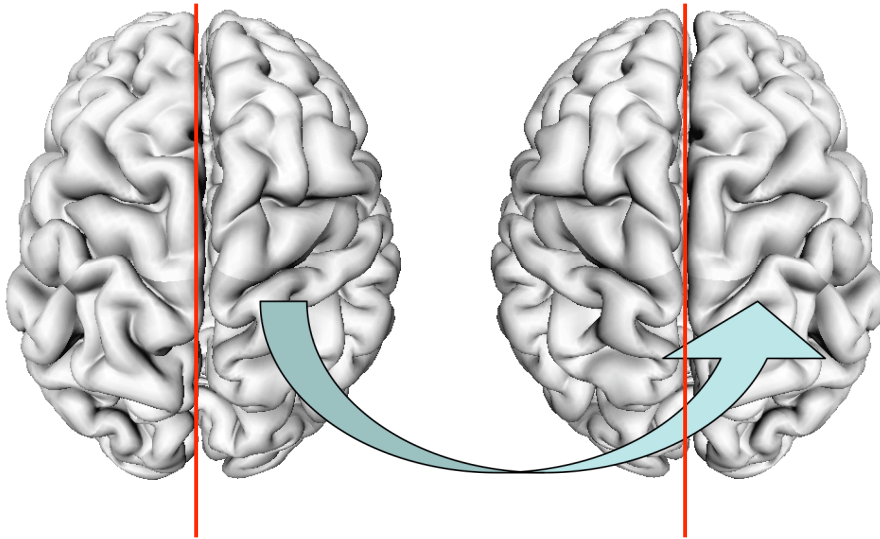
Cortical asymmetry analysis

Establishing hemispheric correspondence algebraically



Mirror reflection: It is done algebraically on WFS

Surface registration



Invariance under
mirror reflection

$$\hat{g}(\theta, \varphi) = \sum_{l=0}^k \sum_{m=-l}^l e^{-l(l+1)\sigma} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi).$$

$$\begin{aligned} \hat{g}(\theta, 2\pi - \varphi) &= \sum_{l=0}^k \sum_{m=-l}^{-1} e^{-l(l+1)\sigma} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi) \\ &\quad - \sum_{l=0}^k \sum_{m=0}^l e^{-l(l+1)\sigma} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi) \end{aligned}$$

Mirror
reflection

Shape decomposition into symmetric and asymmetric parts


Symmetric part

$$S(\theta, \varphi) = \frac{1}{2} \left[\hat{g}(\theta, \varphi) + \hat{g}(\theta, 2\pi - \varphi) \right] = \sum_{l=0}^k \sum_{m=-l}^{-1} e^{-l(l+1)\sigma} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi)$$

Asymmetric part

$$A(\theta, \varphi) = \frac{1}{2} \left[\hat{g}(\theta, \varphi) - \hat{g}(\theta, 2\pi - \varphi) \right] = \sum_{l=0}^k \sum_{m=0}^l e^{-l(l+1)\sigma} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi)$$

Normalized asymmetry index


$$N(\theta, \varphi) = \frac{\hat{g}(\theta, \varphi) - \hat{g}(\theta, 2\pi - \varphi)}{\hat{g}(\theta, \varphi) + \hat{g}(\theta, 2\pi - \varphi)} = \frac{\sum_{l=1}^k \sum_{m=-l}^{-1} e^{-l(l+1)\sigma} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi)}{\sum_{l=0}^k \sum_{m=0}^l e^{-l(l+1)\sigma} \langle f, Y_{lm} \rangle Y_{lm}(\theta, \varphi)}$$

Ratio of negative and positive degree expansions

Asymmetry index

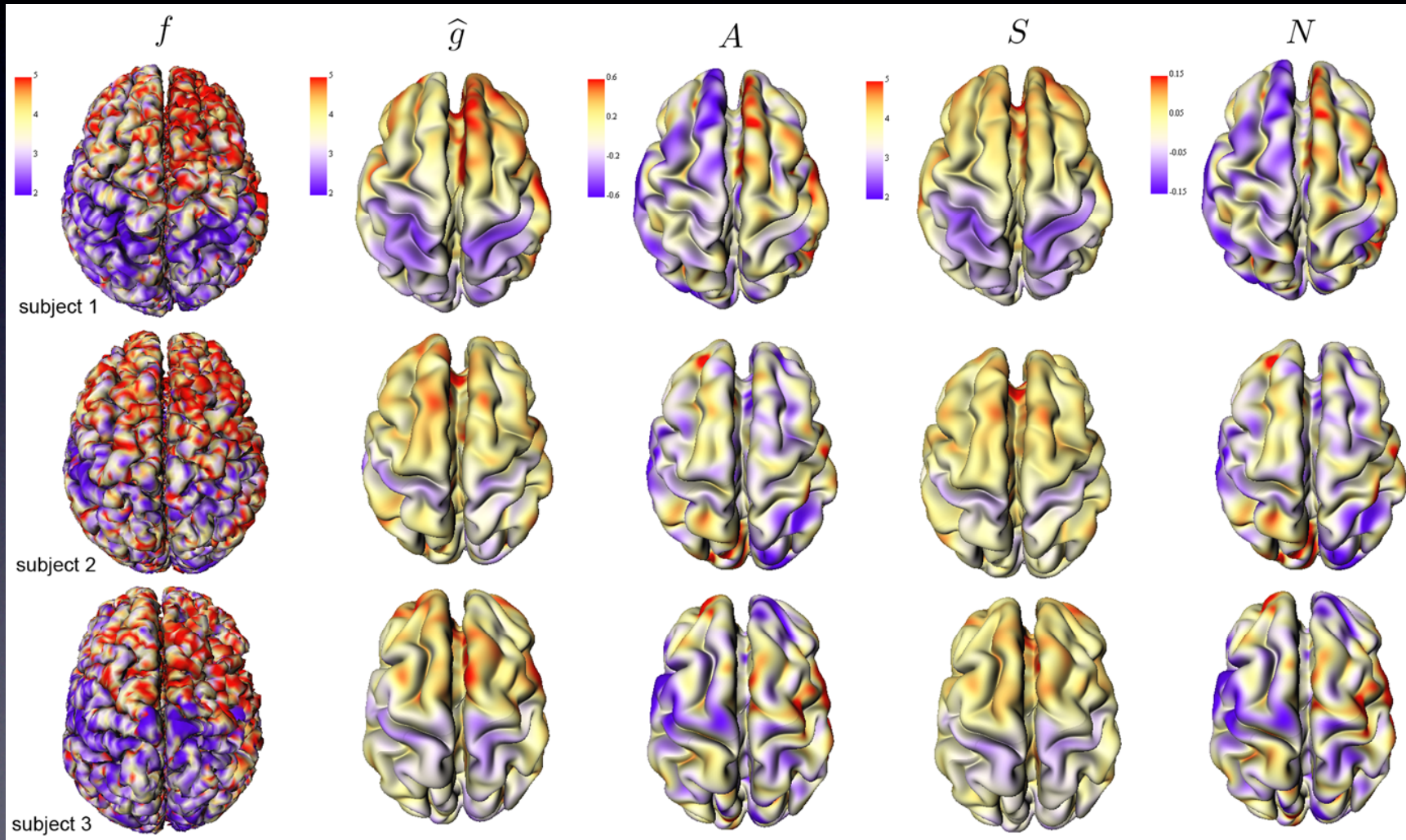
Cortical
thickness

Weighted
SPHARM

Asymmetry
index

Symmetry
index

Normalized
asymmetry
index



Multiple comparisons

$$H_0 : \theta_1(p) = \theta_2(p) \text{ for all } p \in \partial\Omega$$

v.s.

$$H_1 : \theta_1(p) > \theta_2(p) \text{ for some } p \in \partial\Omega.$$

The above null hypothesis is the intersection of collection of hypothesis

$$H_0 = \bigcap_{p \in \partial\Omega} H_0(p)$$

Type I error

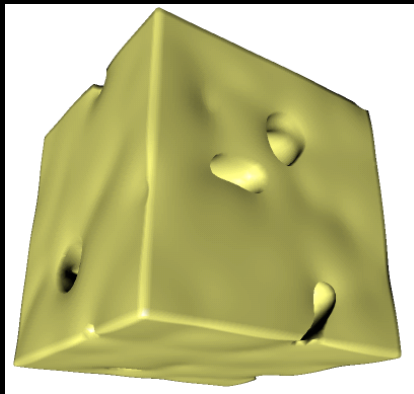
$$\begin{aligned}\alpha &= P(\text{reject at least one } H_0(p) | H_0 \text{ true}) \\ &= P\left(\bigcup_{p \in \partial\Omega} \{T(p) > h\}\right) \\ &= 1 - P\left(\bigcap_{p \in \partial\Omega} T(p) \leq h\right) \\ &= 1 - P(\sup_{p \in \partial\Omega} T(p) \leq h) \\ &= P(\sup_{p \in \partial\Omega} T(p) > h).\end{aligned}$$

t random field

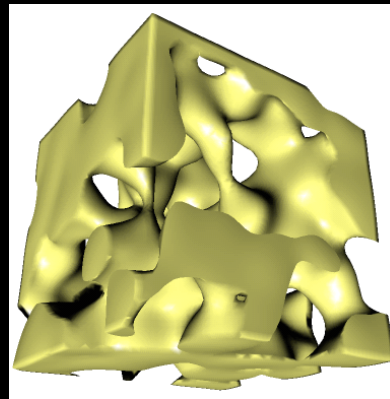
$Z(x)$: Stationary isotropic random field in $x \in \Omega \subset \mathbb{R}^N$

$A_z = \{x : Z(x) > z\}$ excursion set

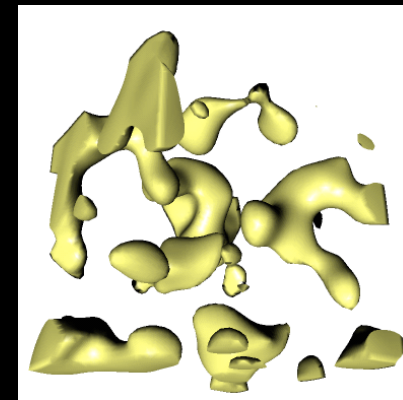
$\chi(A_z)$: Euler characteristic



$z = -10$



$z = 0$



$z = 10$

$$P\left(\max_{x \in \Omega} Z(x) > z\right) \approx \mathbb{E}(\chi(A_z))$$

(Adler, 1984)

T random field on manifolds

$$P\left(\max_{\mathbf{x} \in \partial\Omega_{atlas}} T(\mathbf{x}) \geq y\right) \approx 2\rho_0(y) + \|\partial\Omega_{atlas}\|\rho_2(y)$$

Euler characteristic density

$$\rho_0(y) = \int_y^\infty \frac{\Gamma(\frac{n}{2})}{((n-1)\pi)^{1/2}\Gamma(\frac{n-1}{2})} \left(1 + \frac{y^2}{n-1}\right)^{-n/2} dy,$$

$$\rho_2(y) = \frac{1}{\text{FWHM}^2} \frac{4 \ln 2}{(2\pi)^{3/2}} \frac{\Gamma(\frac{n}{2})}{(\frac{n-1}{2})^{1/2}\Gamma(\frac{n-1}{2})} y \left(1 + \frac{y^2}{n-1}\right)^{-(n-2)/2}$$



Worsley (1995, NeuroImage)

FWHM of smoothing kernel or residual field

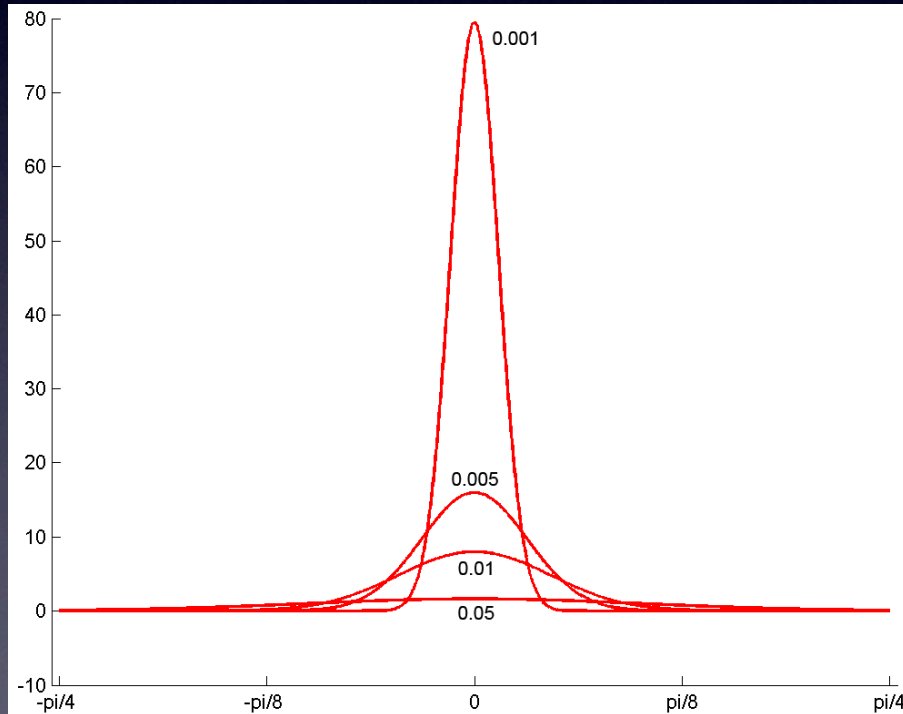
WFS is related to heat kernel smoothing

WFS

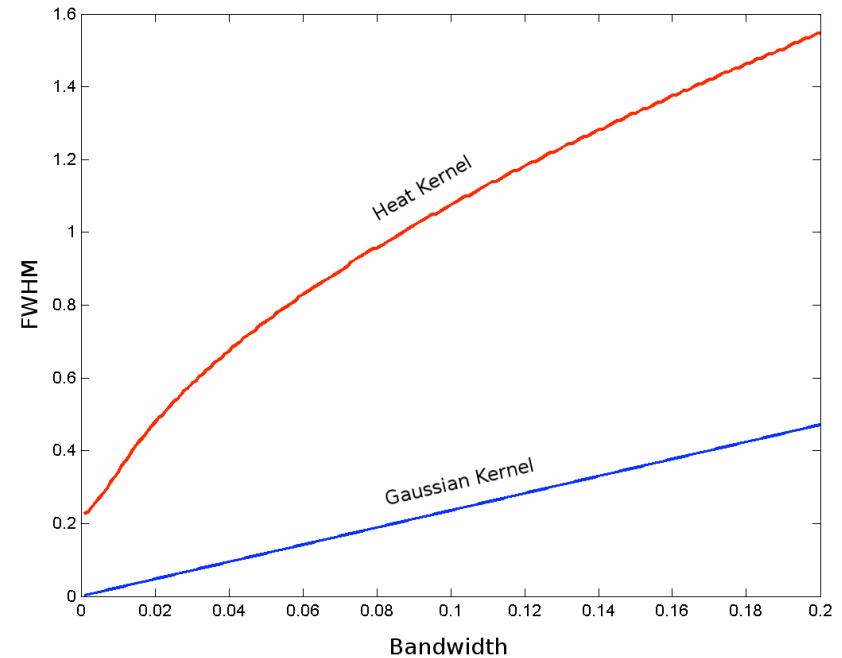
$$g(p, t) = \sum_{j=0}^{\infty} e^{-\lambda_j t} \langle f, \psi_j \rangle \psi_j(p)$$

Heat kernel smoothing

$$= \int_{\mathcal{N}} K_t(p, q) f(q) d\mu(q)$$



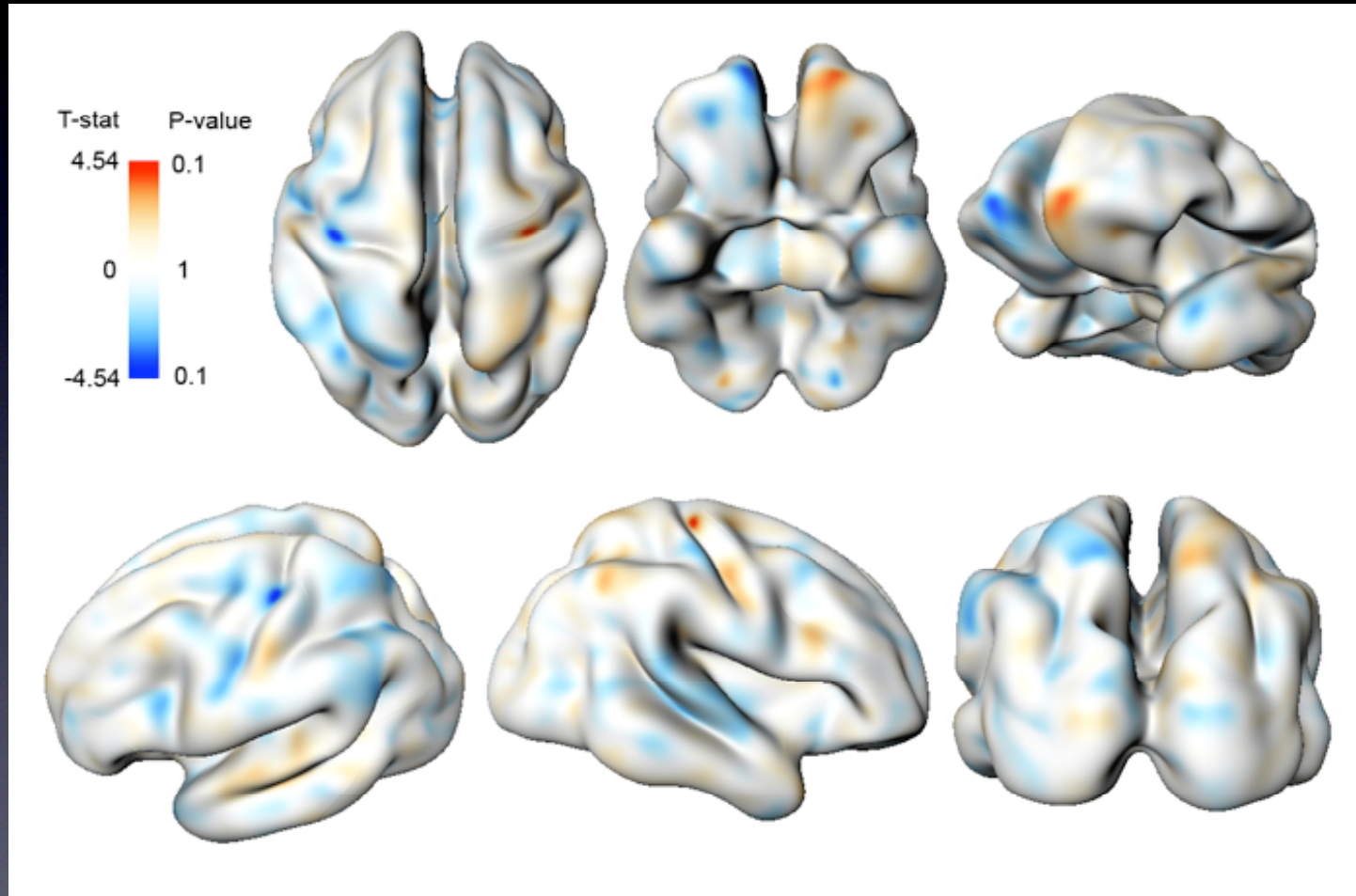
Shape of heat kernel



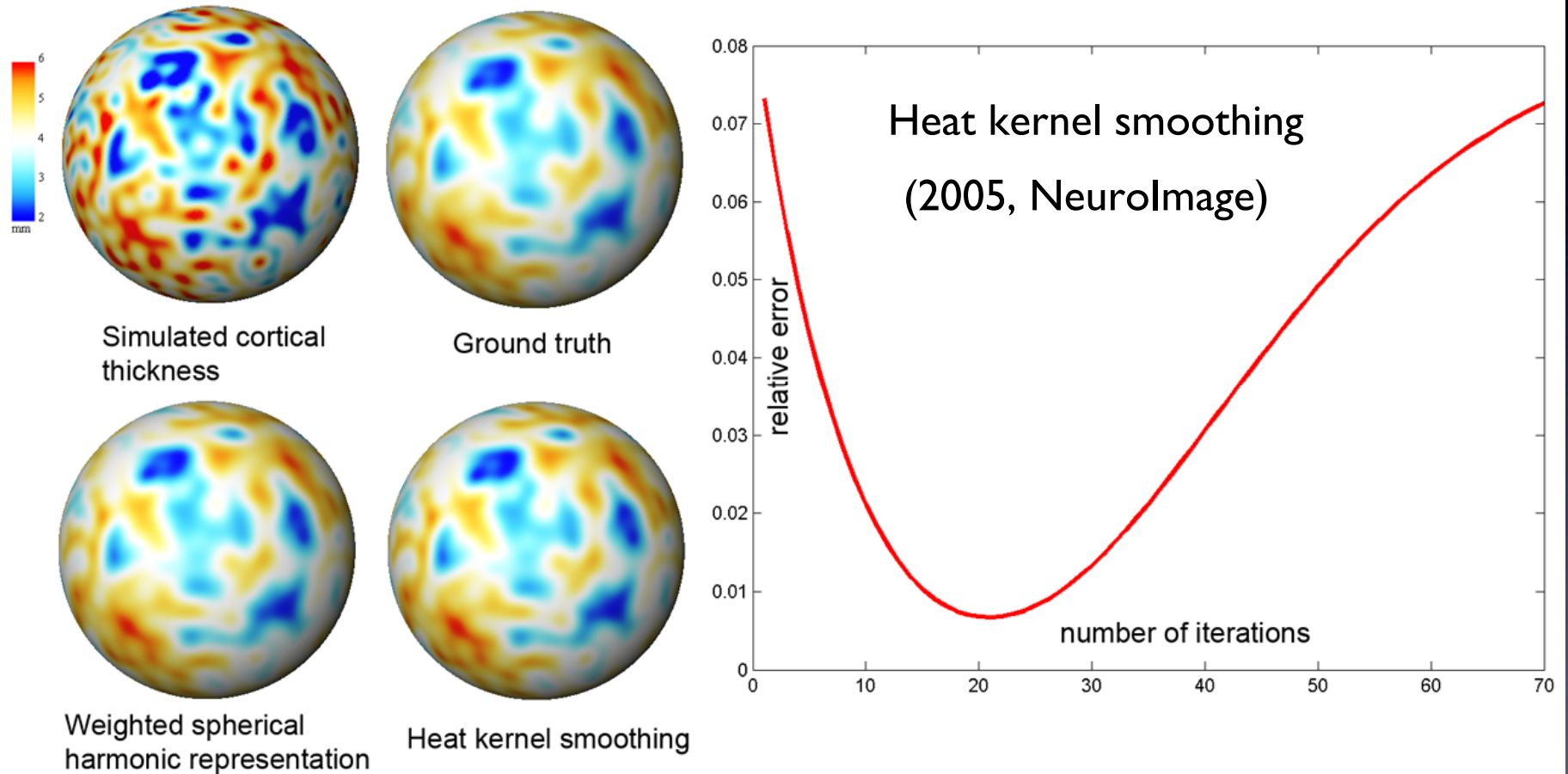
Numerical computation

Statistical parametric map

multiple comparison correction via
the random field theory



Validation of WFS against analytical ground truth



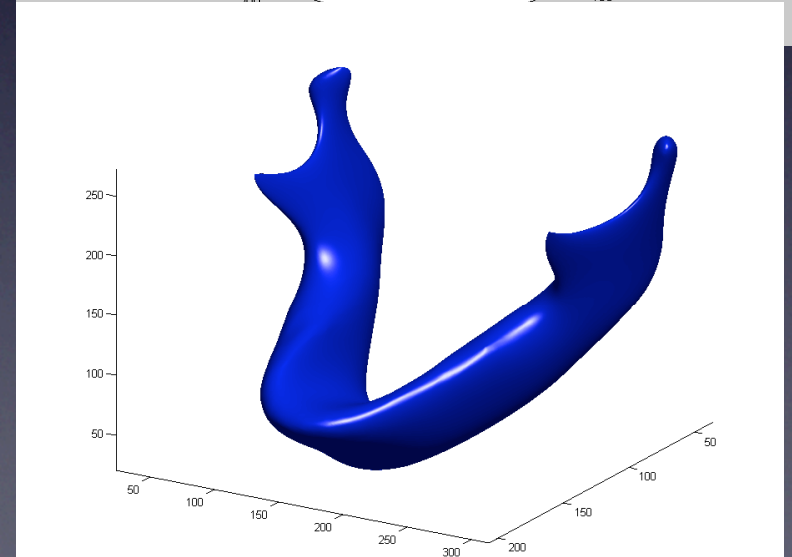
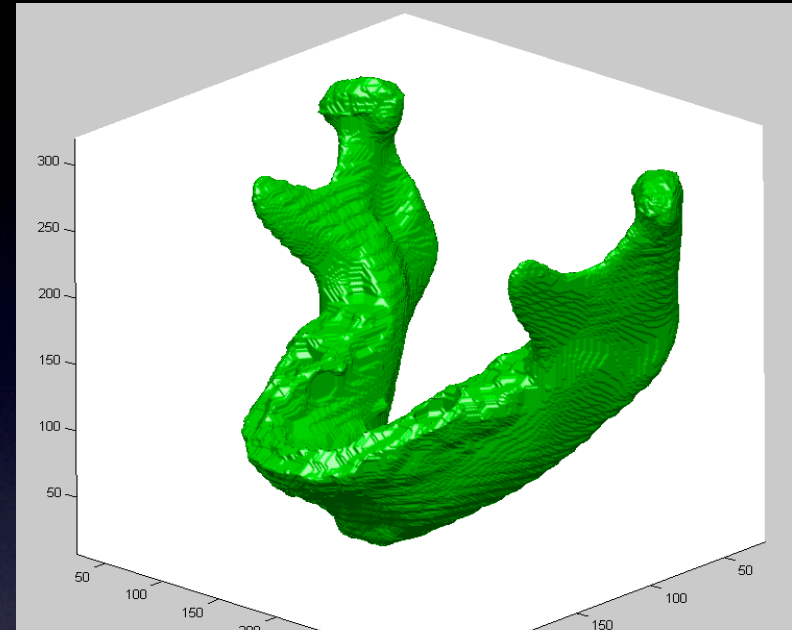
Next project? Mandible surface modeling



Histogram
thresholding



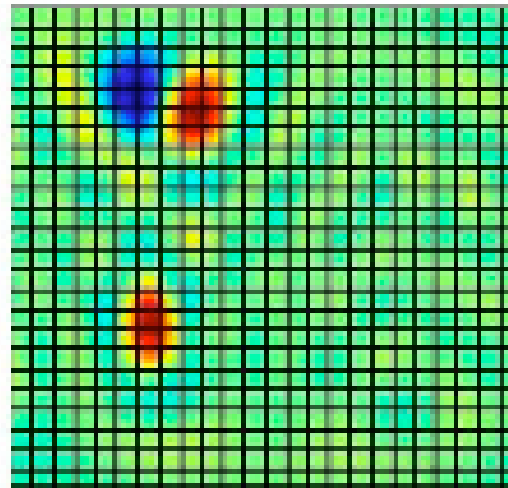
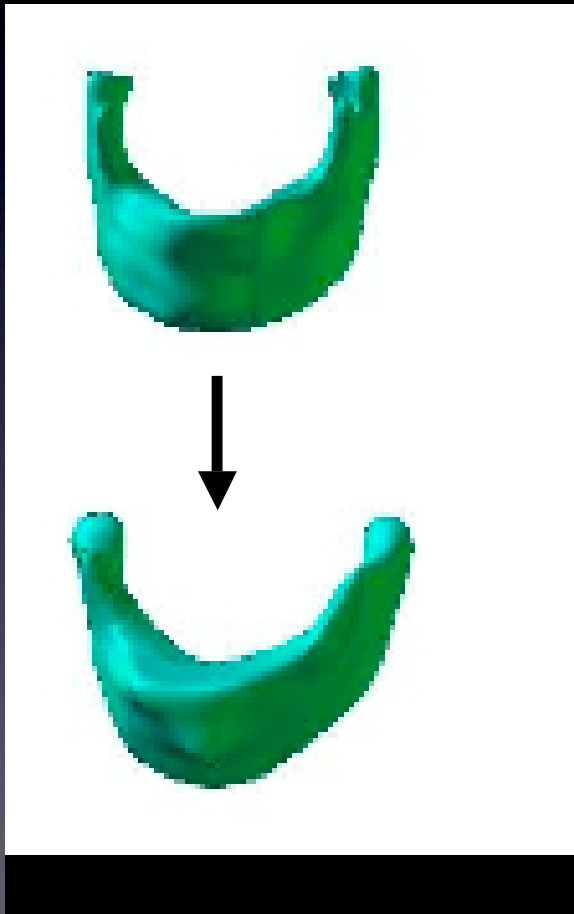
Hole
patching



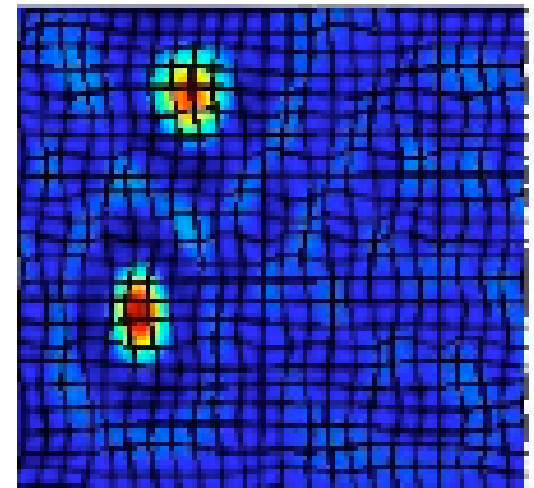
Automatic hole patching is
necessary to construct surface
topologically equivalent to sphere.

Approximately 20,000 triangle
elements

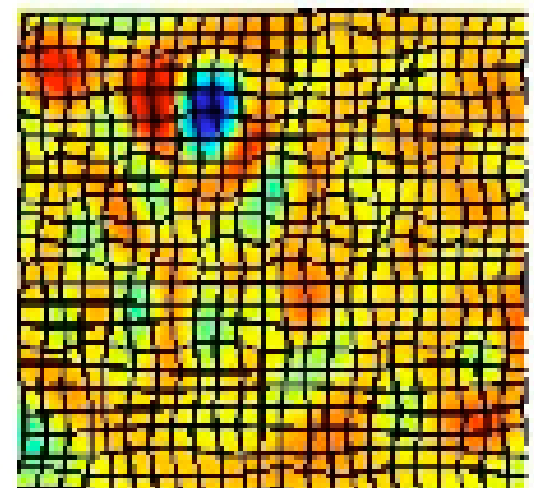
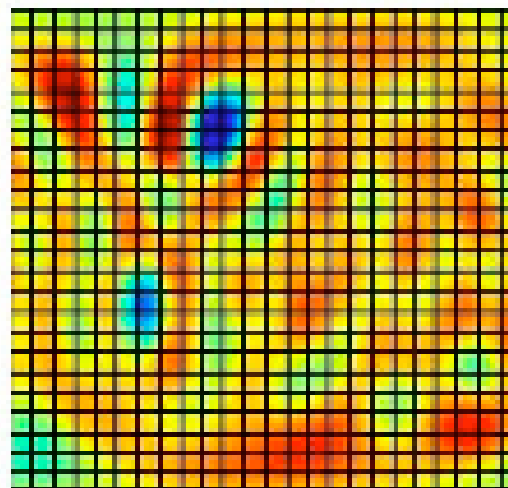
Nonlinear surface registration via curvature matching



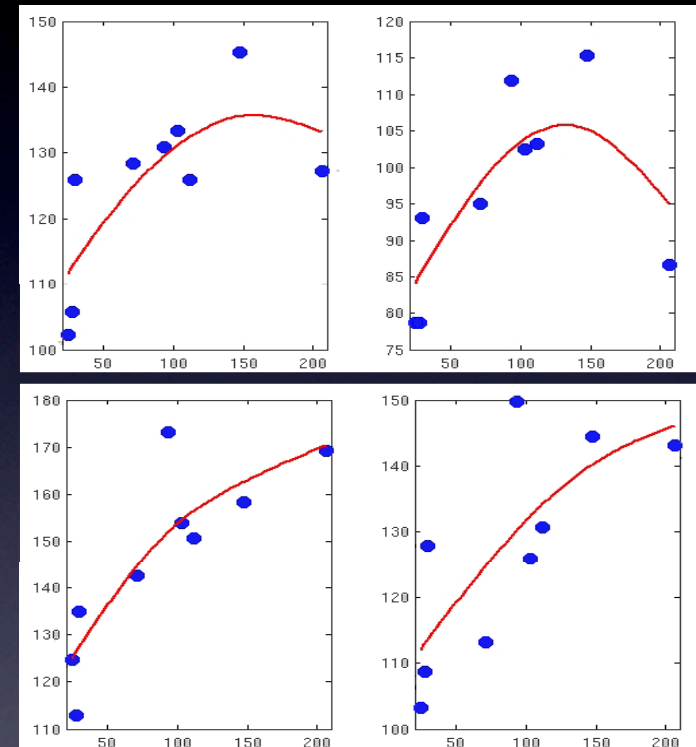
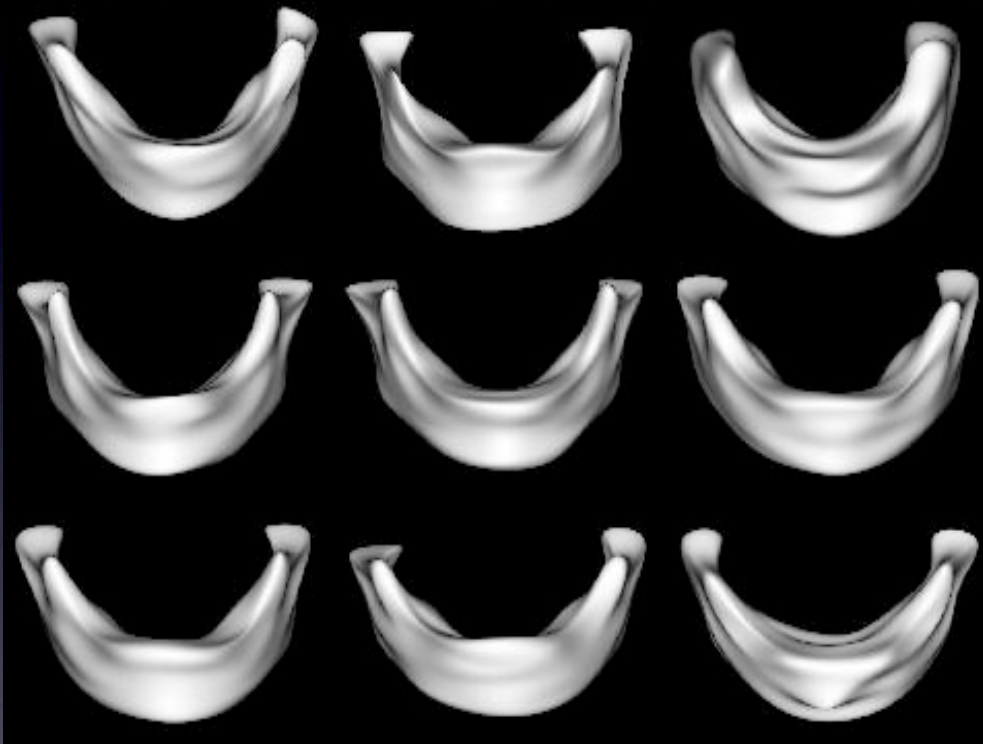
Curvatures



Nonlinear warping grid

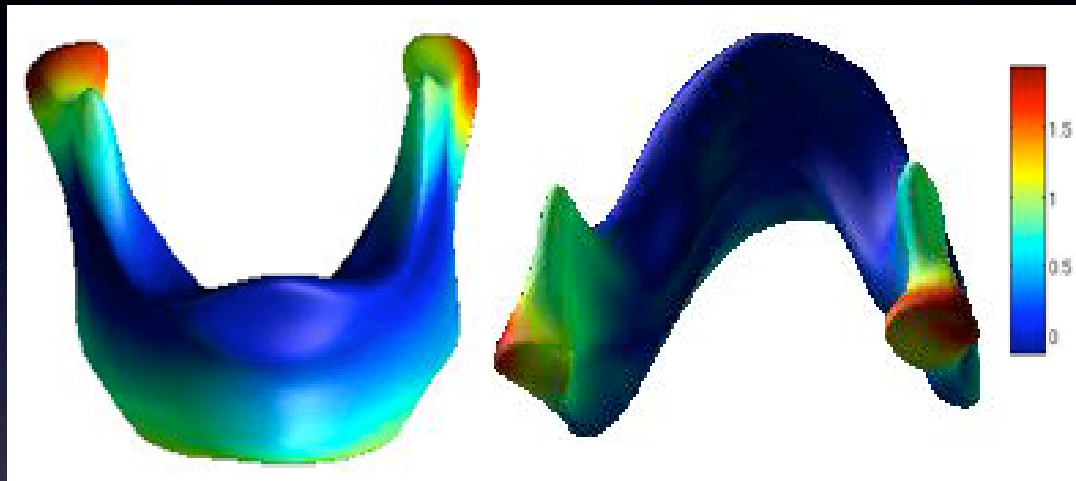


Mandible surface modeling



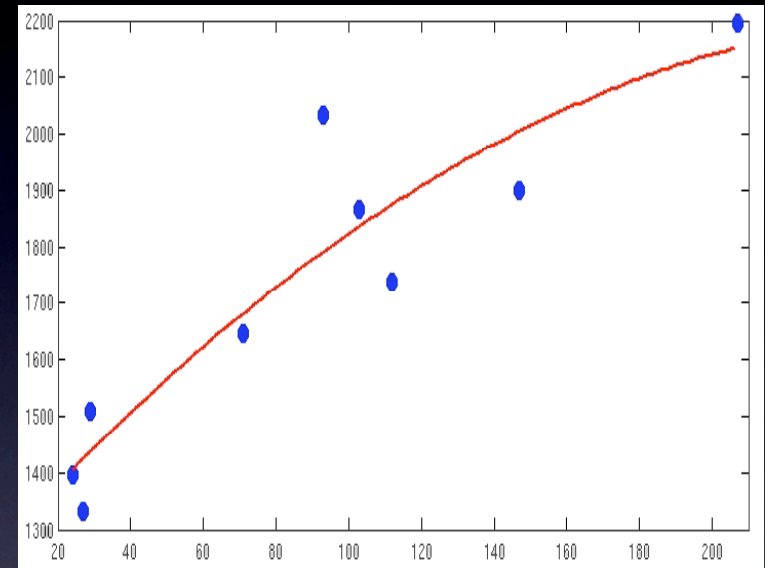
Quadratic fit of 9 male subjects over time in one particular point on the mandible surface **Plan: do this on 300 subjects**

Locally varying growth rate modeling



Growth rate (obtained directly from the regression model) projected on average mandible surface

Plan: incorporate gender and other variables into analysis.



Total surface area growth