

# Connectivity

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AARHUS UNIVERSITY



IMC  
INTERACTING MINDS CENTRE



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# Comments on paper\_1

# Paper\_1 comments

7 pages of 2400 characters (including spaces and references) = 16.800 characters

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- This includes everything:
  - ▶ Abstract
  - ▶ Main text
  - ▶ Footnote/endnotes
  - ▶ etc.

# Paper\_1 comments

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From lecture\_1:

- A selection of three papers with an introduction and discussion/conclusion is to be handed in as one joint submission.
- A paper can be maximum 7 normal pages
  - ▶ code goes in an appendix
  - ▶ Figures does not count.
- Introduction and discussion/conclusion is combined maximum of 7 normal pages.
- A normal page is 2400 characters *including spaces and in-text references*.
- The reference list does *not* count for the pages limits.
- Citation style is APA7

# Connectivity

# What is connectivity?

## Connectivity

The study of connections between brain sites

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Two main questions:

# What is connectivity?

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- What is a brain site?

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The study of connections between brain sites

Two main questions:

- What is a brain site?
- What is a connection?

# What is connectivity?

## Connectivity

The study of connections between brain sites

Two main questions:

- What is a brain site?
  - ▶ Anatomically defined
  - ▶ Functionally defined
- What is a connection?

# What is connectivity?

## Connectivity

The study of connections between brain sites

Two main questions:

- What is a brain site?
  - ▶ Anatomically defined
  - ▶ Functionally defined
- What is a connection?
  - ▶ Anatomical measure (e.g. fiber track)
  - ▶ Functional measure (i.e. statistical)

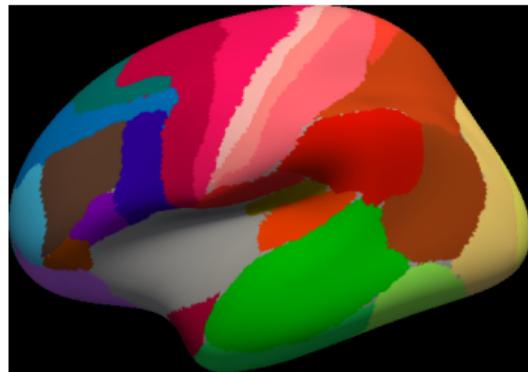
# Brain areas

A priori defined:

# Brain areas

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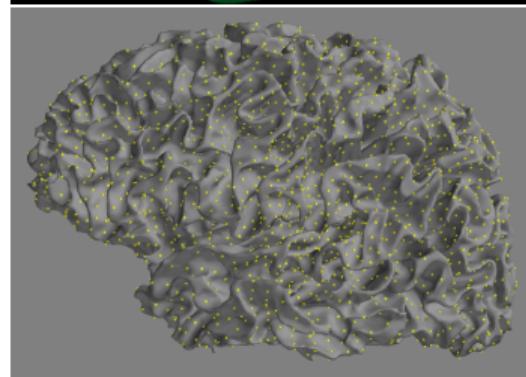
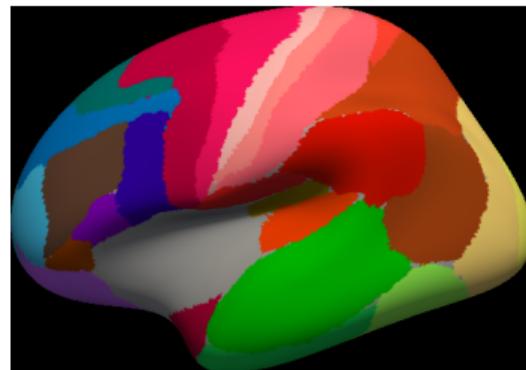
- Parcelations



# Brain areas

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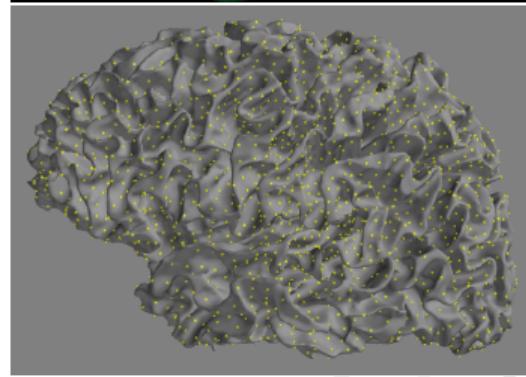
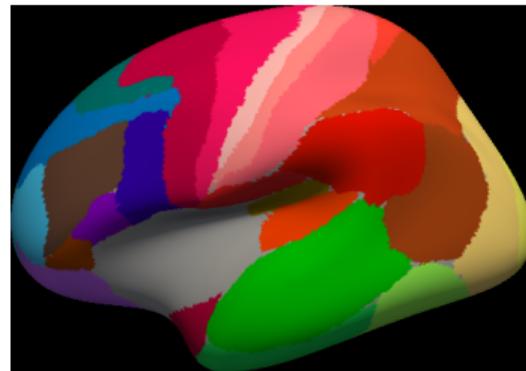
- Parcelations
- Source space points



# Brain areas

A priori defined:

- Parcels
- Source space points
- Voxels

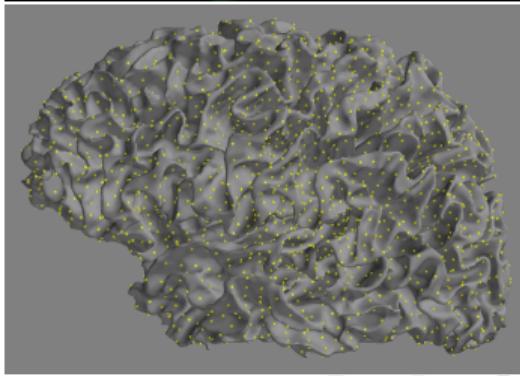
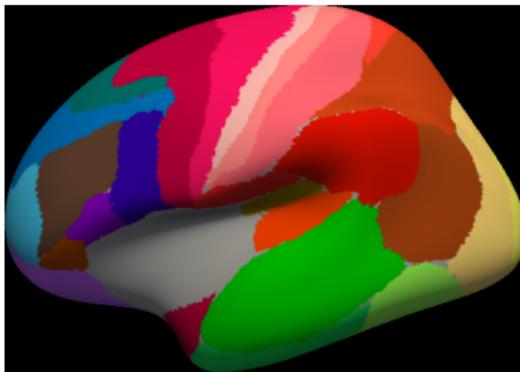


# Brain areas

A priori defined:

- Parcels
- Source space points
- Voxels

Functionally defined:



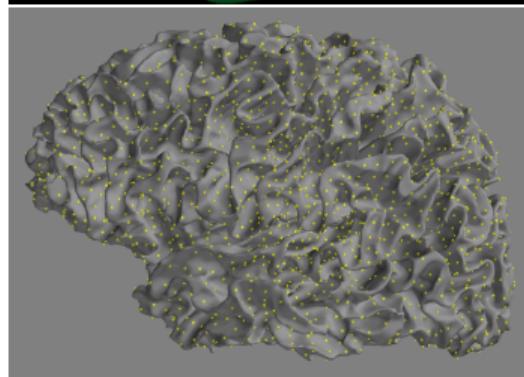
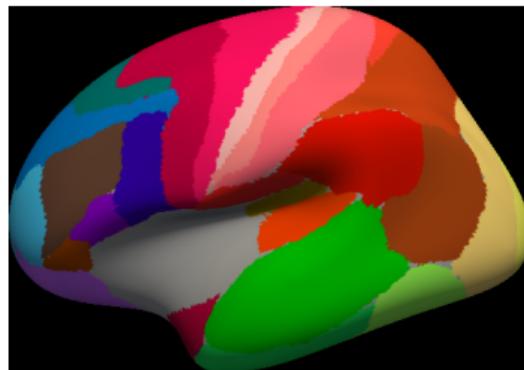
# Brain areas

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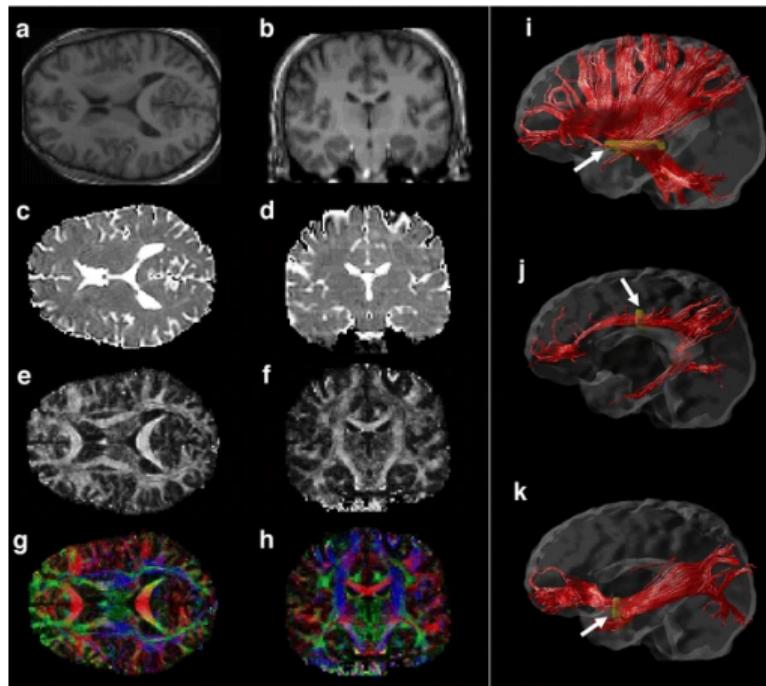
- Parcels
- Source space points
- Voxels

Functionally defined:

- From previous studies
  - ▶ Meta-analyses
  - ▶ Previous data from similar study/studies

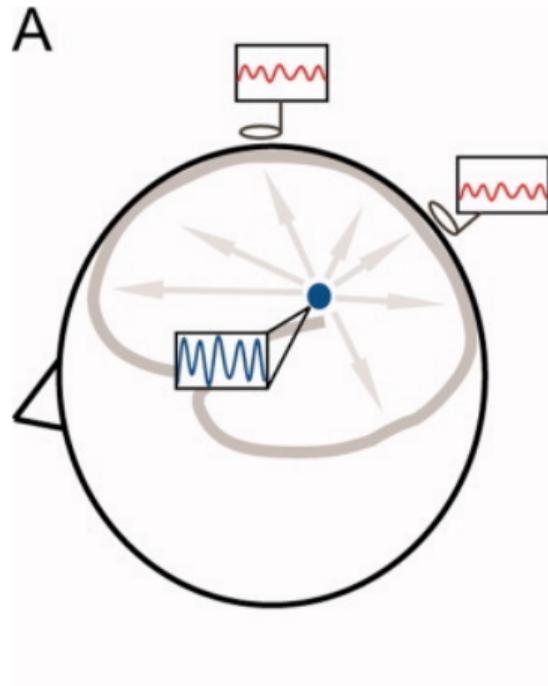


# Measuring connectivity: anatomical measures



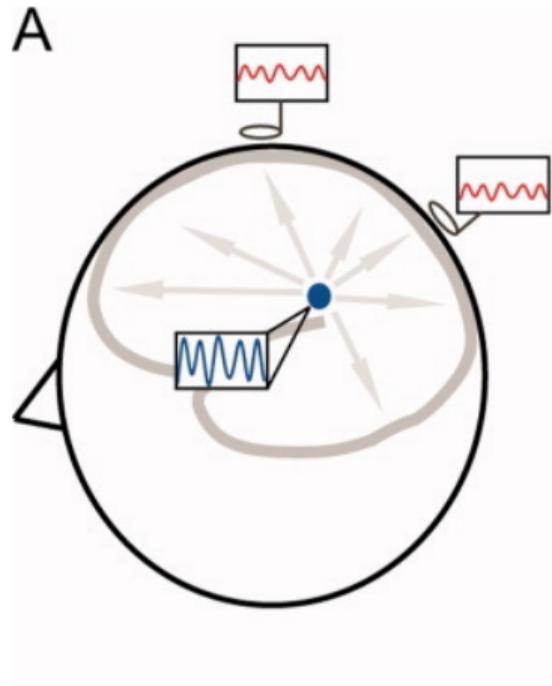
(Figure from Assaf & Pasternak, 2008)

# Measuring connectivity: statistical measures



(Figure from Schoffelen & Gross, 2009)

# Measuring connectivity: statistical measures

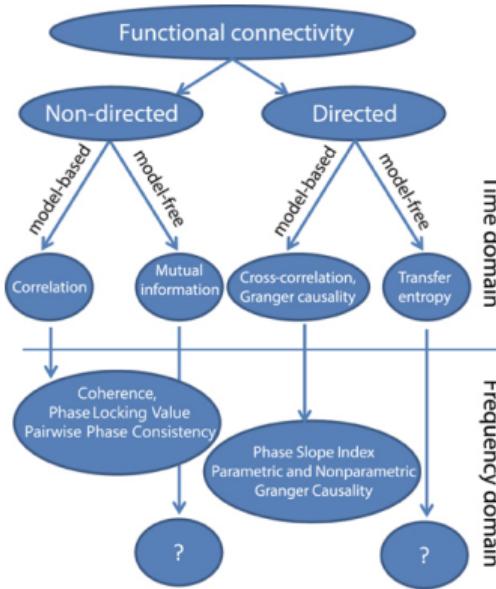


Options in MNE-python:

- Coherence
- Coherency
- Imaginary coherence
- Phase-Locking Value (PLV)
- Corrected imaginary PLV (icPLV)
- Pairwise Phase Consistency (PPC)
- Phase Lag Index (PLI)
- Unbiased estimator of squared PLI
- Weighted Phase Lag Index (WPLI)
- Debiased estimator of squared WPLI
- Mutual information

(Figure from Schoffelen & Gross, 2009)

# Taxonomy of functional connectivity analysis method



(Figure from Bastos & Schoffelen, 2016)

# Granger causality

[A] variable X is said to G[ranger]-cause a variable Y if the past of X contains information that helps predict the future of Y over and above information already in the past of Y.<sup>1</sup>

(Barnett & Seth, 2014, p. 51)

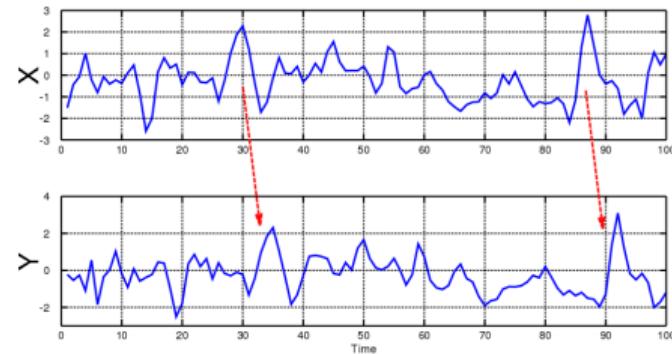
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<sup>1</sup>See also Granger (1969) for the original article.

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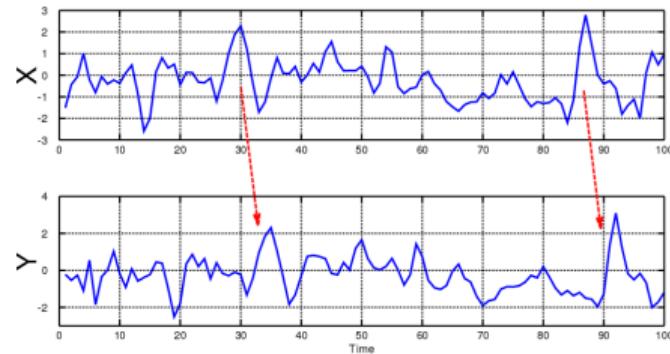


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Granger causality assumes stationarity!

<sup>1</sup>See also Granger (1969) for the original article.

# What is information?

Shannon entropy:

$$H(X) = - \sum_{i=1}^n p(x_i) \log_2 p(x_i)$$

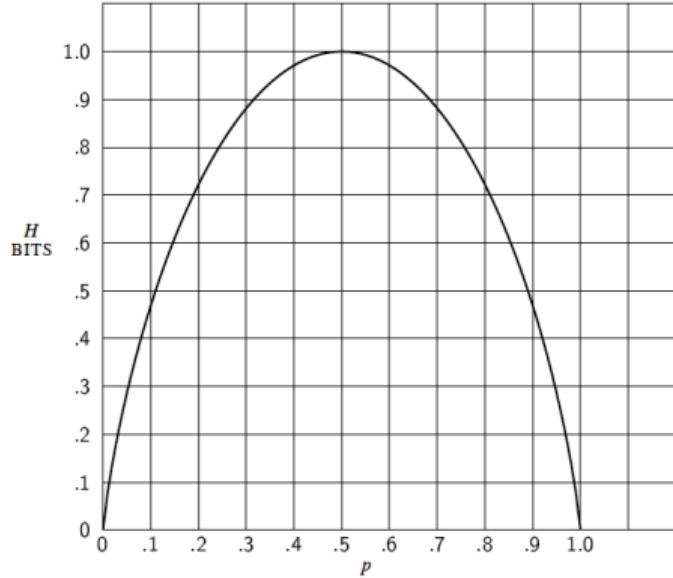
*“[Shannon entropy] is the amount of information, or “surprise,” a variable has” (Cohen, 2014, p. 389)*

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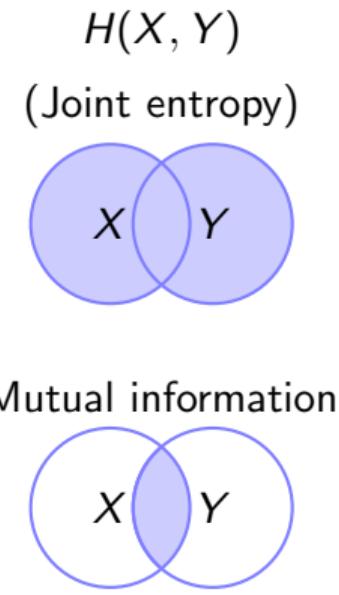
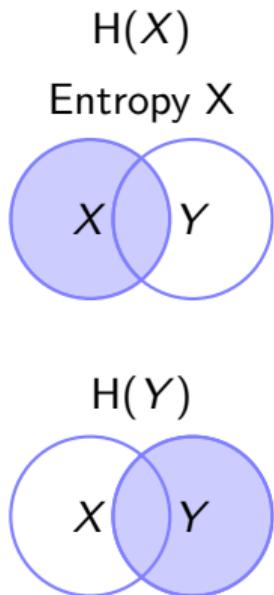
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(Figure from Shannon, 1948)

# What is information?



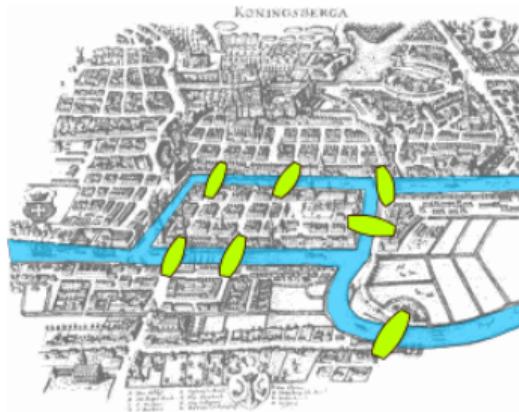
# What is information?

[T]he difference that makes a difference.

(Chalmers, 1996, p. 238)

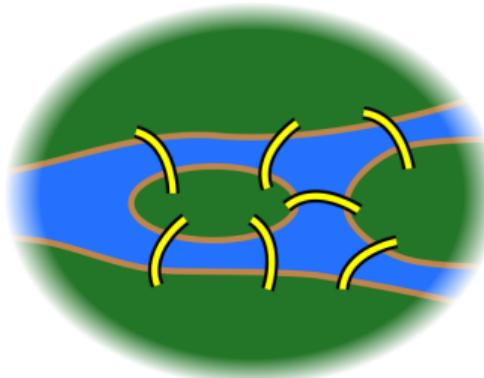
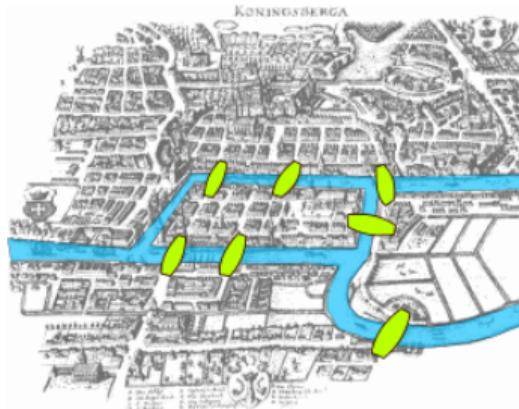
# Graph theory

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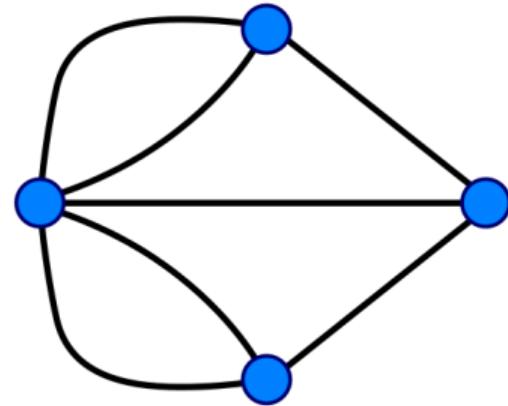
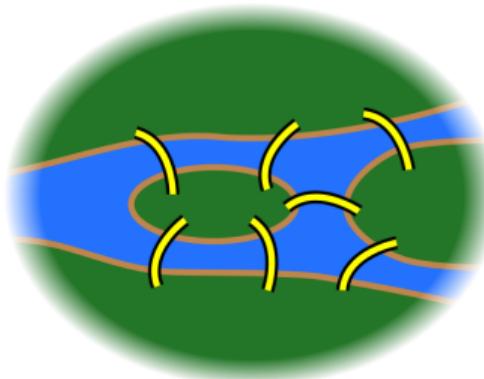
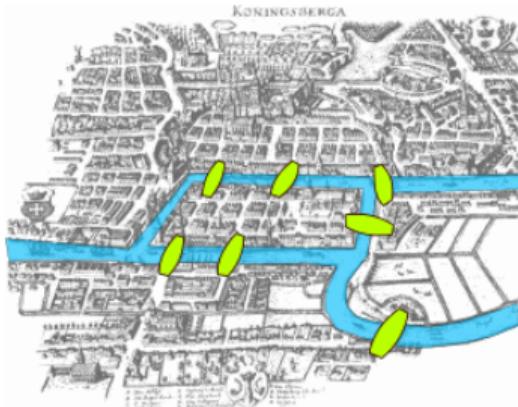
(Figure from [https://en.wikipedia.org/wiki/Seven\\_Bridges\\_of\\_Konigsberg](https://en.wikipedia.org/wiki/Seven_Bridges_of_Konigsberg))

# Graph theory



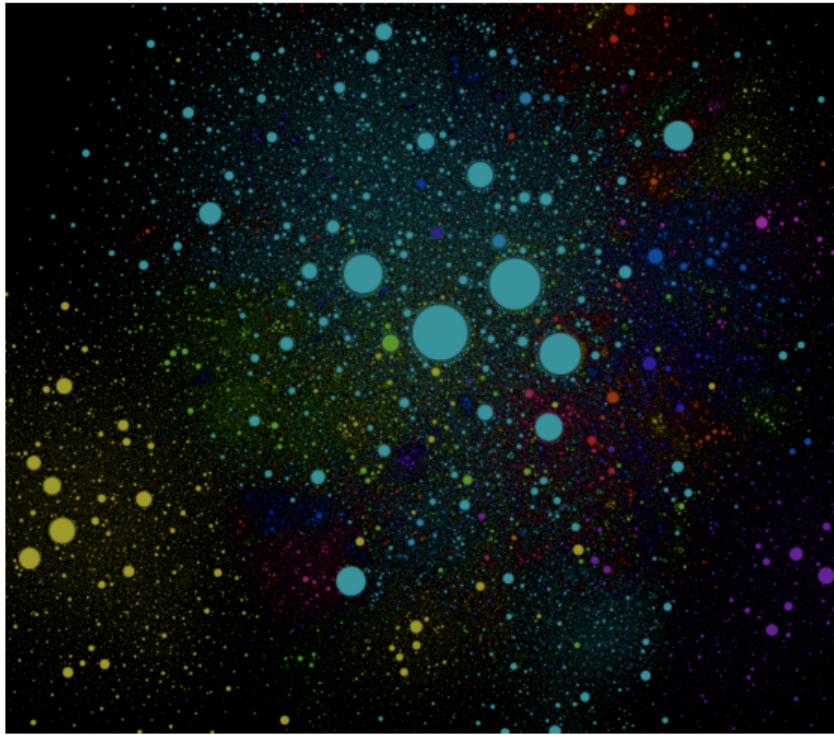
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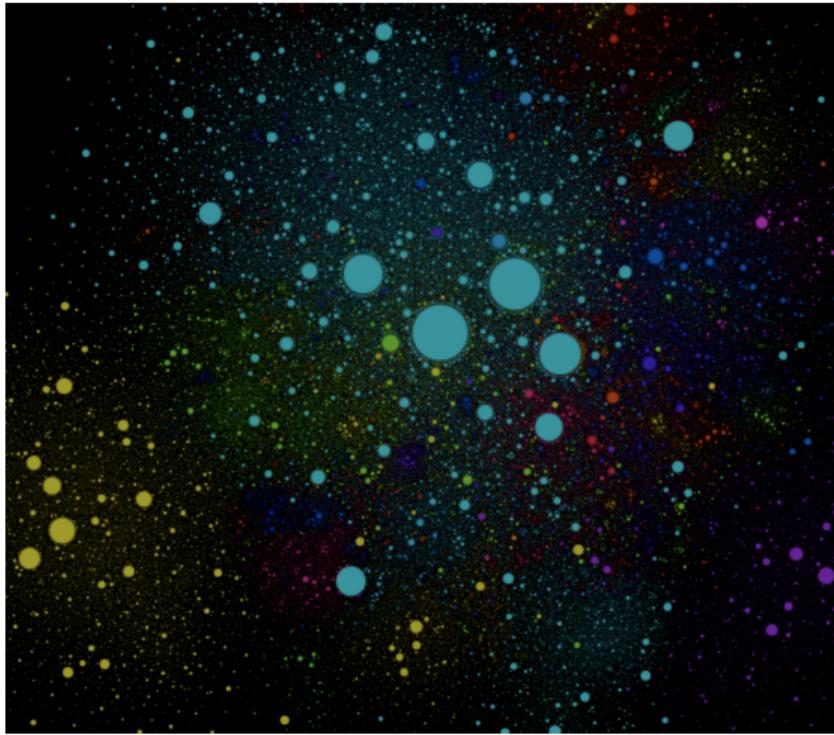


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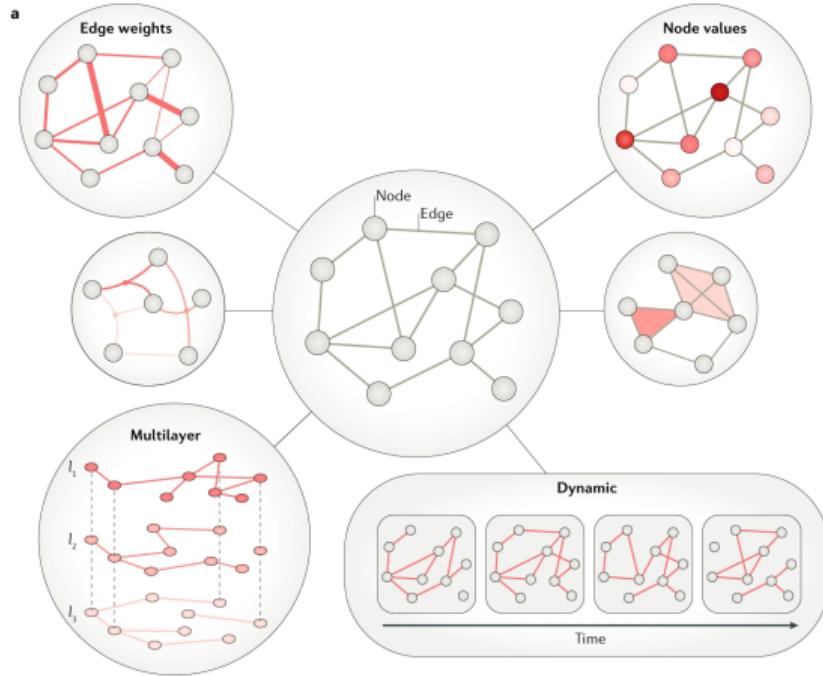
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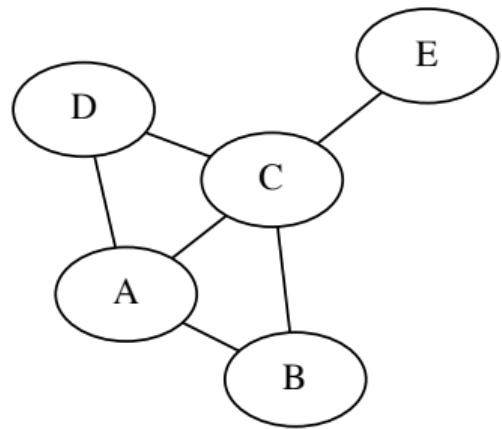
# Graph theory measures



(Figure from Bassett et al., 2018)

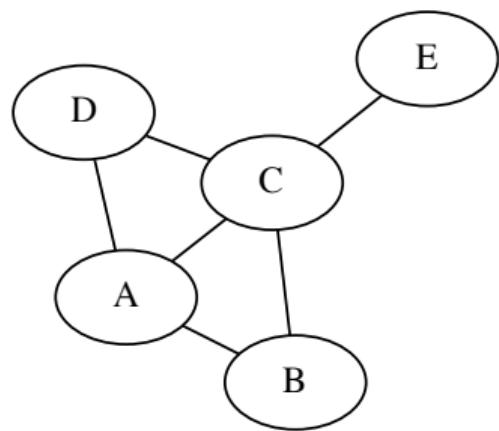
# Representation of graphs

## Graph



# Representation of graphs

Graph



Adjacency matrix

A				
B				
C				
D				
E				

# Graph theory measures

## Measures of functional segregation

“Functional segregation in the brain is the ability for specialized processing to occur within densely interconnected groups of brain regions. Measures of segregation primarily quantify the presence of such groups, known as clusters or modules, within the network.”

(Rubinov & Sporns, 2010, p. 1061)

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## Measures of functional integration

“Functional integration in the brain is the ability to rapidly combine specialized information from distributed brain regions. Measures of integration characterize this concept by estimating the ease with which brain regions communicate and are commonly based on the concept of a path.” (Rubinov & Sporns, 2010, p. 1061)

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## Measures of centrality

“Important brain regions (hubs) often interact with many other regions, facilitate functional integration, and play a key role in network resilience to insult. Measures of node centrality variously assess importance of individual nodes on the above criteria.” (Rubinov & Sporns, 2010, p. 1064)

# Graph theory measures

**Measures of functional  
segregation**

**Measures of functional  
integration**

**Measures of centrality**

# Graph theory measures

## Measures of functional segregation

- Clustering coefficient
- Transitivity

## Measures of functional integration

## Measures of centrality

# Graph theory measures

## Measures of functional segregation

- Clustering coefficient
- Transitivity

## Measures of functional integration

- Characteristic path length
- Global efficiency

## Measures of centrality

# Graph theory measures

## Measures of functional segregation

- Clustering coefficient
- Transitivity

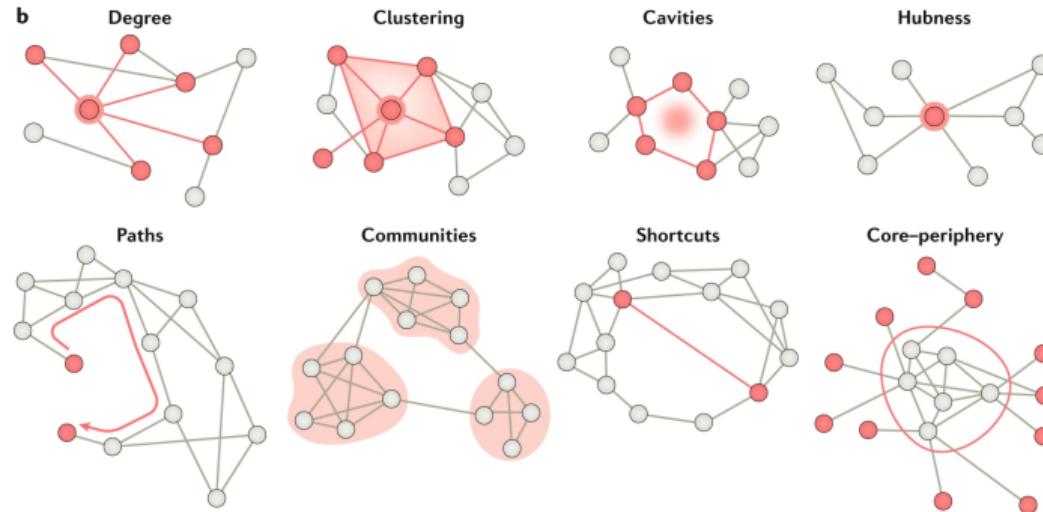
## Measures of functional integration

- Characteristic path length
- Global efficiency

## Measures of centrality

- Degree
- Closeness centrality
- PageRank

# Graph theory measures



(Figure from Bassett et al., 2018)

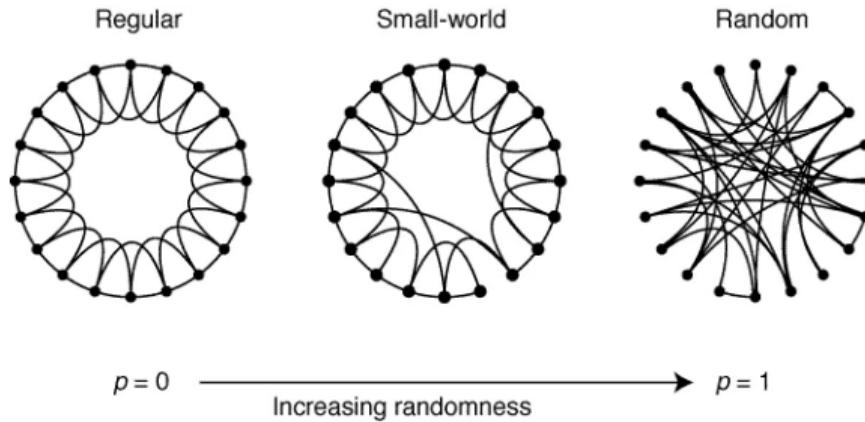
## Collective dynamics of 'small-world' networks

Duncan J. Watts\* & Steven H. Strogatz

*Department of Theoretical and Applied Mechanics, Kimball Hall,  
Cornell University, Ithaca, New York 14853, USA*



# Small-world networks



(Figure from Watts & Strogatz, 1998)

# Small-world networks

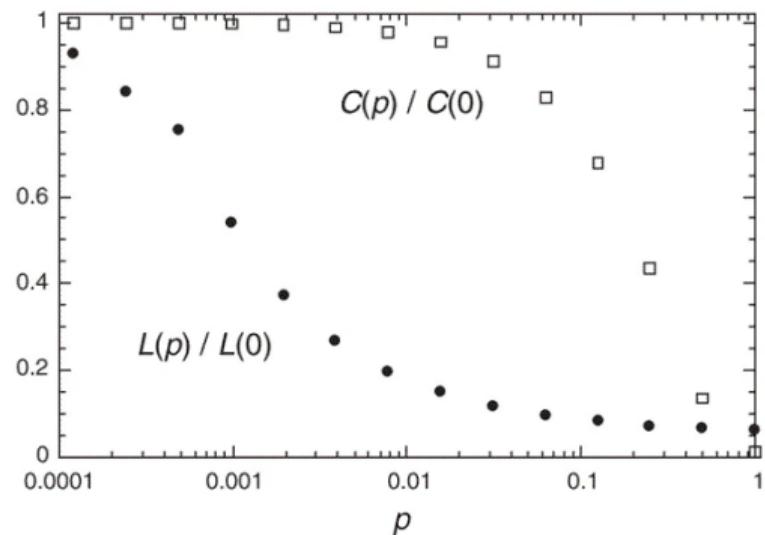
$$swn = \frac{\frac{C}{C_r}}{\frac{L}{L_r}}$$

$C$  is the clustering coefficient,  $L$  is the characteristic path length, subscript  $r$  refer to a random network.

# Small-world networks

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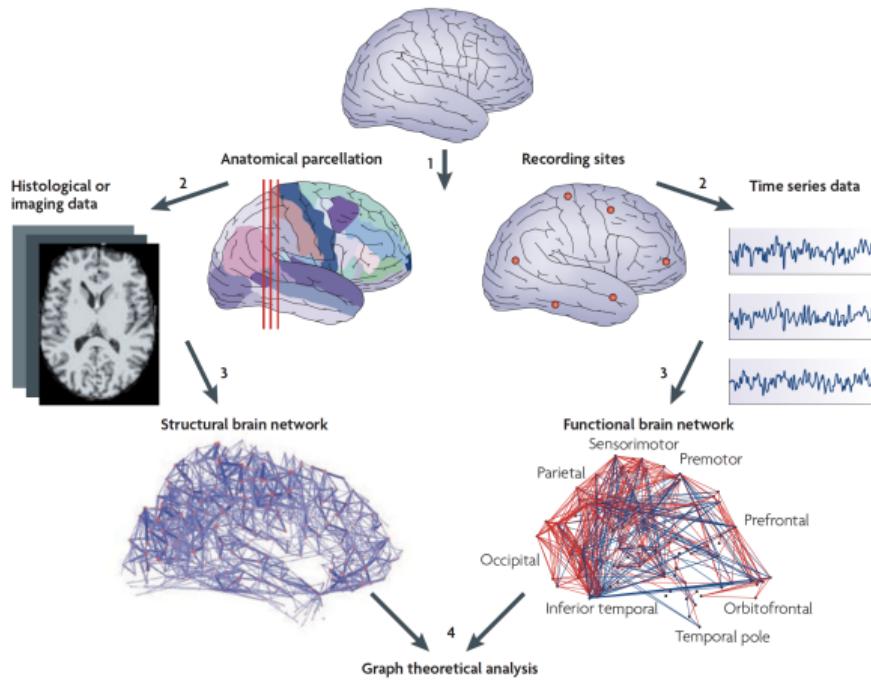
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(Figure from Watts & Strogatz, 1998)

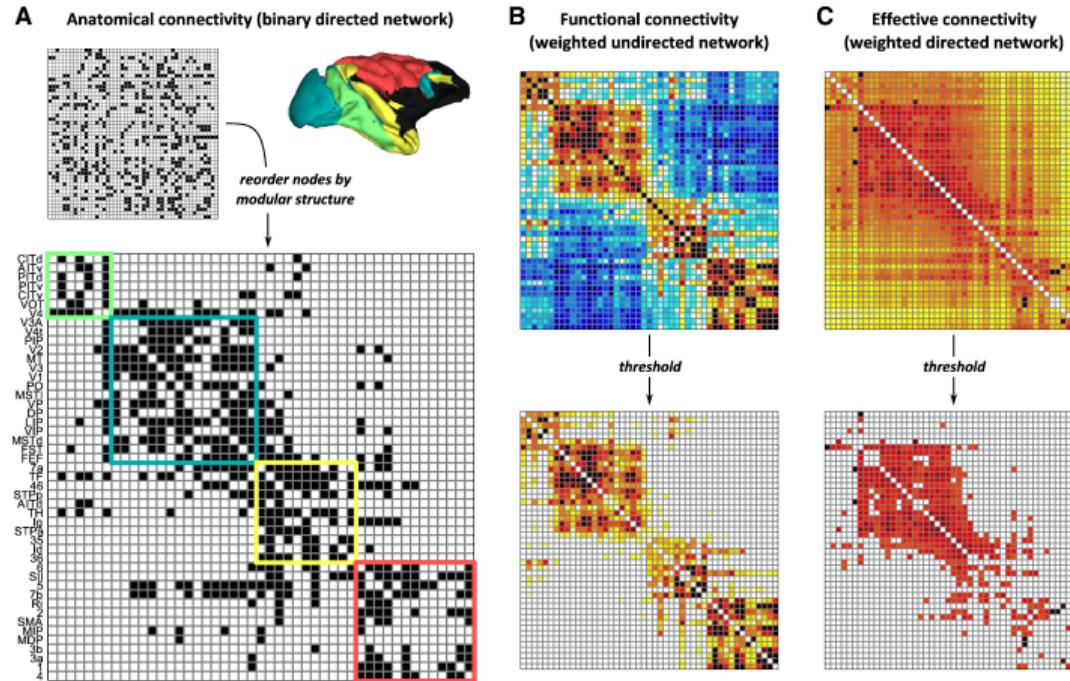
# Graph theory and brain data

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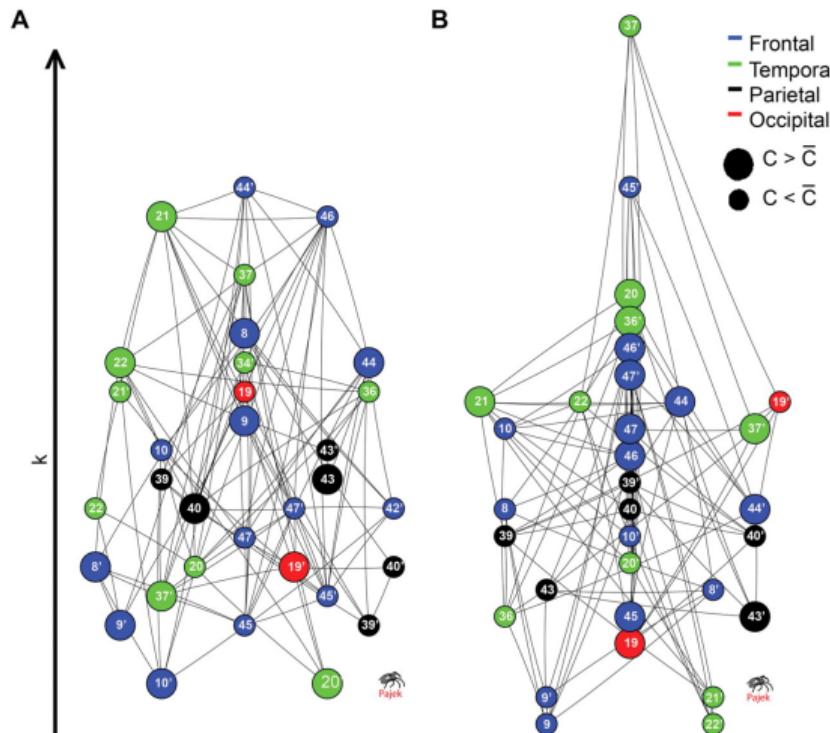
(Figure from Bullmore & Sporns, 2009)

# Graph theory and brain data



(Figure from Rubinov & Sporns, 2010)

# Human cortical networks in health and schizophrenia

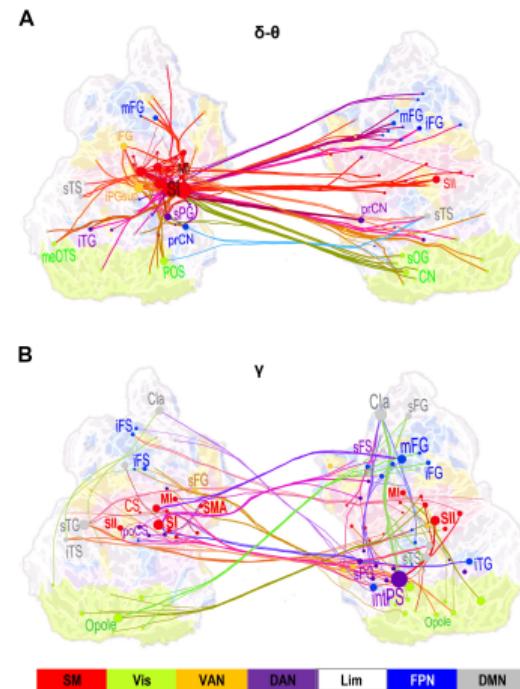


(Figure from Bassett et al., 2008)

Mads Jensen (RFR, IMC, & CFIN)

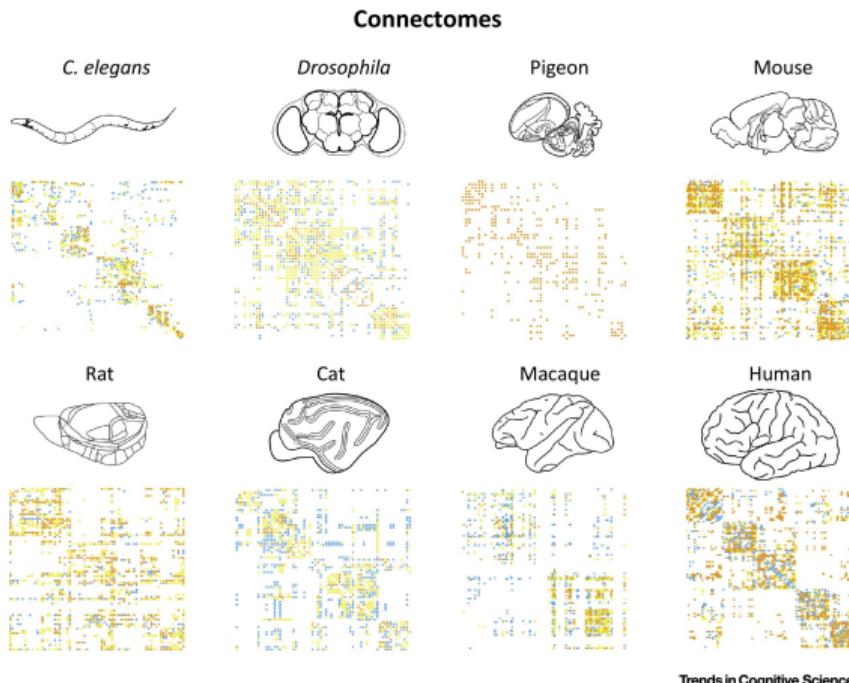
Connectivity

# Dynamic large-scale network synchronization from perception to action



(Figure from Hirvonen et al., 2018)

# Connectome across species



(Figure from van den Heuvel et al., 2016)

# Dynamic representations in networked neural systems

# Dynamic representations in networked neural systems

*“Our goal is to offer a holistic framework for understanding and describing neural information representation and transmission while revealing exciting frontiers for future research.”*

(Ju & Bassett, 2020, p. 908)

# Neural representation

"In studying neural representations, an important recent step has been to measure how a population of neurons or voxels (i.e., volumes of brain tissue) can represent variables by activating in a specific spatial pattern in response to a particular stimulus pattern."

(Ju & Bassett, 2020, p. 908)

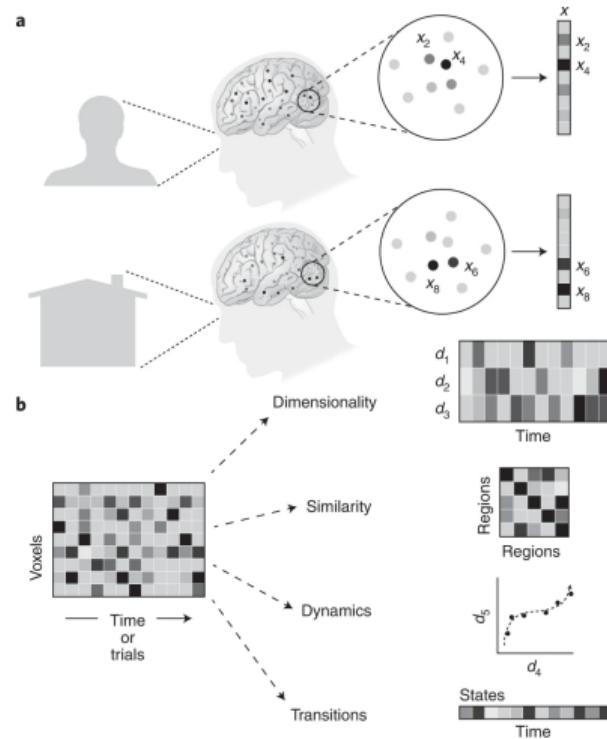
# Neural representation

“In studying neural representations, an important recent step has been to measure how a population of neurons or voxels (i.e., volumes of brain tissue) can represent variables by activating in a specific spatial pattern in response to a particular stimulus pattern.”

“The encoding of representations in neural populations offers a computational advantage over encoding in individual neurons, especially in complex cognitive tasks”

(Ju & Bassett, 2020, p. 908)

# Neural representations and tools to analyze them

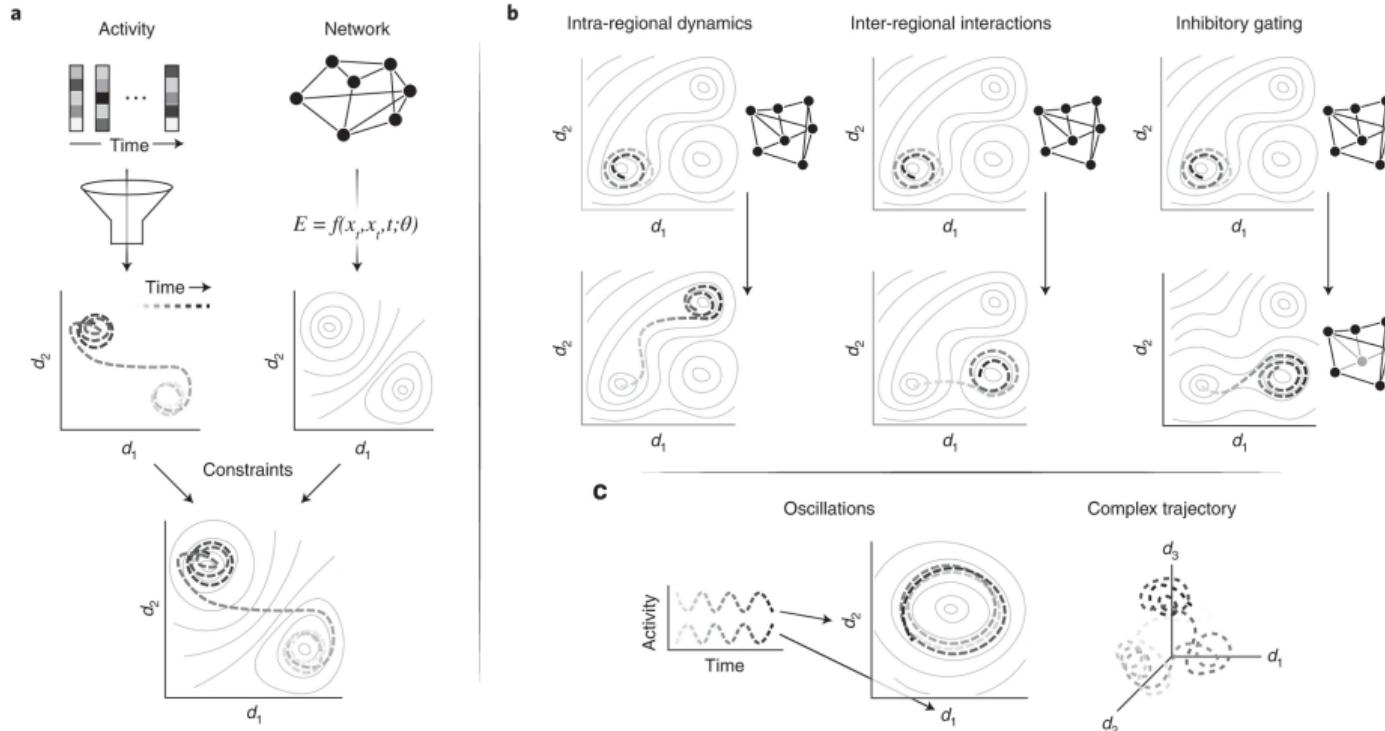


(Figure from Ju & Bassett, 2020)

Mads Jensen (RFR, IMC, & CFIN)

Connectivity

# Network models abstract neural systems

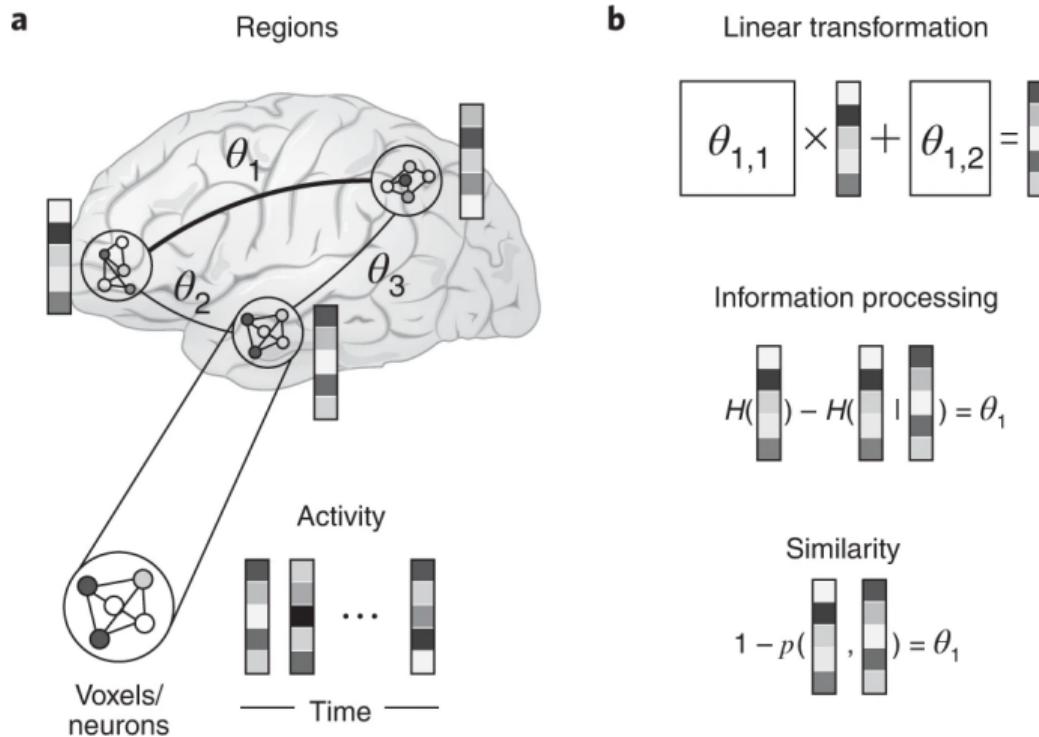


(Figure from Ju & Bassett, 2020)

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Connectivity

# Integrating network models and neural representations

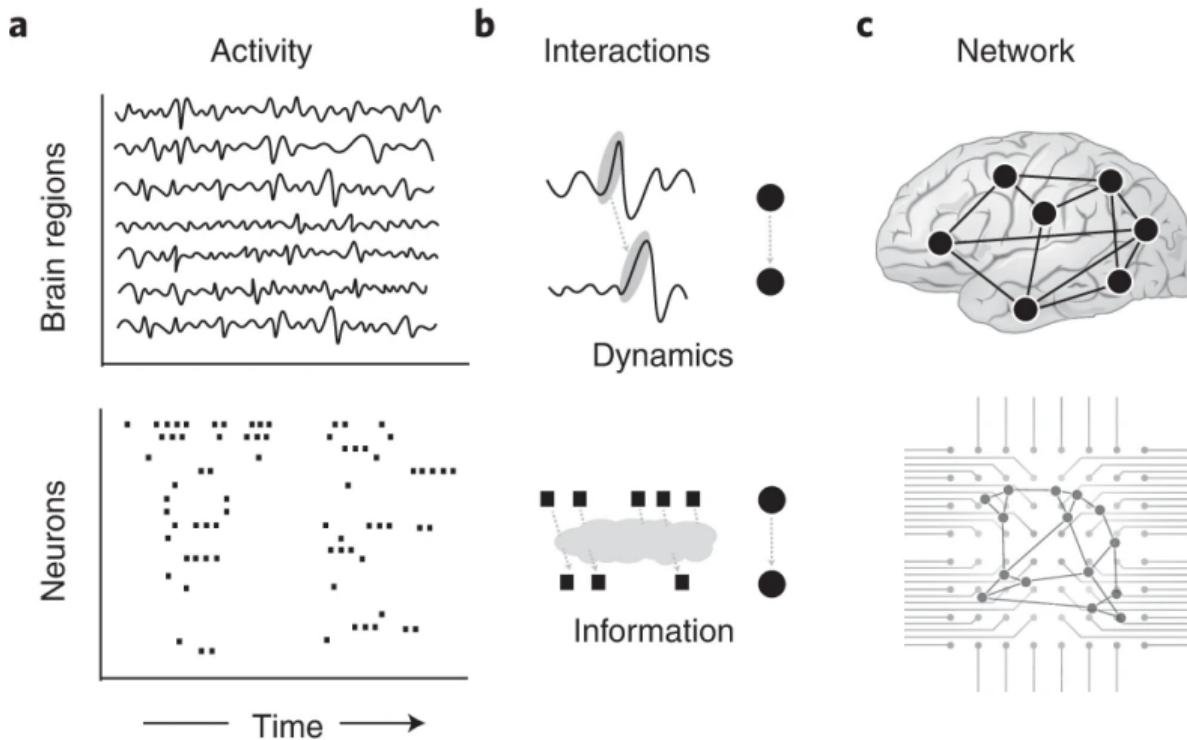


(Figure from Ju & Bassett, 2020)

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# Dynamic representations in networked neural systems



(Figure from Ju & Bassett, 2020)

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Connectivity

# Summary: Dynamic representations in networked neural systems

## References I

- Assaf, Y., & Pasternak, O. (2008). Diffusion Tensor Imaging (DTI)-based White Matter Mapping in Brain Research: A Review. *Journal of Molecular Neuroscience*, 34, 51–61.  
<https://doi.org/10.1007/s12031-007-0029-0>
- Barnett, L., & Seth, A. K. (2014). The MVGC multivariate Granger causality toolbox: A new approach to Granger-causal inference. *Journal of Neuroscience Methods*, 223, 50–68.  
<https://doi.org/10.1016/j.jneumeth.2013.10.018>
- Bassett, D. S., Bullmore, E., Verchinski, B. A., Mattay, V. S., Weinberger, D. R., & Meyer-Lindenberg, A. (2008). Hierarchical organization of human cortical networks in health and schizophrenia. *The Journal of Neuroscience*, 28, 9239–9248.  
<http://www.jneurosci.org/content/jneuro/28/37/9239.full.pdf>
- Bassett, D. S., Zurn, P., & Gold, J. I. (2018). On the nature and use of models in network neuroscience. *Nature Reviews Neuroscience*, 19(9), 566–578.

## References II

- Bastos, A. M., & Schoffelen, J.-M. (2016). A Tutorial Review of Functional Connectivity Analysis Methods and Their Interpretational Pitfalls. *Frontiers in Systems Neuroscience*, 9. <https://doi.org/10.3389/fnsys.2015.00175>
- Bullmore, E., & Sporns, O. (2009). Complex brain networks: Graph theoretical analysis of structural and functional systems. *Nature Reviews Neuroscience*, 10, 186–98. <https://doi.org/10.1038/nrn2575>
- Chalmers, D. J. (1996). *The Conscious Mind – In Search of a Fundamental Theory*. Oxford University Press.
- Cohen, M. X. (2014). *Analyzing neural time series data: Theory and practice*. The MIT Press.
- Granger, C. W. (1969). Investigating causal relations by econometric models and cross-spectral methods. *Econometrica: journal of the Econometric Society*, 424–438.
- Hirvonen, J., Monto, S., Wang, S. H., Palva, J. M., & Palva, S. (2018). Dynamic large-scale network synchronization from perception to action. *Network Neuroscience*, 1–41.

## References III

- Ju, H., & Bassett, D. S. (2020). Dynamic representations in networked neural systems. *Nature Neuroscience*, (8). <https://doi.org/10.1038/s41593-020-0653-3>
- Rubinov, M., & Sporns, O. (2010). Complex network measures of brain connectivity: Uses and interpretations. *NeuroImage*, 52, 1059–1069.  
<https://doi.org/10.1016/j.neuroimage.2009.10.003>
- Schoffelen, J.-M., & Gross, J. (2009). Source connectivity analysis with MEG and EEG. *Human Brain Mapping*, 30, 1857–65. <https://doi.org/10.1002/hbm.20745>
- Shannon, C. E. (1948). A mathematical theory of communication. *The Bell system technical journal*, 27(3), 379–423.
- van den Heuvel, M. P., Bullmore, E. T., & Sporns, O. (2016). Comparative Connectomics. *Trends in cognitive sciences*. <https://doi.org/10.1016/j.tics.2016.03.001>
- Watts, D. J., & Strogatz, S. H. (1998). Collective dynamics of 'small-world' networks. *Nature*, 393, 440–2. <https://doi.org/10.1038/30918>