From Information Flow to Connectomics

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Arrow of time in Physics and in biology?

2. Function and Structure

- Natural correspondence between structural networks and effective networks.
- Physiological reliability.

3. Microconnectome

- Beyond simple statistical properties
- As one many body problem
- Nodes are generally neurons
- The detailed design of the network organization
- Degree, Hub, Cluster, Community, Rich Club

1. Causality and Information flow

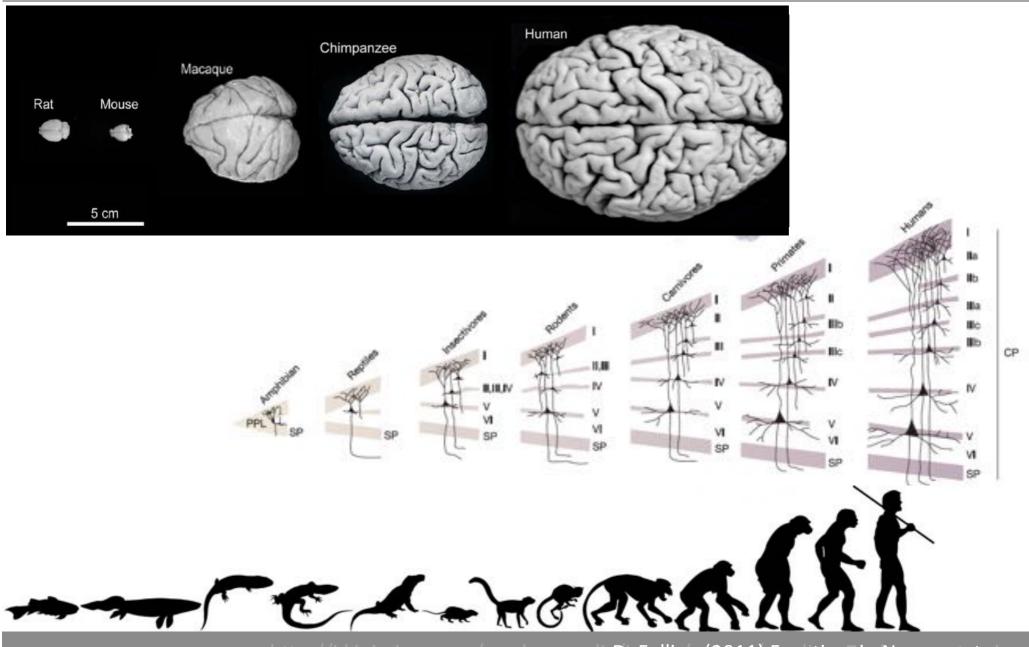
Arrow of time in Physics



Entropy

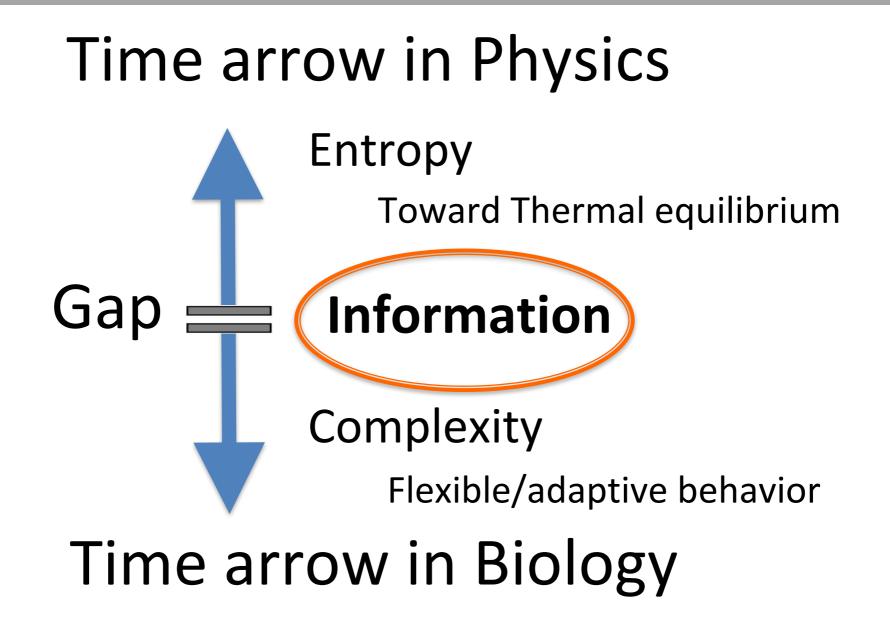
http://i.kinja-img.com/gawker-media/image/upload/ynzdjeav7eakmgzamtpe.jpg

Arrow of time in Biology



http://i.kinja-img.com/gawker-mediaDeFellipe (2011) Frontiers in Neuroanatomy

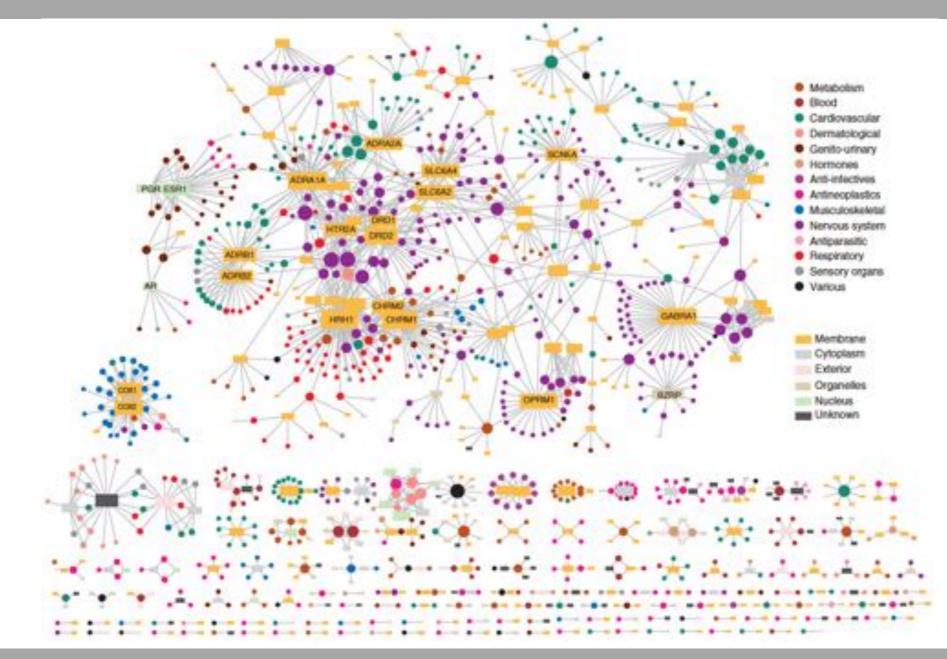
Causality



Causality

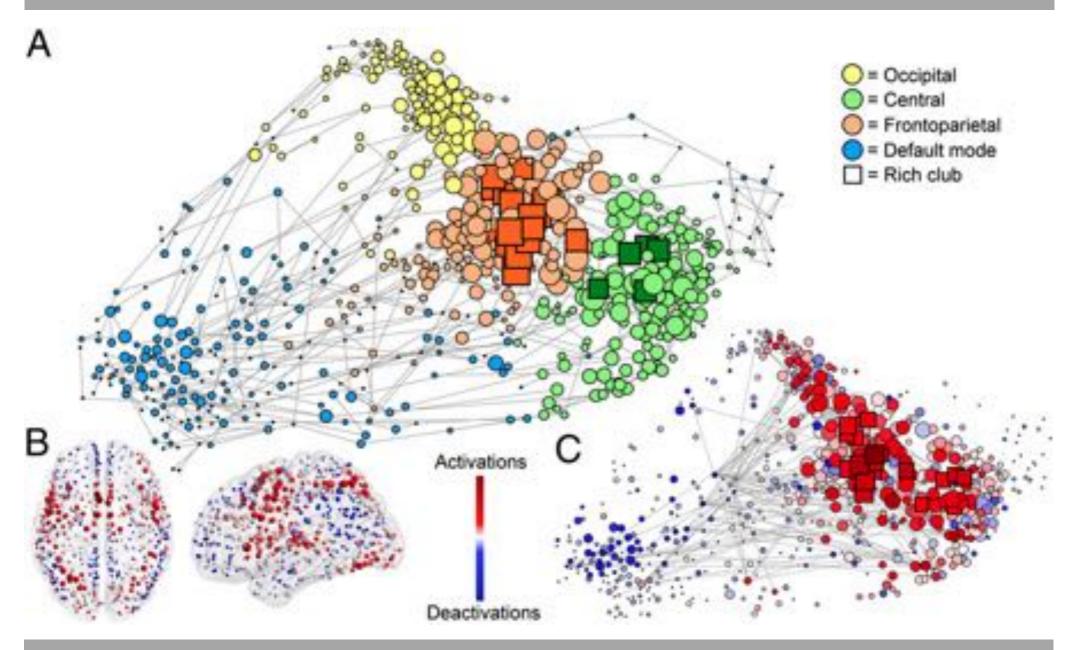


Causalities in biological system is intertwined



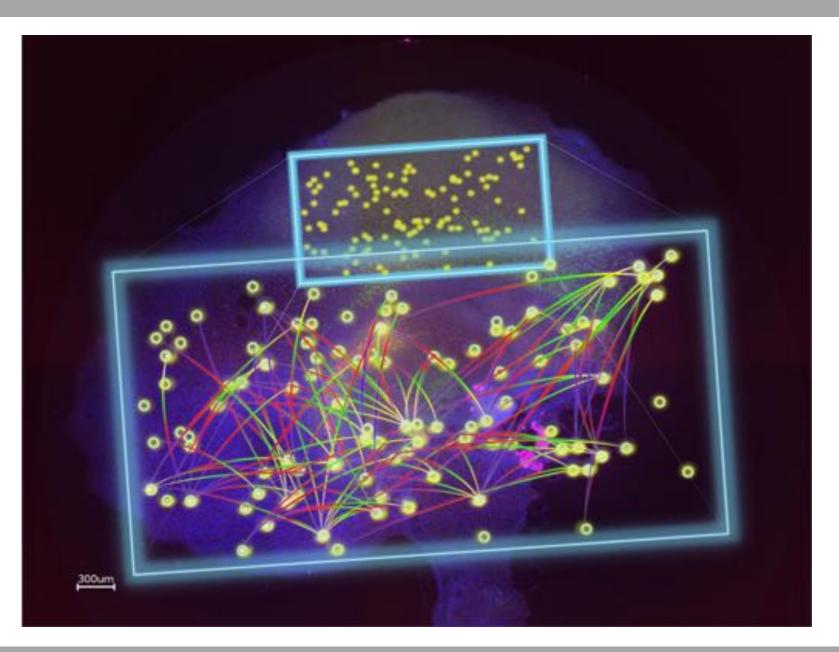
Yildirim et al. (2007) *Nature Biotechnology* – Drug-target network

Causality in brain is complex



Crossley et al. (2013) PNAS

Causal interactions among neurons



2. The non-uniformity- Comparisons with structure -

Three types of networks

Structural networks

A set of physical or structural (anatomical) connections linking neural elements (Cajal, 1905; Fellman and Van Essen, 1991).

Functional networks

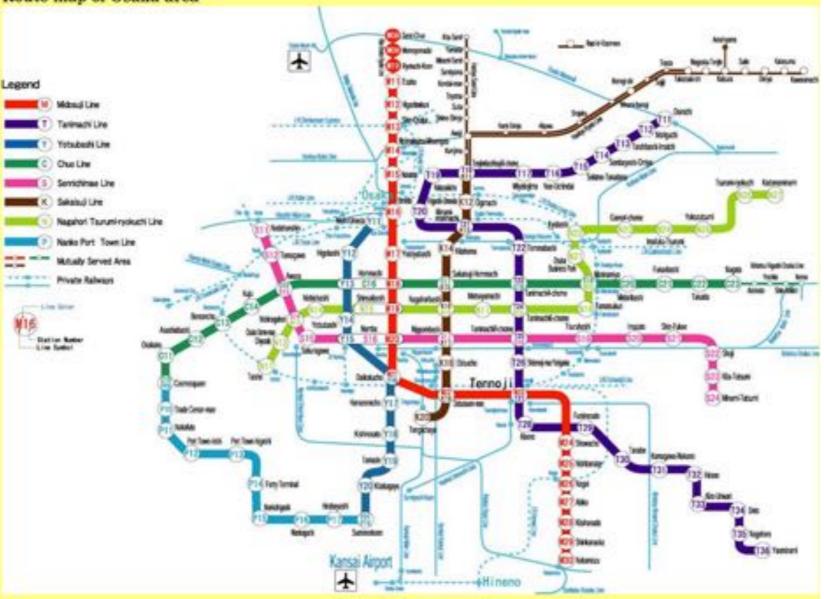
Deviations from statistical independence between distributed and often remote neuronal units (e.g. Gerstein and Perkel, 1969; Singer and Gray, 1995)

Effective networks

The networks of causal effects between neural elements (Adertsen et al., 1989; Friston, 1994)

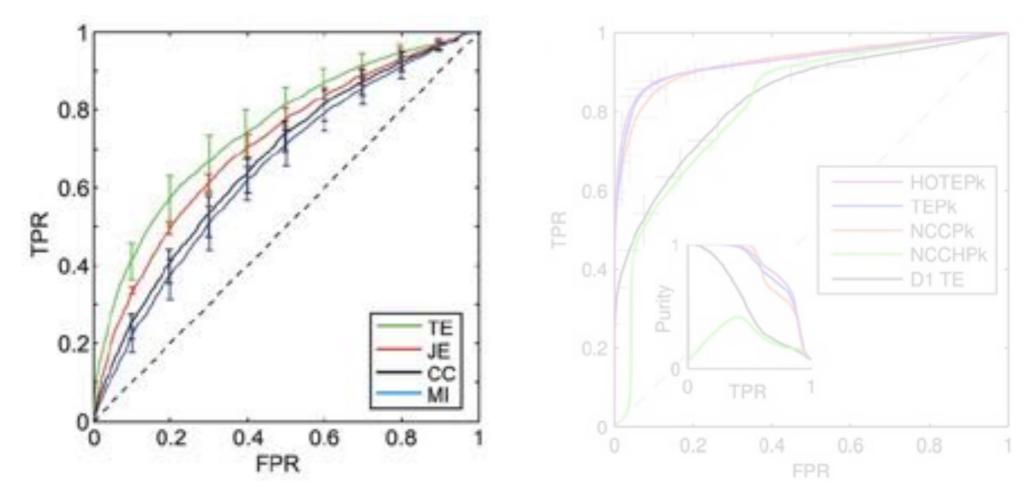
Structural connectivity and effective connectivity

Route map of Osaka area



Evaluation in computational models

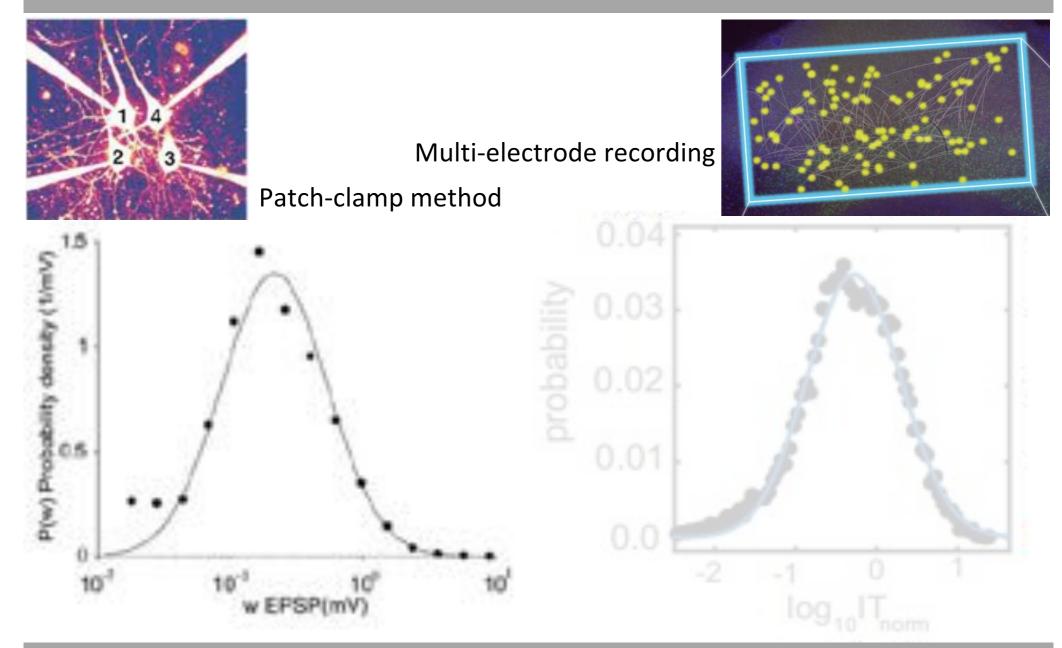
Predicting structural connectivity from effective connectivity



Transfer Entropy is the current champion.

Garofalo et al. (2009) PLOS ONE

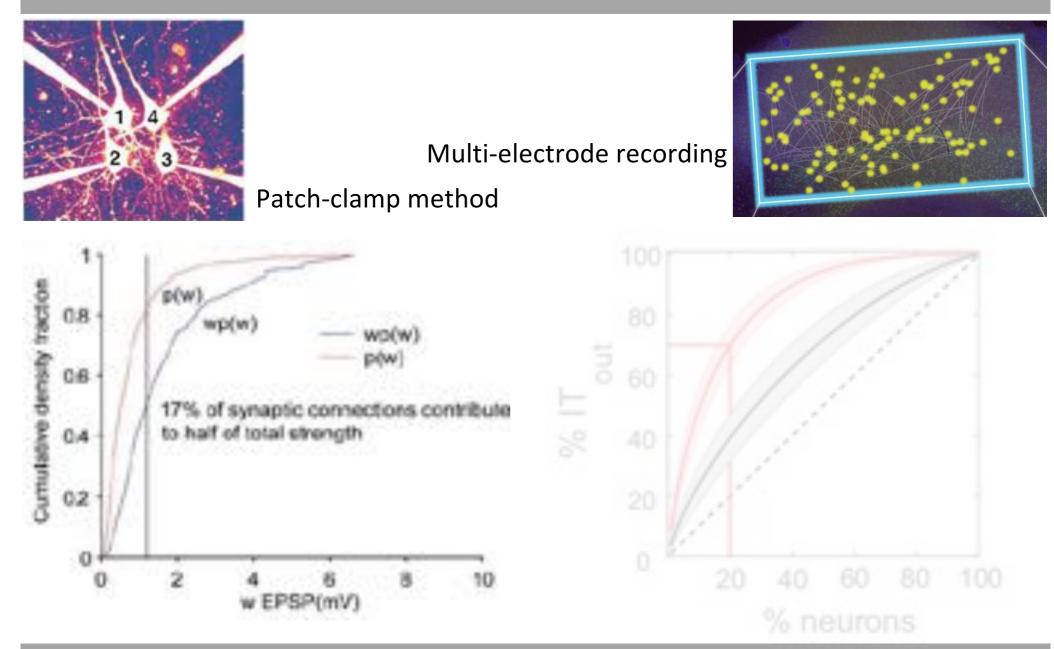
Complex network



Sunny, Shimono,,, Sporns, Beggs (2016)

Song et al. (2009) PLOS Biol.

Complex network



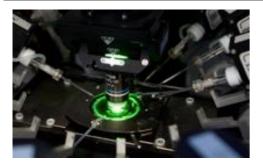
Song et al. (2009) PLOS Biol.

Sunny, Shimono,,, Sporns, Beggs (2016)

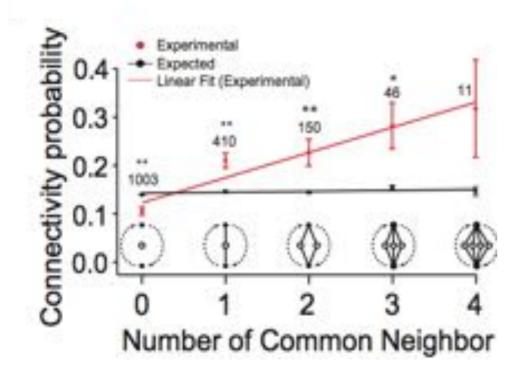
3. Microconnectome

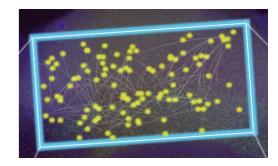
- Beyond simple statistical properties
- The detailed design of the network organization
- Nodes are generally neurons
- As one many body problem

Common neighbor effect

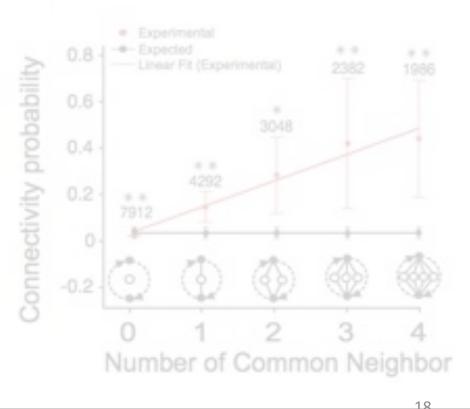


Patch-clamp experiment





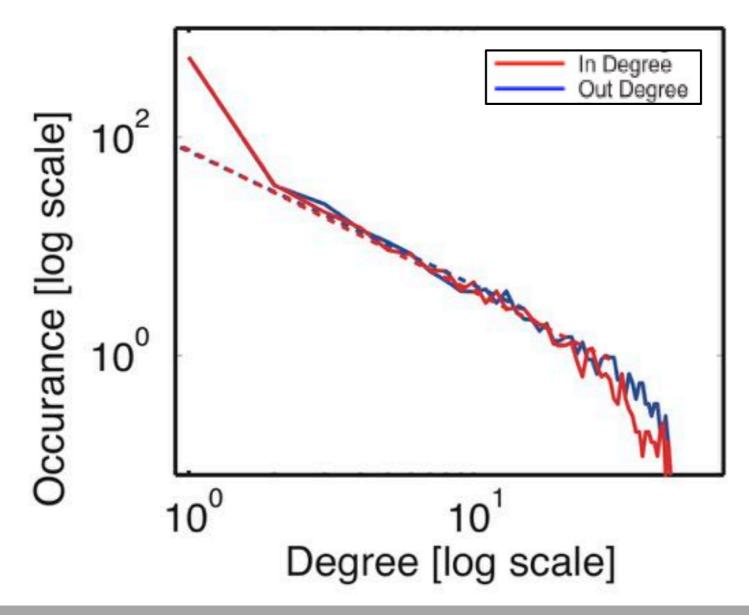
Multi-electrode array



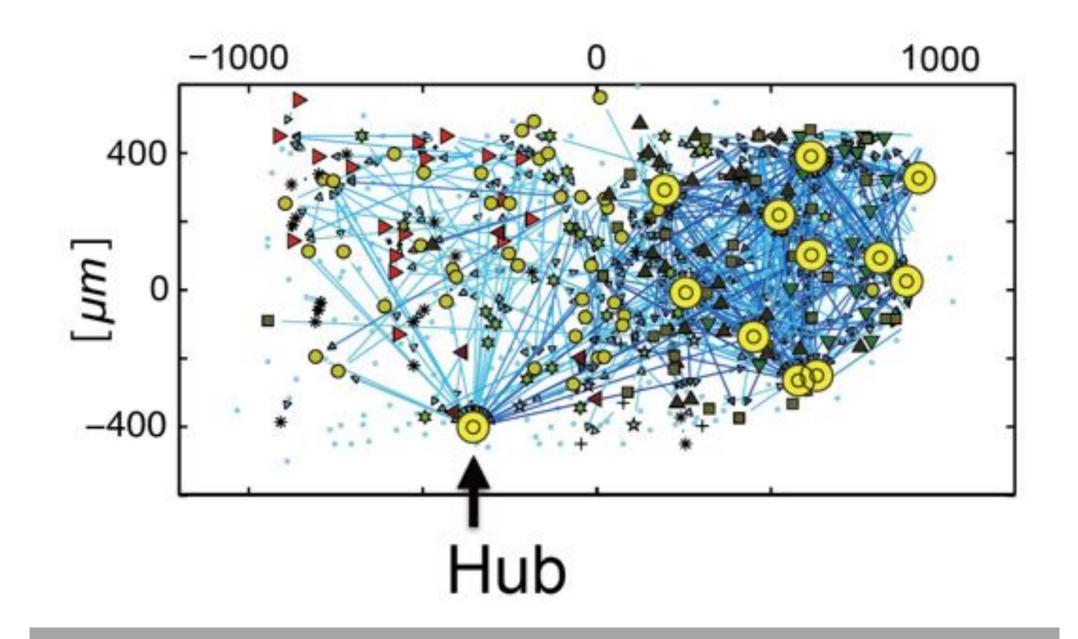
Perin, Berger, Malkram (2009)

Shimono, Beggs (2011)

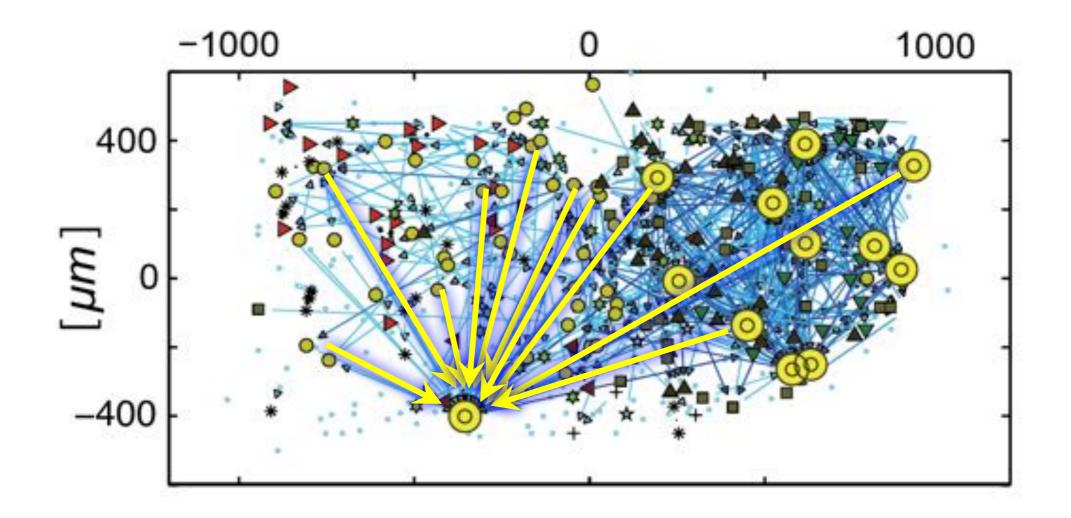
Degree histogram



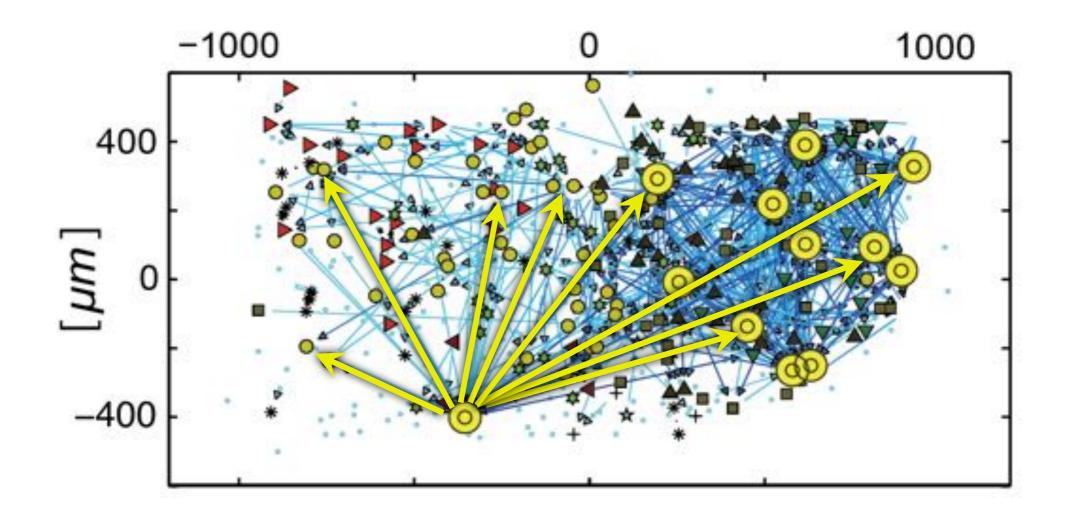
Hubs



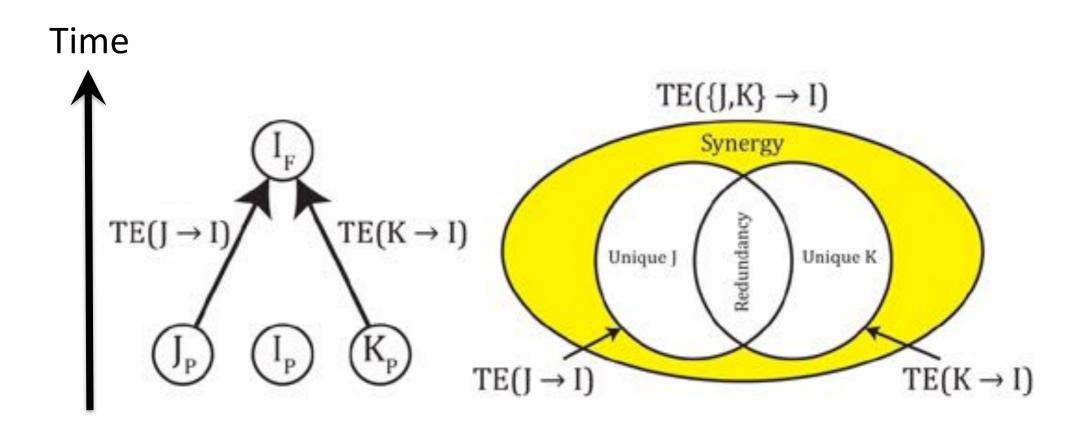
In-degree



Out-degree



Synergy

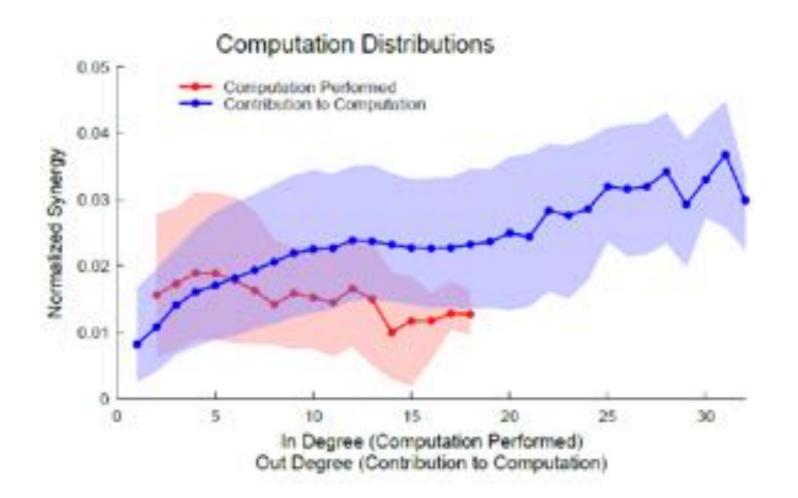


 $TE(\{J,K\} \to I) = Synergy(\{J,K\} \to I) + Unique(K; J \to I)$

 $+Unique(J; K \to I) + Redundancy(\{J, K\} \to I)$

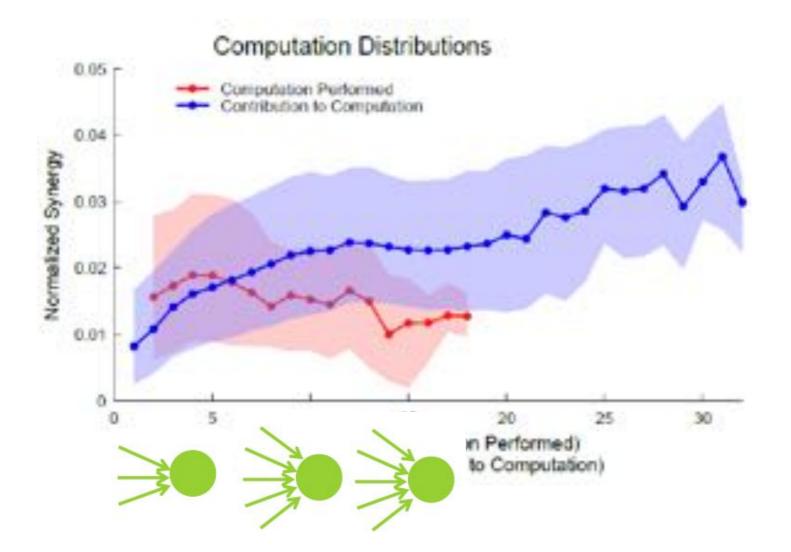
Timme et al. (2016) *PLOS Comput.Biol.*

Degree vs. Synergy



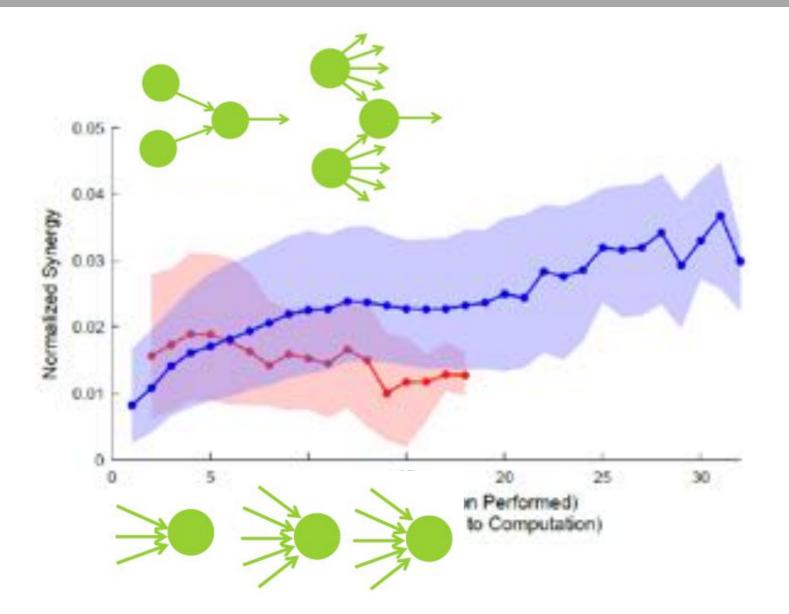
Timme et al. (2016) *PLOS Comput.Biol.*

Degree vs. Synergy



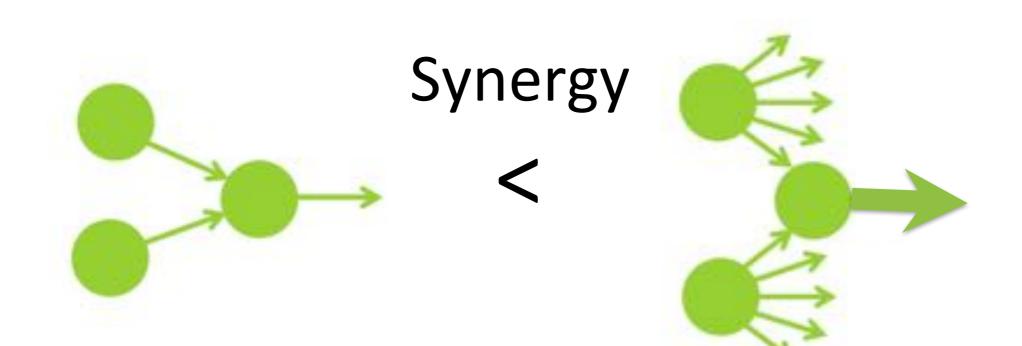
Timme et al. (2016) PLOS Comput. Biol.

Degree vs. Synergy



Timme et al. (2016) *PLOS Comput.Biol.*

Role of nodes receiving inputs from out-put hubs

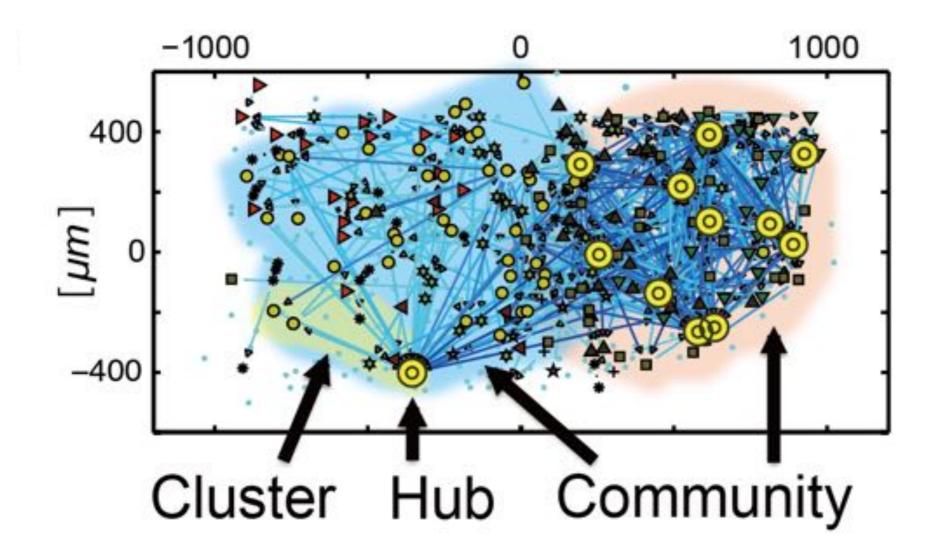


Inputs from less out-degree nodes

Inputs from more out-degree nodes

Timme et al. (2016) *PLOS Comput.Biol.*

Community



Question

50~100 neuron scale

3-5 neuron scale

1 neuron scale

Question

Community

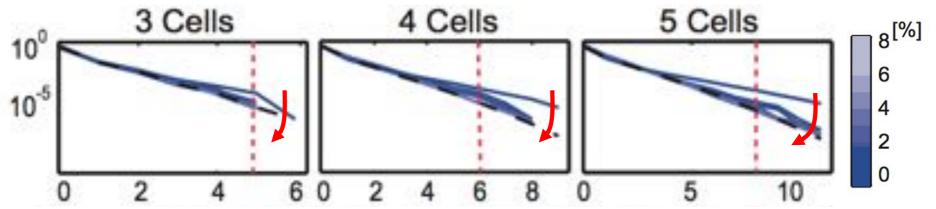
Cluster

Hub

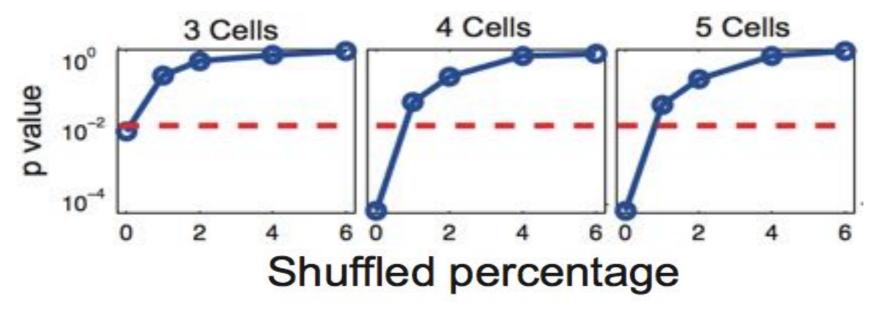
?

Influences of hubs on clusters

Effects of swapping on clusters

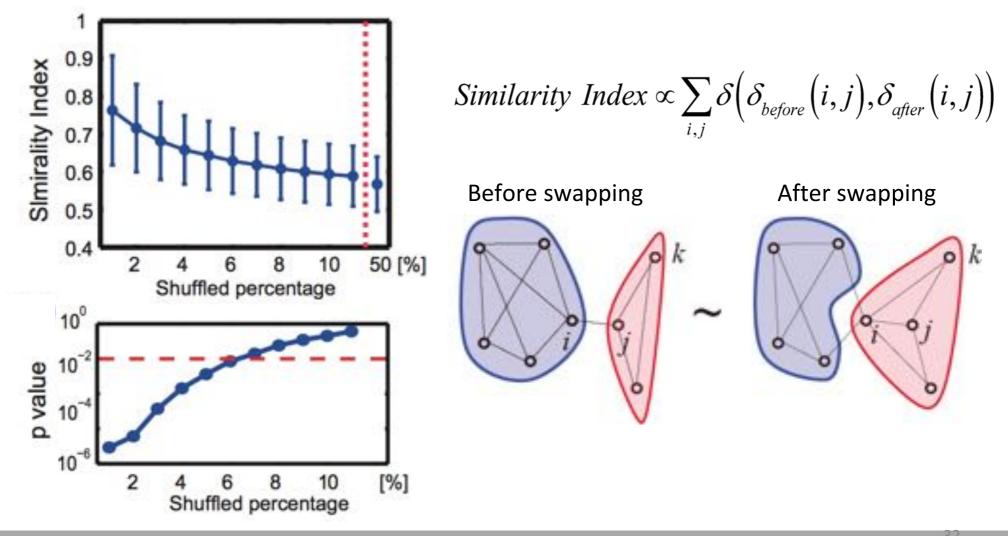


Number of Connections in Clusters



Influences of hubs on communities

Effects of swapping on communities



Summary

Community

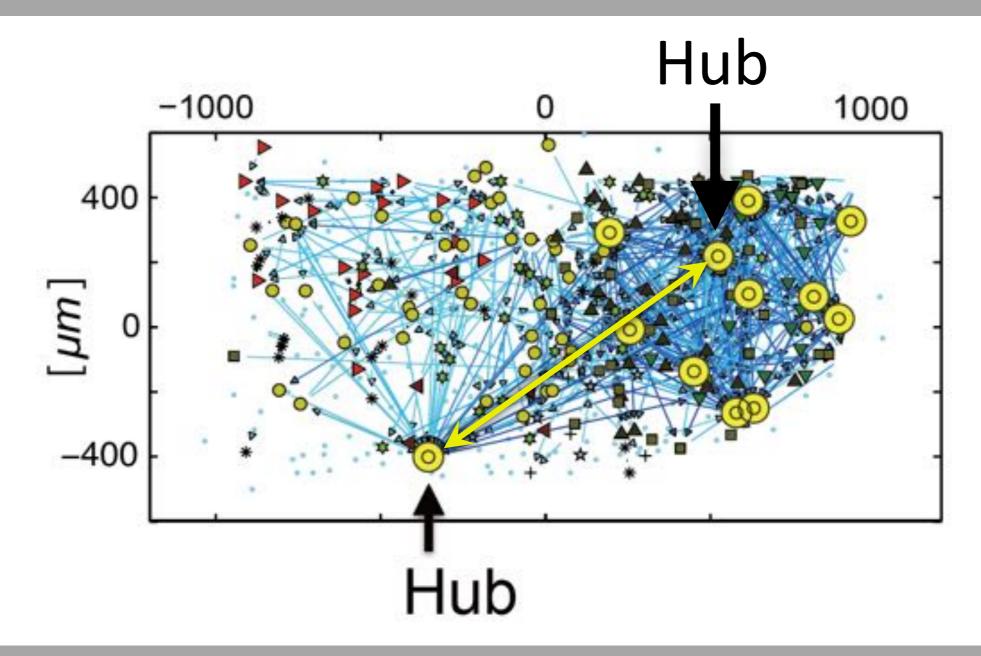
7%

Cluster

hub

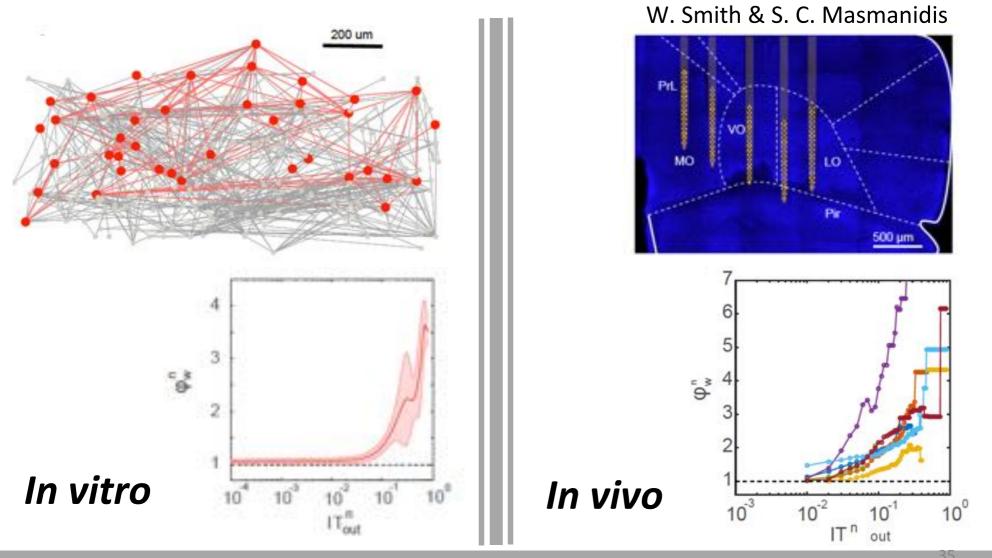
1%

Group of hubs



Rich club effect

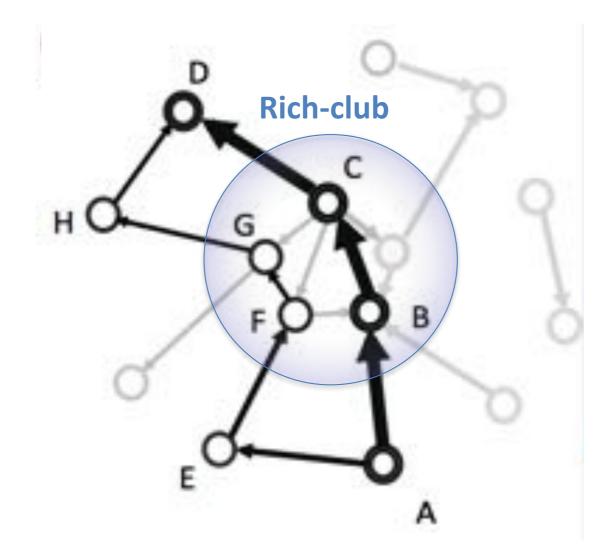
How do hubs connect each other?



Sunny Shimono,,, Sporns, Beggs (2016) J.Neurosci.

Rich club effect

How do hubs connect each other?



Sunny Shimono,,, Sporns, Beggs (2016) J.Neurosci.

Remarks

Basic statistical properties

- Lognormal weights
 - 20% nodes govern 70% information.
- Long-tailed distribution -> Scale-free?

Microconnectome

- Hubs exist in informatic Microconnectome.
- Hubs are surrounded by multiscale structures.

→ Clusters, Communities

- Group of hubs produce Rich-club.
- Commonly driven nodes by hubs gain information

 \rightarrow Synegetic non-hub nodes

