

# Nascent Program in Computational Neuroscience

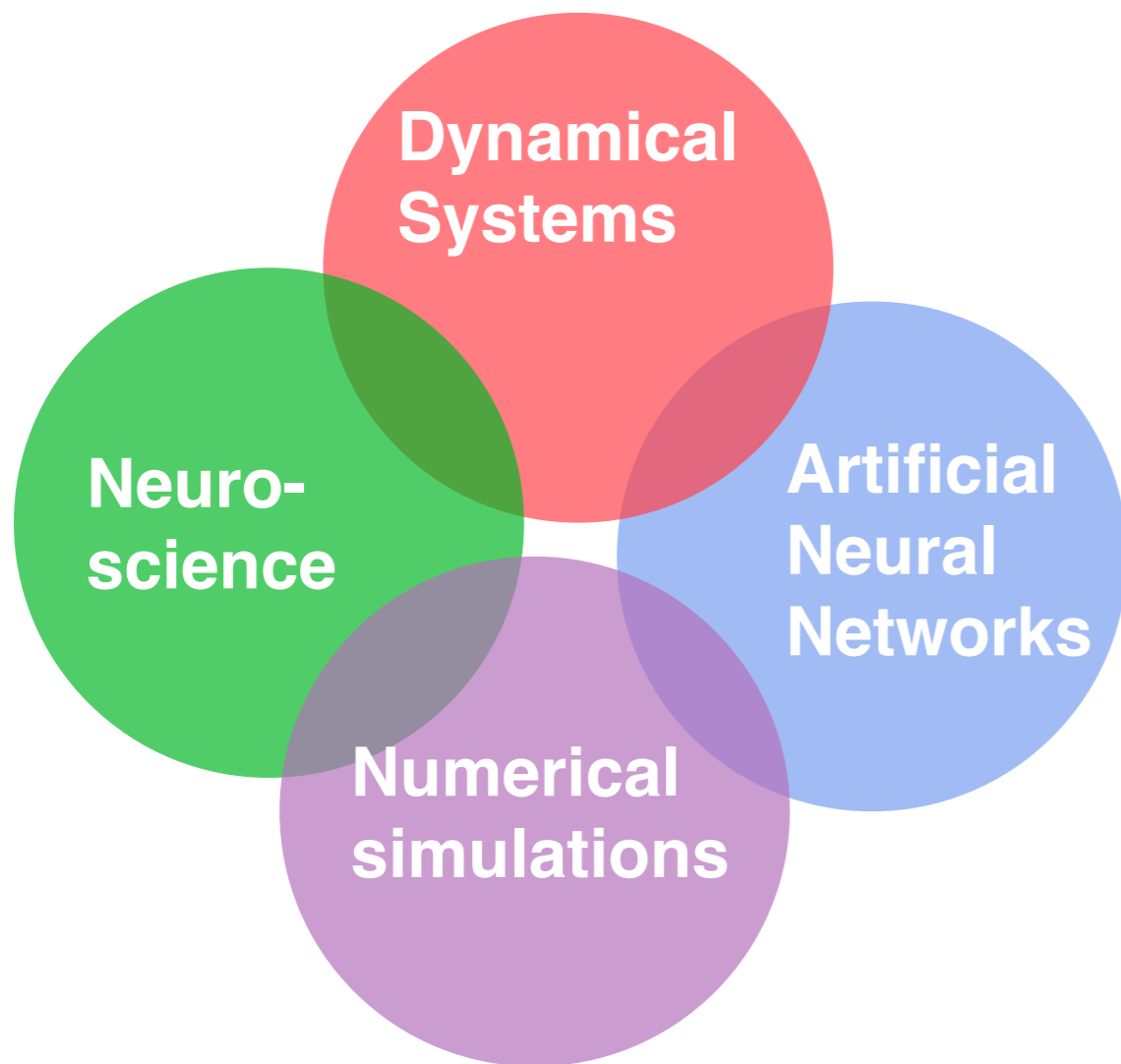
Gordon Erlebacher

# Roadmap

June 2014 - April 2015

- How it all started
- Biology and modeling
- Artificial neural networks
- The learning process
- An algebra for polychrony
- New Collaboration
- Ending thoughts

# Computational Neuroscience



Artificial Neural Networks  
Biological networks

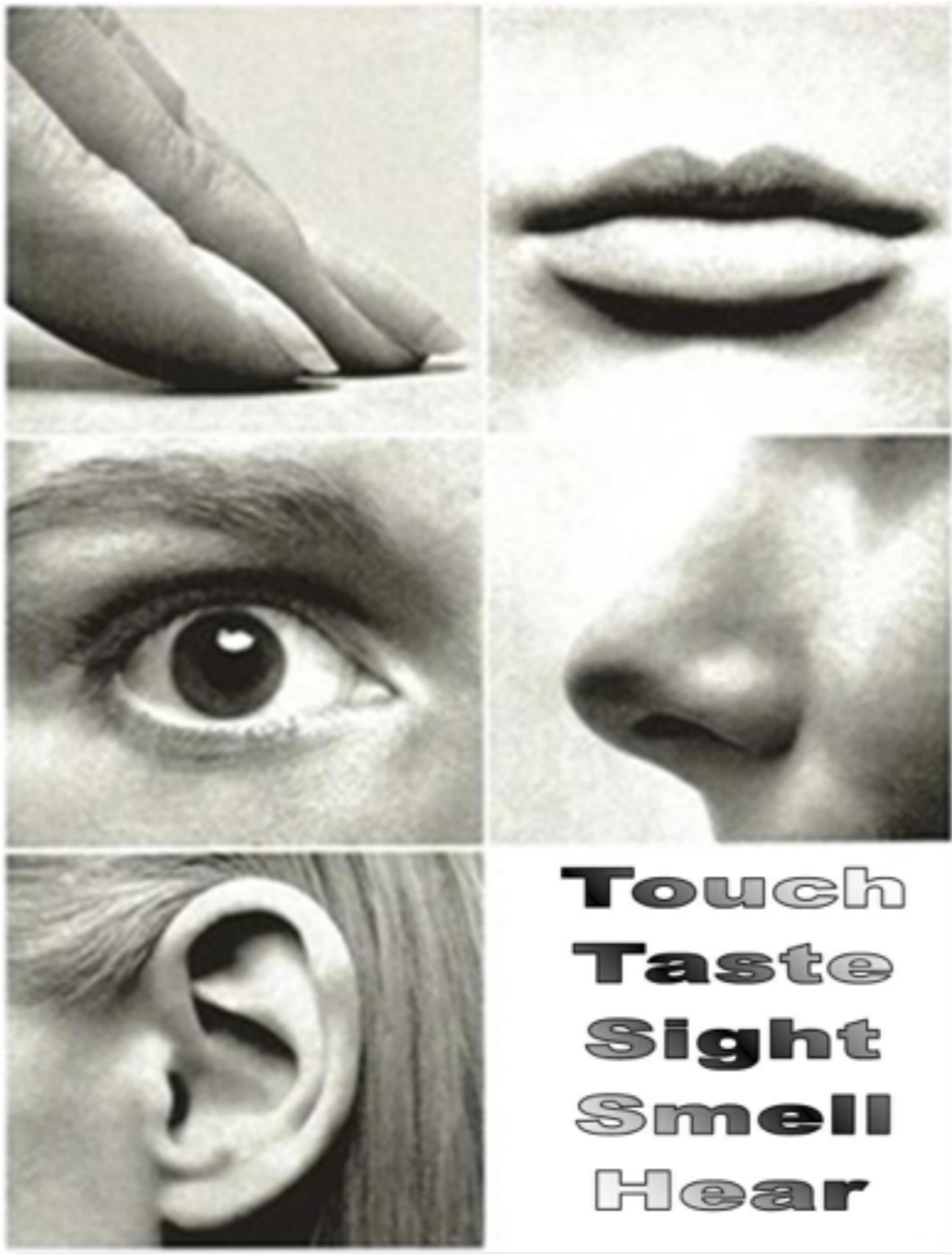
Reduced modeling of neurons  
Population modeling

Dynamical systems  
Mean field equations  
Fokker Planck models  
Stochastics, noise

# Big question

- How does the brain learn?
- If we understood the functioning of the brain, one might create an artificial construct, pose a problem it had not confronted before and it would provide a coherent answer

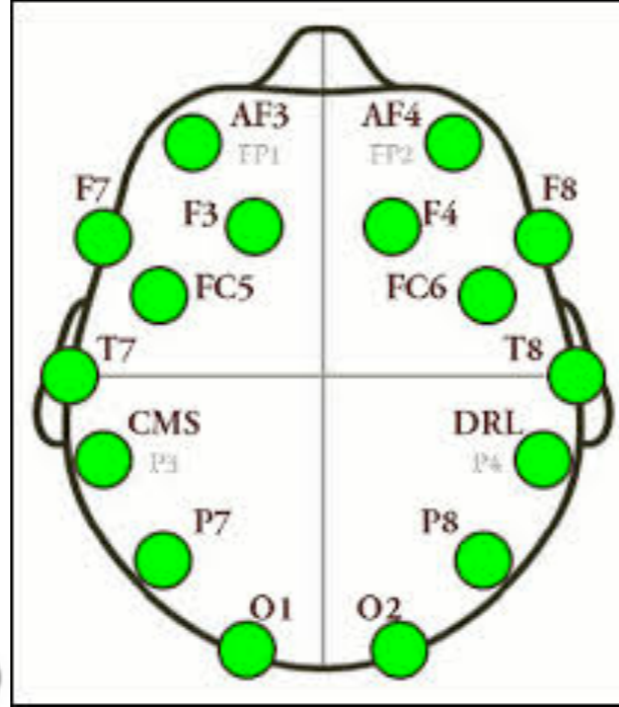
- We interact with the world through our senses
- Our brain emits and receives waves (EEG)

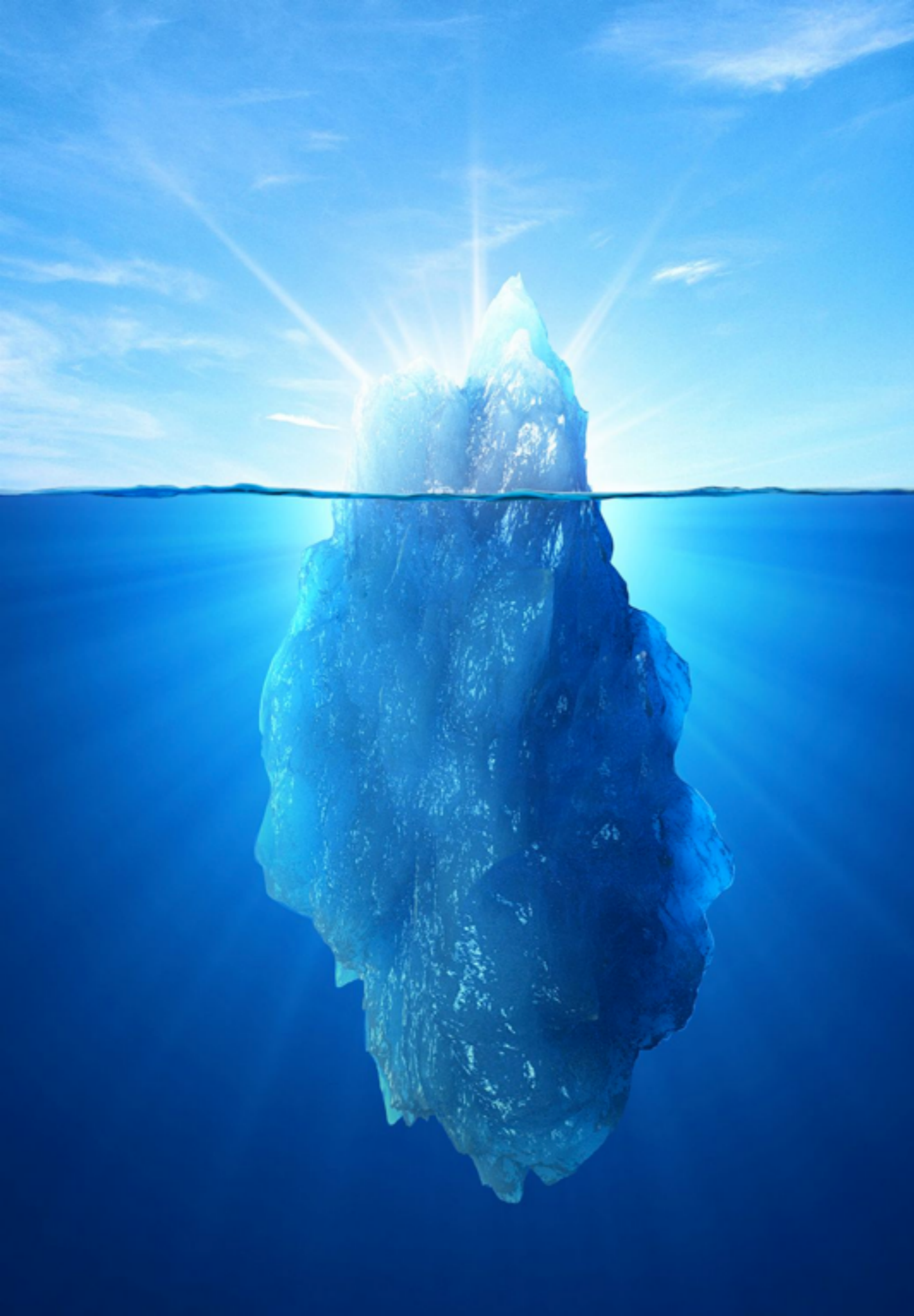


# June-July 2014

- Start work with three Young Scholar Students
- Generated EEG waves with the Emotiv-Epoch
- Used Emotiv to control a simple game with the mind
- Experimented with the [Hierarchical Temporal Memory](#) of Jeff Hawkins (2007)







## Hierarchical Temporal Memory

Convolution networks  
Reservoir networks  
Spiking networks

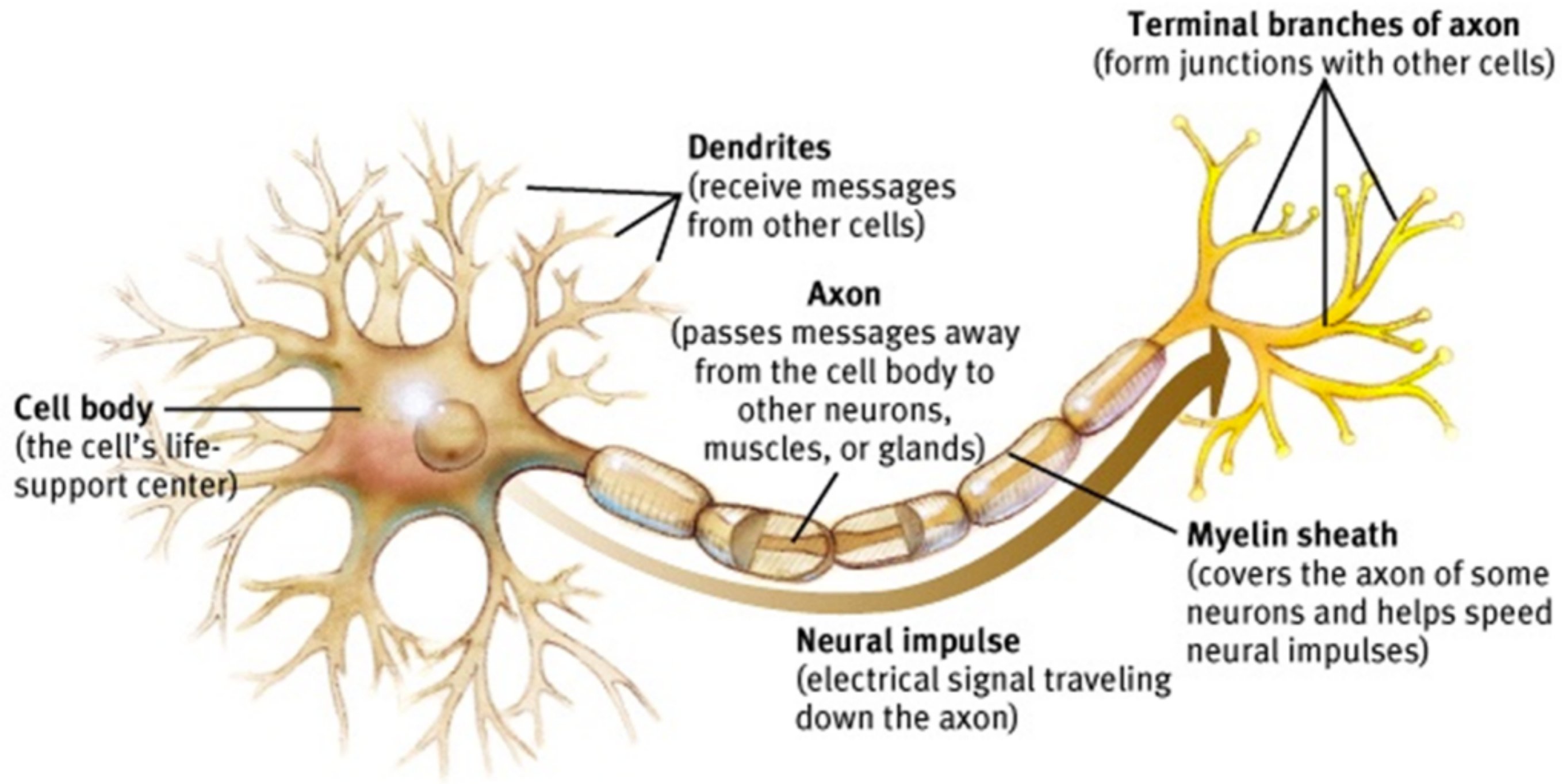
Hodgkin-Huxley model  
Spatial models  
Stochastic models  
Reduced models  
Spiking models

Spiking networks  
Population theory  
Mean equations



# Some Biology

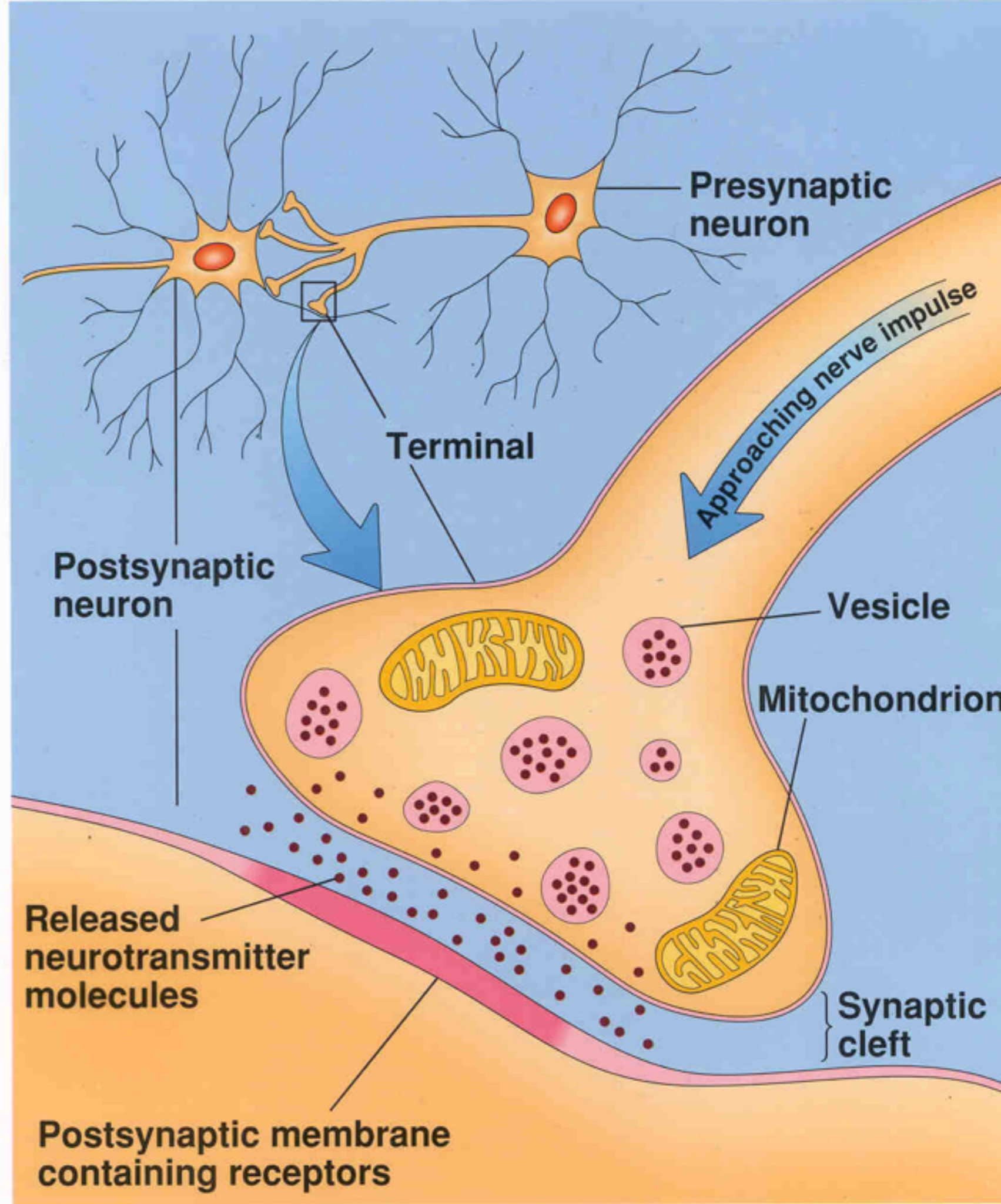
- Neurons
- Synapses
- Astrocytes



# Neuron

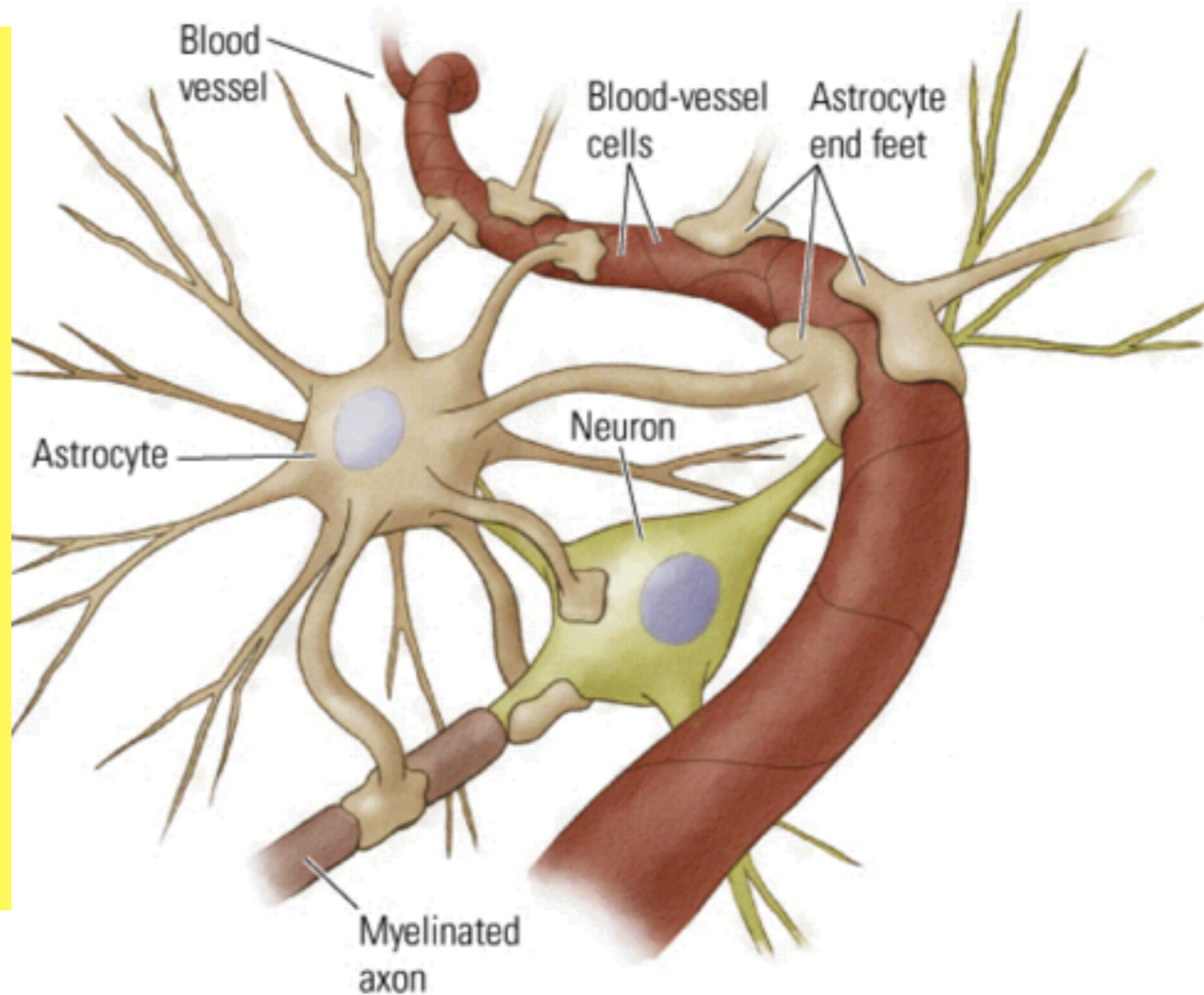
# The Synapse

## Dendrite-Axon Junction



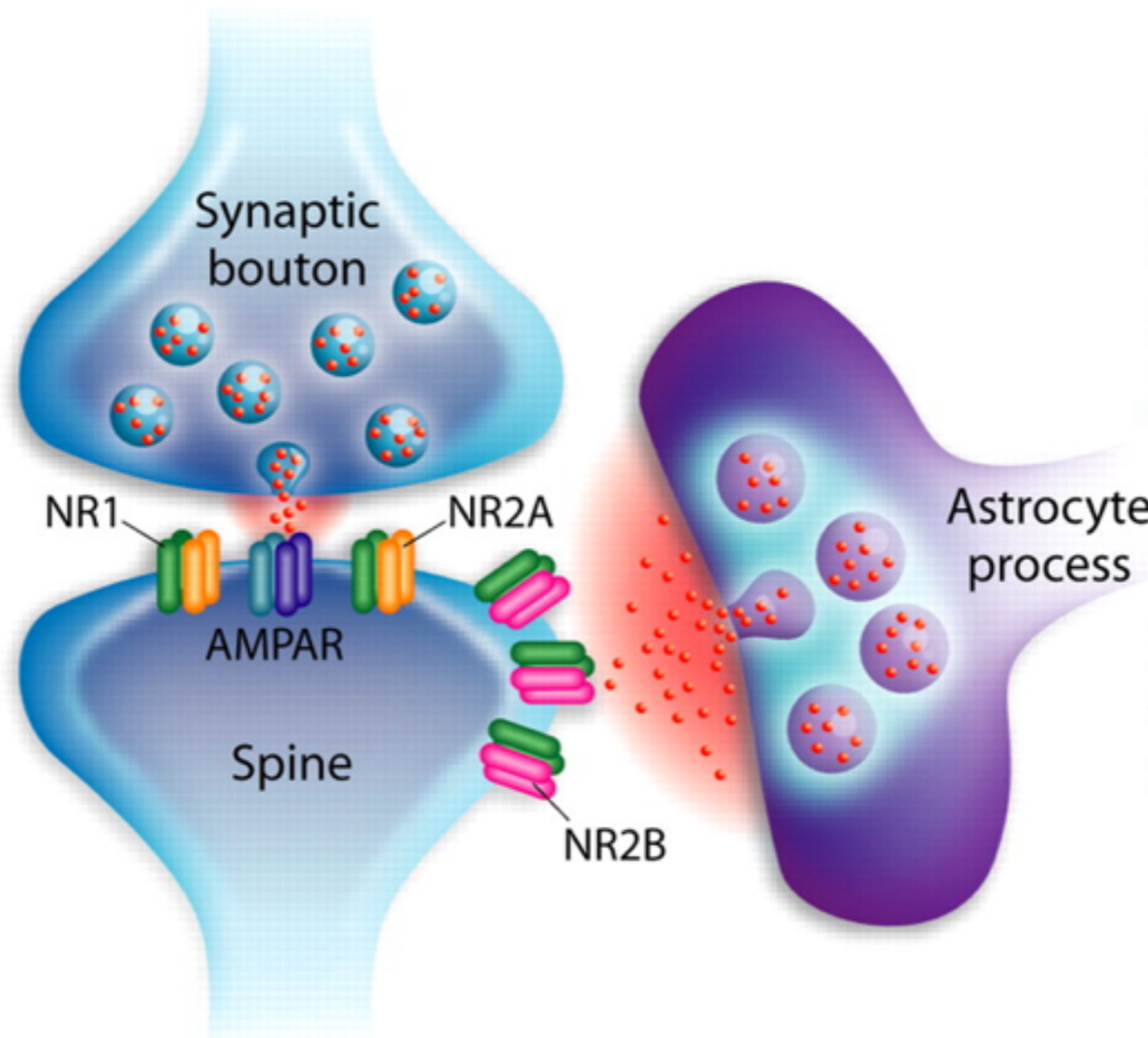
# Astrocytes

- Astrocytes have many functions
  - provide nutrients to neurons
  - regulate calcium flow
  - play a role in various medical disorders (e.g. epilepsy)
  - modulate synaptic strength of neurons



Neuron-to-neuron  
 AMPA/NR1-2A  
 EPSC  
 Rise, ~1ms  
 Decay,  $t_1$ , ~10 ms,  $t_2$ , ~150 ms

LTP  
 CREB activation  
 AMPAR recruitment

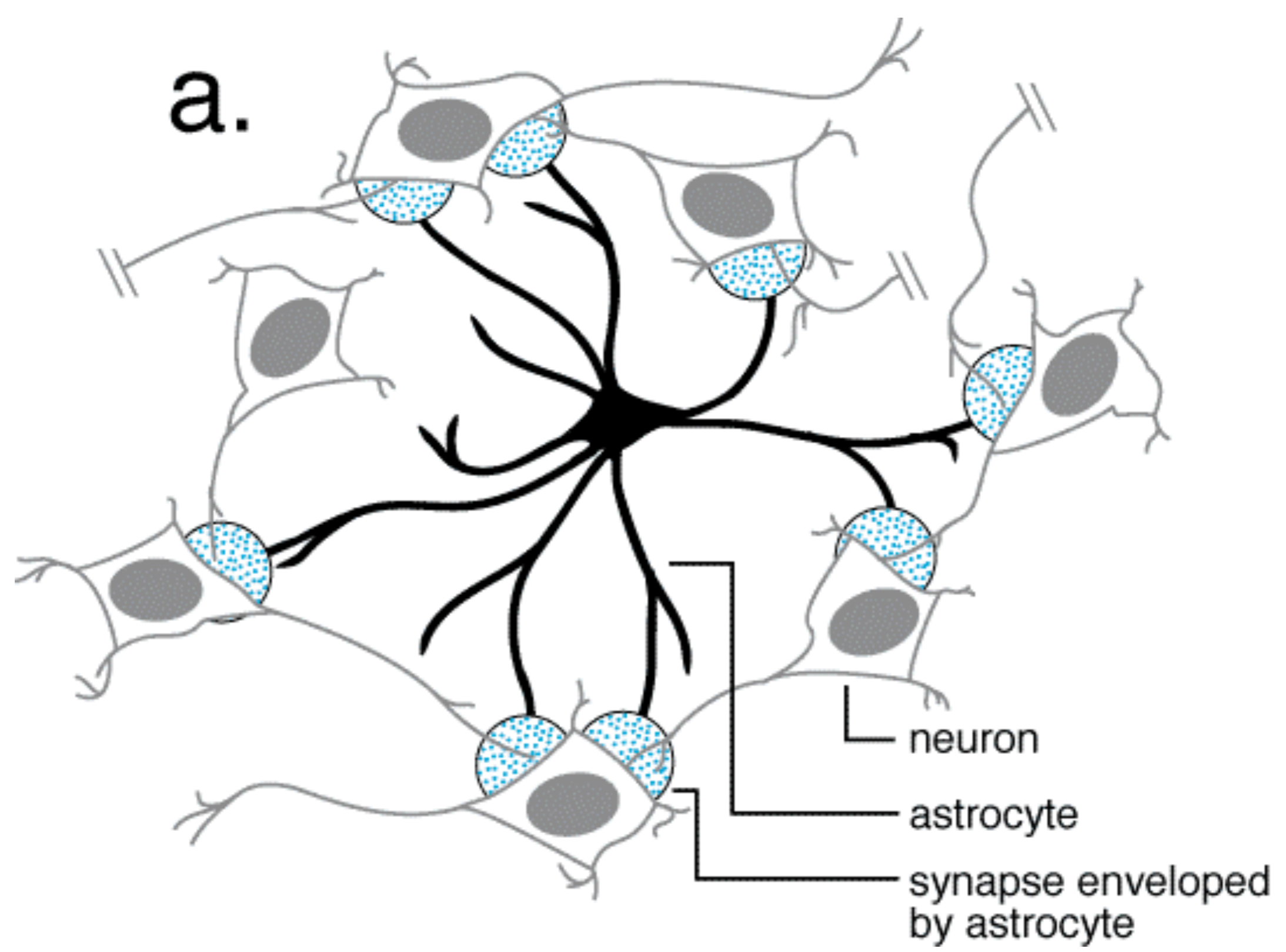


Astrocyte-to-neuron  
 NR1-2B  
 SIC  
 Rise, ~60 ms  
 Decay, ~600 ms

LTD?  
 CREB shut off  
 Neuronal synchrony

# Tripartate Configuration

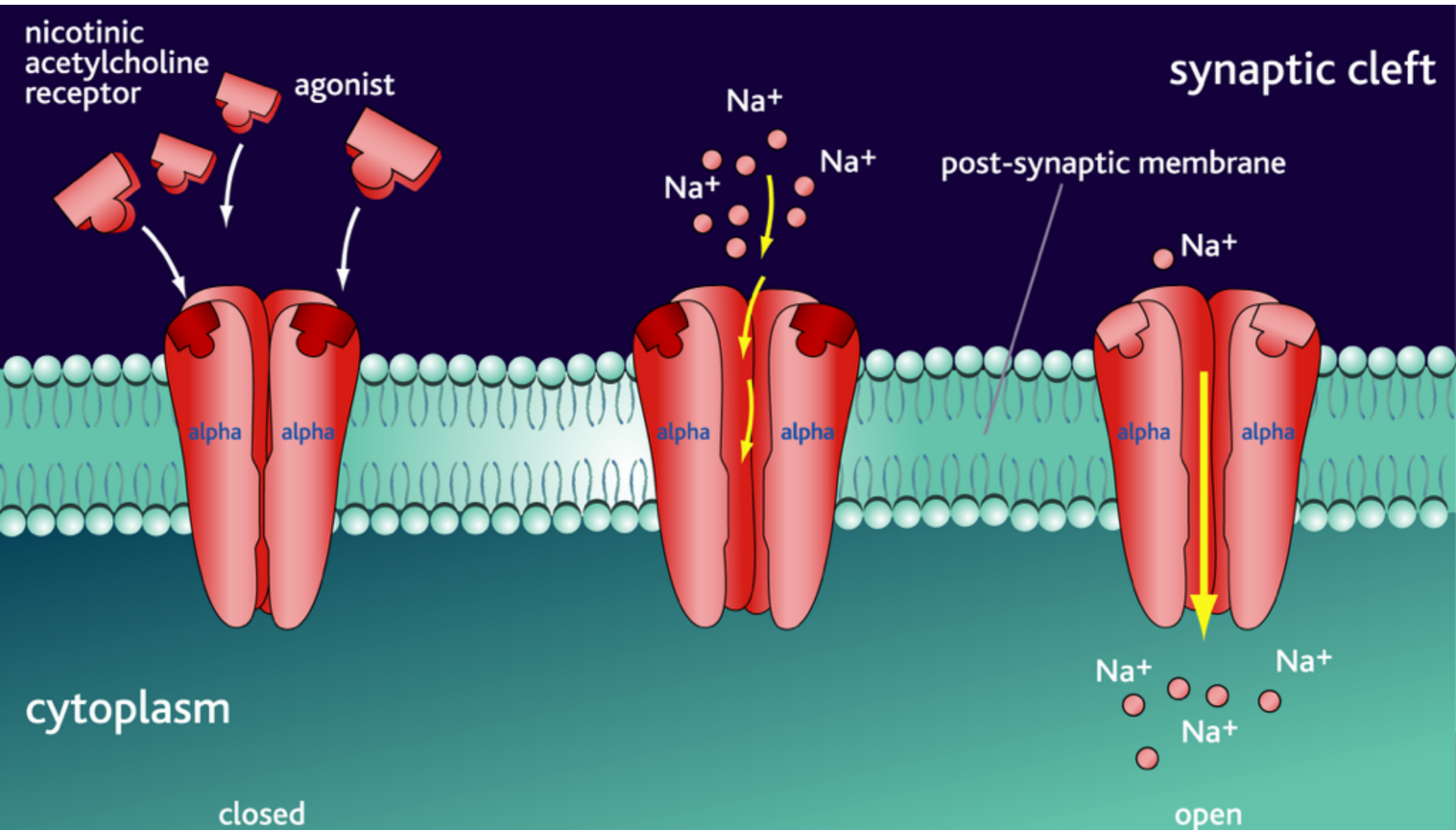
## Astrocyte + Synapse (pre- + post- neuron)



# Prototypical Models

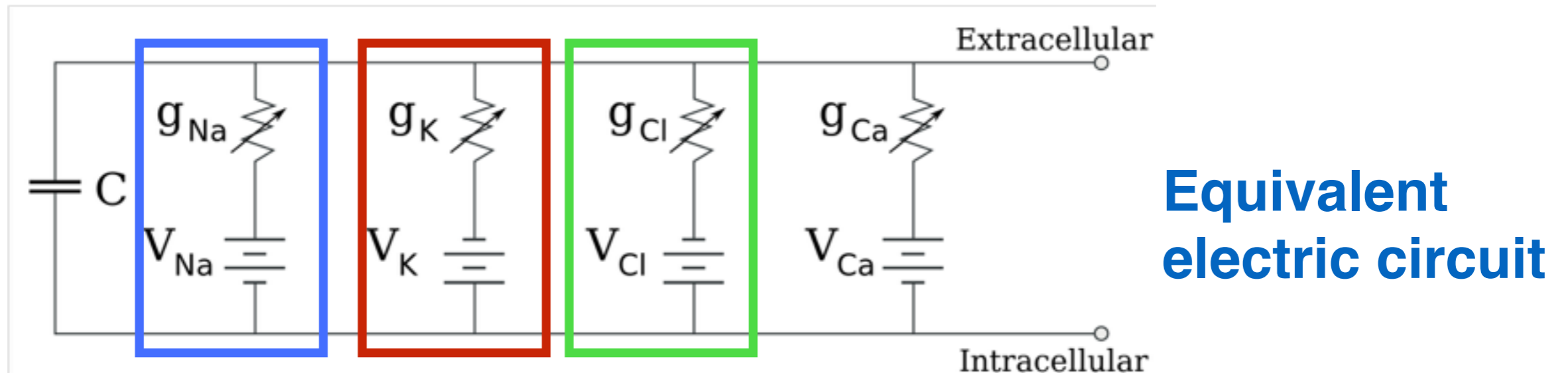
- Hodgkins-Huxley (HH)
  - single compartment
  - modeling of the neuron via electric circuitry
- Multi-compartment models
  - model spatial extent of axon and dendrites

# Ion Channels





# Point Neuron

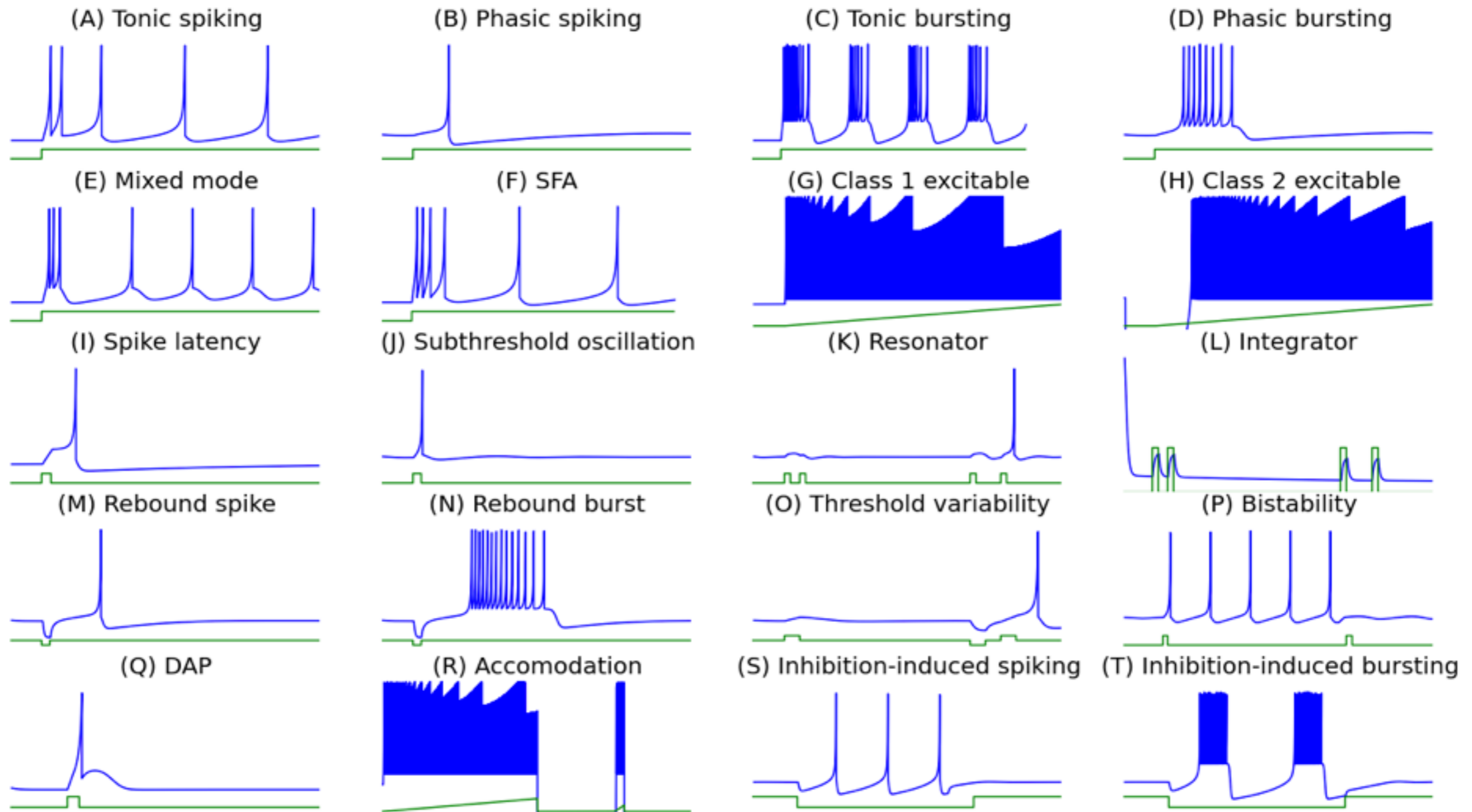


$$I = C_m \frac{dV_m}{dt} + g_K (V_m - V_K) + g_{Na} (V_m - V_{Na}) + g_l (V_m - V_l)$$

**Potassium current**      **Sodium current**      **Leak current**

conductances  $g_K$ ,  $g_{Na}$  and  $g_l$  are functions of voltage and ion channel properties

# Zoo of voltage spiking behavior



(A) Tonic spiking

(B) Phasic spiking

(C) Tonic bursting

(D) Phasic bursting

(E) Mixed mode

(F) SFA

(G) Class 1 excitable

(H) Class 2 excitable

(I) Spike latency

(J) Subthreshold oscillation

(K) Resonator

(L) Integrator

(M) Rebound spike

(N) Rebound burst

(O) Threshold variability

(P) Bistability

(Q) DAP

(R) Accomodation

(S) Inhibition-induced spiking

(T) Inhibition-induced bursting

# Simplified Models

biophysically meaningful  
 tonic spiking  
 phasic spiking  
 tonic bursting  
 phasic bursting  
 mixed mode  
 spike frequency adaptation  
 class 1 excitable  
 class 2 excitable  
 spike latency  
 subthreshold oscillations  
 resonator  
 integrator  
 rebound spike  
 rebound burst  
 threshold burst  
 bistability  
 DAP  
 accommodation  
 inhibition-induced spiking  
 inhibition-induced bursting  
 chaotic spiking

**Flops**

integrate-and-fire	-	+	-	-	-	-	-	+	-	-	-	-	+	-	-	-	-	-	-	-	-	-	5
integrate-and-fire with adapt.	-	+	-	-	-	-	+	+	-	-	-	-	+	-	-	-	-	+	-	-	-	-	10
integrate-and-fire-or-burst	-	+	+		+	-	+	+	-	-	-	-	+	+	+	-	+	+	-	-	-		13
resonate-and-fire	-	+	+	-	-	-	-	+	+	-	+	+	+	+	-	-	+	+	+	-	-	+	10
quadratic integrate-and-fire	-	+	-	-	-	-	-	+	-	+	-	-	+	-	-	+	+	-	-	-	-	-	7
Izhikevich (2003)	-	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	<b>13</b>
FitzHugh-Nagumo	-	+	+	-		-	-	+	-	+	+	+	-	+	-	+	+	-	+	+	-	-	72
Hindmarsh-Rose	-	+	+	+			+	+	+	+	+	+	+	+	+	+	+	+	+	+		+	120
Morris-Lecar	+	+	+	-		-	-	+	+	+	+	+	+	+		+	+	-	+	+	-	-	600
Wilson	-	+	+	+			+	+	+	+	+	+	+	+	+	+		+	+				180
Hodgkin-Huxley	+	+	+	+			+	+	+	+	+	+	+	+	+	+	+	+	+	+		+	<b>1200</b>

# Izhikevich 2003

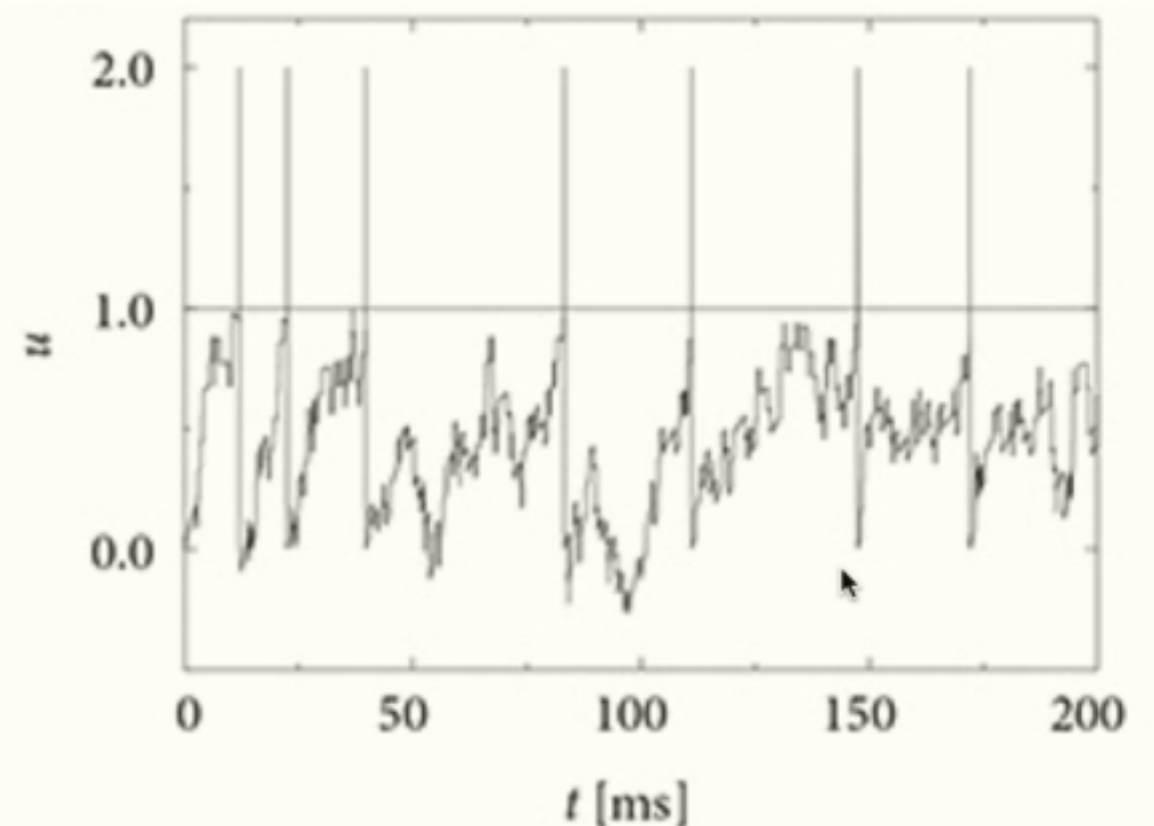
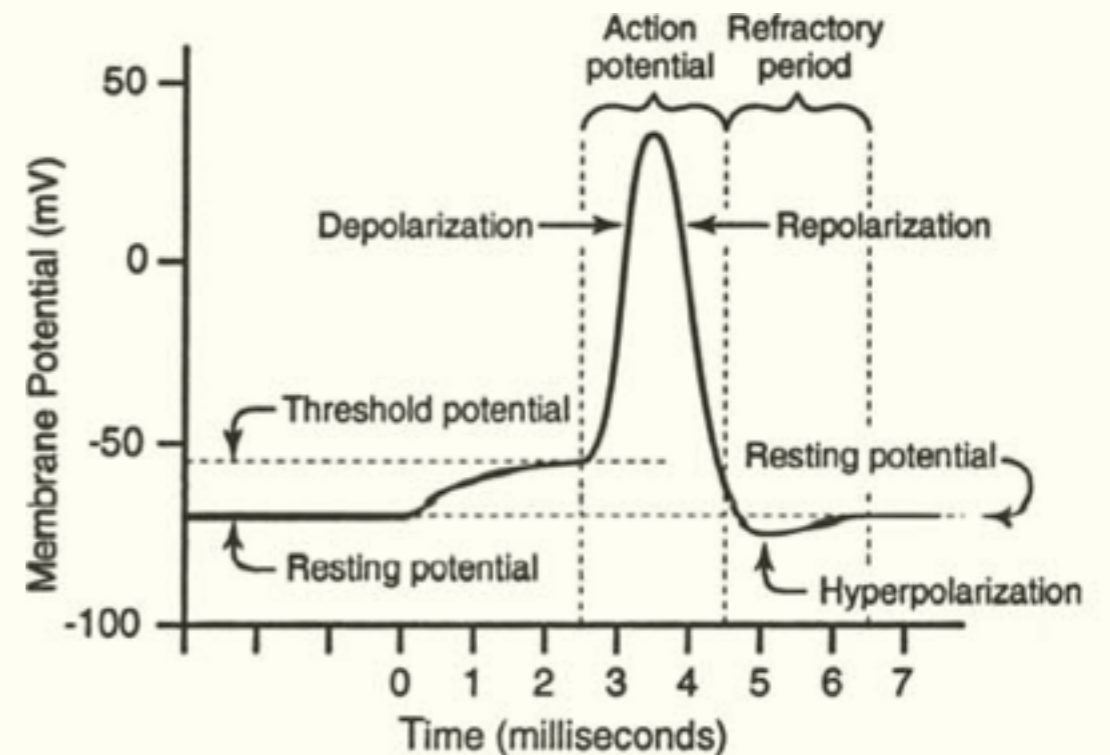
## Integrate & Fire

$$\frac{dV}{dt} = cV^2 + dV - u + I$$

$$\frac{du}{dt} = a(bV - u)$$

if  $V == V_{\text{peak}}$

then  $v \leftarrow V_{\text{reset}}, u = u_0$



# Some important concepts

- **Spiking dynamics**

- how does spiking relate to information content, memory, etc.

- **Propagation speed**

- how long does a single spike take to propagate from neuron to neuron

- **Plasticity**

- change in the synaptic efficacy

# How we Learned

- Online courses
- FSU courses (most notably from R. Bertram in math)
- Several hundred downloaded papers
- Weekly group meetings (open to all)
  - talks and free-flowing conversation
- Question everything
- Coding up interactive demos
- Networking

# Interactive Demo

$$\epsilon \frac{dw}{dt} = (v - v^3/3 - w)$$

$$\frac{dw}{dt} = (v + 1.05 - I)$$

**python FN.py**

# Reality



# Goal



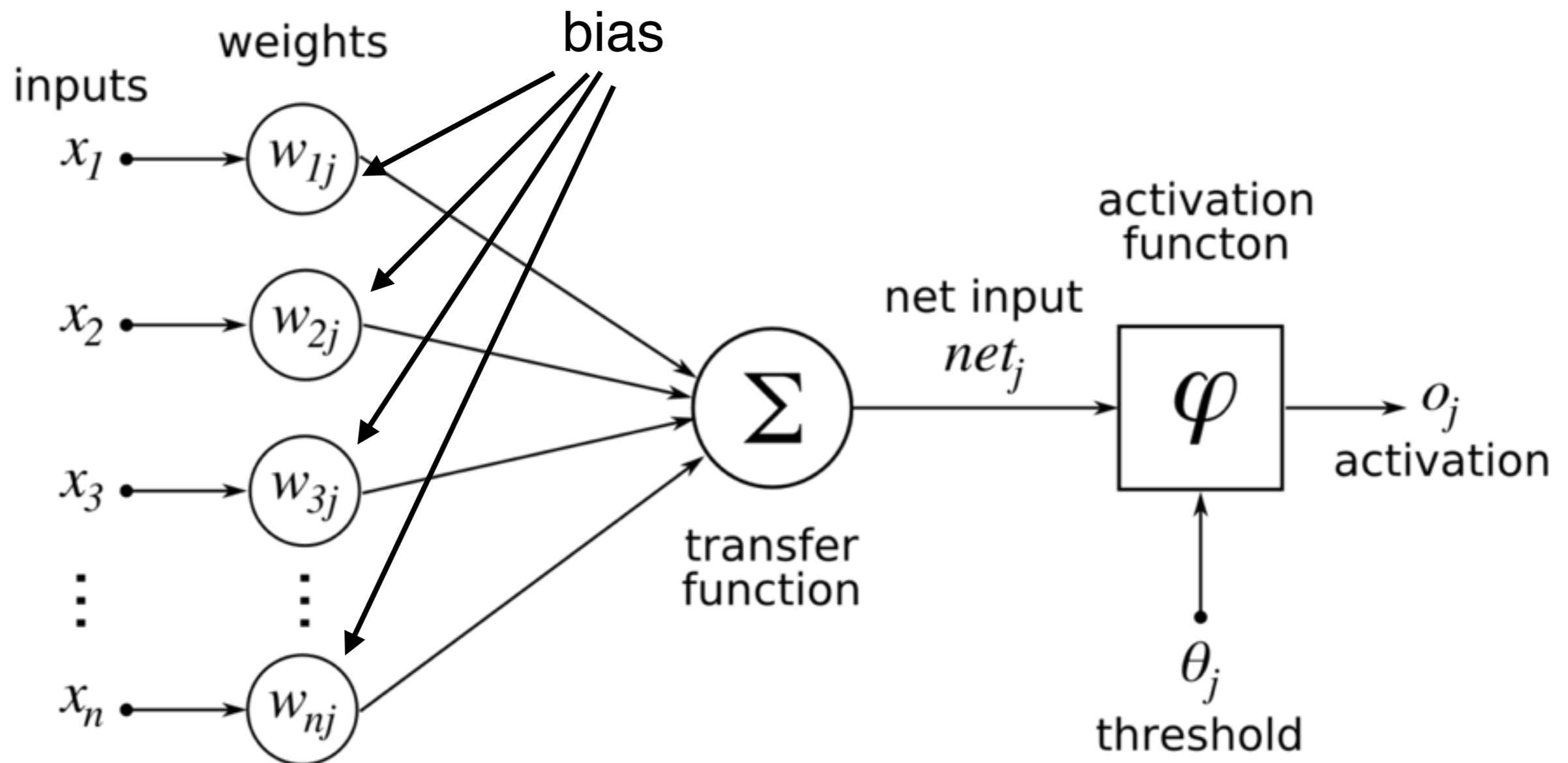


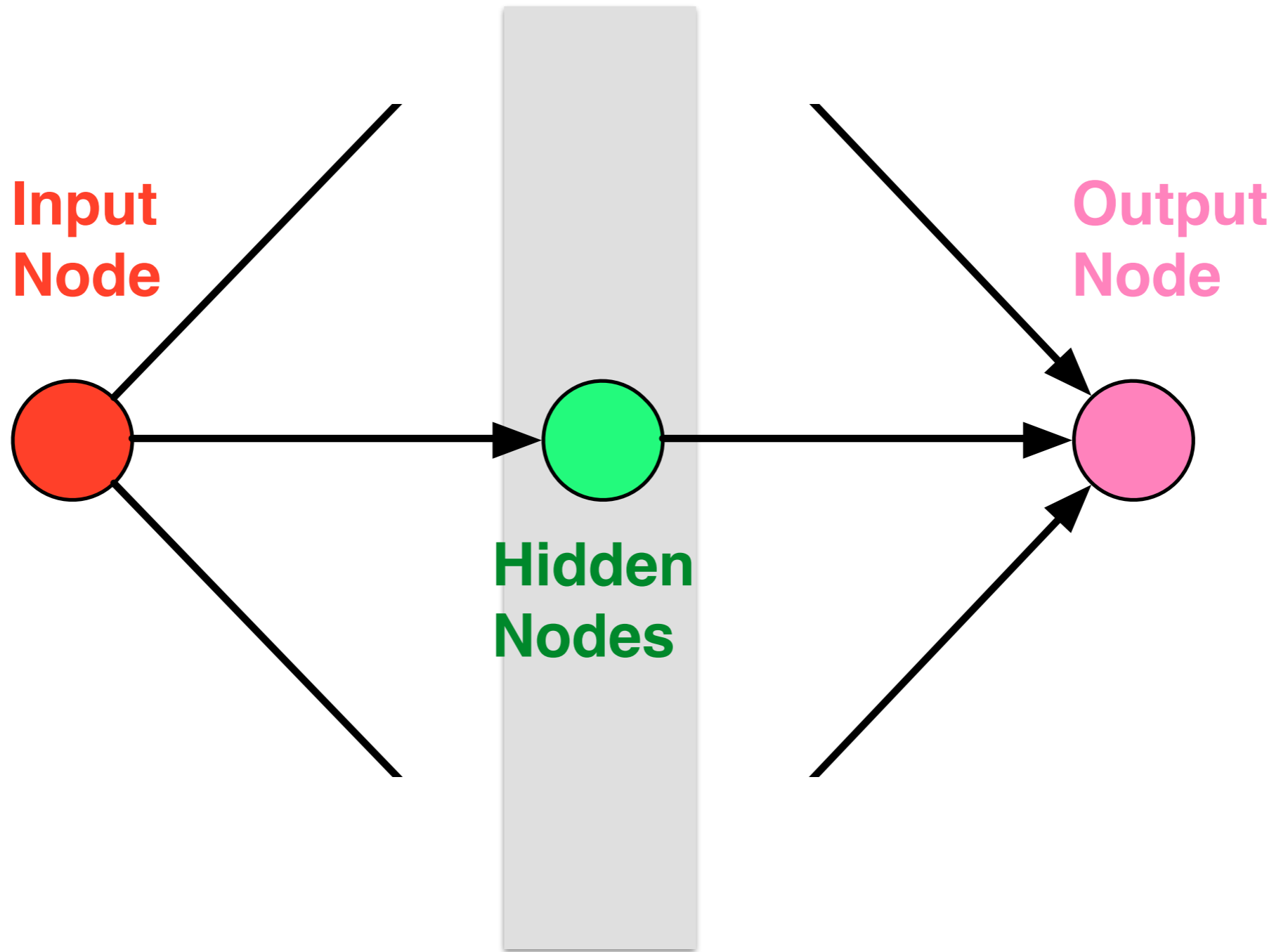
# Most Algorithms (including the Brain)



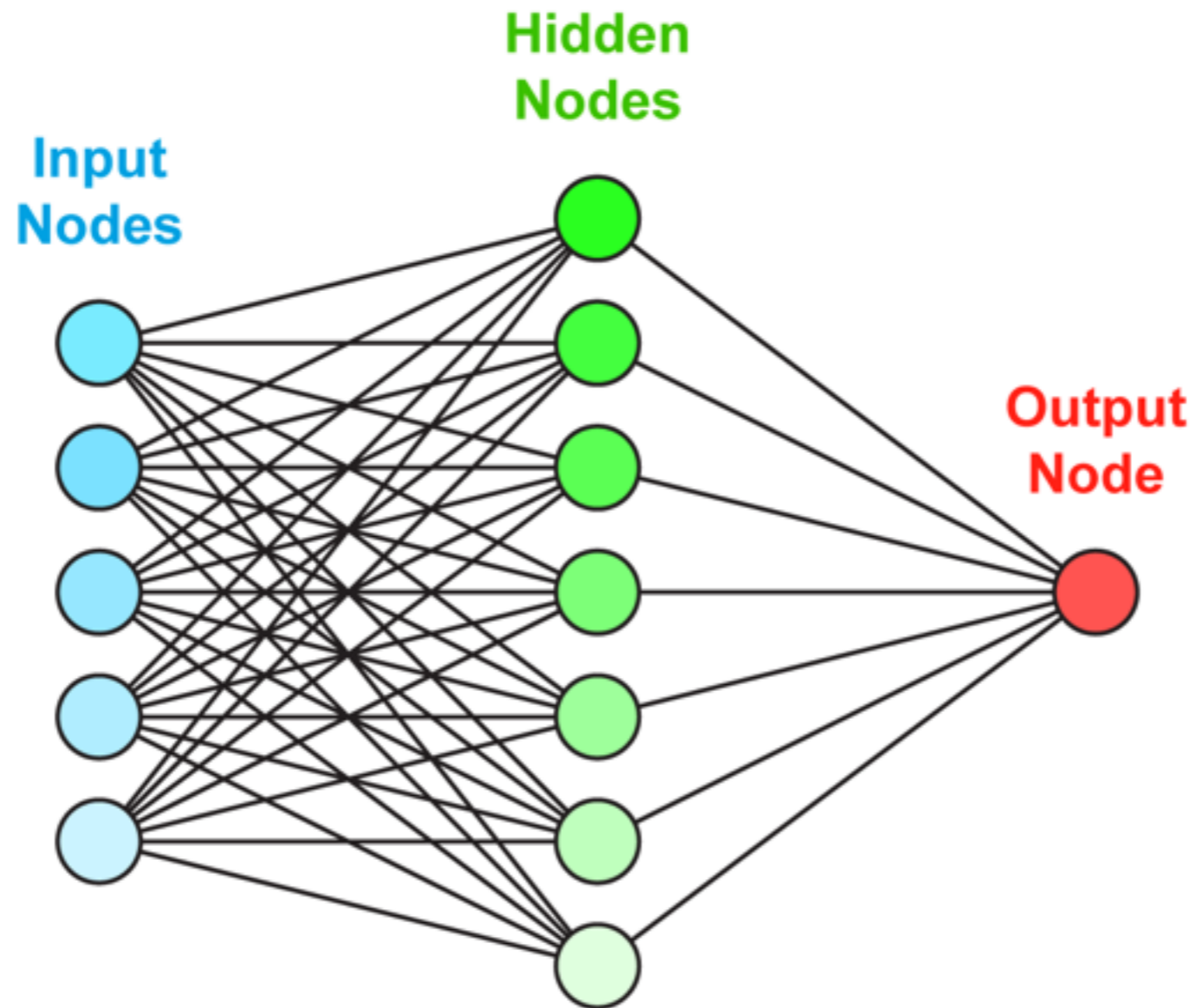
# Artificial Neural Network

## Basic computational unit



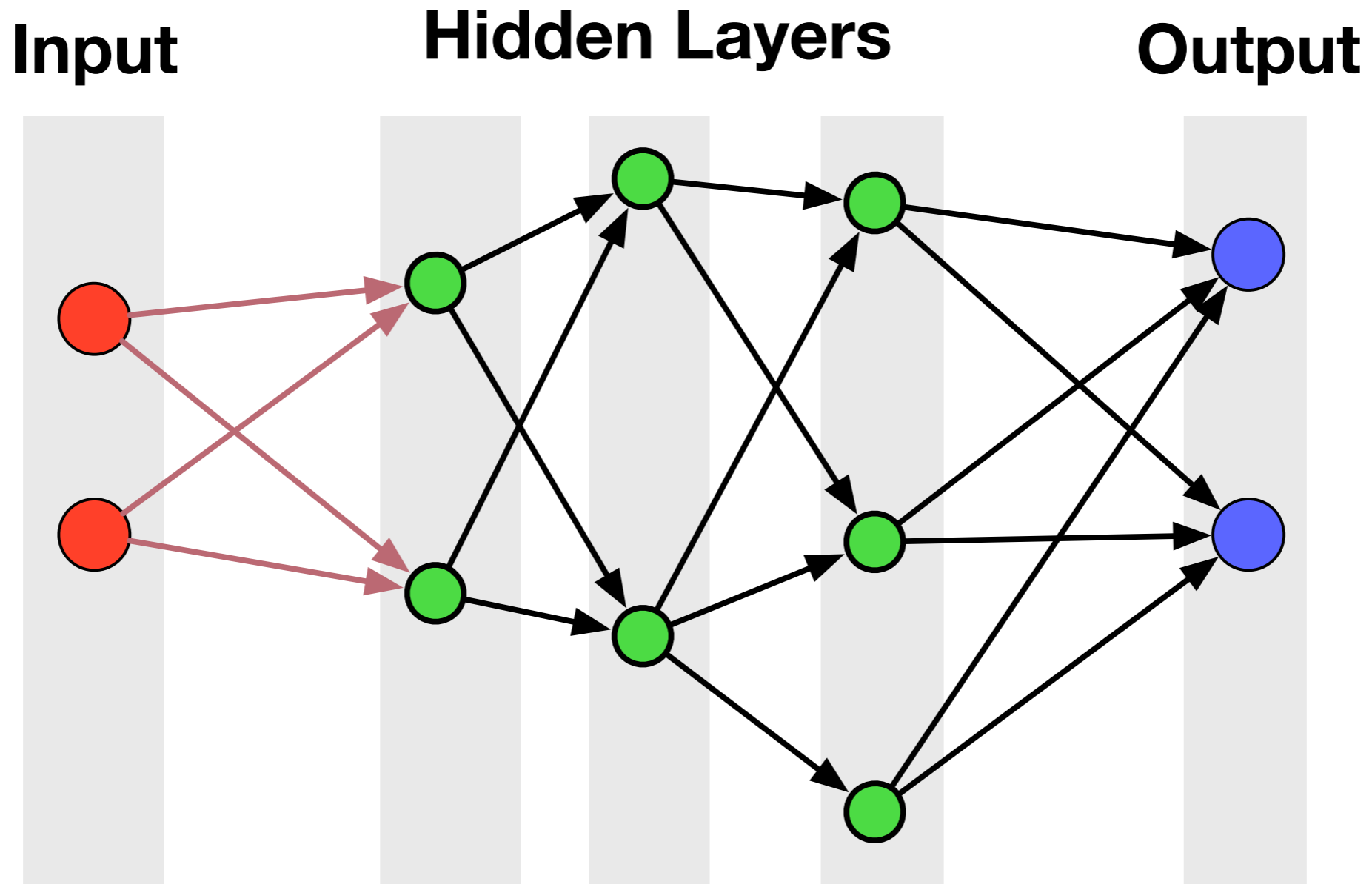


# Single Hidden Layer



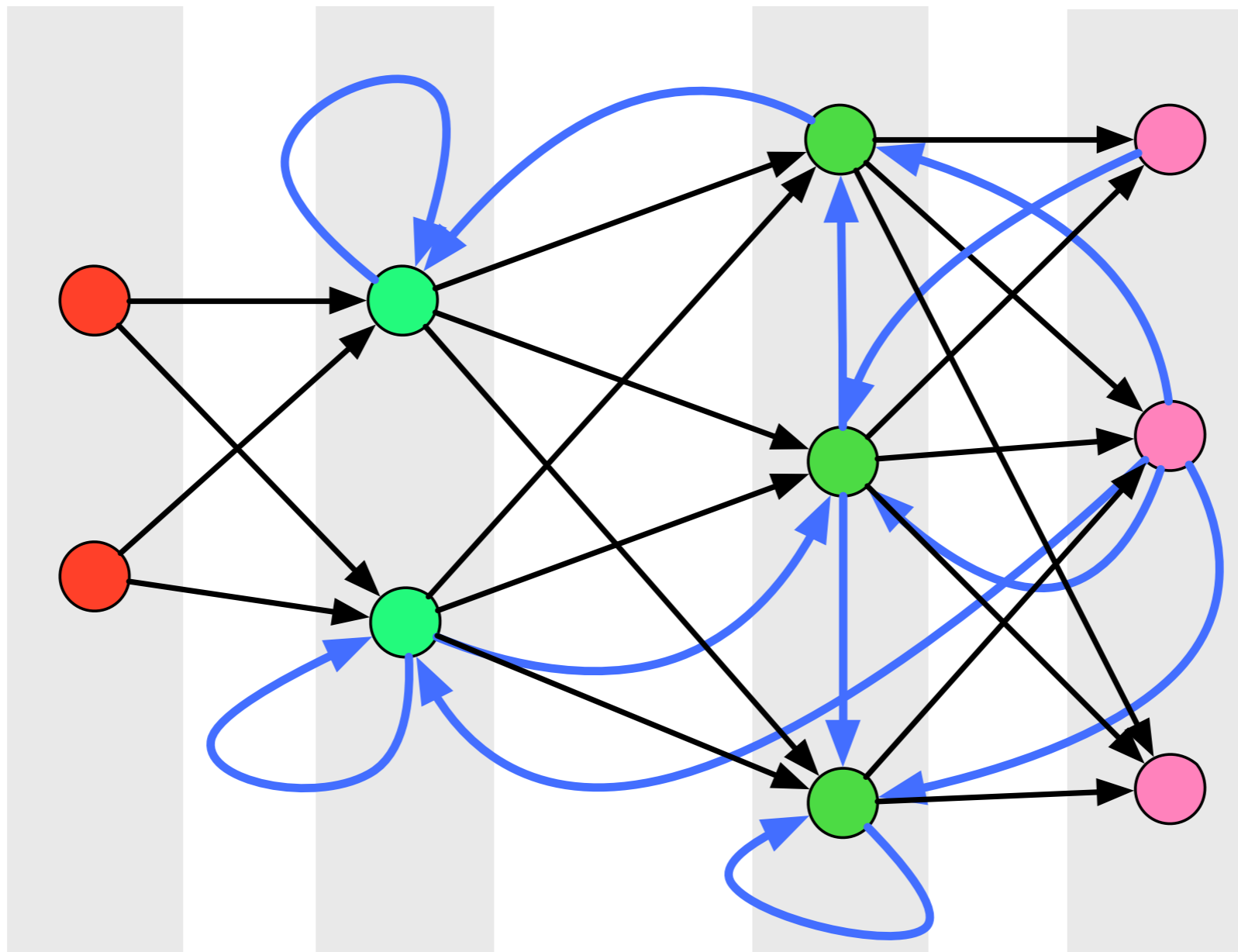
Can approximate any function if there are a sufficient number of nodes in the hidden layer

# Multilayer, Feedforward



Deep Learning

# Multilayer, Recursive



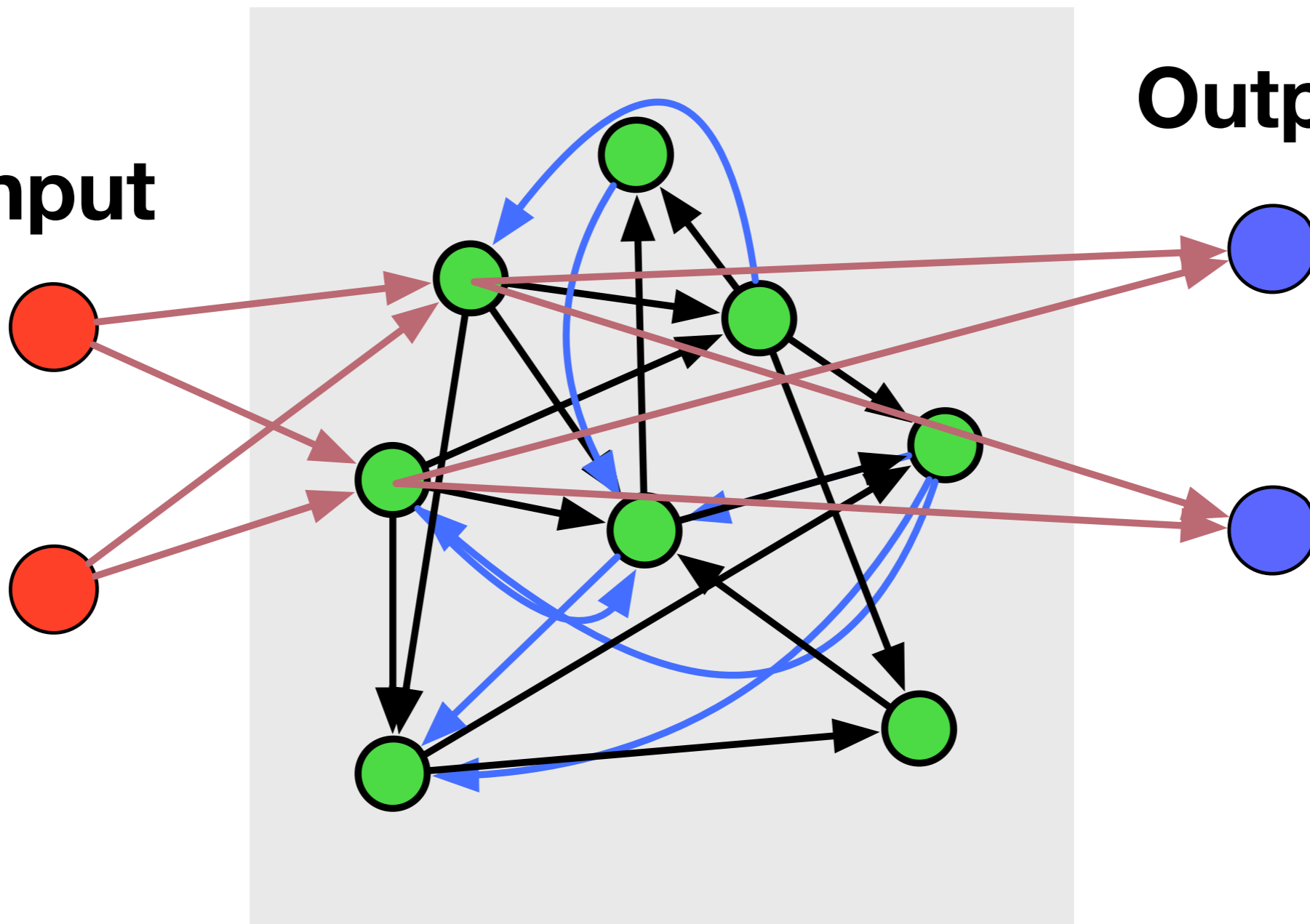
Can model all systems of ODEs  
(i.e., dynamical systems)

# Reservoir Networks

**Reservoir**

**Input**

**Output**



# Reservoir Networks

- Have the potential to store information
- Recursion in networks often translates to the use of past information
- It is possible to control the length of time information is maintained in the network
- Thus, there is the hope of building in memory effects (short, medium, long term) into these reservoirs
- **The average neuron in the brain has 1000 to 10000 recurrent connections**



# Some Remarks

- In all the preceding networks
  - no propagation speed
  - no spiking
  - weights are the solution to a large system of nonlinear equations (one per node), combined with the minimization of some cost function (supervised learning)

# Biologically-Enhanced Artificial Neural Networks

- All the previous networks can be enhanced to add
  - spiking
  - propagations between nodes
  - weight changes via plasticity

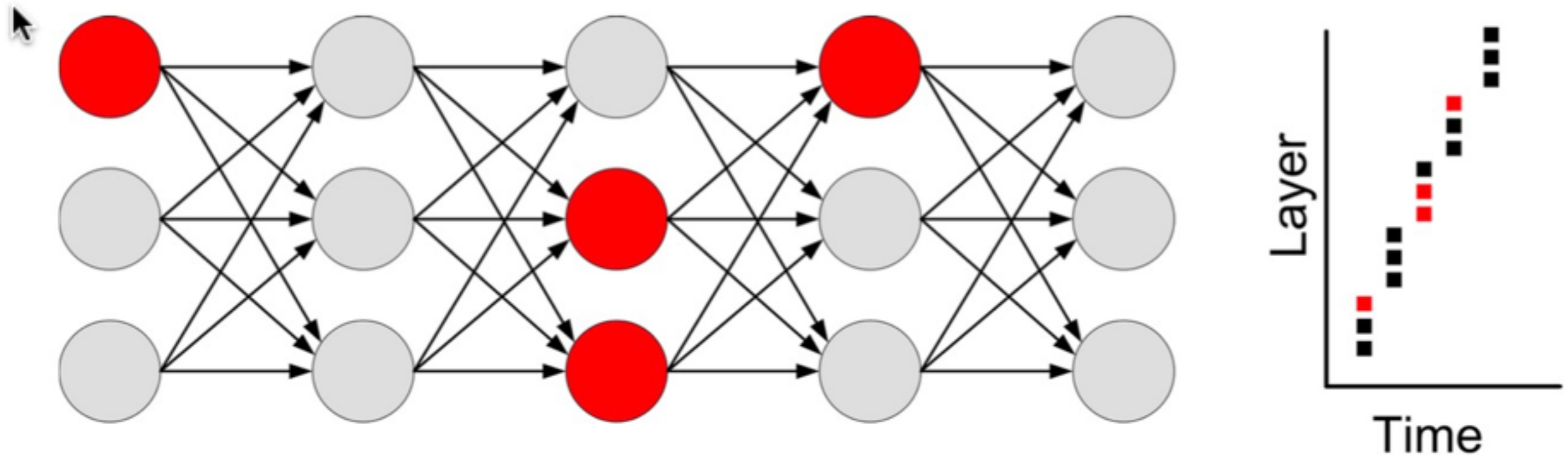
# Important Biological Mechanisms

- Plasticity (multiple forms)
  - mechanisms that affect the strength of synapses
- Synchronization
  - propensity for multiple neurons to fire (i.e., spike) simultaneously
- Balance
  - some neurons excite and some inhibit (**ratio of 5:1 excitatory:inhibitory**), downstream neurons
- Recursion
  - neuron networks are not feedforward

# Synfire Chains

- Under certain conditions, neurons that fire together, will propagate together across a feedforward network
- Synfire chain theory assumes that spike delays are constant
- However, real neuron networks are
  - recursive
  - spike delays are in the range [1ms - 40ms]

# Feedforward Architecture for Synfire Chains

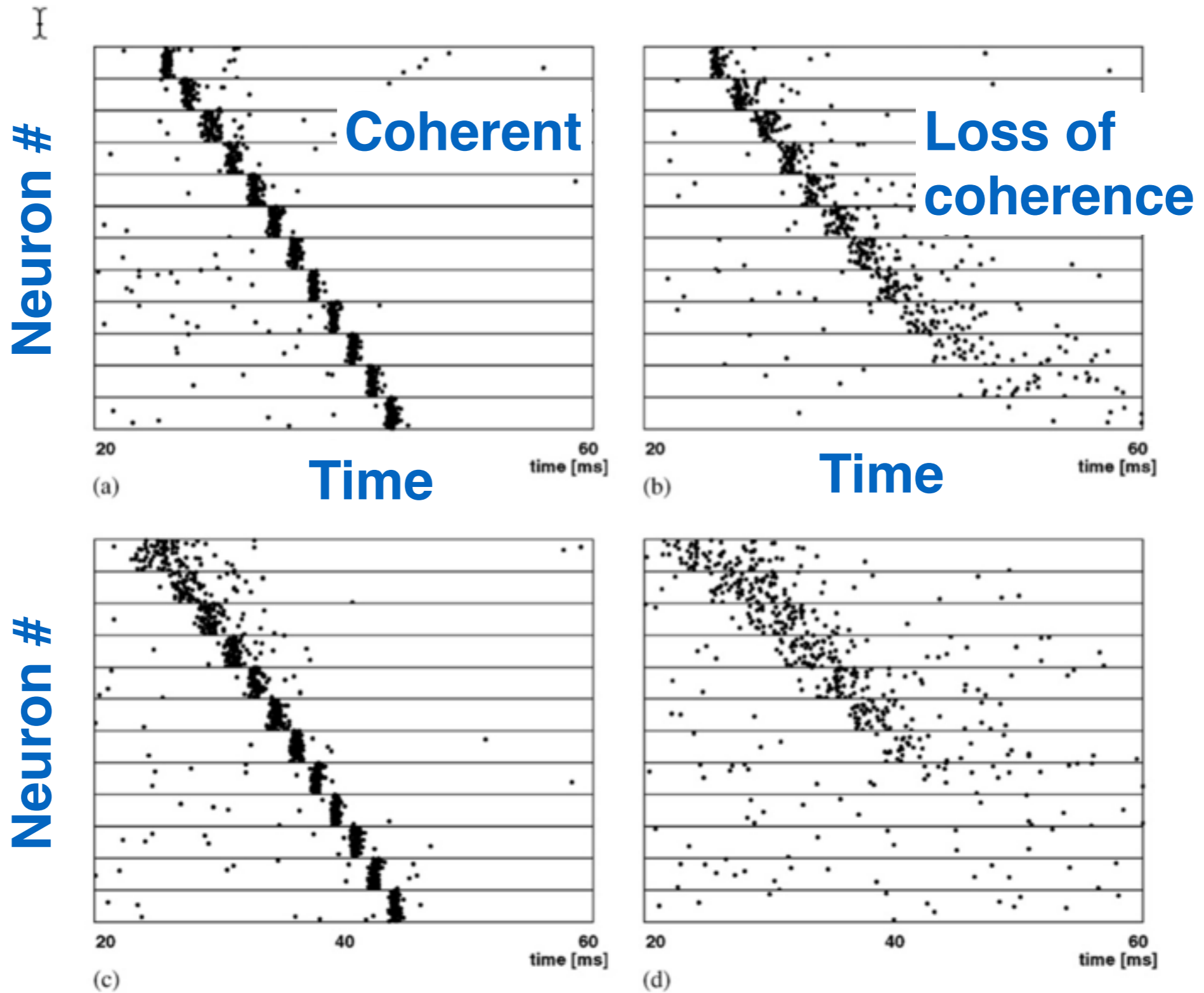


- Neuronal Delays are constant
- Neurons in layer 1 spike within a small time interval
- These spikes propagate across the network, keeping their coherence

# Synfire Chains

M.-O. Gewaltig et al. / Neurocomputing 38-40 (2001) 621-626

623



# Non-constant delays

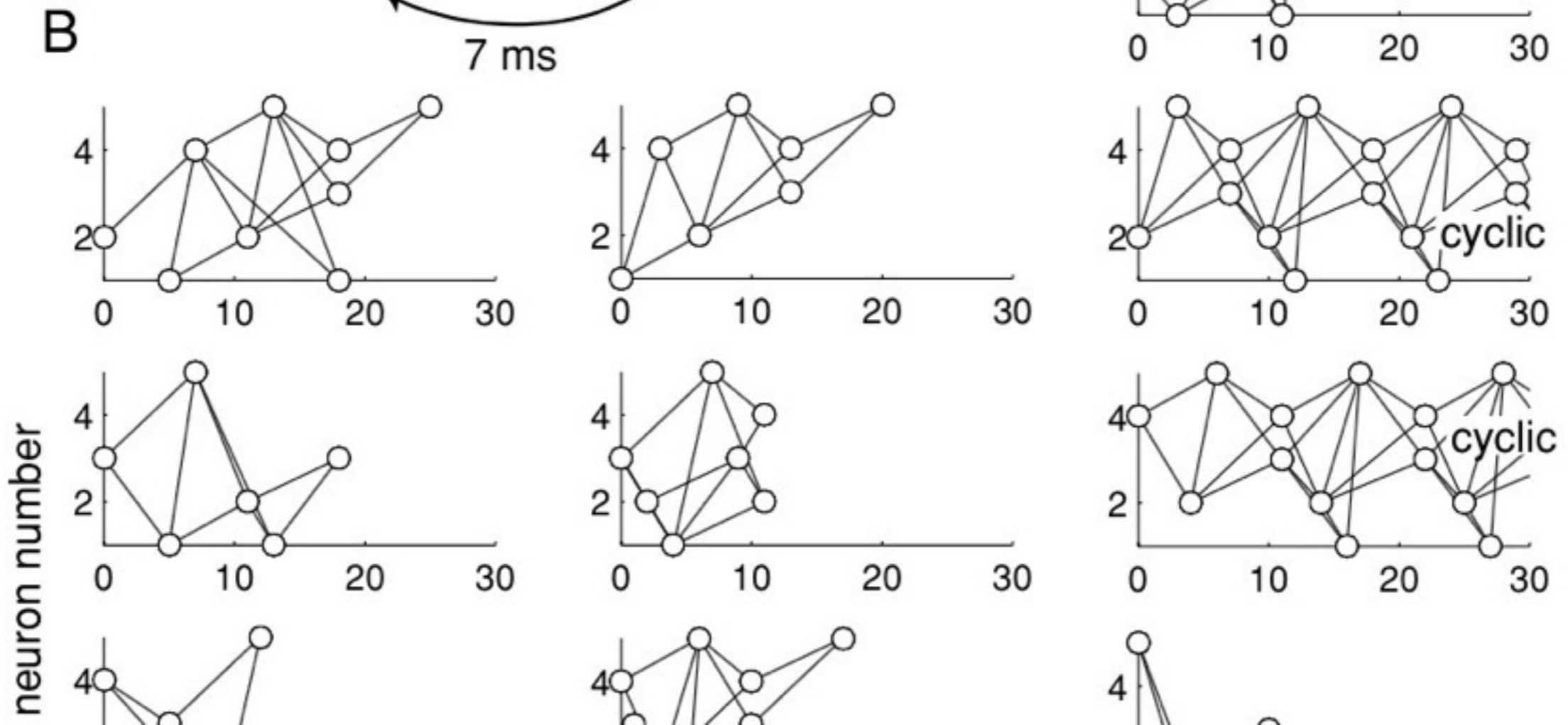
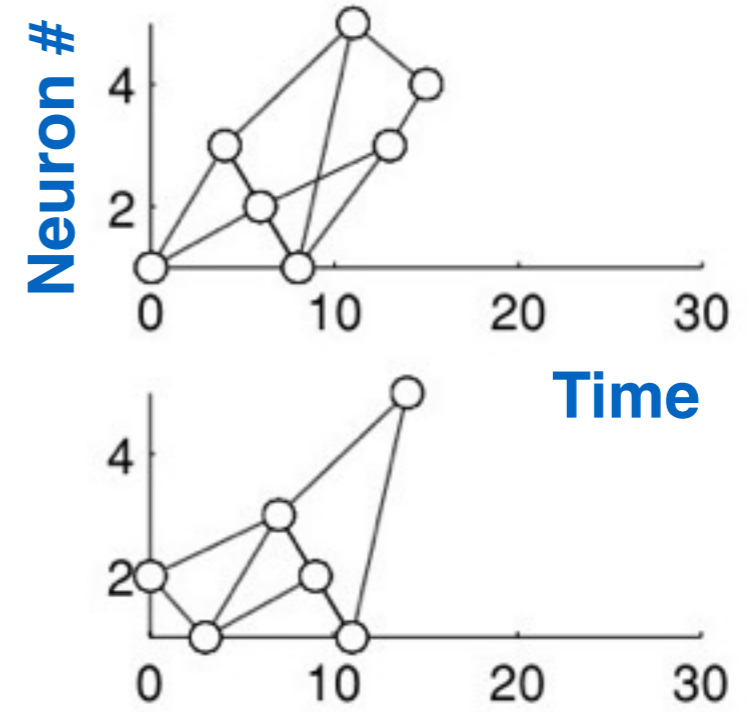
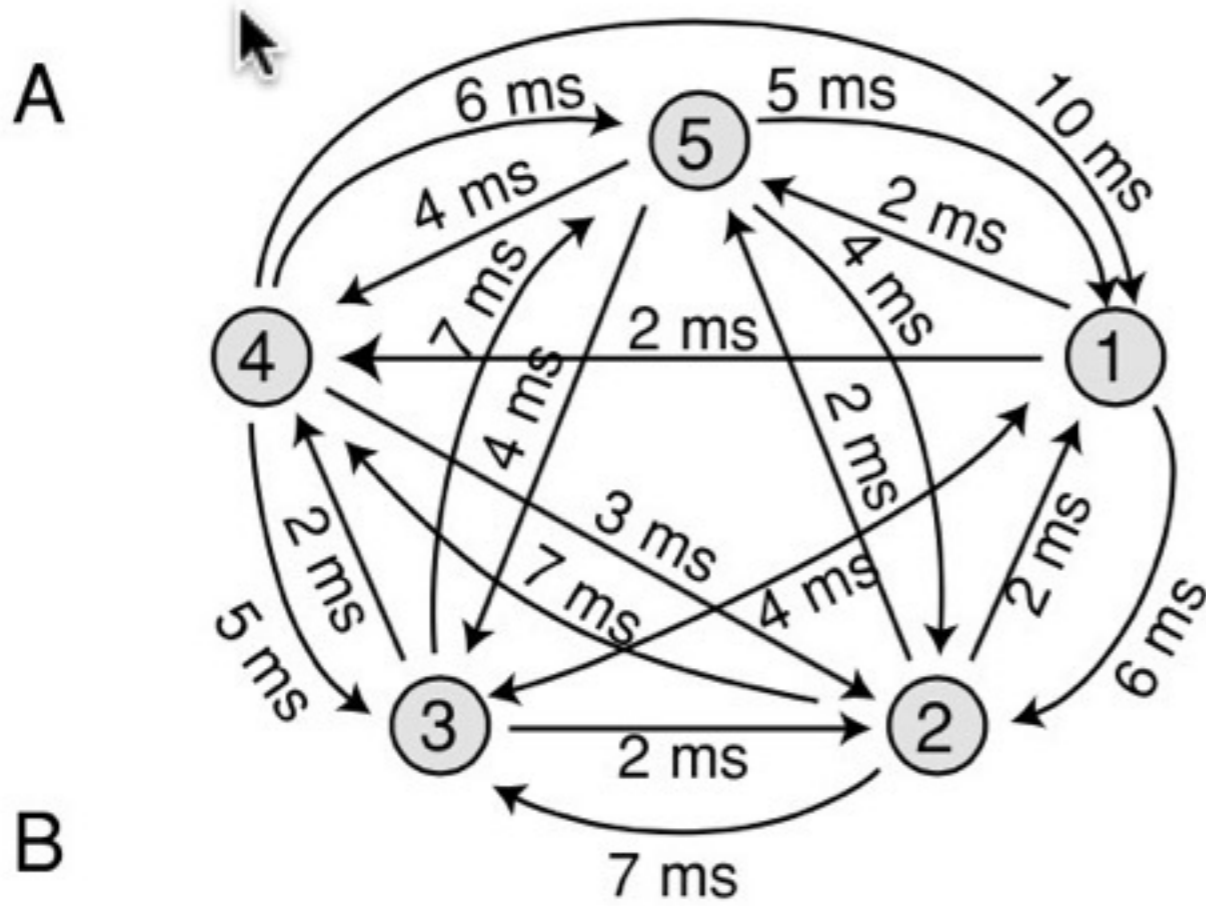
- Synfire chains transforms into a polychronous group
- What is a polychronous group?

A polychronous group is a subset of neurons that fire in a particular space-time sequence that is repeatable given the proper input

# Why are we interested in Polychronous Groups?

Some researchers hypothesize that the sheer number of polychronous groups make them a candidate for the storage of memories





# Questions to answer

- What is the relationship between polychronous groups and synfire chains?
- What are the statistics associated with polychronous groups?
- How are polychronous groups affected by recursion, plasticity, and outside influence (i.e., astrocytes)

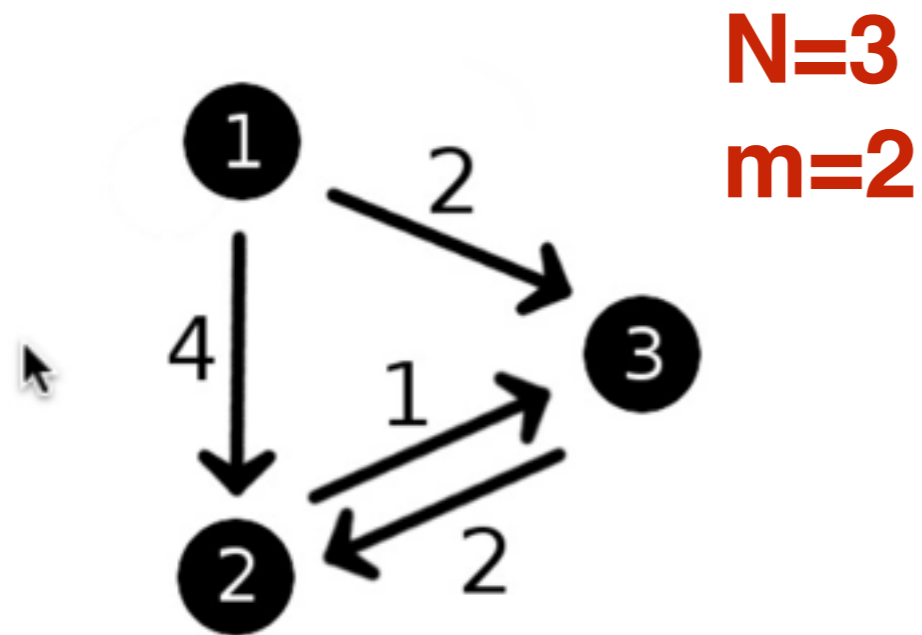
# Some Questions

- Let  $G$  be the number of polychronous groups in a neuron network
- How does  $G(N)$  depend on the number  $N$ , the delay statistics, the number of neuronal connections?
- Dependence of  $G$  on the number of connections  $m$  required to fire?
- What is the affect of recursion on the properties of polychronous groups?
- How robust are polychronous groups to changes the effects of plasticity?
- How many polychronous groups contain a given neuron?

# An Algebra

- Performing simulations to answer these questions is very expensive, even with the simplest models
- We'd like some theoretical results
- Try to create an algebra under very simple assumptions. Hopefully this will provide insight into more realistic situations
- Next: some initial ideas (by [Nathan Crock](#))

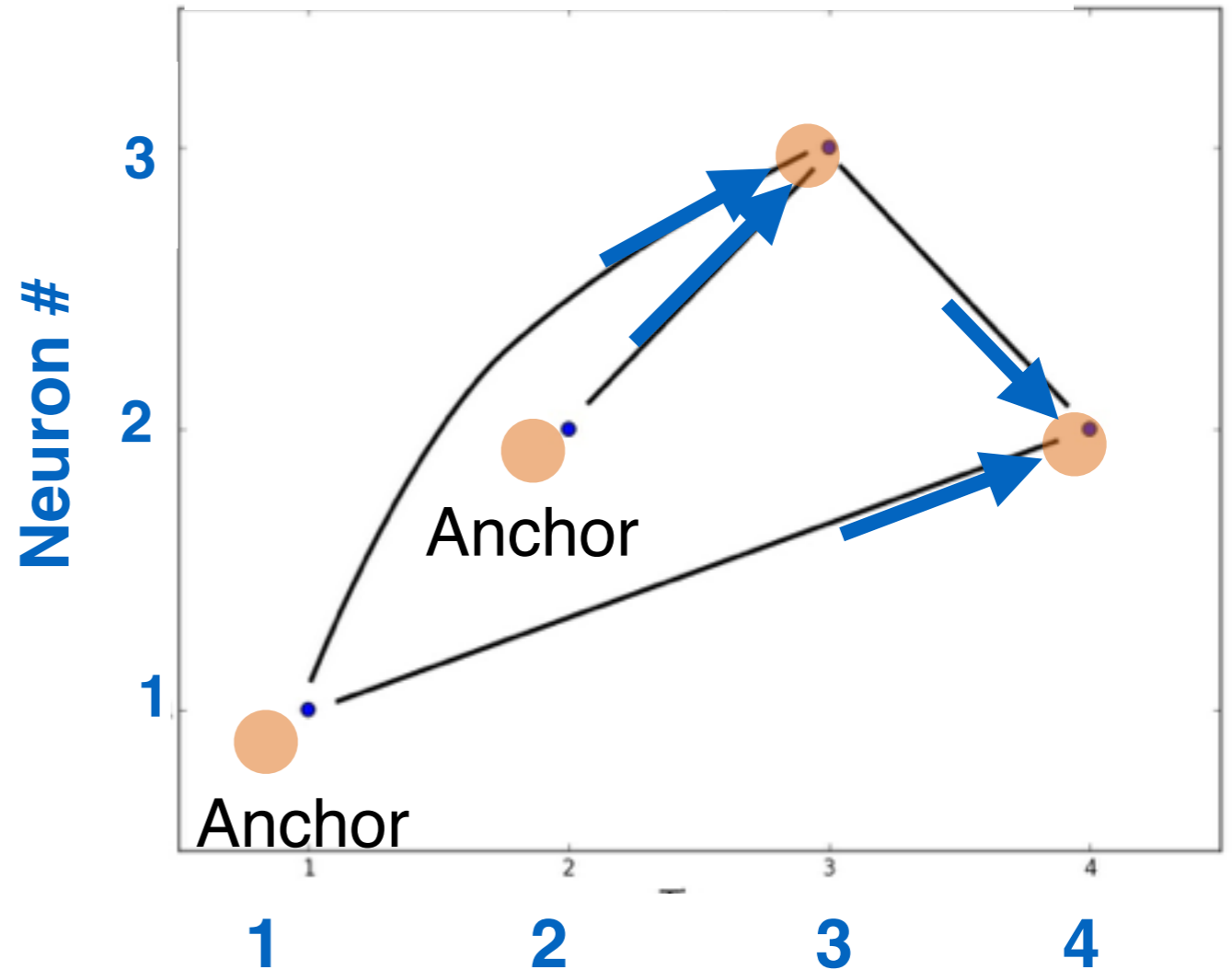
Each neuron requires two input spikes to be activated  
in a biological network, many spikes might be required

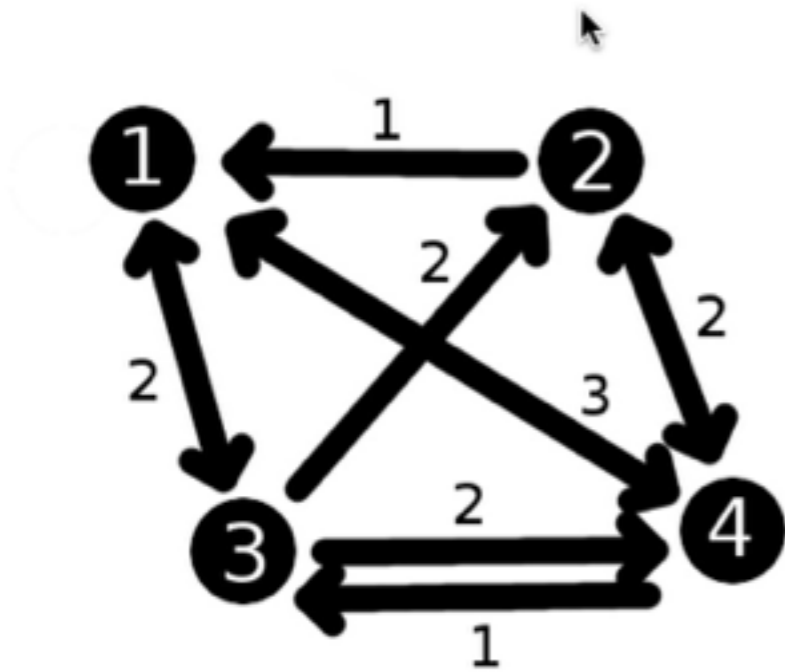


One polychronous group  
labelled by its anchors  
and time of emission

$$P1 = \{(1,1), (2,1)\}$$

### Neuronal Group Activation





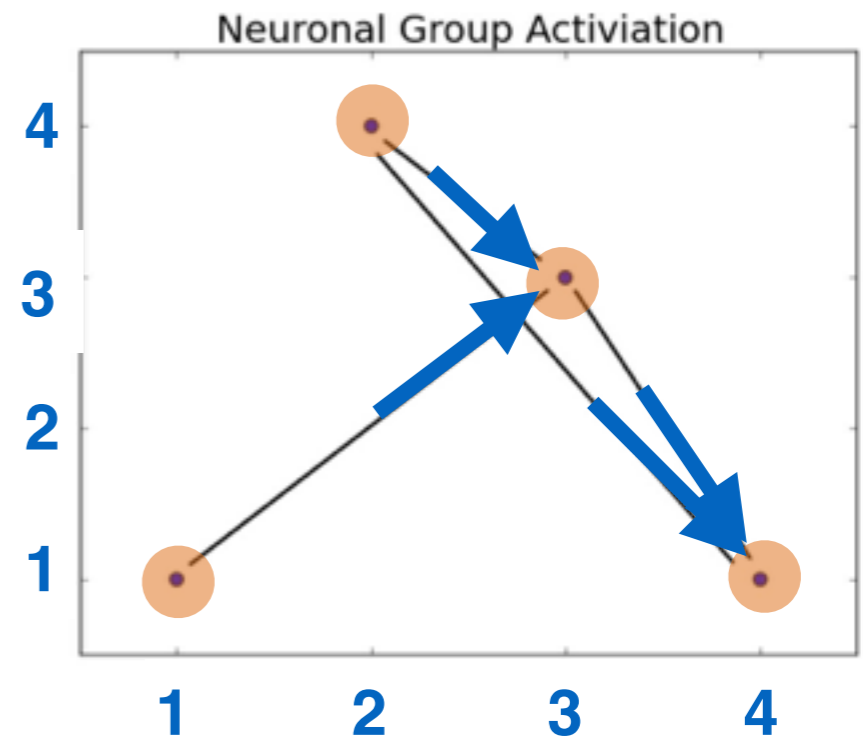
**N=4**  
**m=2**

Two polychronous groups  
labelled by its anchors  
and its firing (spiking)  
time:

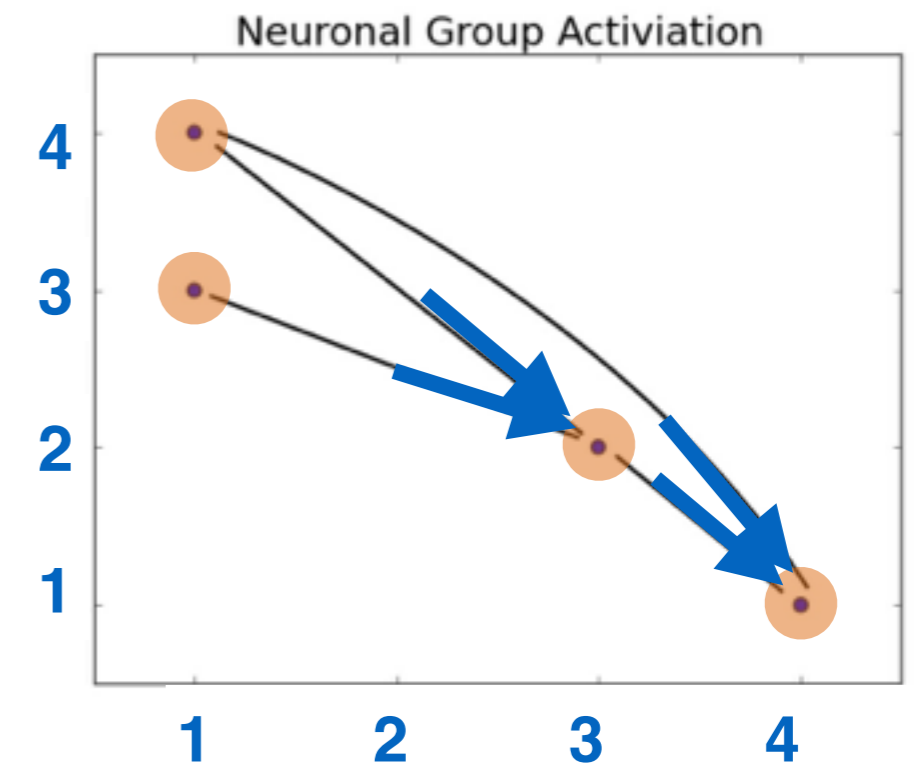
$$P1 = \{(1,1), (4,2)\}$$

$$P2 = \{(3,1), (4,1)\}$$

Neuron #



Neuron #



# Initial properties

- Consider  $P1 = \{(1, 1), (4, 2)\}$
- Assume the group is “activated”  $t_3$  time units earlier
- Rewrite it as  $P1 = \{(1, t_1), (4, t_2)\}$
- The following equality holds
$$P1 = \{(1, t_1 - t_3), (4, t_2 - t_3)\}$$
- A polychronous group is said to exist independently of its time of “activation”

# Network of

$$P1 = \{(1,1), (4,2)\}$$

Time shift invariance of polygroup

(Recall: a polygroup is represented by the neurons that activate it.)

4	0	1	0	0
3	0	0	0	0
2	0	0	0	0
1	1	0	0	0
	1	2	3	4

Neuron #

Time

Matrix-like  
object



# The $i^{\text{th}}$ Neuron

$$\mathbf{n}^{(i)} = \begin{pmatrix} d_1 \\ d_2 \\ \vdots \\ d_N \end{pmatrix}^{(i)}$$

**Neuron index**

**Delay from neuron (i) to neuron 2**

The “elements” of  $\mathbf{n}(i)$  are the delays of its deferent (downstream) connections.

# Summary

- Vector-like objects (neurons)
- Matrix-like objects (networks)
- Basis neurons (not shown)
- Time-shift operators
- Other operators (+, -)

# Some Research Objectives

- Define notion of vector space for neurons and networks (metrics, norms)
- Define equivalence between networks
- Decompose a neuron or network into irreducible representations
- Construct more complex networks from simpler networks
- Pinpoint the notion of “polygroup complexity”
- Use these results as a first approximation of results when simulating more realistic biological configurations

# Sunposium™ 2015: Neural circuits and sunshine

**Please Note:** You can watch the live stream of some of the talks here: <http://www.maxplanckflorida.org/news-and-media/sunposium-live-stream/>.

Max Planck Florida Institute for Neuroscience (MPFI) presents Sunposium™ 2015, the second biennial conference highlighting some of the most complex issues at the forefront of understanding neural circuits.

The two-day conference features world-renowned scientists from the Max Planck Society and research institutes and universities throughout the United States.

**Attended by Nathan Crock and Joel Tabak**



Richard Huganir



Na Ji



Yishi Jin



Erik Jorgensen



## James Schummers, PhD

Research Group Leader

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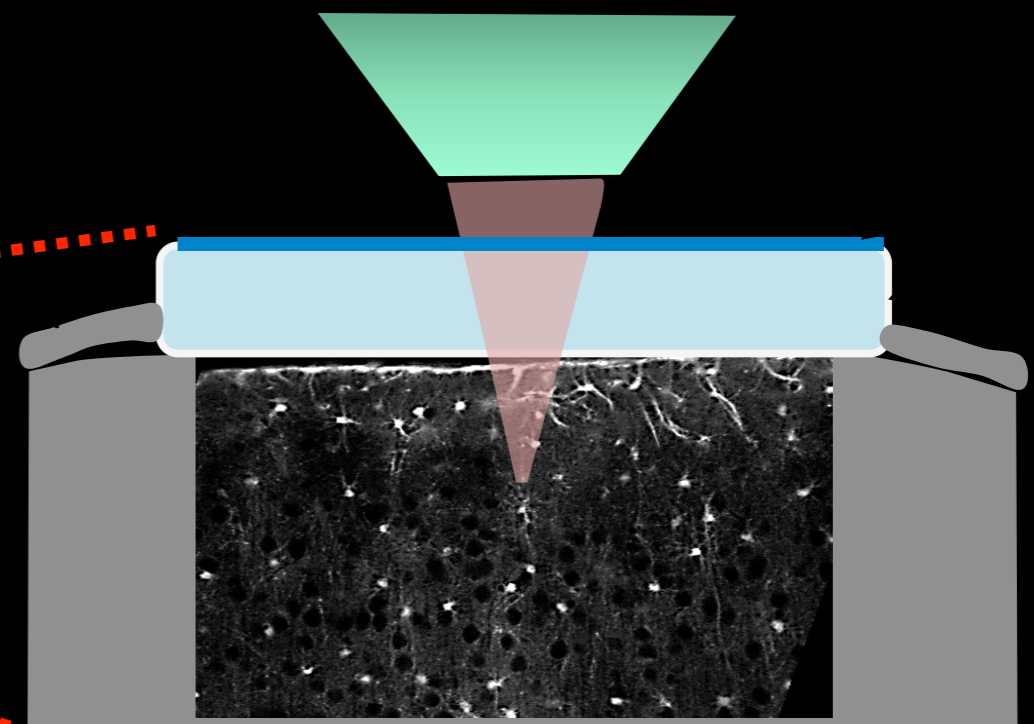
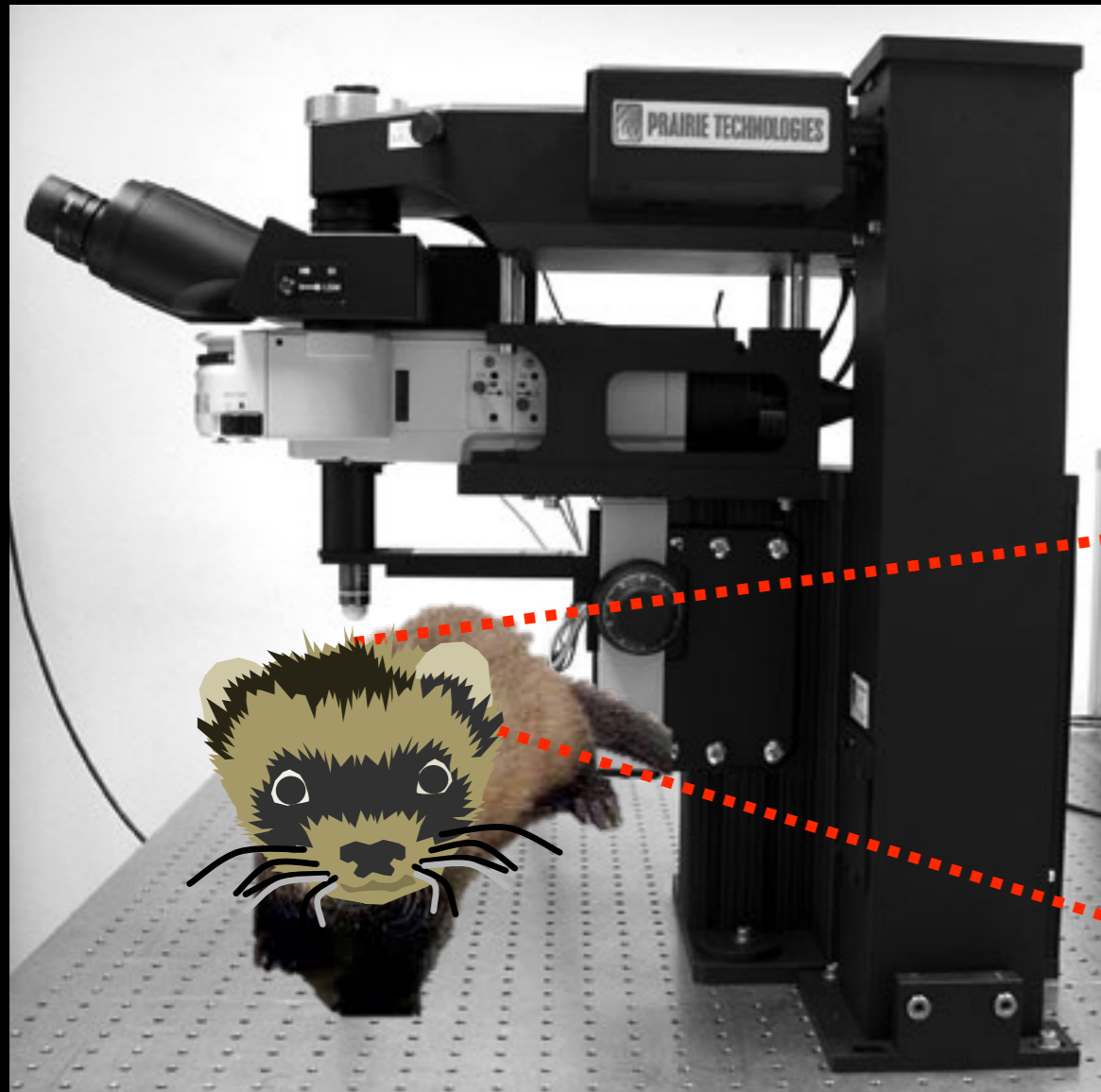
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## Researcher Bio

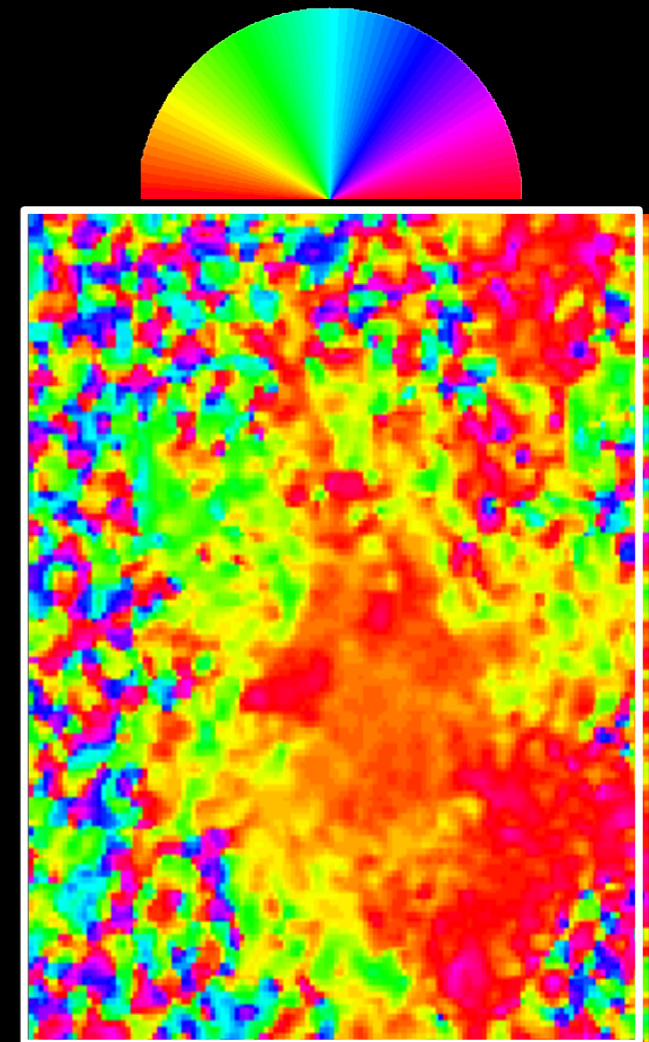
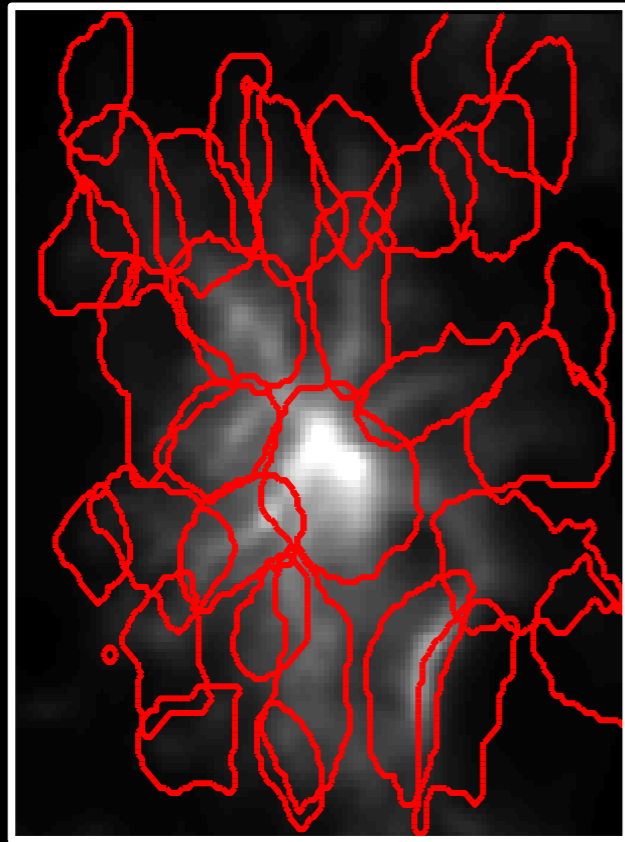
Dr. James Schummers was named an independent Research Group Leader at the Max Planck Florida Institute for Neuroscience in June 2010 and heads the Cellular Organization of Cortical Circuit Function research group. Dr. Schummers received his bachelor's degree in Neuroscience from Oberlin College in Oberlin, OH, where he studied the effects of the neurotransmitter neuropeptide-Y on long-term potentiation (LTP) in the hippocampus. He then moved to Denver CO, where he studied the effects of alcohol on LTP in the Department of Pharmacology at the University of Colorado Health Science Center. He received a PhD in Systems Neuroscience at the Massachusetts Institute of Technology with the support of a Howard Hughes Pre-Doctoral Fellowship. His thesis work combined intracellular and extracellular single neuron recordings with optical imaging approaches to study the integration of synaptic inputs in the context of visual processing. His postdoctoral work, also at MIT, focused on single-cell resolution imaging to study the response properties of different classes of cells, including both neurons and astrocytes, in the visual cortex.

# In vivo 2-photon imaging in ferret visual cortex

- in vivo two photon imaging
- Lightly anesthetized (isoflurane)
- Adult ferrets

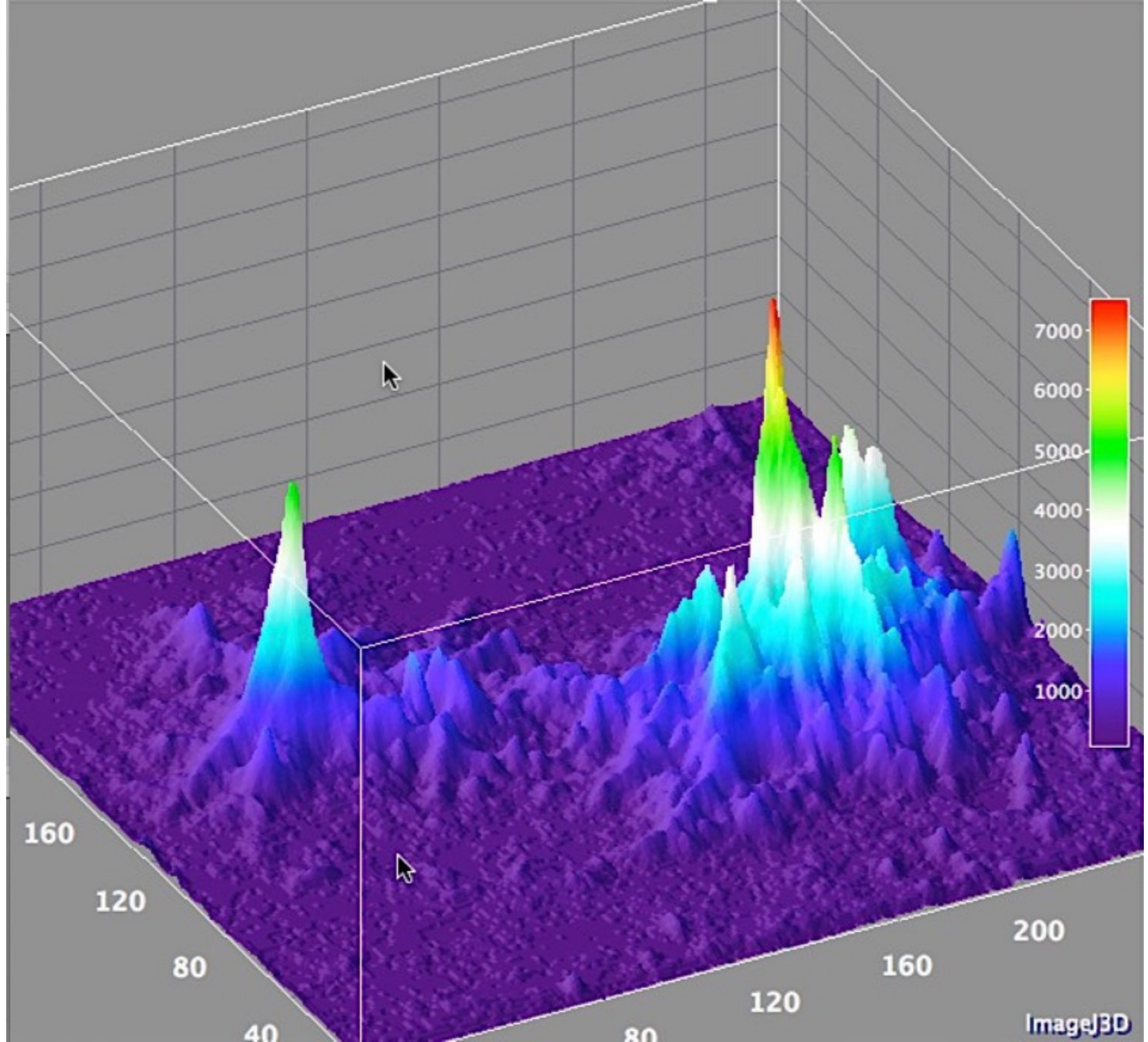


# Orientation tuning in subcellular domains



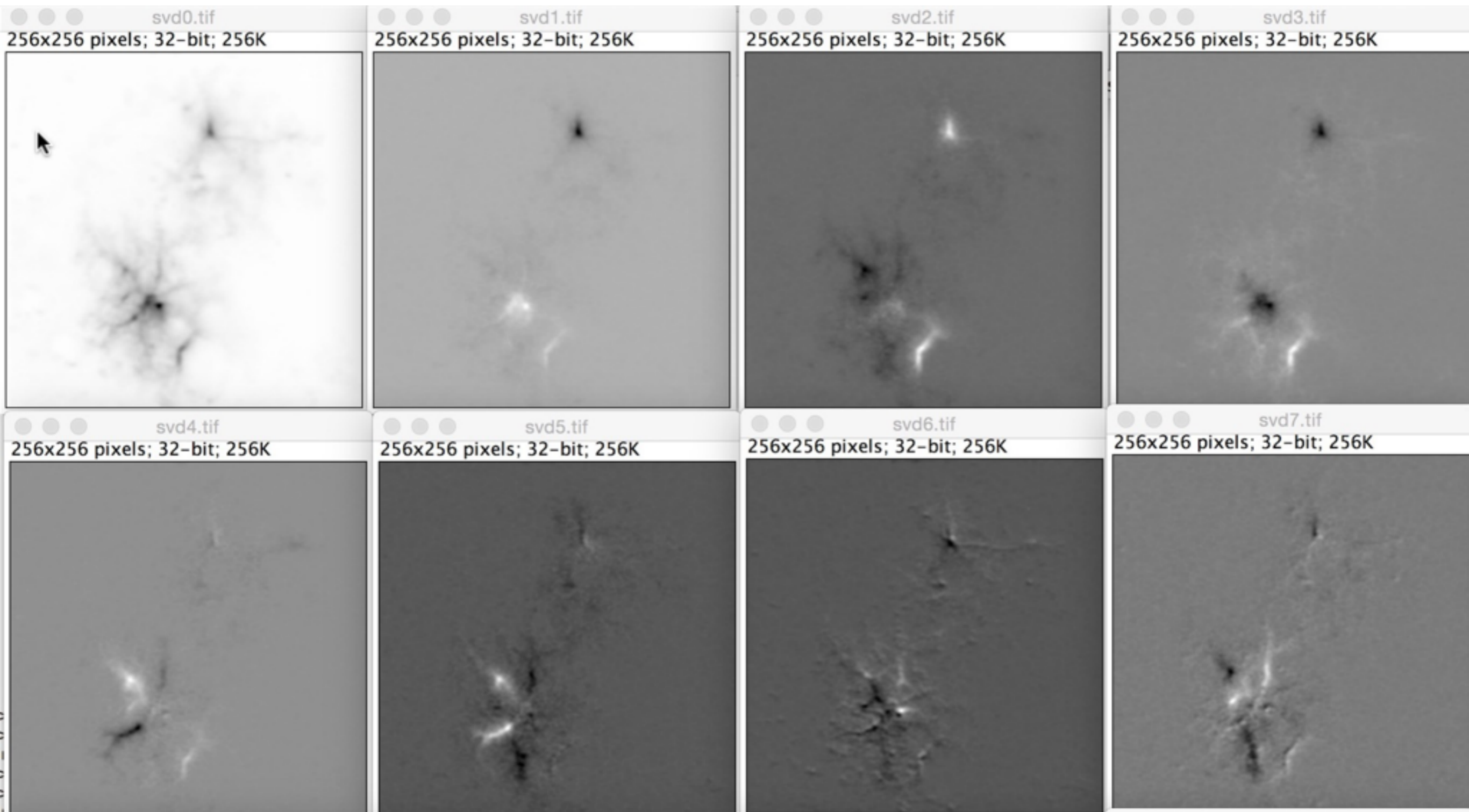
Each sub-domain has similar orientation tuning, with quantitative differences

Does this suggest that they are responding to distinct neural activity?



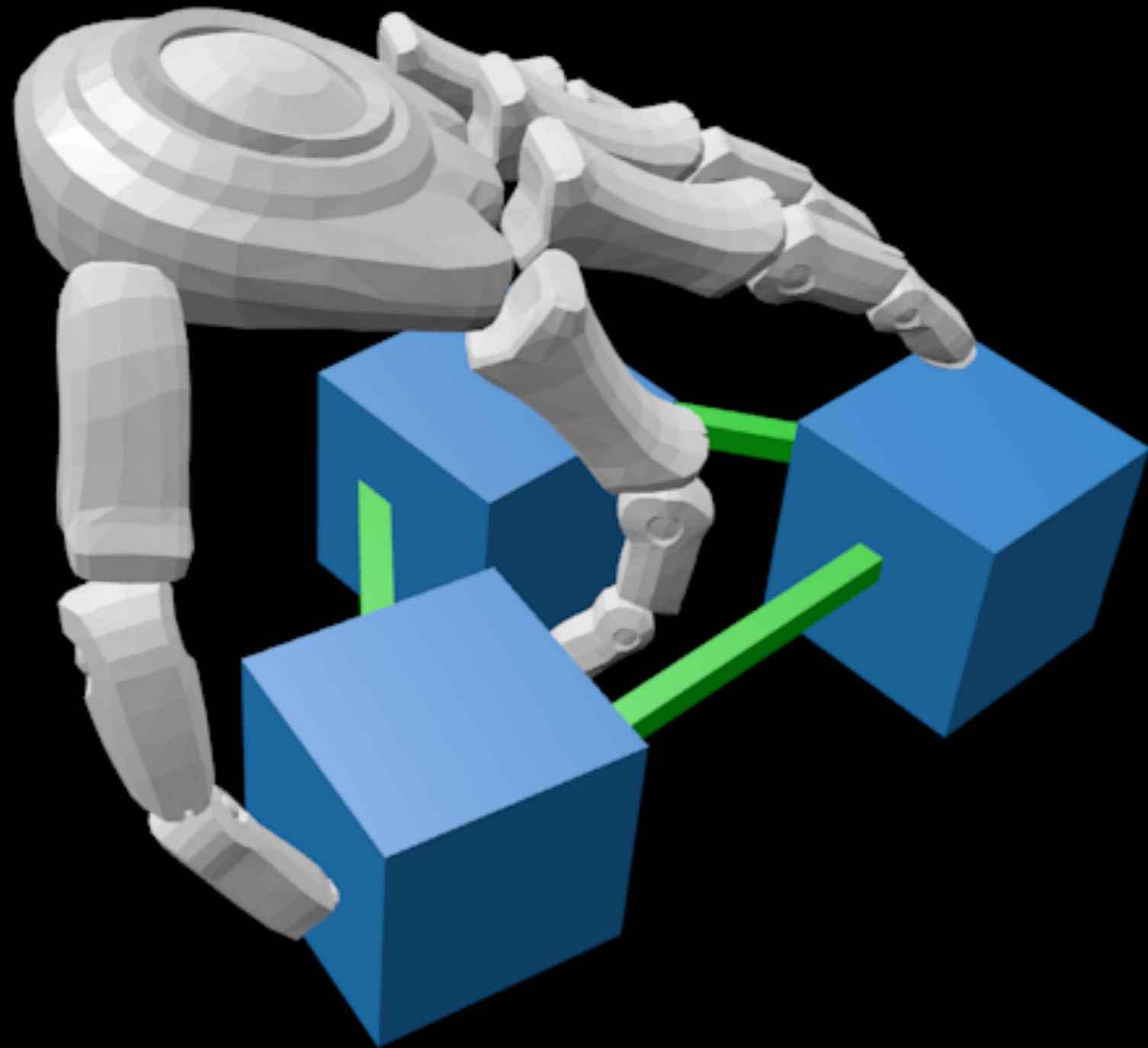


# PCA, first 8 modes



# Oculus Rift + Leap Motion





# Leap + Oculus

- Work of Juan Llanos
- Objectives
  - construct and manipulate a neuron network in 3D space in a natural way
  - develop a backend, interaction computational engine
  - explore the results from this engine as

# Research Group

- Gordon Erlebacher, Lead
- Joel Tabak, FSU Program in Neuroscience
- Nathan Crock (Ph.D.)
  - Astrocyte data analysis, polychrony
- Evan Cresswell (Ms)
  - Astrocyte modeling, plasticity
- Juan Llanos (Ms)
  - Interactive modeling software and visualization