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Aalto University  
School of Science

# Methods for naturalistic neuroimaging

**Enrico Glerean (D.Sc.)**

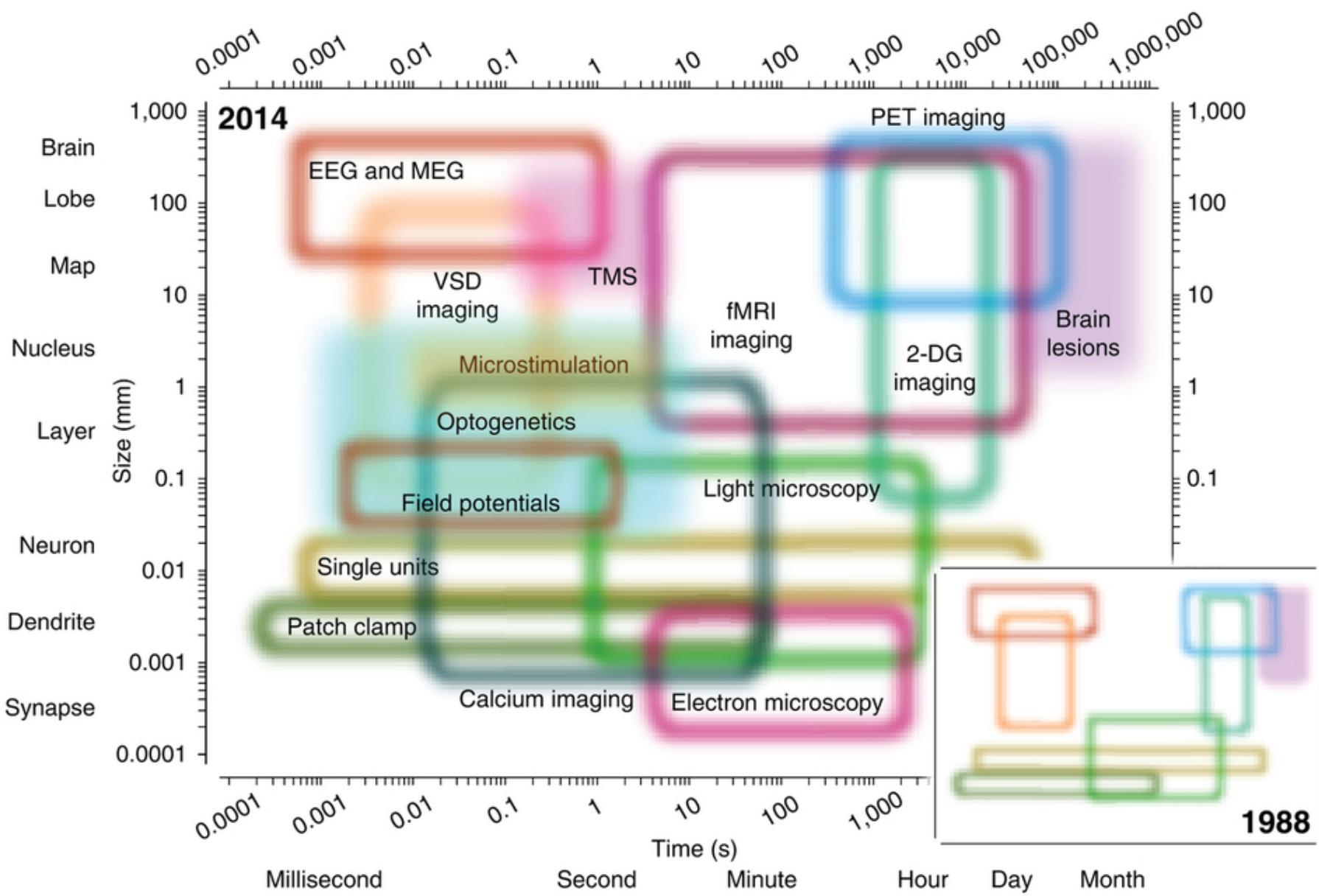
[www.glerean.com](http://www.glerean.com) | @eglerean | [enrico.glerean@aalto.fi](mailto:enrico.glerean@aalto.fi)

# Methods for naturalistic neuroimaging

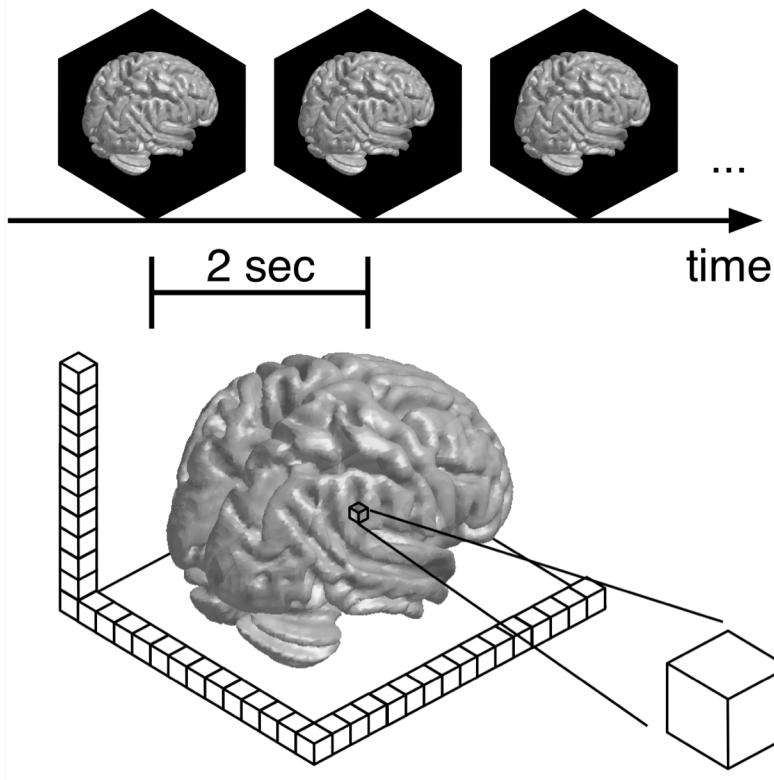
## –Outline

- Why naturalistic neuroimaging?
  - *fMRI + movies*
- Complex stimuli bring new methodological challenges
  - *Intersubject correlation*
  - *Intersubject phase synchronisation*
  - *Intersubject functional correlation*
  - *Intersubject representation similarity analysis*
- Statistical modelling
  - *Statistical considerations*
  - *Brain & behaviour modelling*
- Reflections and future directions

# Space and time in neuro-data



# Functional magnetic resonance imaging (fMRI)



- We measure **multiple time series** at once
- **NOTE:** 1 voxel  $\rightarrow$  5.5e6 neurons 4e10 synapses (density  $\sim$ 1/1000) [Logothetis 2008 Nature]

# Why naturalistic neuroimaging?

- **Naturalistic paradigms:** using movies, audio narratives, music, videogames and other engaging activities during brain scanning (usually fMRI or M/EEG).

[NeuroImage S.I. curated by Vanderwal, Finn and Glerean.](#)



The screenshot shows the homepage of the NeuroImage journal. At the top left is the journal logo featuring a bookshelf icon and the word "NeuroImage". To the right of the logo is the journal title "NeuroImage" and the text "Open access". Below the title are three navigation links: "Articles and issues ▾", "About ▾", and "Publish ▾". A search bar at the bottom of the page contains the placeholder text "Search in this journal".

Naturalistic Imaging: The use of ecologically valid conditions to study brain function

# Why naturalistic neuroimaging?

- Naturalistic paradigms
  - Pro: **closer to real life, only way to study some mental processes** (e.g. processing structure in narratives or movies)
  - Pro: growing number of labs agree this is **the way to go in the future** (see Gallant's lab work in *Nature* 2016)
  - Cons: reviewers complain: **lots of uncontrolled stimulus features**
  - Cons: **stimulus is the same for all subjects**

## Some essential reviews from past months:

- [Sonkusare, S., Breakspear, M., & Guo, C. \(2019\). Naturalistic stimuli in neuroscience: Critically acclaimed. \*Trends in cognitive sciences\*.](#)
- [Eickhoff, S. B., Milham, M., & Vanderwal, T. \(2020\). Towards clinical applications of movie fMRI. \*NeuroImage\*, 116860.](#)

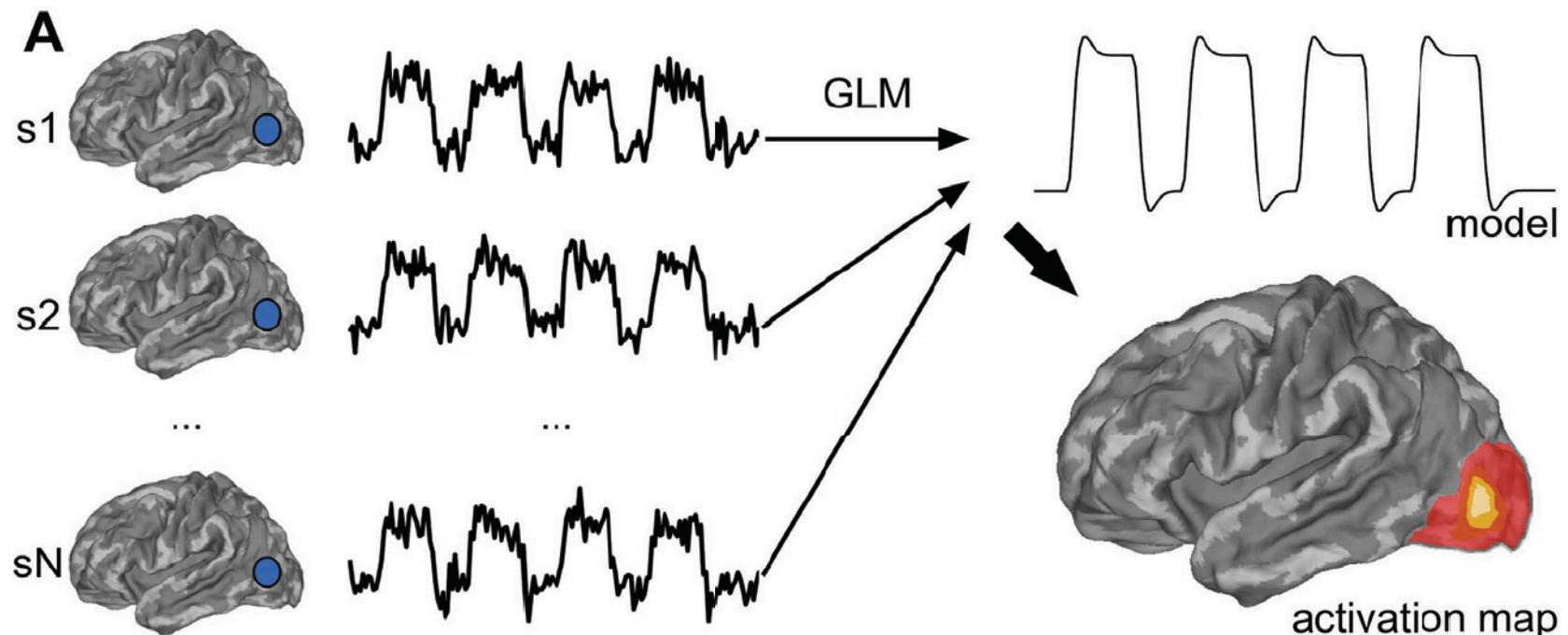
# Analysing naturalistic imaging data

*1) Intersubject correlation*

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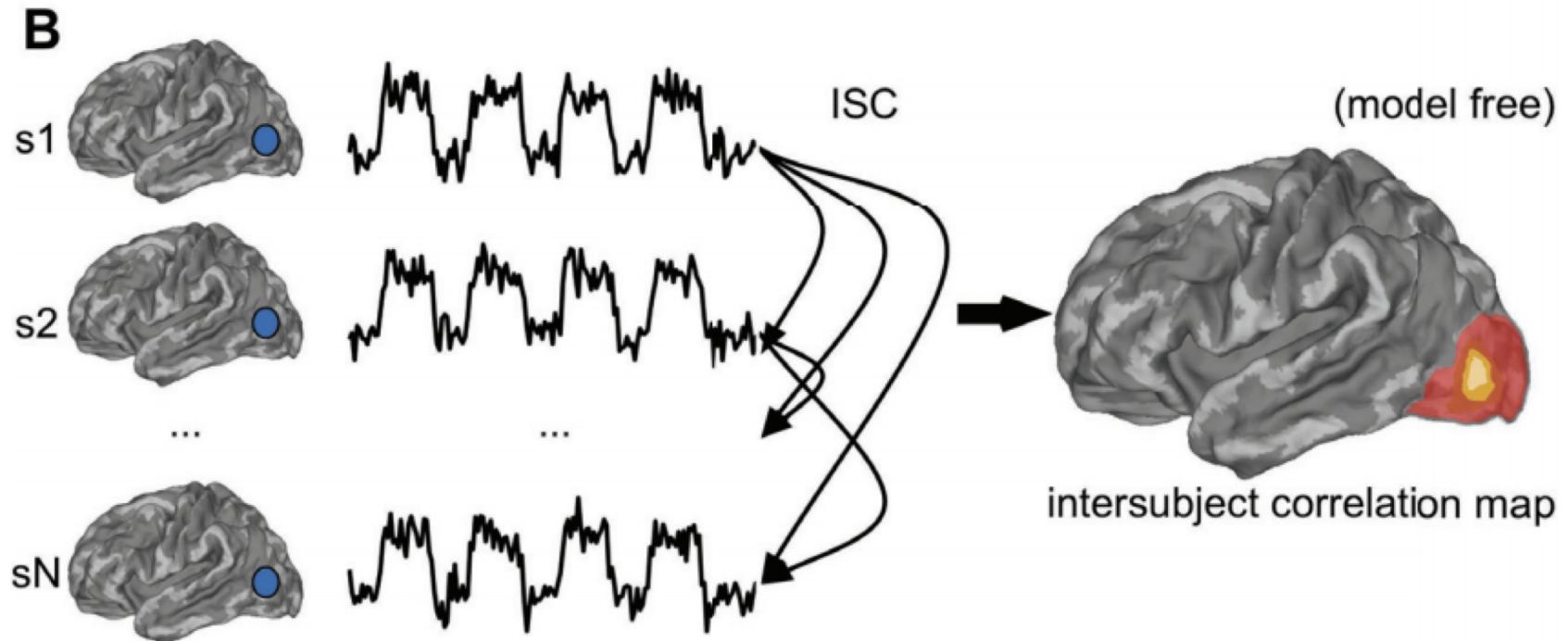
# Individual responses to a model

Controlled experiments with a known model (General linear model)

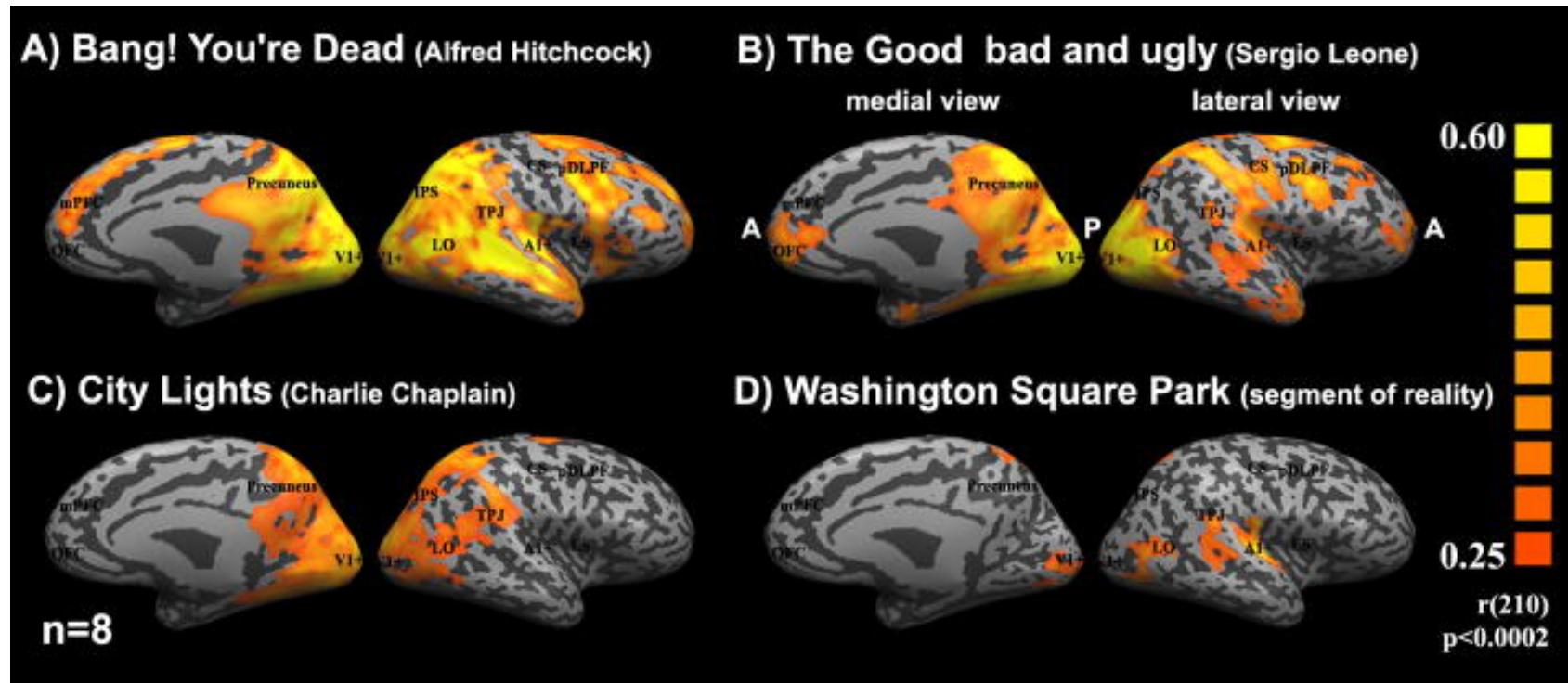


# Intersubject Correlation (ISC)

Model unknown (e.g. complex audio/visual stimulation). Other subjects' activity becomes the model!



# Measuring functional similarity – Intersubject Correlation (ISC)



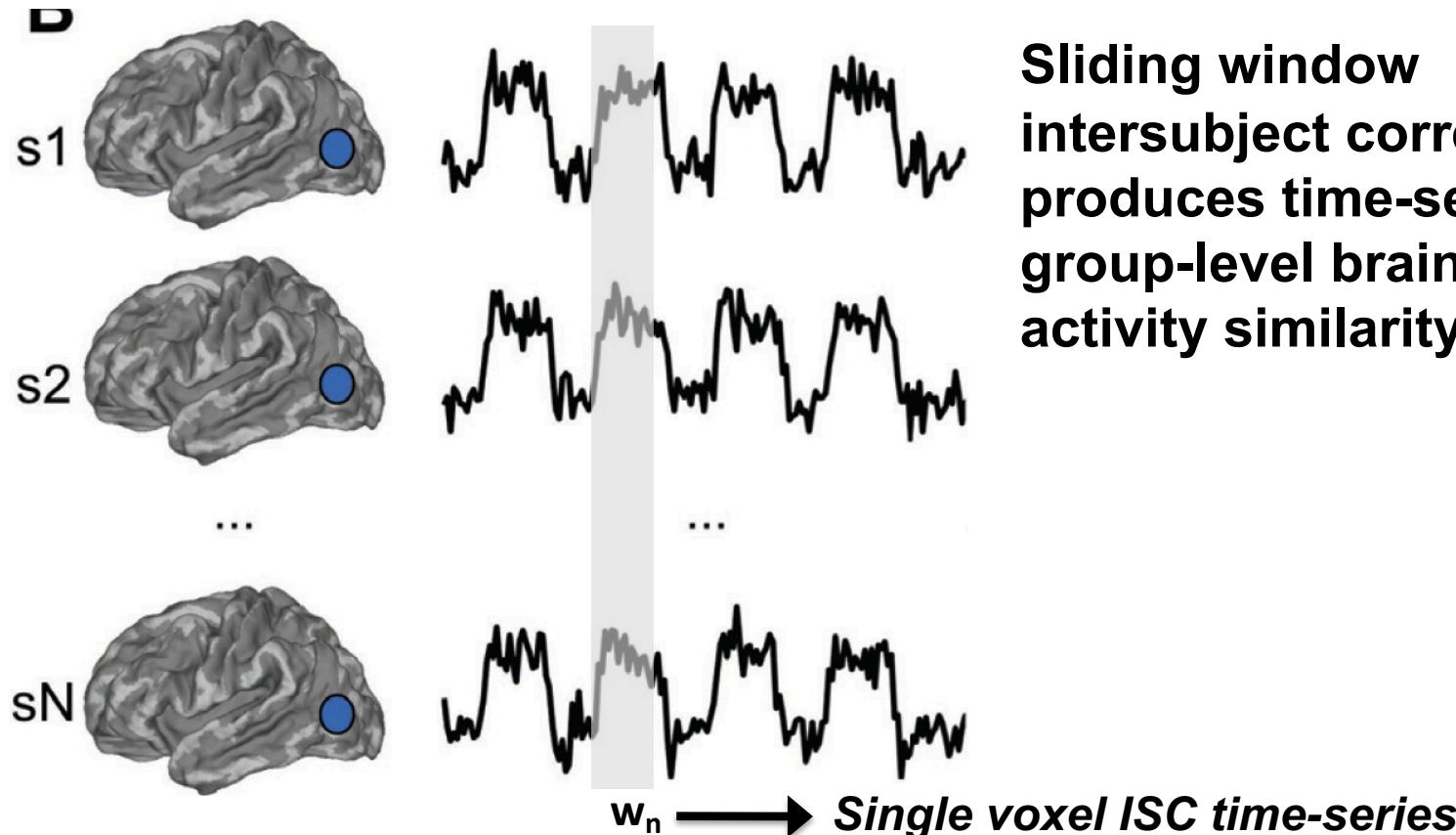
*Hasson et al. (2010) Trends in Cognitive Sciences*

**Complex stimuli  
are constantly  
changing**

**2) *dynamic ISC***

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# Dynamic ISC: functional similarity in time



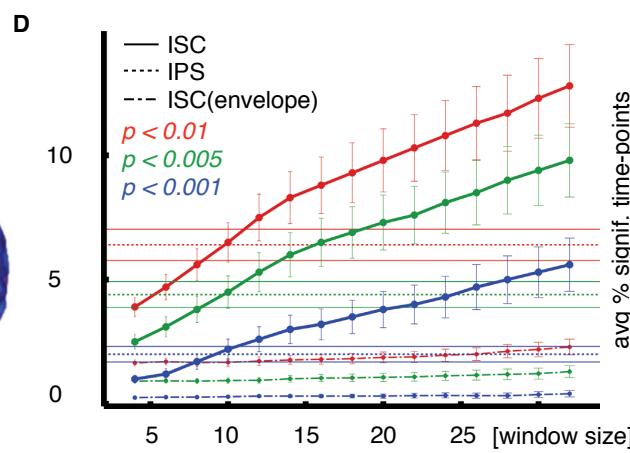
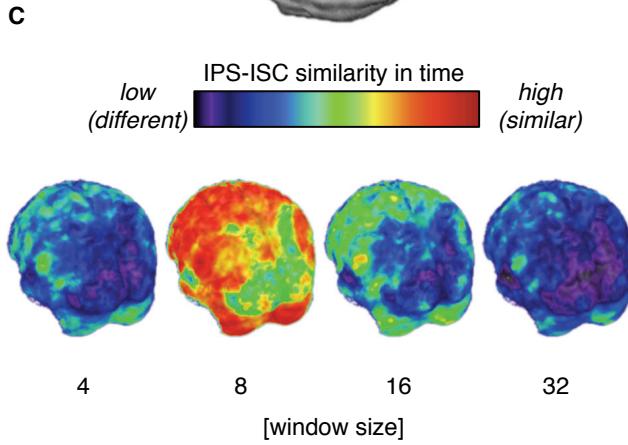
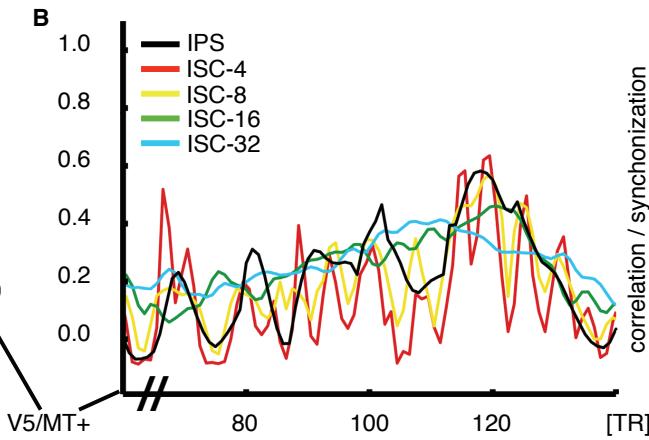
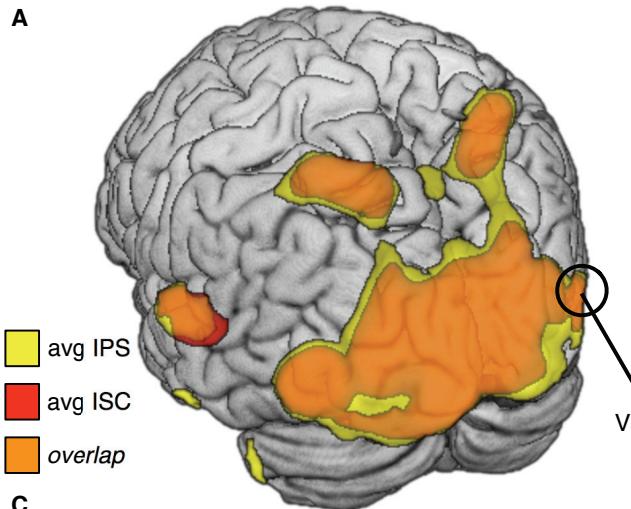
Sliding window intersubject correlation produces time-series of group-level brain activity similarity.

**Complex stimuli  
are constantly  
changing 2**

*3) intersubject phase  
synchronisation*

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# Intersubject phase synchronisation



Average ISC matches temporal average of IPS

Short ISC windows provide noisy ISC time series.

Longer ISC windows are missing the fast changes

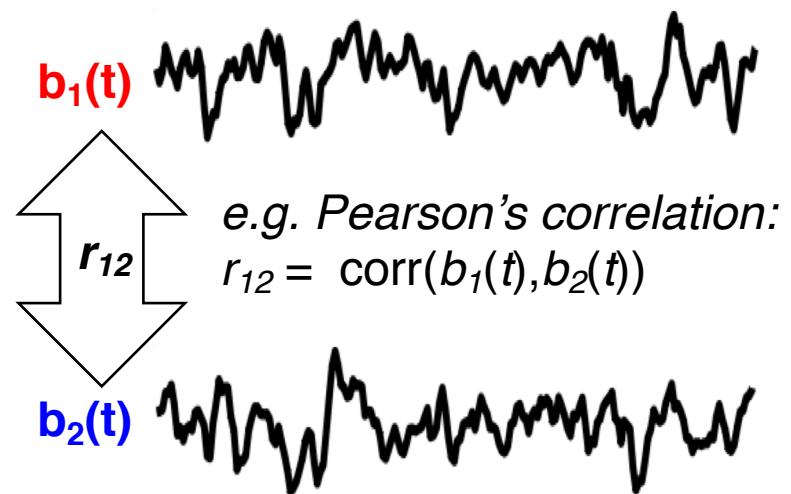
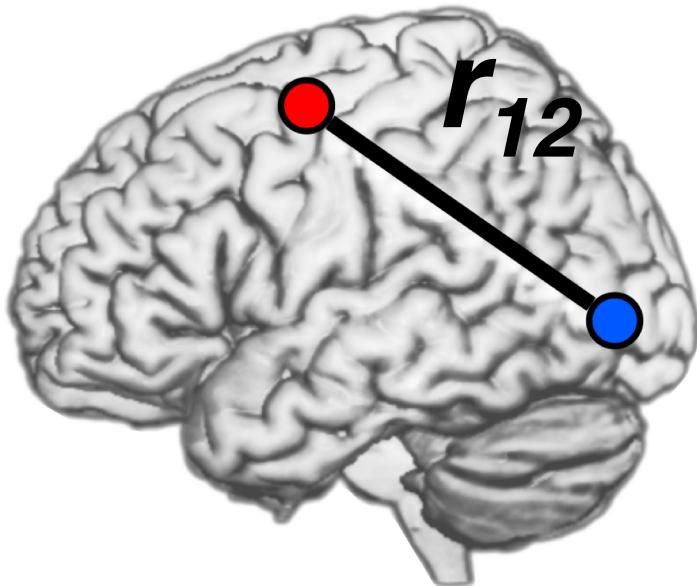
IPS and sliding window ISC seem to match at windows of size 10

# Functional connectivity with naturalistic imaging

*4) functional networks*

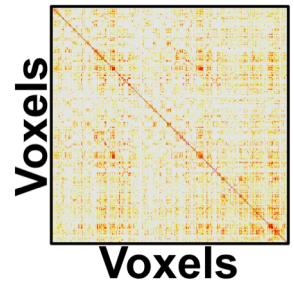
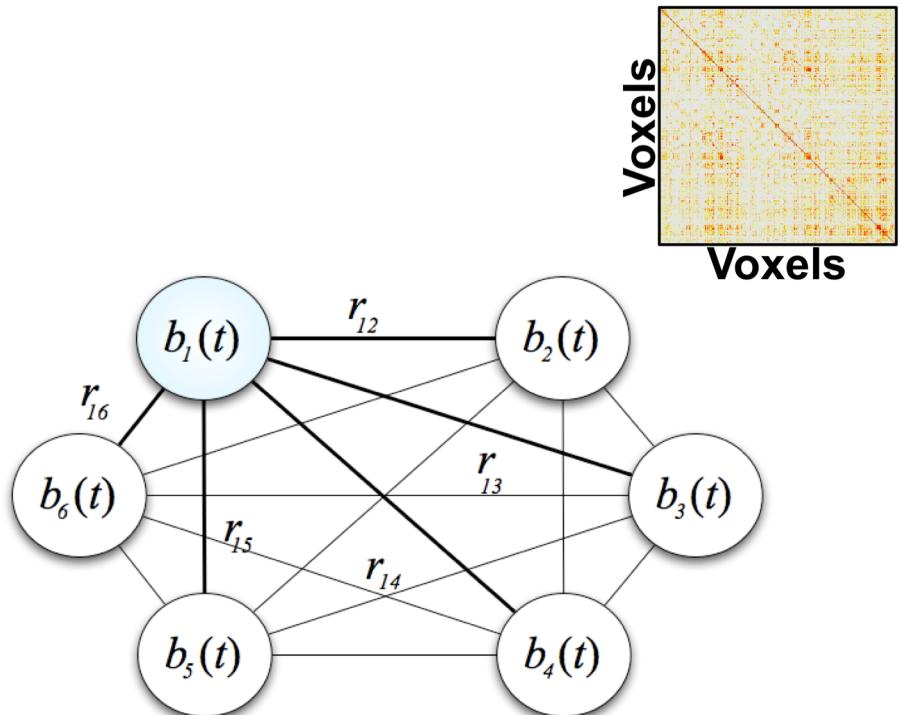
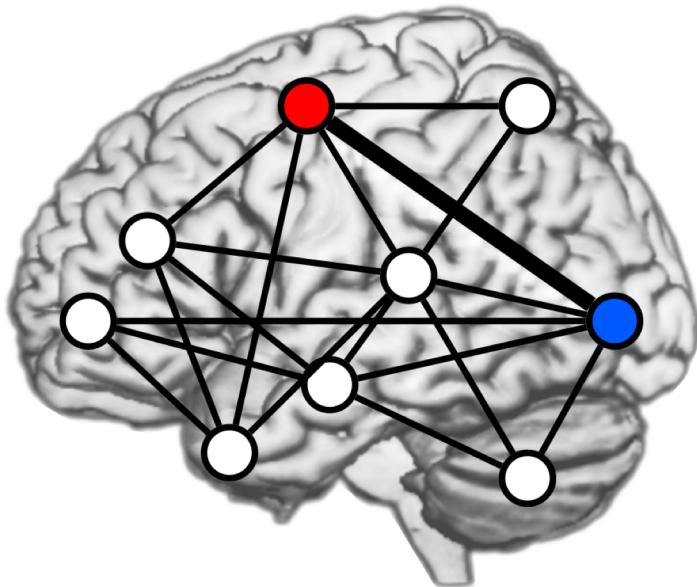
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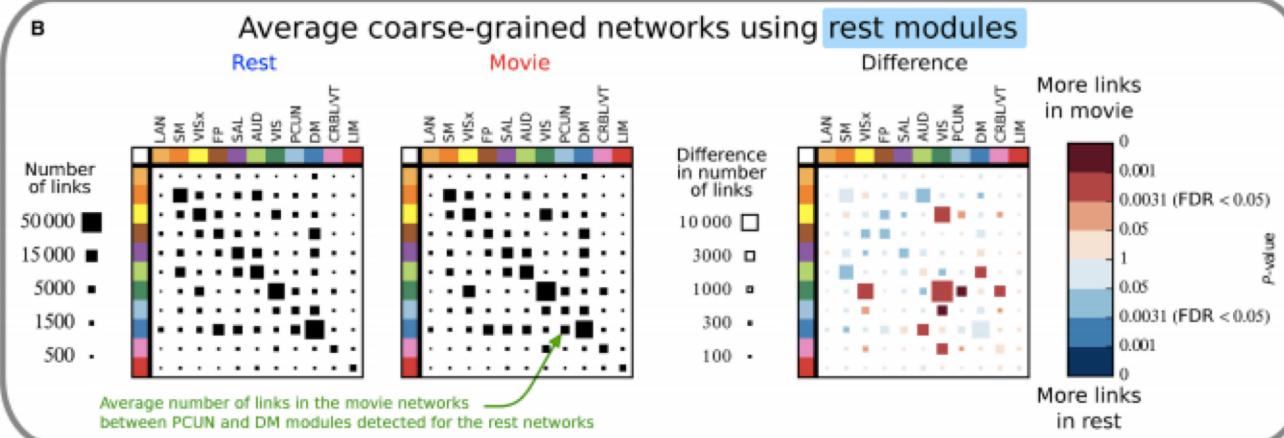
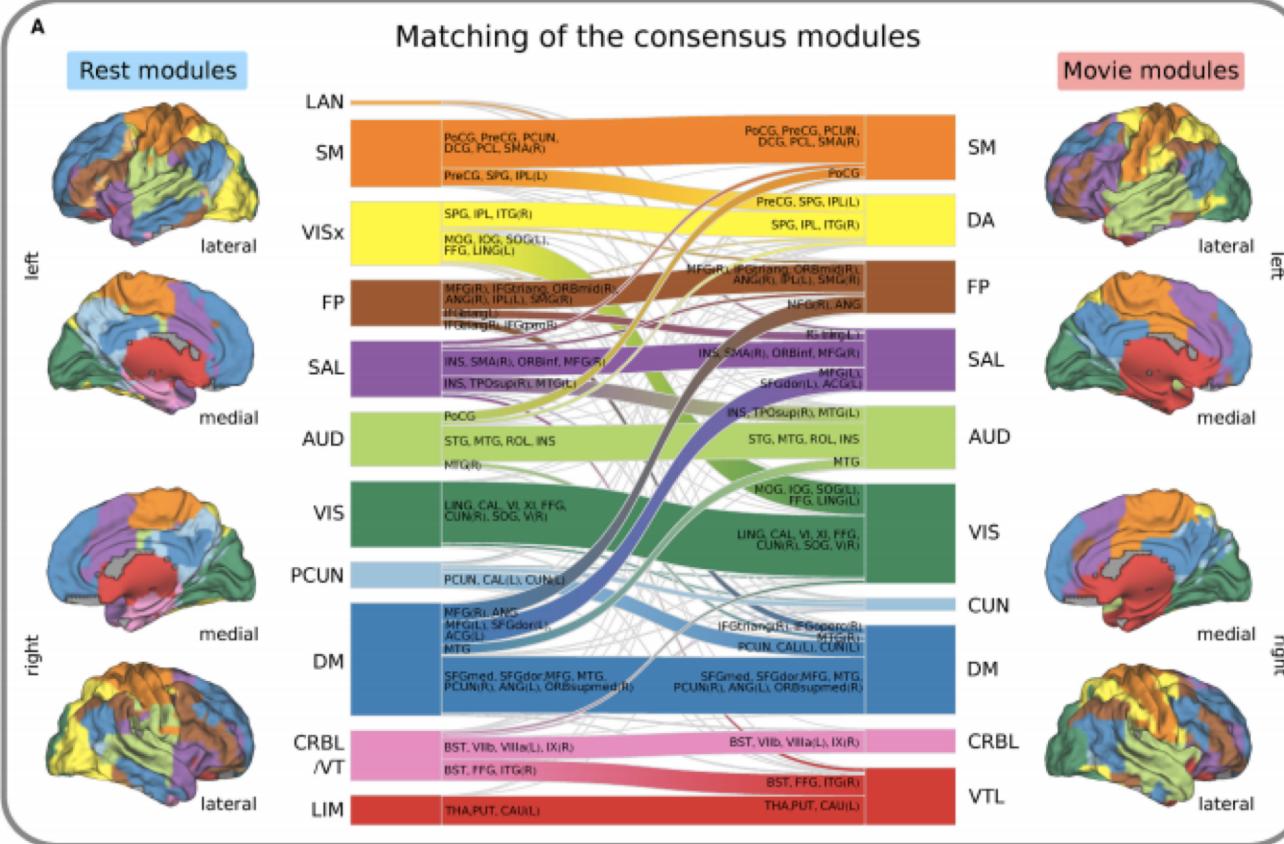
# Functional connectivity and graph theory (network science)



# Building a functional network

Repeat for all pairs of nodes and we get the full functional network





# Functional connectivity with naturalistic imaging 2

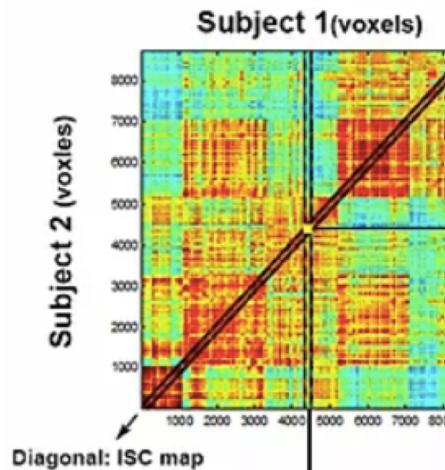
5) *intersubject functional  
connectivity*

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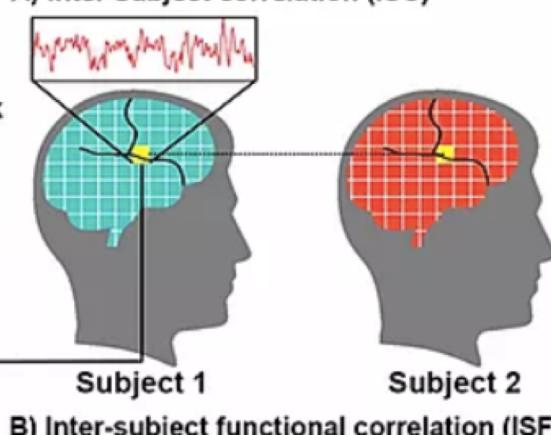
# Separating the functional network that is extrinsic (stimulus dependent) from the intrinsic one

Figure 1

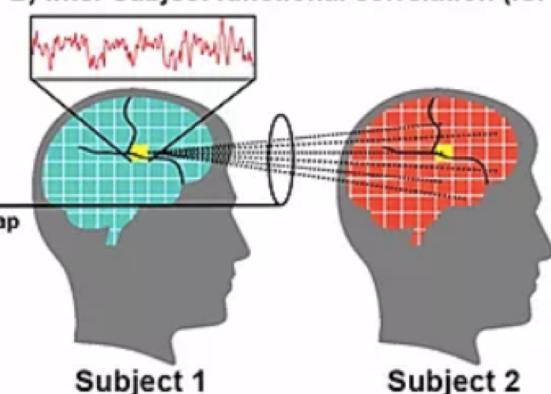
C) Inter-subject functional covariance matrix



A) Inter-subject correlation (ISC)



B) Inter-subject functional correlation (ISFC)



Simony E, Honey CJ,  
Chen J, Lositsky O,  
Yeshurun Y, Wiesel A,  
Hasson U (2016)  
Dynamic  
reconfiguration of  
the default mode  
network during  
narrative  
comprehension. Nat  
Commun 7.

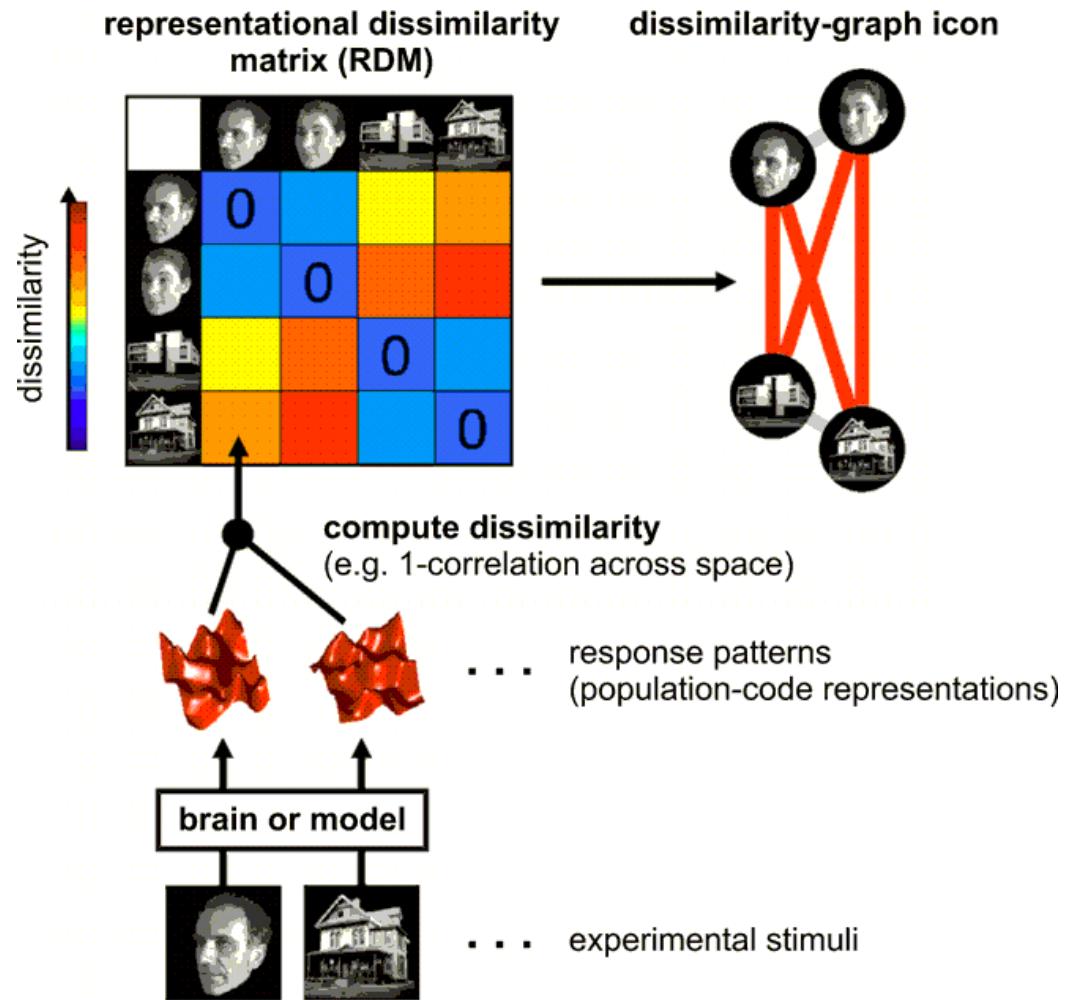
# Inter-individual differences

6) *intersubject representational  
similarity analysis*

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# Representational similarity analysis

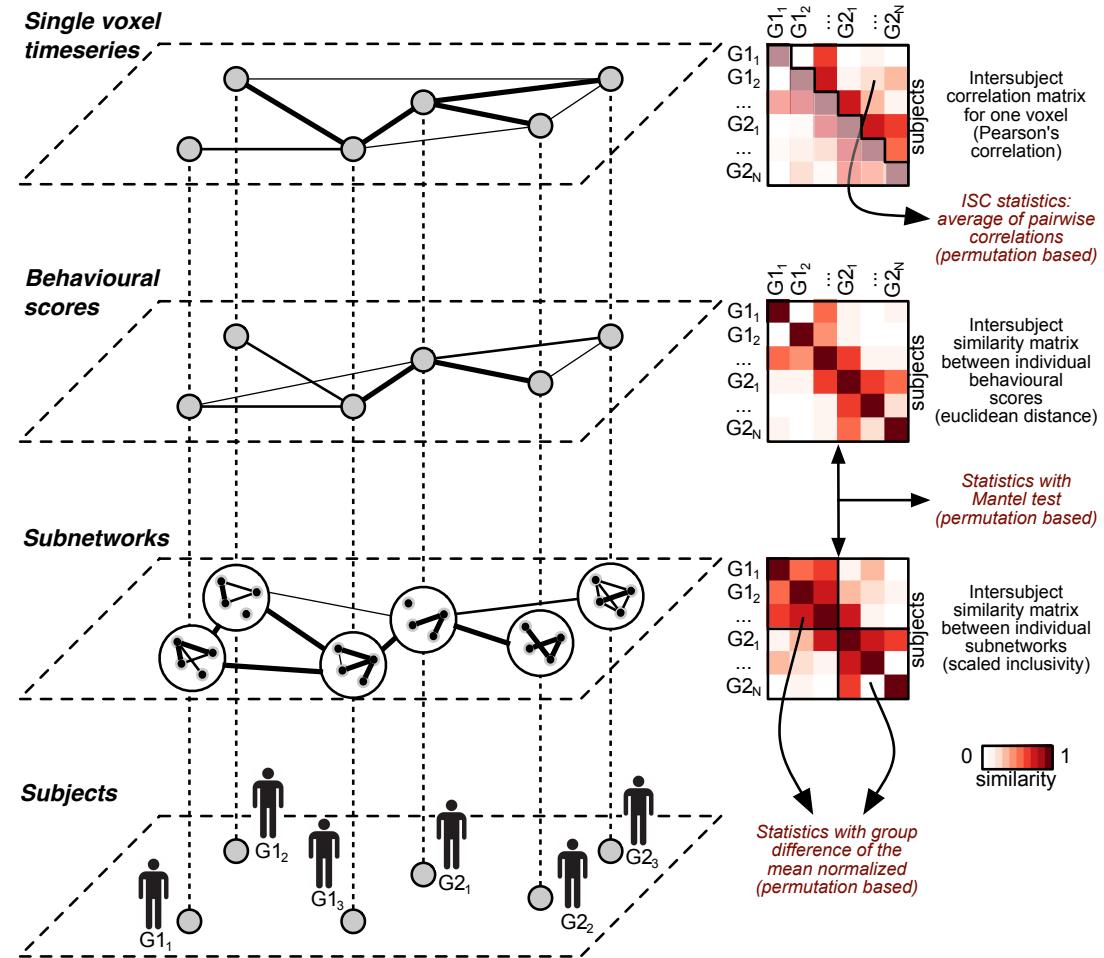
Instead of testing a one-to-one relationship between a stimulus model and brain response (GLM), RSA tests that the **similarity of brain responses** is related to the **similarity of the stimuli presented (Mantel test)**



[Kriegeskorte et al. 2008](#)

# Intersubject analysis framework

- **RSA** can be re-used for similarities across subjects
- **Mantel test** (comparison between similarity matrices)
- **Comparing within groups/conditions similarities**



[Glerean et al. 2016](#)  
[Finn, Glerean, et al. 2020](#)

...and statistics?

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- Statistical modelling
  - ***Statistical considerations***
  - ***Brain & behaviour modelling***
- Reflections and future directions

# Naturalistic paradigms – statistical considerations

**Embrace permutations!**

Due to the nature of ISC, **there is no data independence**.  
The only way to deal with statistical inference is to **adopt permutation based approaches for all statistical analysis**.

Some papers:

<https://pubmed.ncbi.nlm.nih.gov/27195792/>

<https://pubmed.ncbi.nlm.nih.gov/27751943/>

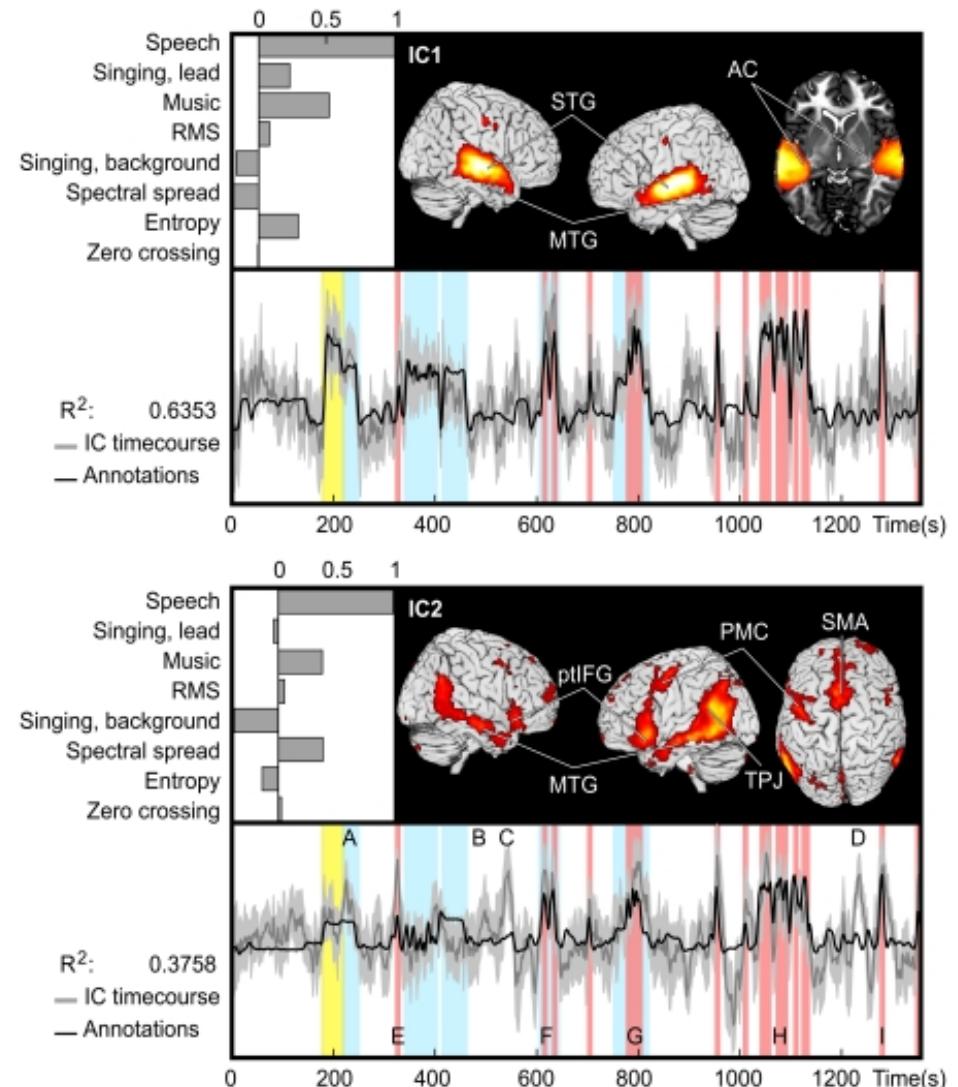
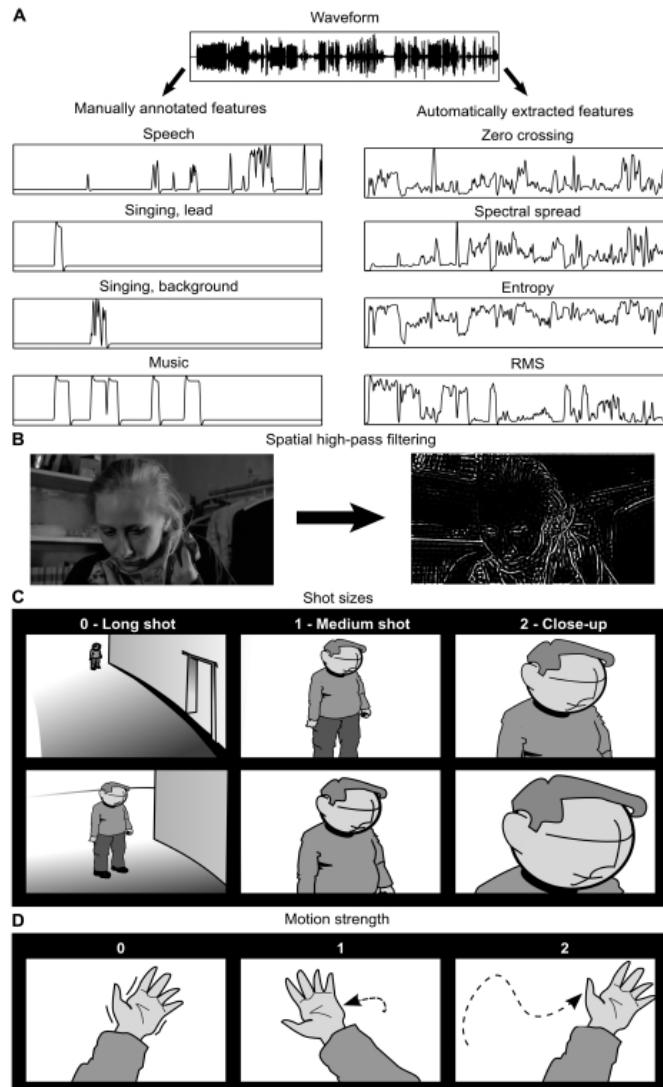
...and behaviour?

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# Model based analysis with automated feature extraction

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# Modelling of naturalistic stimuli and then using GLM



# Model based analysis with behavioural annotations

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**Please annotate the amount of Valence in time by using the mouse**

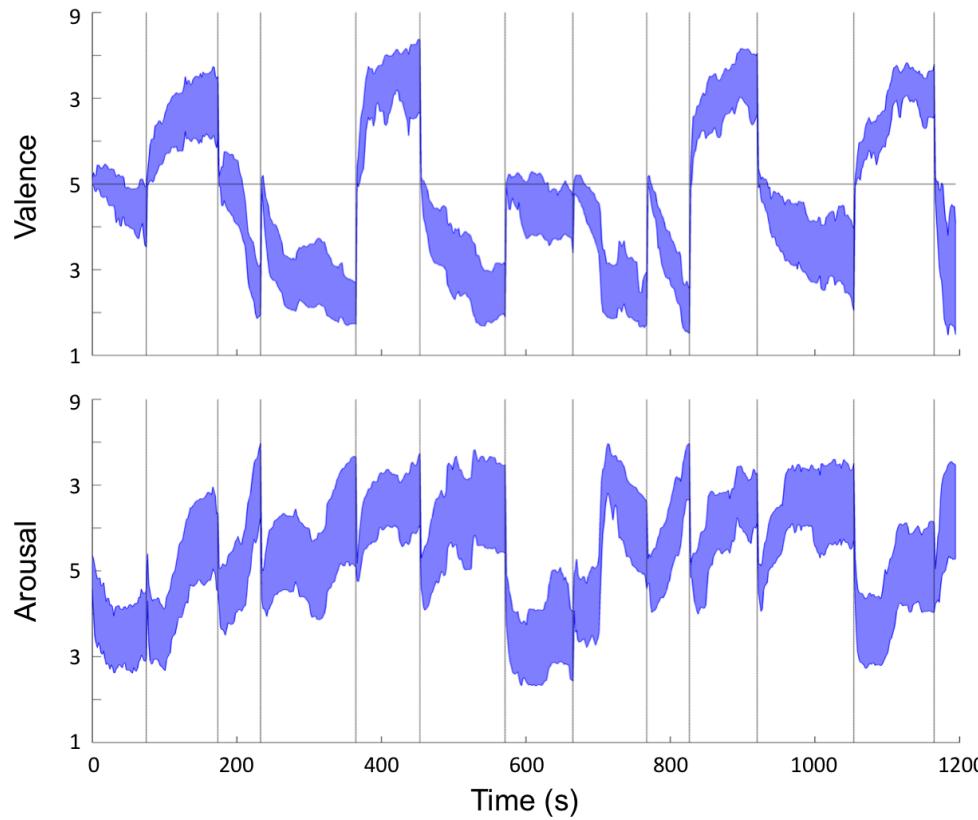
- + positive valence
- negative valence

1.9 of 91.5 seconds



# Participants' annotations used to model brain activity

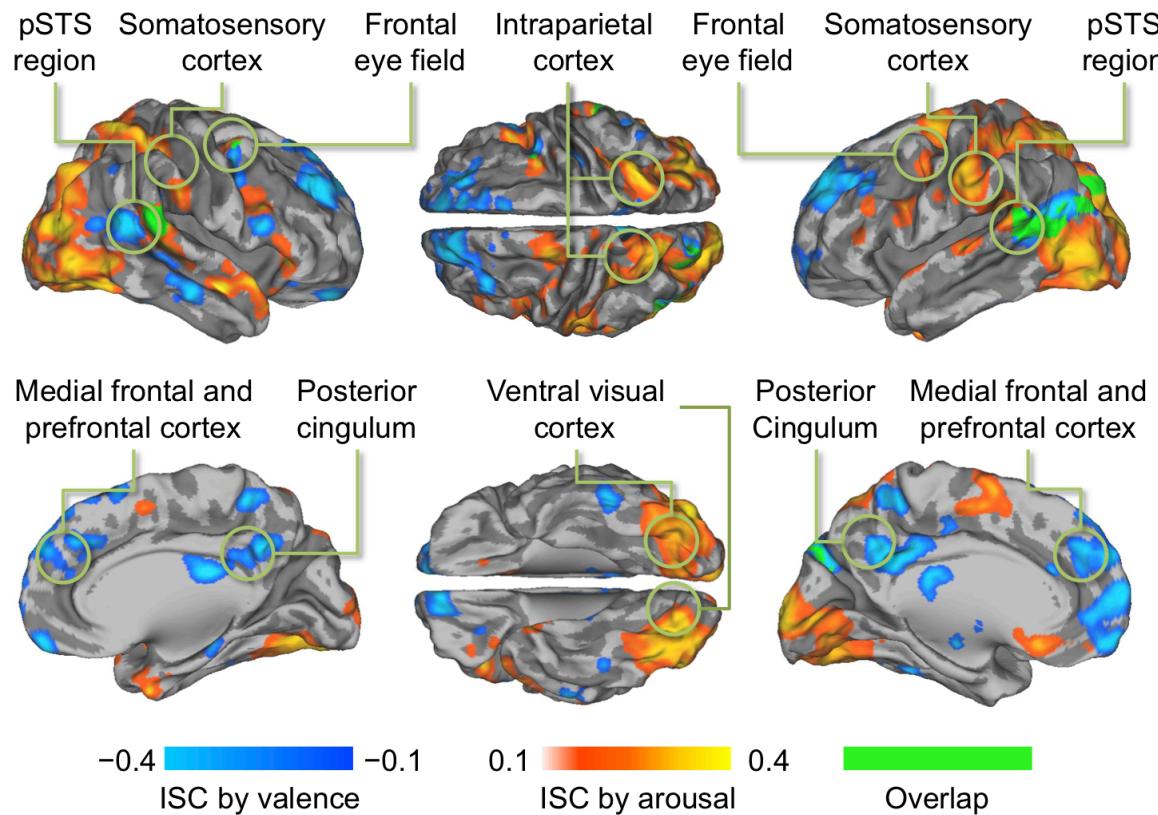
- Average valence/arousal



Nummenmaa, L., Glerean, E. et al. (2012). **Emotions promote social interaction by synchronizing brain activity across individuals.** PNAS 109(24), 9599–604.

# Participants' annotations used to model brain activity

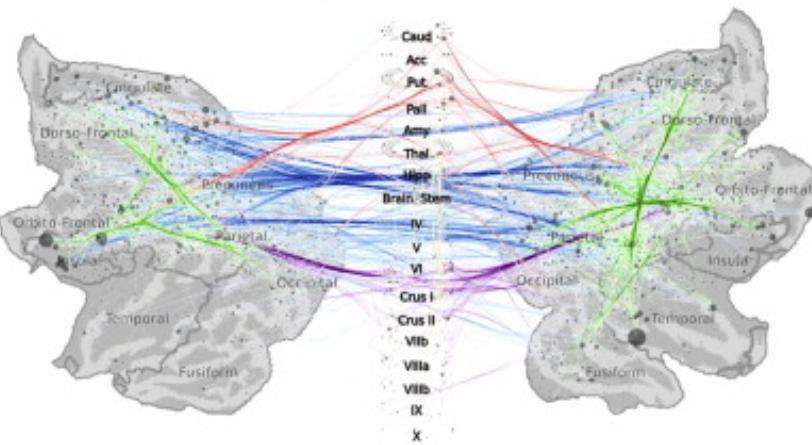
- Intersubject correlation modulated by valence/arousal



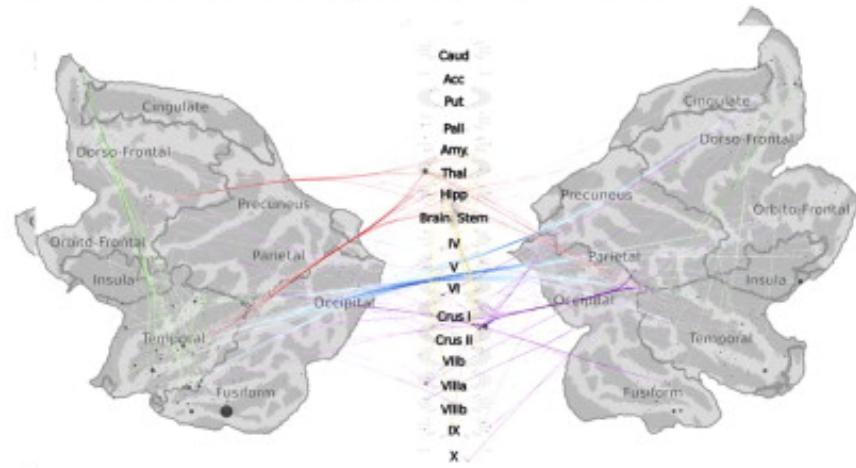
Nummenmaa, L., Glerean, E. et al. (2012). **Emotions promote social interaction by synchronizing brain activity across individuals.** PNAS 109(24), 9599–604.

# Brain networks in emotions

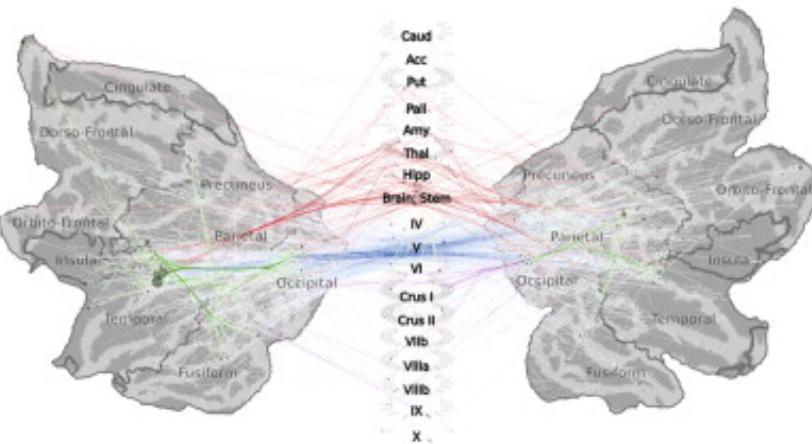
a) Enhanced connectivity due to negative valence



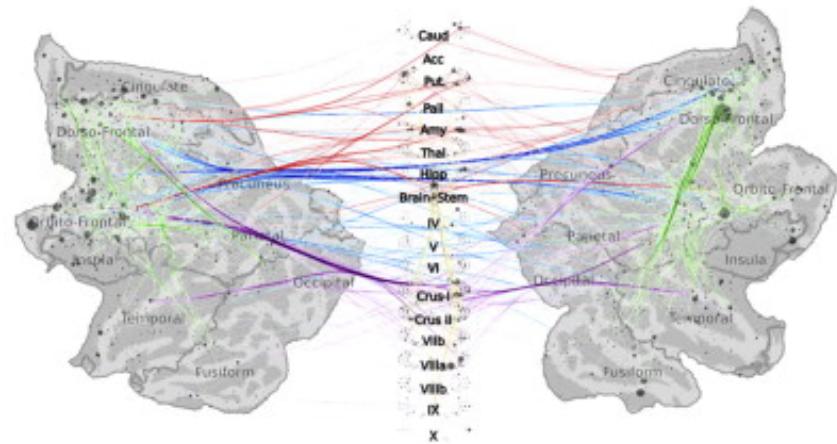
b) Enhanced connectivity due to positive valence



c) Enhanced connectivity due to negative arousal



d) Enhanced connectivity due to positive arousal



Node strength  
• • ● →  
Low High

■ Intrahemispheric cortico-cortical connections   ■ Cerebellar-cortical connections  
■ Interhemispheric cortico-cortical connections   ■ Other connections  
■ Subcortical-cortical connections

Nummenmaa, Saarimäki et al. (2014) Neuroimage

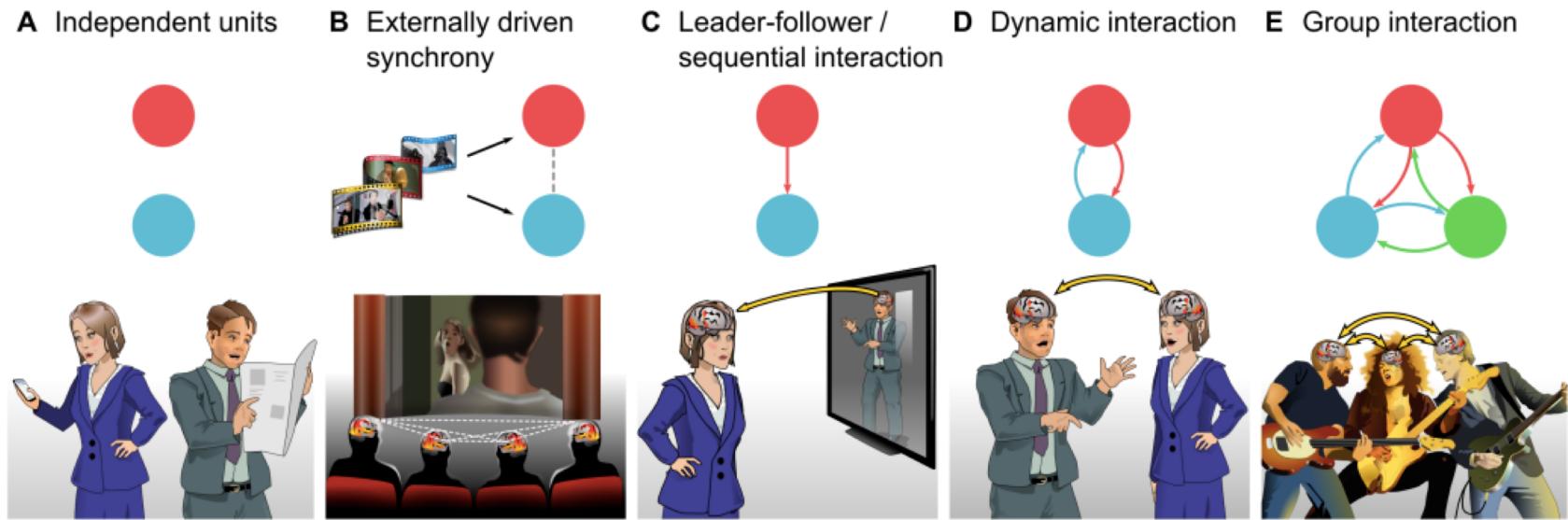
# Reflections and future directions

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# Reflections and future directions

More complex paradigms are being explored

Nummenmaa, Lahnakoski, Glerean 2018



More work when it comes to transparent and open methods [https://figshare.com/articles/Community-driven\\_methods\\_development\\_in\\_naturalistic\\_neuroscience/12366944/1](https://figshare.com/articles/Community-driven_methods_development_in_naturalistic_neuroscience/12366944/1)