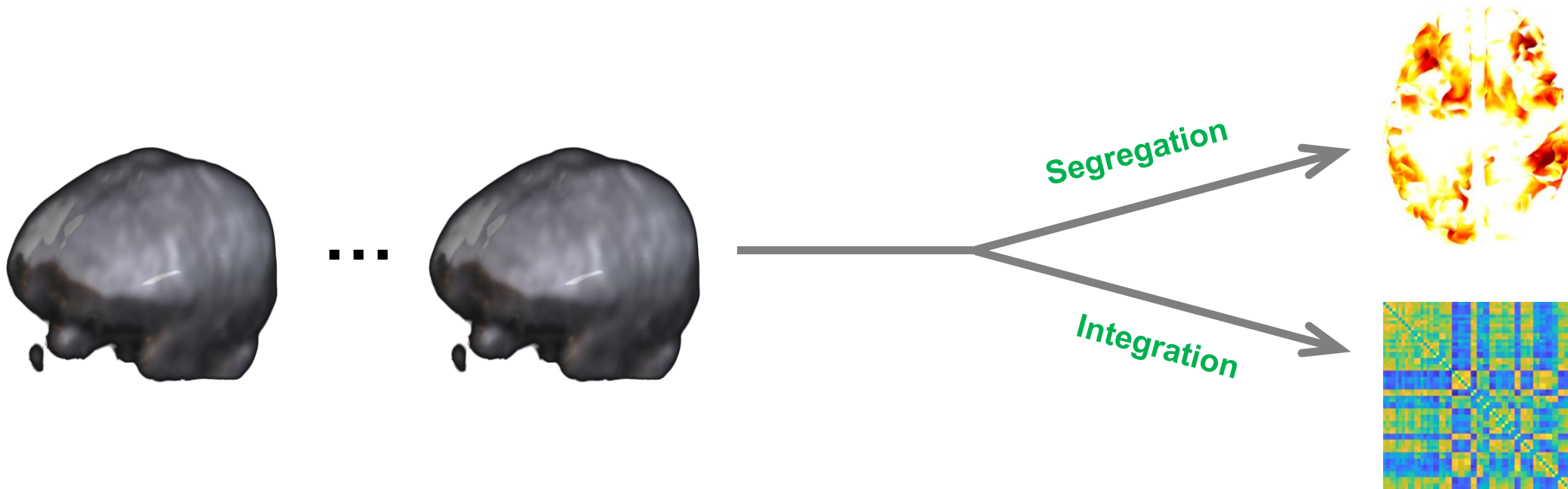


Functional MRI (2): Data Processing

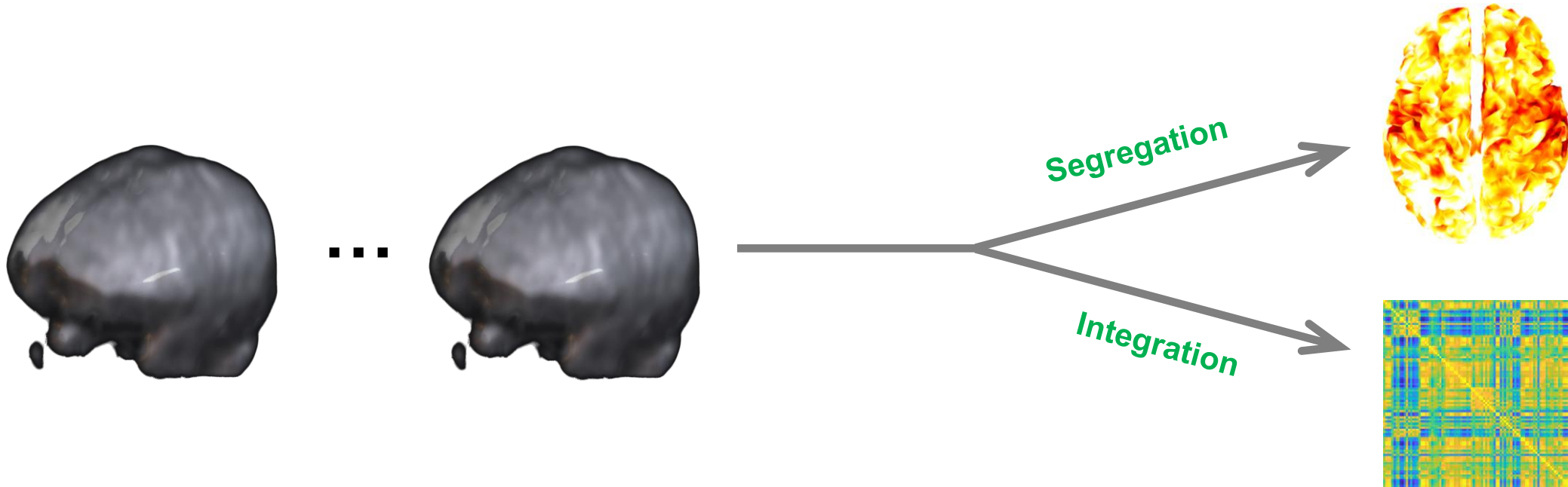
기능 자기공명영상 (2):
데이터 처리 방법

Brain Mapping with functional MRI (fMRI)

- Task-based fMRI

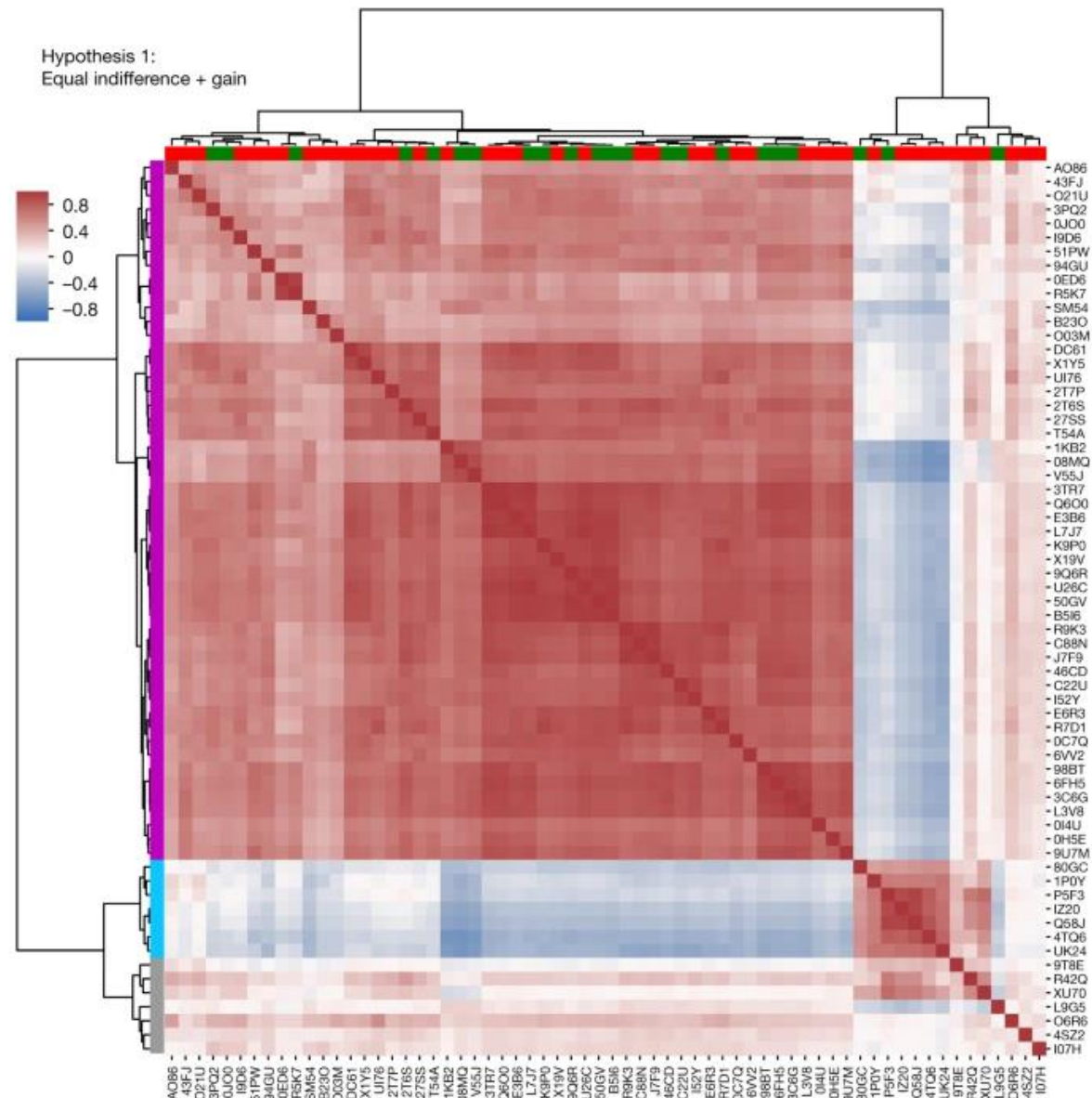


- Resting state fMRI



Analytical Variability in fMRI

- Variability of reported results [\[Botvinik-Nezer et al, 2020\]](#)
 - Resulted from different analysis pipelines
 - As strong factors, spatial smoothness, software package used, and methods of multiple test correction



[Botvinik-Nezer et al, 2020]

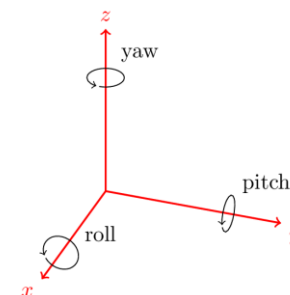
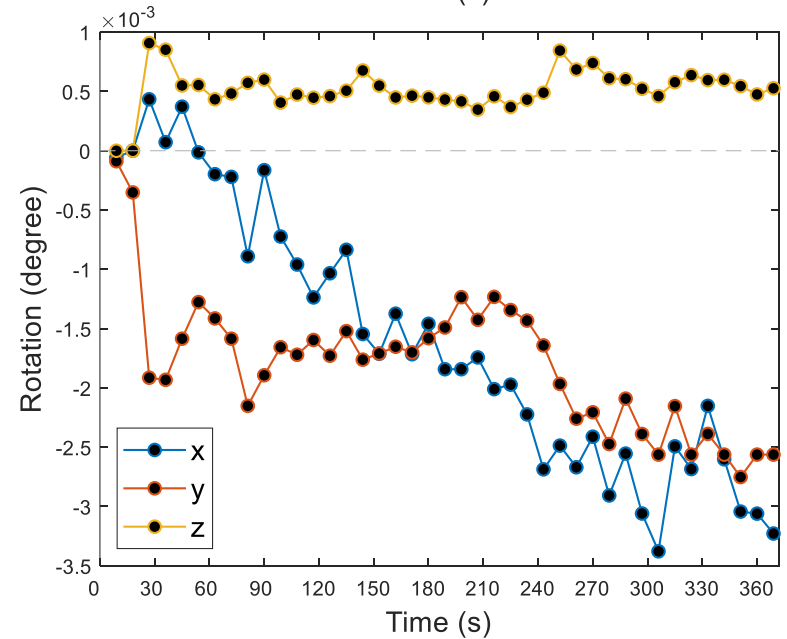
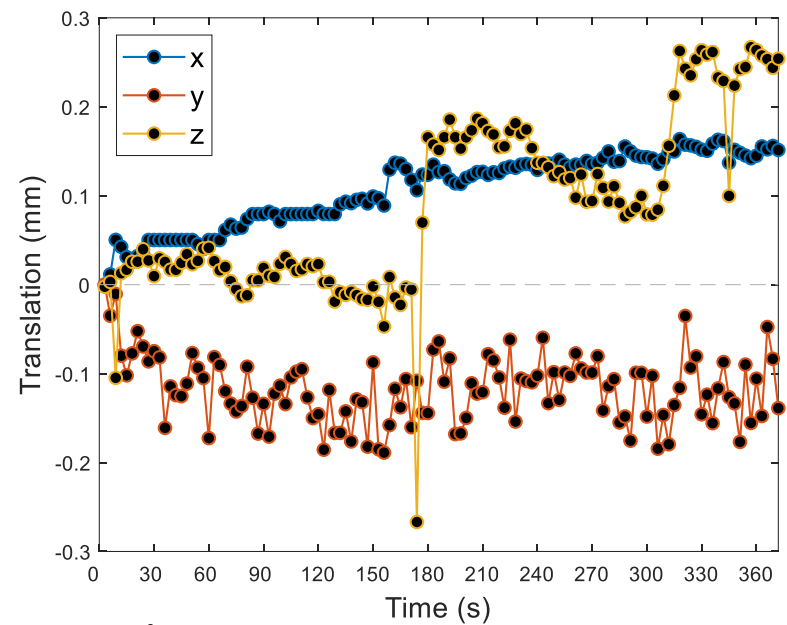
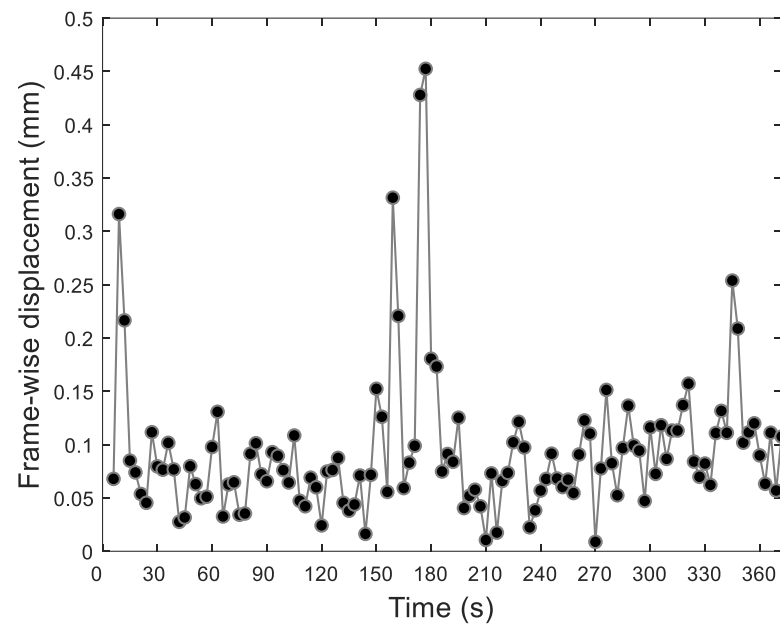
Analytical variability in whole brain statistical results

Preprocessing

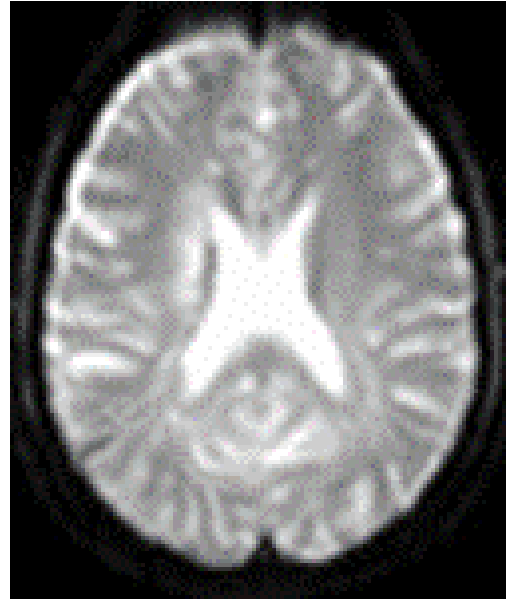
- Numerous steps to clean and standardise fMRI data before modelling and statistical analysis based on the extraction of signals that are faithful to underlying neuronal activity
 - Identifies nuisance (non-neuronal) sources of variability and reduces their effect on fMRI data
 - Addresses particular imaging artifacts and the anatomical localisation of signals

- After preprocessing, it is assumed that fMRI signals are anatomically localised in terms of coordinates
 - Correction for unwanted variation
 - Difference in slice timing
 - Head motion
 - Inhomogeneity (small local inhomogeneities in the magnetic field based on differences in magnetic susceptibility between adjacent tissues or materials (such as air and brain tissue))-induced distortion
 - Normalisation
 - Smoothing

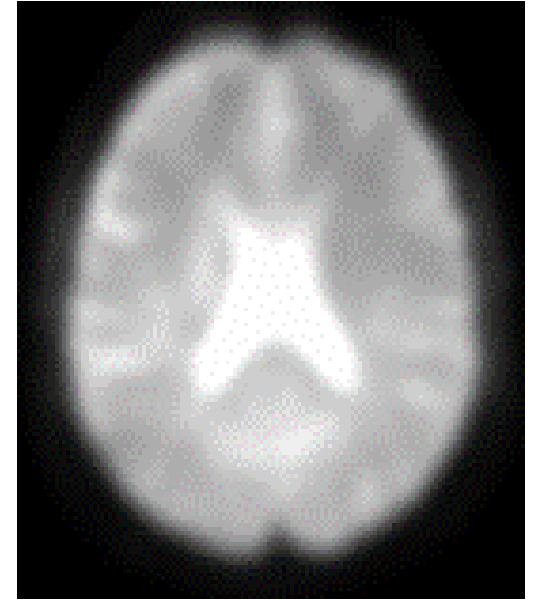
Frame-wise displacement



Estimated head motion



Normalisation

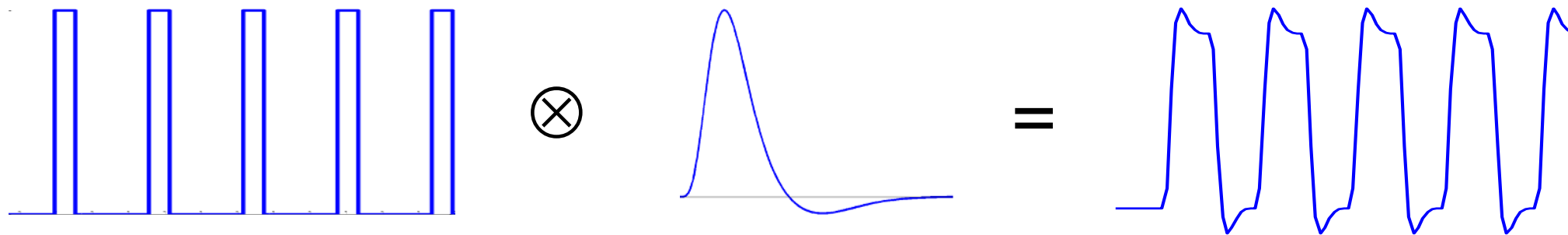


Smoothing

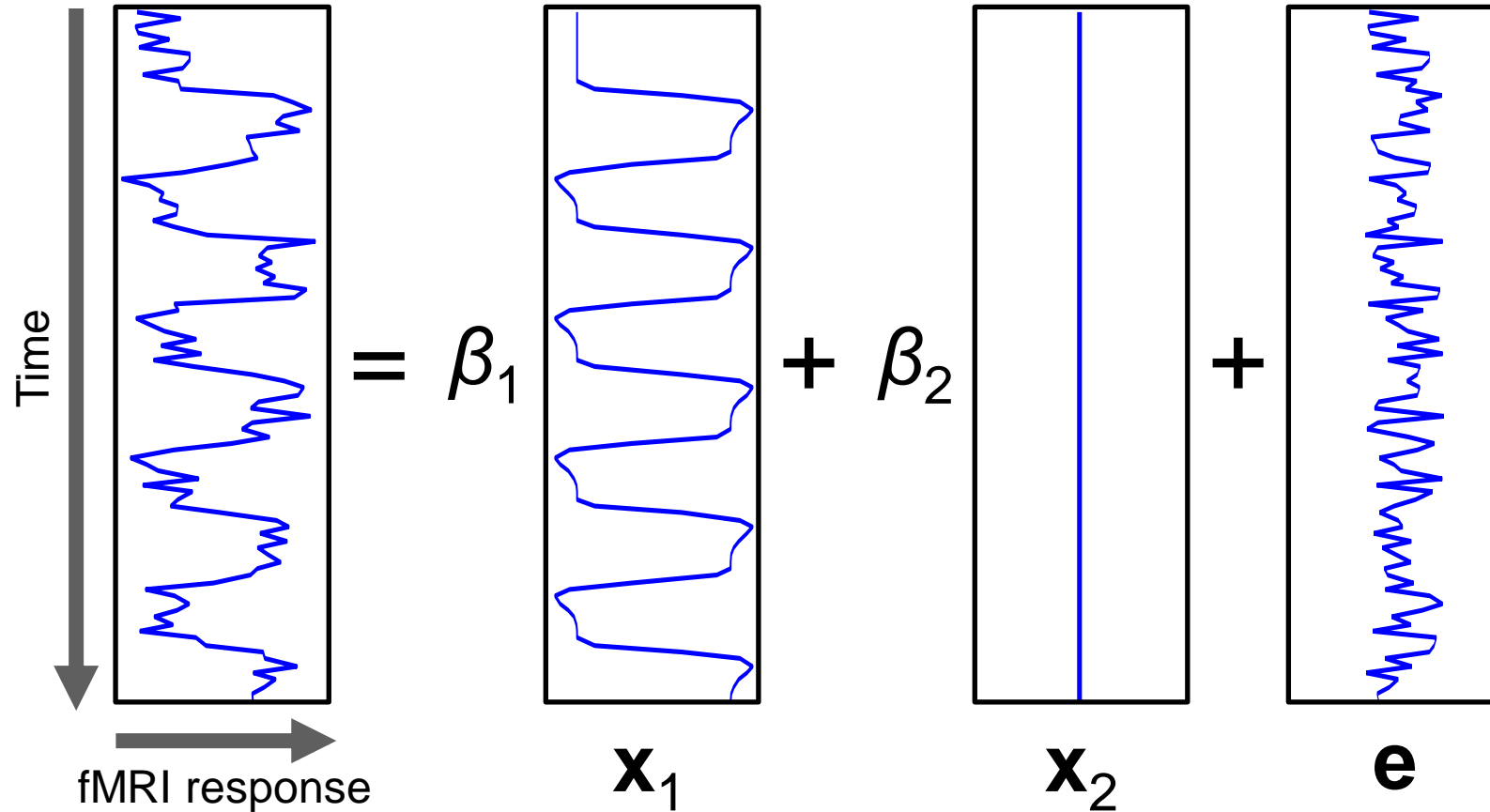
Normalisation and smoothing

Task-based fMRI: Segregation Analysis

- Mass univariate statistical analysis
- General linear model: $\mathbf{y} = \mathbf{X}\boldsymbol{\beta} + \mathbf{e}$
 - Observed fMRI time series \sim predicted fMRI time series + nuisance variables + error
 - \mathbf{y} : observed fMRI time series
 - \mathbf{X} : design matrix
 - $\boldsymbol{\beta}$: parameter estimate
 - \mathbf{e} : error



Predicting fMRI time series

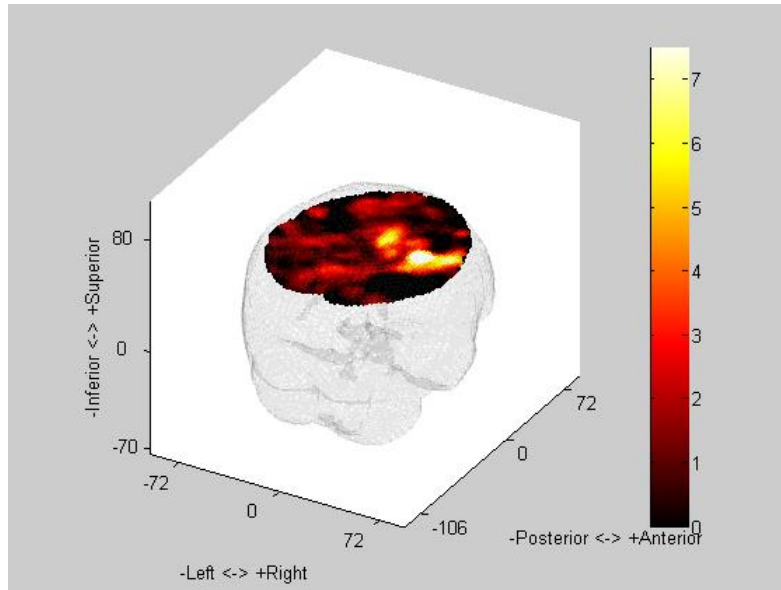


$$y = X\beta + e = x_1\beta_1 + x_2\beta_2 + e$$

General linear model for functional segregation analysis in task-based fMRI

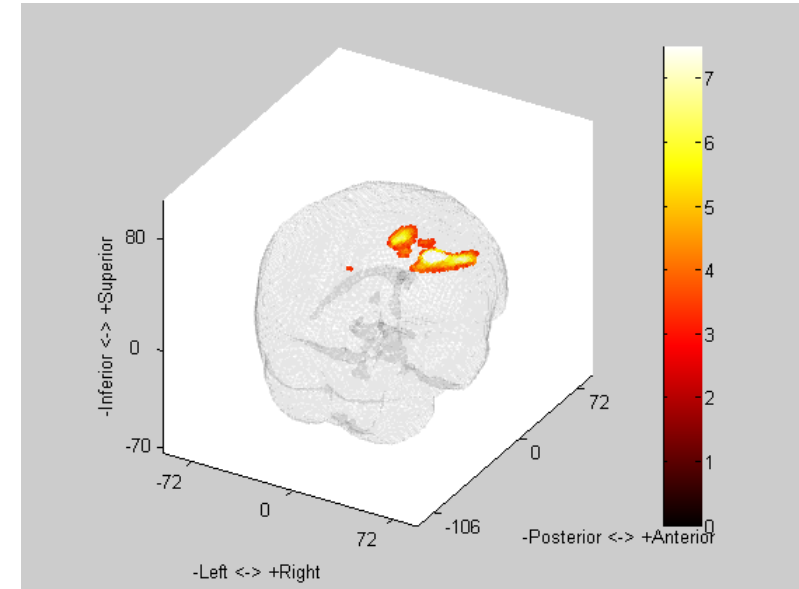
- Search of local brain activity
 - By statistical inferences under the null-hypothesis that predicted fMRI time series are no closer to observed fMRI time series than expected by chance
 - Requires correction for multiple tests

t map



→
Thresholding
at a significance level

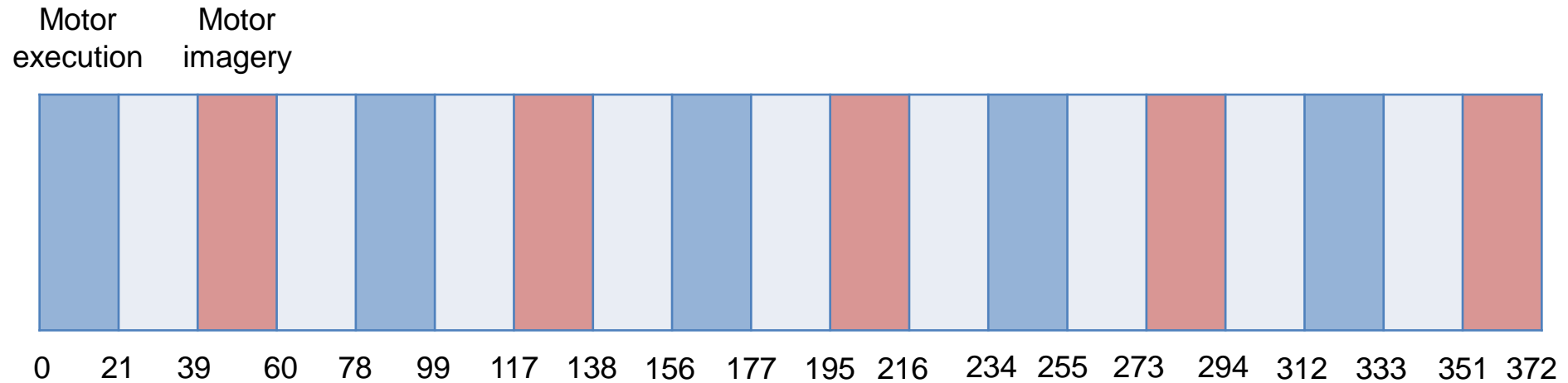
Thresholded t map



$$t = \frac{c^T \hat{\beta}}{\sqrt{\text{var}(e) c^T (X^T X)^{-1} c}}$$

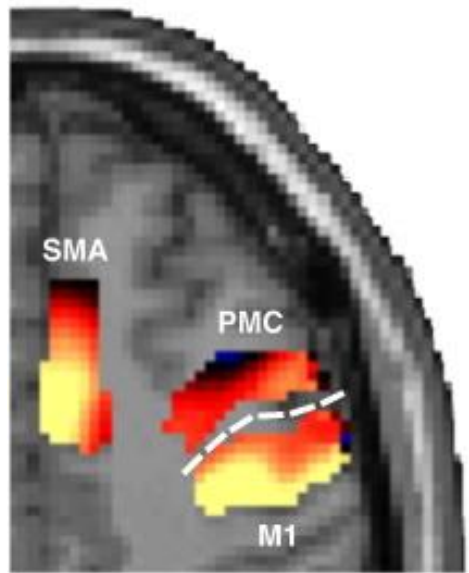
Search for local brain activity by statistical inferences

[Task-based fMRI: Segregation Analysis]

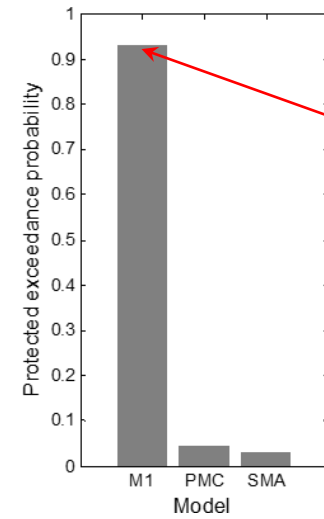
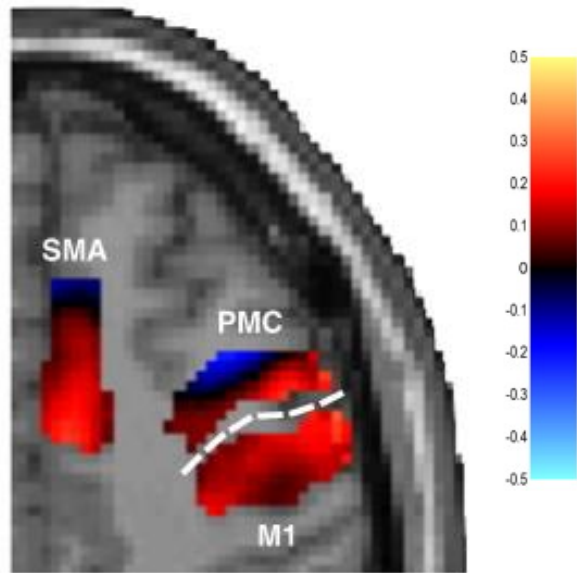


- 124 scans (372 seconds)
- Tasks
 - Motor execution
 - Motor imagery

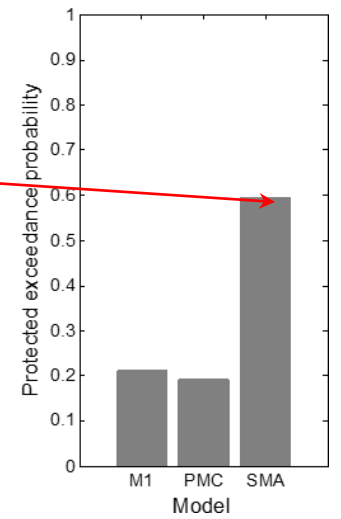
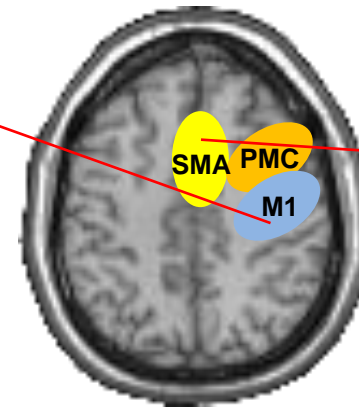
Executed movement of hand grasping



Imagined movement of hand grasping



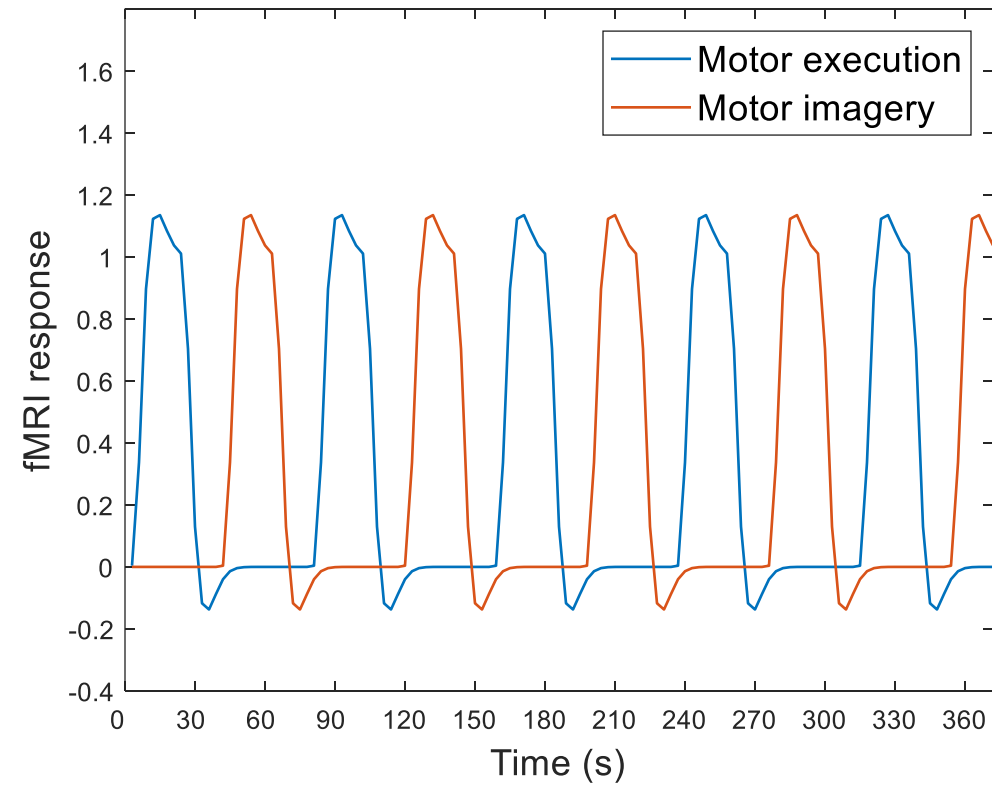
Motor execution



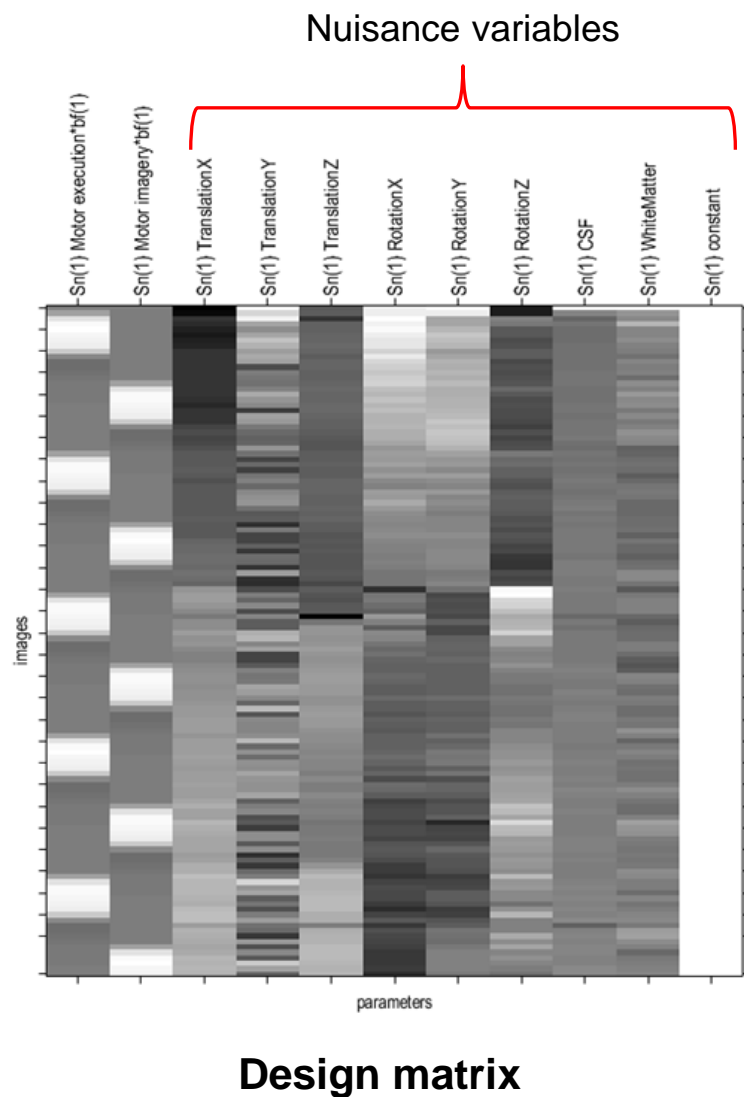
Motor imagery

[Park et al., 2015]

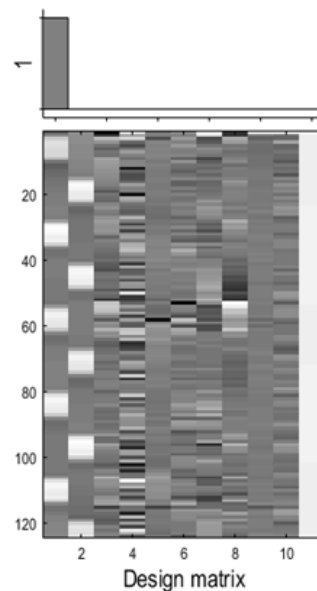
Motor execution vs. motor imagery



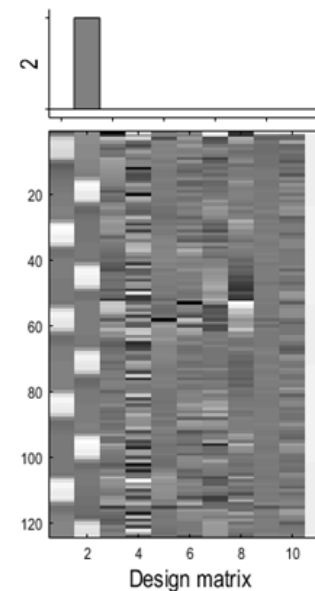
Predicted fMRI time series



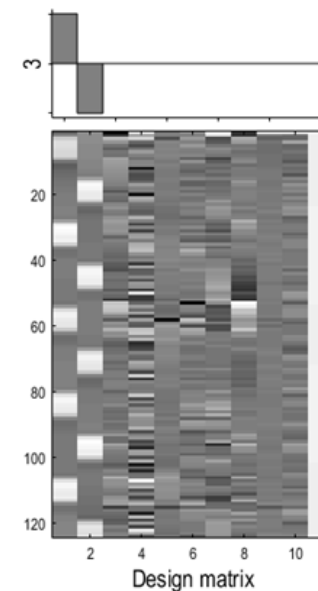
Execution



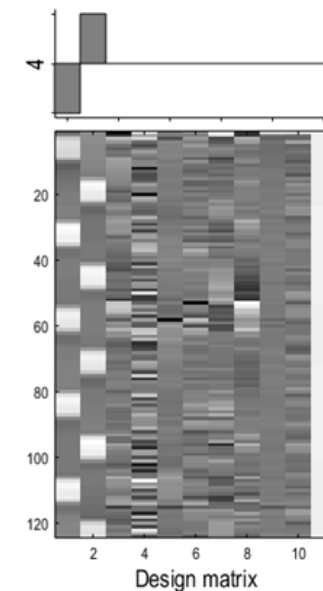
Imagery



Execution > Imagery



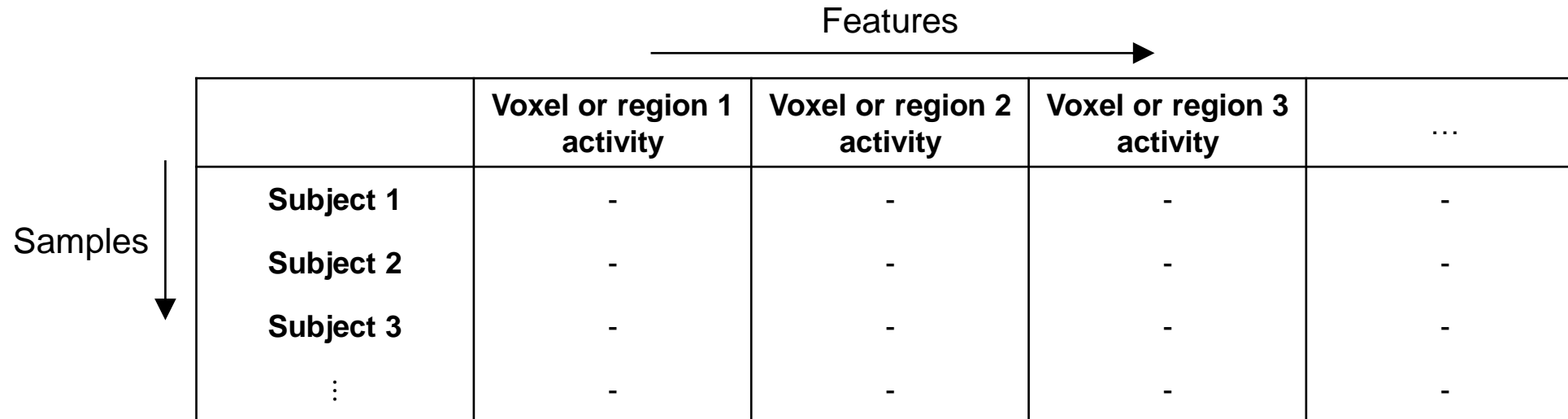
Execution < Imagery



Activity during a task

Activity difference between tasks

- Input to machine learning models
 - Table of voxel-wise or region-wise activity values



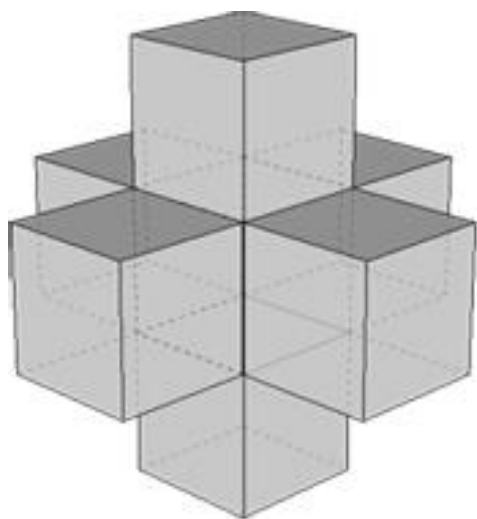
	Features →			
	Voxel or region 1 activity	Voxel or region 2 activity	Voxel or region 3 activity	...
Subject 1	-	-	-	-
Subject 2	-	-	-	-
Subject 3	-	-	-	-
⋮	-	-	-	-

- Brain activity map

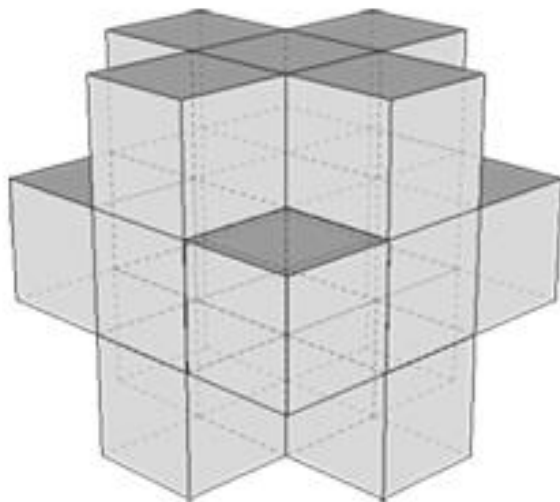
Resting State fMRI: Segregation Analysis

- Regional homogeneity [\[Zang et al., 2004\]](#)
 - Synchronization of time series between a given voxel and its neighbours
 - Neighbours: K nearest neighbours
 - Synchronization: Kendall's coefficient of concordance (KCC)

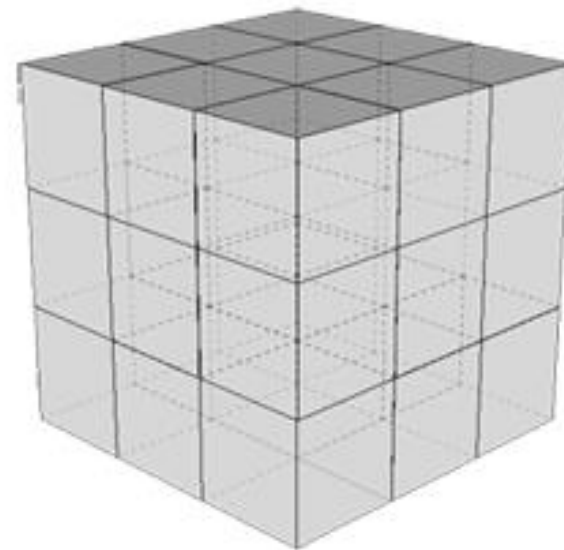
$$\text{KCC} = \frac{\sum_{i=1}^n R_i^2 - n(\bar{R})^2}{\frac{1}{12} K^2 (n^3 - n)} = 12 \frac{\sum_{i=1}^n (\bar{R}_i)^2}{(n^3 - n)} - 3 \frac{(n + 1)}{(n - 1)}$$



Faces
(7 voxels)



Faces + Edges
(19 voxels)

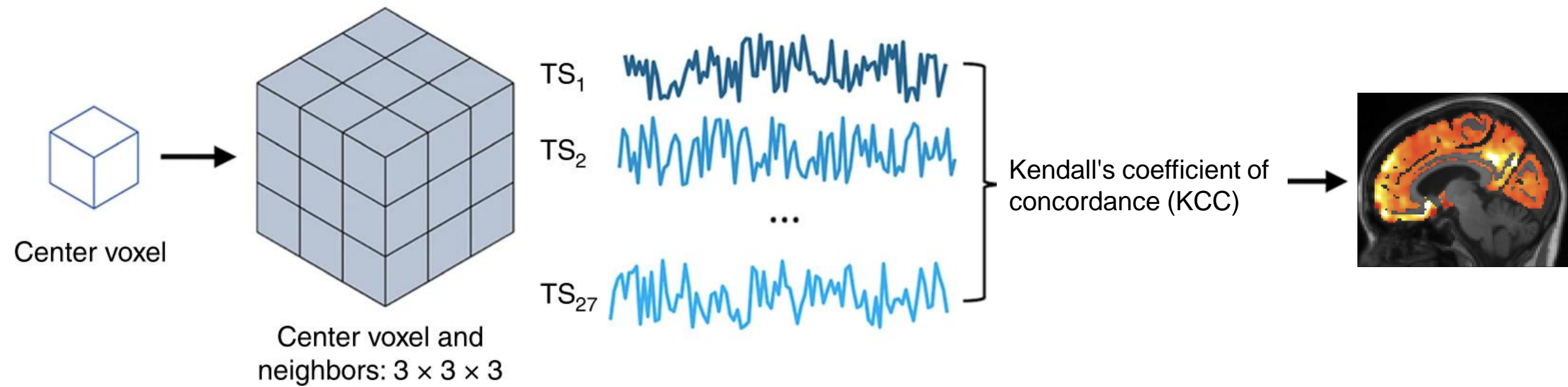


Faces + Edges + Corners
(27 voxels)

[\[https://fcp-indi.github.io/docs/latest/user/reho\]](https://fcp-indi.github.io/docs/latest/user/reho)

Different definitions of nearest neighbours

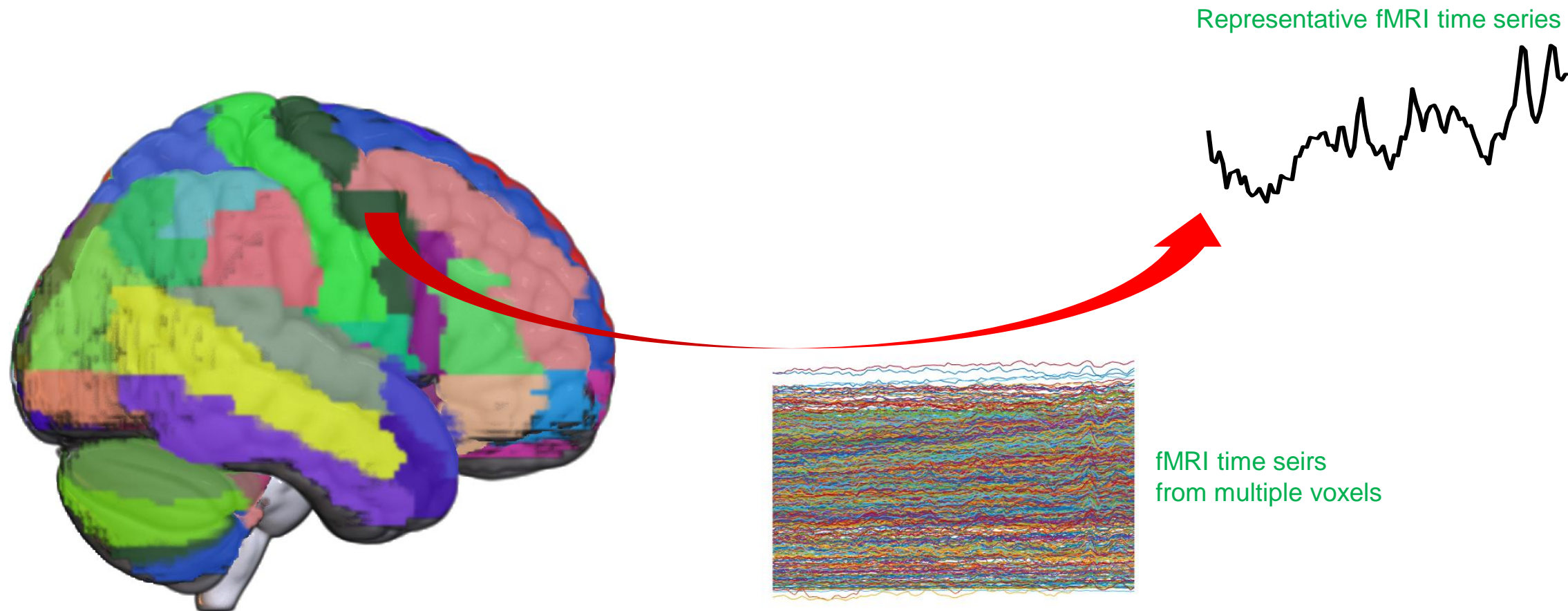
- Based on the hypothesis that intrinsic brain activity is manifested by clusters of voxels rather than single voxels
- Requires no pre-defined voxel or region
- Provides information about the local activity of regions throughout the brain



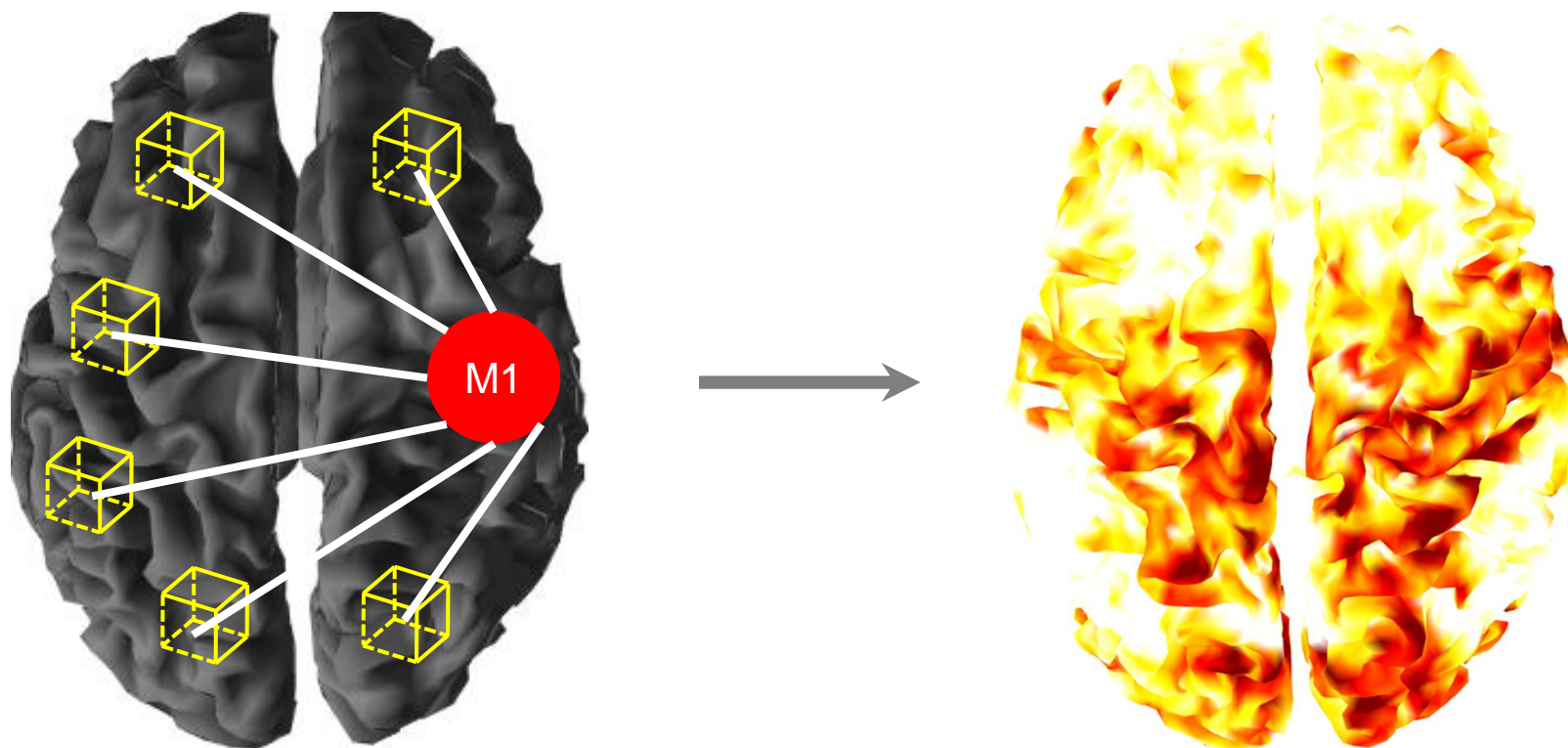
[Harrison et al., 2019]

Regional homogeneity

- Seed-based correlation [\[Biswal et al., 1995\]](#)
 - Synchronization of time series between a seed and all other voxels in the brain
 - Seed: pre-defined voxel or region
 - Synchronization: statistical association, particularly correlation
 - Based on the hypothesis that brain regions with similar activity patterns are likely to be communicating and sharing information
 - Often used to explore a set of brain regions that share similar patterns of activity

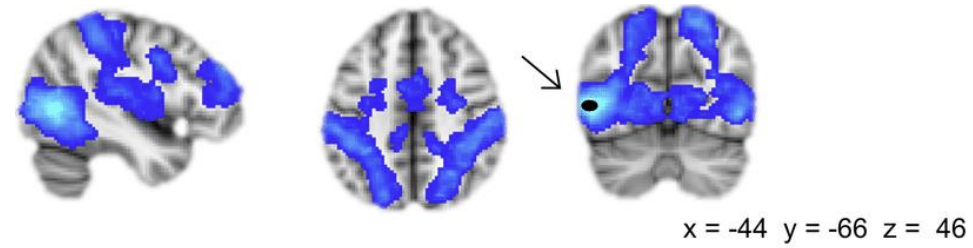


Extracting fMRI time series from the seed (primary motor cortex)

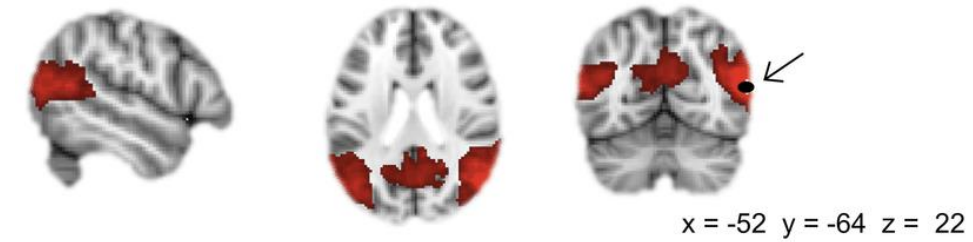


Seed-based correlation for the primary motor cortex

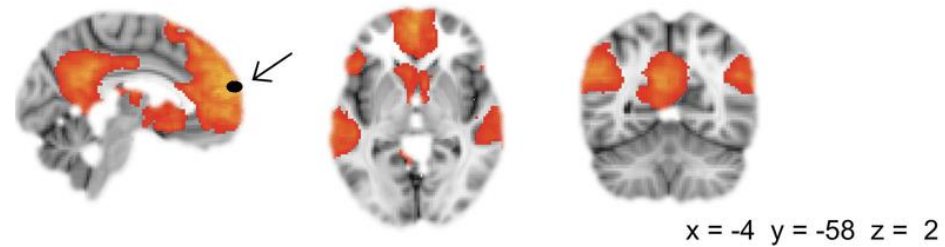
Left middle temporal gyrus



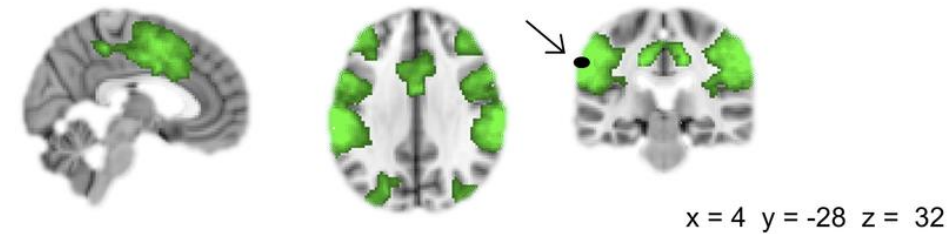
Right middle temporal gyrus



Left medial frontal gyrus



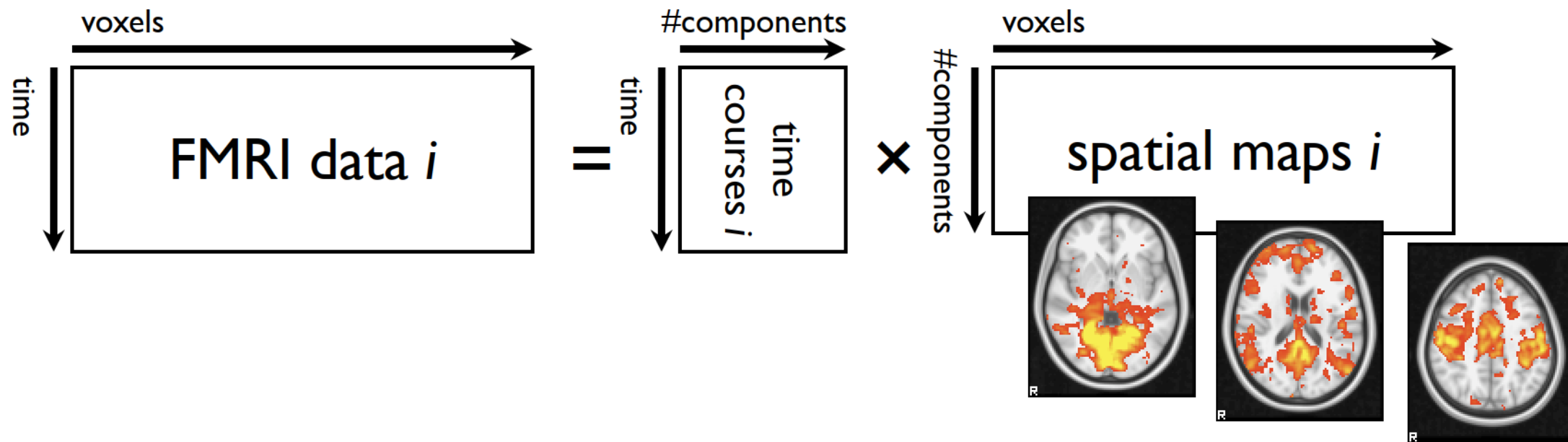
Left supramarginal gyrus

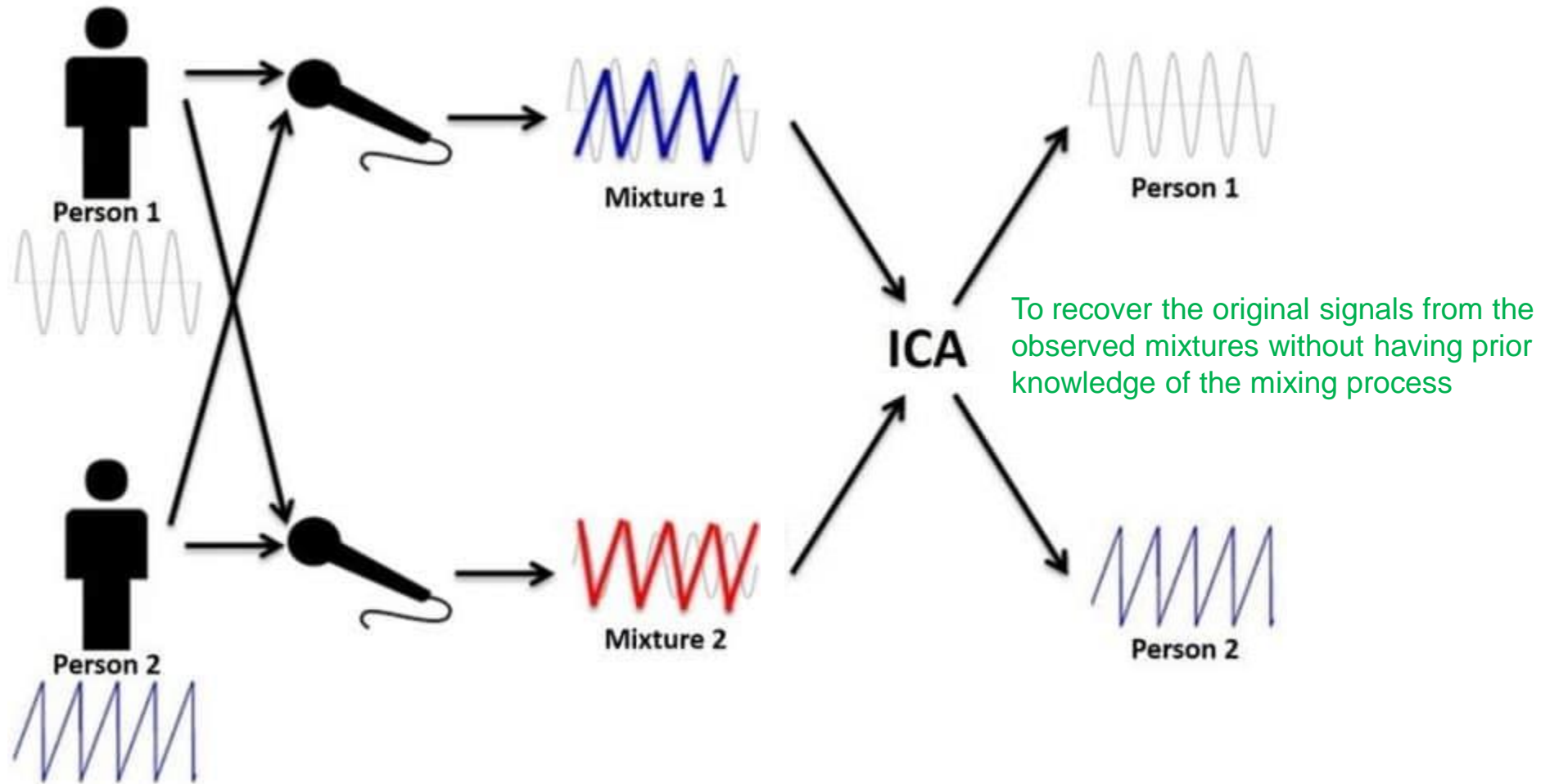


[Cousijn et al., 2014]

Time series synchronization explored by correlation with different seeds

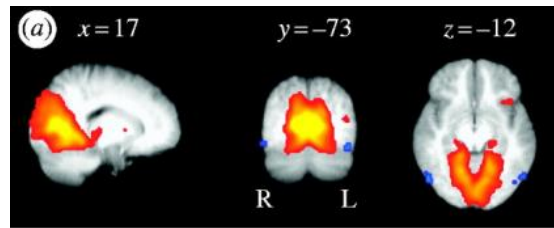
- Independent component
 - Statistical source or factor that independent component analysis (ICA) aims to extract from multivariate data
 - Spatial map and its time course separated from fMRI data
 - Based on the hypothesis that the observed data are linear mixtures of the unknown independent components



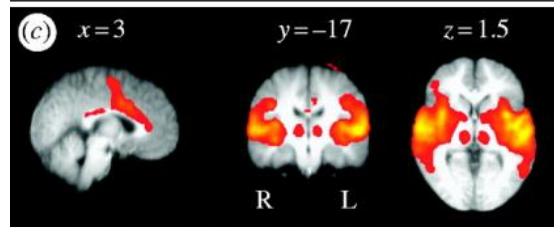


[\[https://vocal.com/blind-signal-separation/independent-component-analysis/\]](https://vocal.com/blind-signal-separation/independent-component-analysis/)

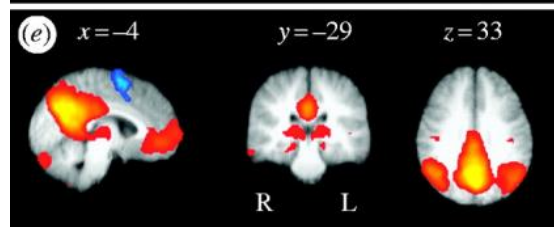
Independent component analysis for the cocktail party problem



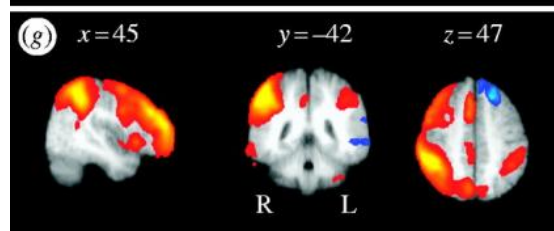
Medial visual
cortical areas



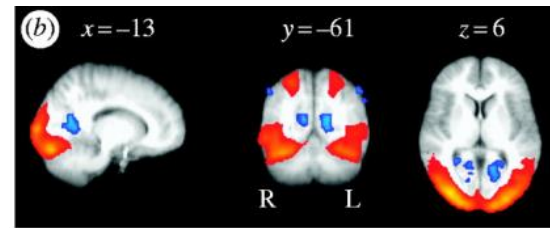
Auditory system



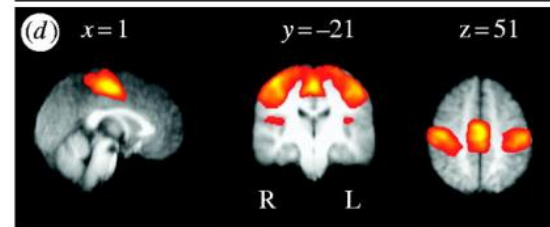
Visuo-spatial
system



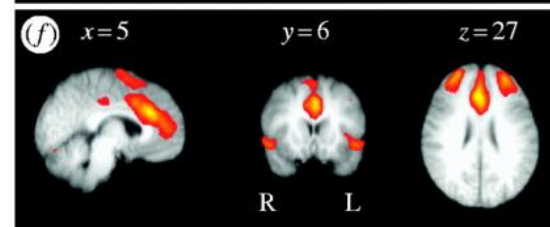
Right dorsal
visual stream



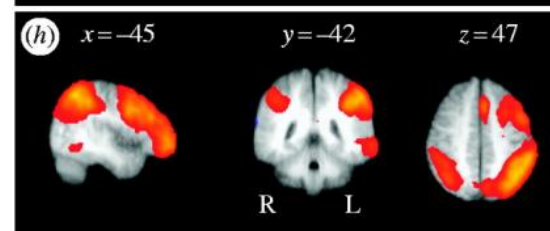
Lateral visual
cortical areas



Sensory-motor
system



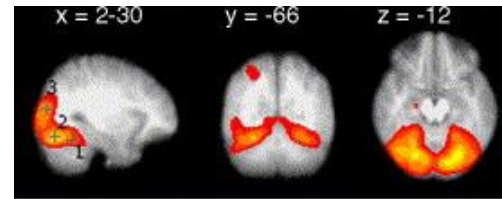
Executive control



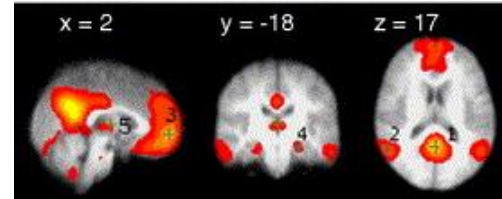
Left dorsal
visual stream

[Beckmann et al., 2005]

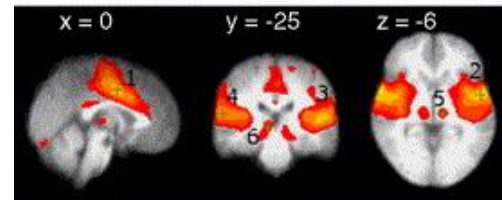
Time series synchronization explored by independent component analysis (1)



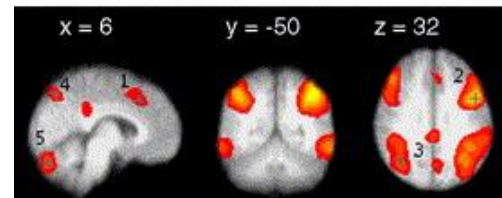
Visual cortical areas



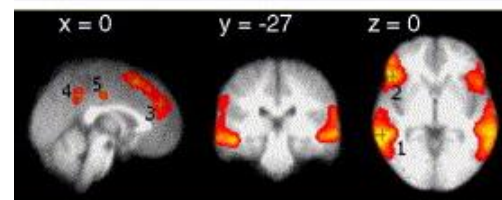
Visuospatial and executive system



Sensory and auditory system



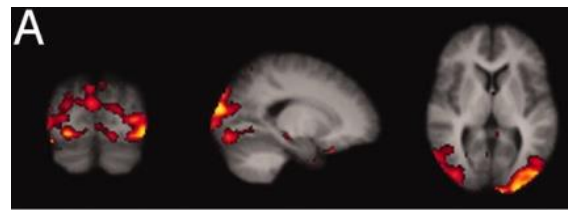
Dorsal pathway



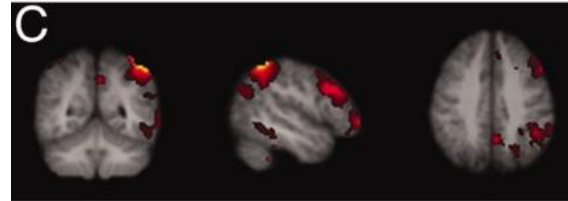
Ventral pathway

[De Luca et al., 2006]

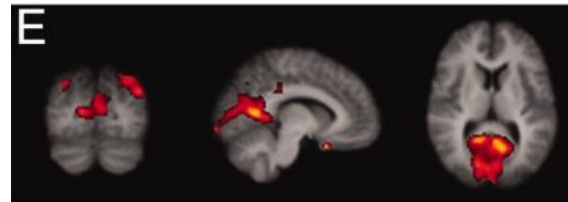
Time series synchronization explored by independent component analysis (2)



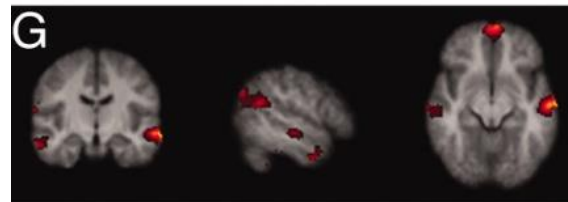
Lateral visual areas



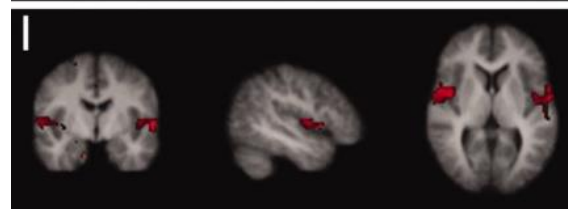
Memory function (left)



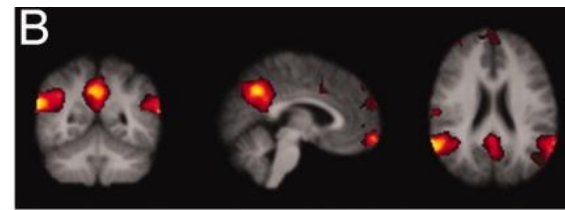
Medial visual areas



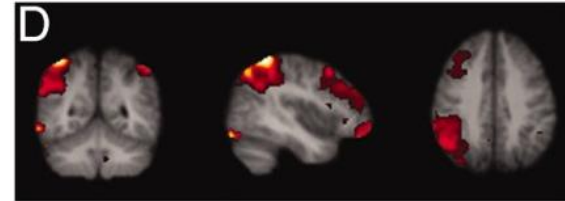
Occipitotemporal pathway (ventral stream)



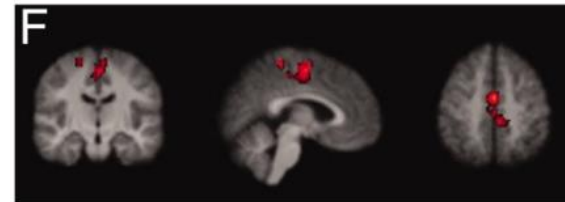
Auditory cortex



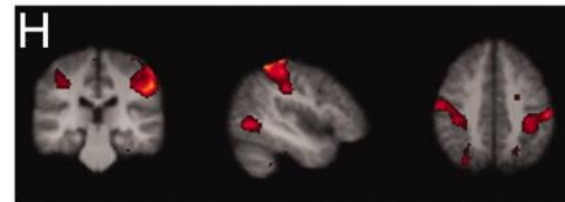
Default-mode network



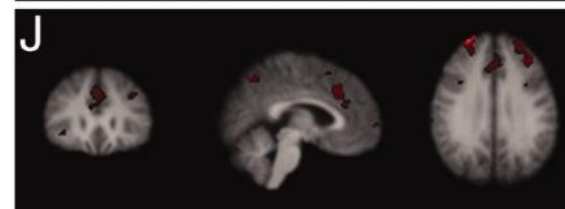
Memory function (right)



Motor and sensory network



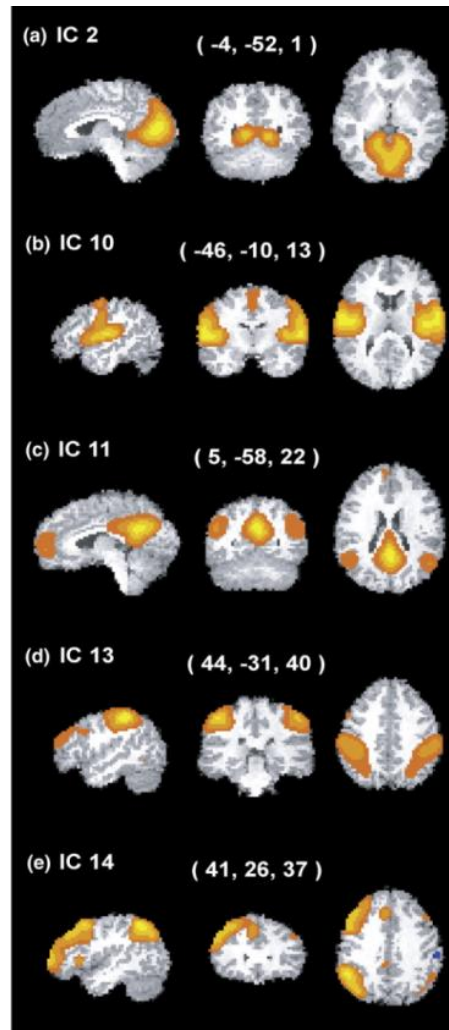
Superior parietal cortex



Executive control and working memory function

[Damoiseaux et al., 2006]

Time series synchronization explored by independent component analysis (3)



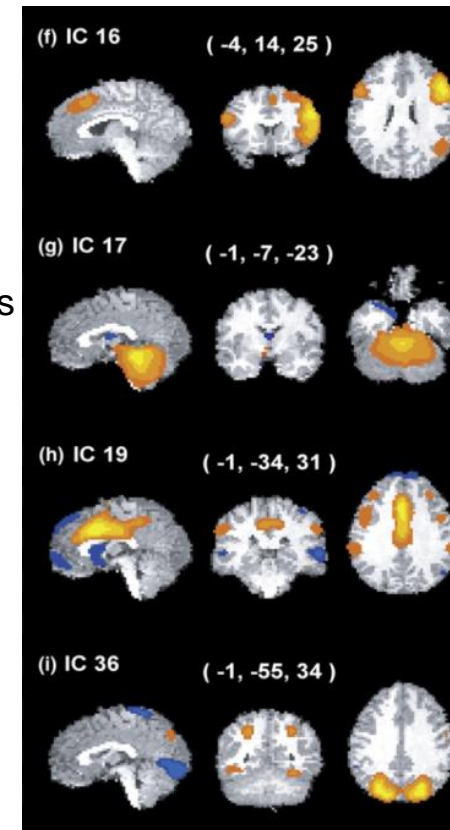
Medial occipital cortices

Bilateral temporal cortices and ACC

PCC, ACC, and bilateral inferior parietal cortex (default-mode network)

Bilateral motor regions

Right medial and lateral frontal cortices and parietal cortex



Left medial and lateral frontal cortices and parietal cortex

Cerebellum

Medial frontal/ACC, bilateral DLPFC, bilateral temporal cortex and striatum

Bilateral lateral occipital cortices

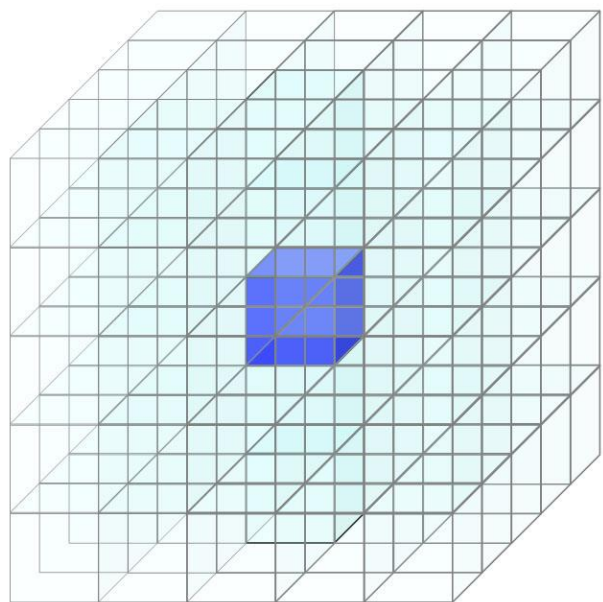
[Chen et al., 2008]

Time series synchronization explored by independent component analysis (4)

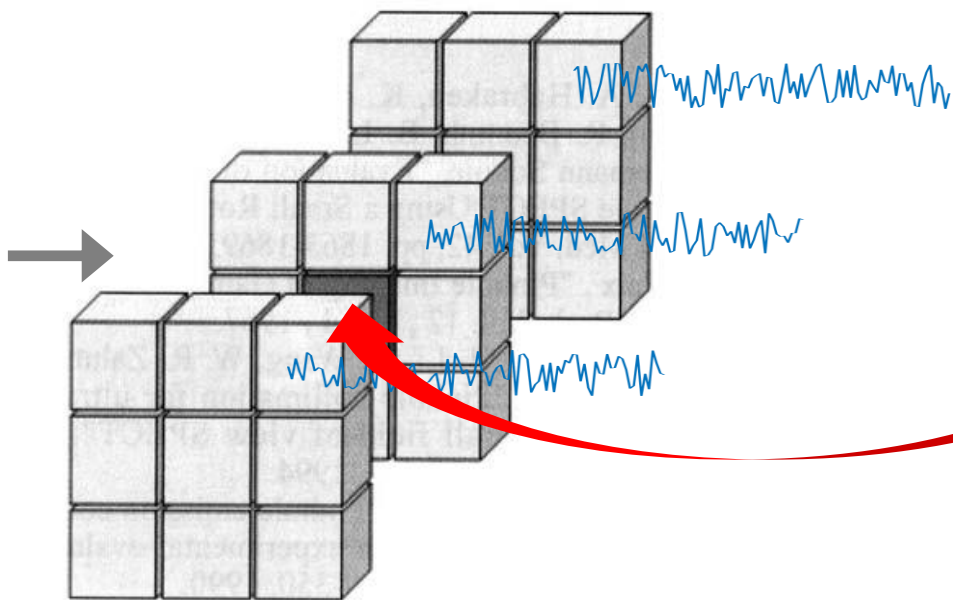
[Resting State fMRI: Segregation Analysis]



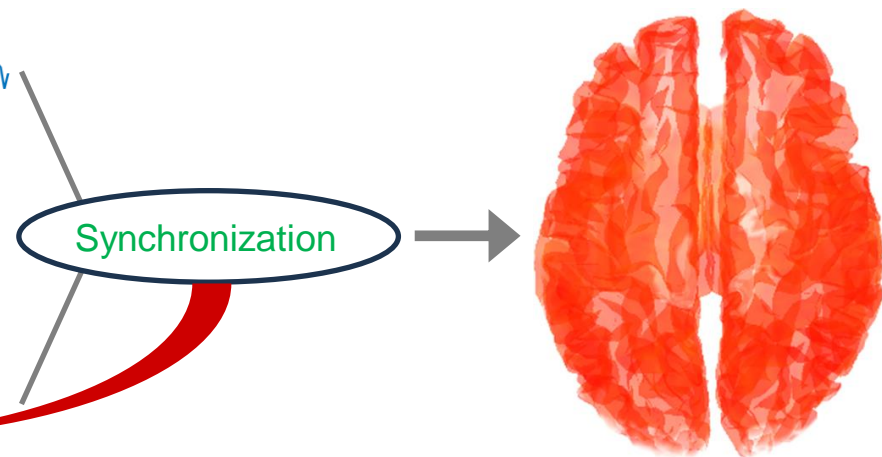
– 100 scans (200 seconds)



Centre voxel

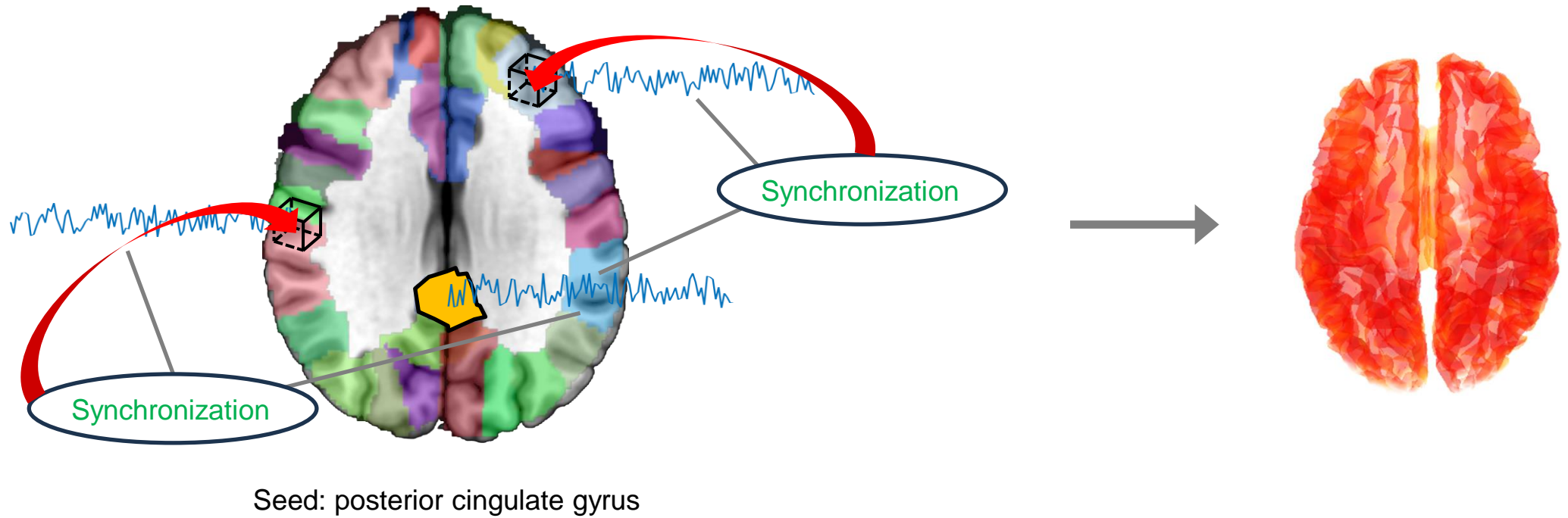


Nearest neighbours



Synchronization

Regional homogeneity

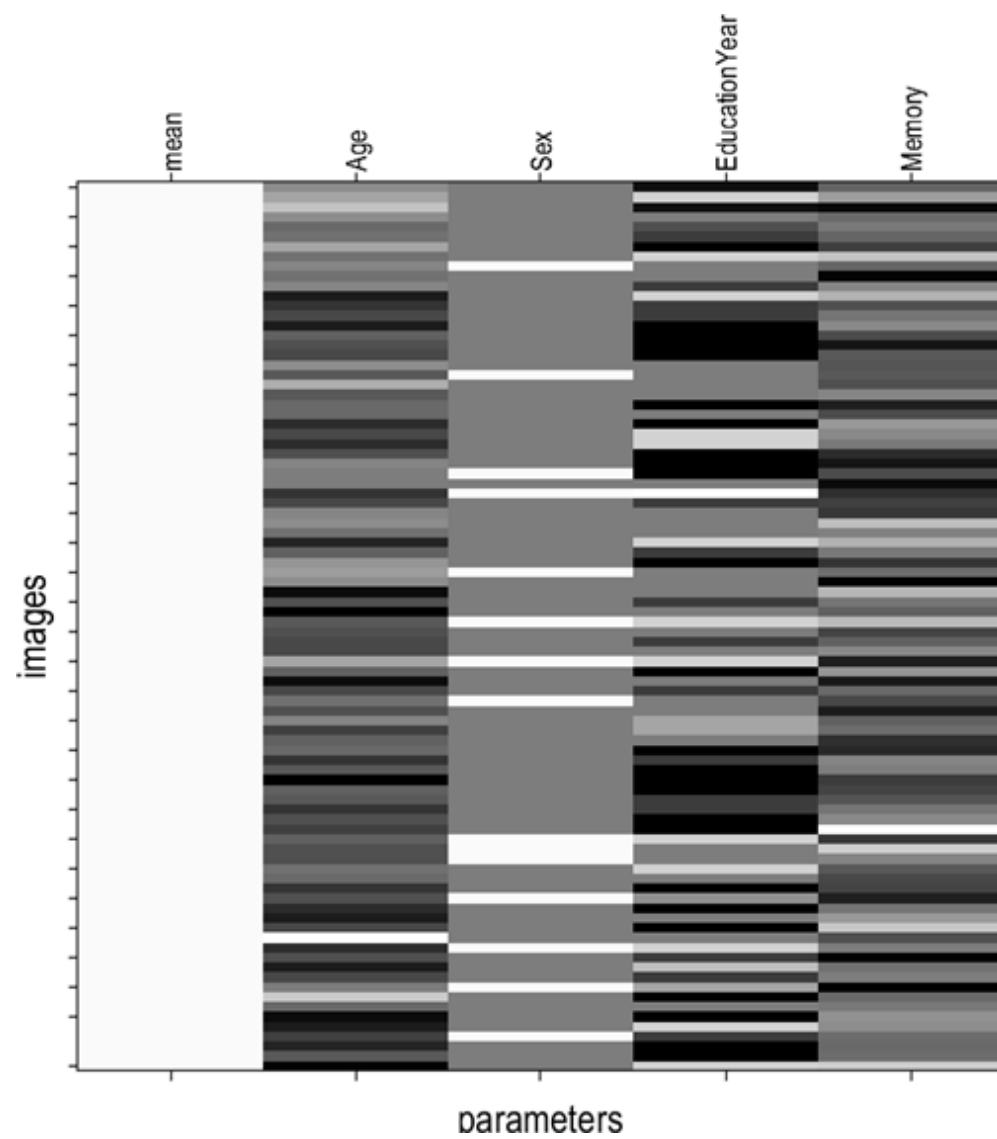


Correlation with the posterior cingulate gyrus: default mode network

[Statistical Analysis of Resting State fMRI]

- Regional homogeneity \sim
Age +
Sex +
Education year +
Memory performance

Design matrix



Output

Regression



Positive correlaton



Negative correlation

- Correlation with the posterior cingulate gyrus ~
Age +
Sex +
Education year +
Memory performance

Output

Regression



Positive correlaton



Negative correlation

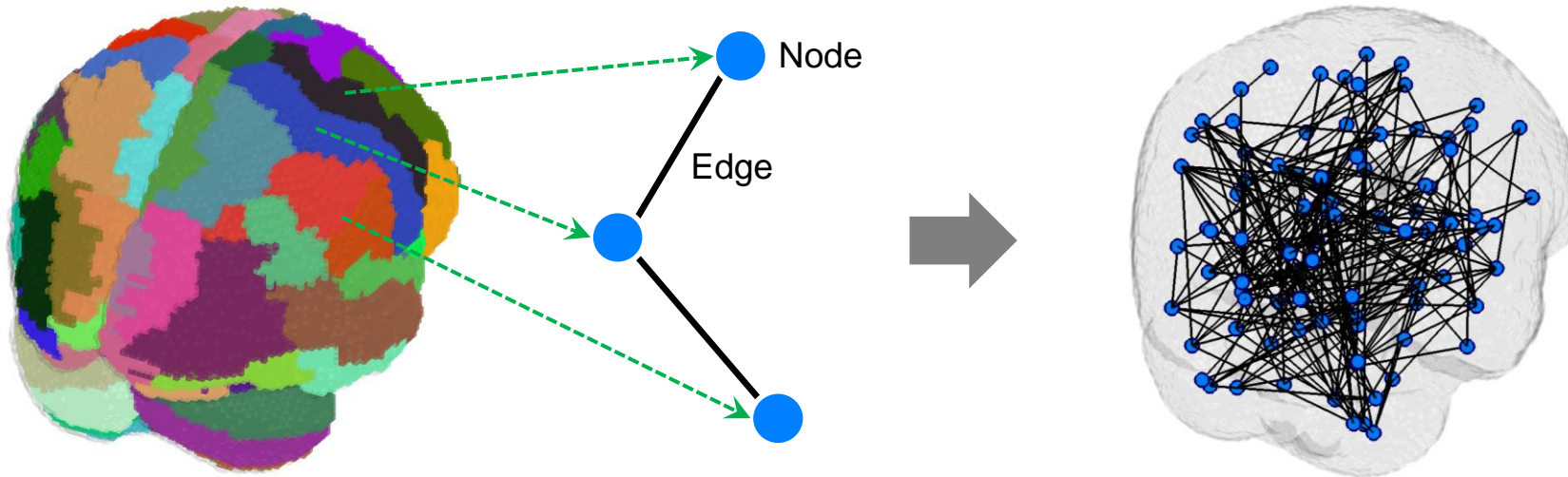
- Input to machine learning models
 - Table of voxel-wise or region-wise synchronization (regional homogeneity, seed-based correlation, or independent component) values

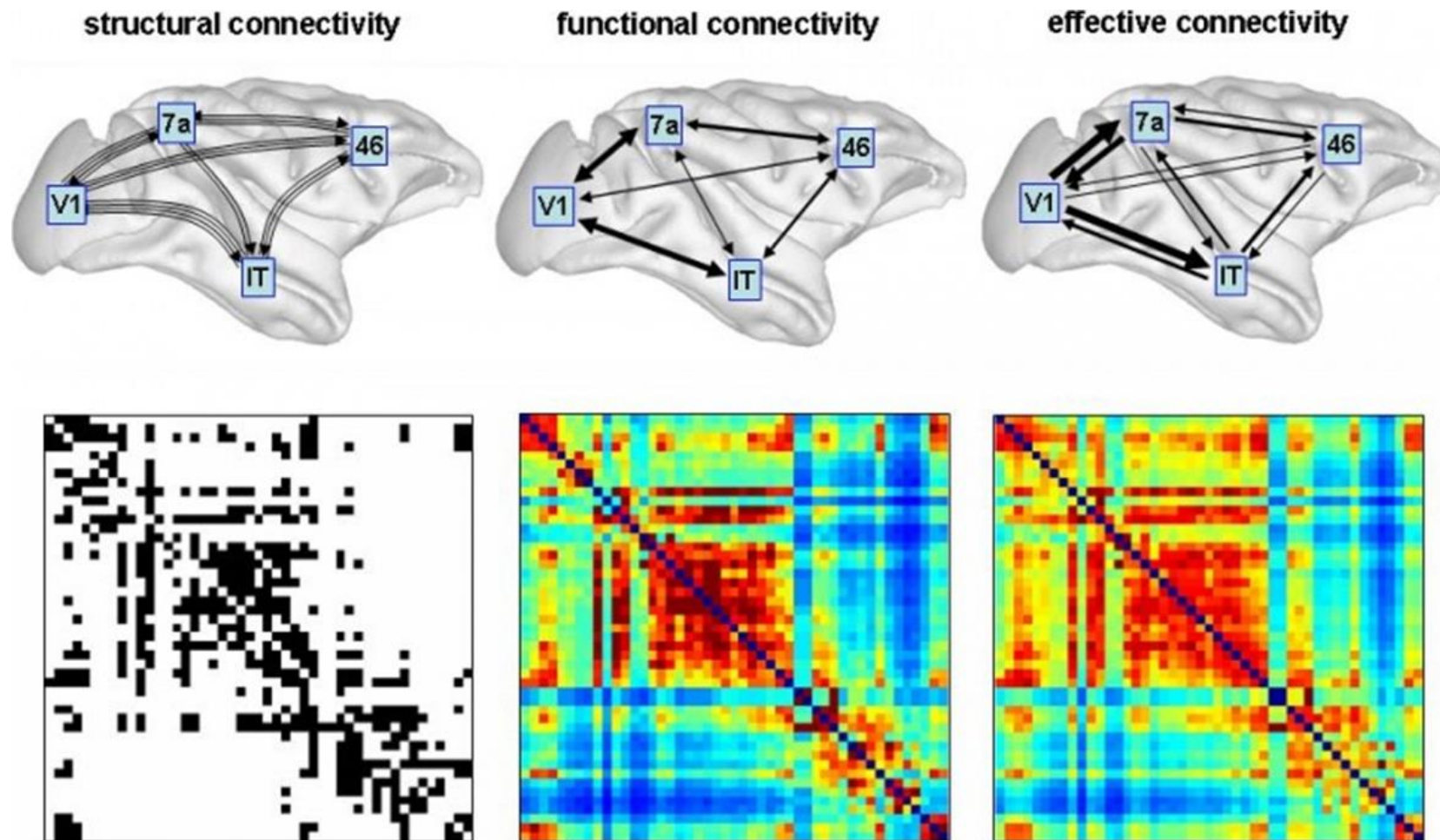
Samples ↓		Features →			
		Voxel or region 1 synchronization	Voxel or region 2 synchronization	Voxel or region 3 synchronization	...
	Subject 1	-	-	-	-
	Subject 2	-	-	-	-
	Subject 3	-	-	-	-
	⋮	-	-	-	-

- Time series synchronization (regional homogeneity, seed-based correlation, or independent component) map

Resting State fMRI: Integration Analysis

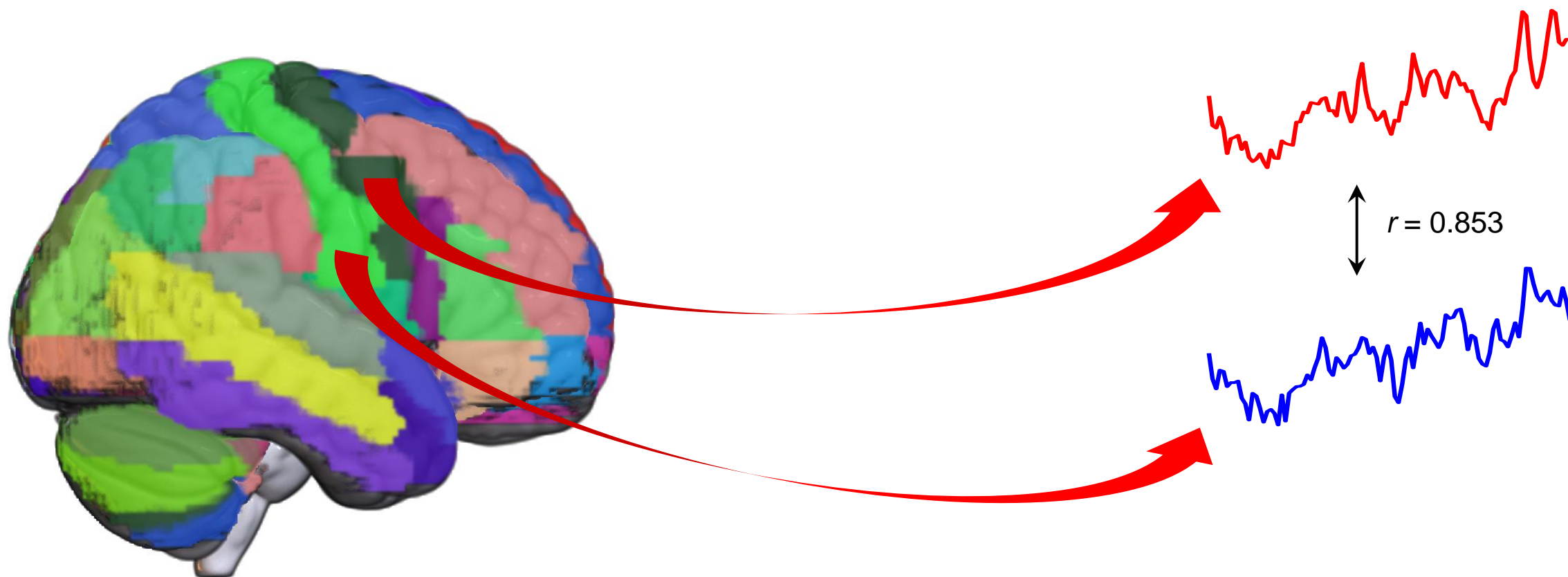
- Network
 - Set of nodes and edges
 - Nodes: pre-defined regions
 - Edges: connectivity (correlation or causality) between regions



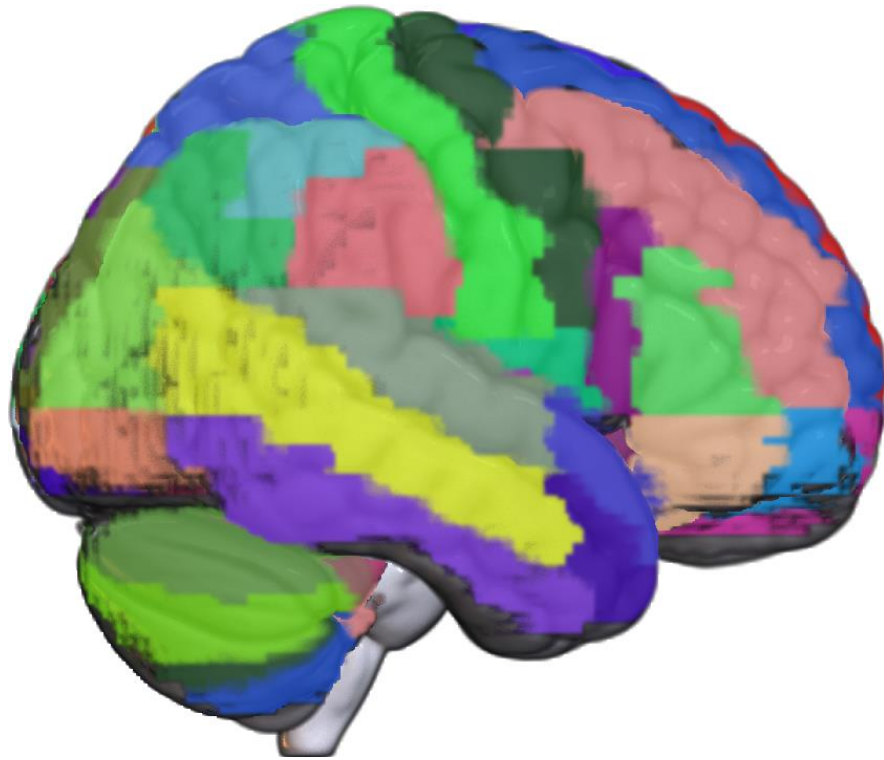


[Honey et al., 2007]

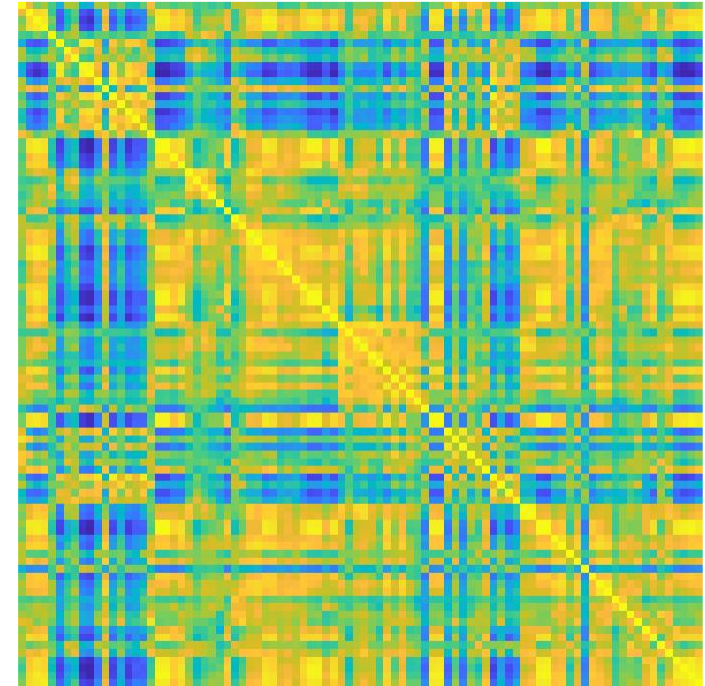
Modes of brain connectivity



Pair-wise correlation of time series



Time series correlation



Functional network or connectome

- Input to machine learning models
 - Table of region-to-region connectivity (correlation or causality) values

		Features			
		Brain regions 1 – 2 connectivity	Brain regions 1 – 3 connectivity	Brain regions 1 – 4 connectivity	...
Samples	Subject 1	-	-	-	-
	Subject 2	-	-	-	-
	Subject 3	-	-	-	-
	⋮	-	-	-	-

- Functional network map