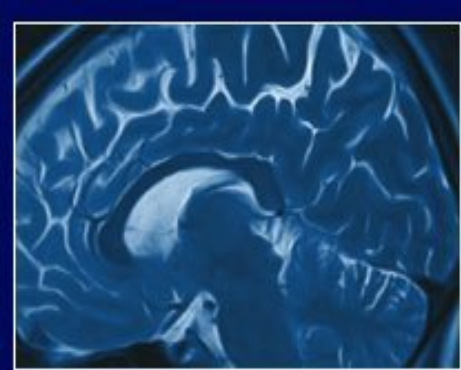




*The Waisman Laboratory
for Brain Imaging and Behavior*

University of Wisconsin
**SCHOOL OF MEDICINE
AND PUBLIC HEALTH**

Random Walks in the Permutation Group

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Abstract

The permutation test is an often used test procedure in brain imaging. Unfortunately, generating every possible permutation for large-scale brain image datasets such as HCP and ADNI with thousands images is not practical. Many previous attempts at speeding up the permutation test rely on various approximation strategies such as estimating the tail distribution with known parametric distributions. In this study, we show how to rapidly accelerate the permutation test without any type of approximate strategies by exploiting the underlying topological structure of the permutation group via random walks. The method is applied to large number of magnetic resonance images in two applications: (1) localizing the male and female differences and (2) localizing the regions of high genetic heritability in the sulcal and gyral graph patterns of the human cortical brain. The talk is based on [arXiv:1812.06696](https://arxiv.org/abs/1812.06696).

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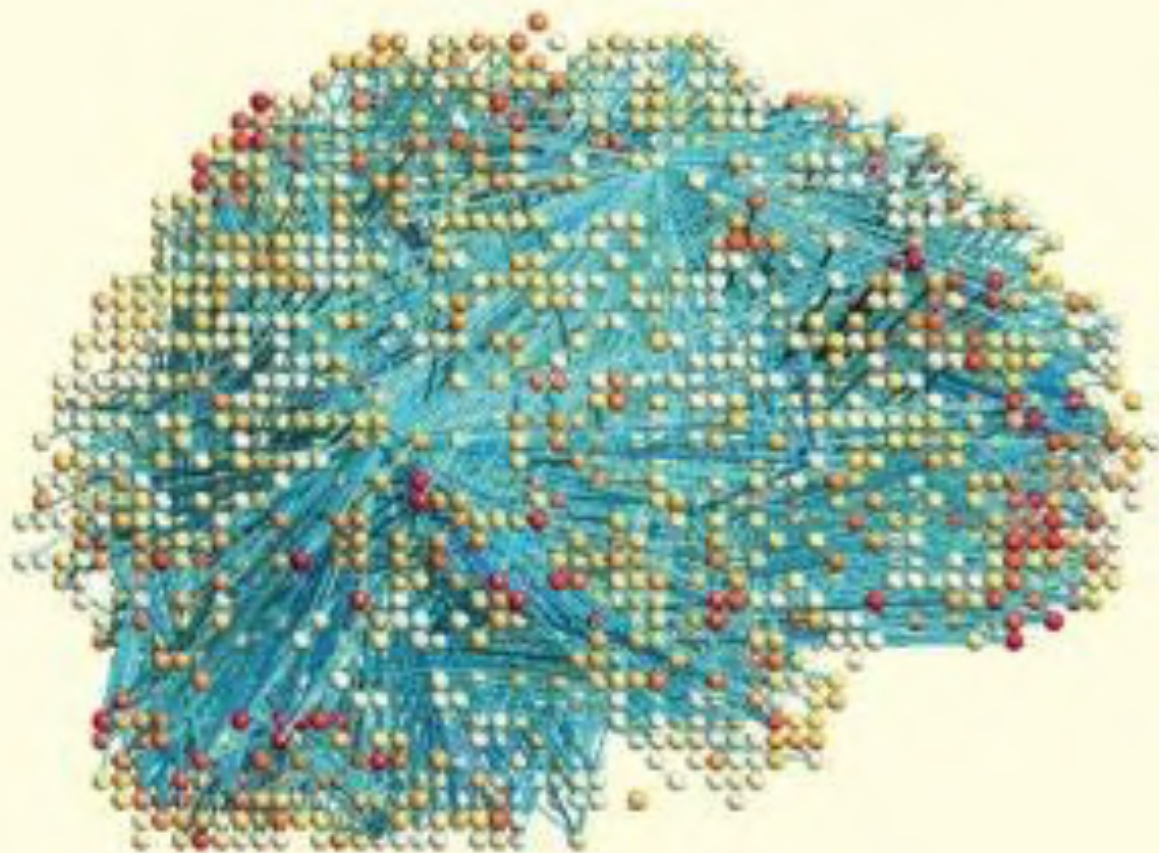
Full day course
**Topological and Object
Oriented Data Analysis** in
International Biometric
Conference (IBC2020)
COEX Seoul, Korea

Sunday July 5, 2020

OODA: Steve Marron (UNC)
TDA: Yuan Wang (USC)
Computation: Moo K. Chung
(UW-Madison)



BRAIN NETWORK ANALYSIS



Moo K. CHUNG

Date: August 31, 2019

Publisher: Cambridge
University Press

One chapter deals with
TDA based network
analysis

TDA MATLAB Codes, Data & Talk Slides

www.stat.wisc.edu/~mchung/TDA

What is permutation test?

$$\mathbf{x} = (x_1, x_2, \cdots, x_m)$$

$$\mathbf{y} = (y_1, y_2, \cdots, y_n)$$

$$(\mathbf{x}, \mathbf{y}) = (x_1, \cdots, x_m, y_1, \cdots, y_n)$$

$$\pi(\mathbf{x}, \mathbf{y}) \in \mathbb{S}_{m+n}$$

Permutation group of order $m+n$

$$p\text{-value} = \frac{1}{(m+n)!} \sum_{\pi \in \mathbb{S}_{m+n}} \mathcal{I}\left(f(\mathbf{x}, \mathbf{y}) \geq f(\pi(\mathbf{x}, \mathbf{y}))\right)$$

History of permutation test

Fisher 1935, The Design of Experiment

$$\binom{8}{4} = 70$$

Thompson et al. 2001, Nature Neuroscience

$$\binom{40}{20} = 1.34 \cdot 10^{11}$$

*Supercomputer for
1 million permutations*

Nichols et al. 2002, Human Brain Mapping

4279 citations

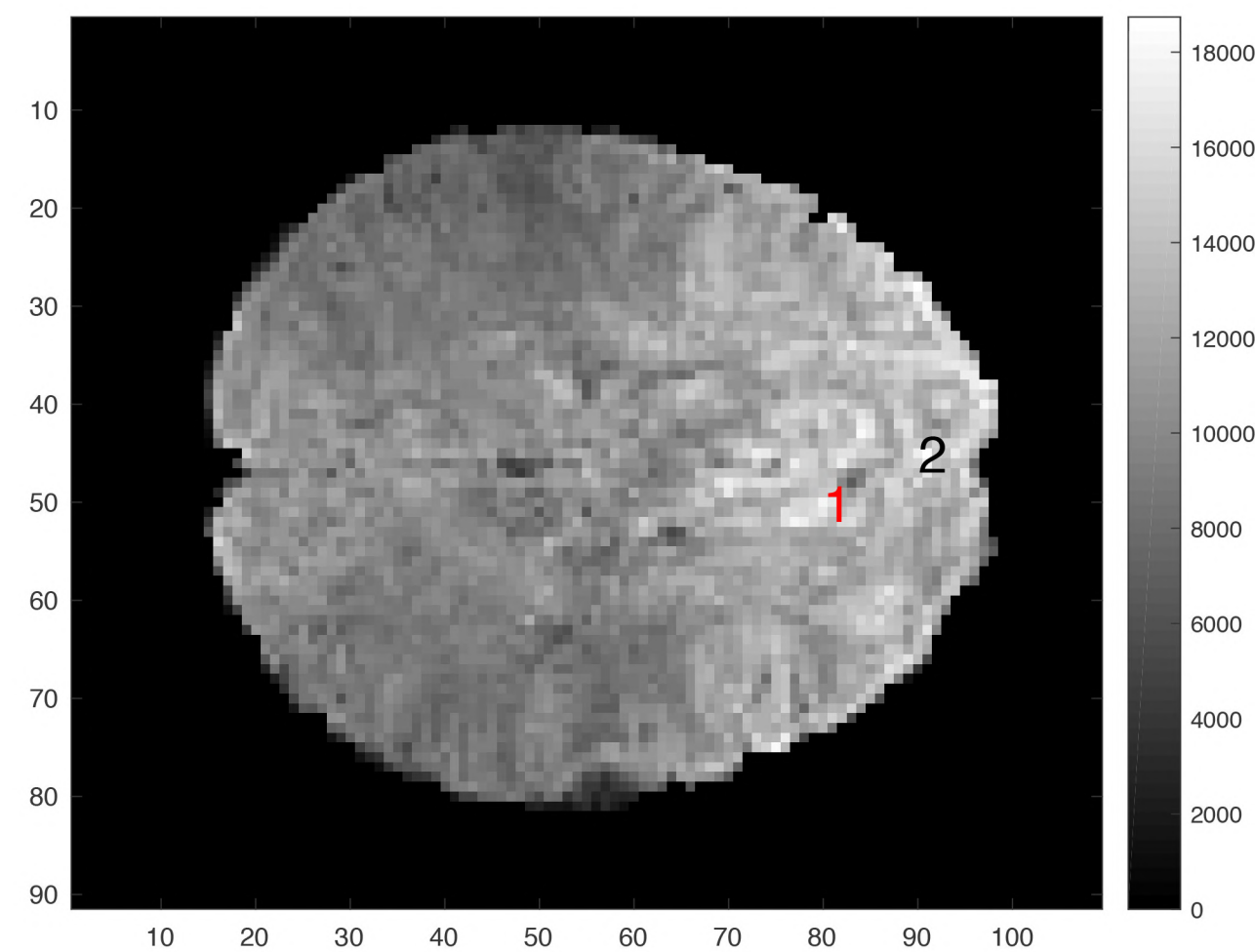
$$\binom{6}{3} = 20$$

Previous method: Exact topological inference

Chung et al. 2017 Information Processing in Medical Imaging

Chung et al. 2018 MICCAI

Chung et al. 2019 Network Neuroscience



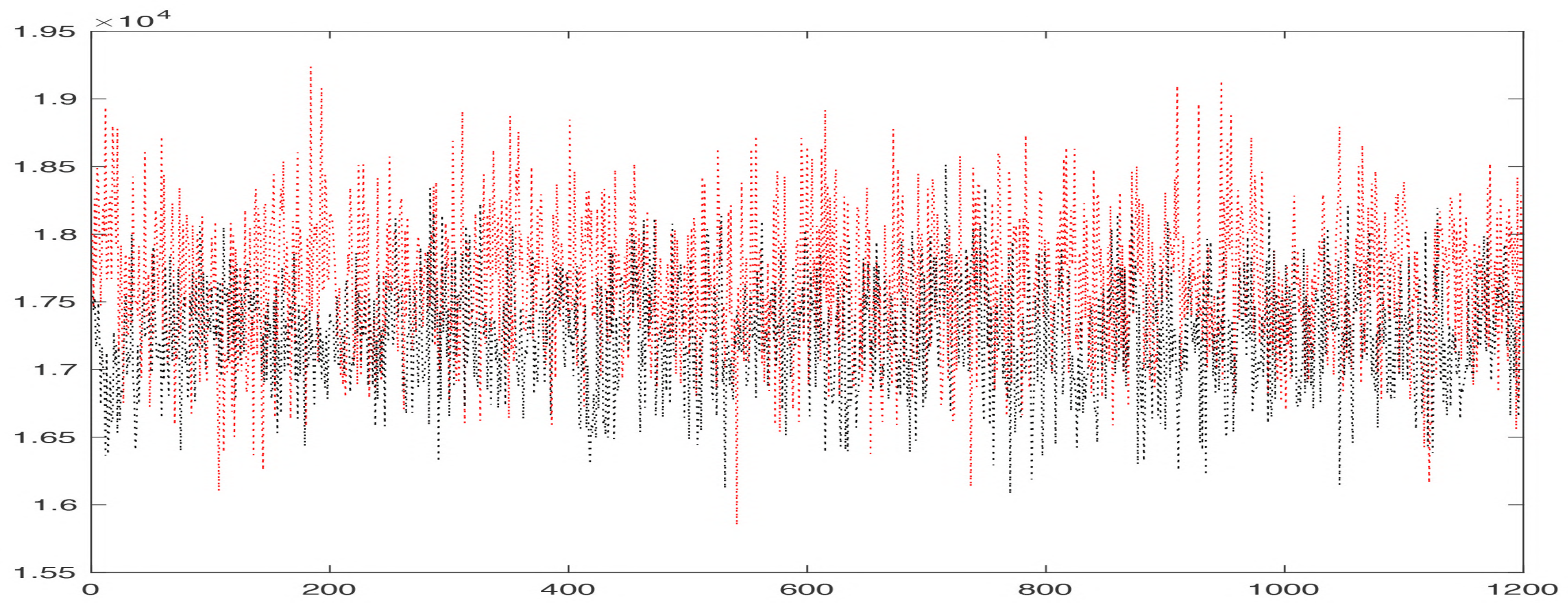
Resting-state functional-MRI at
1200 time measured over 14min
33 seconds.

416 subjects

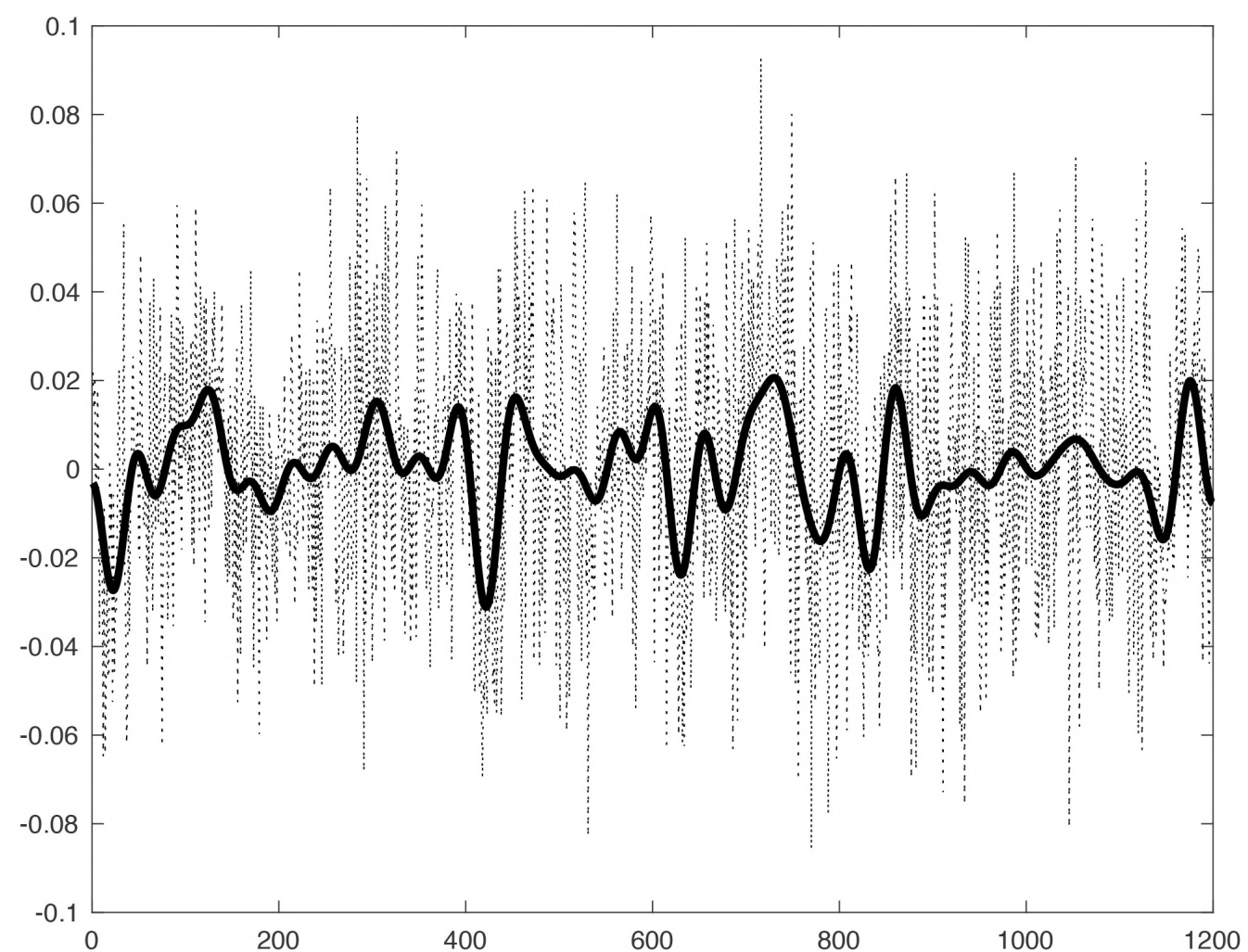
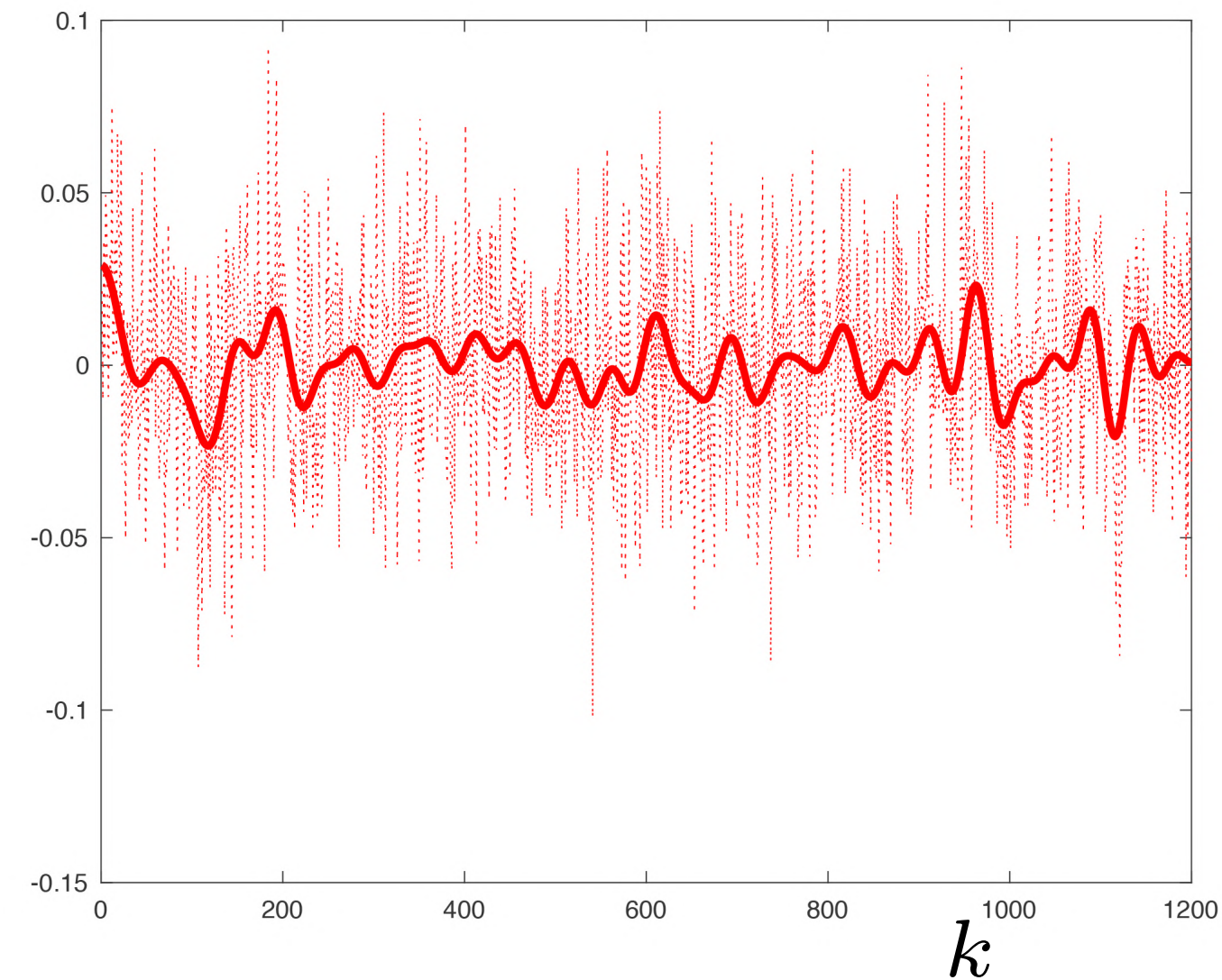
= 131 Monozygotic (MZ) twins

77 Dizygotic (DZ) twins

= **2GB x 416 = 832GB data**



Cosine Series Representation

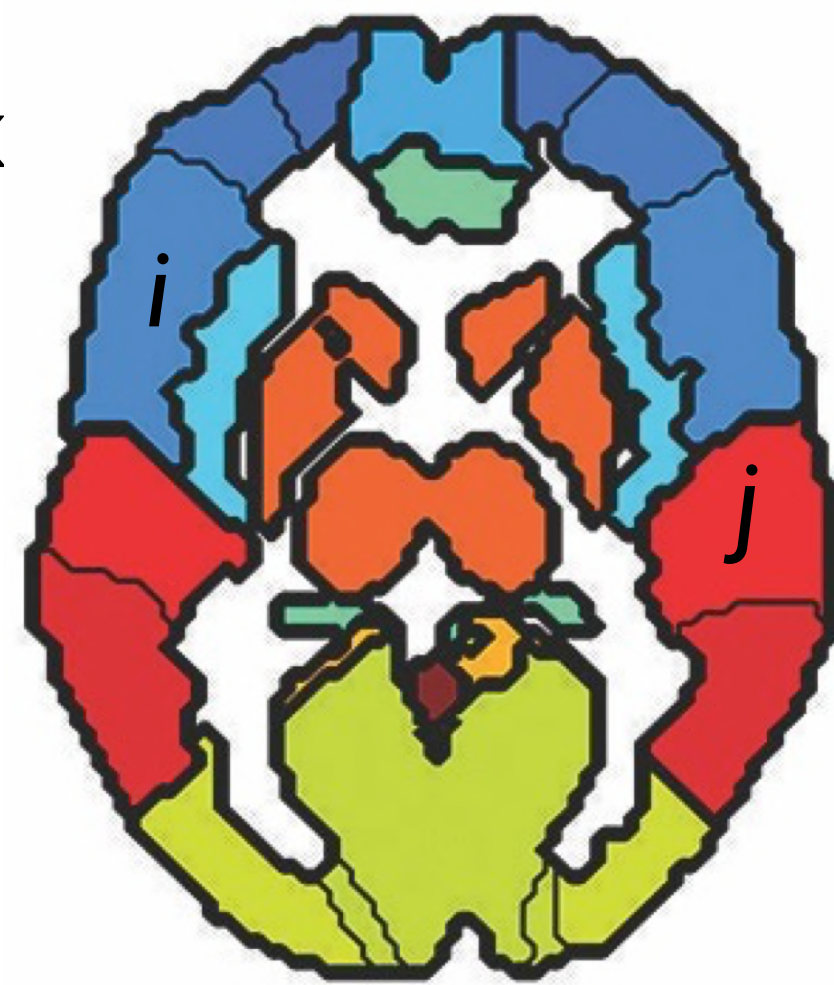


$$\zeta_i(t) = \sum_{l=0}^k d_{li} \psi_l(t), \quad t \in [0, 1]$$

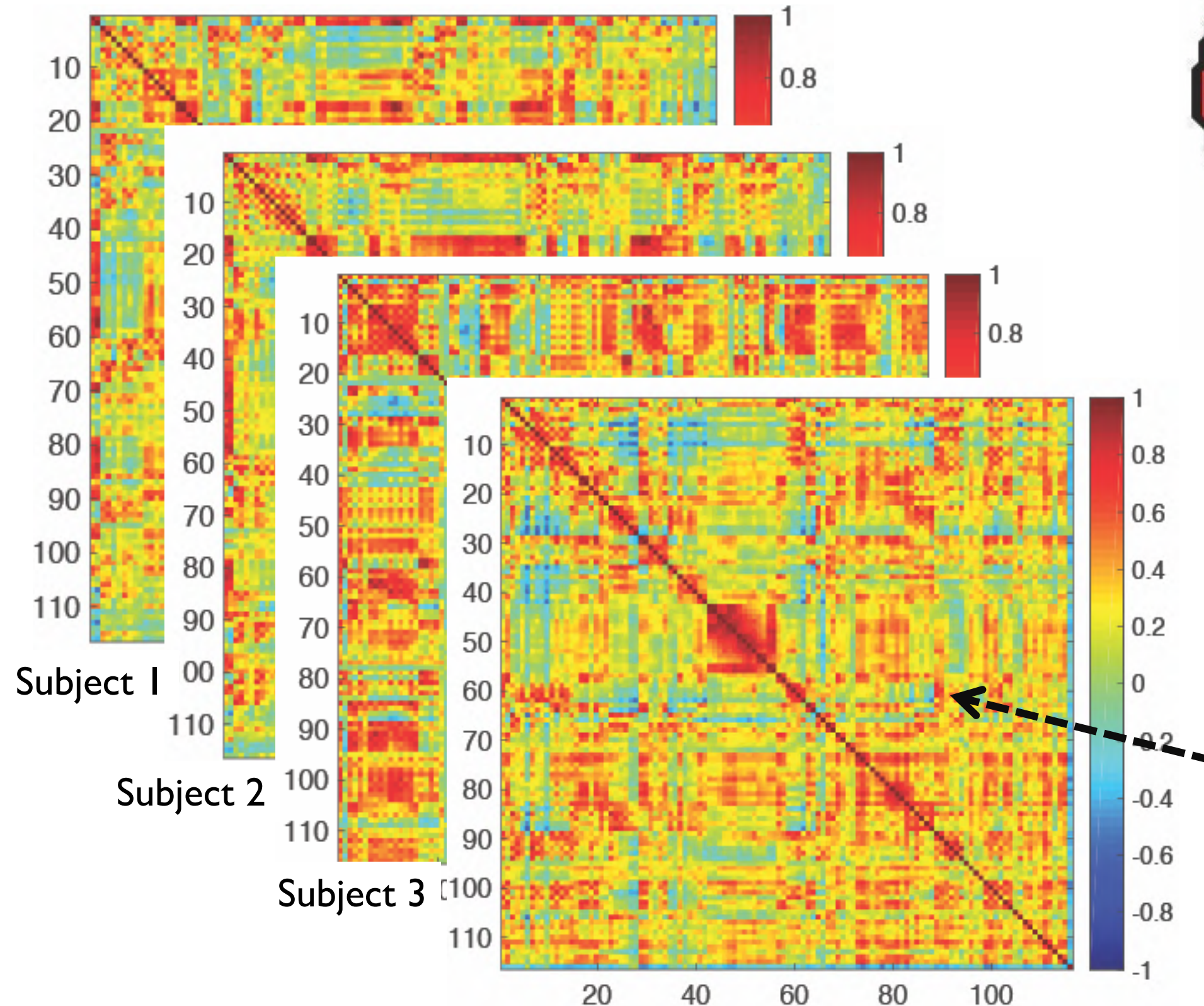
$$\psi_0(t) = 1, \psi_l(t) = \sqrt{2} \cos(l\pi t)$$

120 features $\longrightarrow \mathbf{d}_i = (d_{0i}, d_{1i}, \dots, d_{ki})$

Subject level brain connectivity matrix

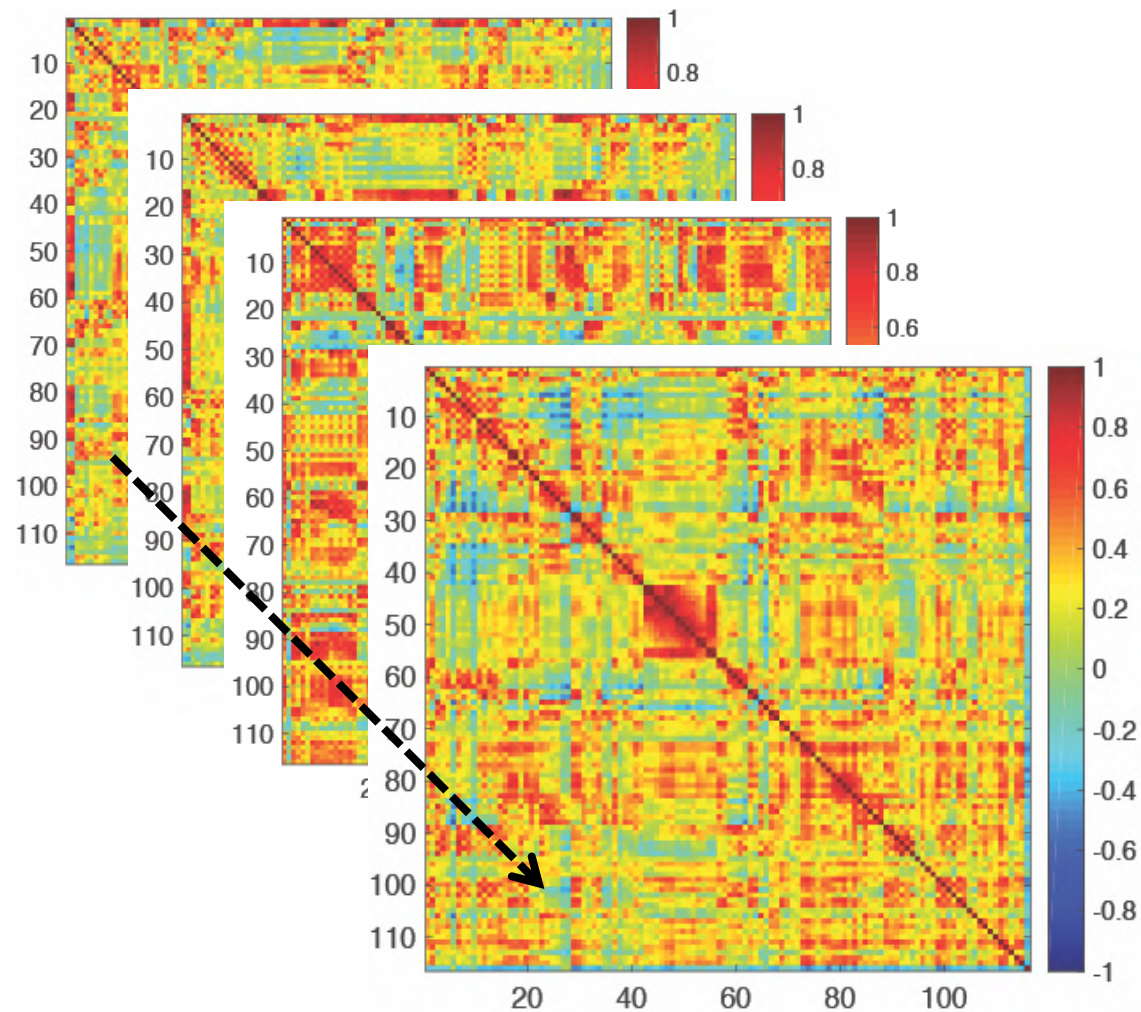


$$c_{ij} = \text{corr}(\mathbf{d}_i, \mathbf{d}_j)$$

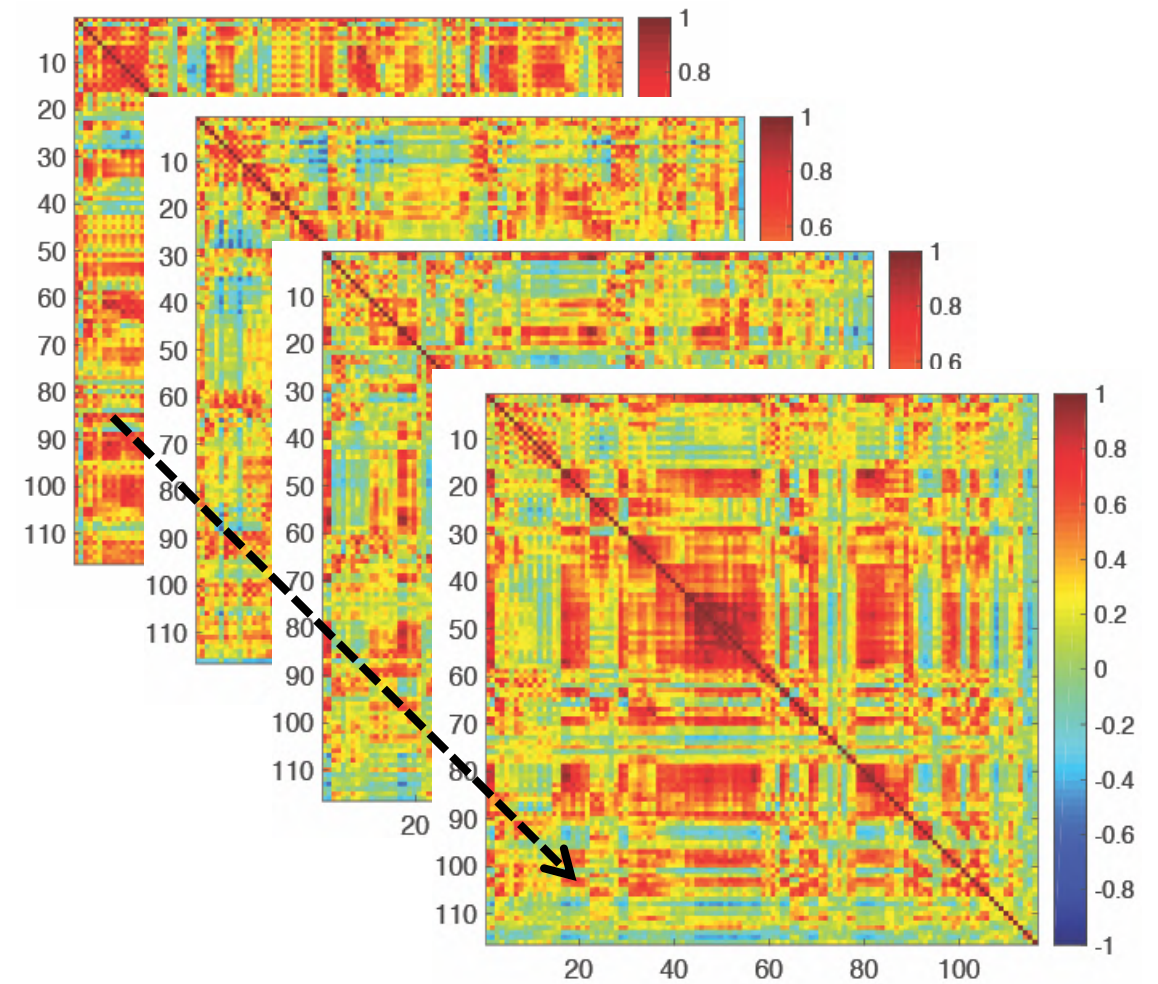


Correlation of
Fourier
coefficients

Correlation (group) of correlation (subject)



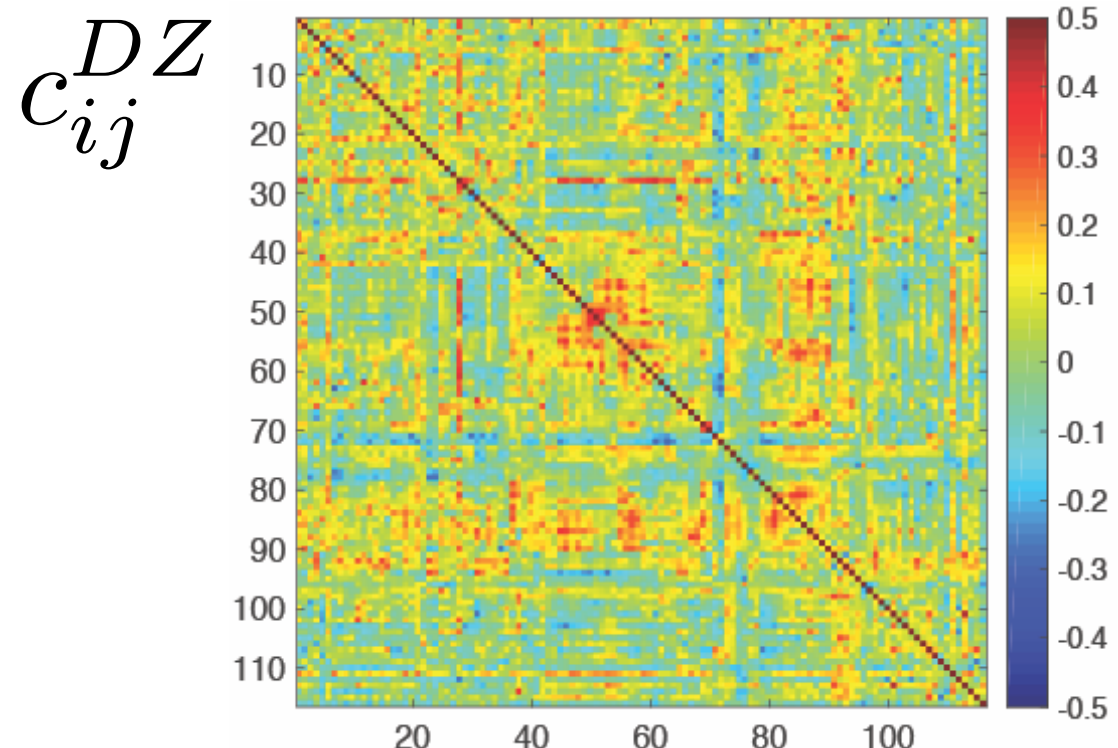
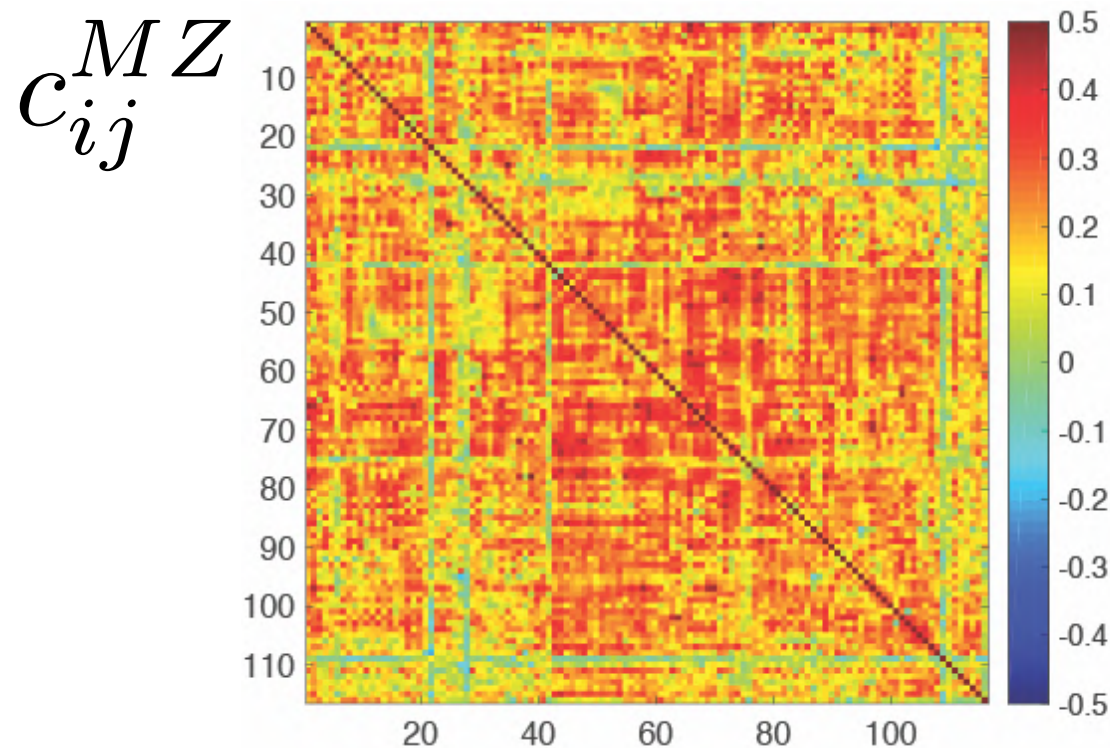
$$\mathbf{c}_{ij}^1 = (c_{ij}^{11}, \dots, c_{ij}^{1m})$$



$$\mathbf{c}_{ij}^2 = (c_{ij}^{21}, \dots, c_{ij}^{2m})$$

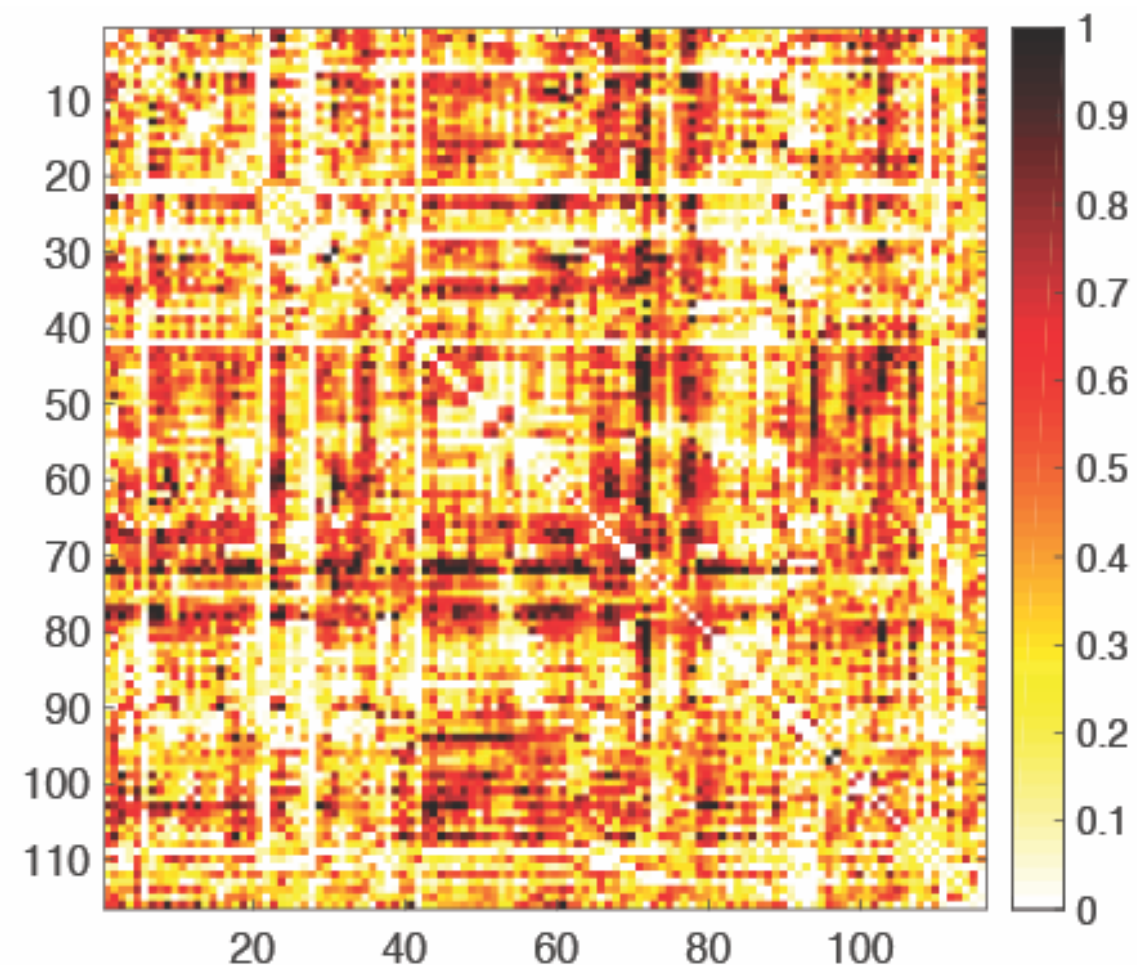
$$c_{ij}^{MZ} = \text{corr}(\mathbf{c}_{ij}^1, \mathbf{c}_{ij}^2)$$

MZ- and DZ-twin correlation difference

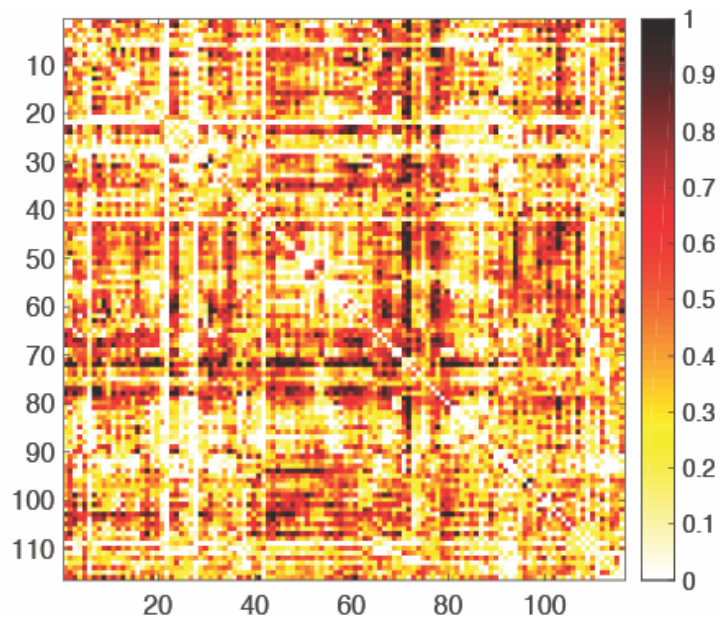


$$h_{ij} = 2(c_{ij}^{MZ} - c_{ij}^{DZ})$$

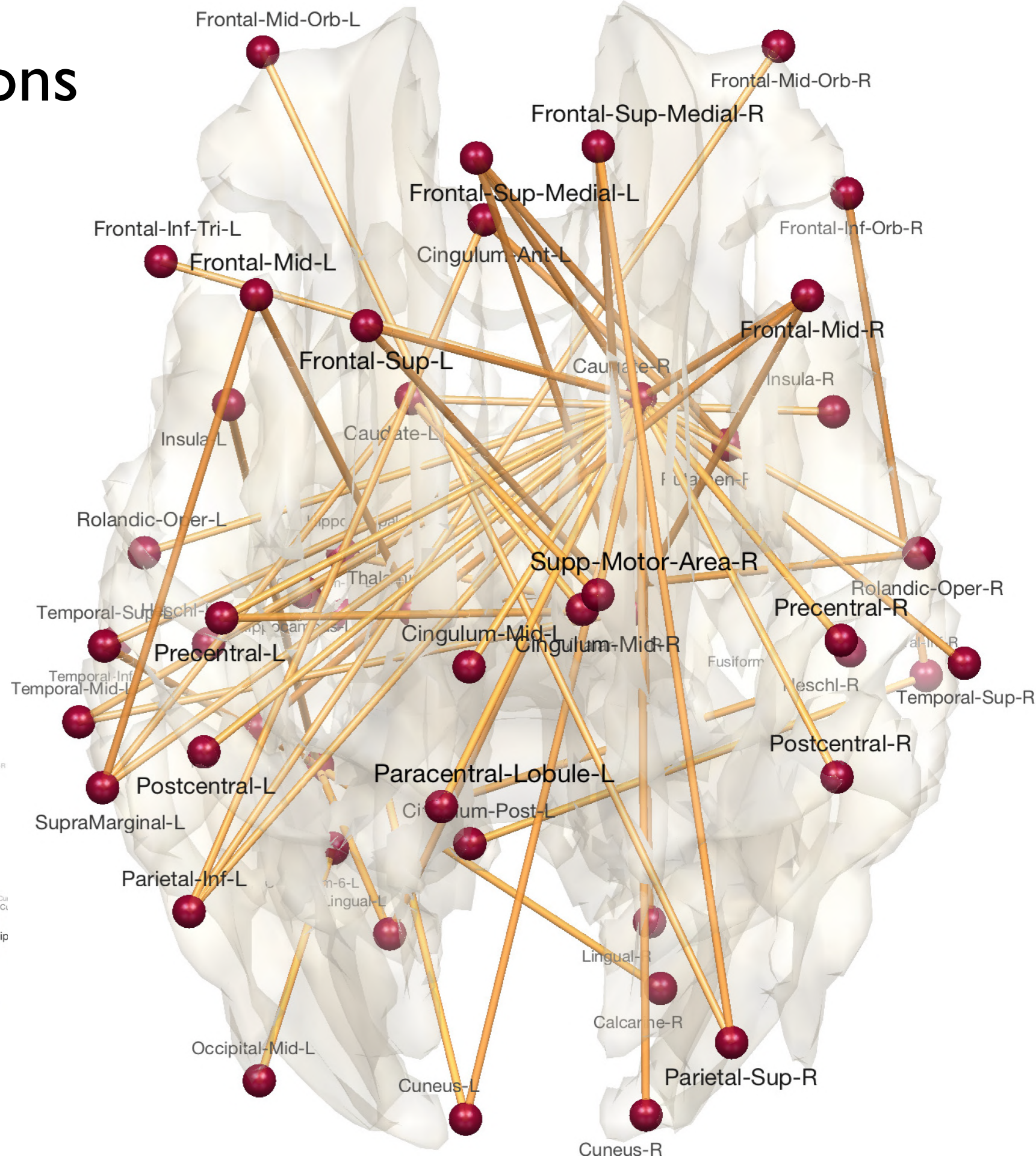
Heritability index = amount
of genetic contribution
(MZ- and DZ-twins)



Heritable brain regions

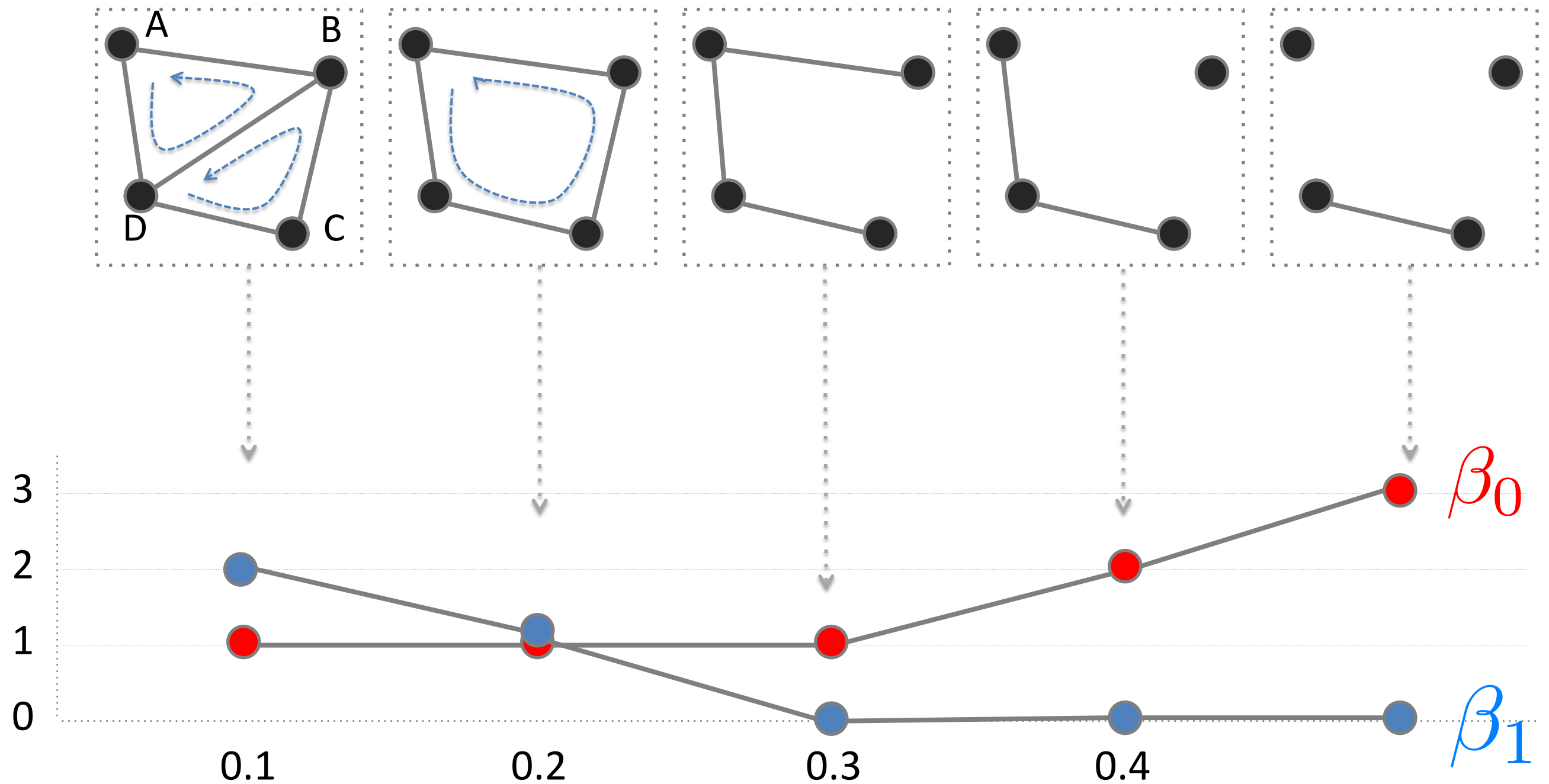


$$h_{ij} \geq 1$$



Statistical significance?

Betti-plots



filtration value = correlation

Theorem 1. β_0 and β_1 are monotone over graph filtration.

Monotonicity of β_0 : Deletion of edge increases the number of connected components by at most 1. β_0 increases by 0 or 1.

Monotonicity of β_1 :

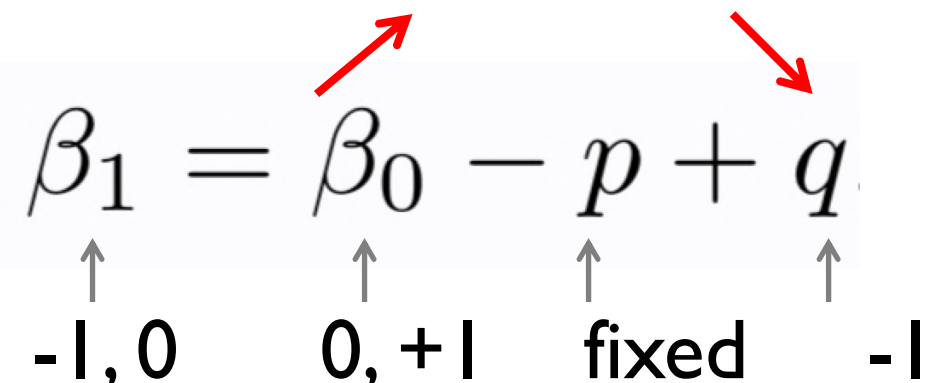
Euler characteristic:

$$\chi = \beta_0 - \beta_1 = p - q$$

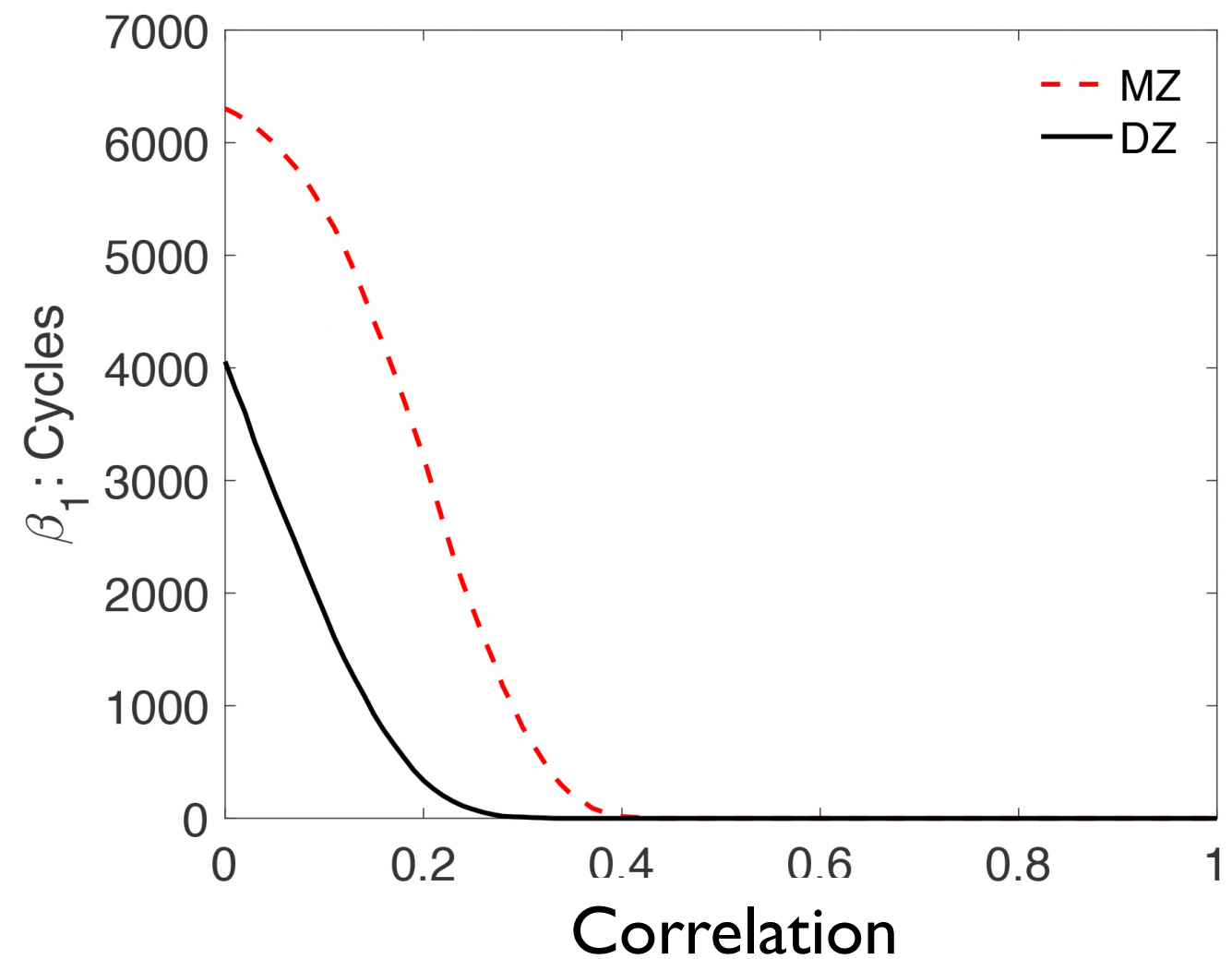
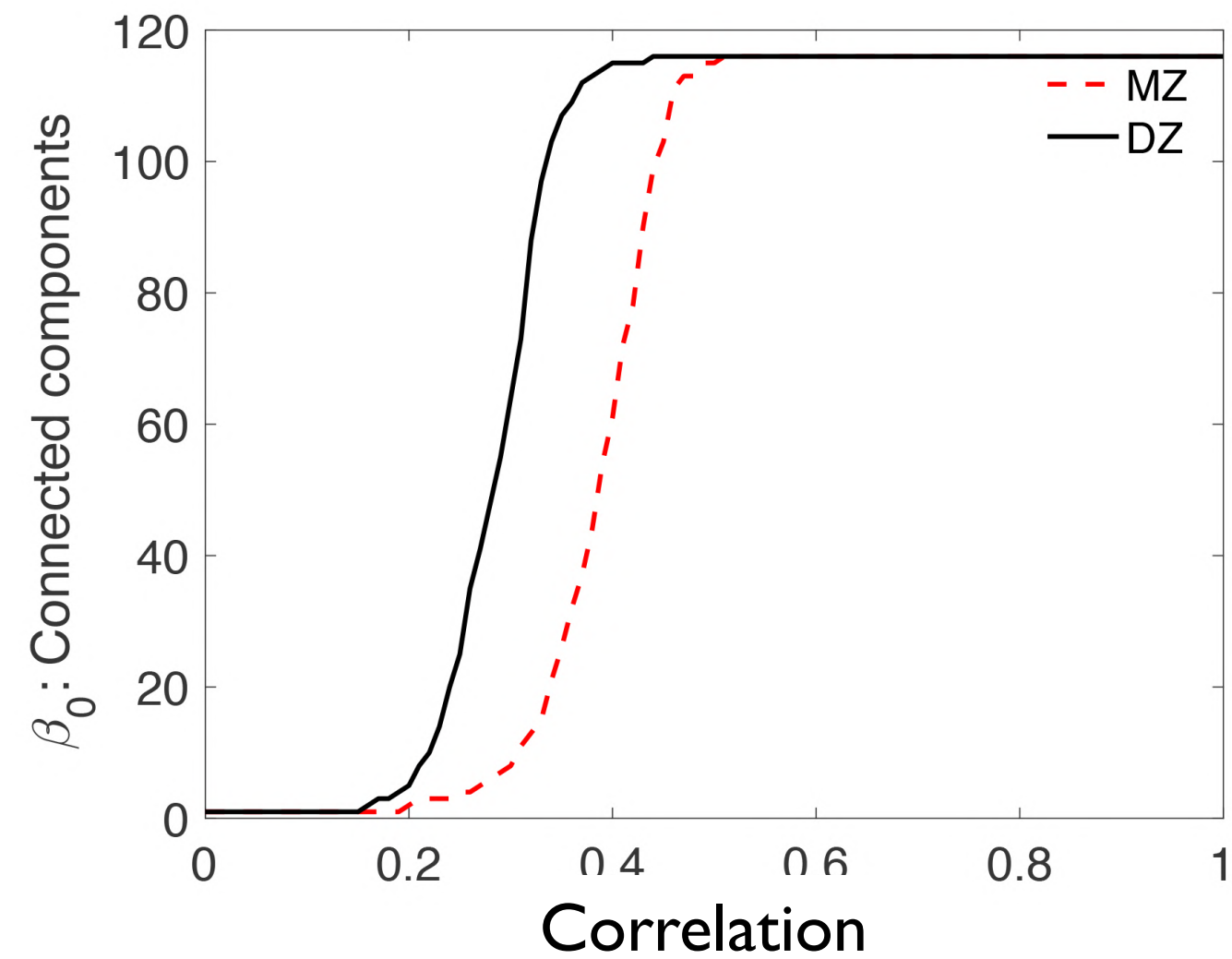
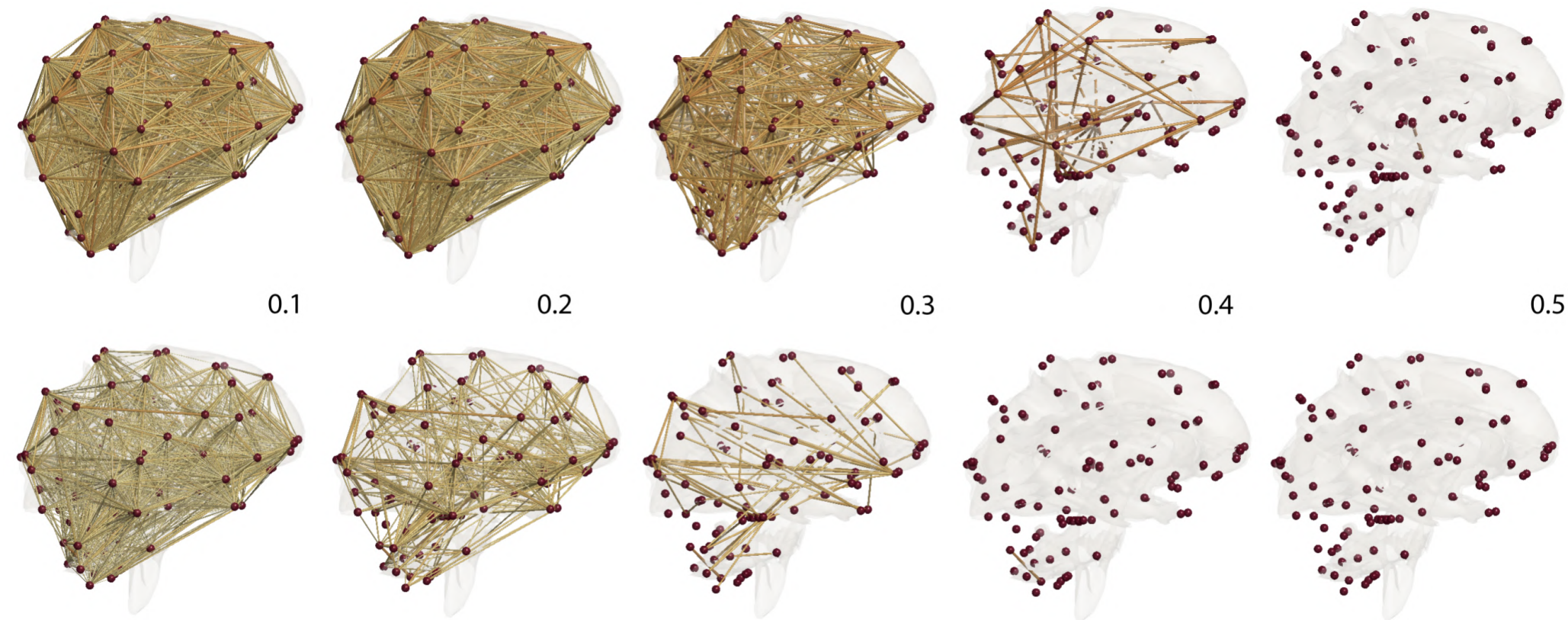
↑ nodes ↑ edges

$$\beta_1 = \beta_0 - p + q$$

↑ ↑ ↑ ↑
-1, 0 0, +1 fixed -1



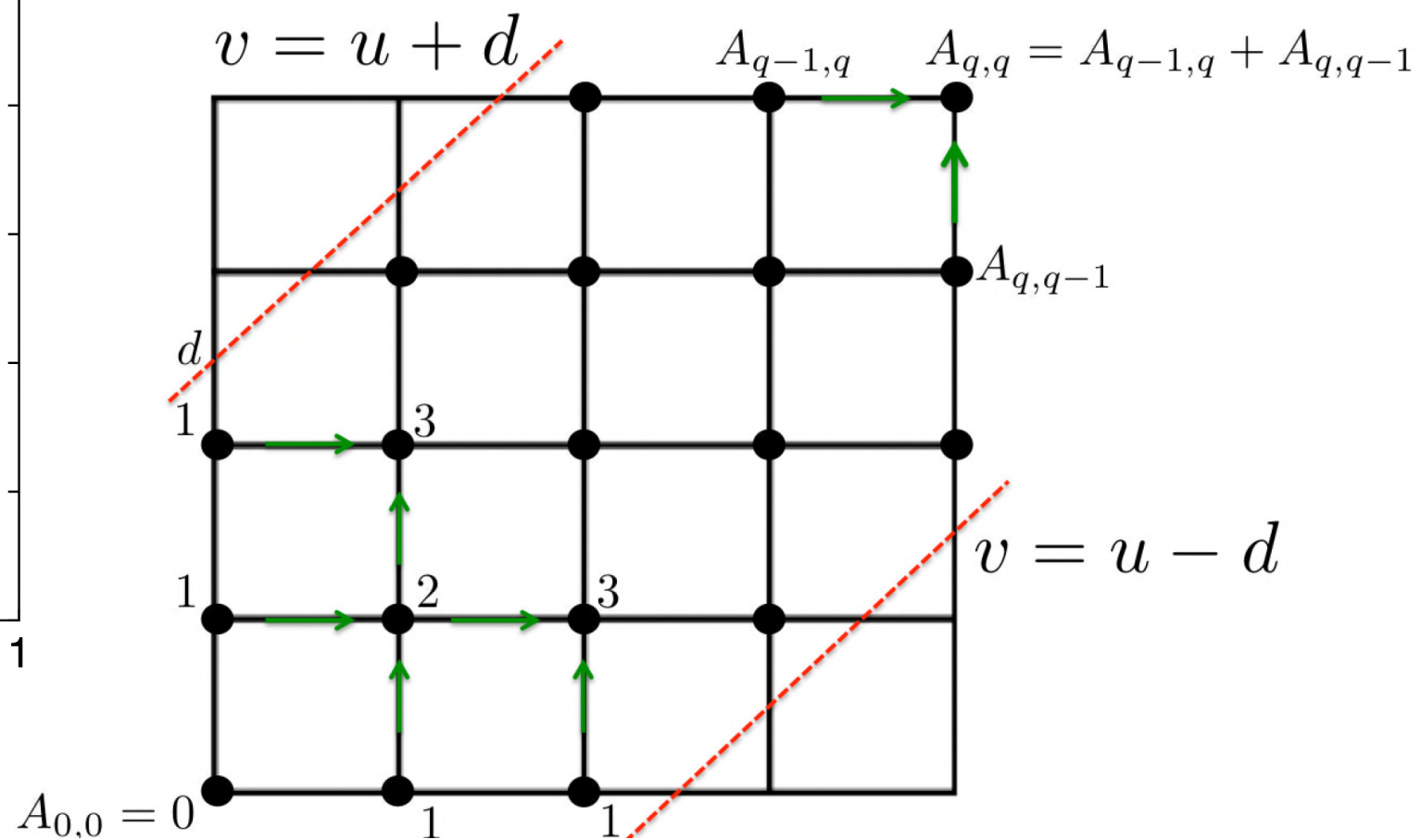
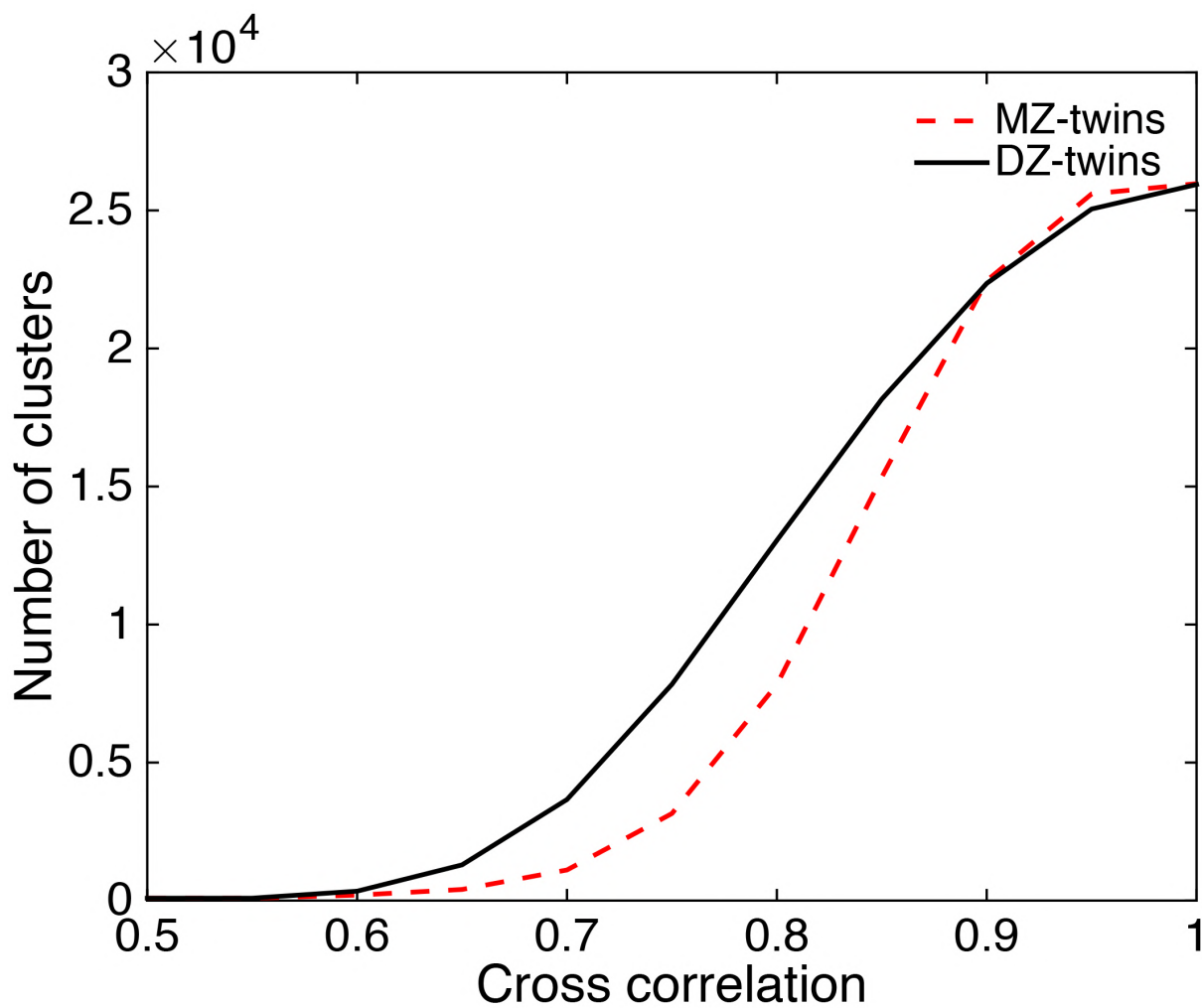
Betti-plots on graph filtration



Exact topological inference

Theorem 2.
$$D_q = \sup_{1 \leq j \leq q} |\beta_i(G_{\lambda_j}^1) - \beta_i(G_{\lambda_j}^2)|$$

$$P(D_q \geq d) = 1 - \frac{A_{q,q}}{\binom{2q}{q}}$$



$$\lim_{q \rightarrow \infty} P\left(D_q / \sqrt{2q} \geq d\right) = 2 \sum_{i=1}^{\infty} (-1)^{i-1} e^{-2i^2 d^2}$$

New approach Rapid acceleration via slow random walks

Chung et al. 2019 [arXiv:1812.06696](https://arxiv.org/abs/1812.06696)

Random walk on the permutation group

$$\mathbf{X} = (x_1, x_2, \dots, x_{i-1}, \boxed{x_i}, x_{i+1}, \dots, x_m)$$

transpose i -th and j -th data

$$\mathbf{y} = (y_1, y_2, \dots, y_{j-1}, \boxed{y_j}, y_{j+1}, \dots, y_n)$$



$$\pi_{ij}(\mathbf{X}) = (x_1, x_2, \dots, x_{i-1}, \boxed{y_j}, x_{i+1}, \dots, x_m)$$

$$\pi_{ij}(\mathbf{y}) = (y_1, y_2, \dots, y_{j-1}, \boxed{x_i}, y_{j+1}, \dots, y_n)$$

Online computation of statistic over transposition

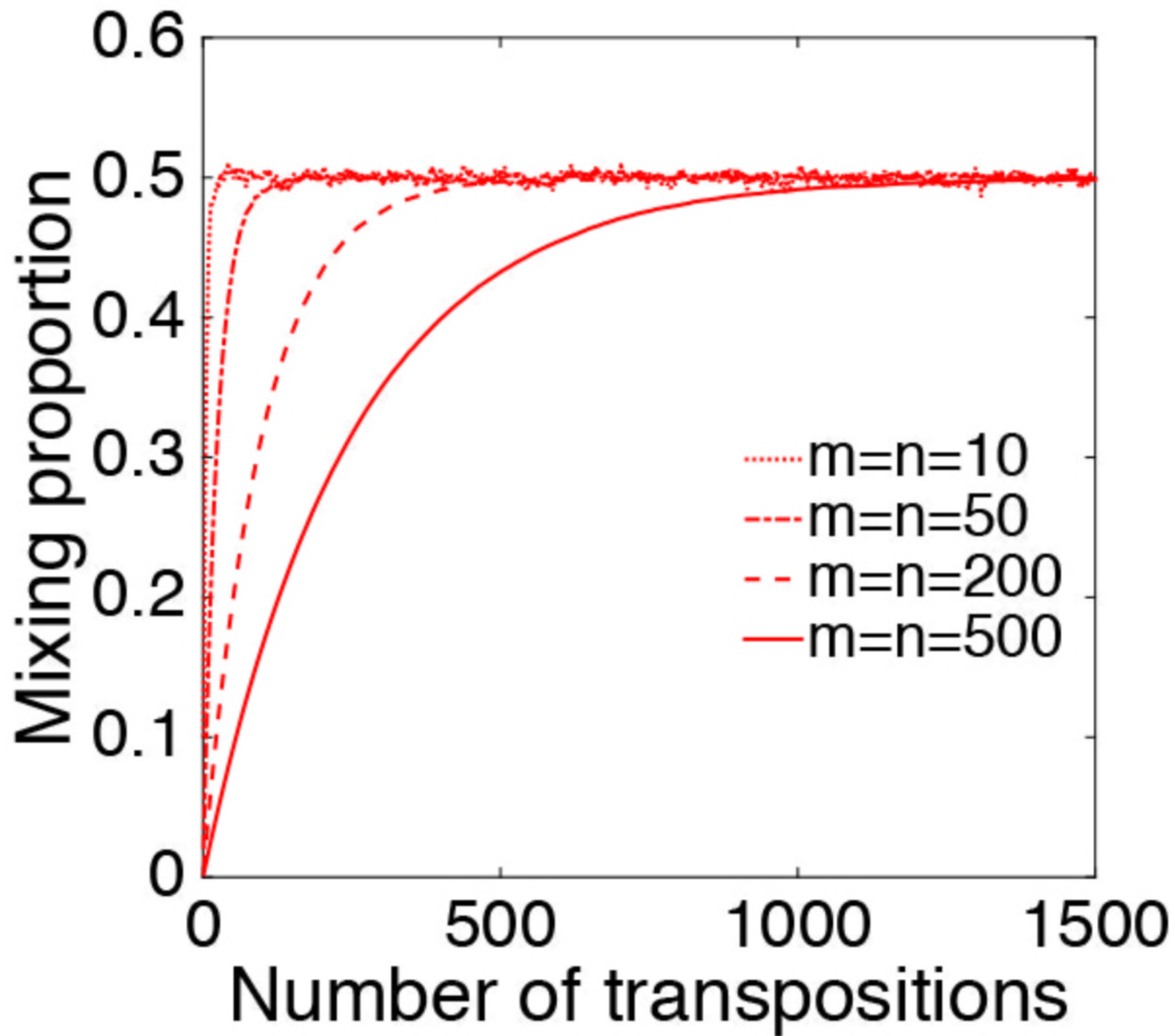
$$f(\mathbf{x}, \mathbf{y}) \dashrightarrow f(\pi_{ij}(\mathbf{x}), \pi_{ij}(\mathbf{y}))$$

Theorem 3. If the test statistic is algebraic, there exists a function g such that

$$f(\pi_{ij}(\mathbf{x}), \pi_{ij}(\mathbf{y})) = g(f(\mathbf{x}, \mathbf{y}), x_i, y_i)$$

where computational complexity of g is constant.

Theorem 4. Any permutation in S_{m+n} can be reachable by a sequence of walks.



Online computation

$$\mu(\mathbf{x}) = \frac{1}{m} \sum_{k=1}^m x_k \quad \text{Transpose } i\text{-th and } j\text{-th} \quad \mu(\mathbf{y}) = \frac{1}{m} \sum_{k=1}^n y_k \quad \mathcal{O}(n)$$

←-----→

$$m\mu(\pi_{ij}(\mathbf{x})) = m\mu(\mathbf{x}) + y_j - x_i \quad \mathcal{O}(2)$$

Transpose
 i -th and j -th

$$\sigma^2(\mathbf{x}) = \frac{1}{m-1} \sum_{k=1}^m (x_k - \mu(\mathbf{x}))^2$$

$$\sigma^2(\mathbf{y}) = \frac{1}{m-1} \sum_{k=1}^n (y_k - \mu(\mathbf{y}))^2 \quad \mathcal{O}(3m+2)$$

$$(m-1)\sigma^2(\pi_{ij}(\mathbf{x})) = \quad \mathcal{O}(9)$$

$$(m-1)\sigma^2(\mathbf{x}) + \left((m\mu(\mathbf{x}))^2 - (m\mu(\pi_{ij}(\mathbf{x})))^2 \right) / m + y_j^2 - x_i^2$$

Computational Complexity

$$T(\pi_{ij}(\mathbf{x}), \pi_{ij}(\mathbf{y})) = \frac{\left(\frac{\nu(\pi_{ij}(\mathbf{x}))}{m} - \frac{\nu(\pi_{ij}(\mathbf{y}))}{n} \right)}{\sqrt{\frac{\omega^2(\pi_{ij}(\mathbf{x})) + \omega^2(\pi_{ij}(\mathbf{y}))}{m+n-2} \left(\frac{1}{m} + \frac{1}{n} \right)}}$$

T-stat computation per permutation

Standard method: $O(4m+4n+20)$

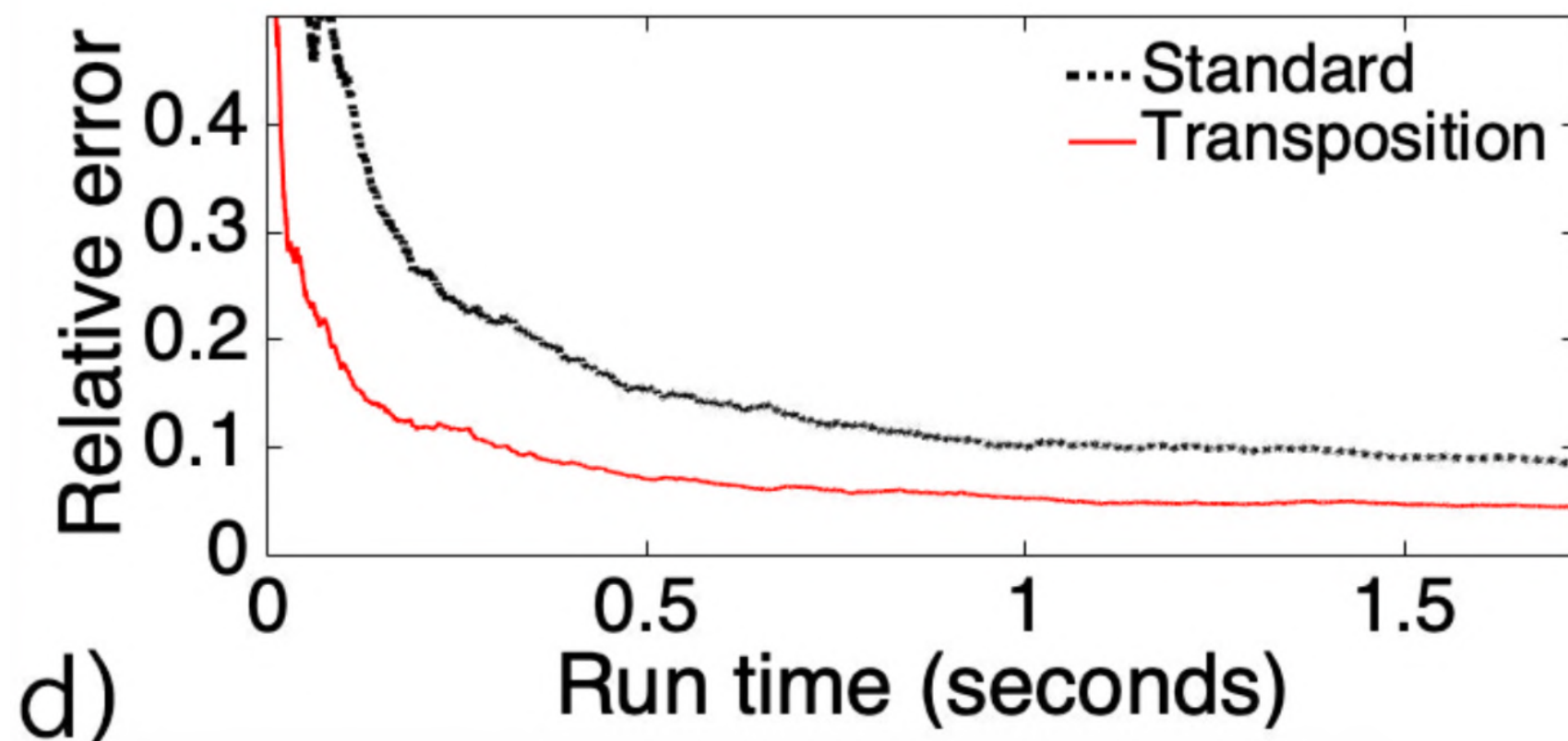
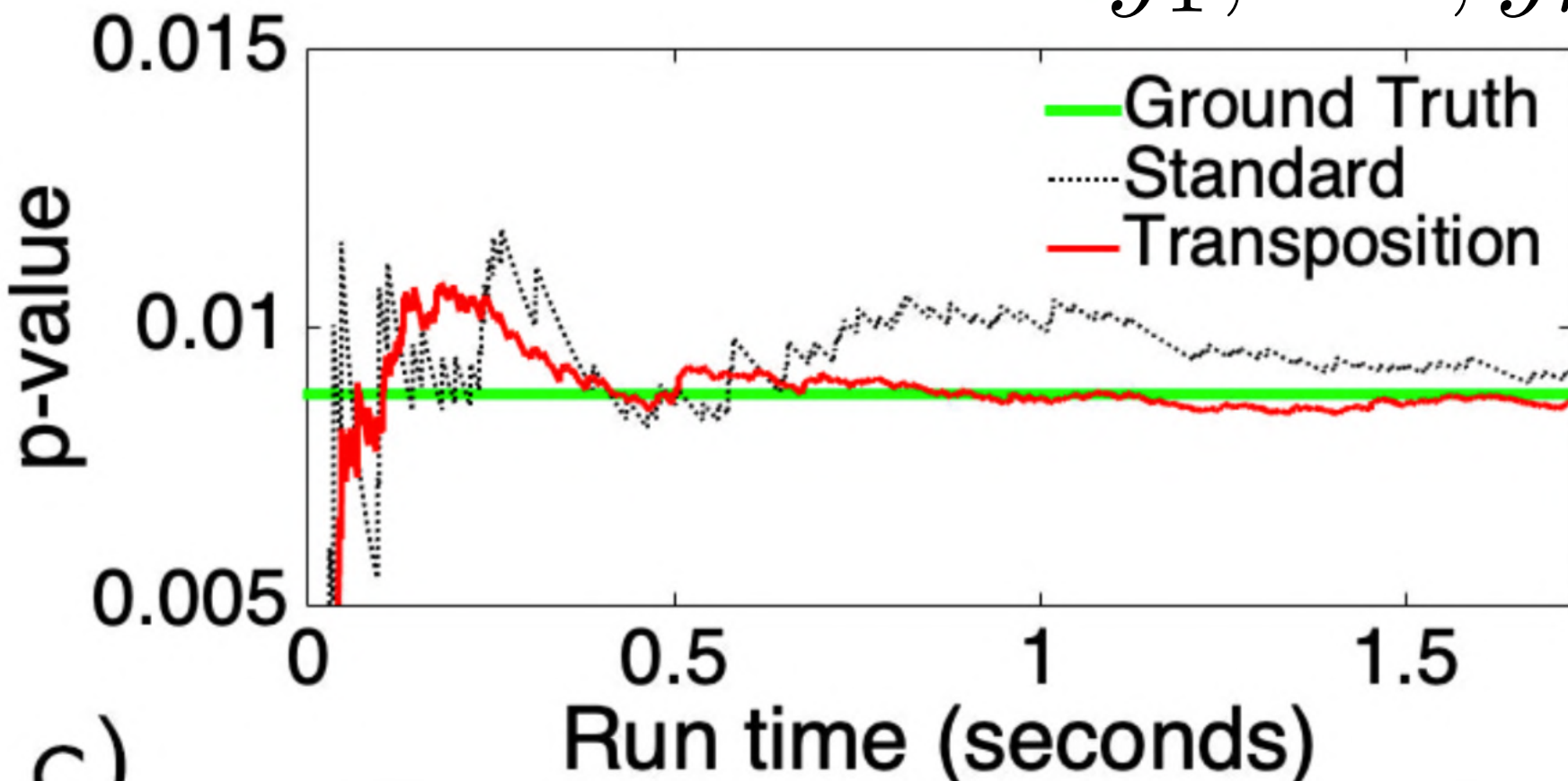
Our approach: $O(35)$

Simulation

$$x_1, \dots, x_m \sim N(0, 1)$$

$$y_1, \dots, y_n \sim 0.1 + N(0, 1)$$

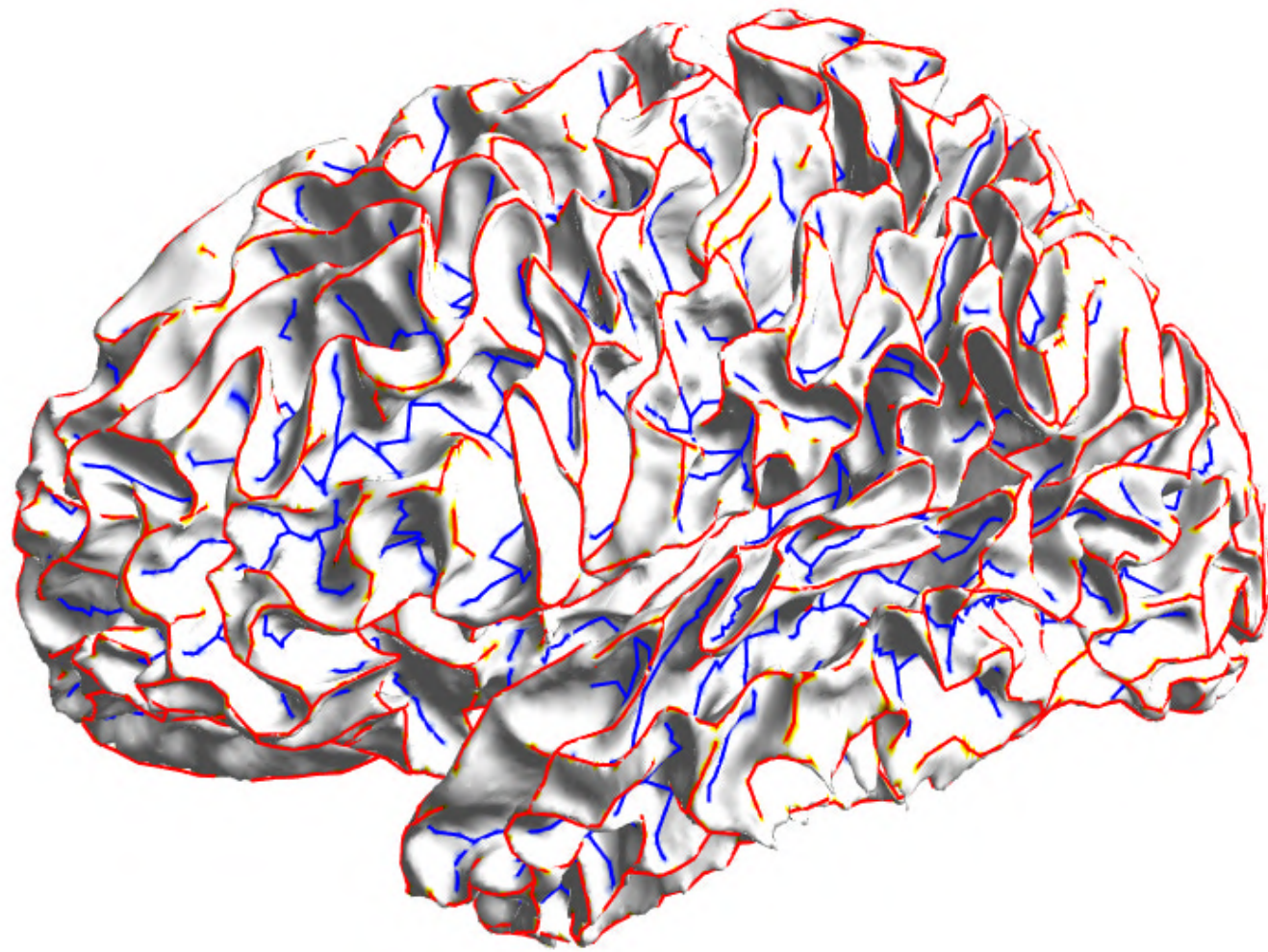
$$m=n=100$$



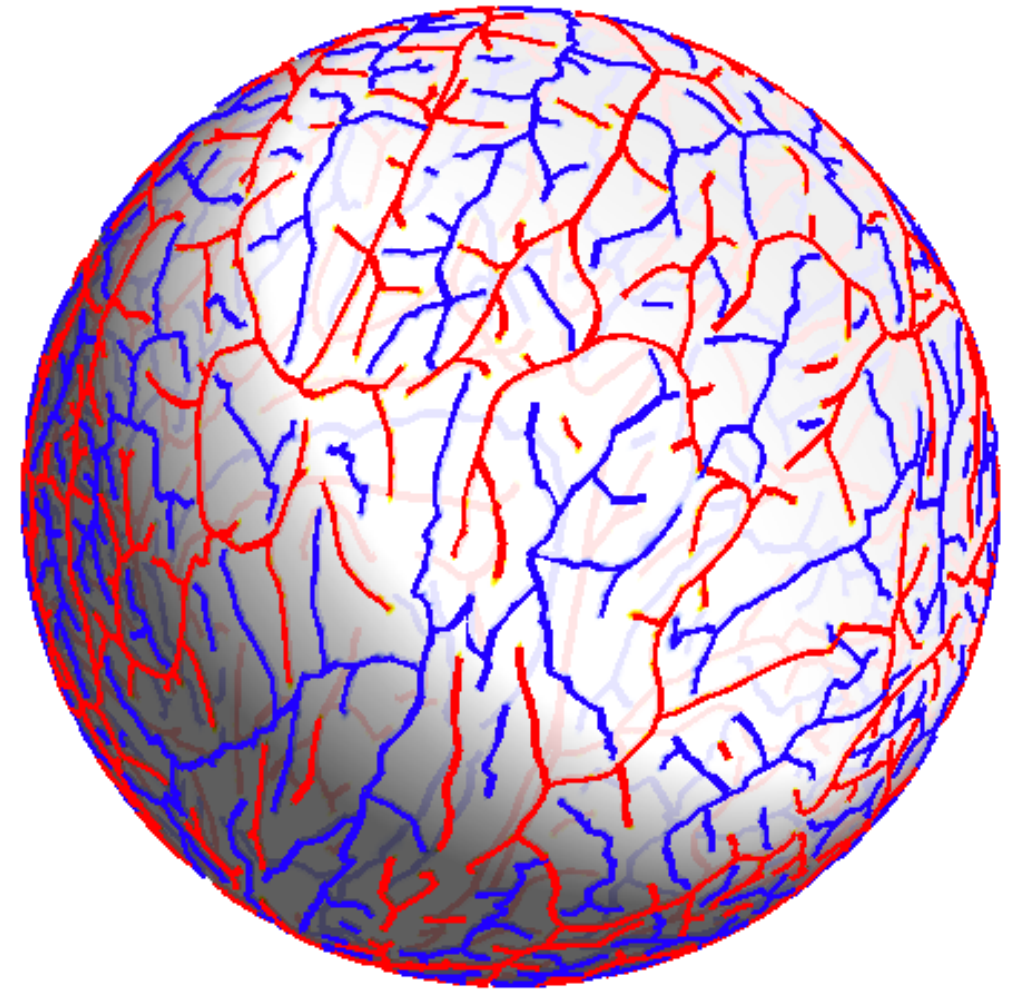
Sulcal and gyral
graphs on brain
cortical surface

Sulci = mountain regions of the brain (**red**)

Gyri = valley regions of the brain (**blue**)



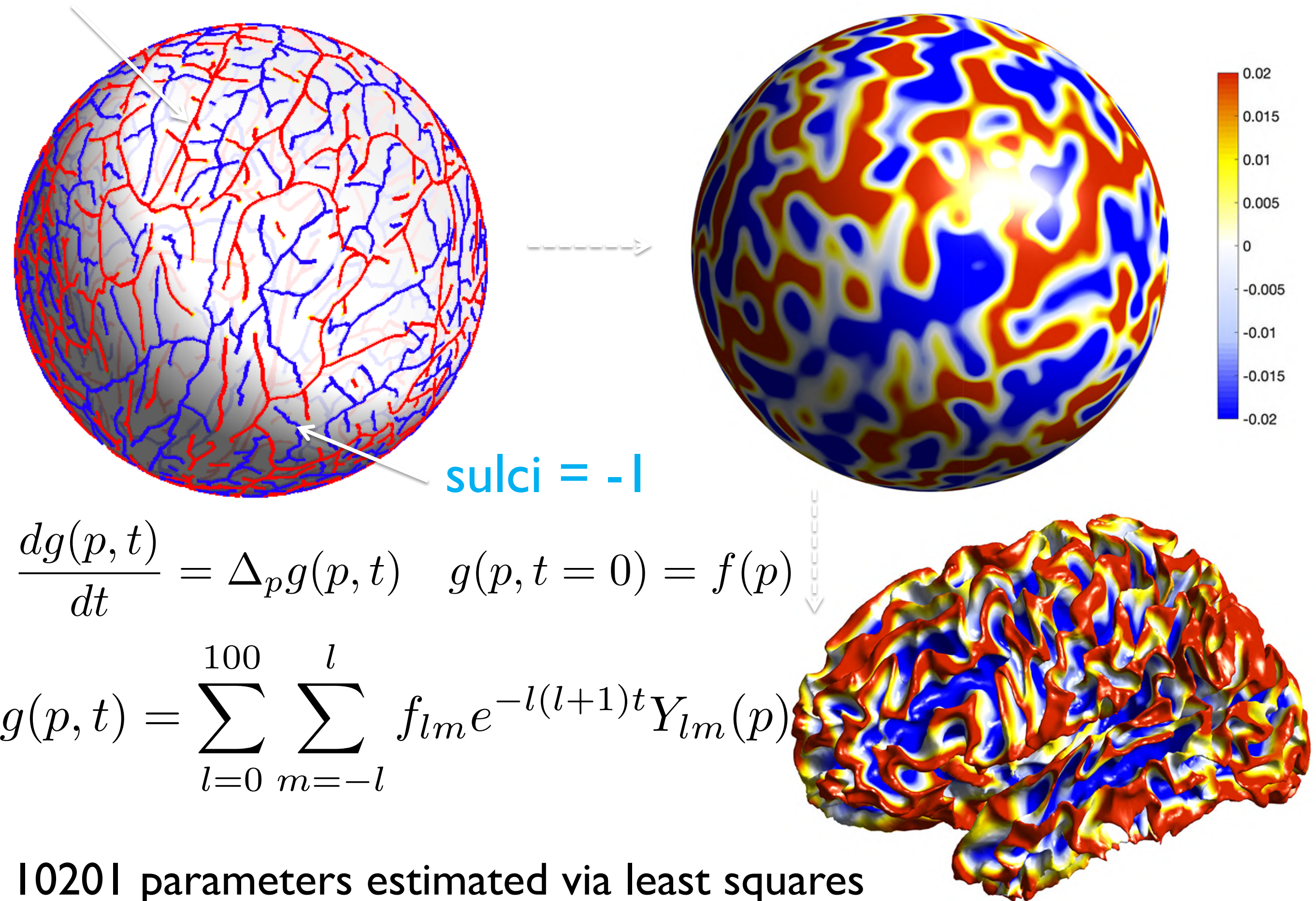
White matter
surface



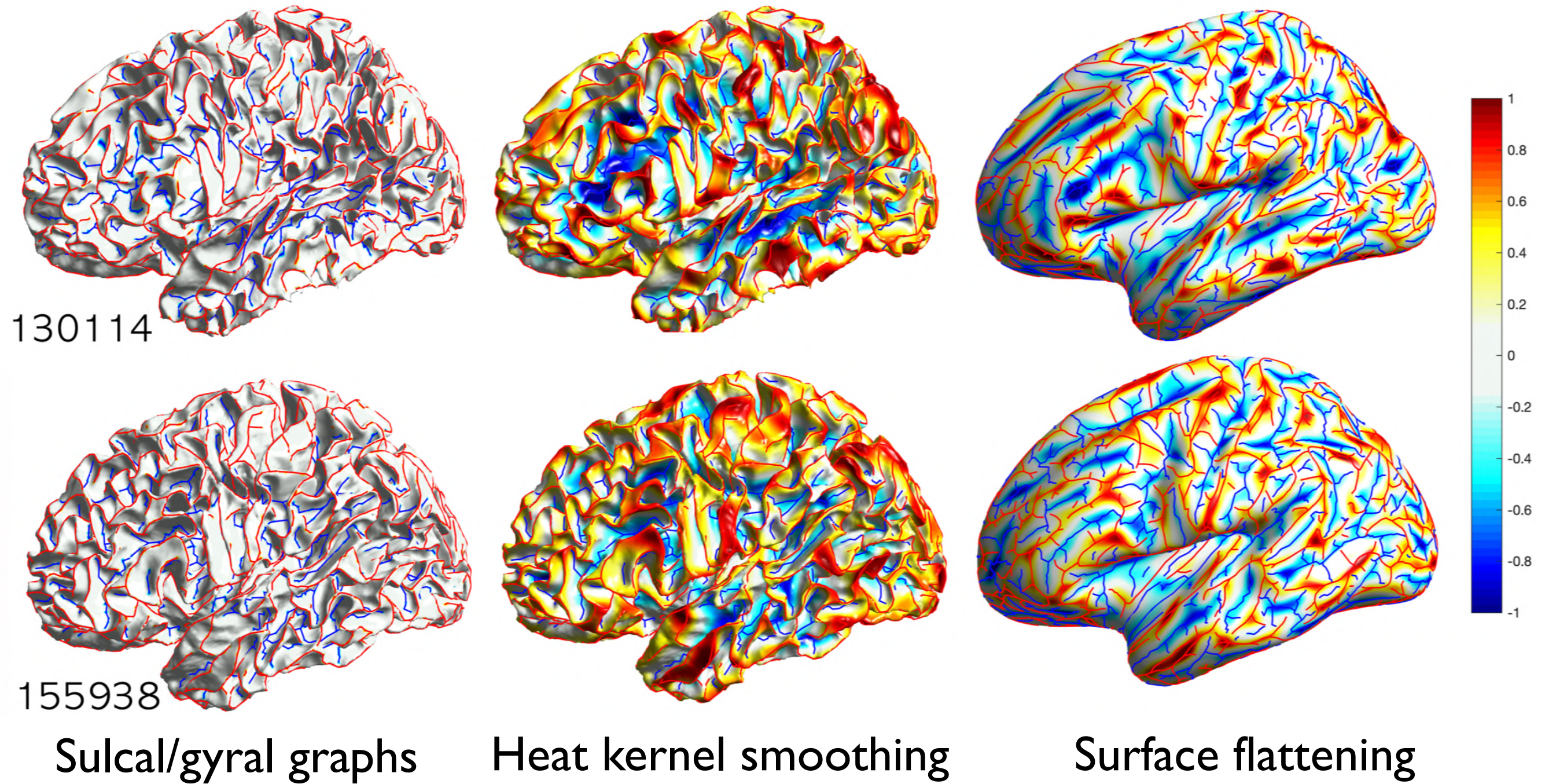
Sulcal/gyral graphs
on manifold

gyri=+1

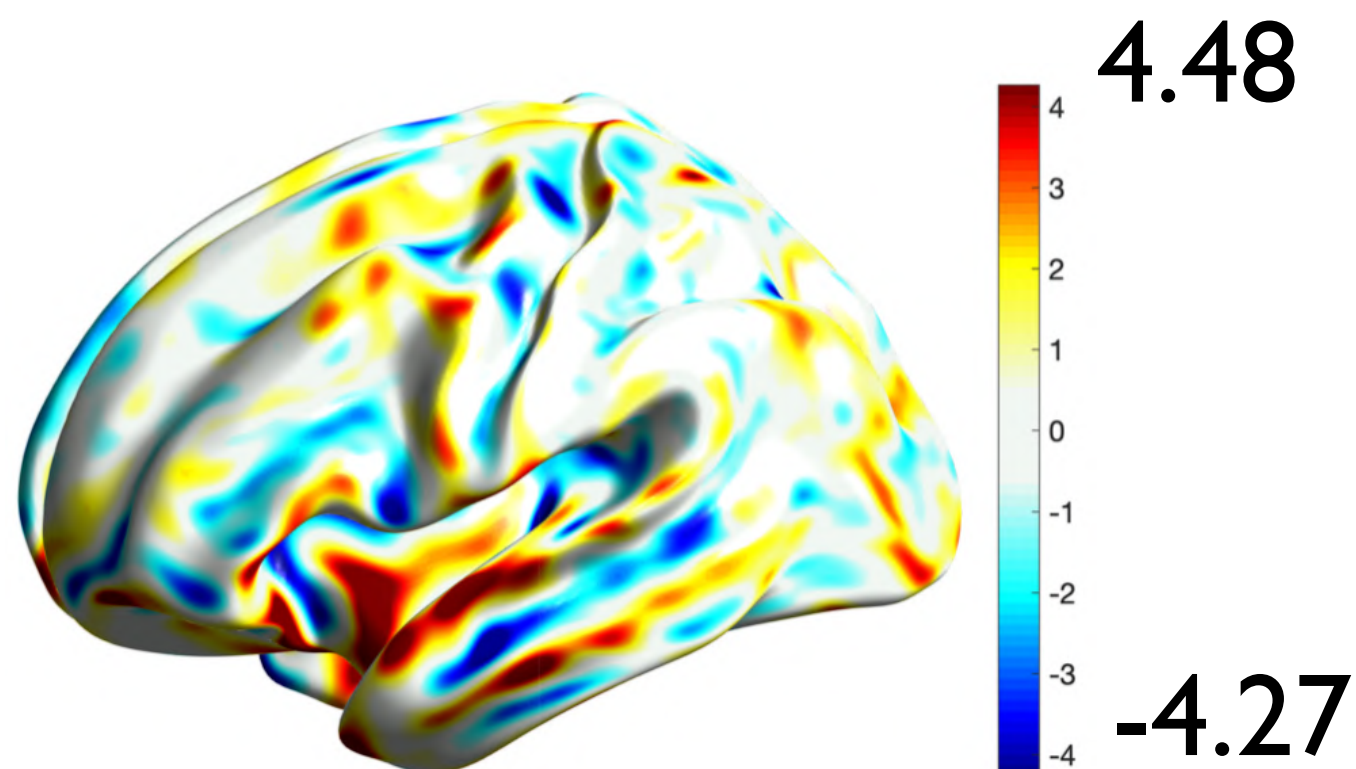
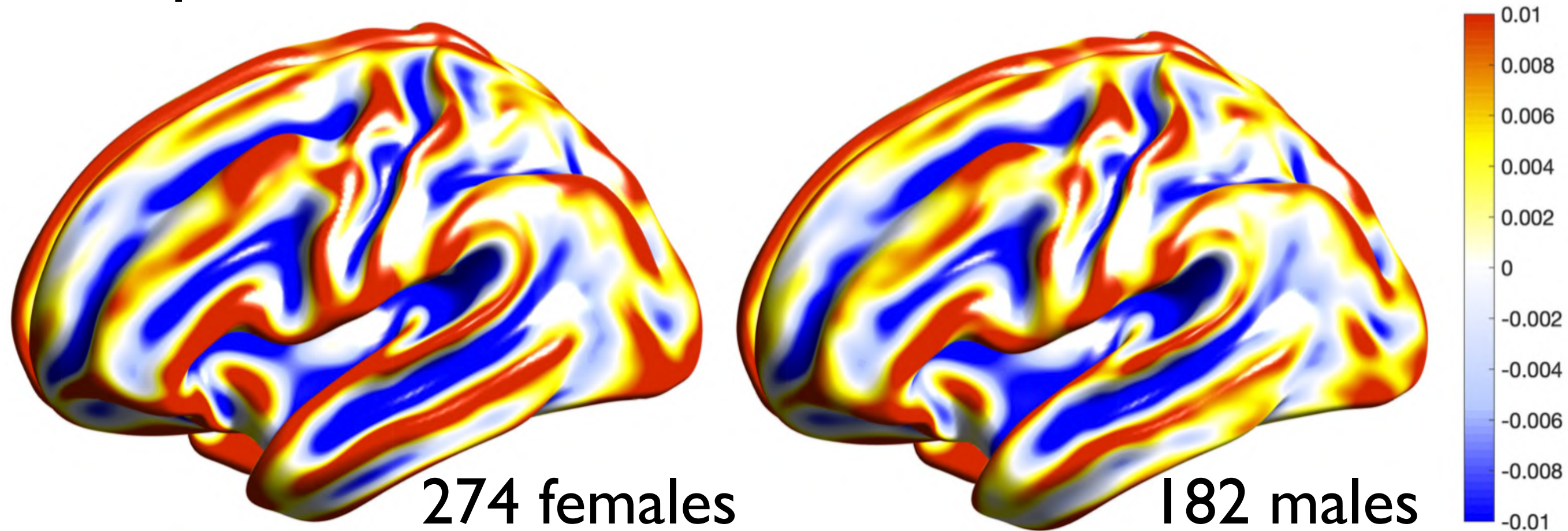
Heat kernel smoothing (*Chung et al. 2007 IEEE TMI*)



Heat kernel smoothing



Group mean differences



t-statistic (thresholds at
corrected pvalue of 0.05)

Our method: **40min.**

Standard methods: **18 days**



Thank you!

Question? mkchung@wisc.edu