

# Flexible Computation in Neuronal Networks



Christoph Kirst

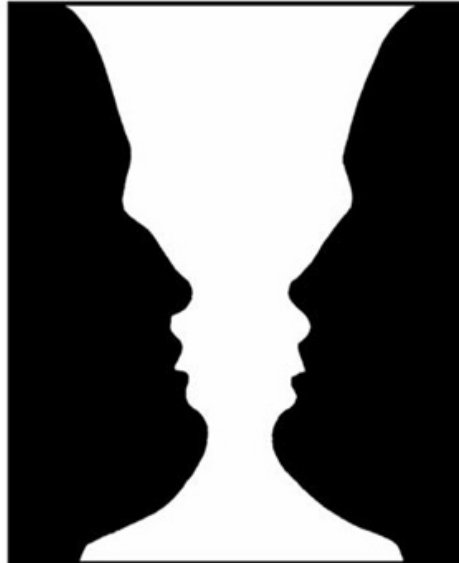
Fellow for Physics and Biology, Center for Theoretical Studies

Kavli Fellow, Kavli Neural Systems Institute

The Rockefeller University, New York City



# Brains dynamically route and process information

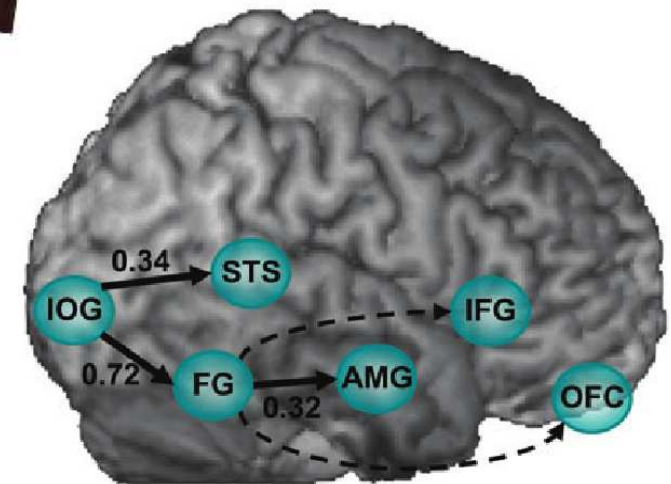


- How does the brain flexibly control information flow and processing?
- What are potential mechanisms ?

# Face Recognition is a Neuronal Network Task



[Steeves et al, Neuropsychologica 2006]



[Fairhall Ishai, Cerebral Cortex 2007]

- Functional brain network necessary for higher level face recognition

# The Face Recognition Network is Context Dependent

IOG: inferior occipital gyrus - core system

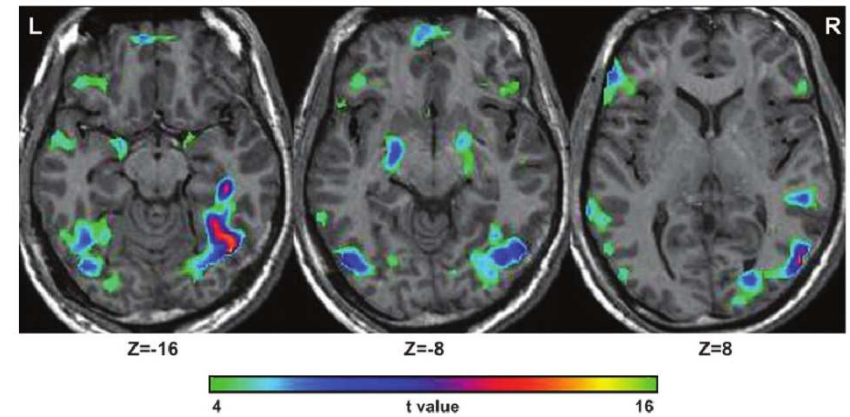
IFG: inferior frontal gyrus - semantic aspects

OFC: orbitofrontal cortex - facial beauty

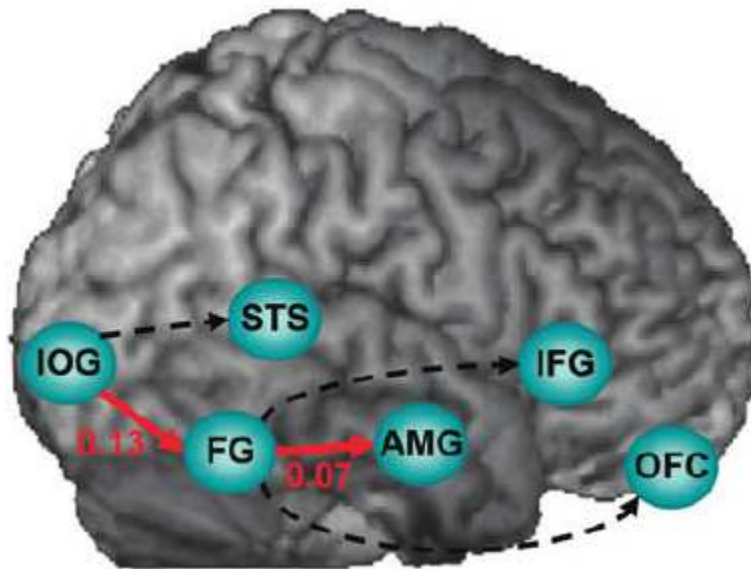
FG: fusiform gyrus - identification of individuals

AMG: amygdala - emotions

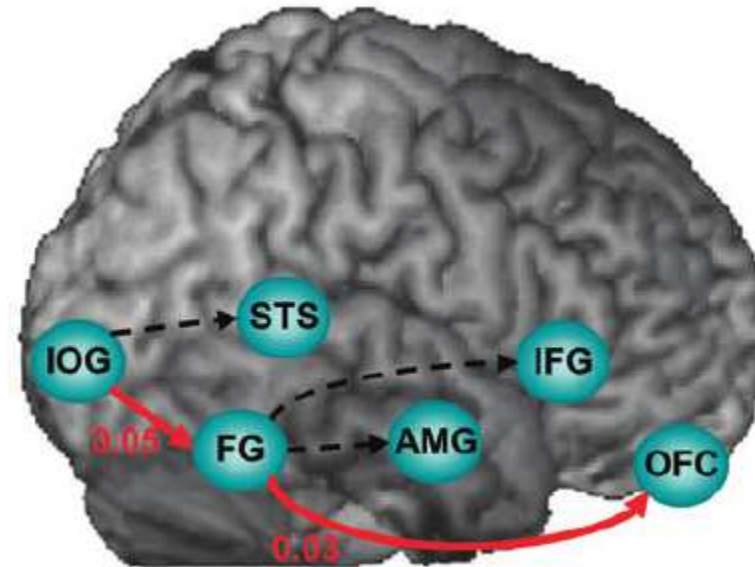
STS: superior temporal sulcus - gaze direction



emotional faces

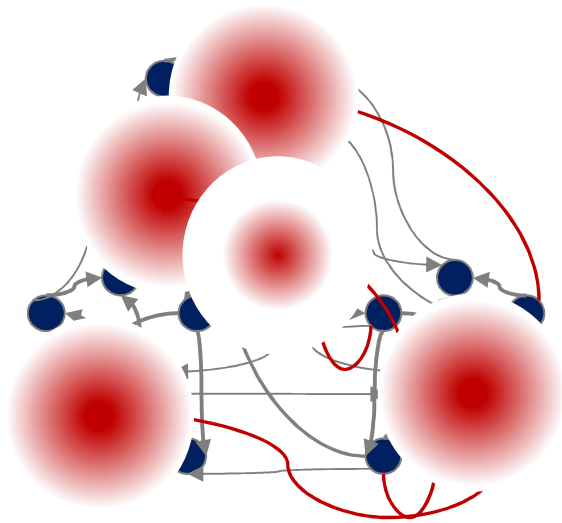
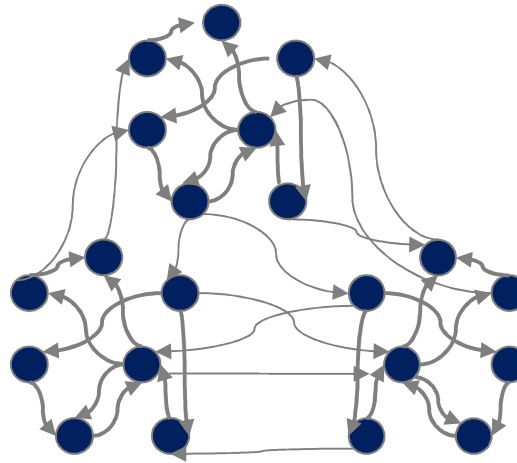


famous faces

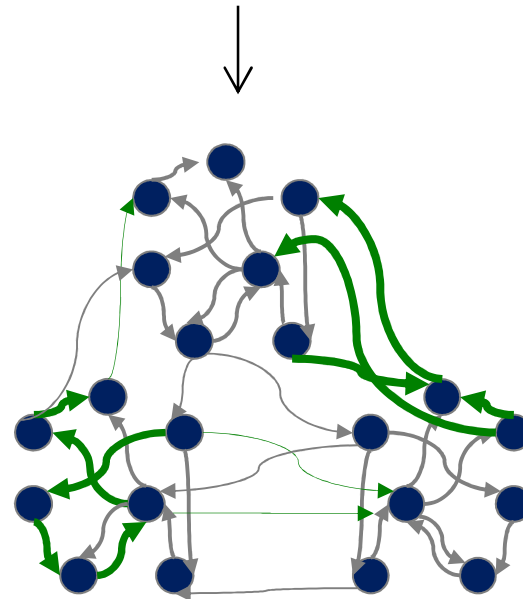


[Fairhall Ishai, Cerebral Cortex 2007]

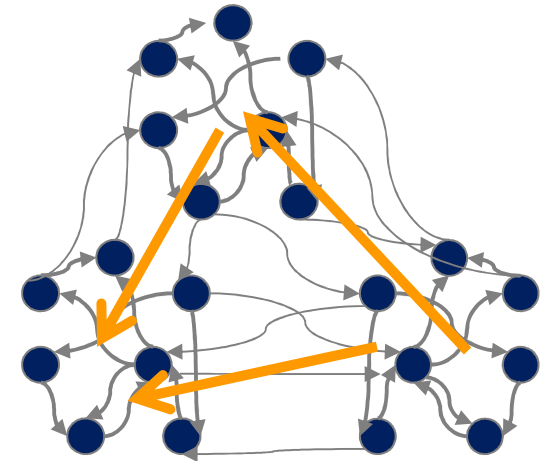
# Felxible Function of Neuronal Circuits



Neuro Modulation



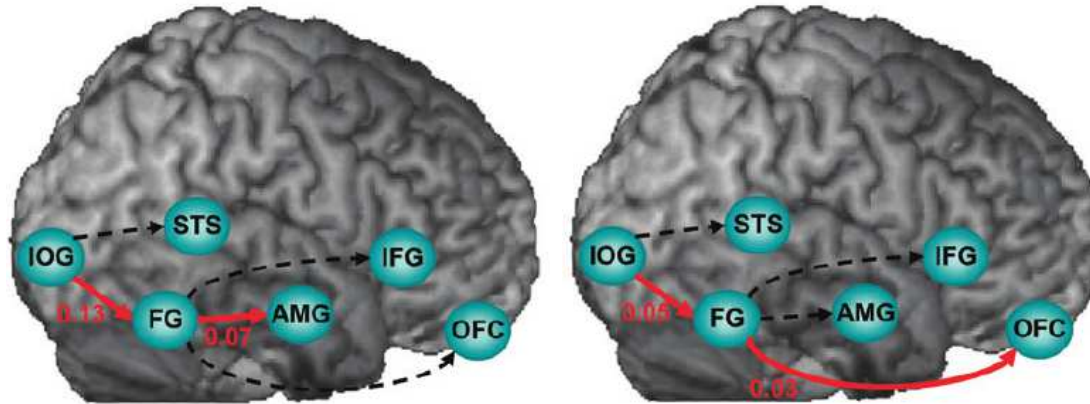
Plasticity



State Dependence

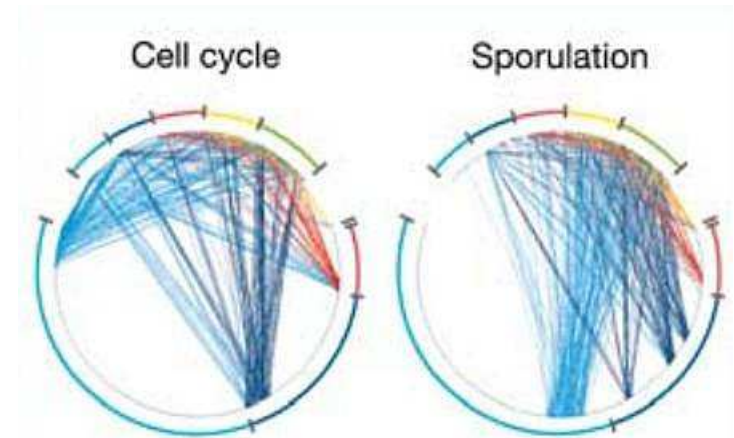
# Flexible Communication in Complex Networks

- Biological Networks: Neuronal networks



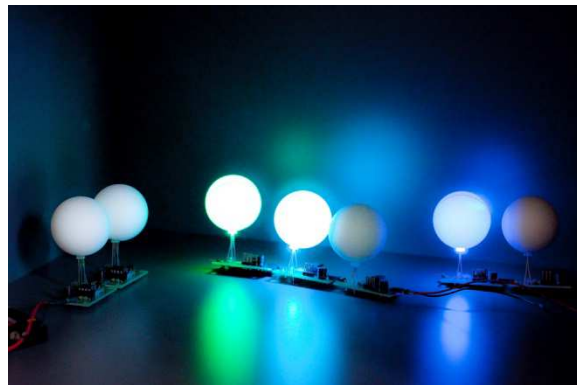
[Fairhall Ishai, Cerebral Cortex, 2007]

- Gene regulatory circuits



[Luscombe, et al. Nature 2004]

- Artificial Networks: Self-organizing distributed sensor networks

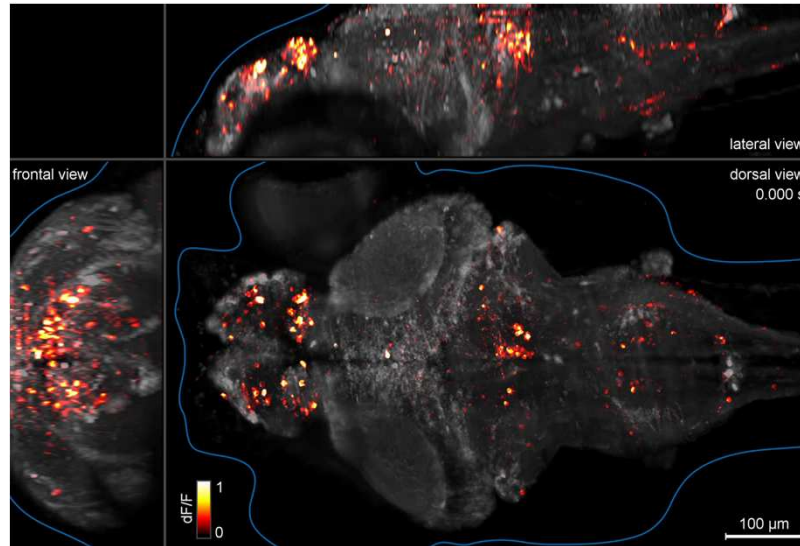


[Klinglmayr, Kirst, Timme, Bettstetter, 2012,  
Klinglmayr, Bettstetter, Timme, Kirst IEEE TAC, 2016]

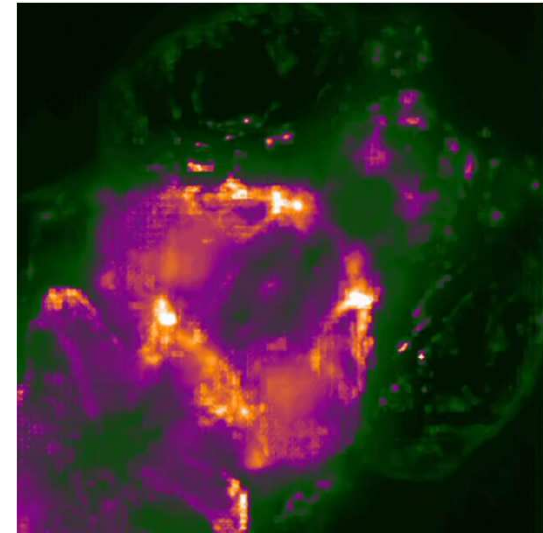


Mechanisms for  
flexible communication ?

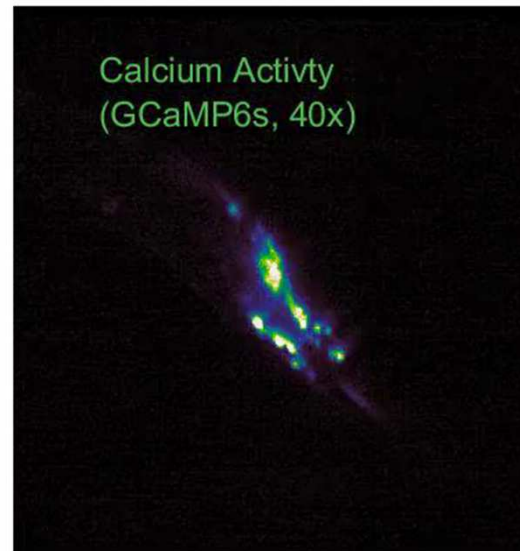
# Collective Dynamics in Neuronal Networks



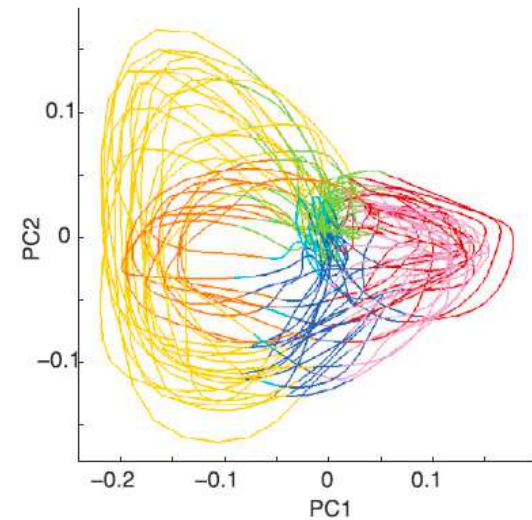
[Vladimirov, ..., Ahrens, Nat Meth 2014]



[Prevedel, ..., Vaziri, Nat Meth 2014]



[Nguyen, ..., Leifer, PNAS 2014]



[Kato et al, Cell 2015]

# Collective Neuronal Oscillations

## Über das Elektrenkephalogramm des Menschen.

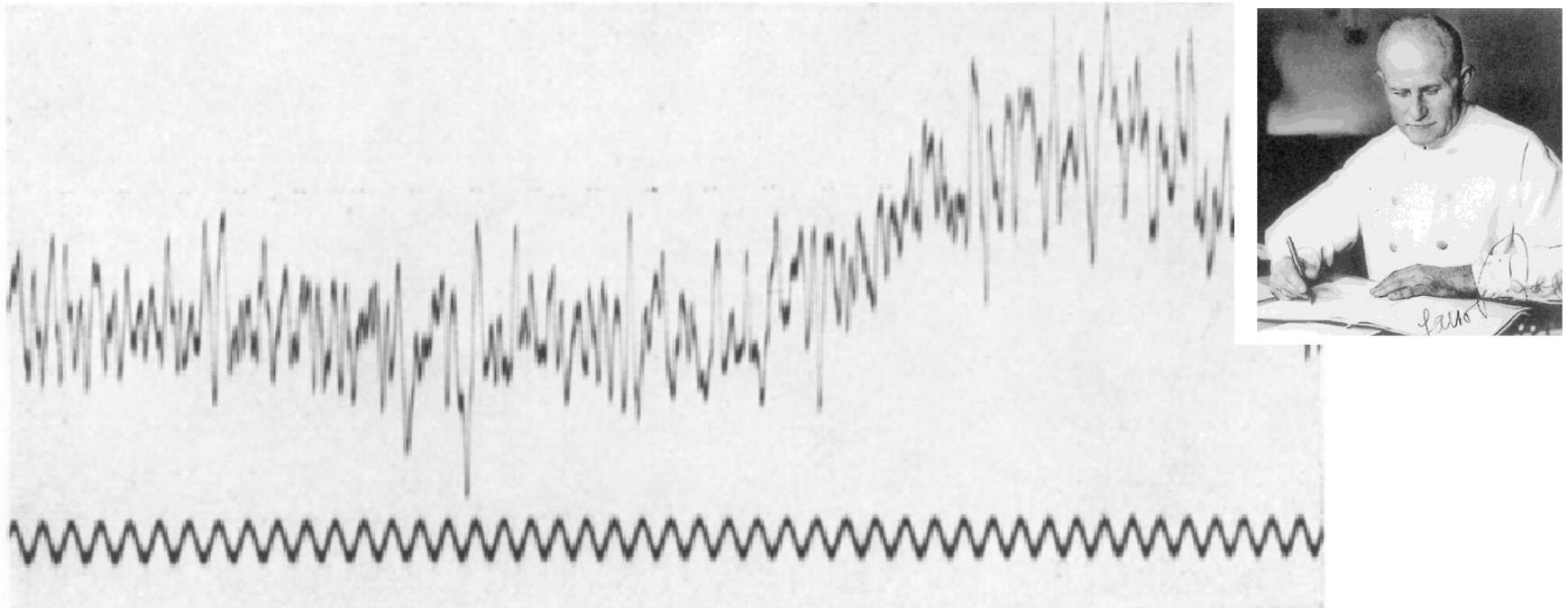


Abb. 2. W. H., 37 Jahre alt, an Dementia paralytica leidend. E.E.G. bei Nadelableitung von der Rinde des linken Stirn- und des rechten Scheitellappens. Zeit in  $\frac{1}{10}$  Sek.

[Berger, Arch Psychiatr Nervenkr. 1929 – 1938 (14 papers + 1 book)]

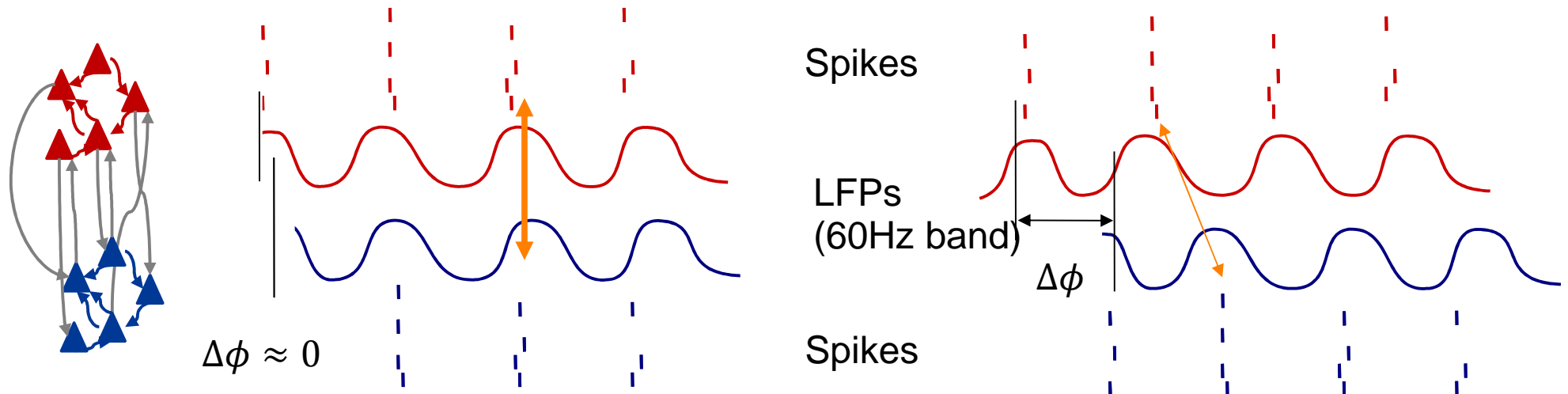
→ Epi-phenomenon or functional property ?



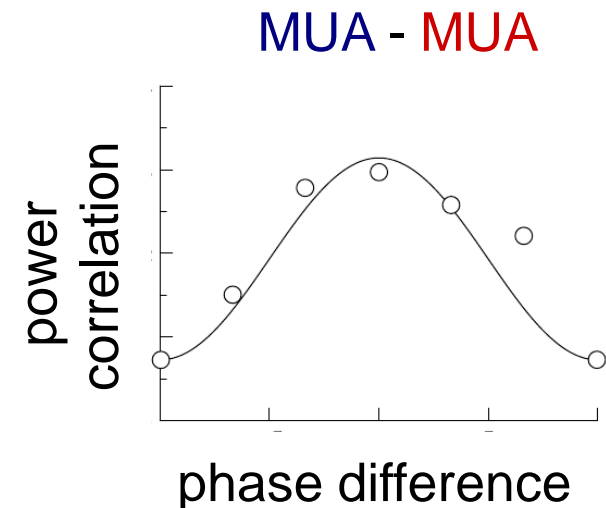
# Information Flow and Neuronal Oscillations

- communication through coherence hypothesis

[Fries, TICS, 2005]



[Womelsdorf et al. Science, 2007]



# Outline: Flexible Function in Neuronal Networks

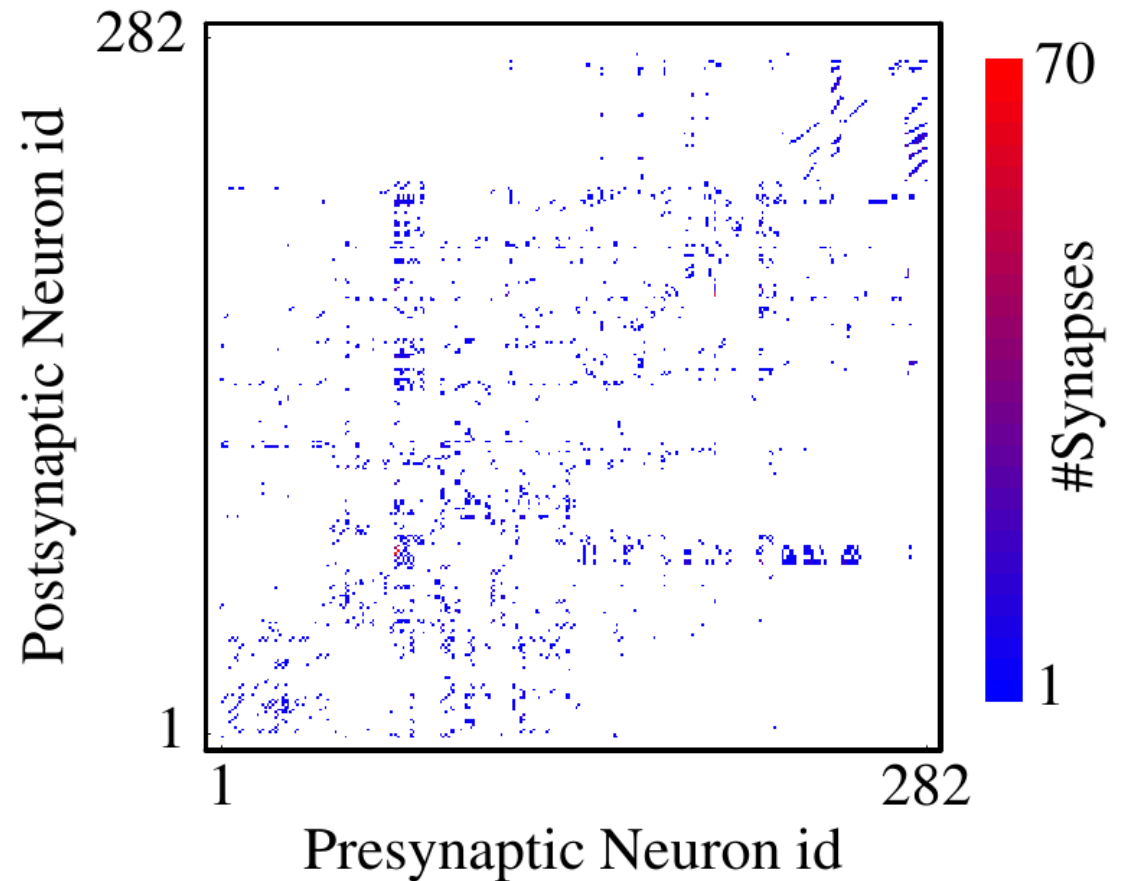
- dynamic information routing in complex networks
  - structural vs. effective connectivity
  - flexible information routing in oscillator networks
  - spiking networks, transient dynamics
- flexible information processing in complex networks
  - oscillatory Hopfield networks
  - self-organized pattern recognition
- learning flexible function in neuronal networks
- connections to experiments
  - brain state identification in zebrafish
  - complete brain activity mapping in mouse
- conclusions

# Outline: Flexible Function in Neuronal Networks

- dynamic information routing in complex networks
  - structural vs. effective connectivity
  - flexible information routing in oscillator networks
  - spiking networks, transient dynamics
- flexible information processing in complex networks
  - oscillatory Hopfield networks
  - self-organized pattern recognition
- learning flexible function in neuronal networks
- connections to experiments
  - brain state identification in zebrafish
  - complete brain activity mapping in mouse
- conclusions

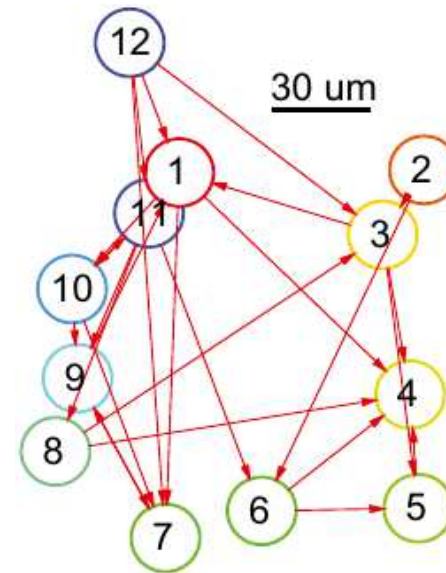
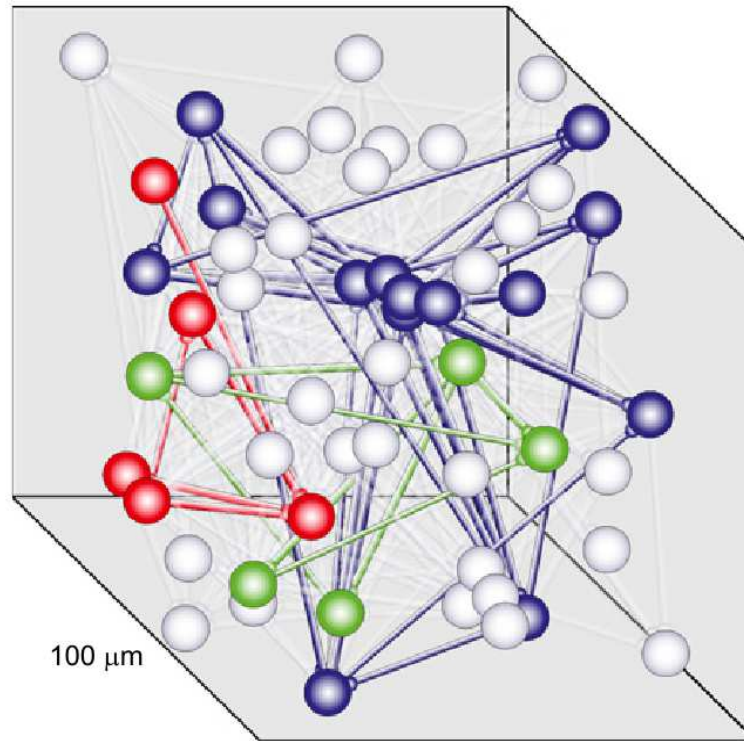
# Structural Connectivity

structural connectivity = physical wiring (e.g. via synapses)



[J. G. White, E. Southgate, J. N. Thomson, S. Brenner, 1986]

# Clustering / Modular Network Structure



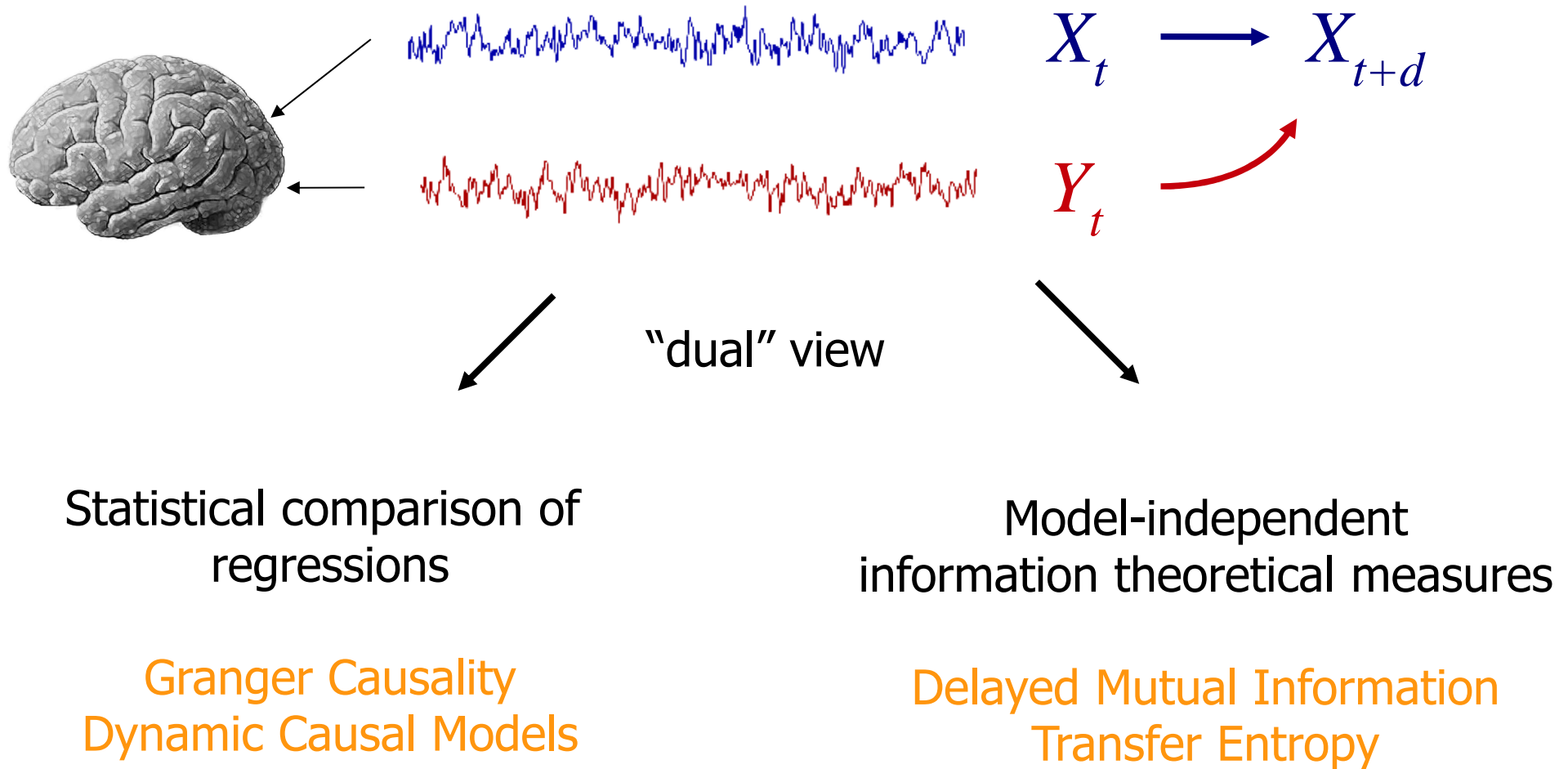
[Perin et al. PNAS 2010]

[see also: Song, .., Chklovskii, 2005]

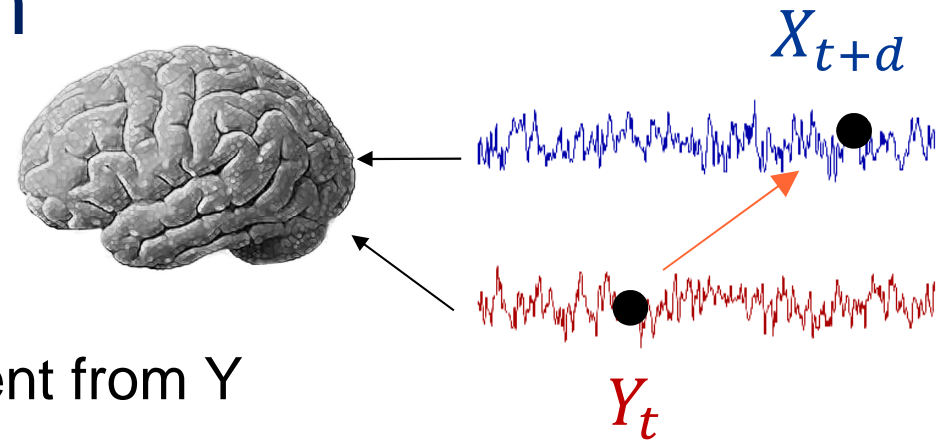
- interlaced strongly connected groups (color)  
on top of a sea of weak connections (gray)

# Effective connectivity

- functional connectivity as measured from dynamics
- focus on forecast: improvement of forecast of  $X_{t+d}$  by history of  $Y_t$



# Delayed mutual information



joint probability

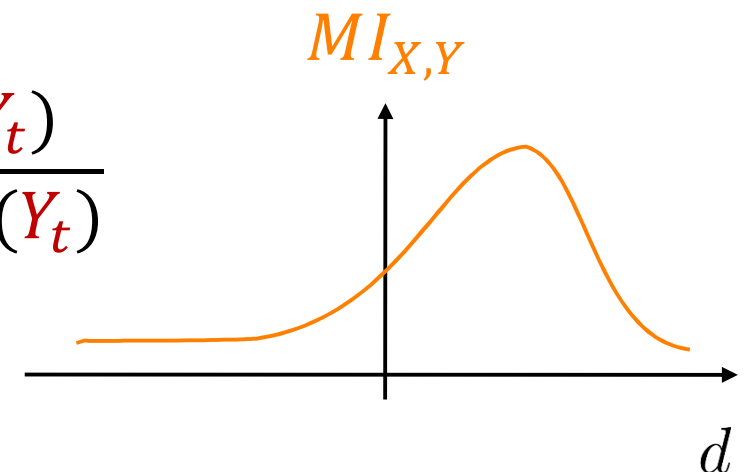
$$p(X_{t+d}, Y_t)$$

X independent from Y

$$p(X_{t+d})p(Y_t)$$

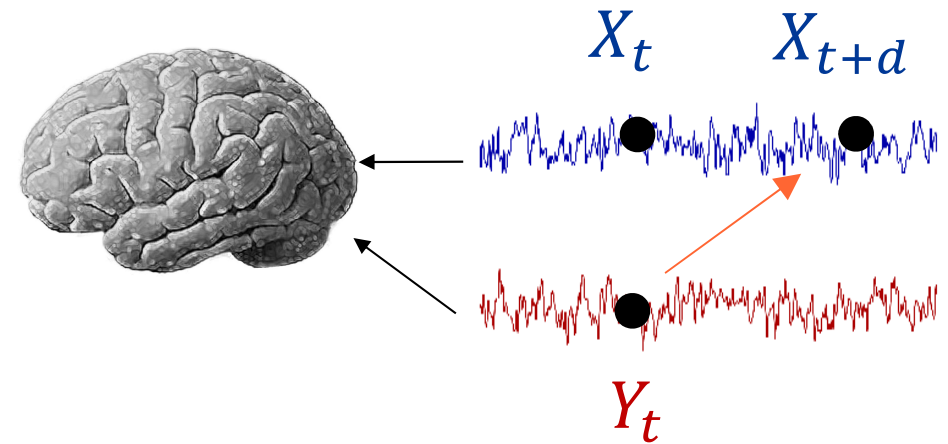
KL divergence

$$MI_{X,Y}(d) = \int p(X_{t+d}, Y_t) \log \frac{p(X_{t+d}, Y_t)}{p(X_{t+d})p(Y_t)}$$
$$= H(X_{t+d}) - H(X_{t+d}|Y_t)$$

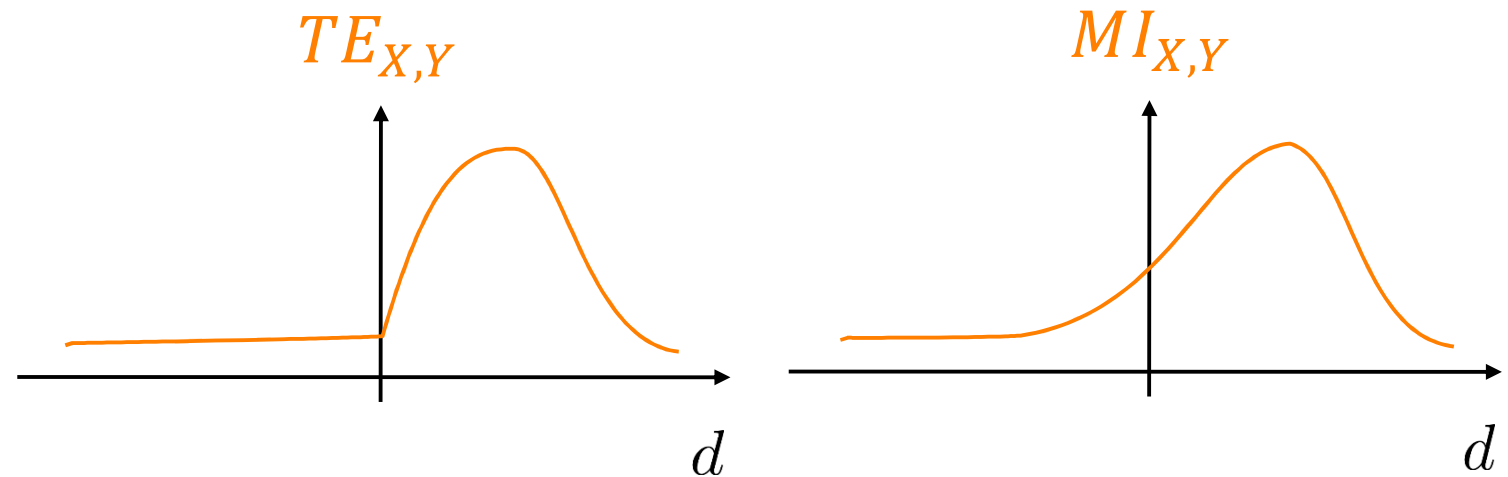


- $d > 0 \rightarrow$  'shared' information from the present of Y to the future of X
- $d < 0 \rightarrow$  information 'shared' from X to Y
- in general non-symmetric !

# Transfer Entropy



$$TE_{X,Y}(d) = H(X_{t+d}|X_t) - H(X_{t+d}|X_t, Y_t)$$

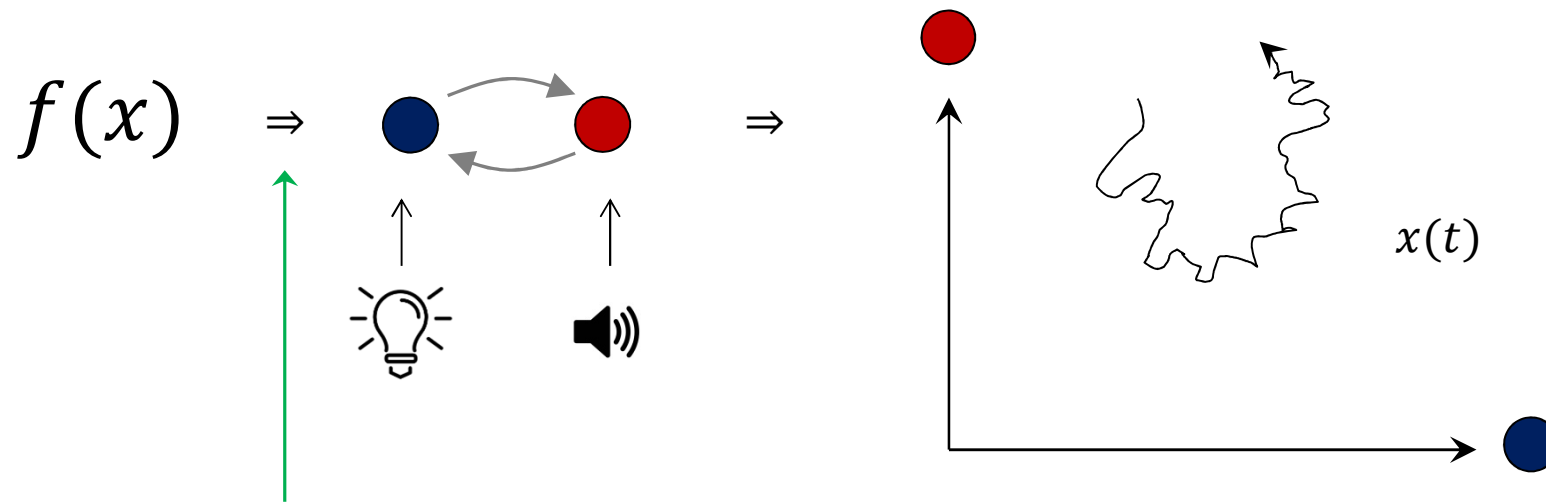


- non-symmetric by definition
- takes into account auto-correlations in time
- in general non-symmetric !



# Dynamics from Function

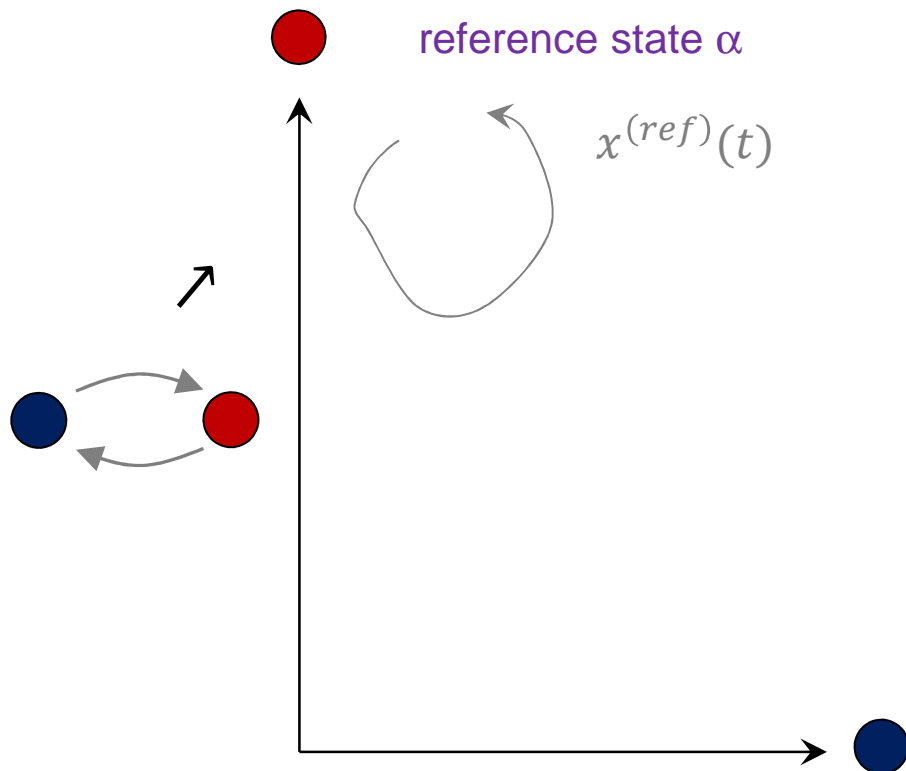
- Function  $\Rightarrow$  Neuronal Network  $\Rightarrow$  Dynamics
  - early sensory processing
  - 'direct information encoding and processing'



- artificial networks + learning [Hertz Korgh Palmer, 1991, ....]
- neuronal compiler [Eliasmith, Science 2012]
- ...

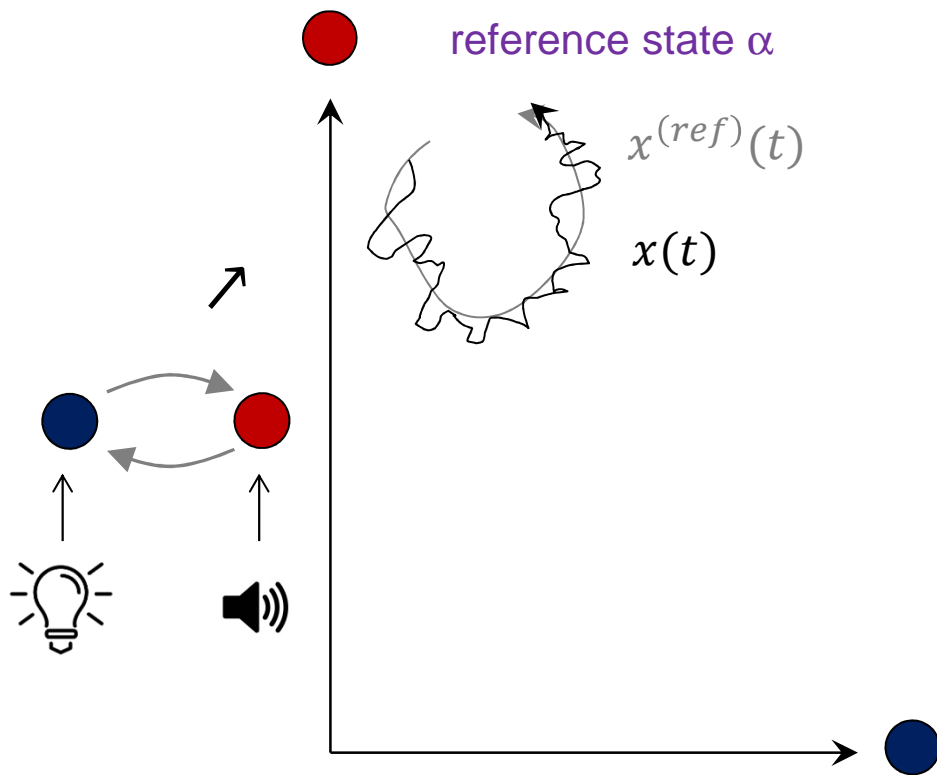
# Information Routed on top of Dynamical Reference States

- Function  $\Rightarrow$  Neuronal Network  $\Rightarrow$  Dynamics
  - early sensory processing
  - 'direct information encoding and processing'
- Reference Dynamics  $\Rightarrow$  Effective Network  $\Rightarrow$  Function
  - larger scale communication / self-organized information routing
  - 'information and processing in fluctuations around reference state'



# Information Routed on top of Dynamical Reference States

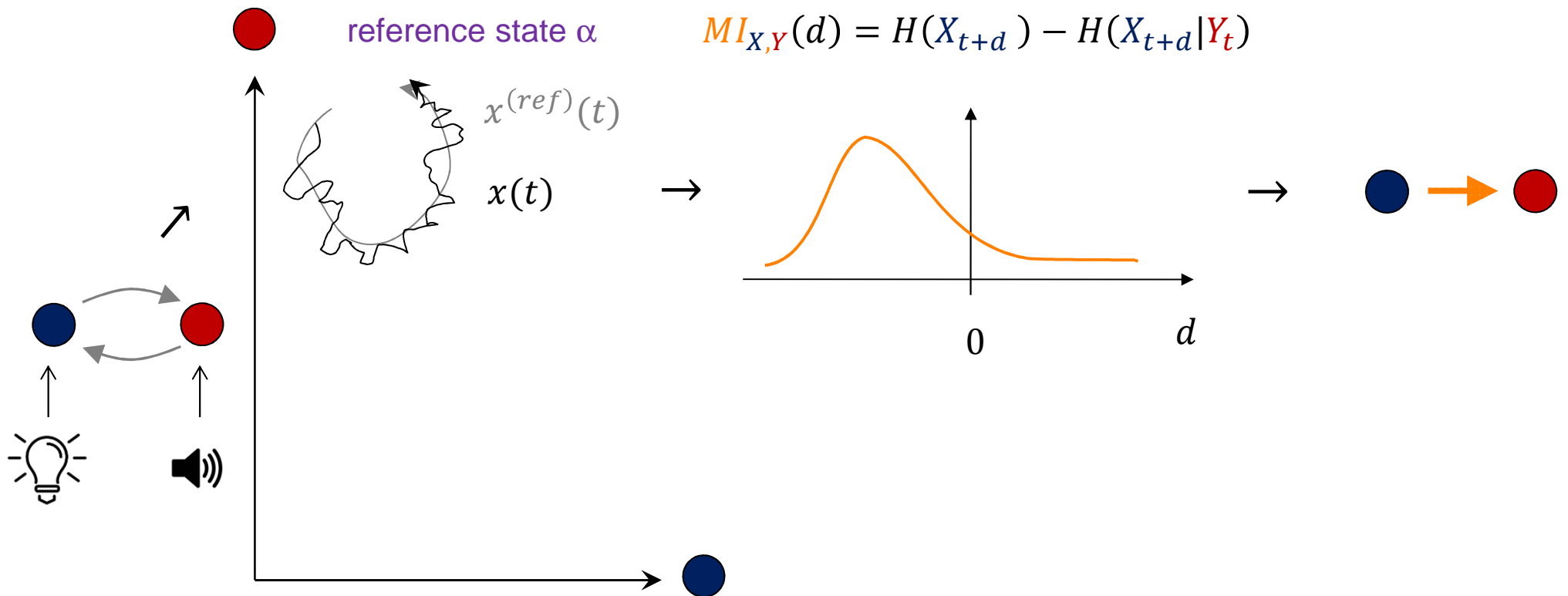
- Function  $\Rightarrow$  Neuronal Network  $\Rightarrow$  Dynamics
  - early sensory processing
  - 'direct information encoding and processing'
- Reference Dynamics  $\Rightarrow$  Effective Network  $\Rightarrow$  Function
  - larger scale communication
  - 'information and processing in fluctuations around reference state'



[Kirst, Time, Battaglia, Nature Communications, 2016, in press]

# Information Routed on top of Dynamical Reference States

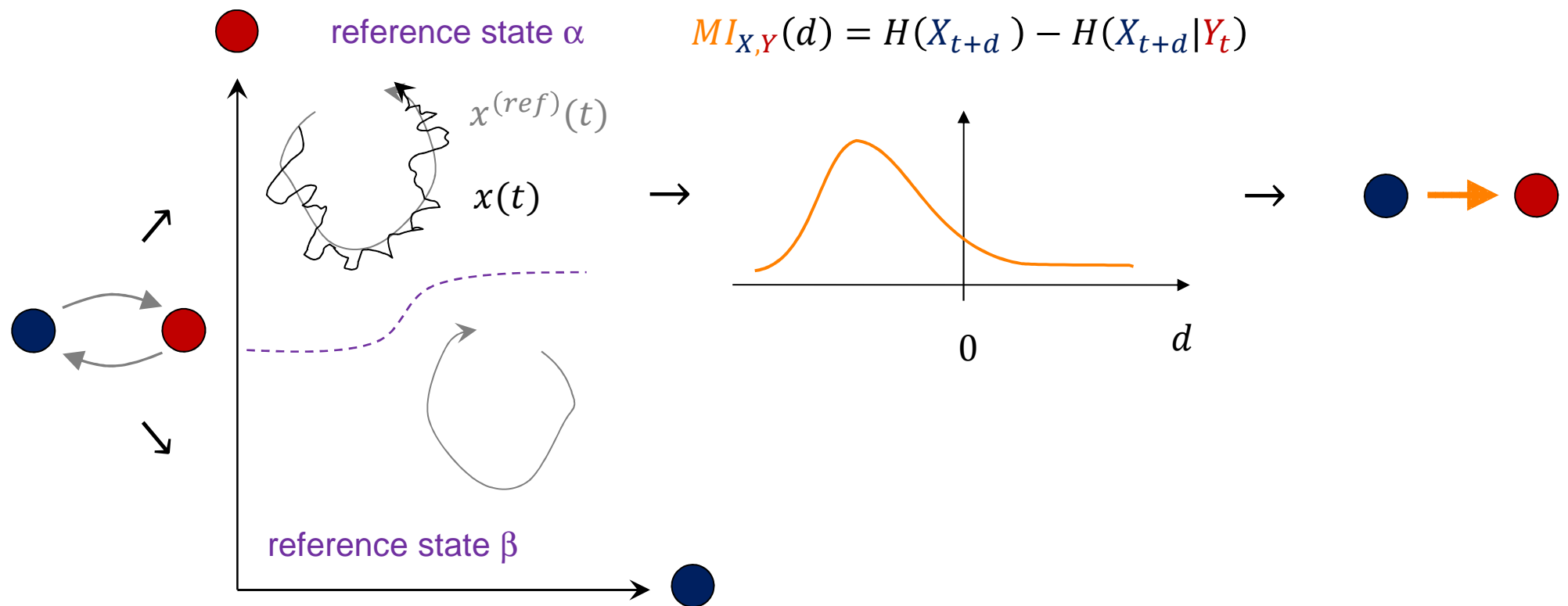
- Function  $\Rightarrow$  Neuronal Network  $\Rightarrow$  Dynamics
  - early sensory processing
  - 'direct information encoding and processing'
- Reference Dynamics  $\Rightarrow$  Effective Network  $\Rightarrow$  Function
  - larger scale communication
  - 'information and processing in fluctuations around reference state'



[Kirst, Time, Battaglia, Nature Communications, 2016, in press]

# Information Routed on top of Dynamical Reference States

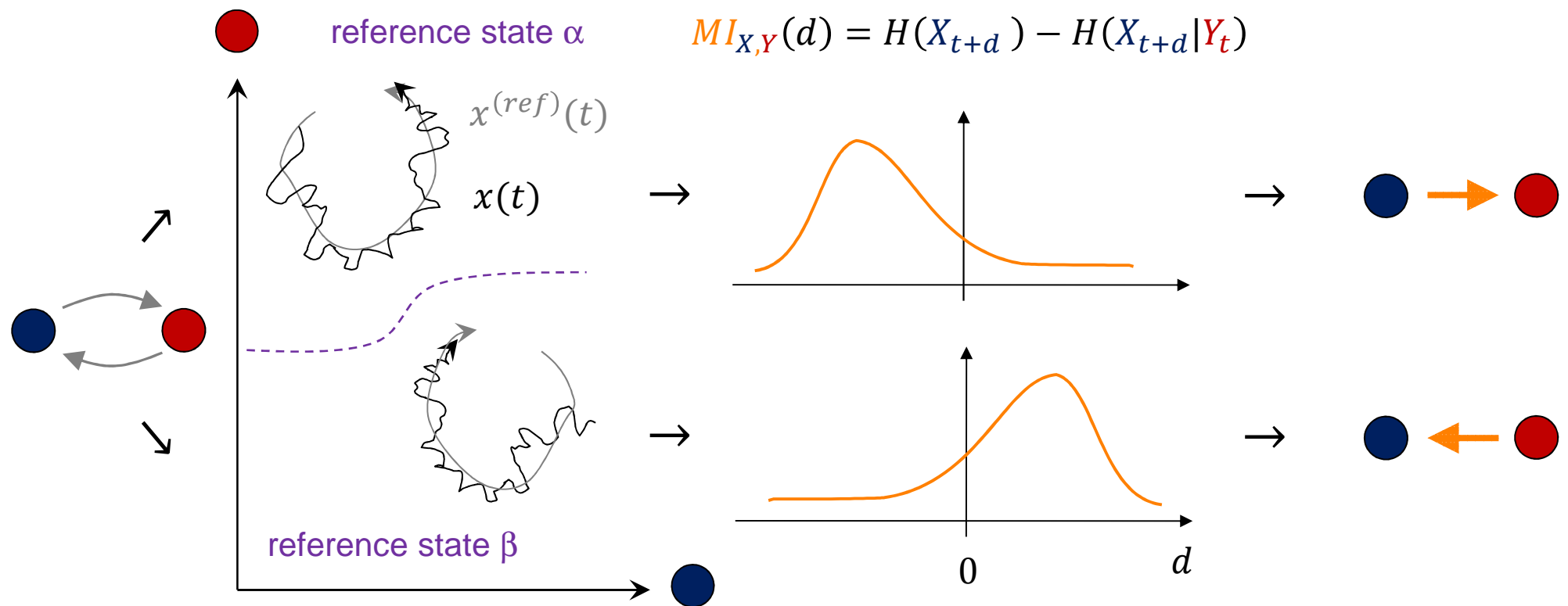
- Function  $\Rightarrow$  Neuronal Network  $\Rightarrow$  Dynamics
  - early sensory processing
  - 'direct information encoding and processing'
- Reference Dynamics  $\Rightarrow$  Effective Network  $\Rightarrow$  Function
  - larger scale communication
  - 'information and processing in fluctuations around reference state'



[Kirst, Time, Battaglia, Nature Communications, 2016, in press]

# Information Routed on top of Dynamical Reference States

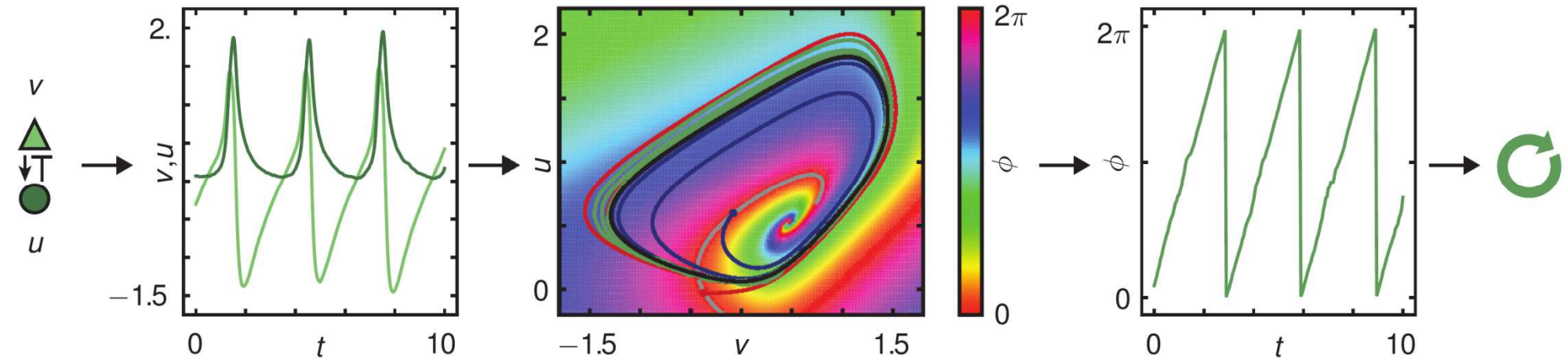
- Function  $\Rightarrow$  Neuronal Network  $\Rightarrow$  Dynamics
  - early sensory processing
  - 'direct information encoding and processing'
- Reference Dynamics  $\Rightarrow$  Effective Network  $\Rightarrow$  Function
  - larger scale communication
  - 'information and processing in fluctuations around reference state'



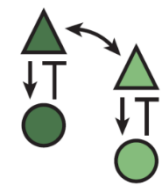
[Kirst, Time, Battaglia, Nature Communications, 2016, in press]

# Phase Reduction and Kuramoto Oscillators

Phase reduction:



Phase dynamics (of weakly coupled oscillators):



$$\frac{d}{dt}\phi_i = \omega_i + \sum_j \gamma_{ij} (\phi_i - \phi_j) + \zeta_i$$

[Kuramoto, Springer 1984]  
[Acebron et al. 2005]

- 'standard model' for synchronization
- 'phase part' of discrete Ginzburg -Landau / multiple Hopf bifurcation
- common dynamics: phase locking

# Information Routing Measures in Kuramoto Networks

- $$\frac{d}{dt}\phi_i = \omega_i + \sum_j \gamma_{ij} (\phi_i - \phi_j) + \zeta_i$$

- joint probability distribution:  $p(\phi_t, \phi_{t+d}) = p(\phi_t) p(\phi_{t+d}|\phi_t)$

- no noise  $\zeta_i = 0$  stable phase-locked state:  $\Omega = \omega_i + \sum_j \gamma_{ij} (\Delta\phi_{ij})$

- small noise expansion around phase locked state  $\phi_i = \psi_i + \sigma\varphi_i + \dots$

- $\Rightarrow p(\phi_{t+d}|\phi_t)$

- phase rotation symmetry / phase variables live on circle

- $\Rightarrow p(\phi_t)$

- integration to marginals

→ **Theorem**

delayed mutual information  $MI_{ij}(d)$  and transfer entropy  $TE_{i \rightarrow j}(d)$

as function of *network structure* and *dynamical reference state*



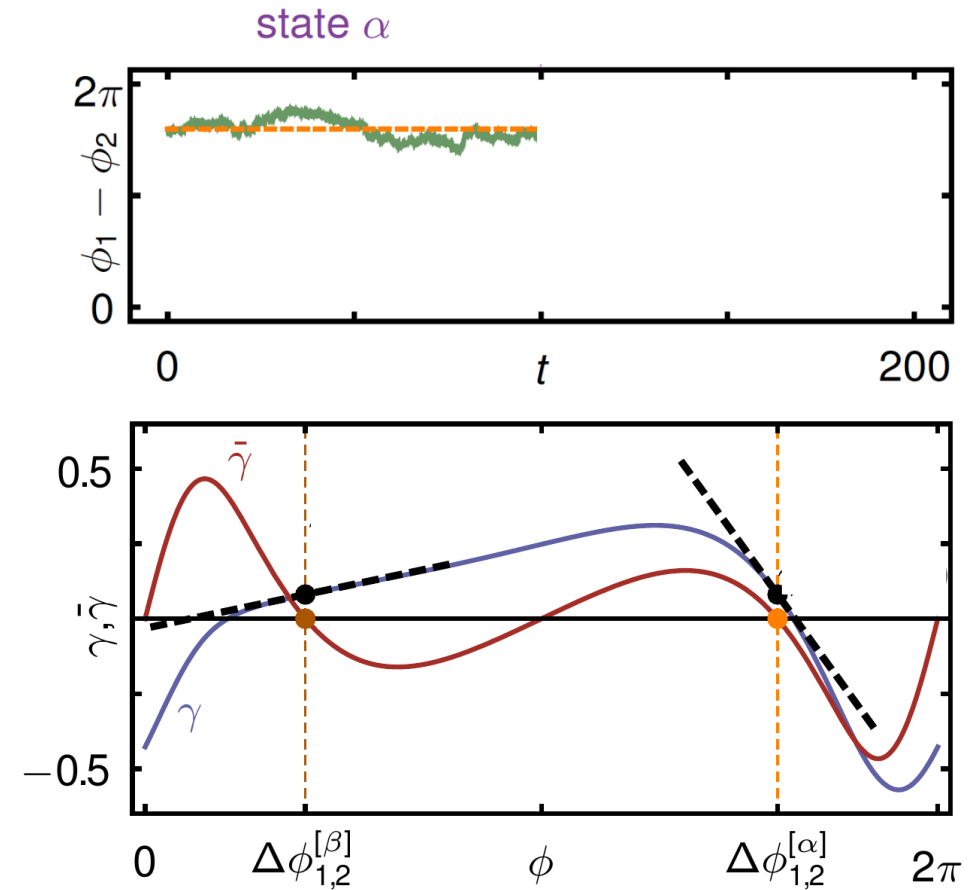
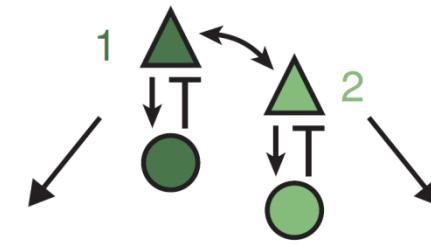
# Dynamic Information Routing

- phase dynamics:

$$\dot{\phi}_i = \omega + \gamma (\phi_i - \phi_j) + \sigma \zeta_i(t)$$

- phase locking:  $\sigma = 0$

$$\Delta \dot{\phi} = \gamma (\Delta \phi) - \gamma (-\Delta \phi) = \bar{\gamma} (\Delta \phi)$$



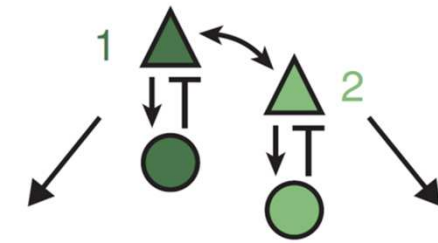
# Dynamic Information Routing

- phase dynamics:

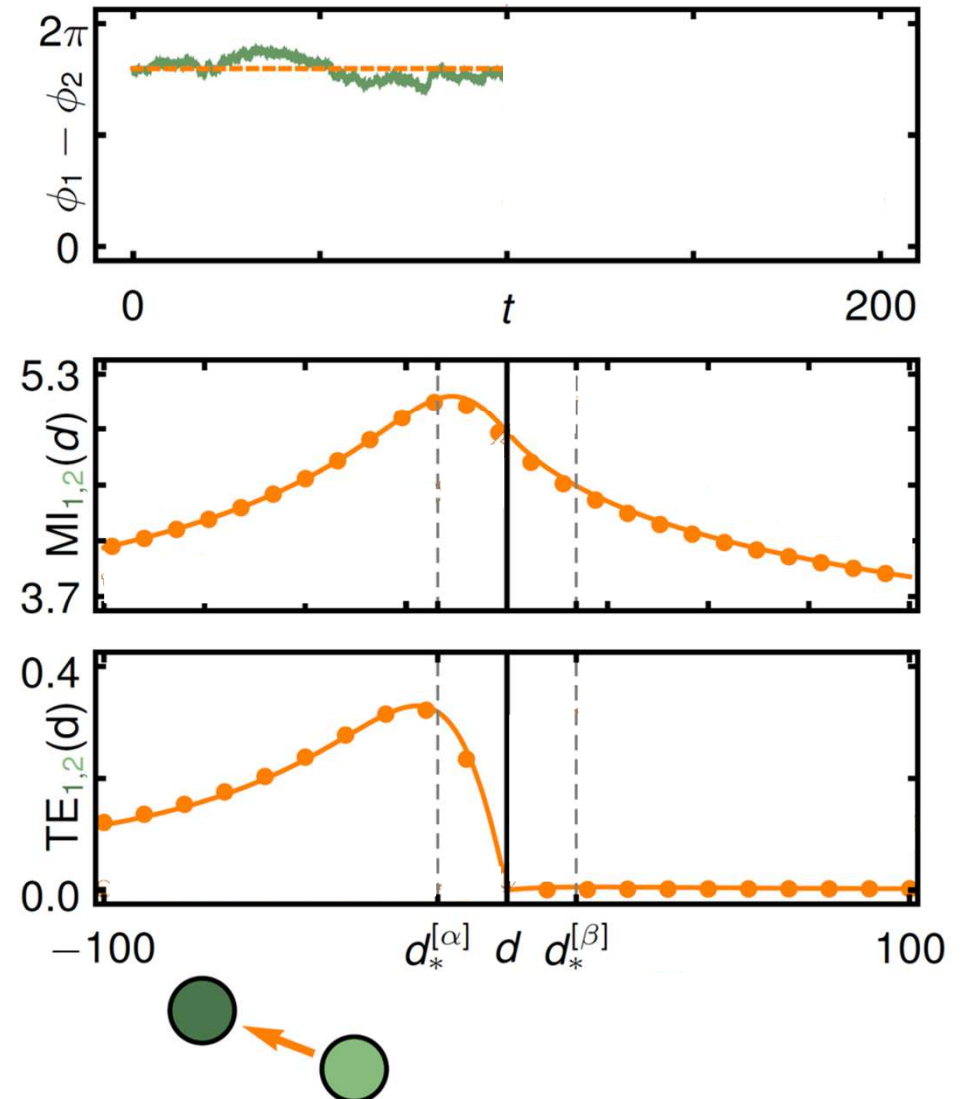
$$\dot{\phi}_i = \omega + \gamma (\phi_i - \phi_j) + \sigma \zeta_i(t)$$

- phase locking:  $\sigma = 0$

$$\Delta \dot{\phi} = \gamma (\Delta \phi) - \gamma (-\Delta \phi) = \bar{\gamma} (\Delta \phi)$$



state  $\alpha$



# Dynamic Information Routing

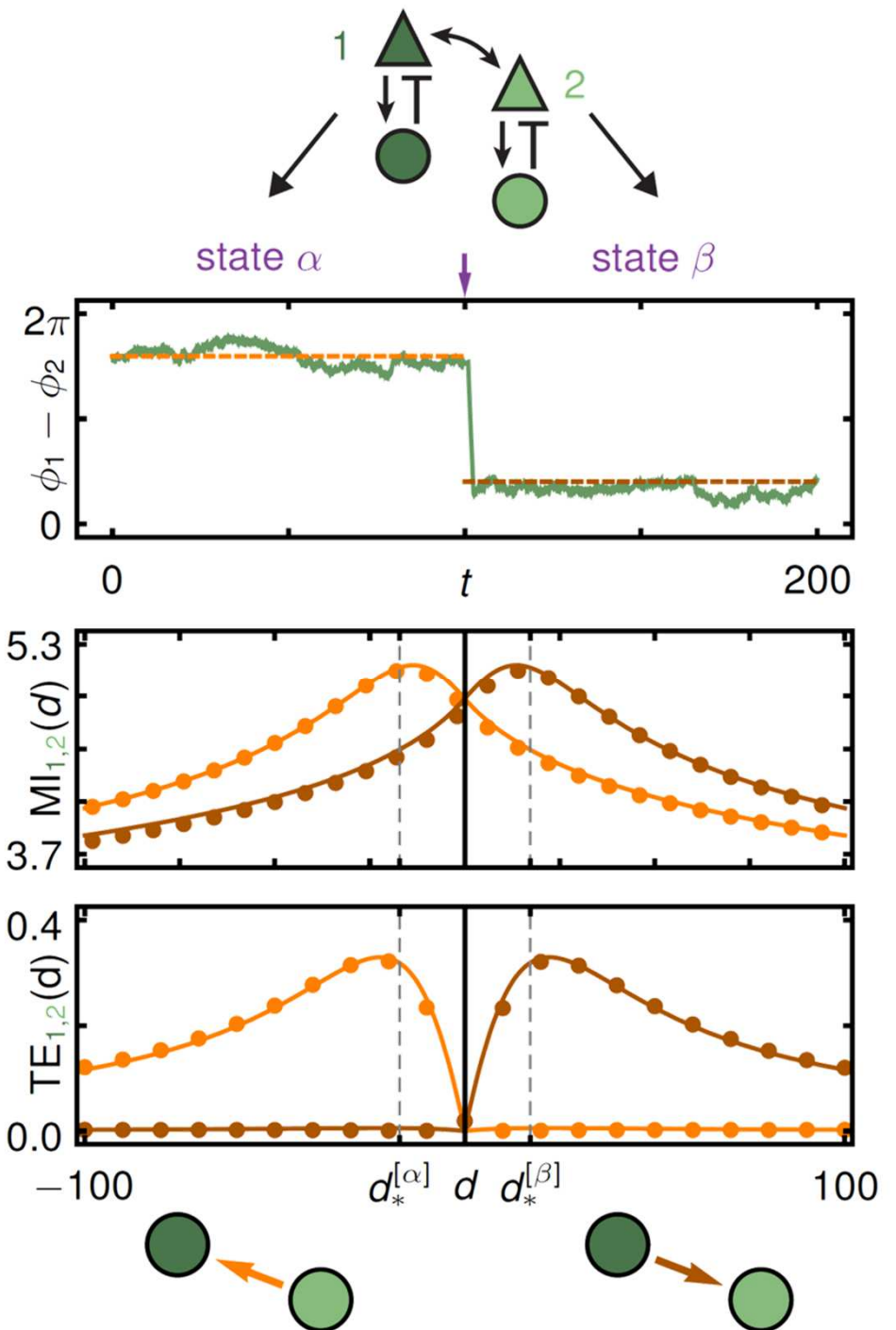
- phase dynamics:

$$\dot{\phi}_i = \omega + \gamma (\phi_i - \phi_j) + \sigma \zeta_i(t)$$

$$\sigma = 0$$

$$\Delta \dot{\phi} = \gamma (\Delta \phi) - \gamma (-\Delta \phi) = \bar{\gamma} (\Delta \phi)$$

→ switching reference states switches network communication function



# Dynamic Information Routing

- phase dynamics:

$$\dot{\phi}_i = \omega + \gamma (\phi_i - \phi_j) + \sigma \zeta_i(t)$$

- phase locking:  $\sigma = 0$

$$\Delta \dot{\phi} = \gamma (\Delta \phi) - \gamma (-\Delta \phi) = \bar{\gamma} (\Delta \phi)$$

- small noise expansion:

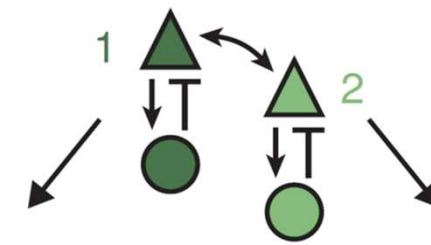
$$\phi_i = \psi_i + \sigma \varphi_i + \dots$$

$$\dot{\psi}_i = \omega + \gamma (\Delta \phi_0) = \Omega$$

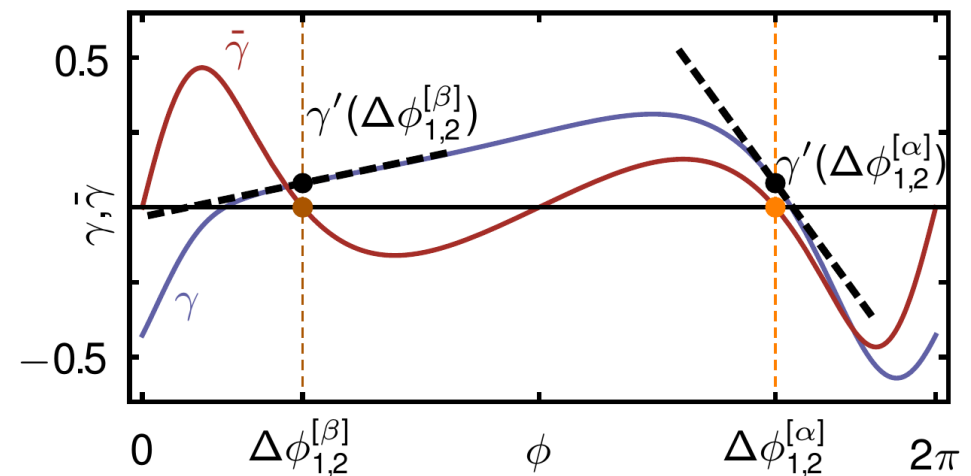
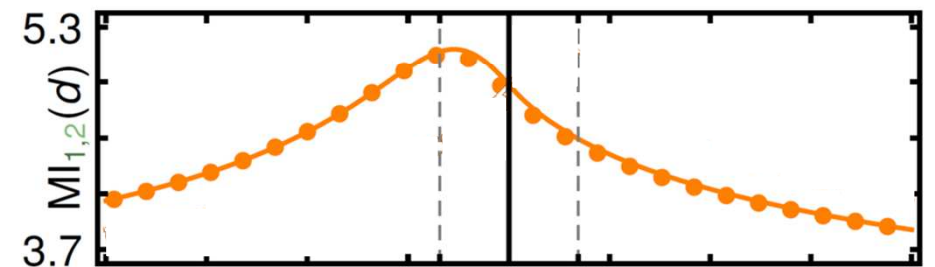
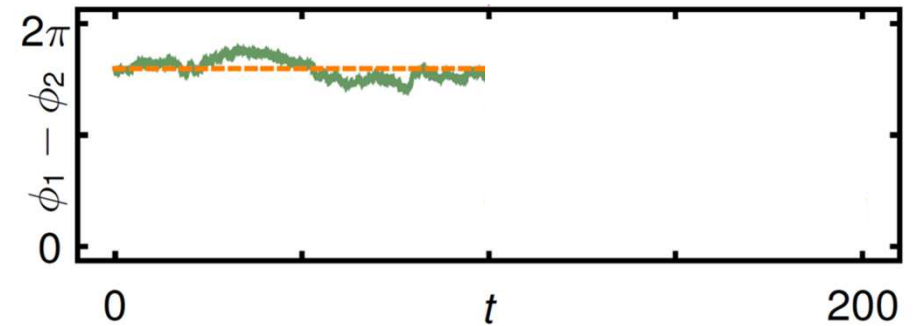
$$\dot{\varphi}_1 = \gamma' (\Delta \phi_0) (\varphi_1 - \varphi_2) + \zeta_1$$

$$\dot{\varphi}_2 = \gamma' (-\Delta \phi_0) (\varphi_2 - \varphi_1) + \zeta_2$$

[Kirst et al, Nat Comm, 2016]



state  $\alpha$



# Dynamic Information Routing

- phase dynamics:

$$\dot{\phi}_i = \omega + \gamma (\phi_i - \phi_j) + \sigma \zeta_i(t)$$

- phase locking:  $\sigma = 0$

$$\Delta \dot{\phi} = \gamma (\Delta \phi) - \gamma (-\Delta \phi) = \bar{\gamma} (\Delta \phi)$$

- small noise expansion:

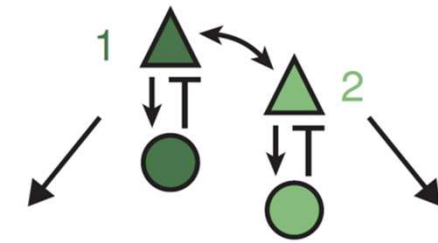
$$\phi_i = \psi_i + \sigma \varphi_i + \dots$$

$$\dot{\psi}_i = \omega + \gamma (\Delta \phi_0) = \Omega$$

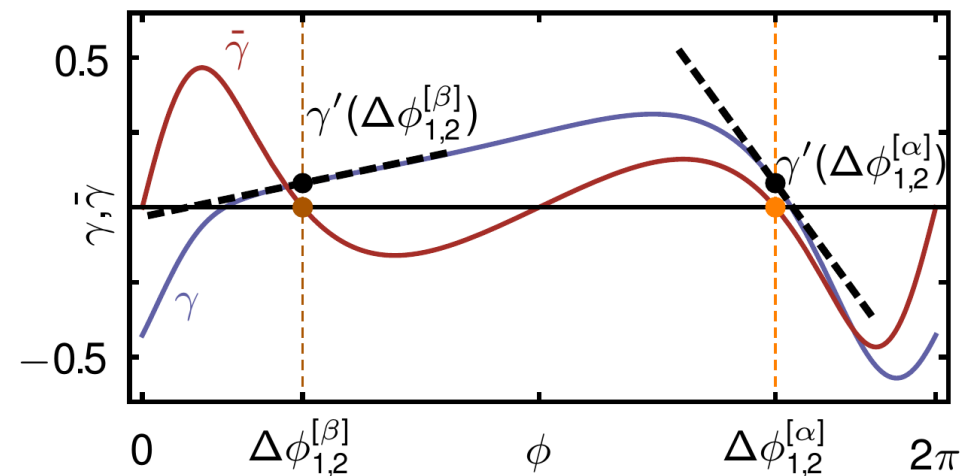
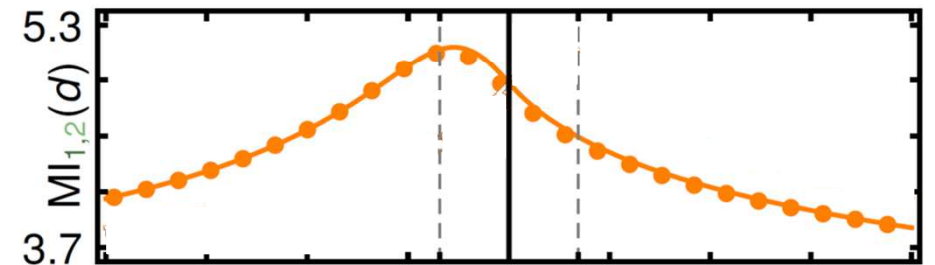
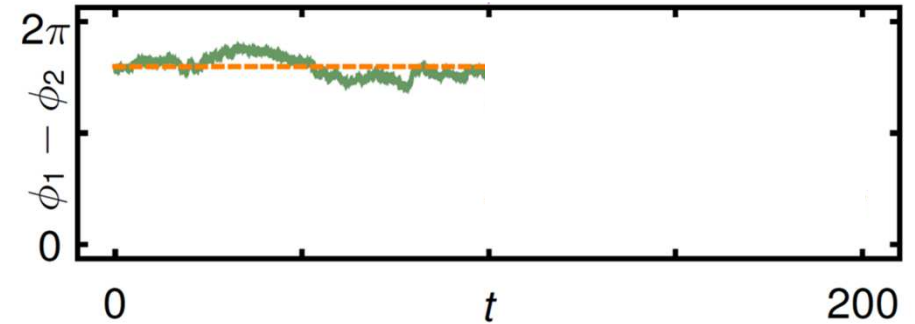
~~$$\dot{\phi}_1 = \gamma'(\Delta \phi_0) (\varphi_1 - \varphi_2) + \zeta_1$$~~

~~$$\dot{\phi}_2 = \gamma'(-\Delta \phi_0) (\varphi_2 - \varphi_1) + \zeta_2$$~~

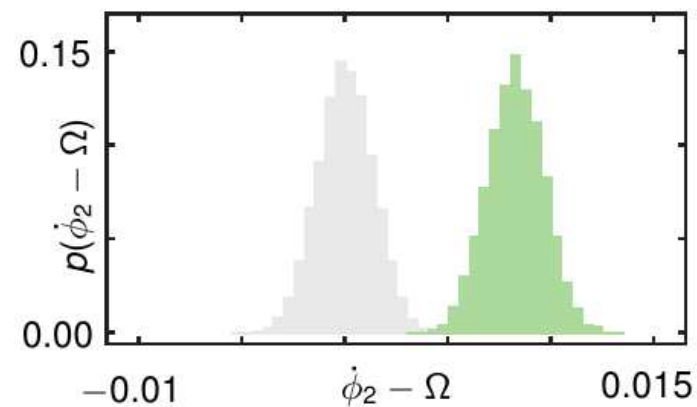
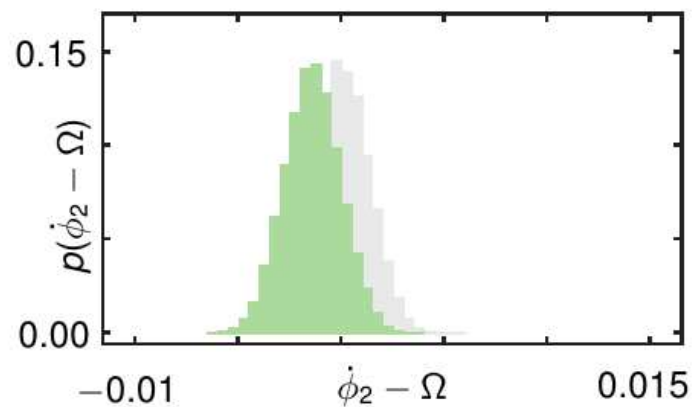
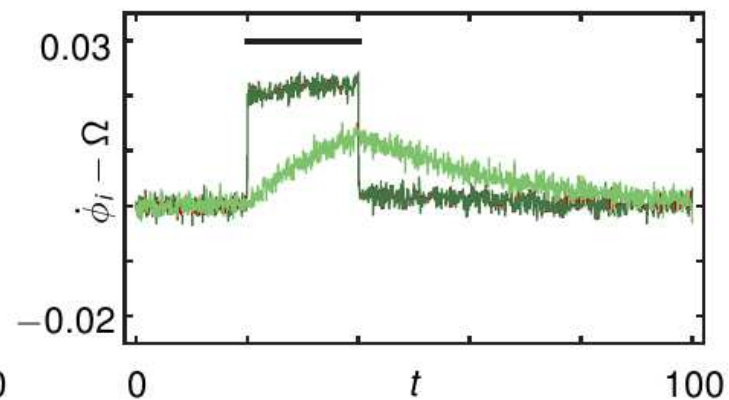
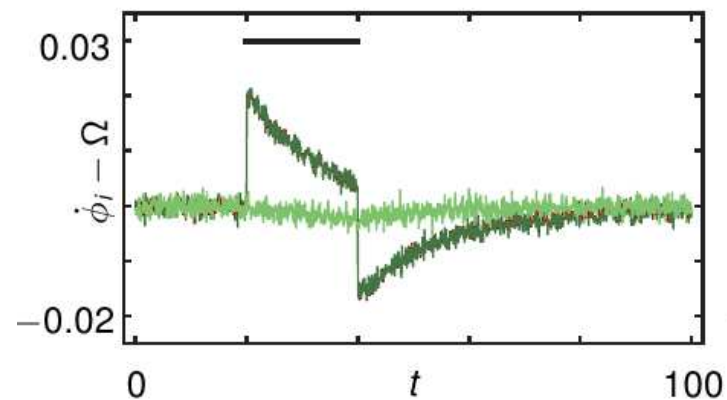
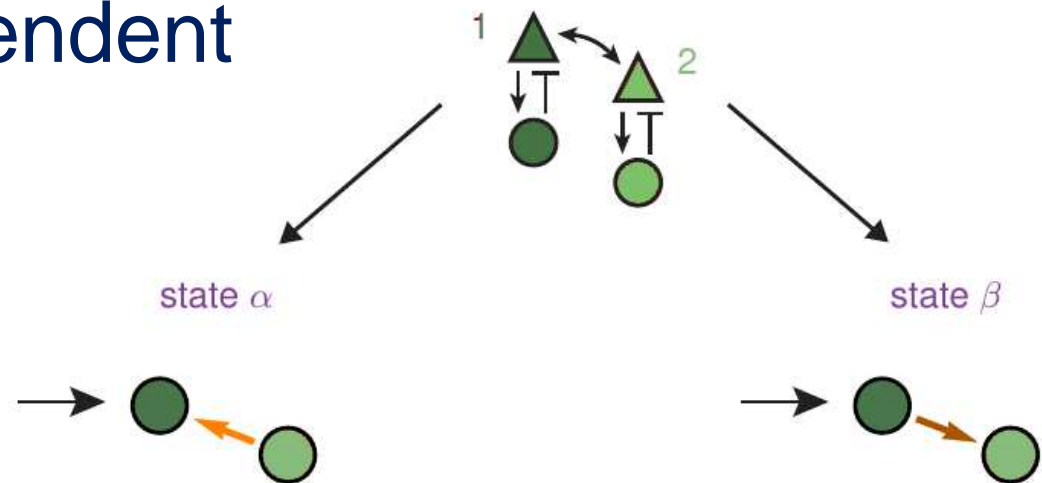
[Kirst et al, Nat Comm, 2016]



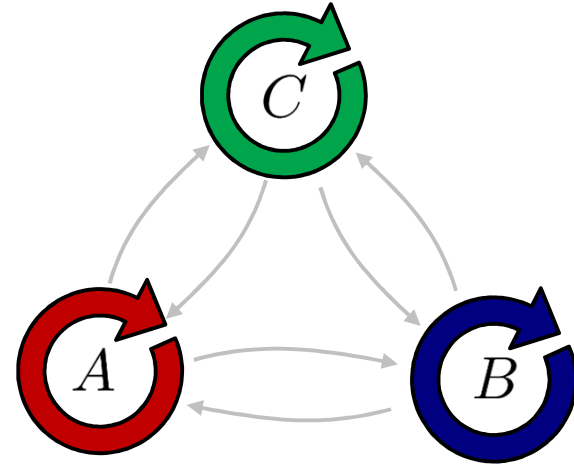
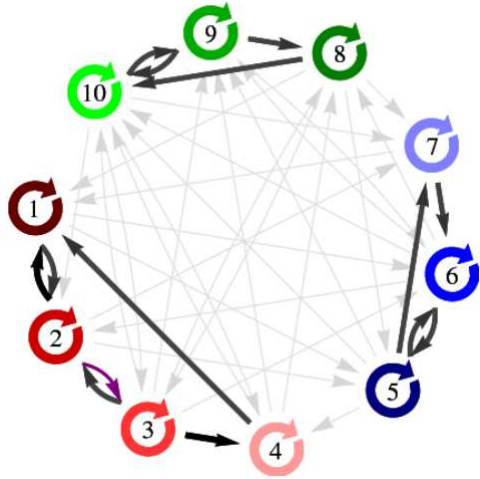
state  $\alpha$



# Dynamical State Dependent Signal Transmission



# Information Routing in Modular Networks



- $\dot{\phi}_i = \omega_i + \sum_j \gamma_{ij} (\phi_i - \phi_j) + \zeta_i$

⇒ Analytic expression  
for delayed mutual information

$MI_{ij}(d)$

as function of

- dynamical state and
- network parameters

- second phase reduction step on clusters

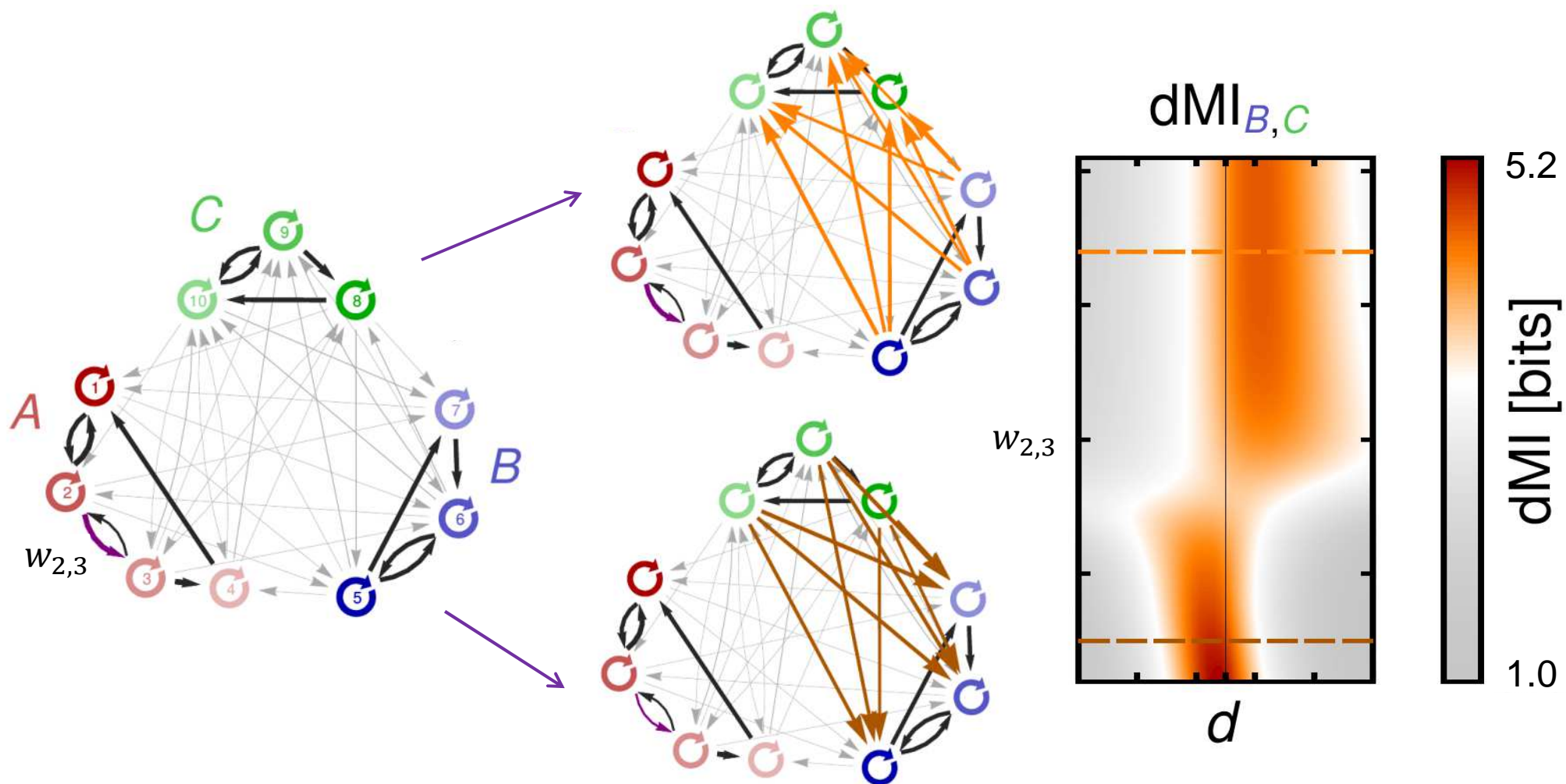
$$\dot{\Phi}_X = \Omega_X + \sum_Y Z_X(\Phi_X) G_{XY}(\Phi_X, \Phi_Y) + \xi_X$$

- stochastic averaging

$$\dot{\Phi}_X = \Omega_X + \sum_Y \Gamma_{XY}(\Phi_X - \Phi_Y) + \Xi_X$$

⇒ cluster phase response  $Z_X$  and thus  $\Gamma_{XY}$   
and **delayed mutual information**  
depend on **local clusters properties**

# Hierarchical Networks: Action at a Distance

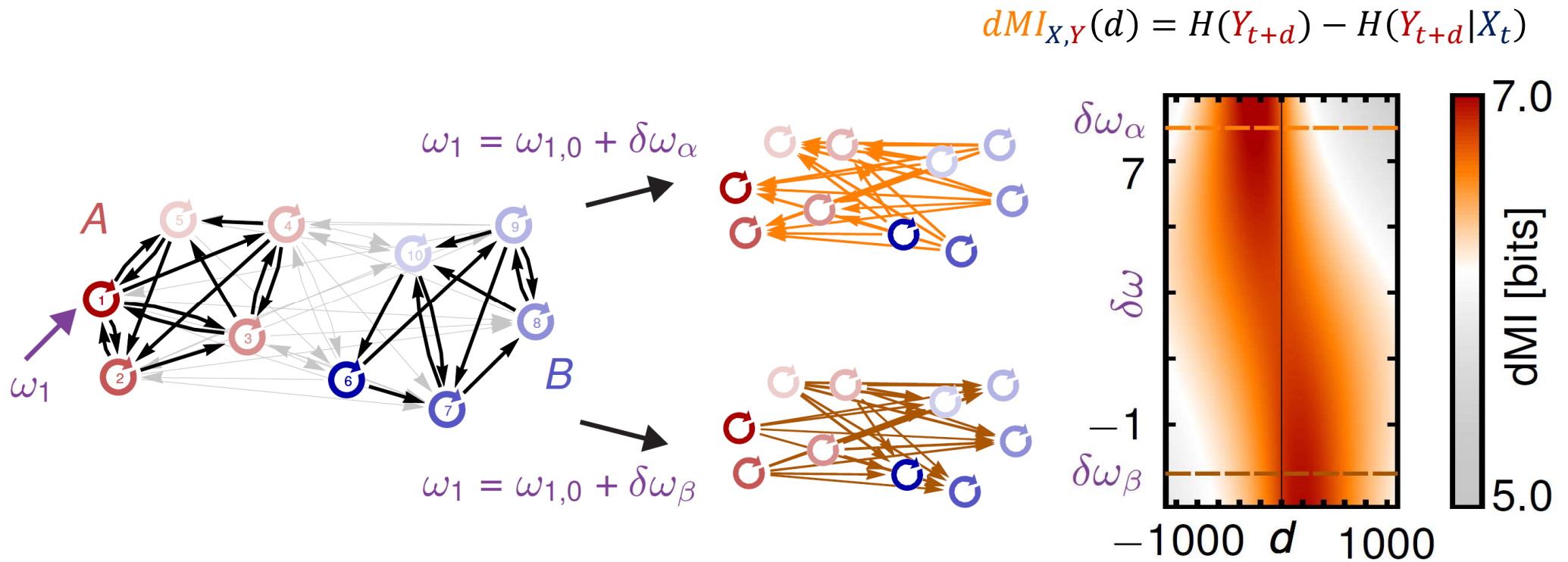


- local control of non-local functional connectivity motifs



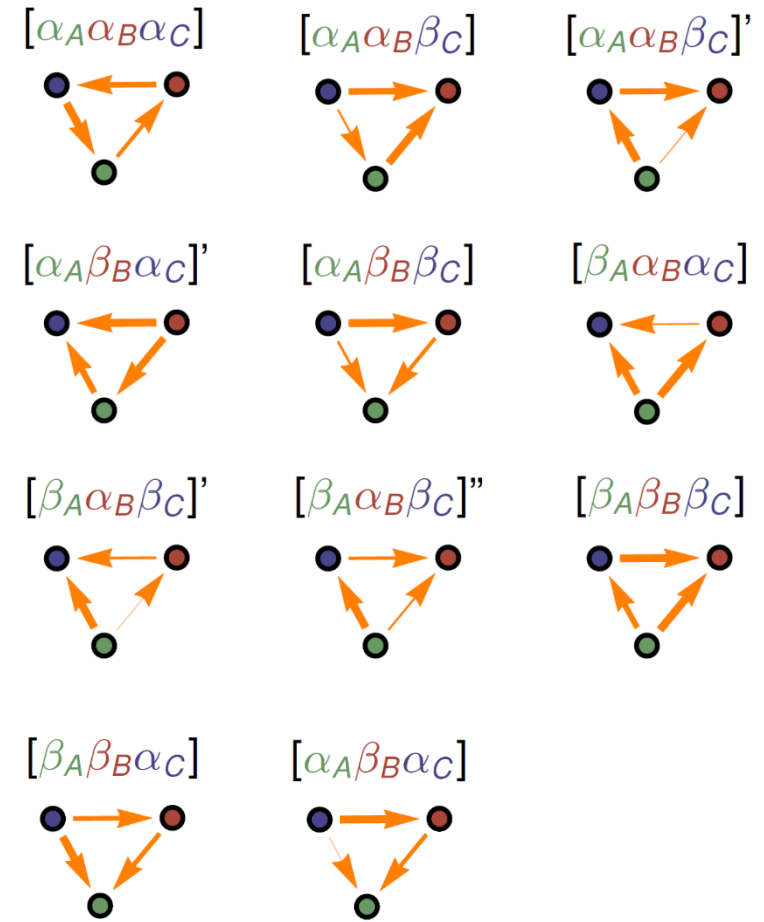
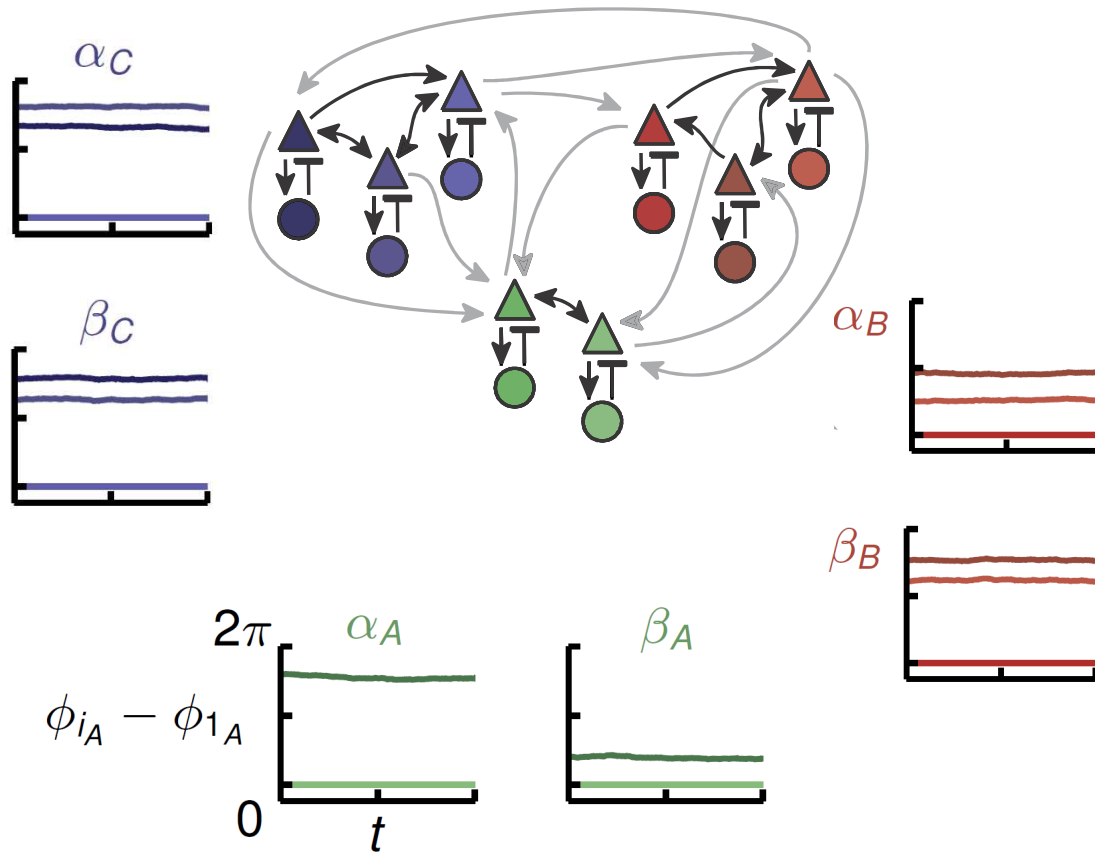
# Local Control of Non-Local Information Routing

- Reference Dynamics  $\Rightarrow$  Effective Network  $\Rightarrow$  Function
- action at a distance: local control of non-local communication



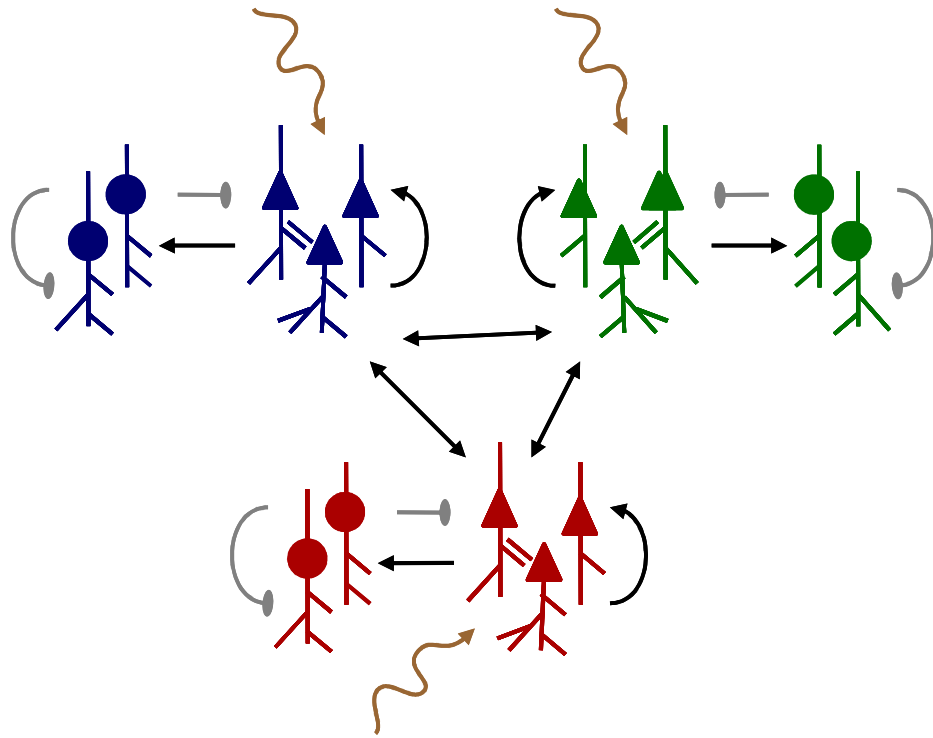
# Combinatorial Information Routing

- multi-stable local dynamical states

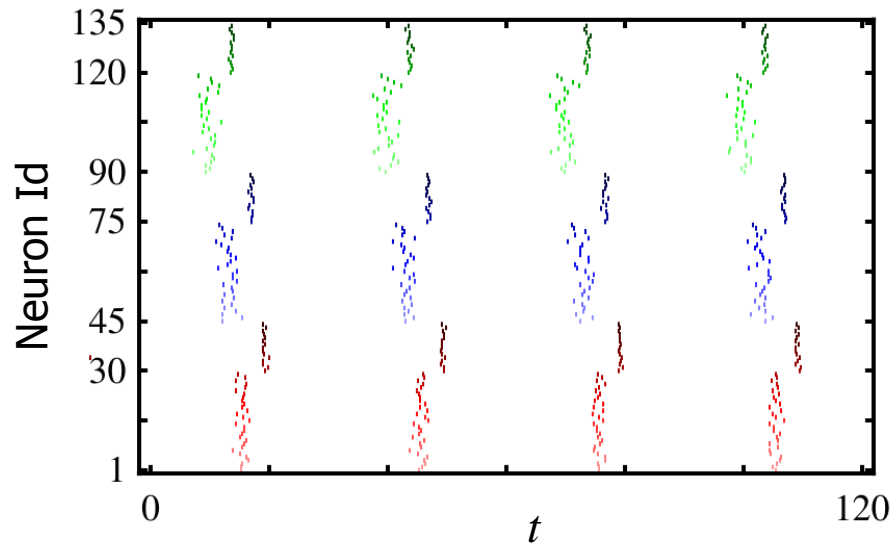
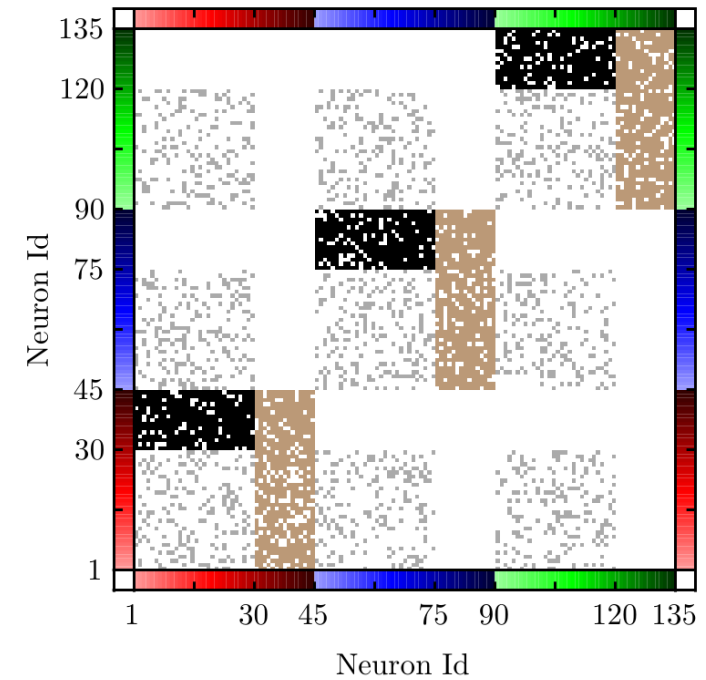


→ combination of local dynamical states determines non-local information flow pattern

# Networks of PING Clusters



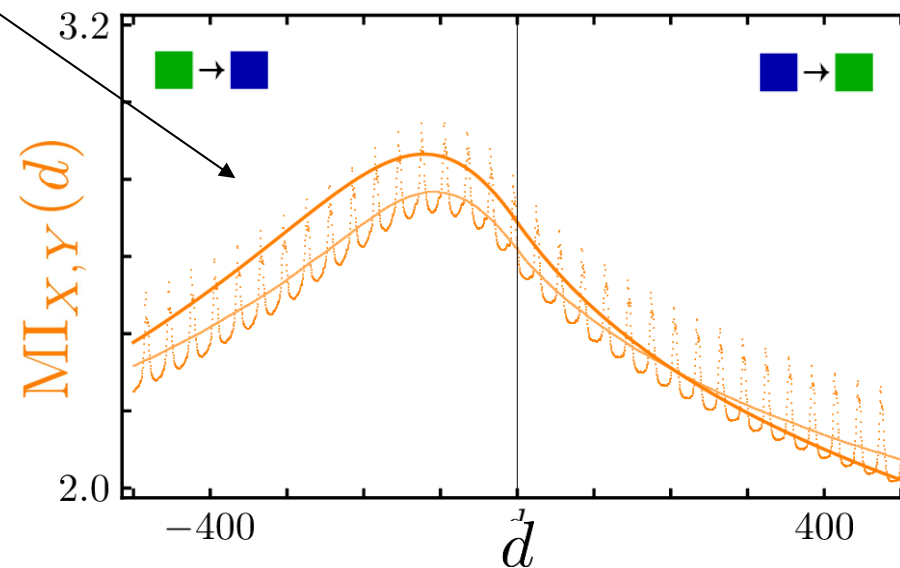
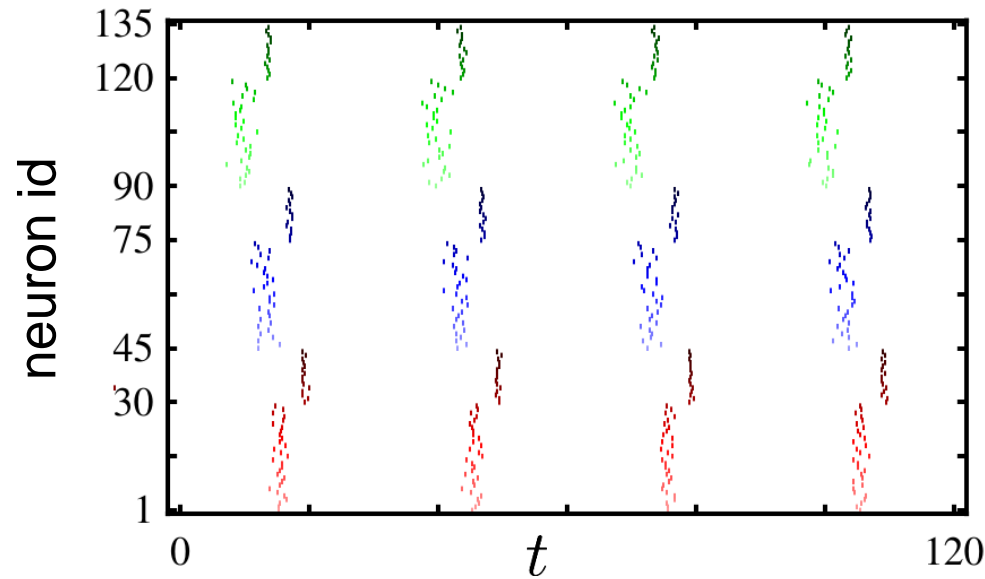
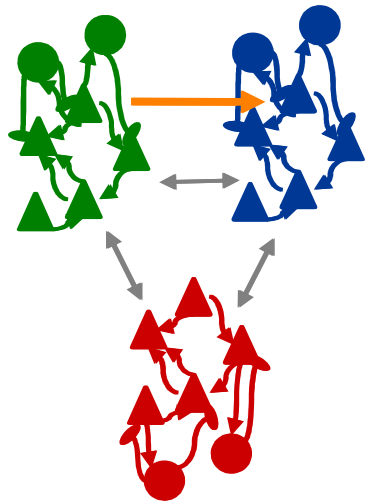
coupling matrix



phase locking between PING clusters

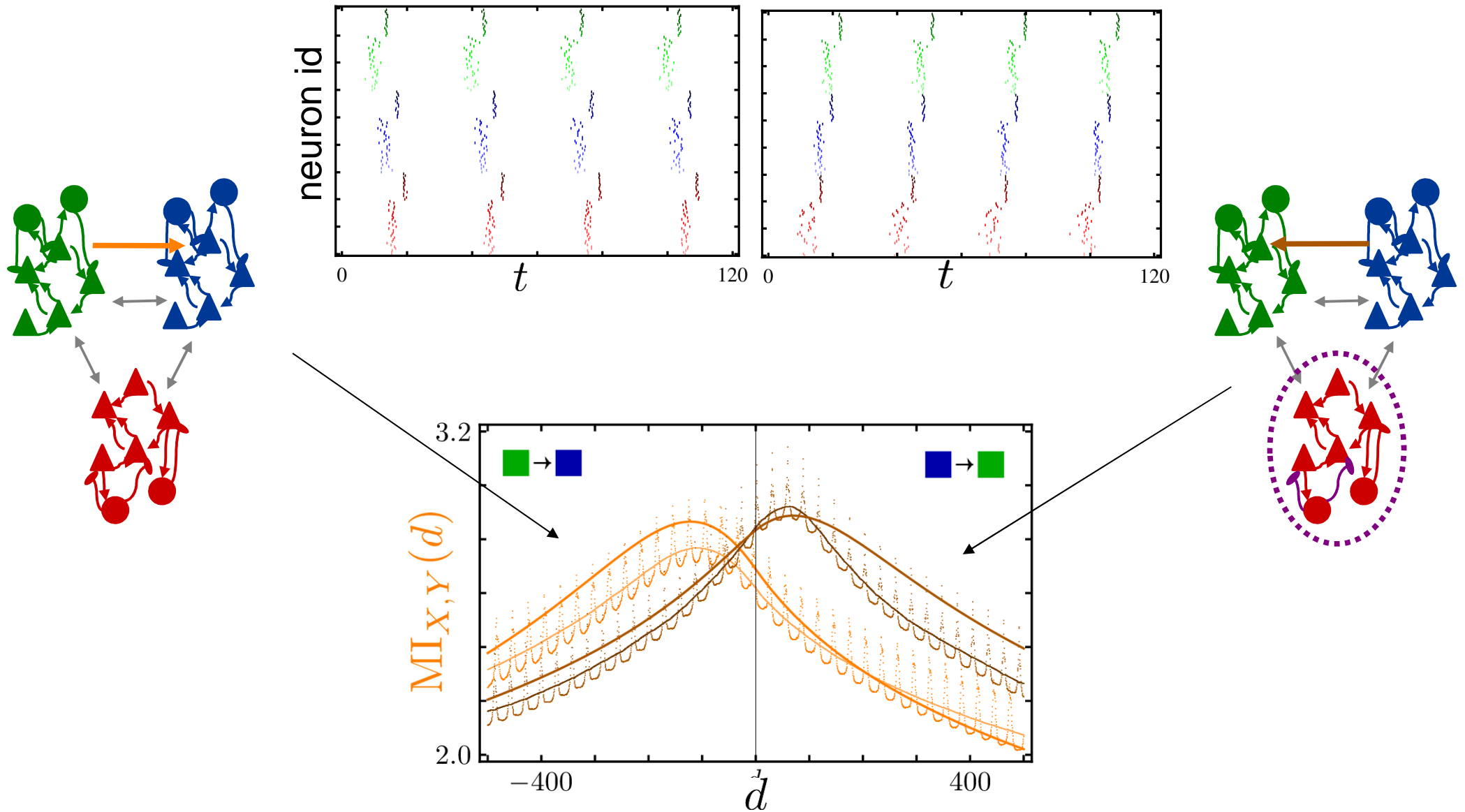
# Local Control of Non-Local Functional Connectivity

- clusters of **Pyramidal Inter-Neuron Gamma** networks [Börger & Kopell, 2005]



# Local Control of Non-Local Functional Connectivity

- clusters of **Pyramidal Inter-Neuron Gamma** networks [Börger & Kopell, 2005]



# Information Flow in Spike Patterns

- phase channel:

- limitations on precise readout / max capacity

- amplitude channel / spike patterns:

- limited by cluster size only

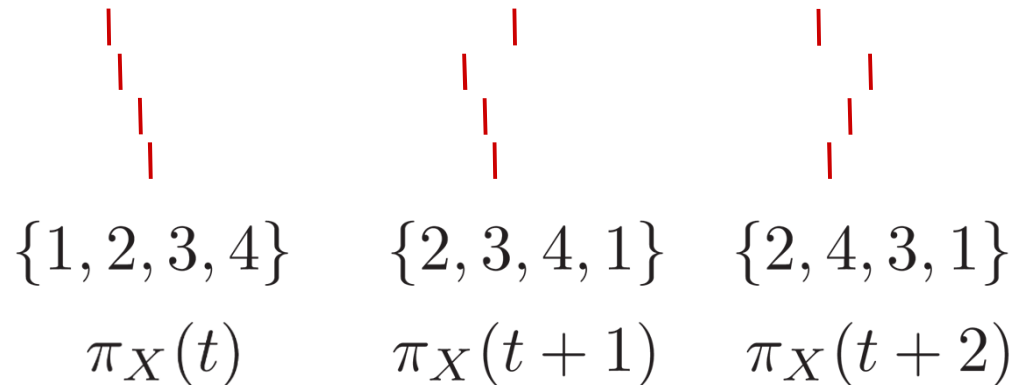
- PING: spike-to-spike oscillation

- clocked sequence of codewords (ordering of spikes)

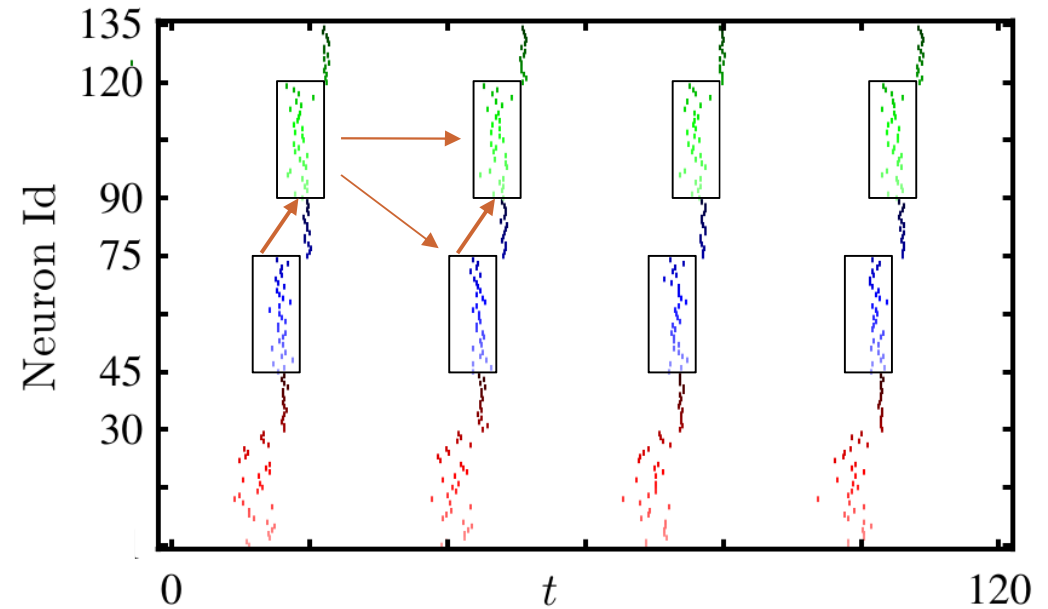
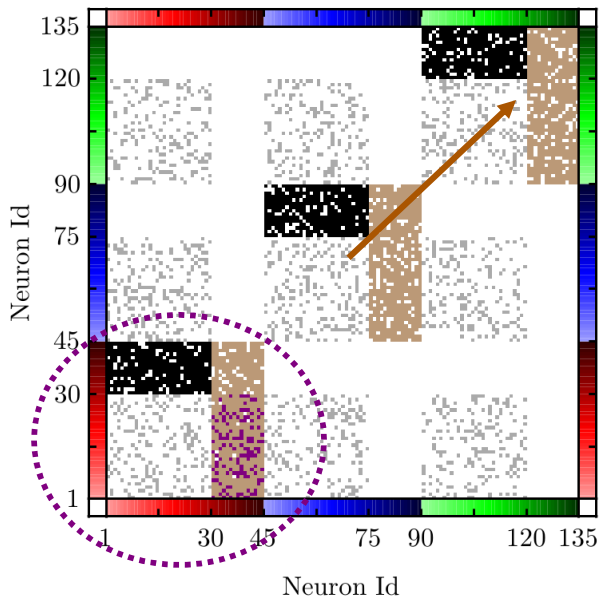
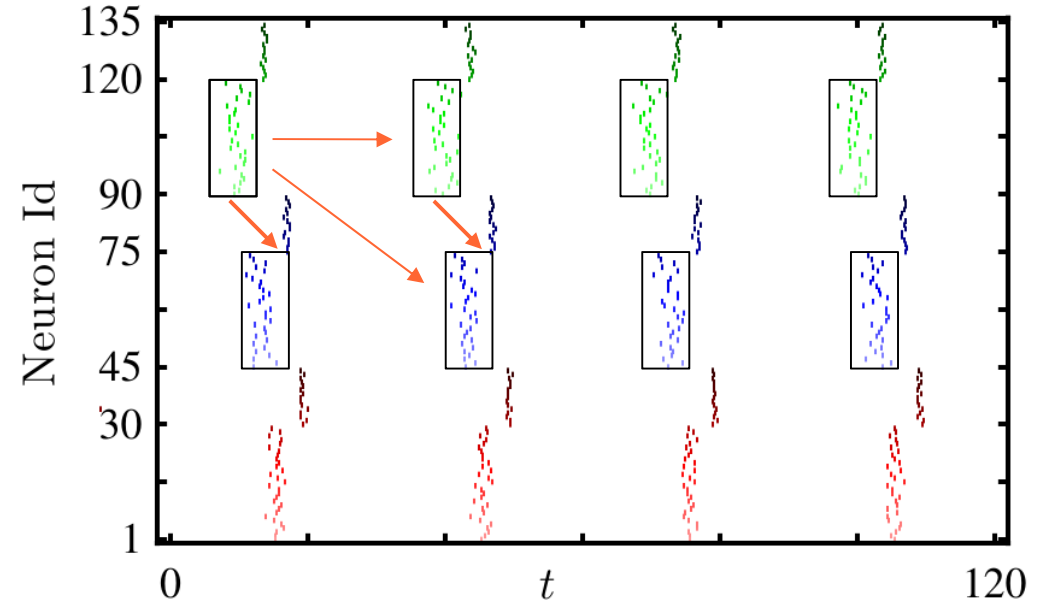
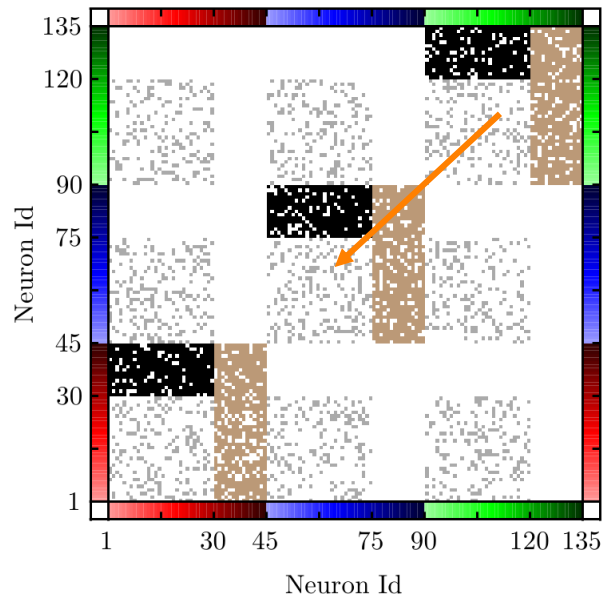
$$\pi \in S_N$$

- delayed mutual information

$$MI_{XY}^{\pi}(d)$$



# Local Control of Non-Local Info Flow



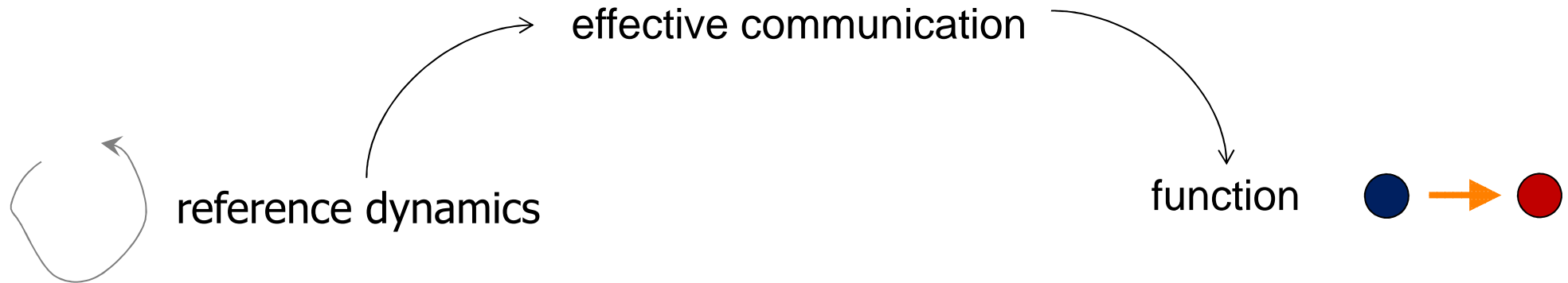
# Outline: Flexible Function in Neuronal Networks

- dynamic information routing in complex networks
  - structural vs. effective connectivity
  - flexible information routing in oscillator networks
  - spiking networks, transient dynamics
- flexible information processing in complex networks
  - oscillatory Hopfield networks
  - self-organized pattern recognition
- learning flexible function in neuronal networks
- connections to experiments
  - brain state identification in zebrafish
  - complete brain activity mapping in mouse
- conclusions



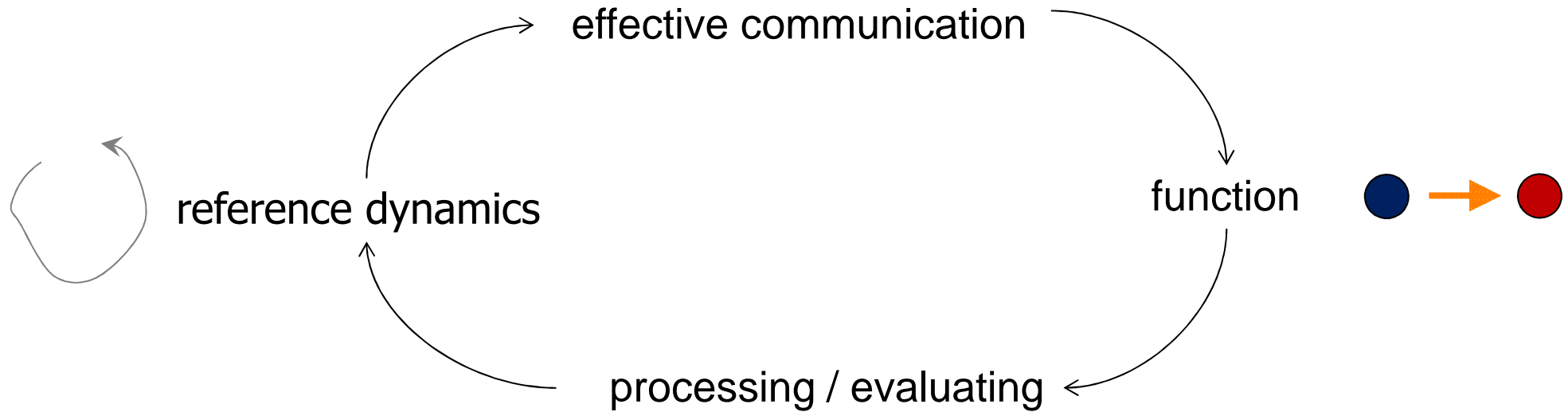
# Self-Reprogramming Neuronal Networks via Dynamics

- Closing the loop from functional dynamics to dynamic function:



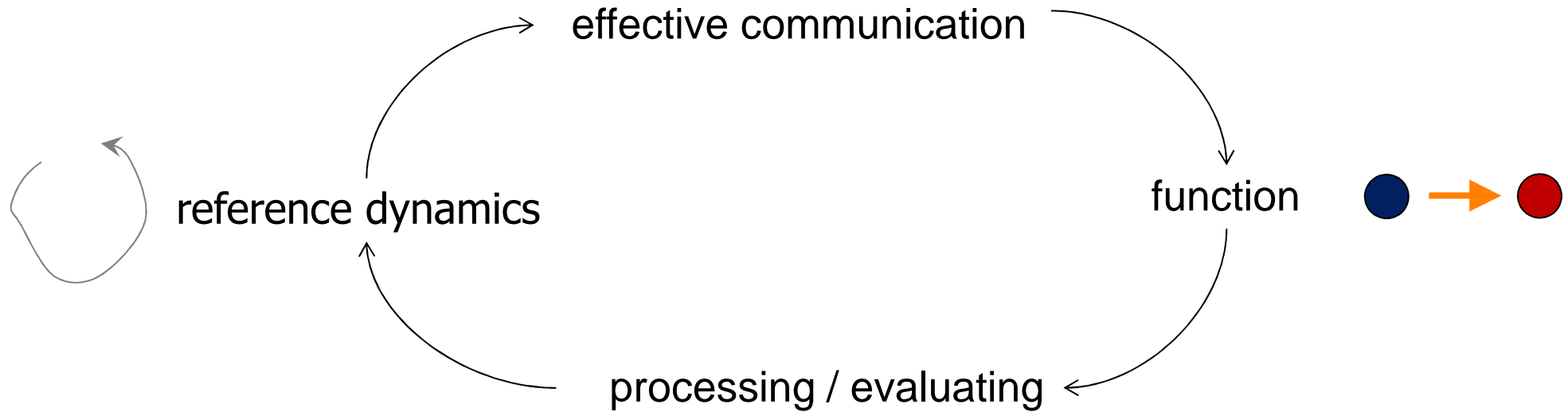
# Self-Reprogramming of Neuronal Networks via Dynamics

- Closing the loop from functional dynamics to dynamic function:

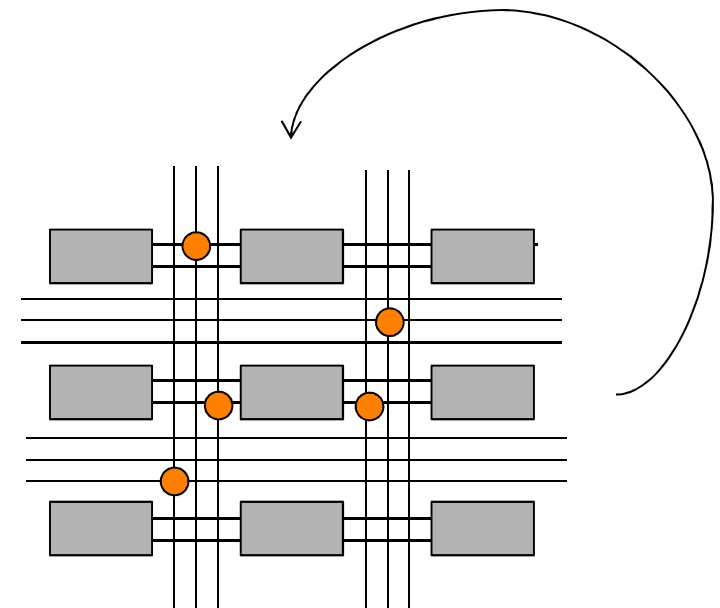


# Self-Reprogramming of Neuronal Networks via Dynamics

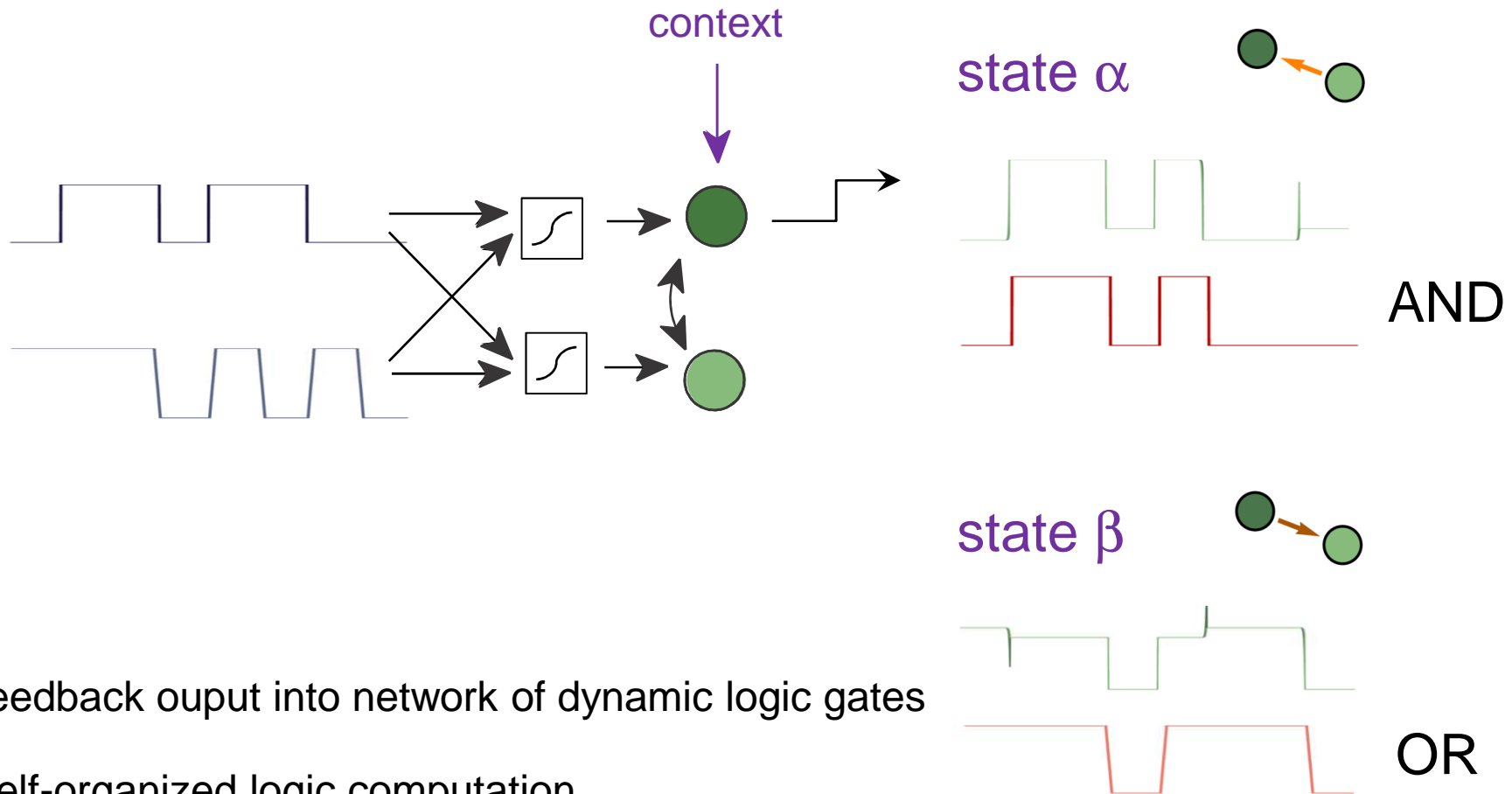
- Closing the loop from functional dynamics to dynamic function:



- Field-Programmable Gate Arrays (FPGAs)



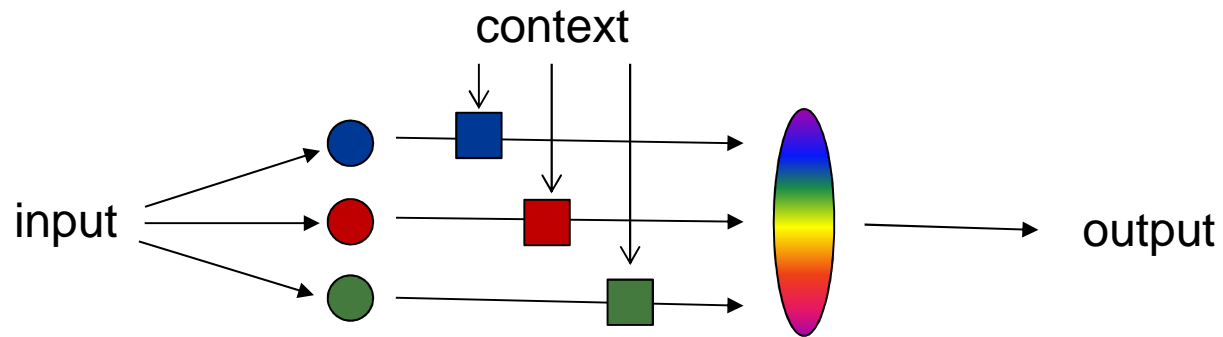
# Dynamic Logic Gates



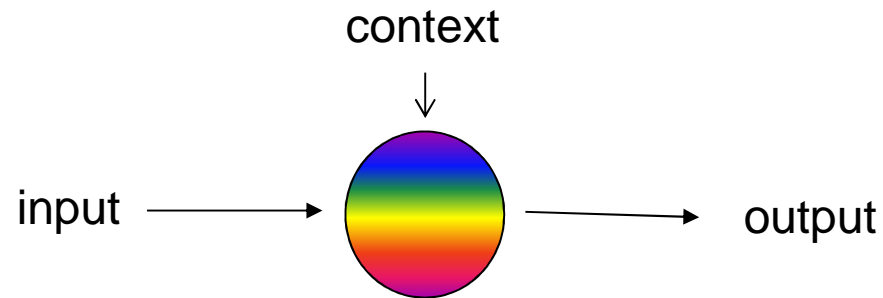
- feedback output into network of dynamic logic gates
- self-organized logic computation
- thermalization of logic computation

# Innate Dynamic Functionality

- Dynamic functionality via appropriate gating of multiple pre-computed functions



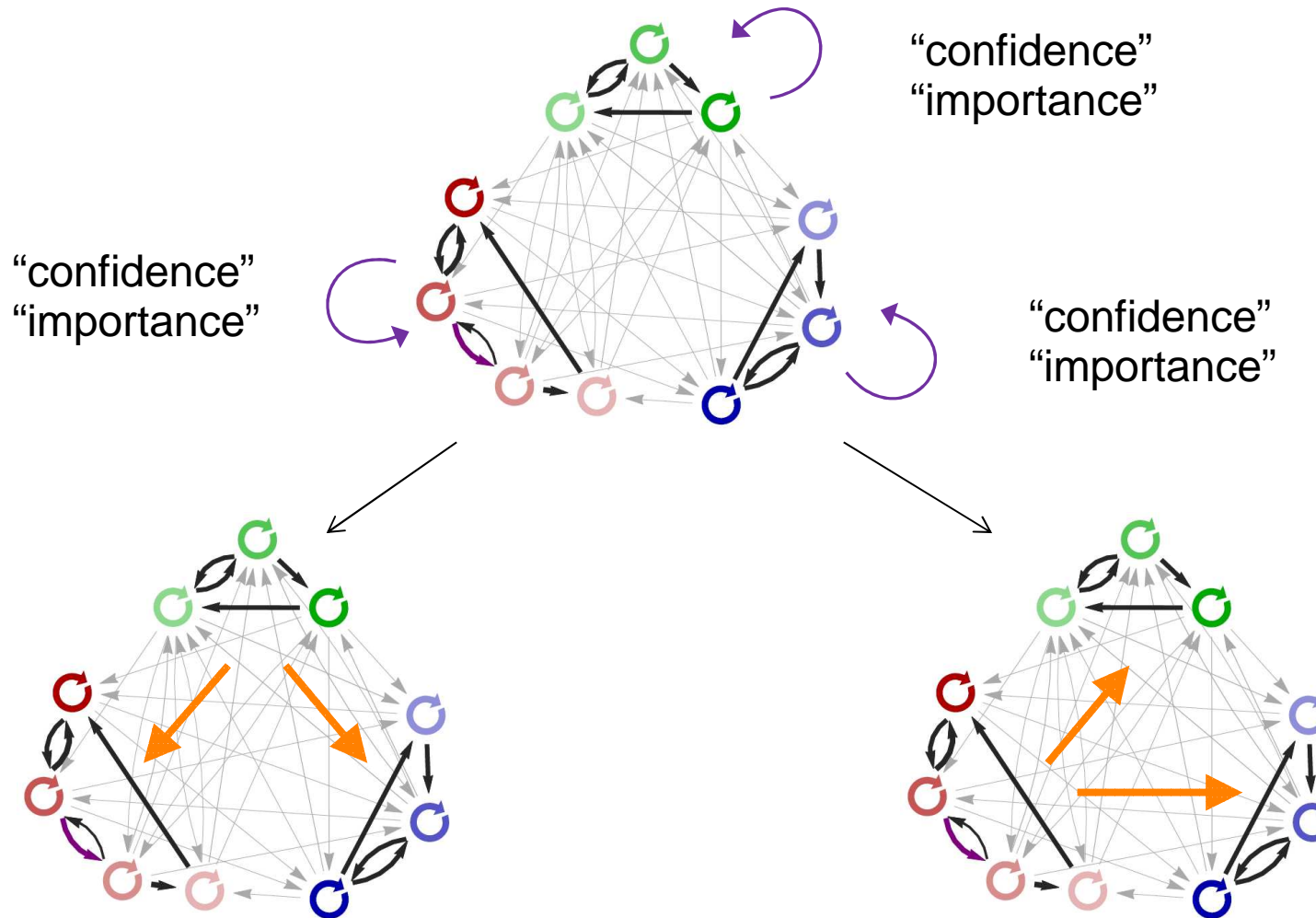
- Can we achieve direct flexible functionality ?



- oscillatory Hopfield networks
- learning innate flexible function in firing rate models

# Self-Organized Information Processing

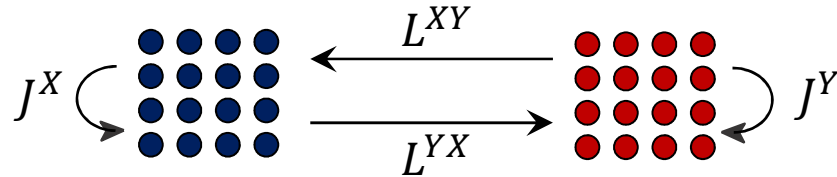
- Oscillatory drive from local importance/confidence signals



- Dynamics self-organize distribution of context information / propagate believe

# Self-Organized Contextual Pattern Recognition

- hierarchical Hopfield network

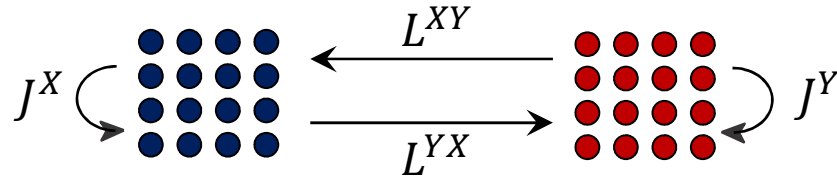


- memory patterns  $J^X$ ,  $J^Y$

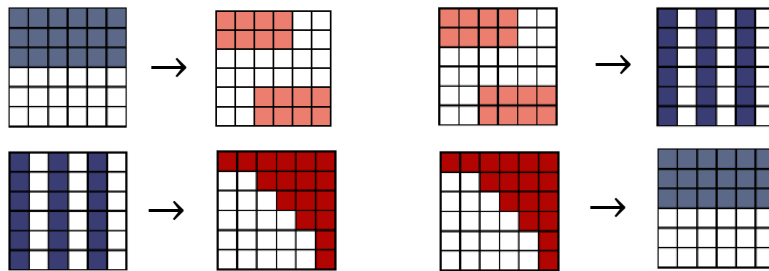


# Self-Organized Contextual Pattern Recognition

- hierarchical Hopfield network



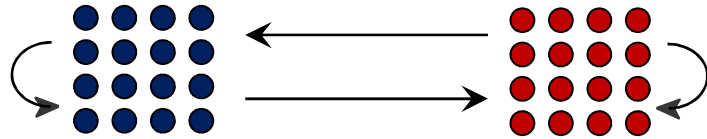
- memory patterns + context rules  $L^{XY}$ ,  $L^{YX}$



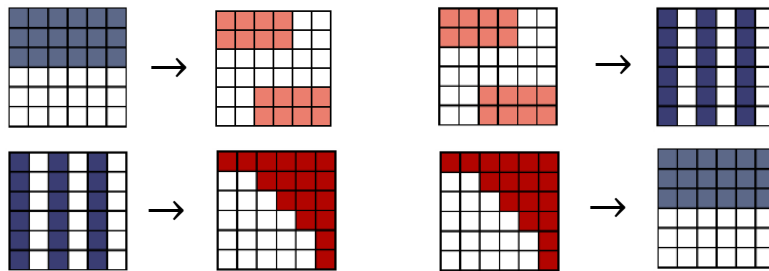


# Self-Organized Contextual Pattern Recognition

- hierarchical Hopfield network



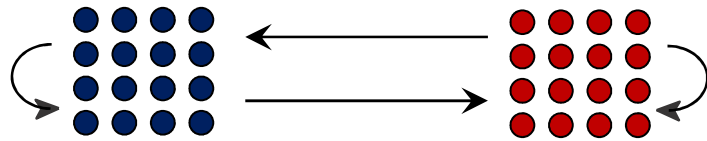
- memory patterns + context rules



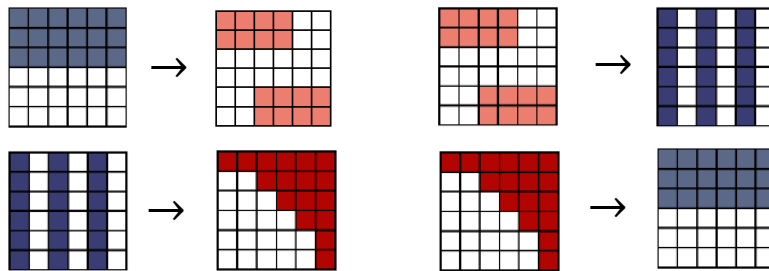
- confidence =  $K - \min(\text{distance to memory patterns}) = \eta$

# Self-Organized Contextual Pattern Recognition

- hierarchical Hopfield network

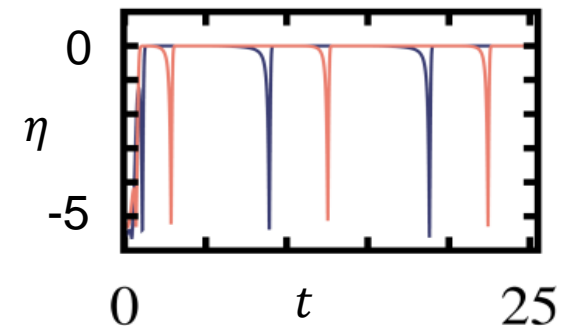
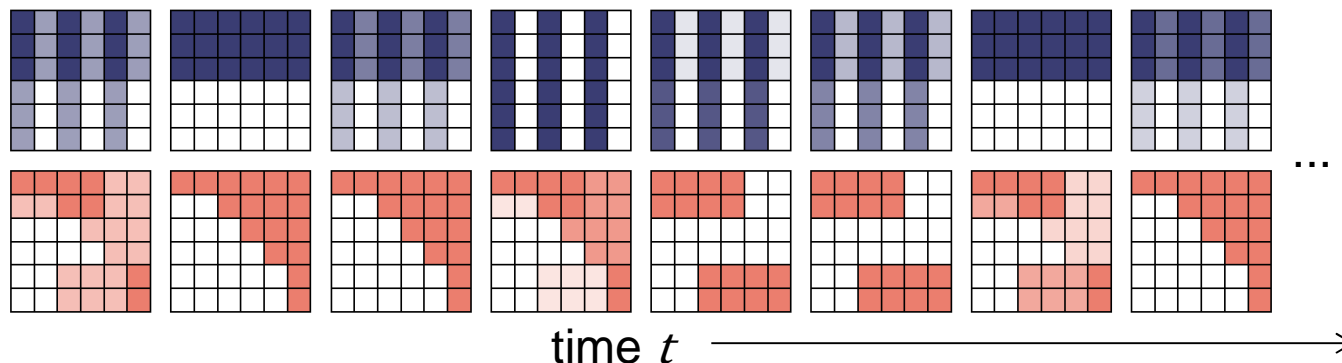


- memory patterns + context rules



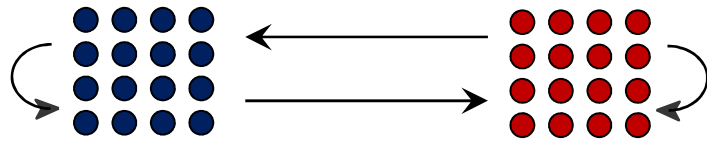
- confidence =  $K - \min(\text{distance to memory patterns}) = \eta$

- classical network (context rules permanently active):

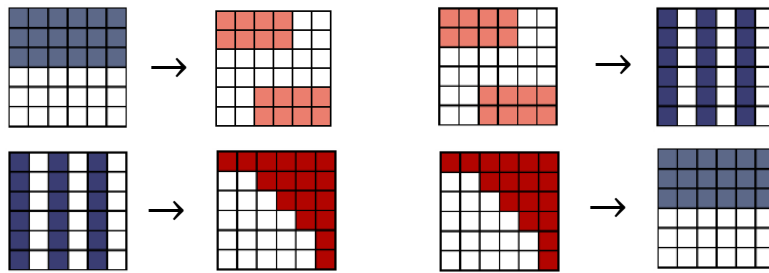


# Self-Organized Contextual Pattern Recognition

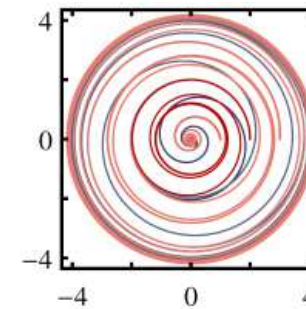
- hierarchical oscillatory Hopfield network



- memory patterns + context rules

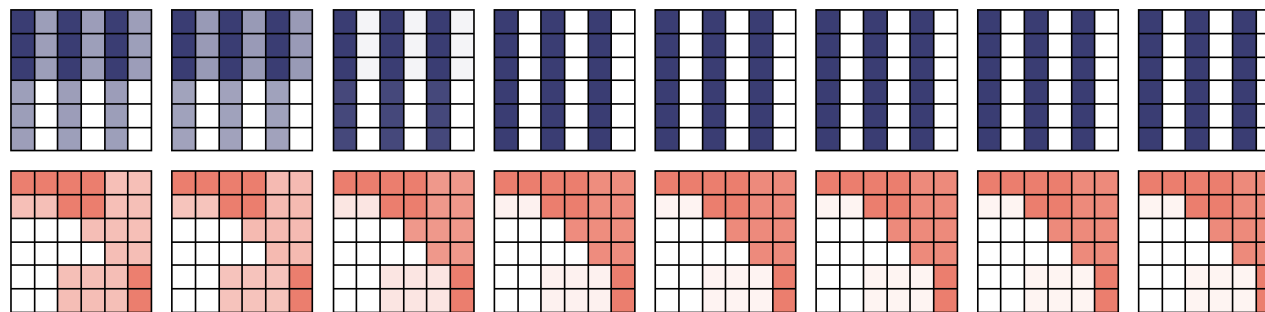


- oscillatory dynamics

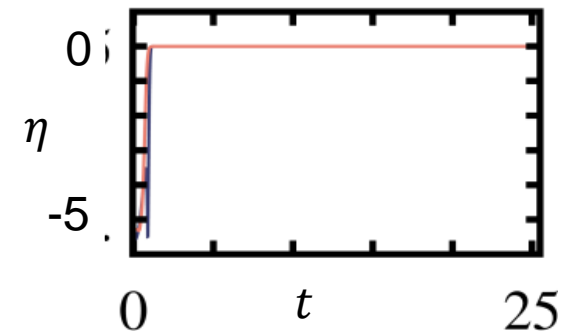


- confidence =  $K - \min(\text{distance to memory patterns}) = \eta$

- self-organized context distribution

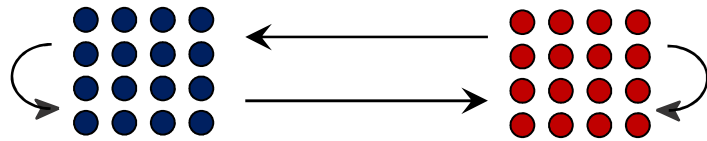


time  $t$  →

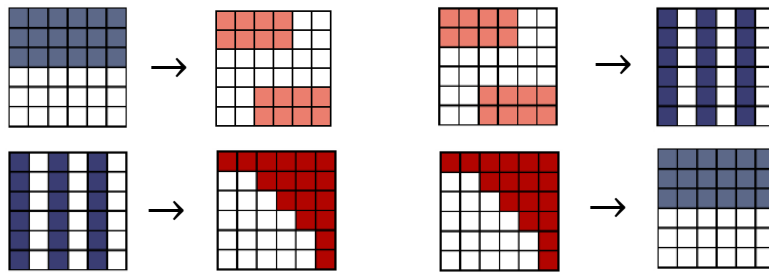


# Self-Organized Contextual Pattern Recognition

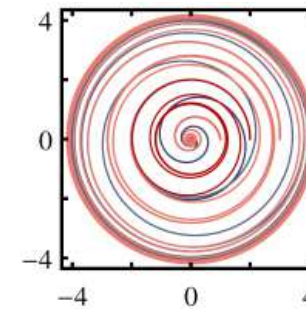
- hierarchical oscillatory Hopfield network



- memory patterns + context rules

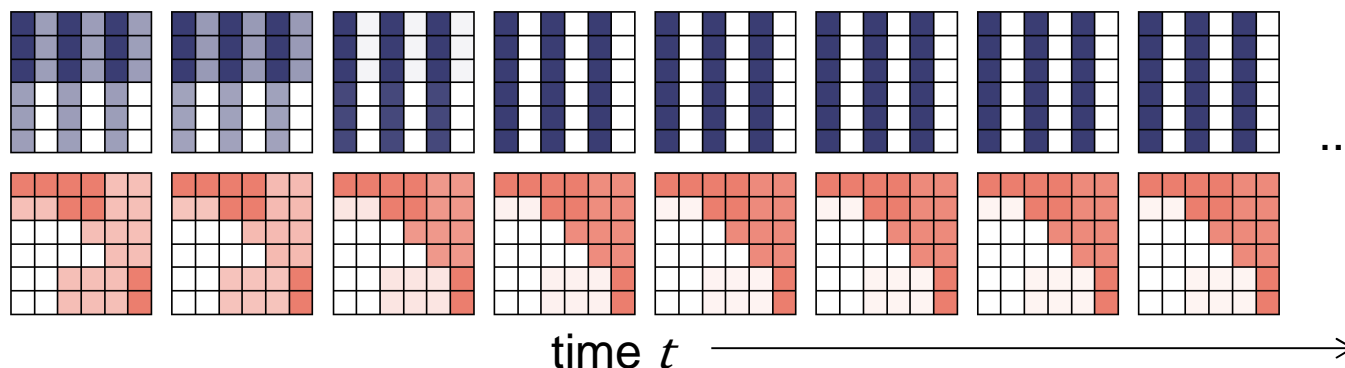


- oscillatory dynamics

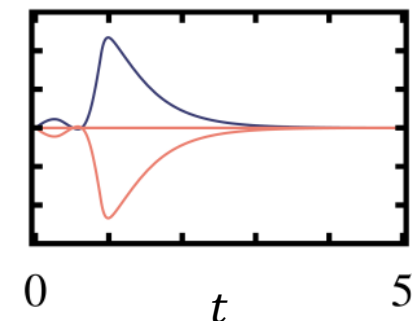


- confidence =  $K - \min(\text{distance to memory patterns}) = \eta$

- self-organized context distribution



effective coupling

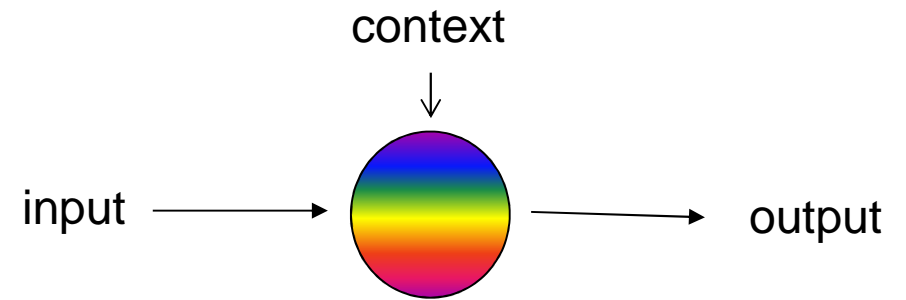


# Outline: Flexible Function in Neuronal Networks

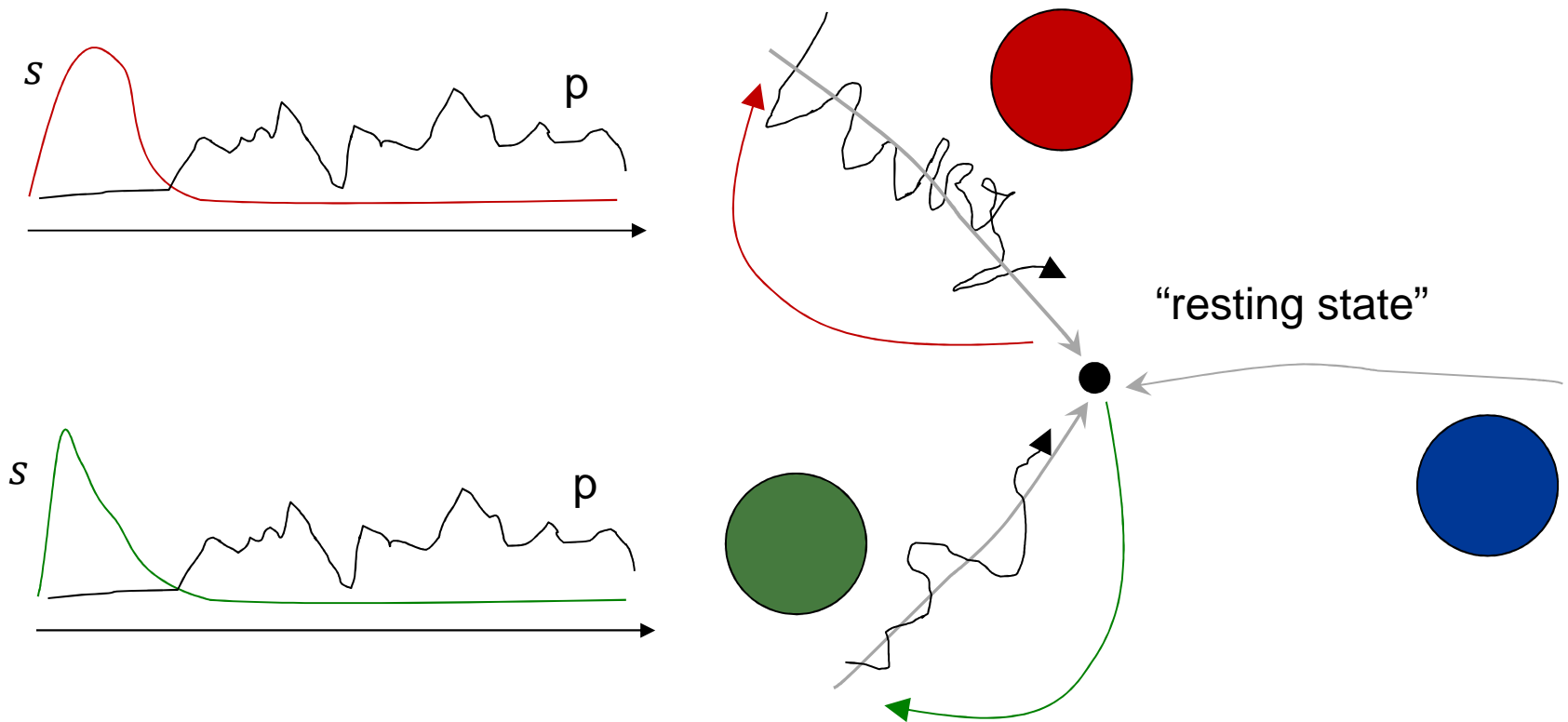
- dynamic information routing in complex networks
  - structural vs. effective connectivity
  - flexible information routing in oscillator networks
  - spiking networks, transient dynamics
- flexible information processing in complex networks
  - oscillatory Hopfield networks
  - self-organized pattern recognition
- **learning flexible function in neuronal networks**
- connections to experiments
  - brain state identification in zebrafish
  - complete brain activity mapping in mouse
- conclusions

# Innate Dynamic Functionality using Transients

- Can we achieve direct flexible functionality ?



- ✓ Flexible computation on top of transients:



# Innate Dynamic Functionality using Transients

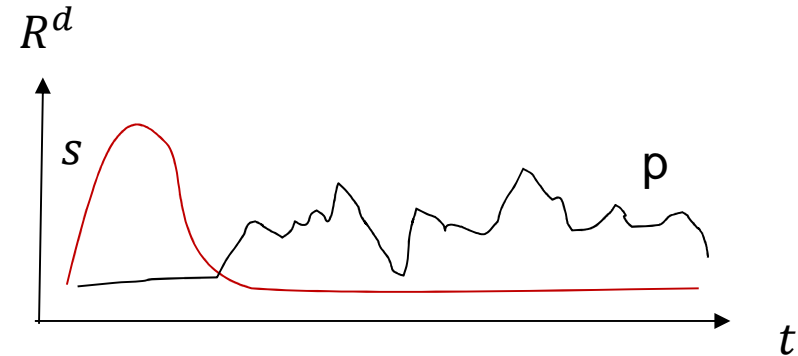
- network dynamics:

$$\dot{x}(t) = f(x(t)) + s(t) + p(t)$$

$x$ : state

$s$ : contextual / steering input

$p$ : input to process



- reference transient  $x_0(t)$ :

$$s = s^{(0)}, p = 0:$$

$$\dot{x}_0(t) = f(x_0(t)) + s^{(0)}(t)$$

- processing along transient ( $p \ll 1$ ):

$$\dot{x}(t) = f(x(t)) + s^{(0)}(t) + p(t)$$

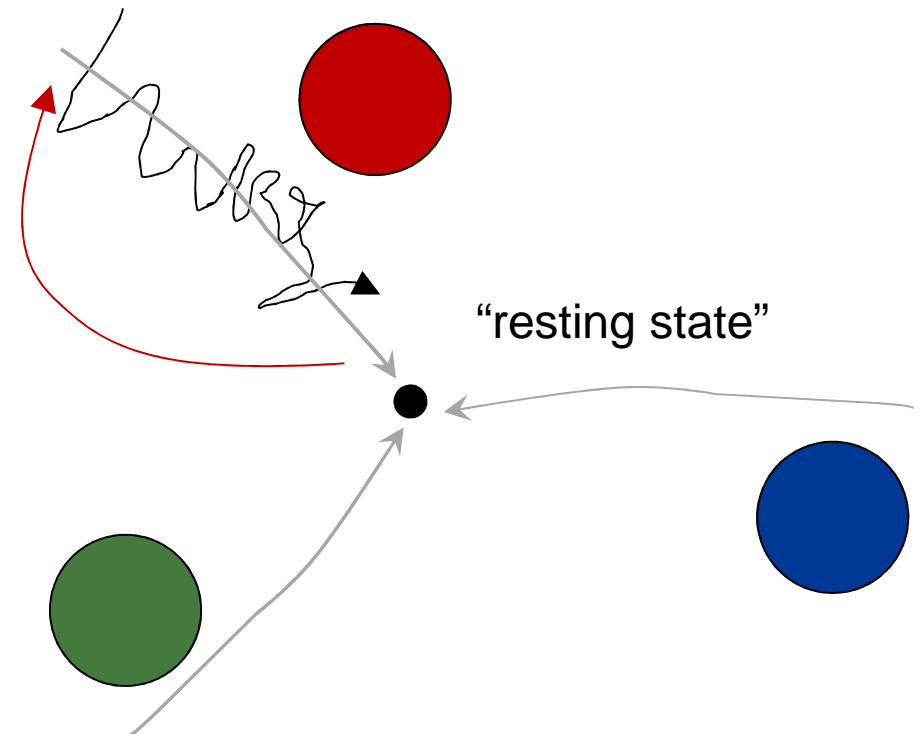
$$x(t) = x^{(0)}(t) + \delta x(t)$$

$$\dot{\delta x}(t) \approx Df(x^{(0)}(t))\delta x + p(t)$$

→ linear time varying system:

$$\delta x(t) = \int_0^t \Phi^{(0)}(t, \tau) p(\tau) d\tau$$

$\Phi^{(0)}$   $x^{(0)}$  dependent



# Taming Chaos in Neuronal Networks

- firing rate network:

$$\dot{x}_i = -x_i + g \sum_j w_{ij} r(x_j)$$

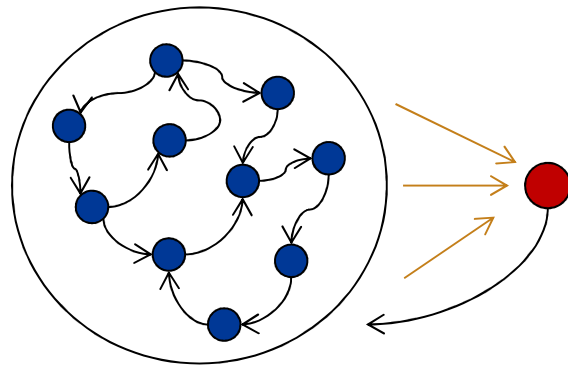
- network transitions to chaos when gain  $g$  is increased

[Sompolinsky, Crisanti, Sommers, PRL 1988]

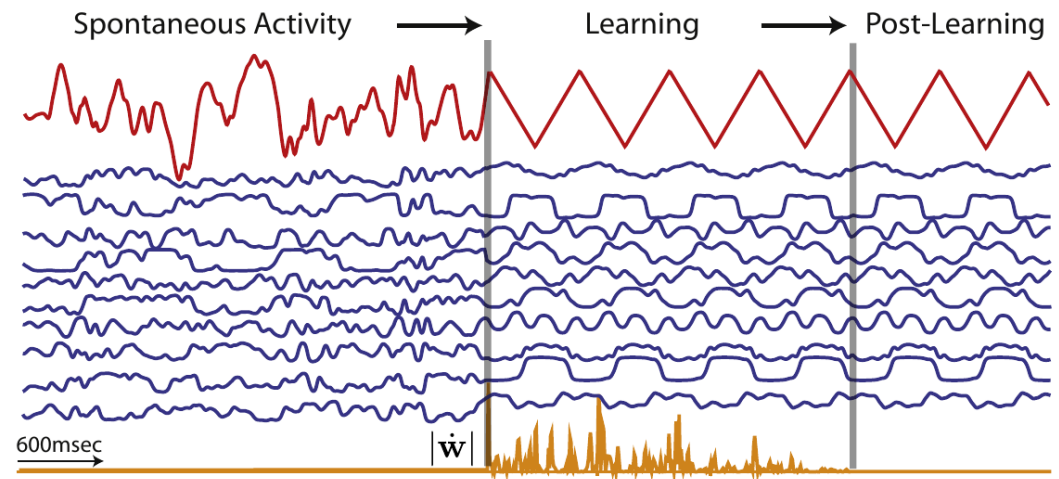
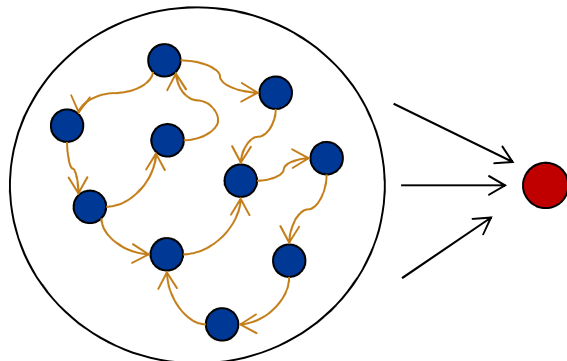
- learning output or recurrent weights to stabilize transients (FORCE learning)

[Sussillo, Abbott, Neuron 2009]

[Laje, Buonomano, Nat Neuro, 2013]



or

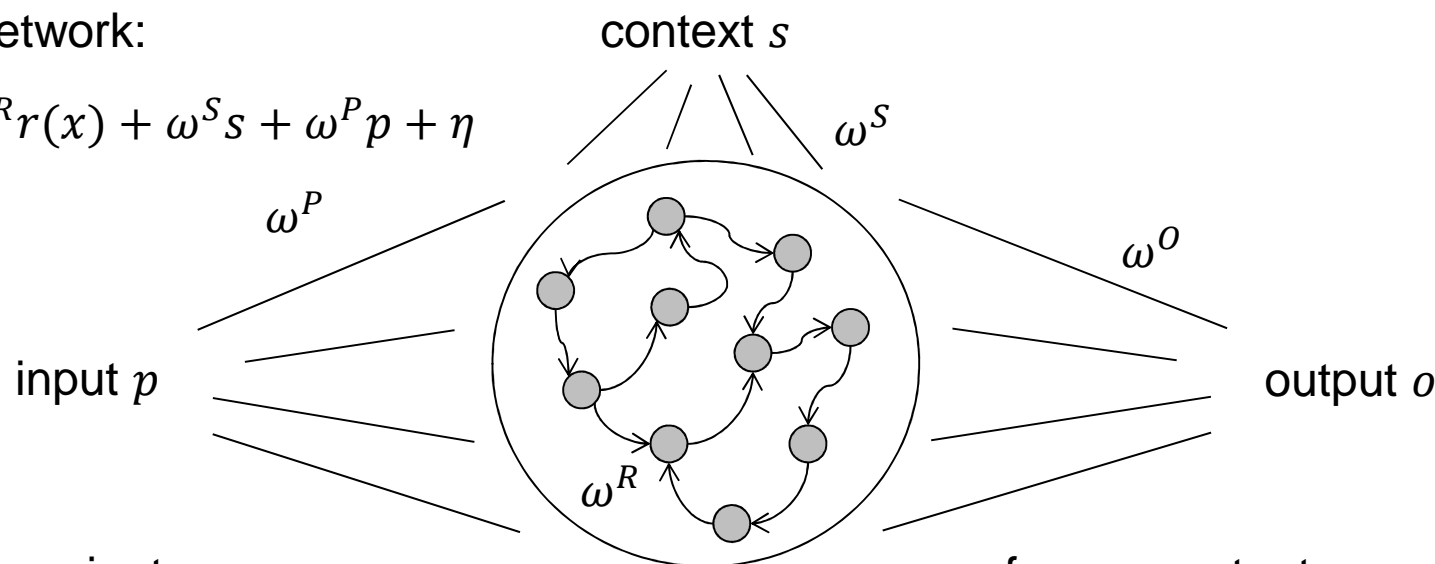




# Flexible Network Function Along Stabilized Transients

- firing rate network:

$$\dot{x} = -x + \omega^R r(x) + \omega^S s + \omega^P p + \eta$$



- reference transients:

$$\dot{x}^{(0)} = -x^{(0)} + \omega^R r(x^{(0)}) + \omega^S s^{(0)}$$

trained to stabilize transients

- reference output:

$$o(t) = \omega^O r(x) = o^{(0)} = \text{const}$$

trained to stabilize output

- small inputs:

$$\delta \dot{x}(t) = \omega^R DR(x^{(0)}(t)) \delta x(t) + \omega^P p$$

$$\delta o(t) = \omega^O DR(x^{(0)}(t)) \delta x(t)$$

- reference state dependent filtering:

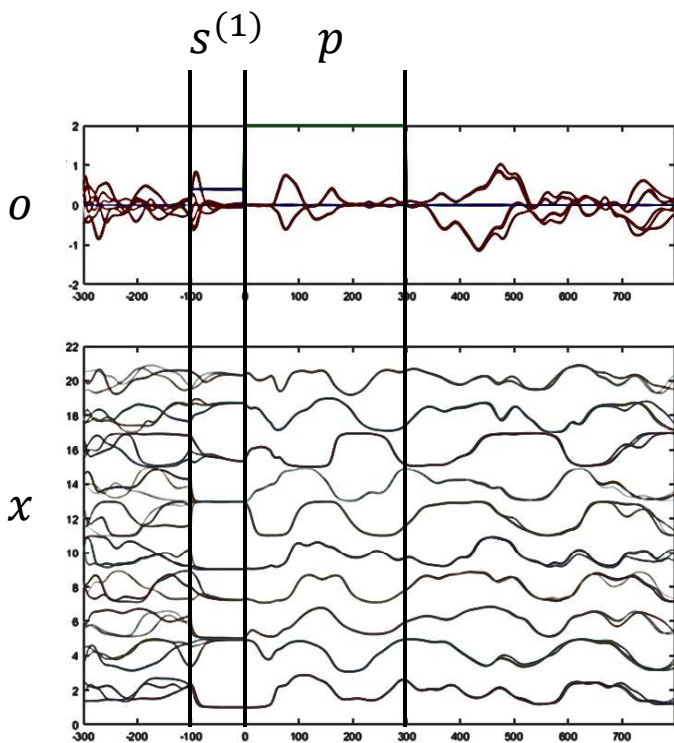
$$\delta o = \omega^O DR(x^{(0)}(t)) \int_0^t \Phi^{(0)}(t, \tau) \omega^P p(\tau) d\tau$$

train to achieve specific function

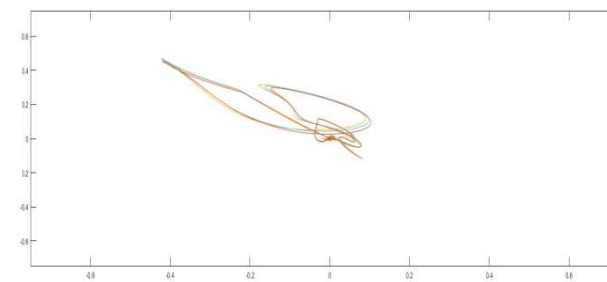
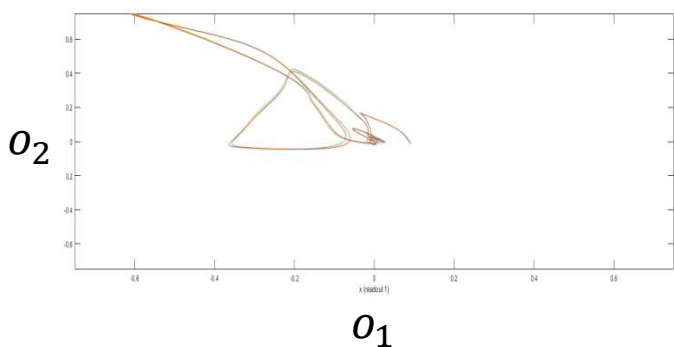
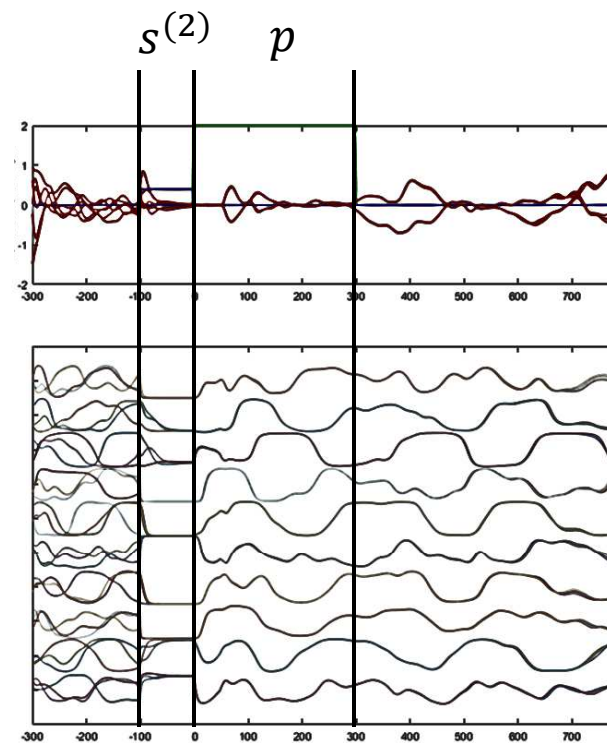
# Flexible Network Function Along Stabilized Transients

- example:

– reference transient 1



– reference transient 2



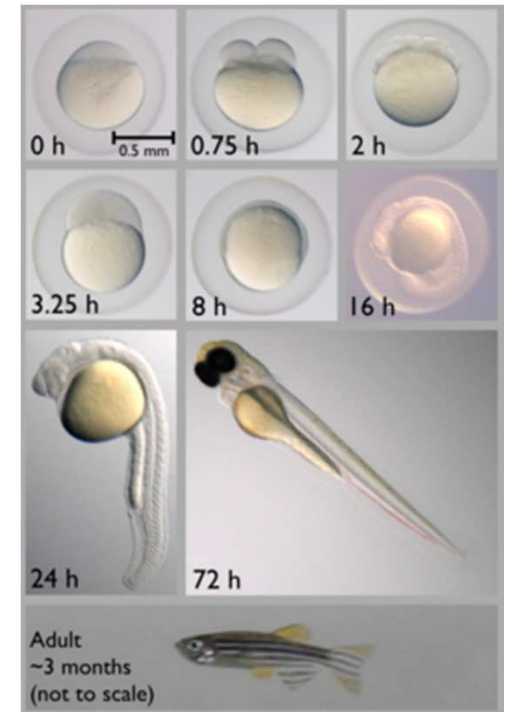
# Outline: Flexible Function in Neuronal Networks

- dynamic information routing in complex networks
  - structural vs. effective connectivity
  - flexible information routing in oscillator networks
  - spiking networks, transient dynamics
- flexible information processing in complex networks
  - oscillatory Hopfield networks
  - self-organized pattern recognition
- learning flexible function in neuronal networks
- **connections to experiments**
  - **brain state identification in zebrafish**
  - **complete brain activity mapping in mouse**
- conclusions

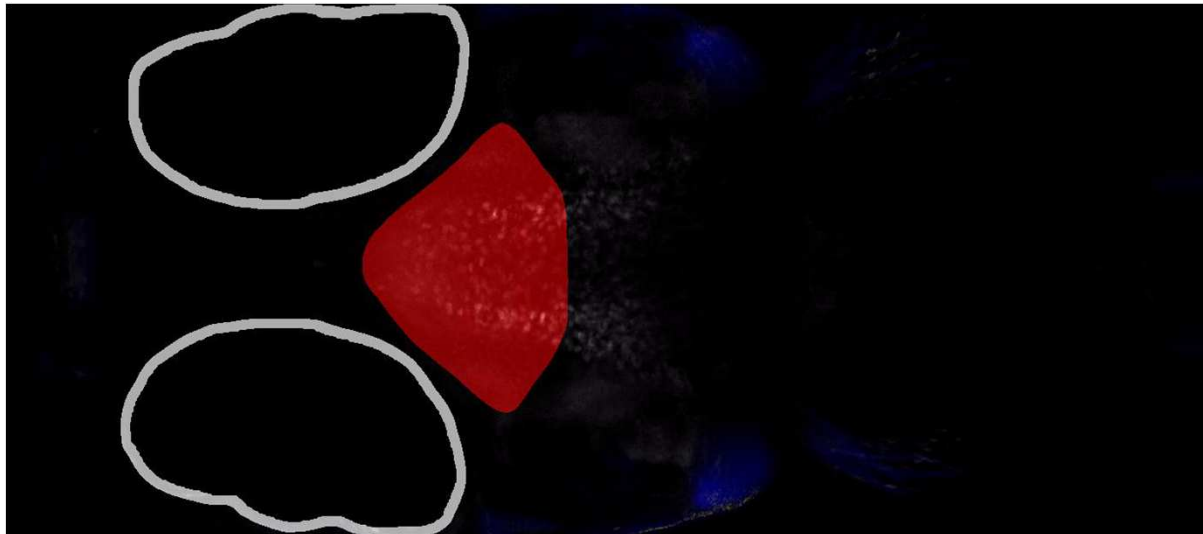
# The Zebrafish Larvae Brain

Zebrafish:

- tropical fish from Ganges River in East India and Burma
- used to study development (clear eggs, vertebrate)
- transparent larvae – imaging ‘easy’
- Z-brain: zebrafish brain atlas

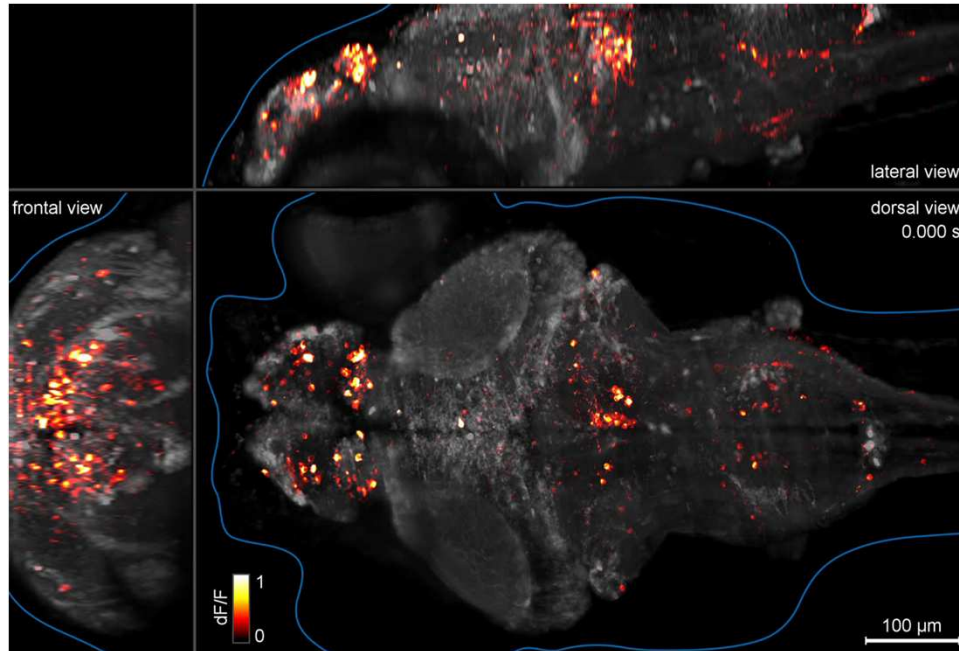


[Randlett, et al, Nature Methods 2015]

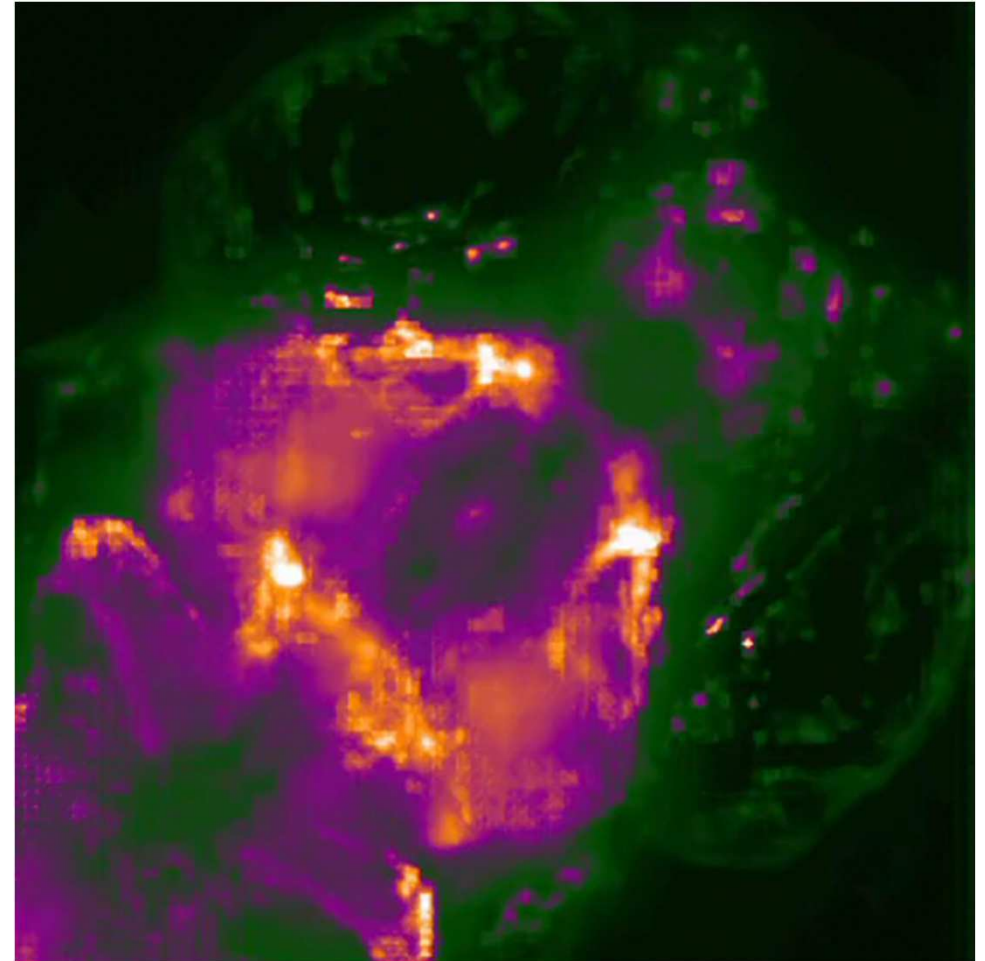


# Zebrafish Whole Brain Imaging

- light sheet imaging

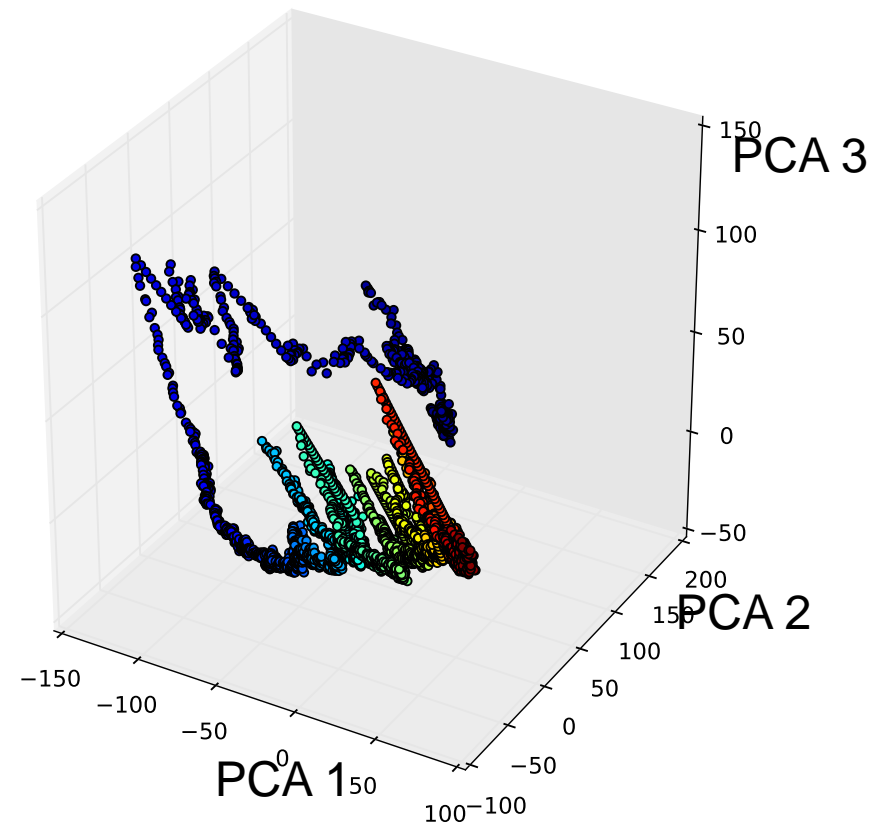
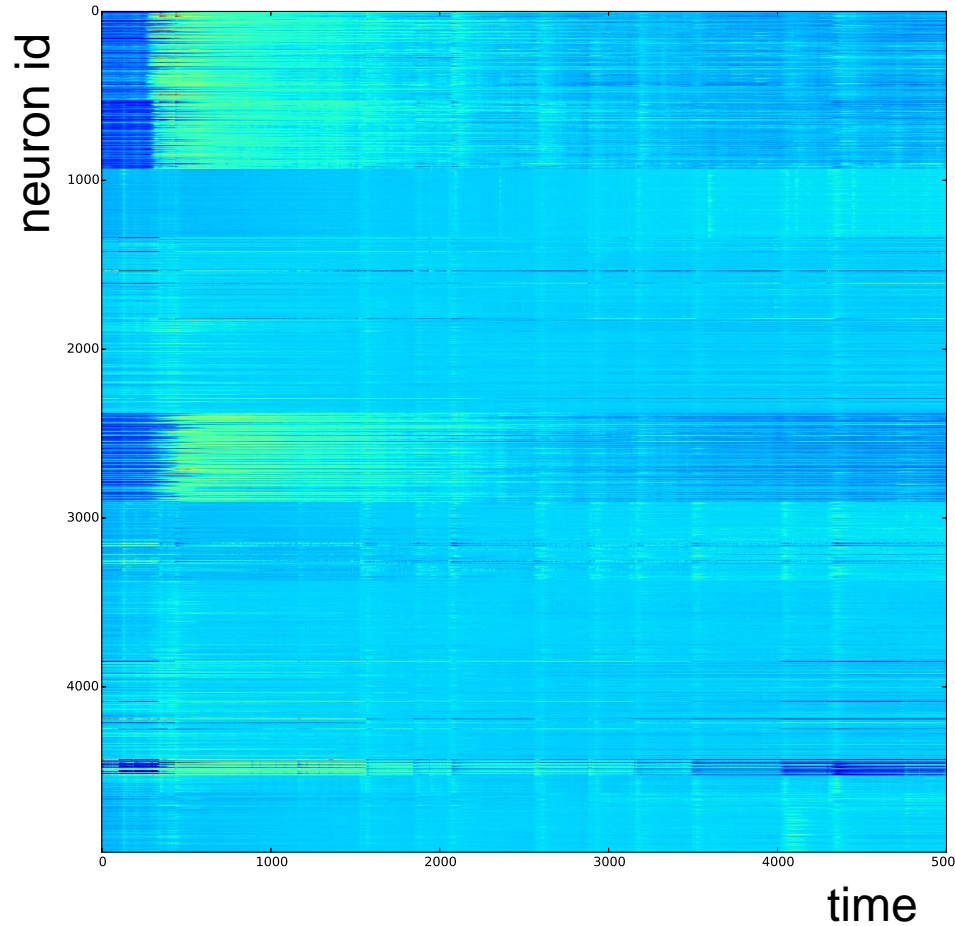


- light field deconvolution imaging



[Vladimirov, ..., Ahrens, Nature Methods 2014] [Prevendel, ..., Vaziri et al, Nature Methods 2014]

# Zebrafish Whole Brain Dynamics - PCA



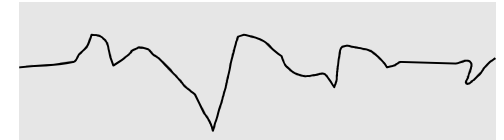
Data: [Prevedel,...,Vaziri et al, Nature Methods 2014]

# Dynamical Sate Identification

- single auto-regressive (AR) processes  $\mathbf{M}_1$

$$x_t = a_0 + a_1x_{t-1} + a_2x_{t-2} + \dots + a_Kx_{t-K} + \varepsilon_t$$

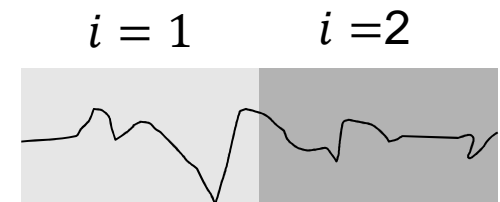
- log likelihood:  $l_1 = -\frac{N}{2}(1 + \log(2\pi)) - \frac{1}{2}N \log(\hat{\sigma})^2$
- #parameter:  $k_1 = K + 1$



- multiple AR processes for non-stationary time series  $\mathbf{M}_2$

$$x_t = a_{0,i} + \sum_m a_{m,i}x_{t-m} + \varepsilon_{i,t} \quad \begin{array}{l} 0 \leq t < t_1, i = 1 \\ t_1 \leq t < t_2, i = 2 \end{array}$$

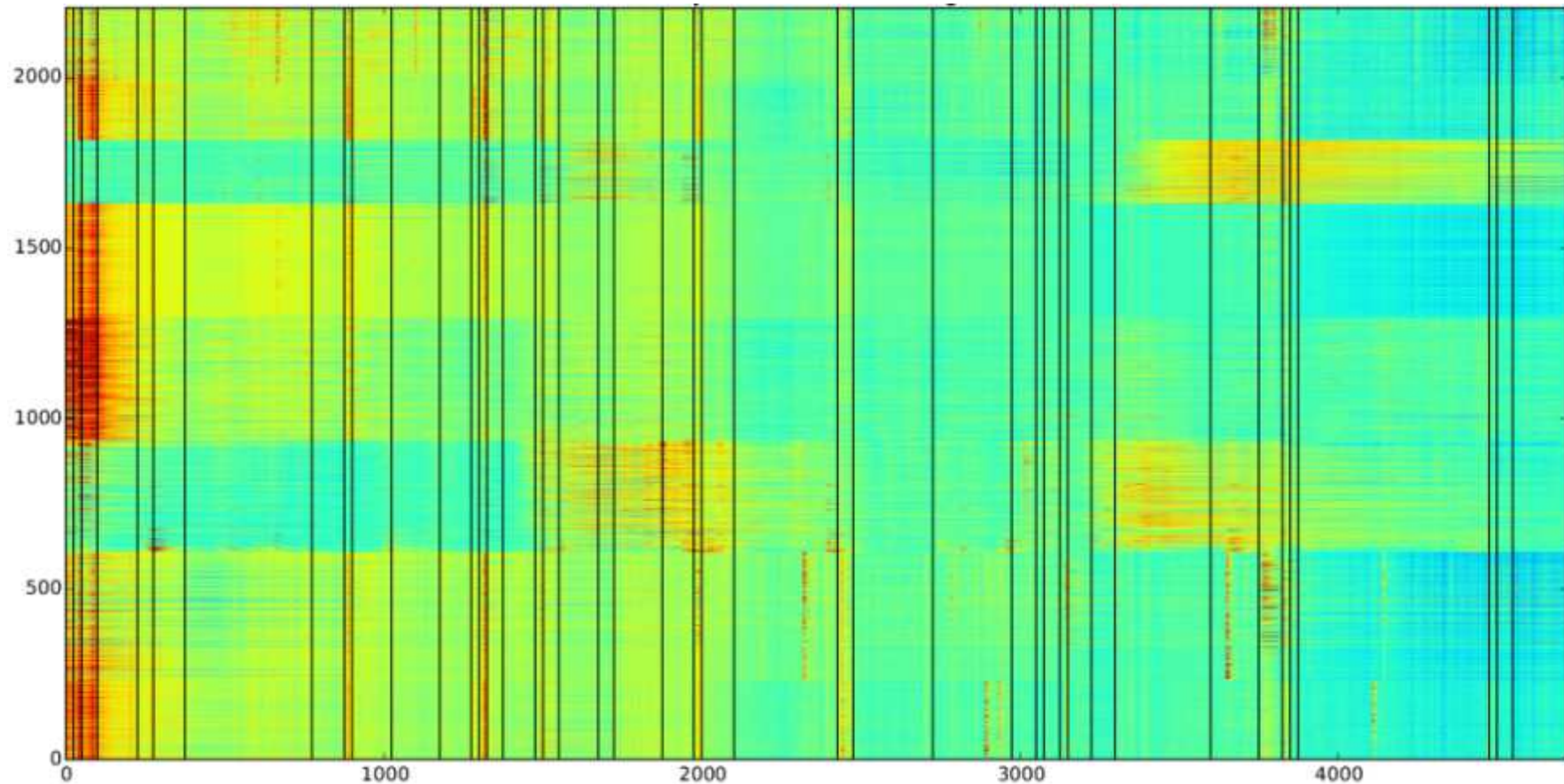
- log likelihood:  $l_2 = -\frac{N}{2}(1 + \log(2\pi)) - \frac{1}{2}\sum N_i \log(\hat{\sigma}_i)^2$
- #parameter:  $k_2 = 2K + 2$



- max AIC to select if a switch has occurred:

$$\text{AIC}_j = l_j - k_j$$

# Dynamical State Identification in Whole Brain Zebrafish Recordings



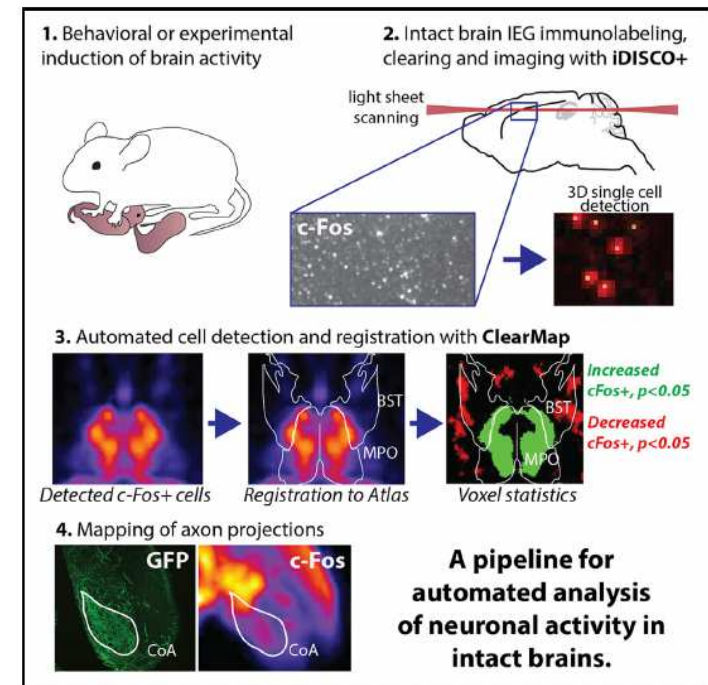
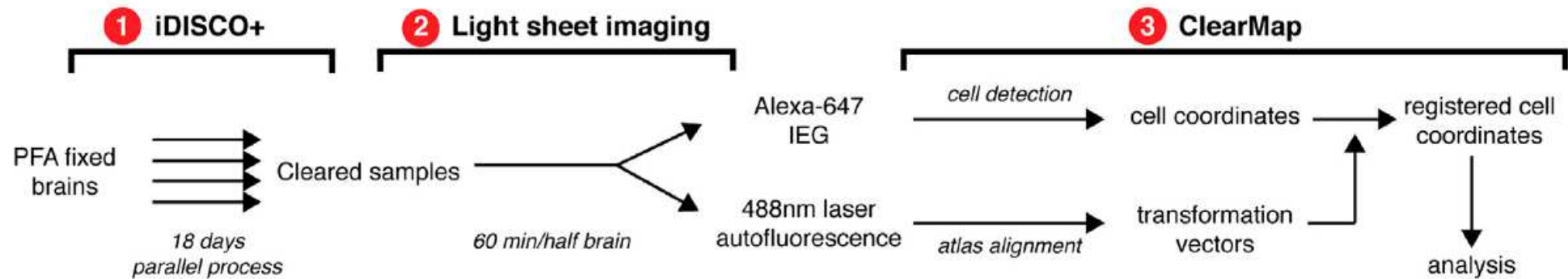
- fast
- takes into account all neurons equally
- extensions to groups of neurons (multi-dimensional ARs)



# Outline: Flexible Function in Neuronal Networks

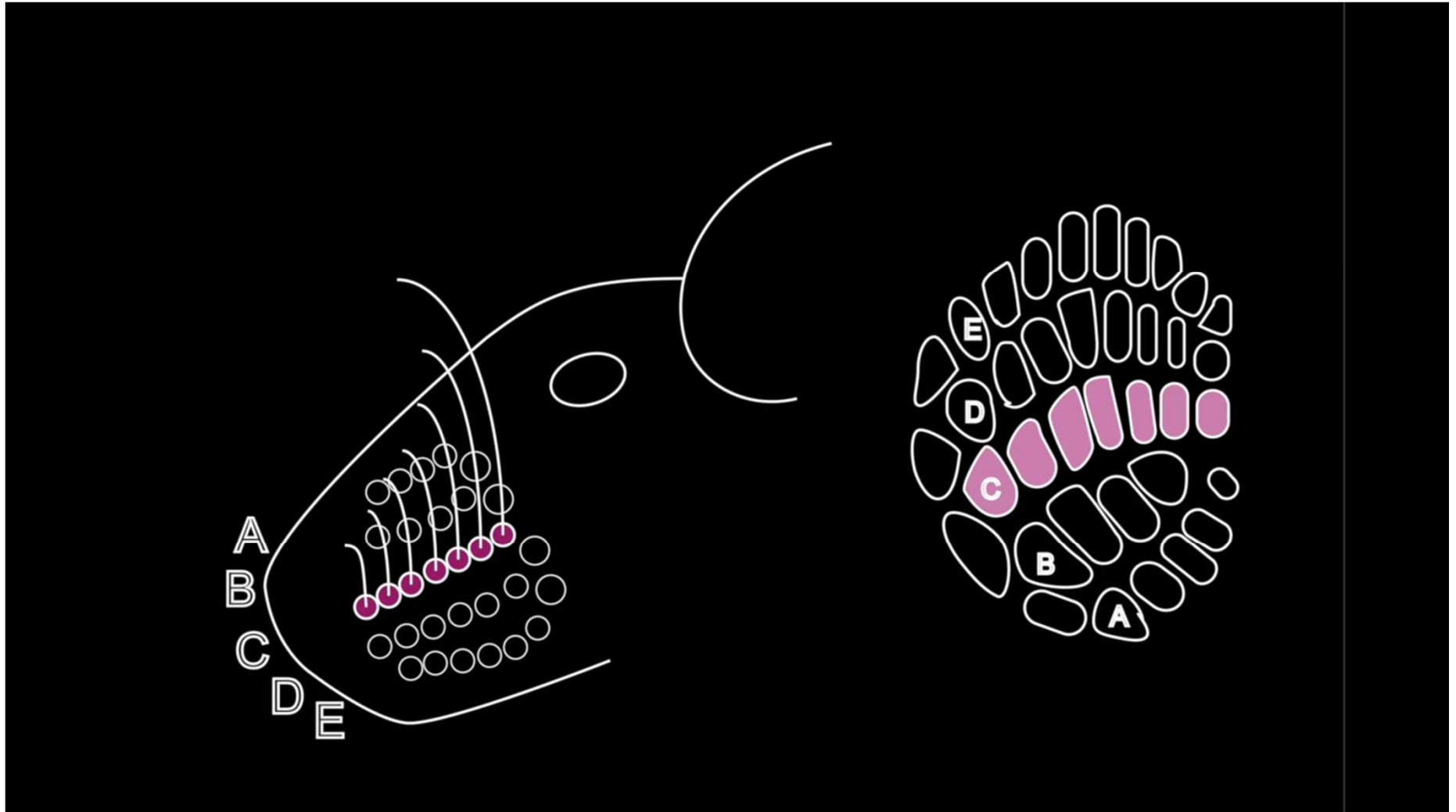
- dynamic information routing in complex networks
  - structural vs. effective connectivity
  - flexible information routing in oscillator networks
  - spiking networks, transient dynamics
- flexible information processing in complex networks
  - oscillatory Hopfield networks
  - self-organized pattern recognition
- learning flexible function in neuronal networks
- **connections to experiments**
  - **brain state identification in zebrafish**
  - **complete brain activity mapping in mouse**
- conclusions

# Complete mapping of cellular brain activity by automated volume analysis of immediate early genes



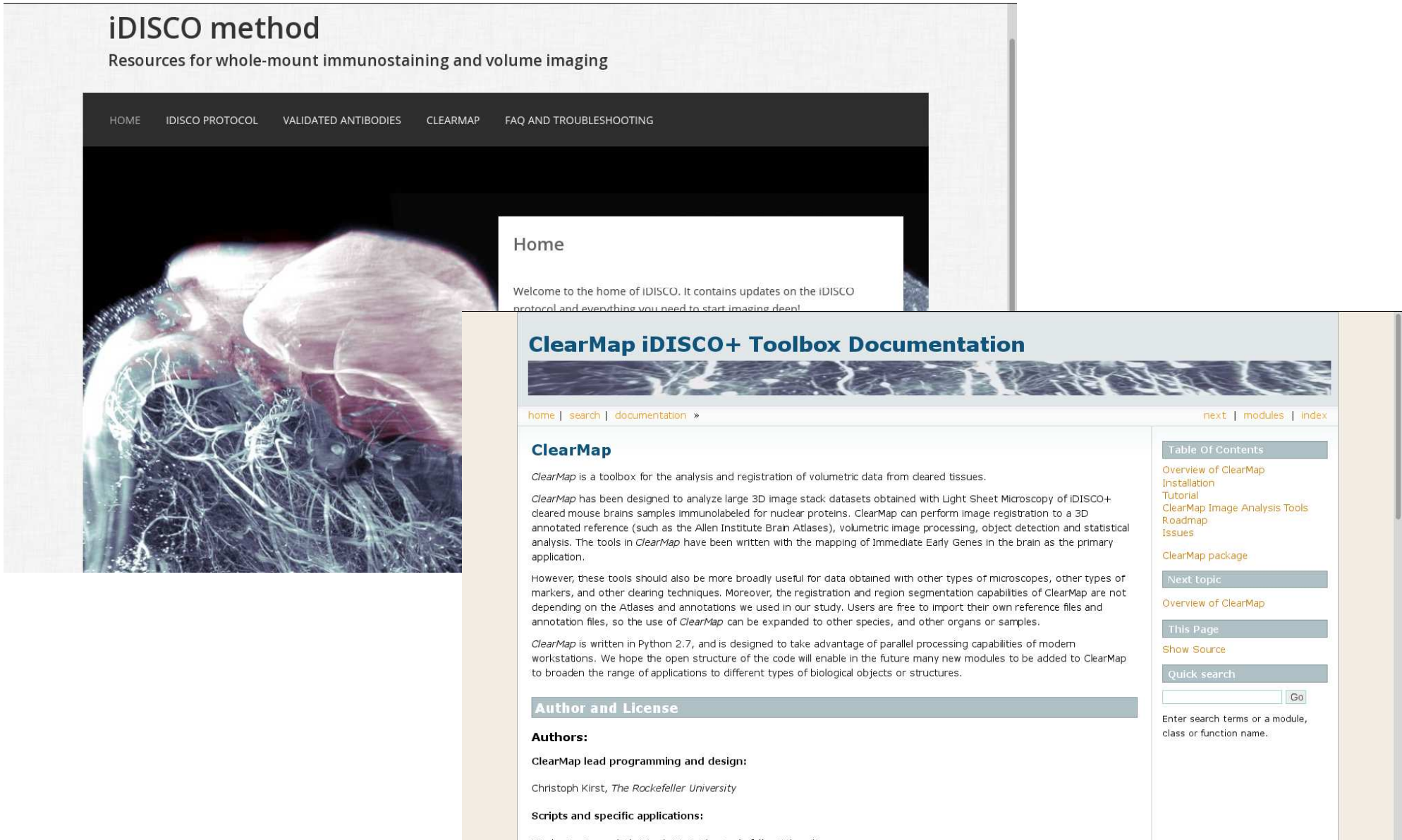
[Renier\*, . Adams\*, Kirst\*, Wu\*, ..., Dulac, Osten, Tessier-Lavigne, Cell 2016]

# Complete mapping of cellular brain activity by automated volume analysis of immediate early genes



[Renier\*, . Adams\*, Kirst\*, Wu\*,..., Dulac, Osten, Tessier-Lavigne, Cell 2016]

https://idisco.info/



The image shows a composite of two web pages. The top page is the iDISCO method website, featuring a navigation menu with links for HOME, IDISCO PROTOCOL, VALIDATED ANTIBODIES, CLEARMAP, and FAQ AND TROUBLESHOOTING. Below the menu is a large image of a cleared mouse brain with a 3D reconstruction of its neural network. A white box on the page says "Home" and "Welcome to the home of iDISCO. It contains updates on the iDISCO protocol and everything you need to start imaging deep!".

The bottom page is the "ClearMap iDISCO+ Toolbox Documentation" page. It has a navigation bar with "home | search | documentation" and "next | modules | index". The main content area is titled "ClearMap" and contains the following text:

**ClearMap**

*ClearMap* is a toolbox for the analysis and registration of volumetric data from cleared tissues.

*ClearMap* has been designed to analyze large 3D image stacks obtained with Light Sheet Microscopy of iDISCO+ cleared mouse brains samples immunolabeled for nuclear proteins. *ClearMap* can perform image registration to a 3D annotated reference (such as the Allen Institute Brain Atlases), volumetric image processing, object detection and statistical analysis. The tools in *ClearMap* have been written with the mapping of Immediate Early Genes in the brain as the primary application.

However, these tools should also be more broadly useful for data obtained with other types of microscopes, other types of markers, and other clearing techniques. Moreover, the registration and region segmentation capabilities of *ClearMap* are not depending on the Atlases and annotations we used in our study. Users are free to import their own reference files and annotation files, so the use of *ClearMap* can be expanded to other species, and other organs or samples.

*ClearMap* is written in Python 2.7, and is designed to take advantage of parallel processing capabilities of modern workstations. We hope the open structure of the code will enable in the future many new modules to be added to *ClearMap* to broaden the range of applications to different types of biological objects or structures.

**Author and License**

**Authors:**

**ClearMap lead programming and design:**

Christoph Kirst, *The Rockefeller University*

**Scripts and specific applications:**

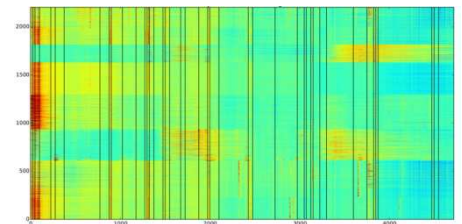
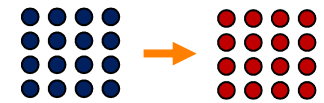
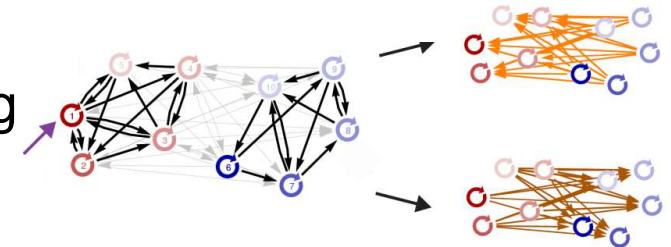
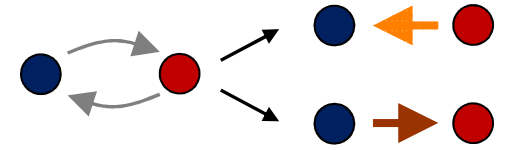
The right sidebar of the documentation page contains a "Table of Contents" with links to Overview of ClearMap, Installation, Tutorial, ClearMap Image Analysis Tools, Roadmap, and Issues. Below that are buttons for "ClearMap package", "Next topic", "Overview of ClearMap", "This Page", and "Show Source". At the bottom is a "Quick search" section with an input field and a "Go" button.

[Renier\*, . Adams\*, Kirst\*, Wu\* ...,  
Dulac, Osten, Tessier-Lavigne, Cell 2016]

https://github.com/ChristophKirst/ClearMap

# Conclusion

- reference dynamics  $\Rightarrow$  effective network  $\Rightarrow$  function
- theory for information routing in phase oscillator networks
- flexible dynamic information routing on top of dynamics:
  - multi-stable dynamical states  
→ fast and self-organized information re-routing
  - hierarchical networks  
→ action at a distance, combinatorial IRPs
- self-organized information processing by closing the loop
  - contextual pattern recognition in oscillatory Hopfield nets
- learning flexible function on top of dynamical reference states
- novel approaches to the analysis of large scale neural activity



# Acknowledgments

Marc Timme

Max Planck for Dynamics & self-Organization, Göttingen

Demian Battaglia

University Université Aix-Marseille, Institut de Neurosciences des Systèmes

Alipasha Vaziri

Marc Tessier-Lavigne

Marcello Magnasco

Cori Bargmann

Gaby Maimon

Nicolas Reiner

Shay Stern

Jonathan Green

Carl Modes

Center for Physics and Biology and Kavli Neural Systems Institute  
The Rockefeller University, New York City, USA

## You for your Attention !