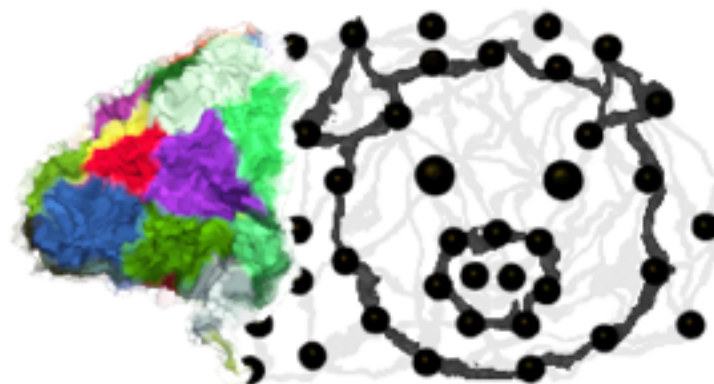


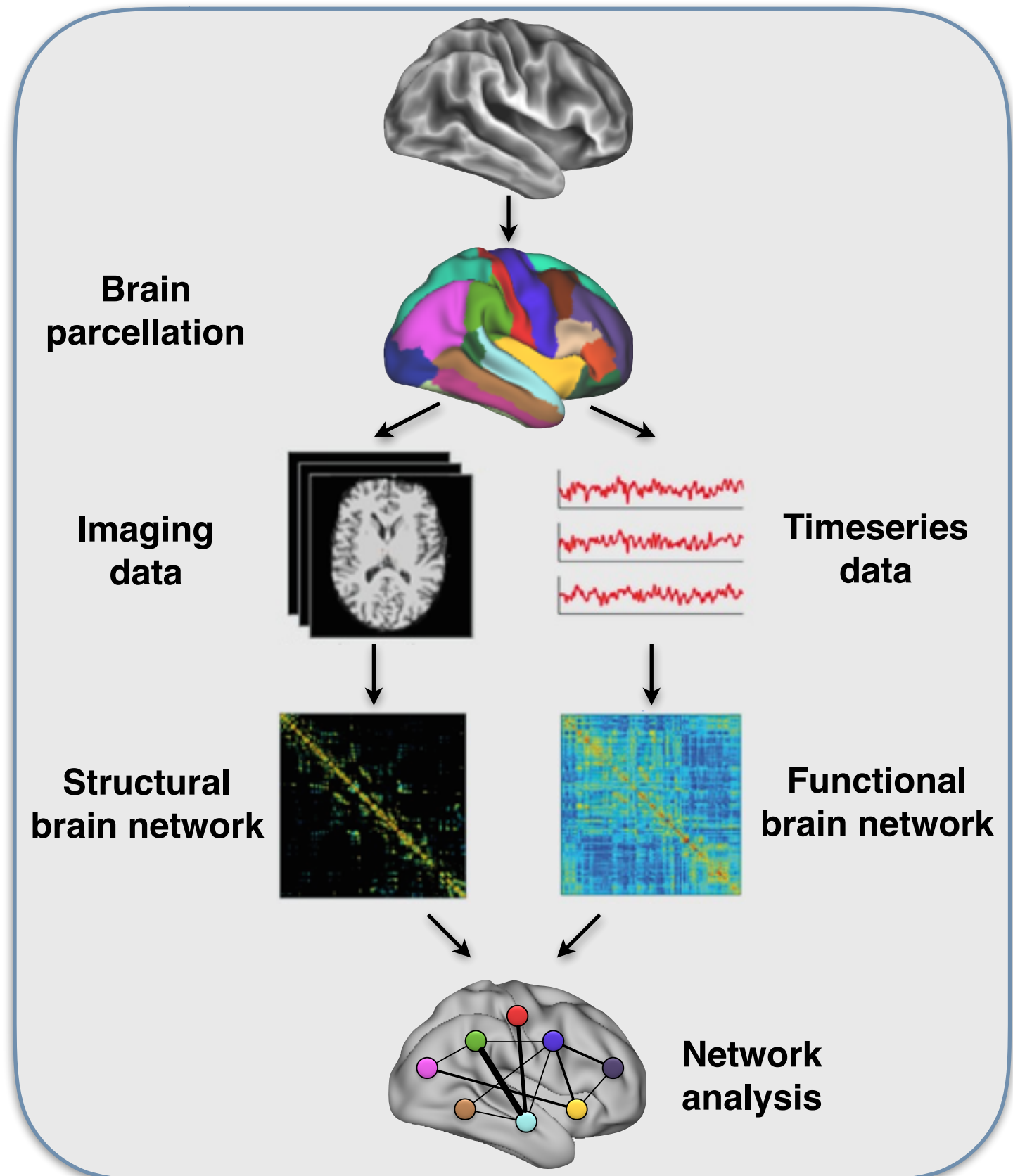
# COMPARISON OF BRAIN NETWORKS WITH UNKNOWN CORRESPONDENCES

Sl Ktena, S Parisot, J Passerat-Palmbach and D Rueckert

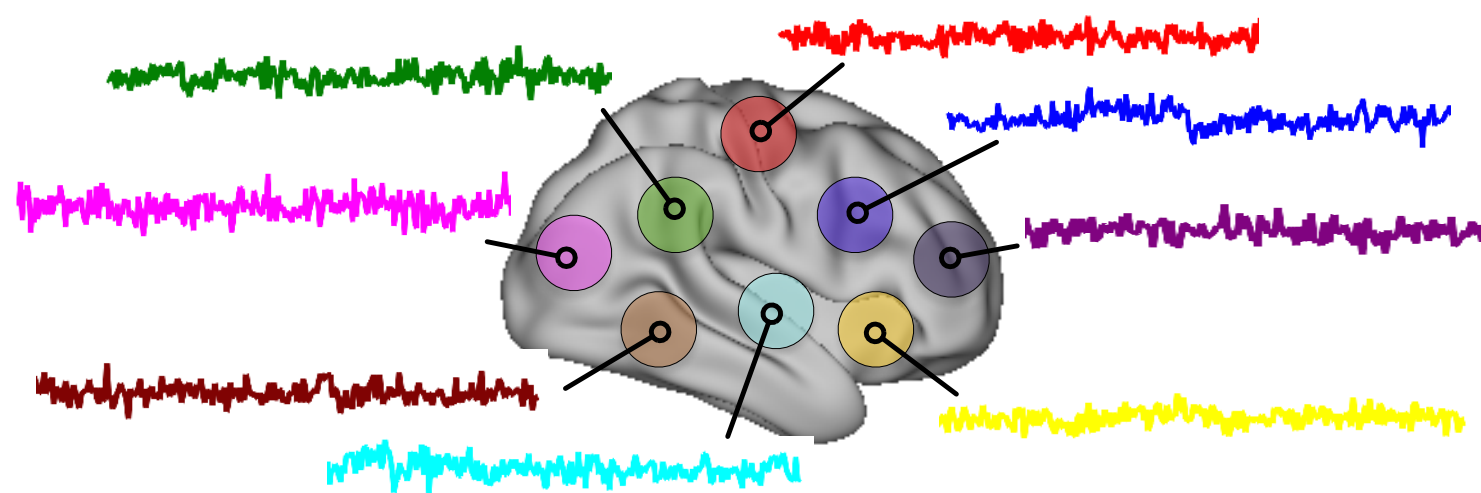


# The brain from a network perspective

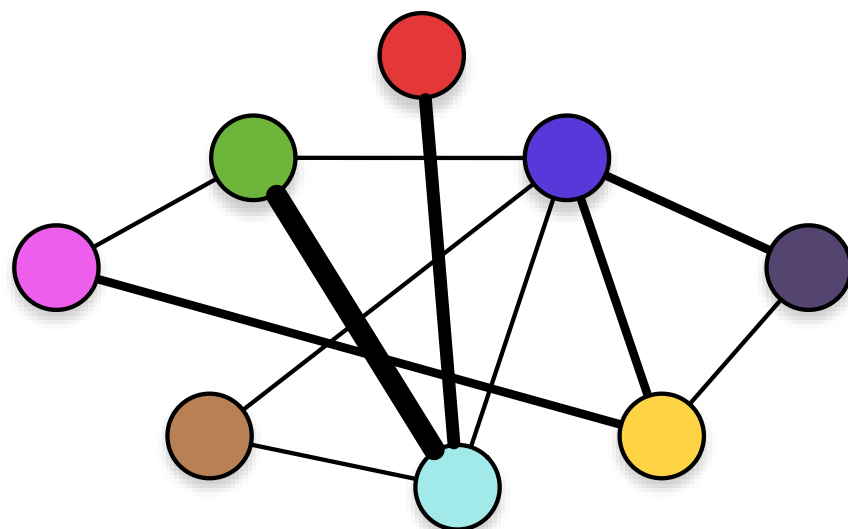
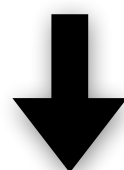
- cognition is a network phenomenon [Sporns, *Dial. Clin. Neurosc.* (2013)]
- no physical trace of certain diseases, only changes in the physical wiring and strength of connections
- given two brain graphs representing connectivity, how similar are they?  
(*within/between subjects, between modalities etc.*)



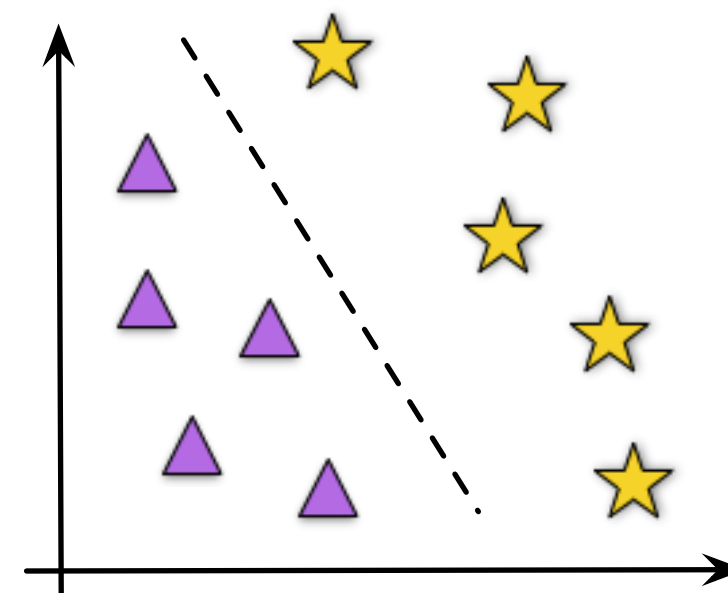
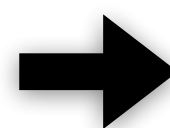
# Inference: Naive approach



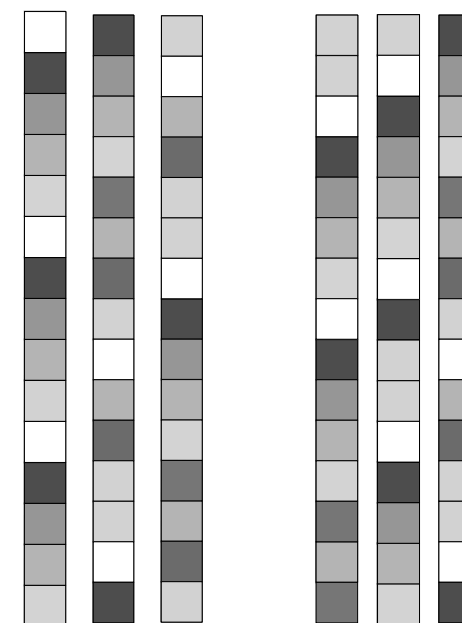
Graph  
construction





Graph  
embedding



Model training  
and inference



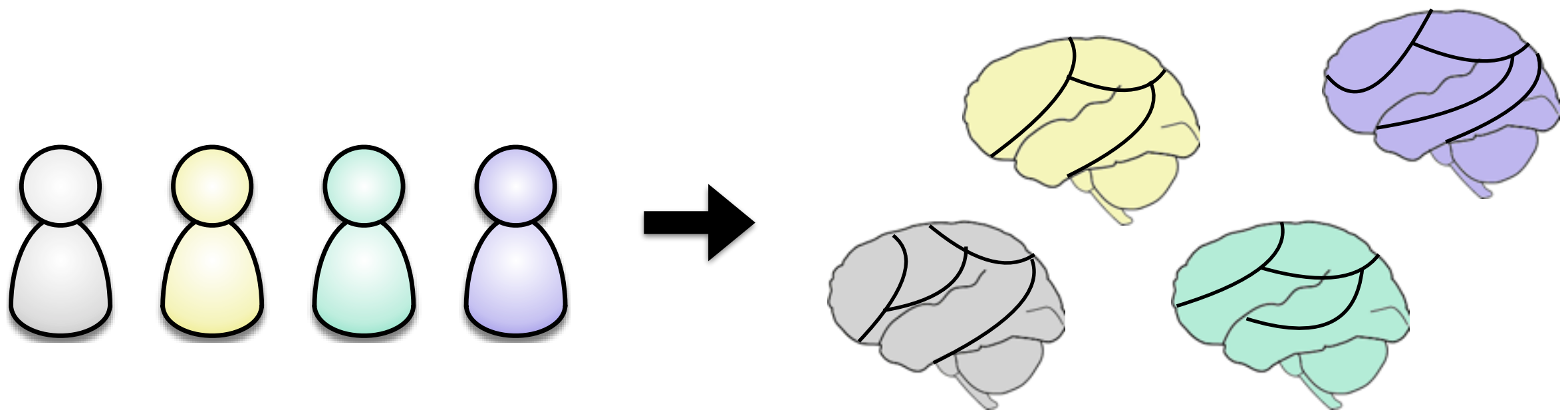
class 

class 

Richiardi et al., *IEEE Signal Processing Magazine* (2013)  
Varoquaux and Craddock, *Neuroimage* (2013)

# Why individual parcellations?

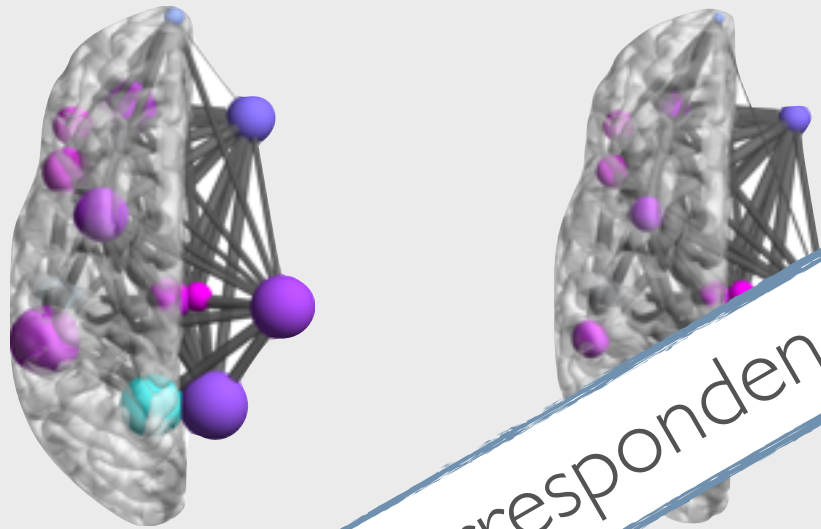
- Standard anatomical atlases subdivide the brain based on cytoarchitecture (e.g. Brodmann) or anatomical landmarks (e.g. Desikan-Killiany)
- Individual variability in terms of anatomy or function due to maturation or brain injury are not accounted for [Timofiyeva, *PLoS One* (2014)]
- Data-driven single subject parcellations capture this variability



# Inexact graph matching

- Evaluate how much two graphs share

## Graph embedding

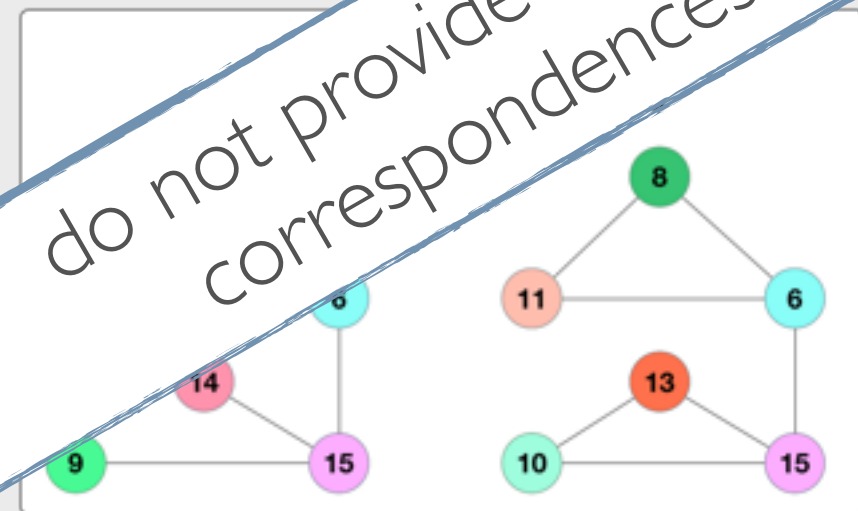
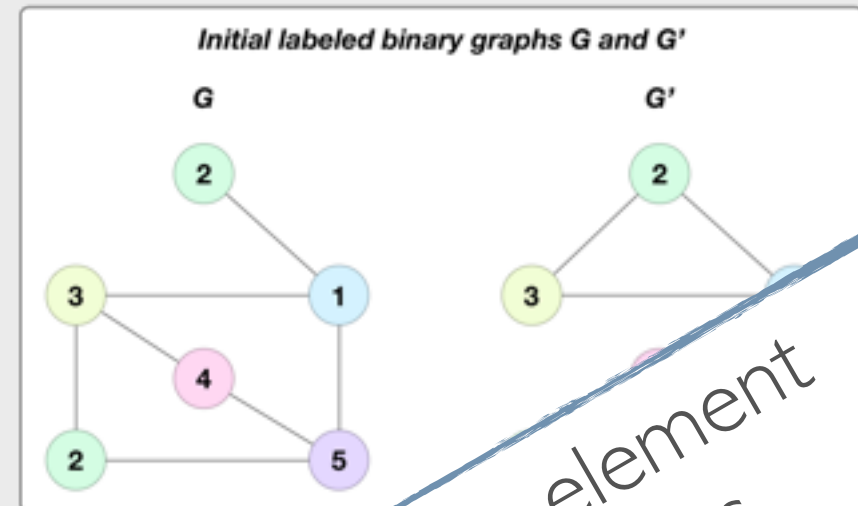


requires node correspondences



Craddock et al., *Nature Methods* (2013)

## Graph kernels



Feature vector representations of  $G$  and  $G'$

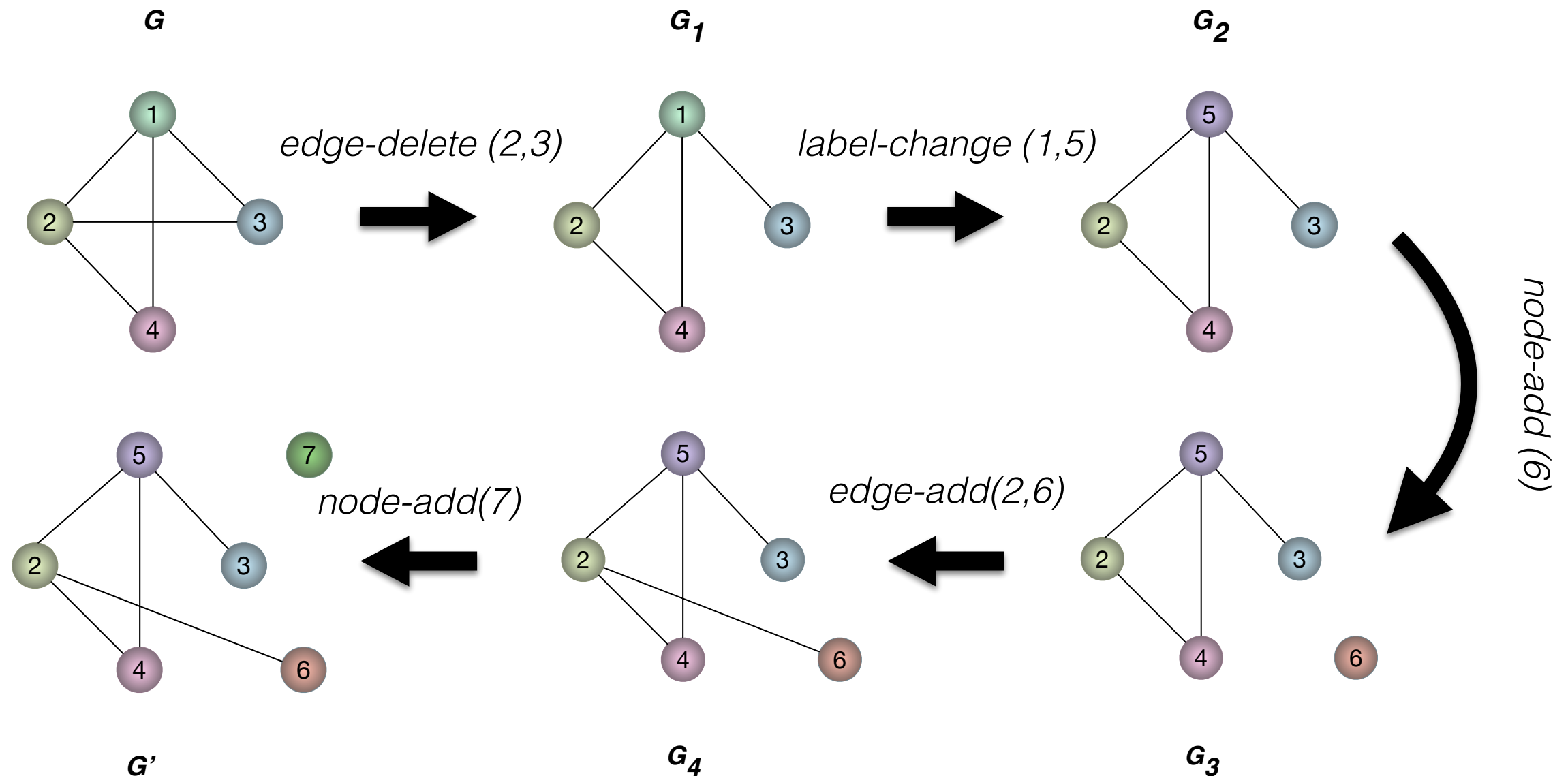
$$\phi_{WL}(G) = (1, 2, 1, 1, 1, 1, 1, 0, 1, 0, 0, 1, 0, 1, 1)$$
$$\phi_{WL}(G') = (1, 2, 1, 1, 1, 1, 0, 1, 0, 1, 1, 0, 1, 0, 1)$$
$$k_{WL}(G, G') = \langle \phi_{WL}(G), \phi_{WL}(G') \rangle = 8$$

Jie et al. *Human Brain Mapping* (2014)



# Graph edit distance

- Measure of dissimilarity between graphs, defined directly in their domain  $\mathcal{G}$  as a nonnegative function  $d : \mathcal{G} \times \mathcal{G} \rightarrow \mathbb{R}^+$ . Able to model structural variation in a very intuitive and illustrative way.



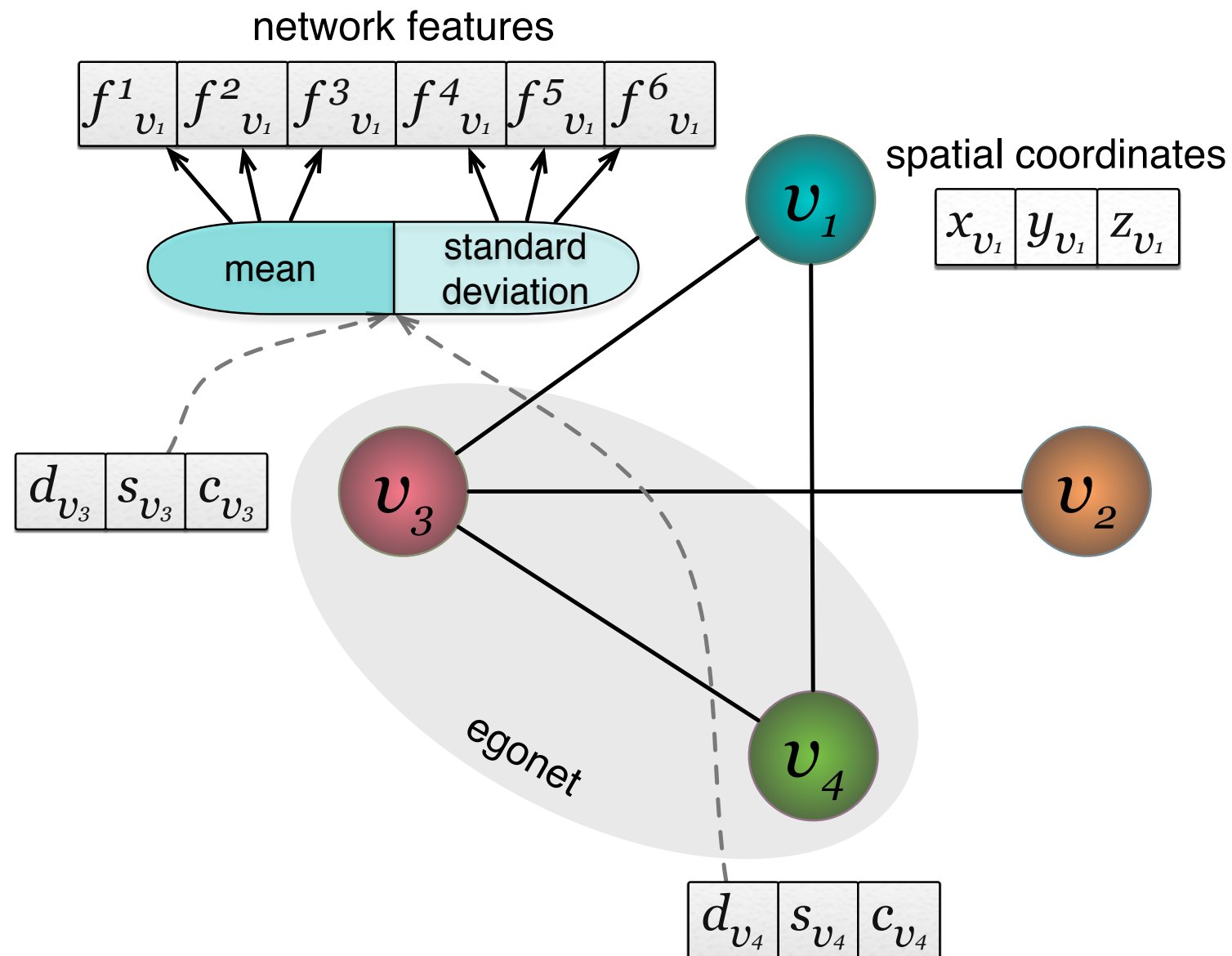
# GED computation

- The Hungarian algorithm provides a fast approximate solution to the GED computation [Riesen & Bunke, *Img & Vis. Comp.*, (2009)]
- Given two labeled graphs  $G_1, G_2$  with  $|\mathcal{V}(G_1)| = n$  and  $|\mathcal{V}(G_2)| = m$  a square cost matrix  $\mathbf{C}$  of order  $n + m$  is defined, which encodes all the possible edit operation costs

$$\mathbf{C} = \begin{bmatrix} \boxed{\begin{array}{ccc} c_{1,1} & \dots & c_{1,m} \\ \text{substitutions} \\ c_{n,1} & \dots & c_{n,m} \end{array}} & \boxed{\begin{array}{ccc} c_{1,\varepsilon} & \dots & \infty \\ \text{deletions} \\ \infty & \dots & c_{n,\varepsilon} \end{array}} \\ \boxed{\begin{array}{ccc} c_{\varepsilon,1} & \dots & \infty \\ \text{insertions} \\ \infty & \dots & c_{\varepsilon,m} \end{array}} & \begin{array}{ccc} 0 & \dots & 0 \\ \vdots & \ddots & \vdots \\ 0 & \dots & 0 \end{array} \end{bmatrix}$$

# Node features

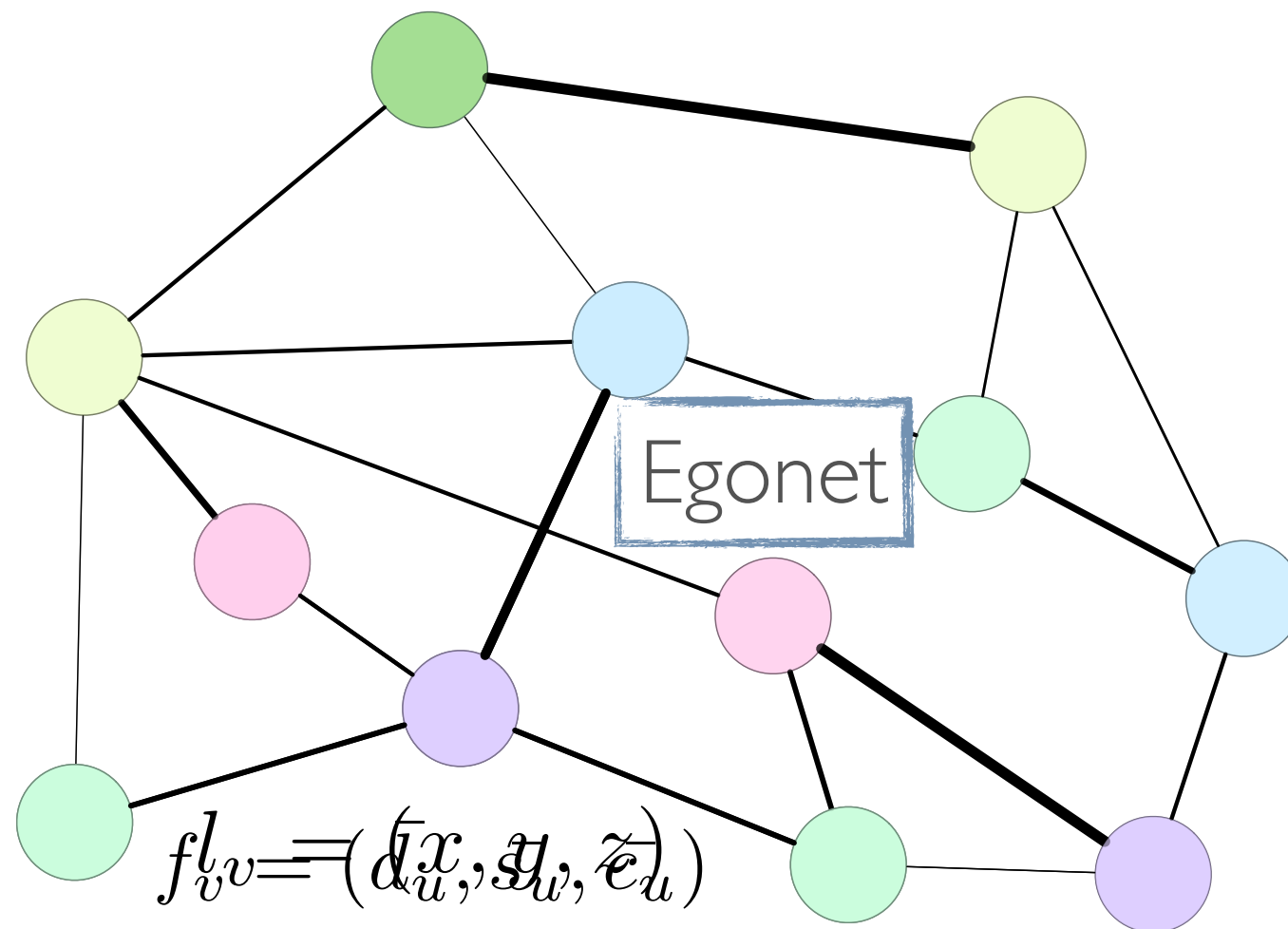
- Spatial information (coordinates in standard brain space)
- Feature information (network measures, egonet based)





# Node distance

- Spatial distance
- Feature distance



$$d_l = d(v_1, v_2) = d_{\text{eucl}}(v_1, v_2) + d_{\text{canb}}(v_1, v_2) = \sum_{i=1}^d \frac{|f_{v_{1i}} - f_{v_{2i}}|}{|f_{v_{1i}}| + |f_{v_{2i}}|}$$

# Tailoring GED for brain graphs

- In order to achieve better approximation of the true edit distance, edge operations need to be involved.
- Constrain substitutions to the nearest  $N$  nodes - set cost to  $Inf$  for the rest of the substitutions

**if**  $u_j \in neigh(v_i)$  **then**

$$c_{i,j} = \alpha * d_{euclidean}(v_i, u_j) + (1 - \alpha) * d_{canberra}(v_i, u_j) + d_{edge}(v_i, u_j)$$

**else**

$$c_{i,j} = \infty$$

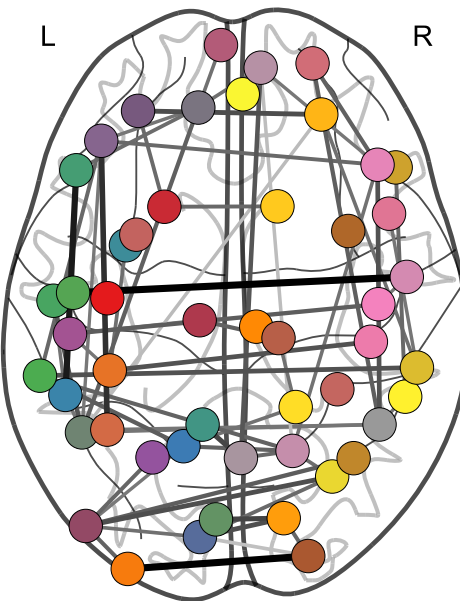
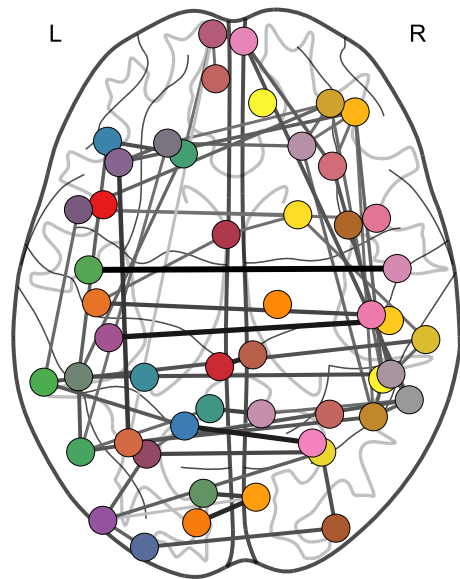
- Take into account node betweenness centrality  $g$  for the cost of node insertion/deletion

$$c_{\epsilon,j} = \alpha + (1 - \alpha) * g(u_j)$$

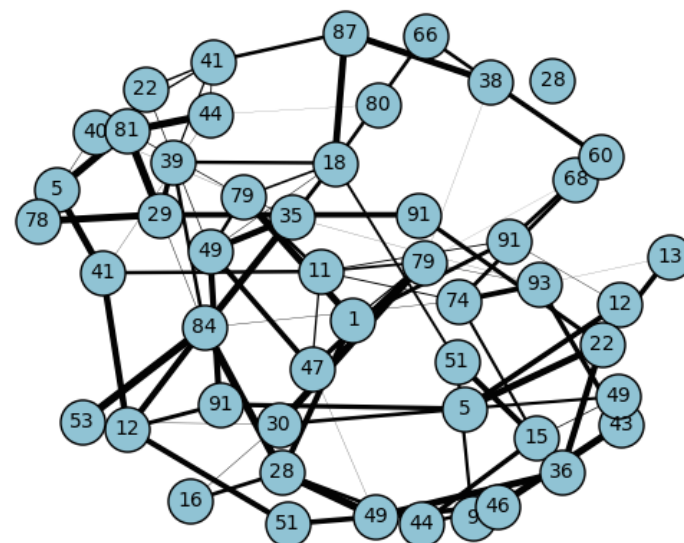
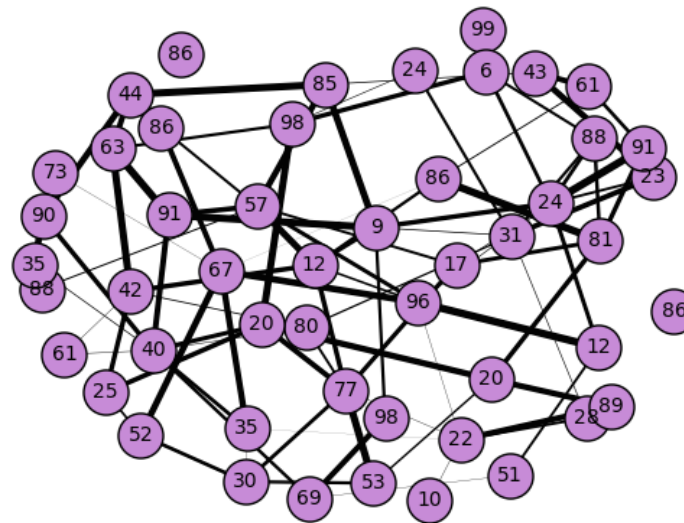
$$c_{i,\epsilon} = \alpha + (1 - \alpha) * g(v_i)$$

# Summary

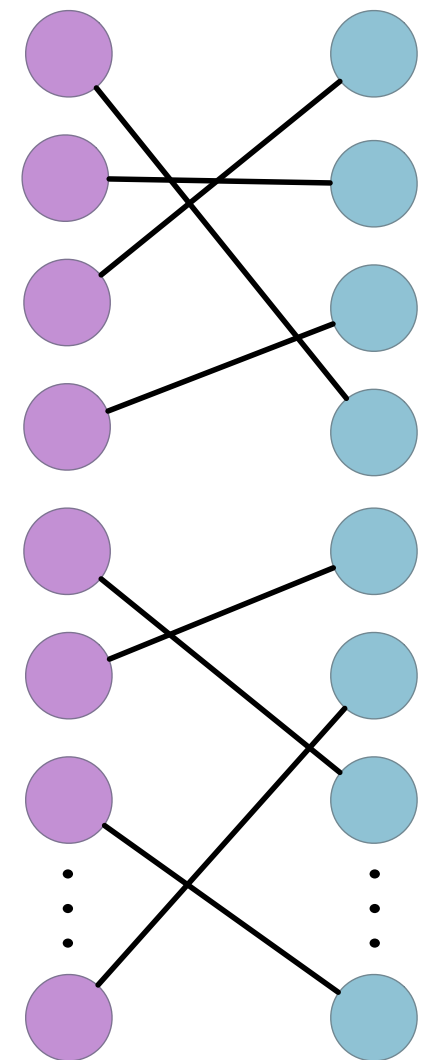
## Brain connectivity networks



## Labeled graphs

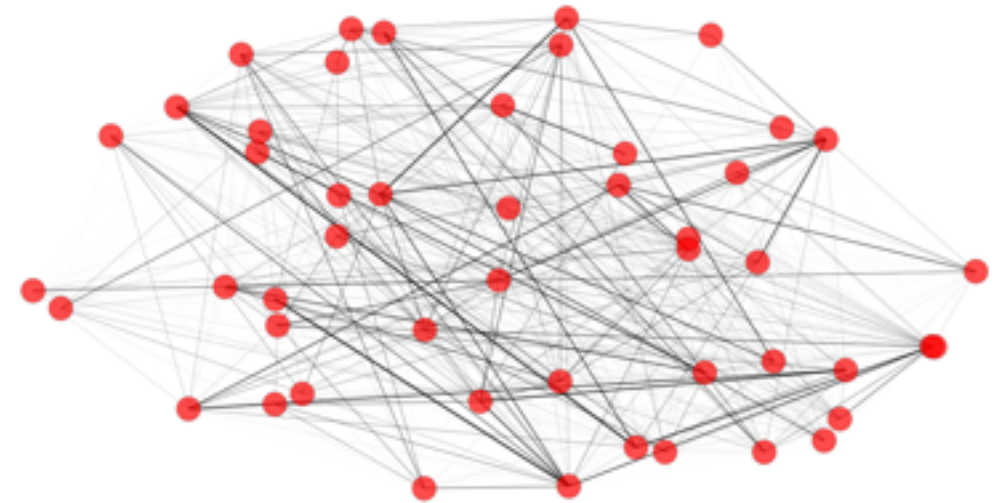
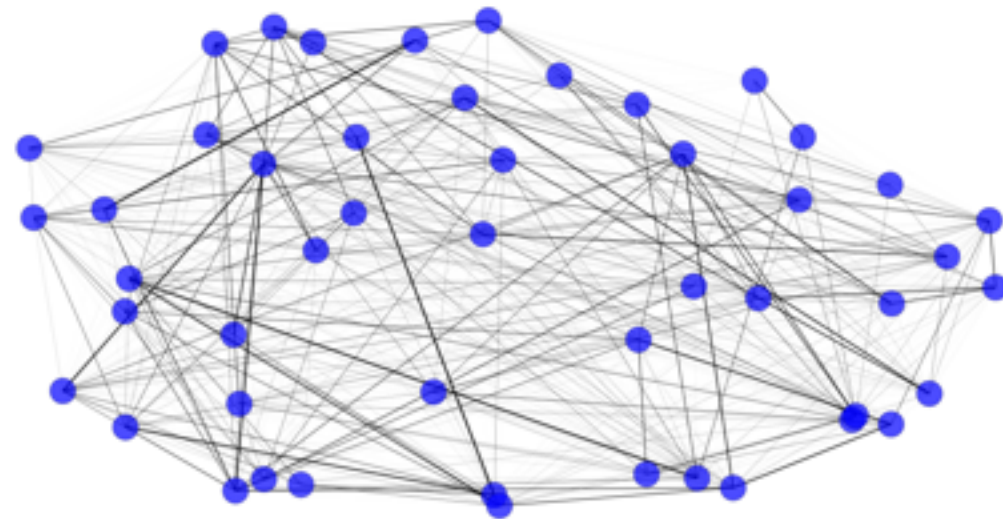


## Graph Edit Distance (Hungarian algorithm)



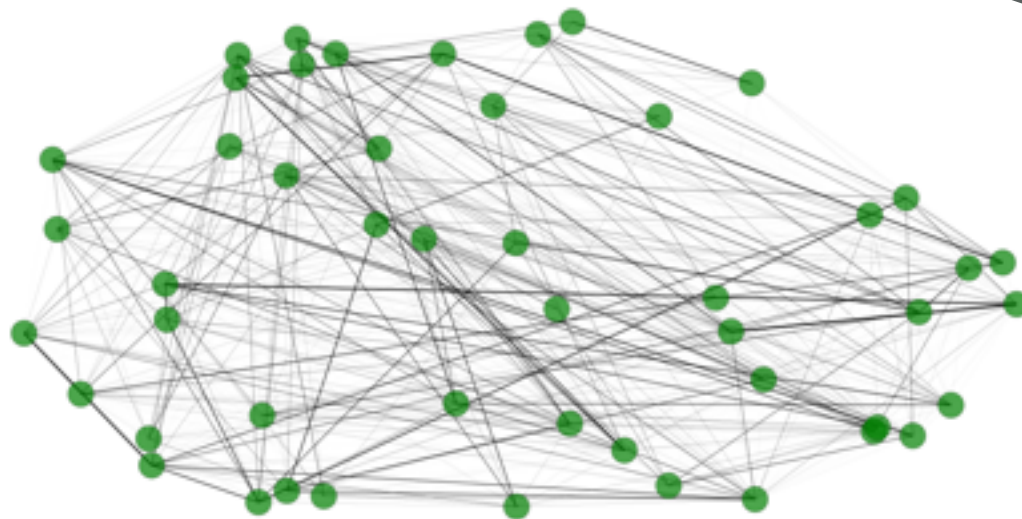
- Diffusion and functional MRI data from the Human Connectome Project
- 30 healthy unrelated subjects as well as 20 monozygotic and 20 dizygotic female twin pairs (MZ twins share 100% of genetic information, while DZ share only 50%)
- Connectivity driven **single-subject** parcellations [Parisot et al., *MICCAI*, (2016)]
- Structural networks derived with probabilistic tractography
- Functional networks estimated using partial correlation

# Single-subject parcellations



same  
subject

different  
subjects

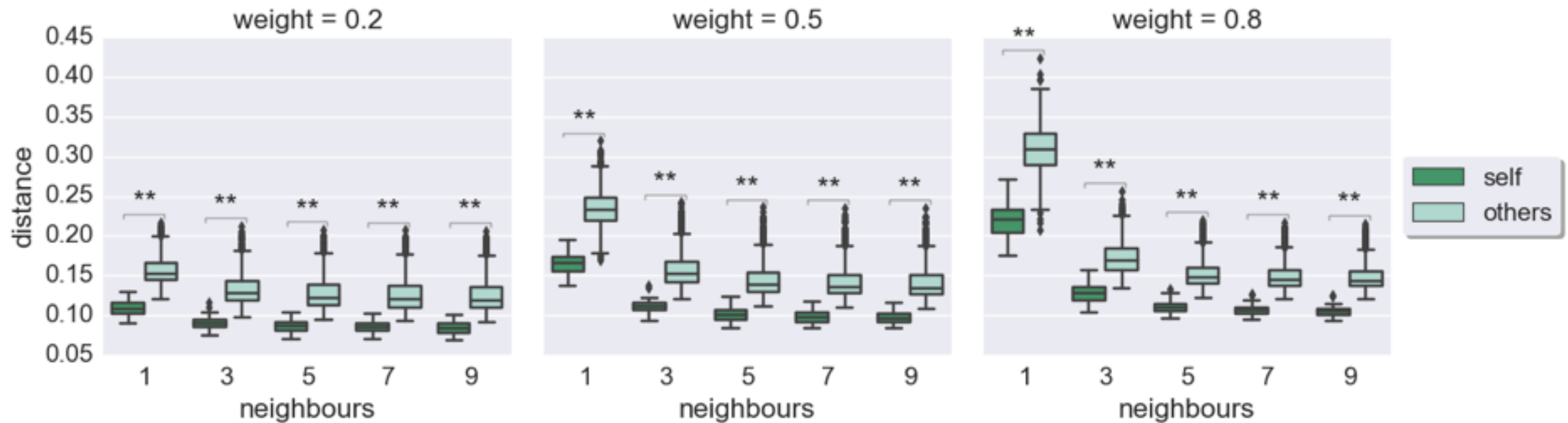


**$\text{GED}(\text{same subject}) < \text{GED}(\text{different subjects})$**

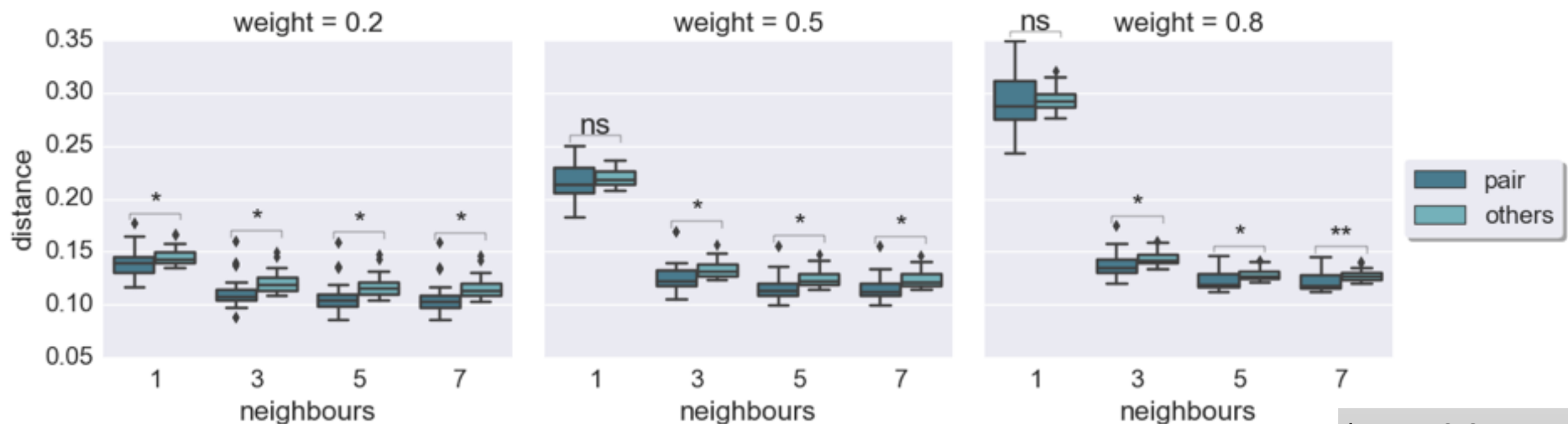


# Structural networks

## Single subject structural networks (50 nodes)



## Structural networks for MZ twins

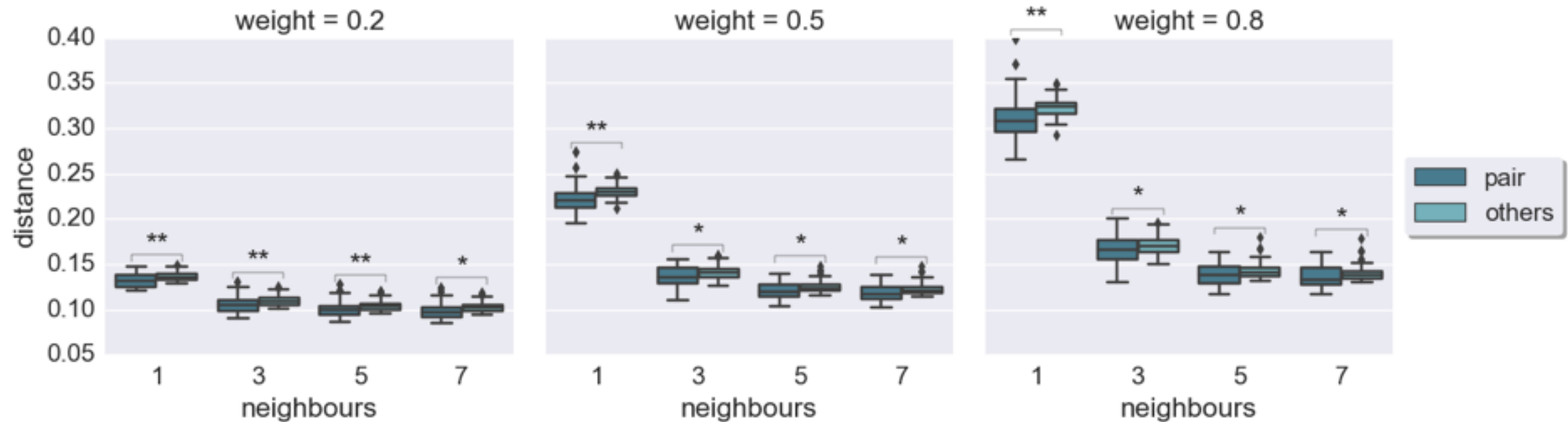


\*  $p < 0.05$   
 \*\*  $p < 0.001$   
 ns non-significant

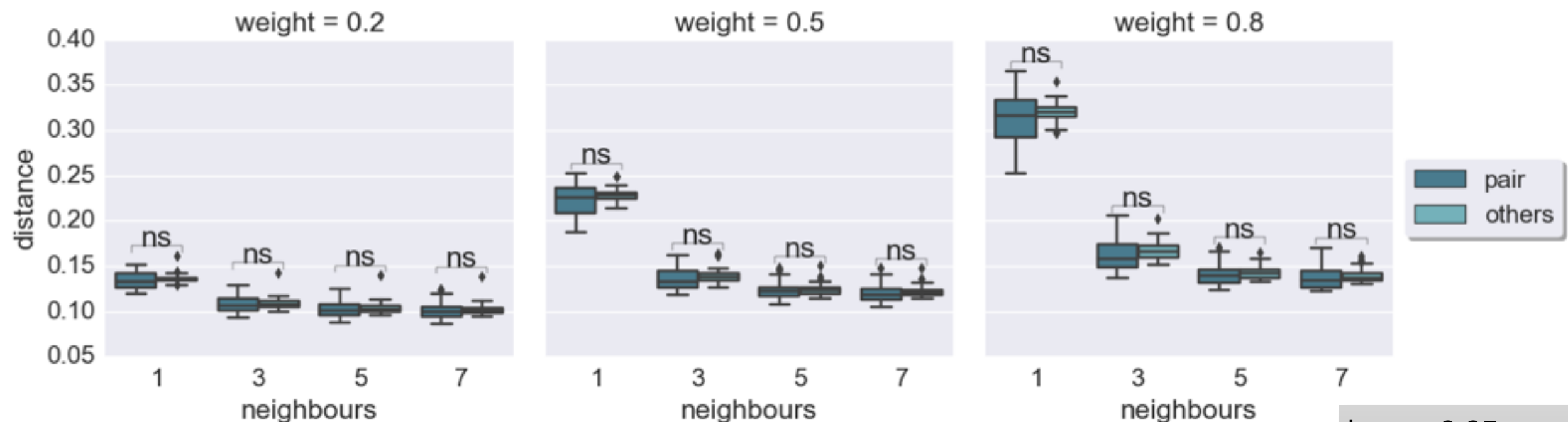


# Functional networks

## Functional networks for MZ twins

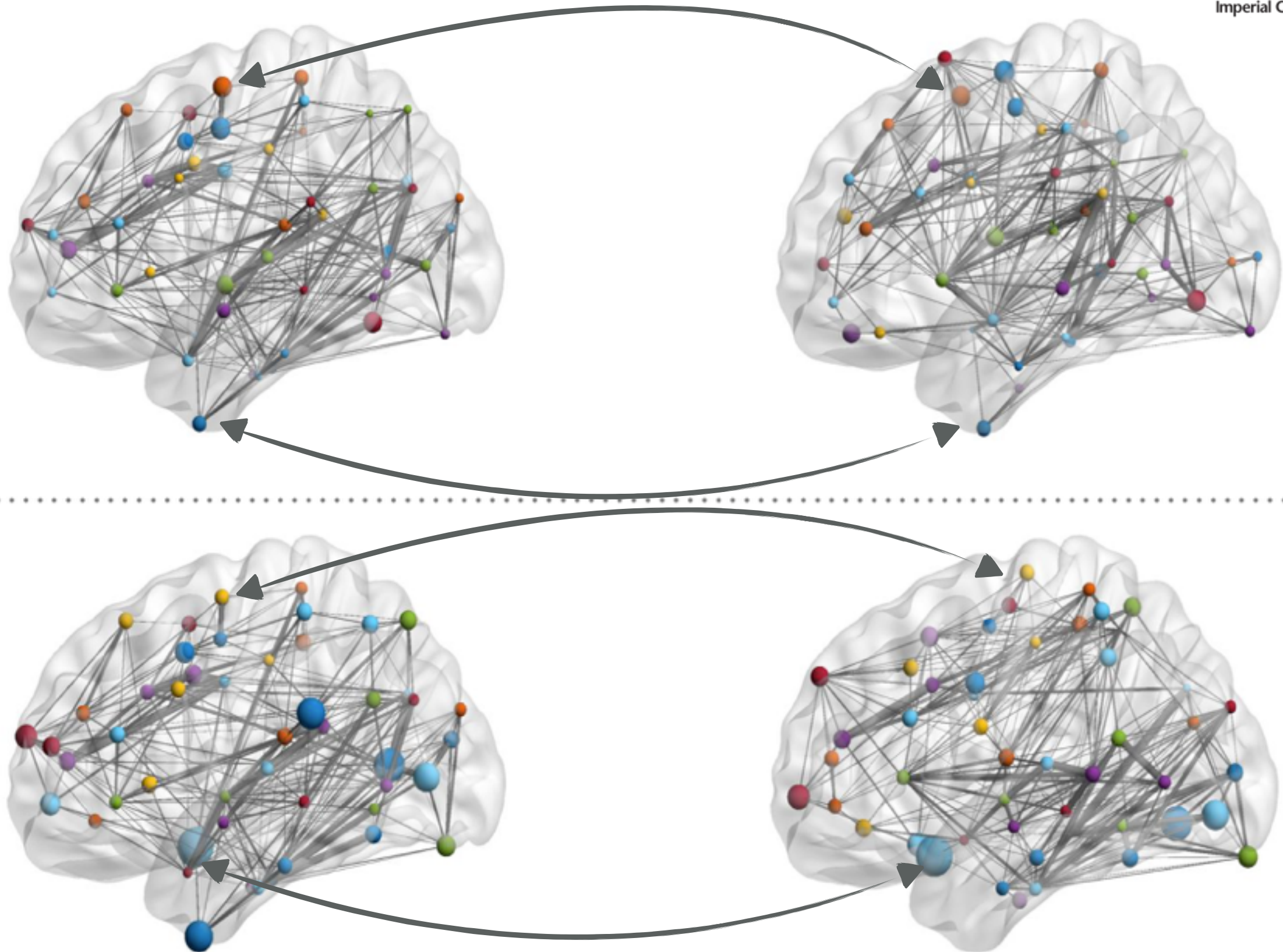


## Functional networks for DZ twins



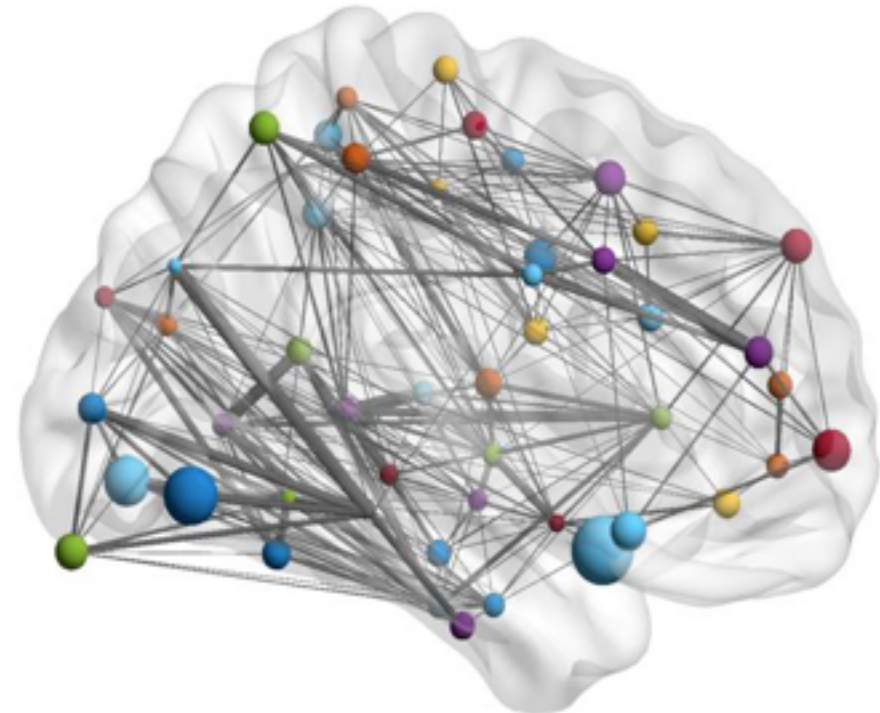
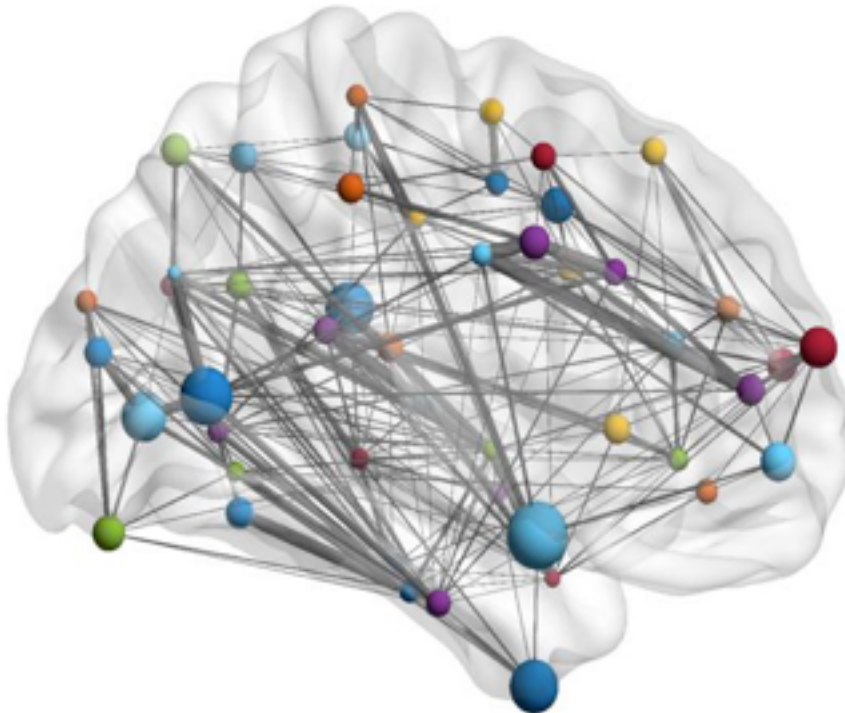
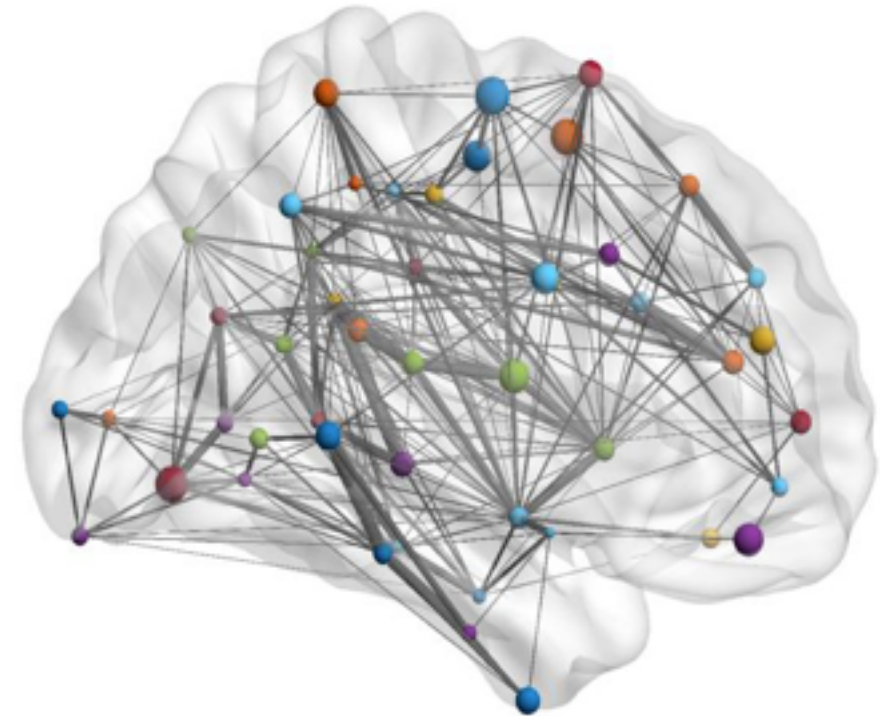
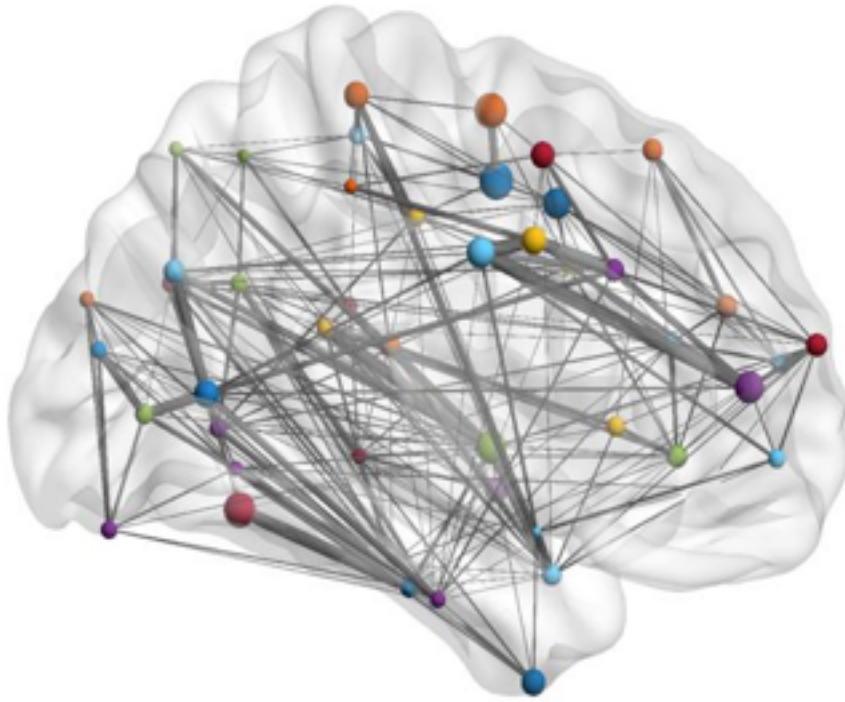
\*  $p < 0.05$   
 \*\*  $p < 0.001$   
 ns non-significant

# Monozygotic vs. unrelated pair





# Monozygotic vs. unrelated pair



- Novel way of evaluating graph similarity between brain networks based on graph edit distance
- Enforces spatial constraints and incorporates feature information
- Applied on healthy unrelated subjects and twin pairs and was able to reflect similarities between corresponding networks
- Future steps:
  - ▶ Predicting phenotype using GED distance matrix
  - ▶ Network dynamics (brain development, disease progression, brain plasticity)

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