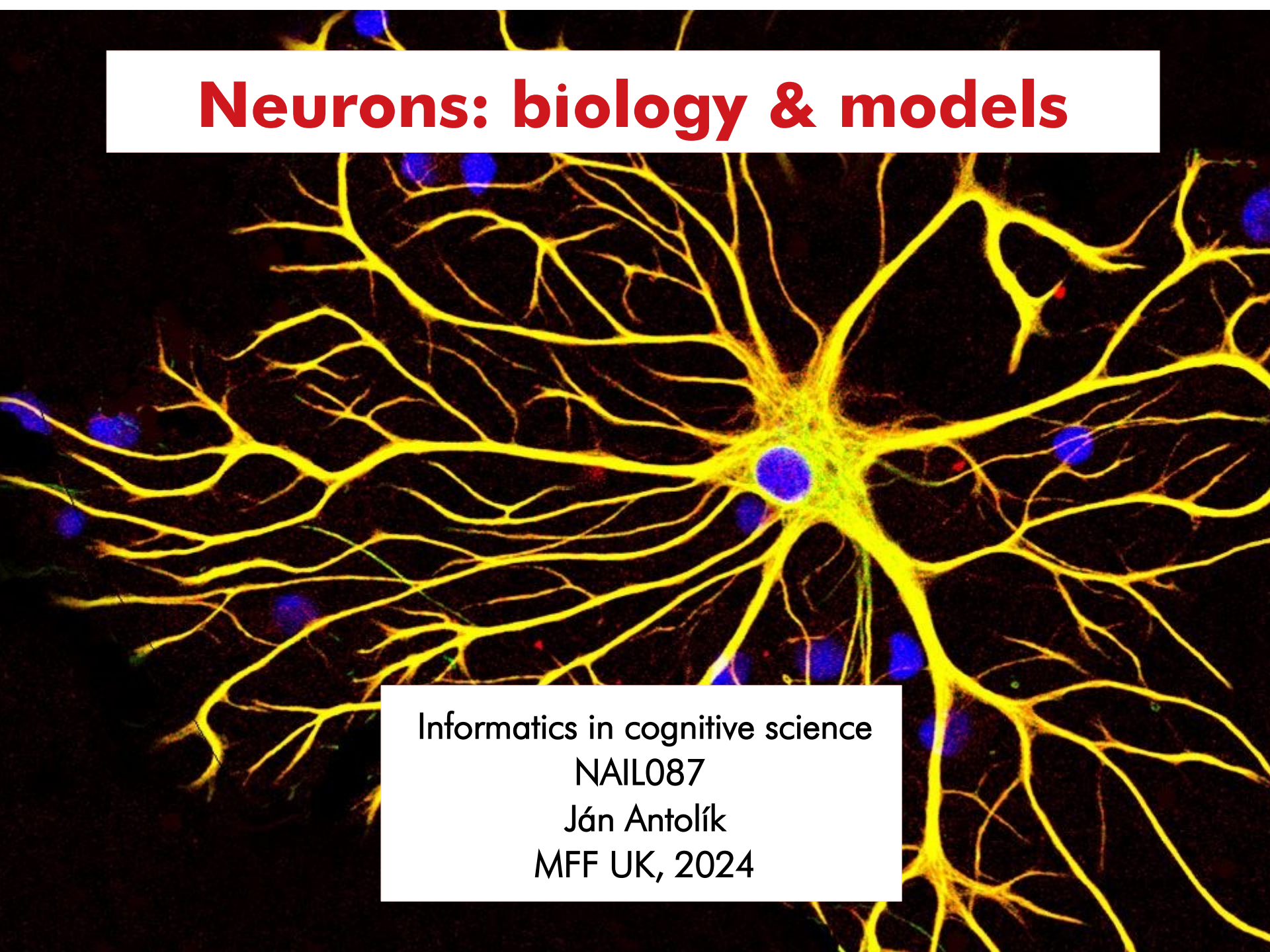


Neurons: biology & models



Informatics in cognitive science
NAIL087
Ján Antolík
MFF UK, 2024

Lecture Outline

- Little recap
- Axons, dendrites, soma
- Neuronal Membrane
- Synapses and synaptic transmission
- Neuron as a signal integration element
- Integrate & Fire abstraction of neural operation
- Outlook

Little recap



How did we simplify brain?

Little recap



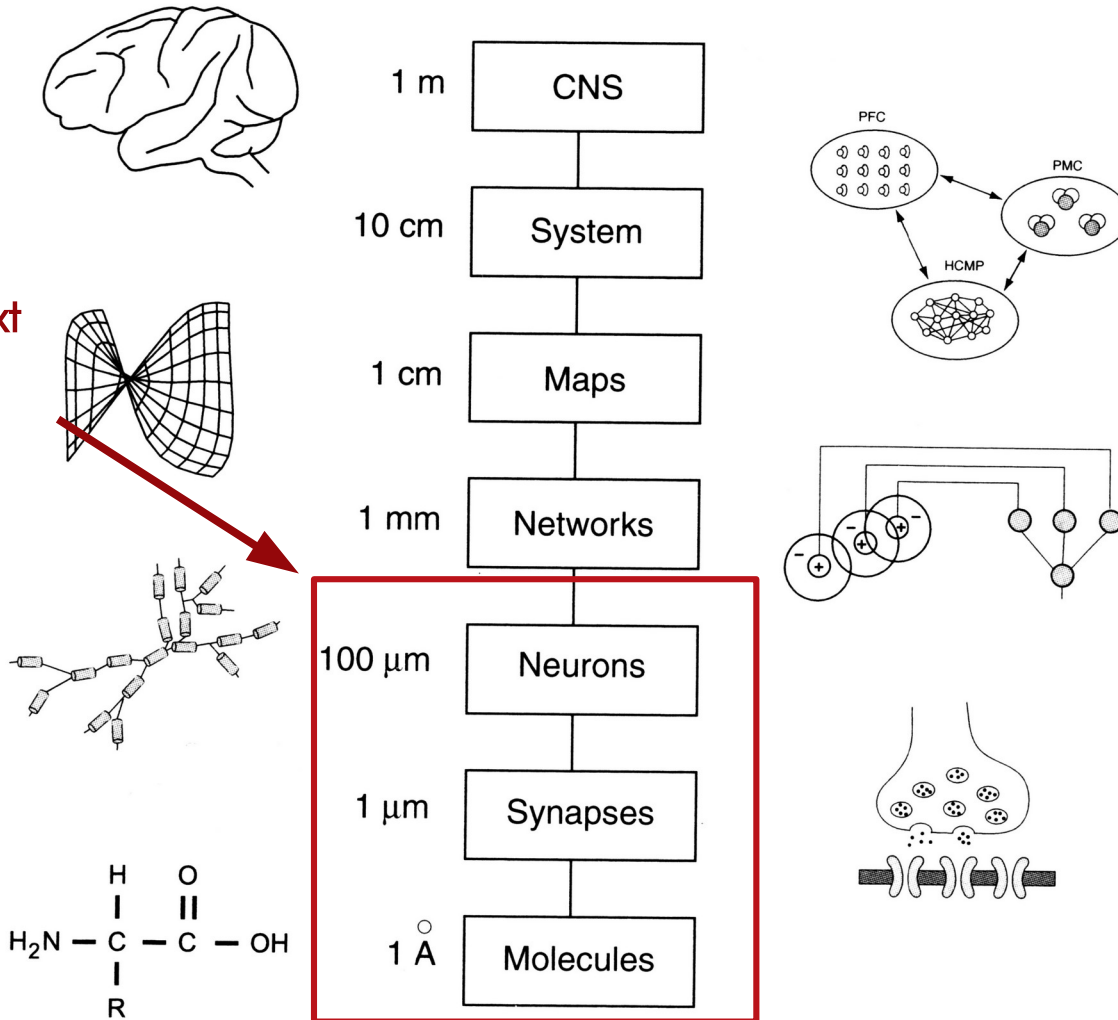
What could be the elementary units from which to build our understanding of neural computation?

Why do we care about neurons?

Neurons, axons, dendrites and synapses are to neuroscience what atoms and electrons are to physics.

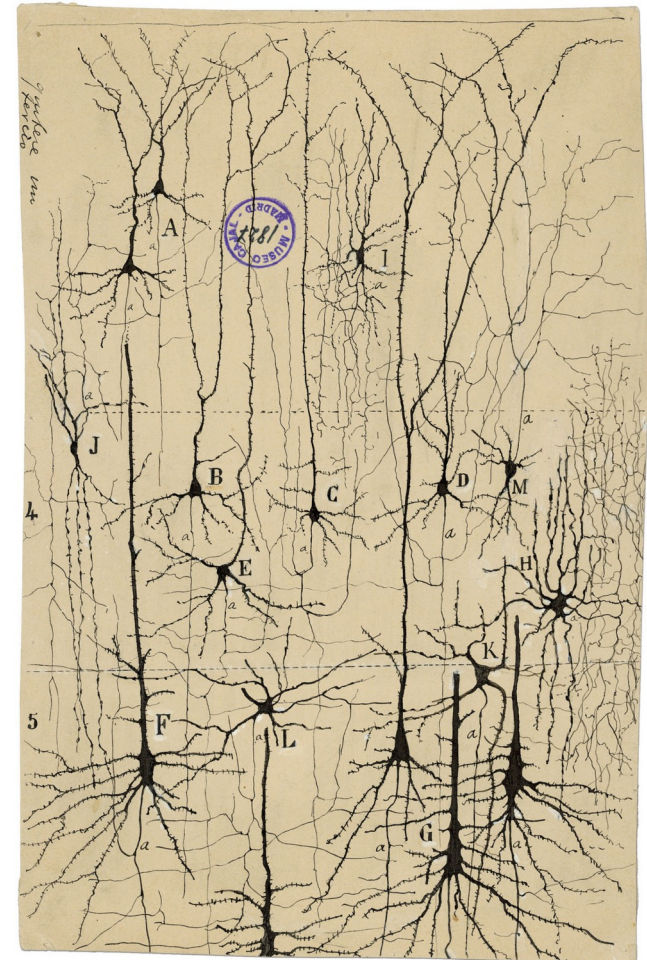
Brain as a multi-scale system

In this and next
lecture we will
work here



The discovery of neuron

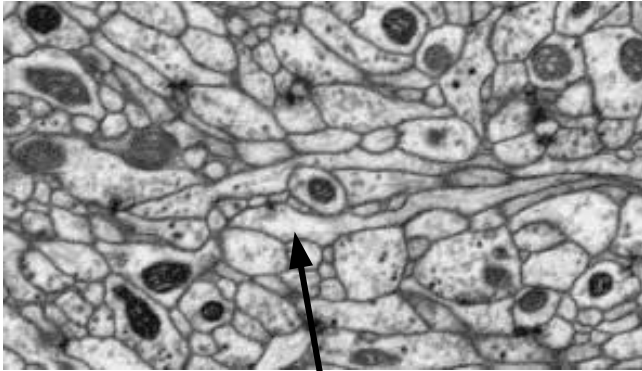
- Glogi's method of silver staining
- Discovered by S.R. Cajal
- The dye is injected in the neural tissue
- It stains the neurons which can then be viewed under optical microscope



Santiago Ramón y Cajal

Ways to look at it

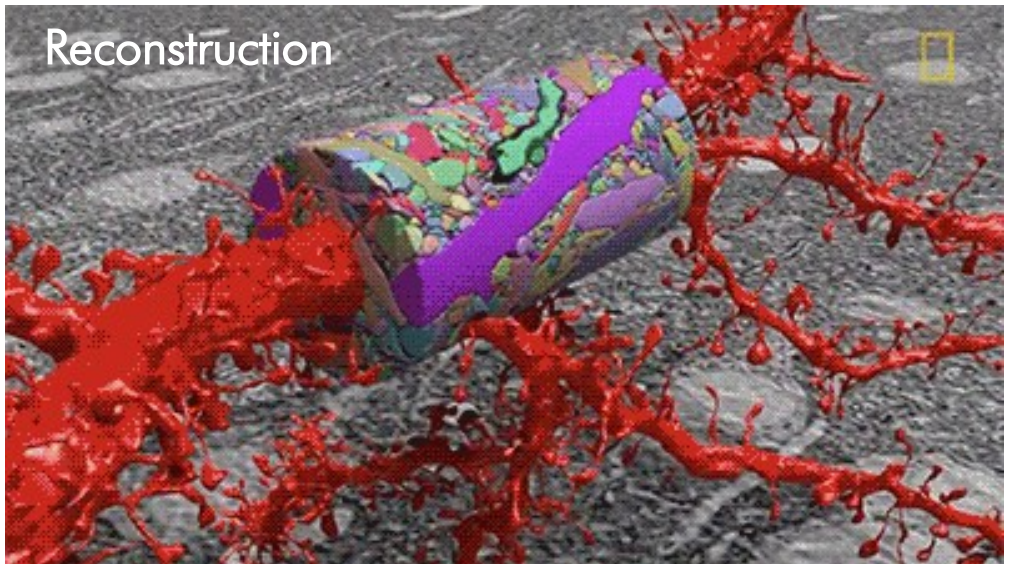
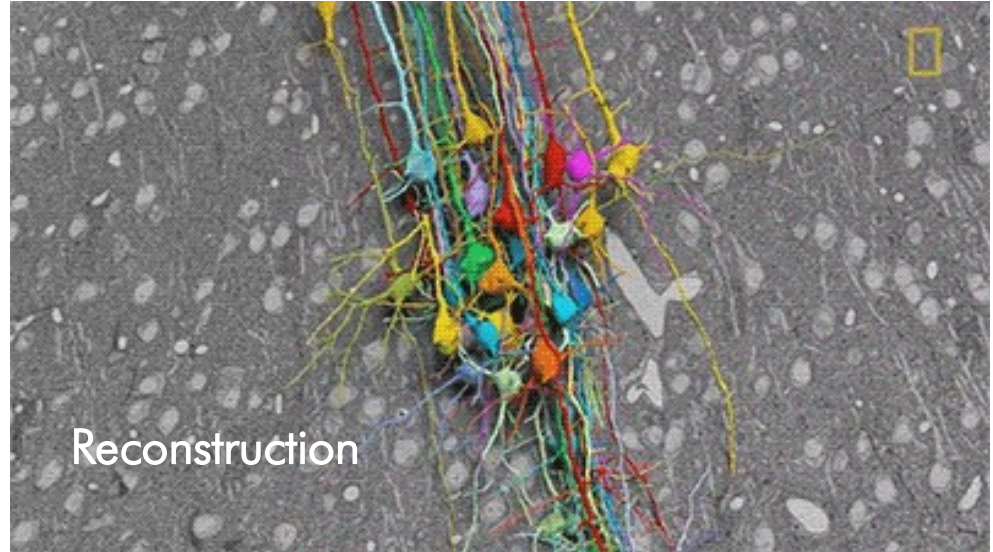
Electron microscope view



Apropos 3km of wires: each delineated region here here belongs to a different cell.

Compromises between different methods of studying neurons:

- * spatial resolution
- * recording length
- * fixed/in-vitro/in-vivo tissue

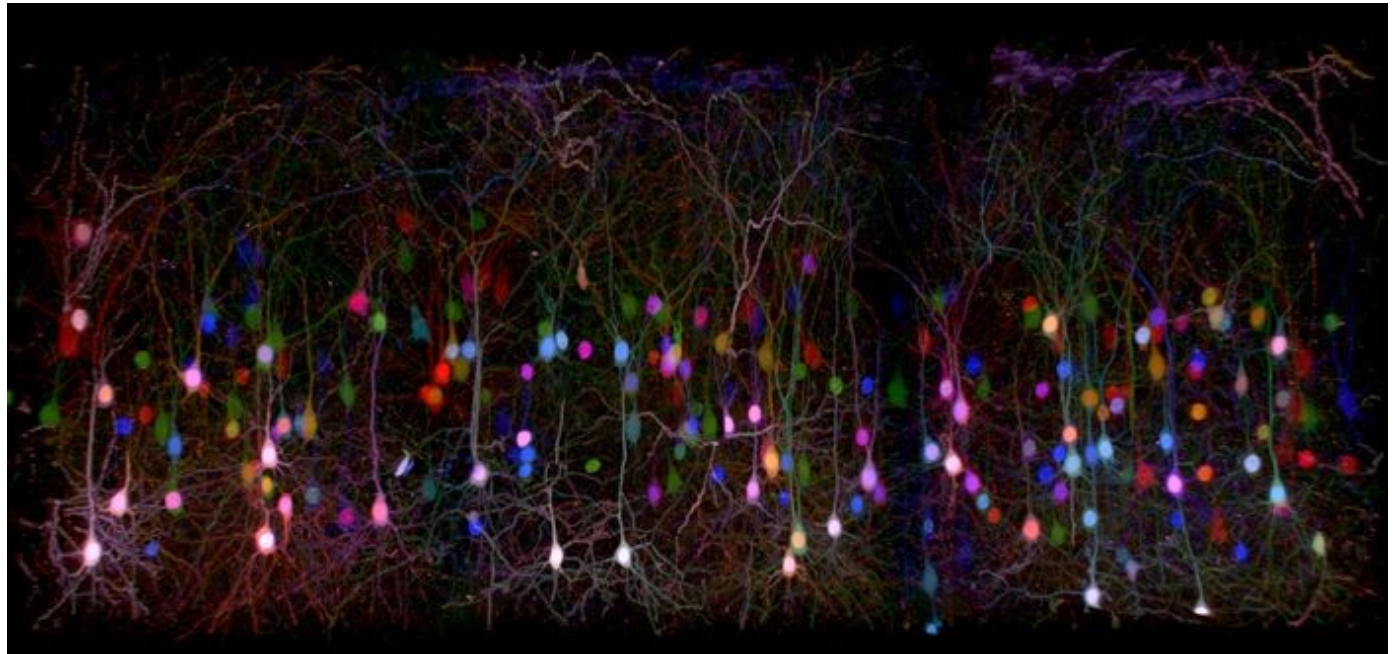
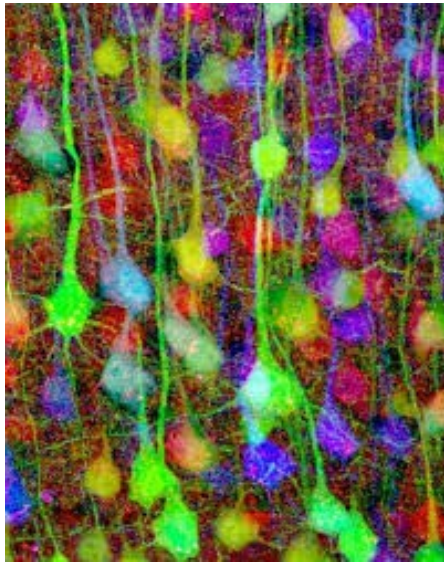
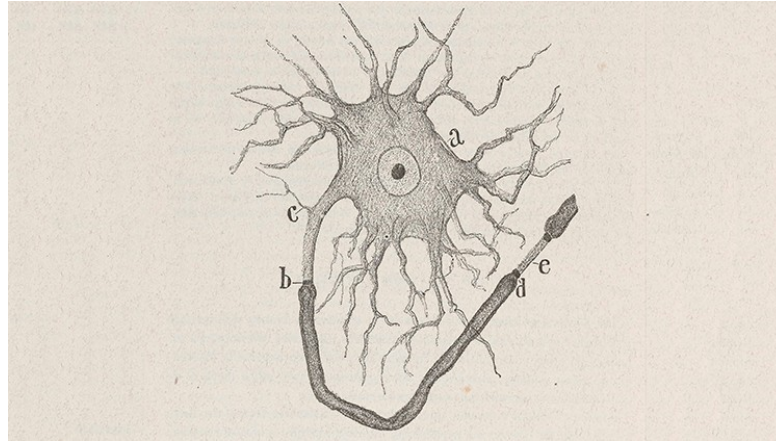


in-vitro vs. in-vivo



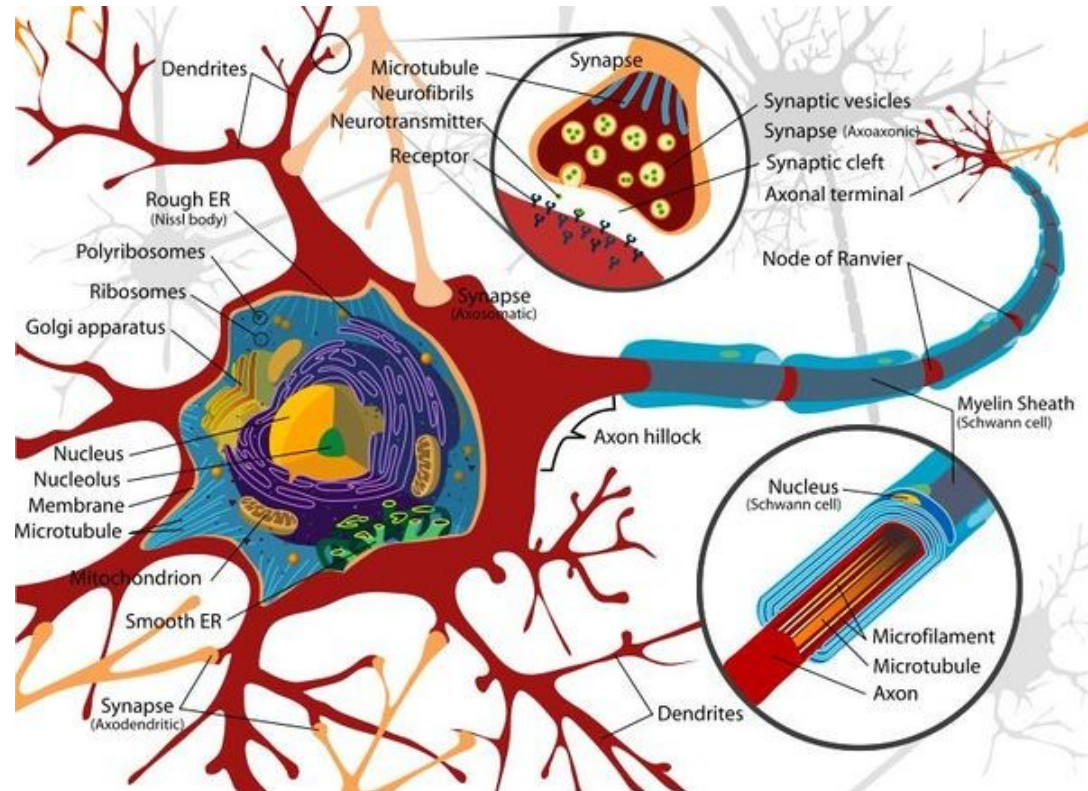
What's the difference?

Ways to look at it



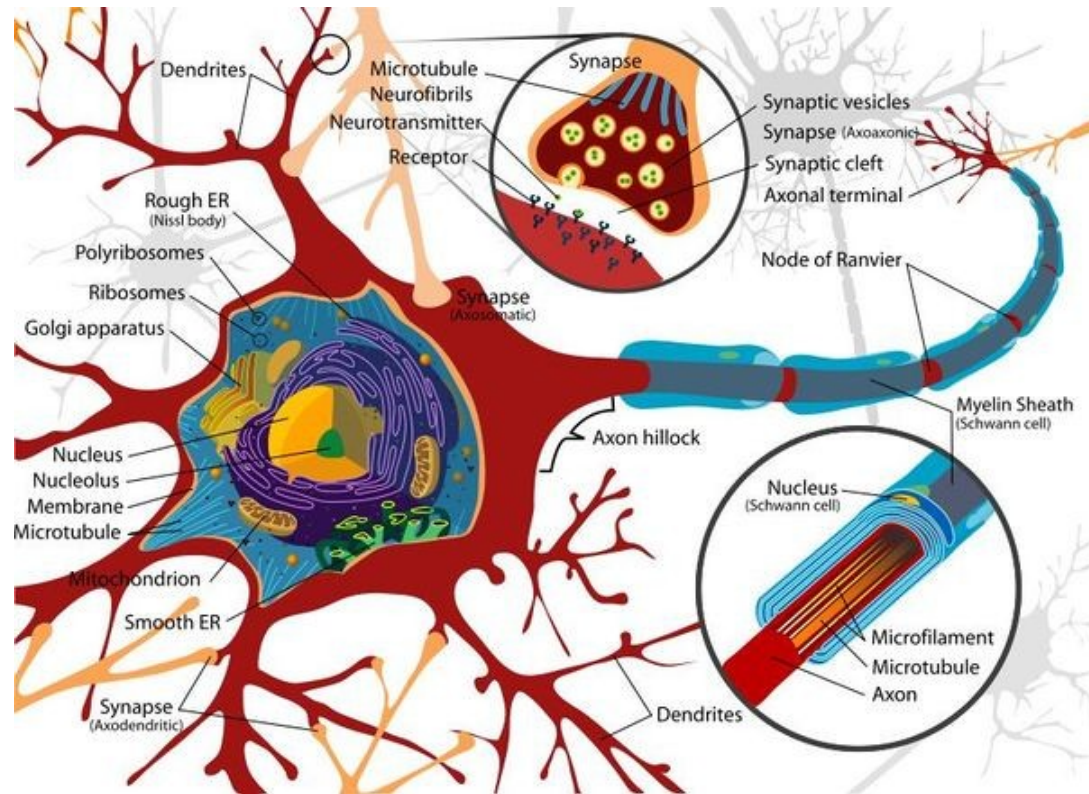
Neuron is just another cell

- Nucleus
- Ribosome
- Mitochondrias
- Cytoskeleton
- Endopl. reticulum

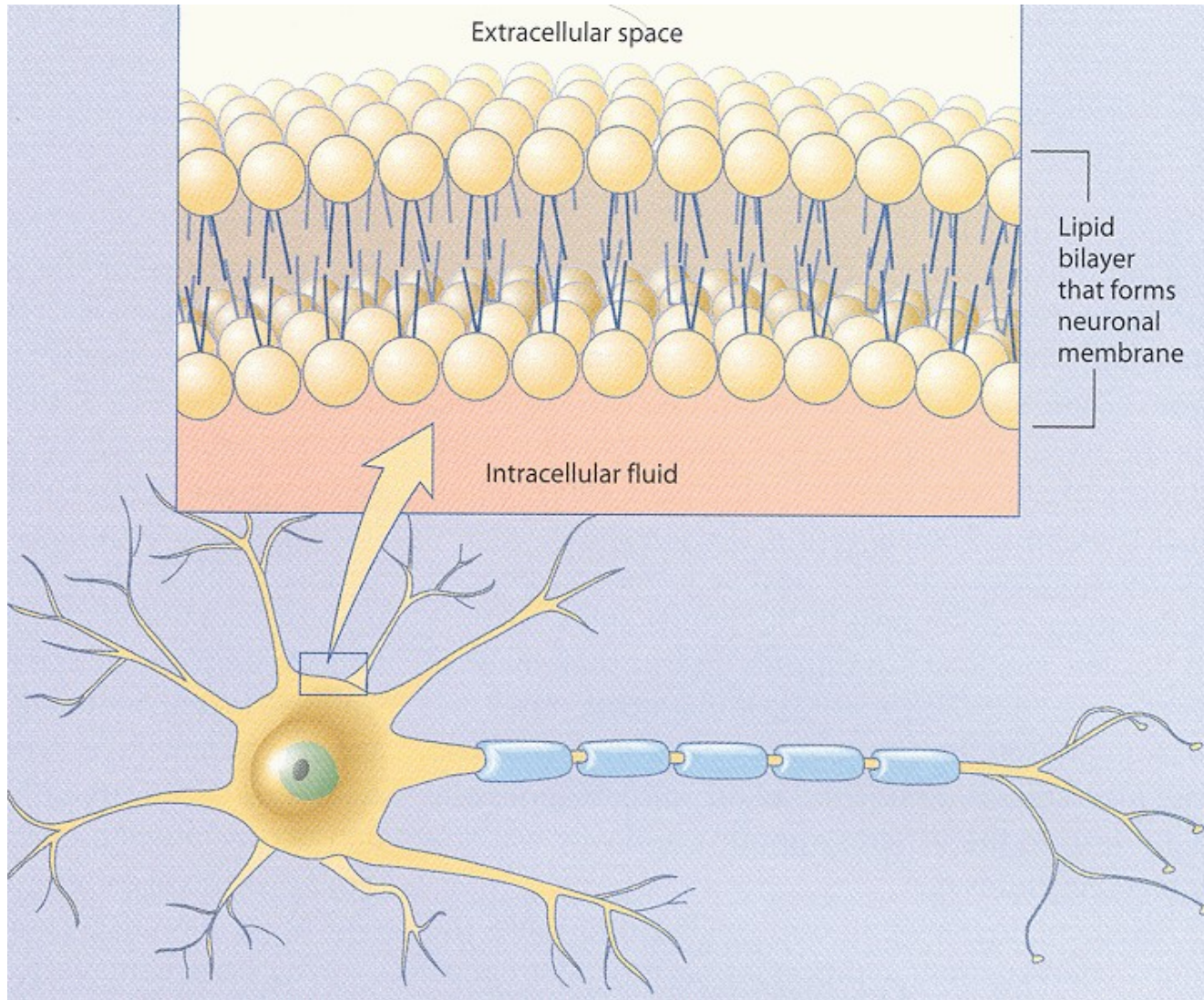


The key components

- Membrane
- Dendrites
- Soma
- Axon
- Synapses
- Spines



Neural membrane



Membrane potential (single ion)

Blackboard time

Nernst equation

$$E_X = \frac{RT}{zF} \ln \frac{[X]_{\text{out}}}{[X]_{\text{in}}}$$

$$E_X = \frac{58}{z} \log \frac{[X]_{\text{out}}}{[X]_{\text{in}}}$$

R : Gas constant
T : Absolute temperature
F : Faraday constant
z : Valence

Equilibrium potential for K^+

- Concentration inside: 150 mMol/L
- Concentration outside: 5 mMol/L

$$[K^+]_{in} = 130 \text{ mM} \quad [K^+]_{out} = 5 \text{ mM} \quad z = +1$$

$$E_{K^+} = \frac{58 \text{ mV}}{1} \log \frac{5}{130} = -82 \text{ mV}$$

Membrane potential (multiple ions)

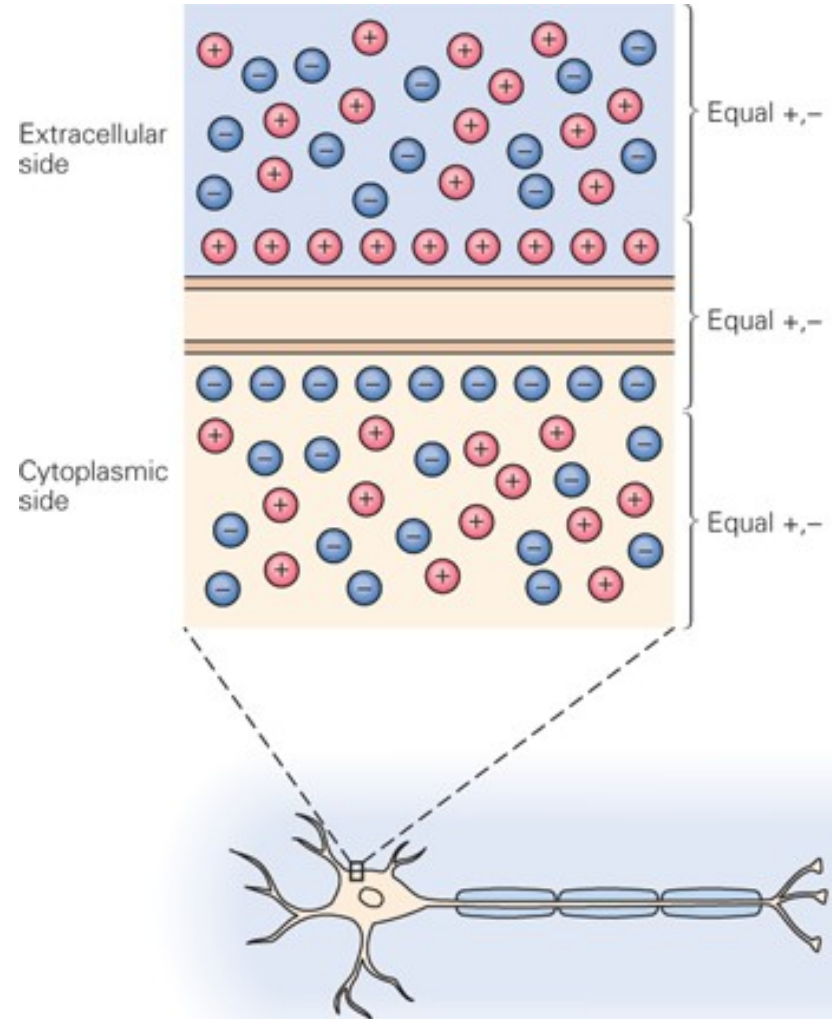
Blackboard time

**How can neurons change
their membrane potential?**



Membrane potential summary

- Equilibrium potential
 - single ion
 - concentration gradient
 - remains constant
 - difference of charge
 - permeability
- Membrane potential
 - Equilibrium potential across all ions

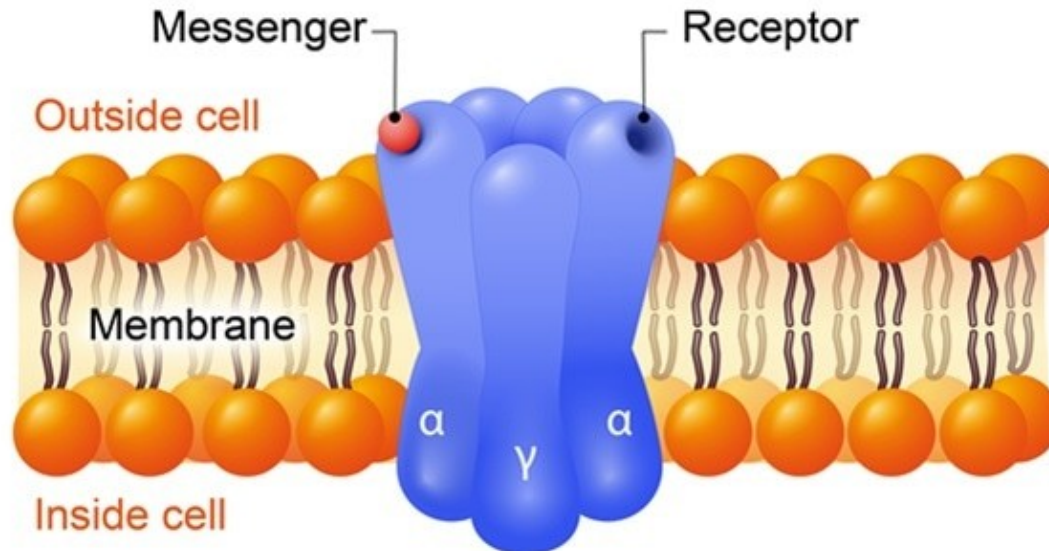
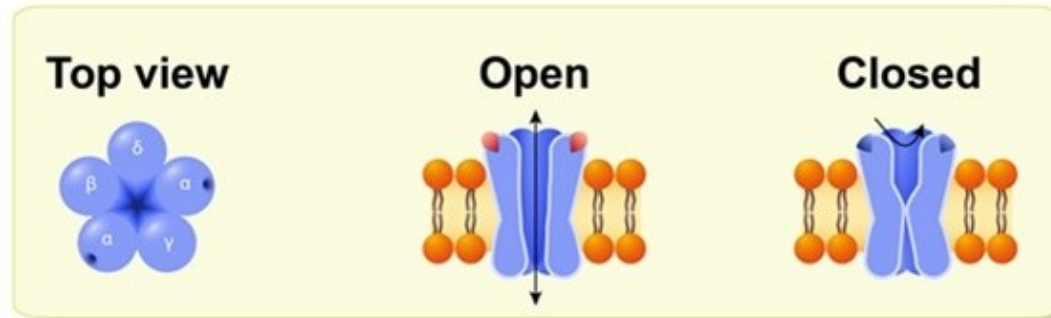


**How can cell change its
membrane permeability?**



Membrane Channels

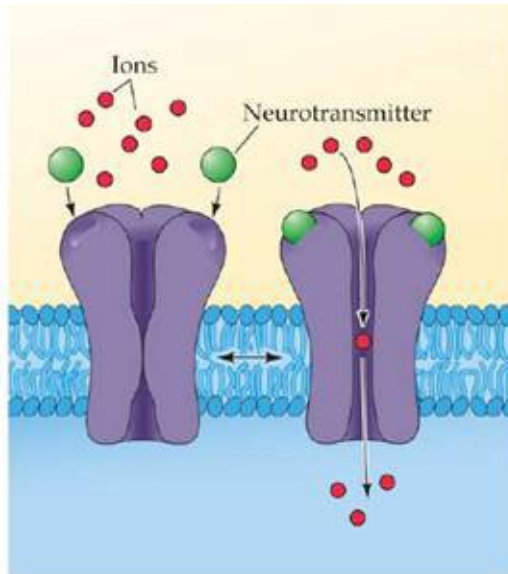
ION CHANNEL



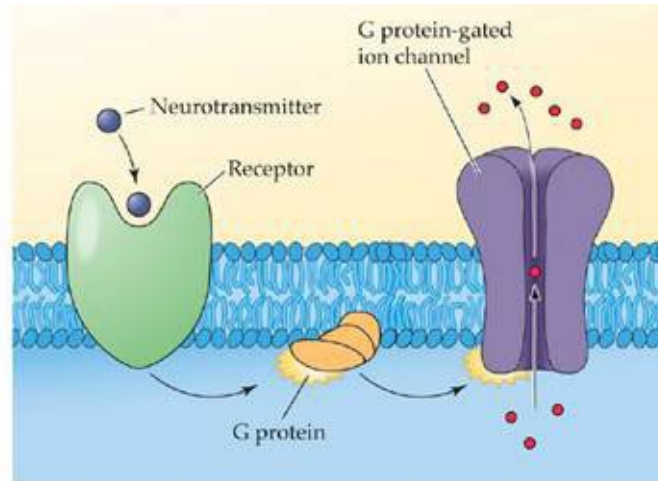
**Design a channel that will
depolarize a cell?**



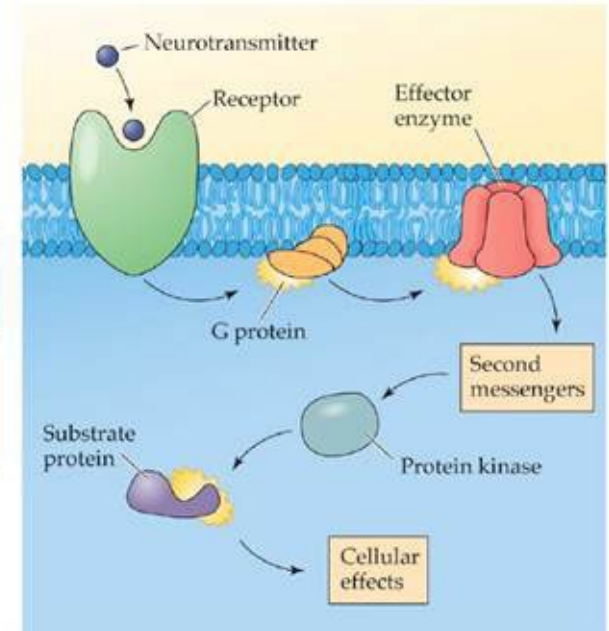
Channel Types



(A) An ionotropic receptor, which allows ions into the cell directly

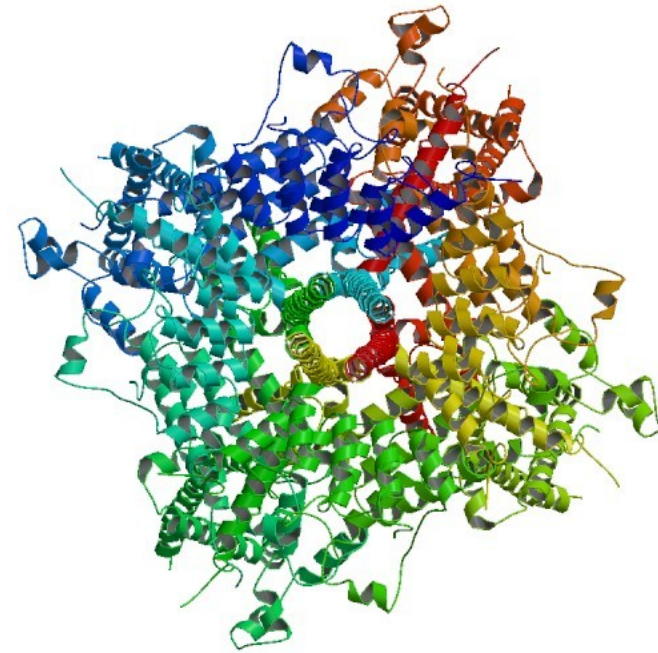
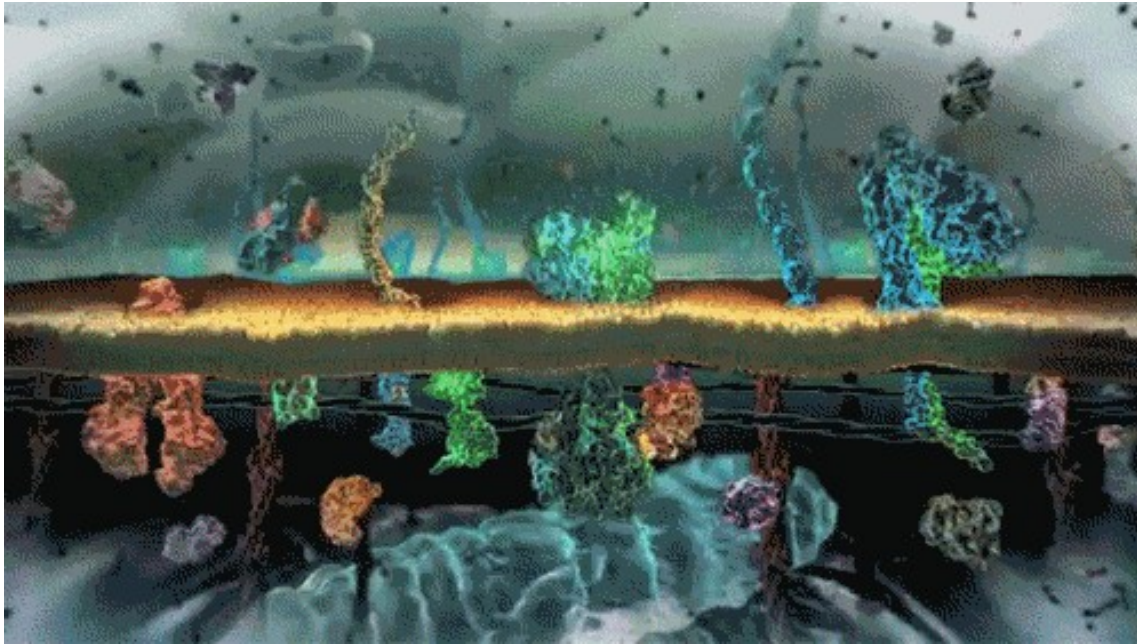


(B) A metabotropic receptor, which uses a G protein to activate an ion channel and let ions into the cell

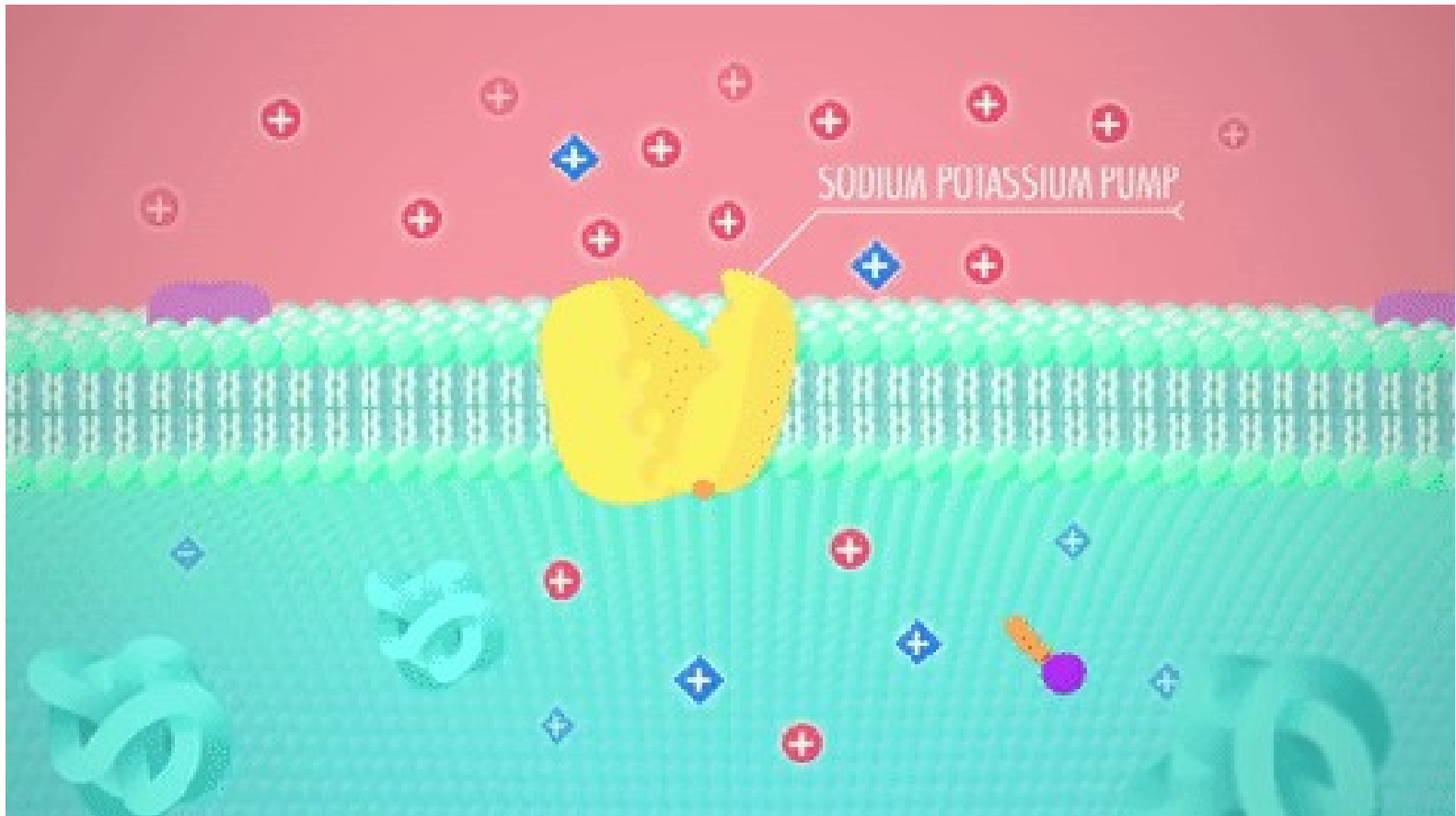


(C) A metabotropic receptor which uses G proteins to create a signalling cascade

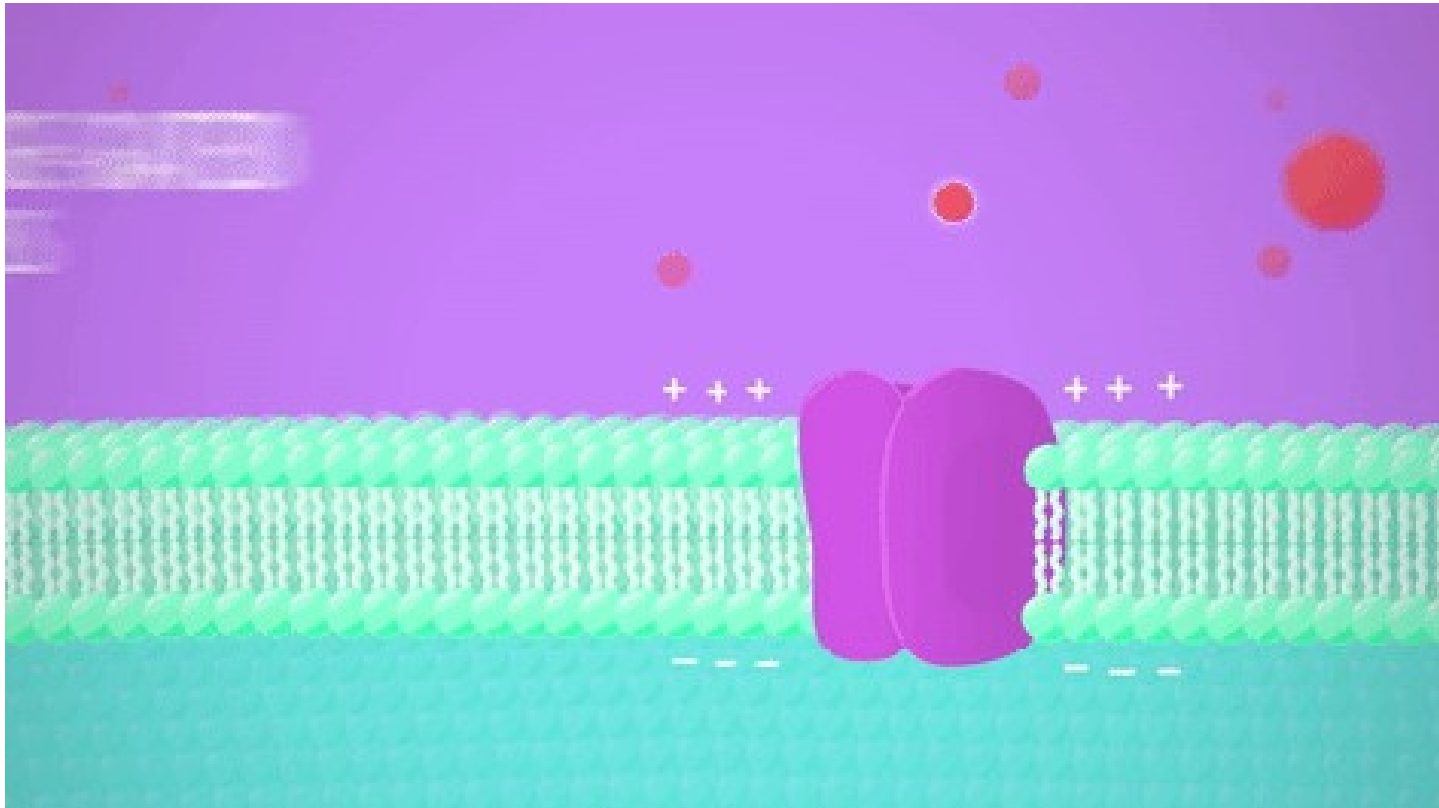
Channel structure



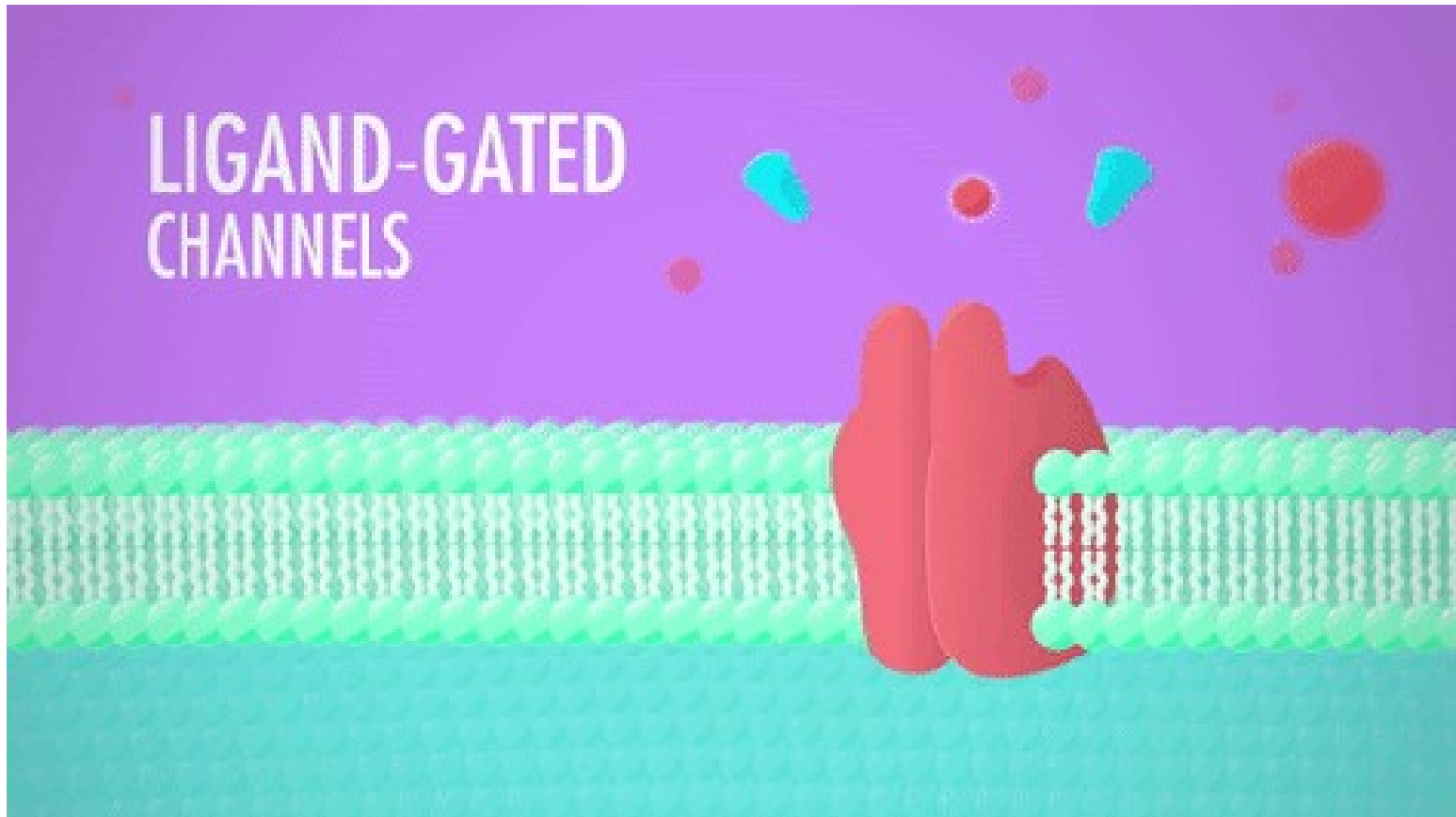
Sodium–Potassium Channels



Voltage gated channels



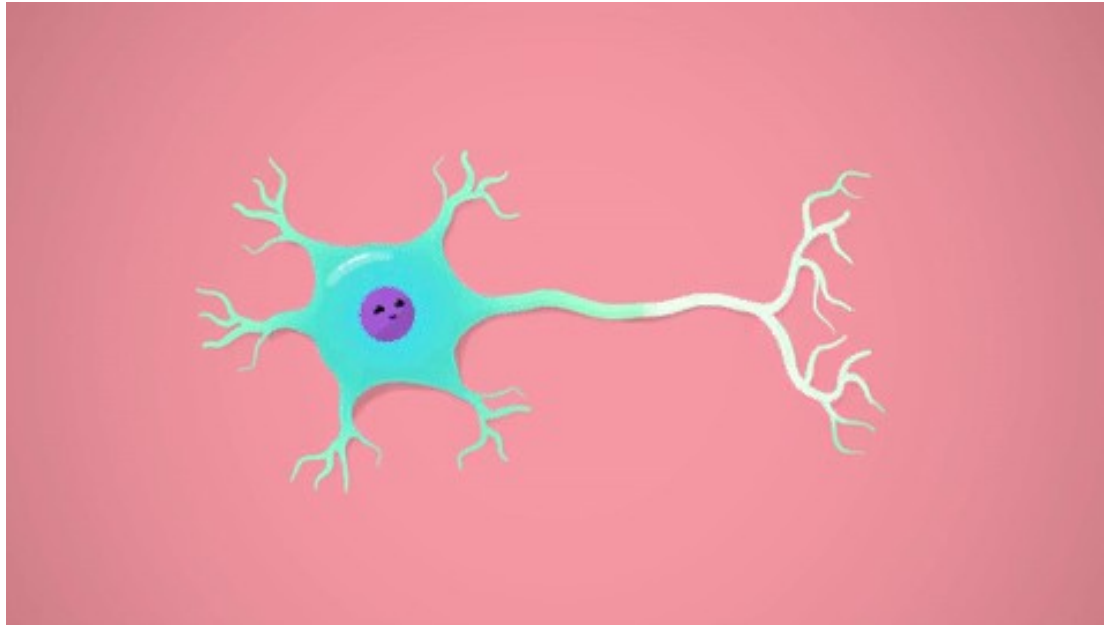
Ligand Gated Channels



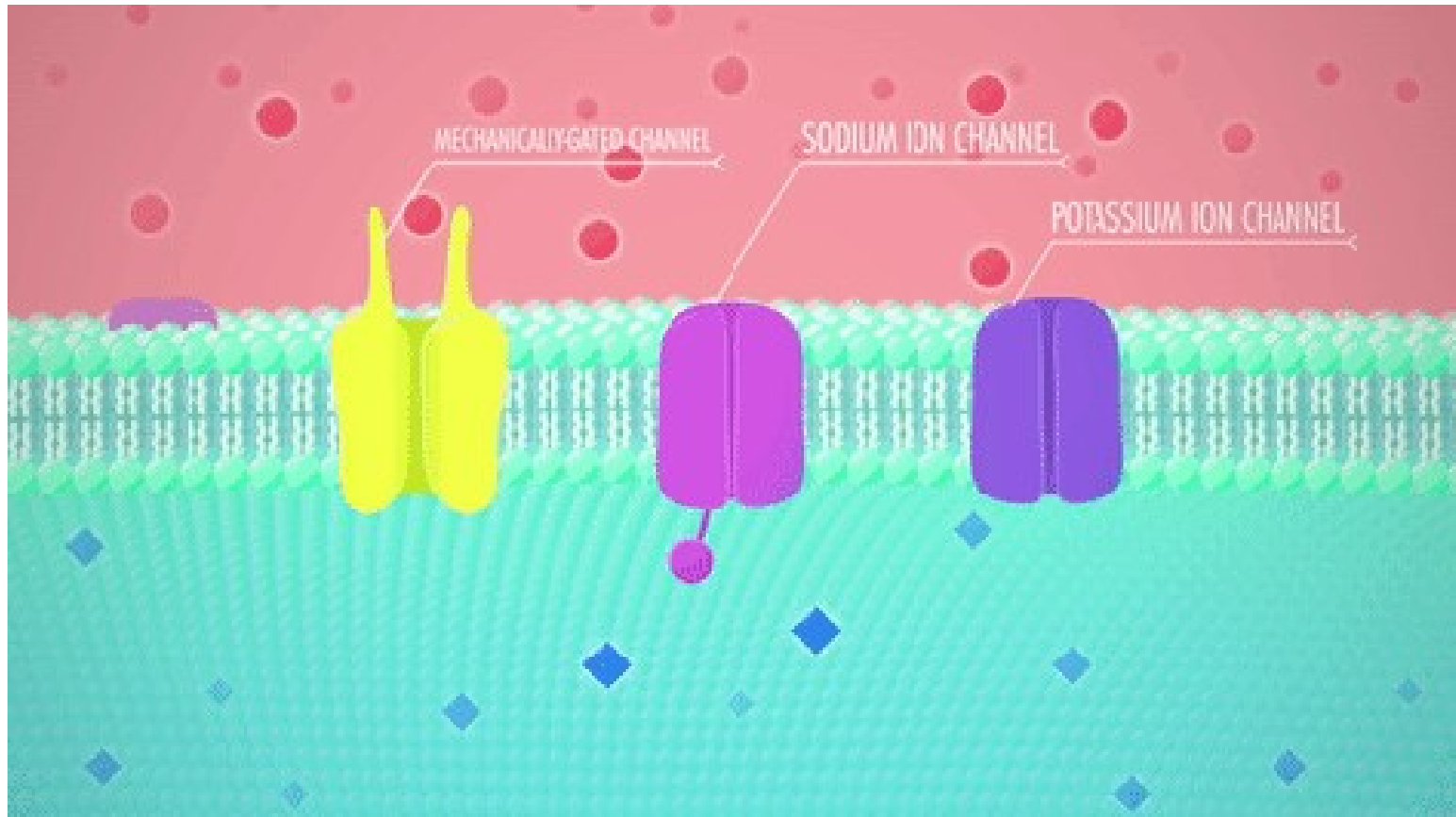
How to transmit information using membrane potential?



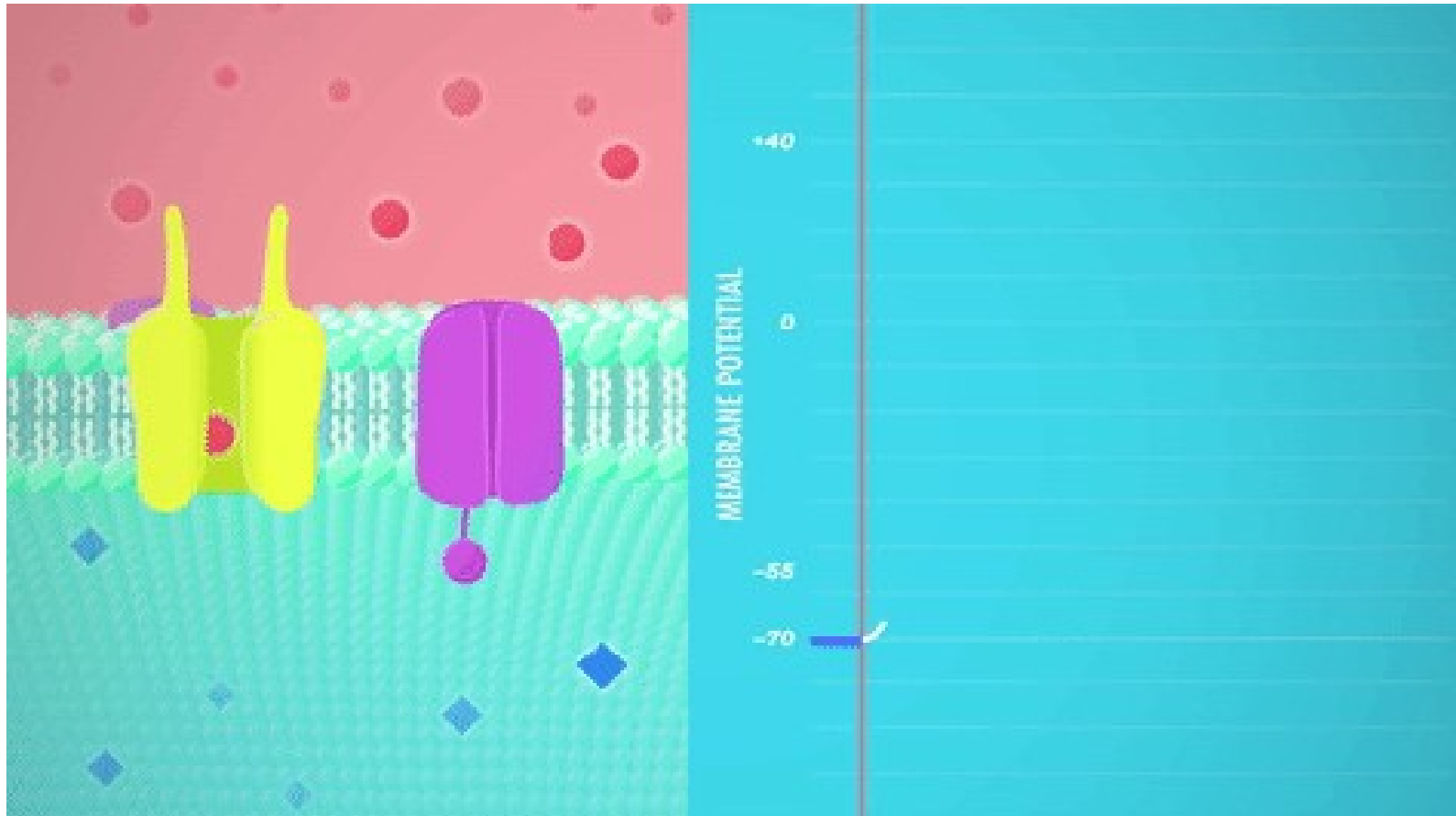
Action Potential



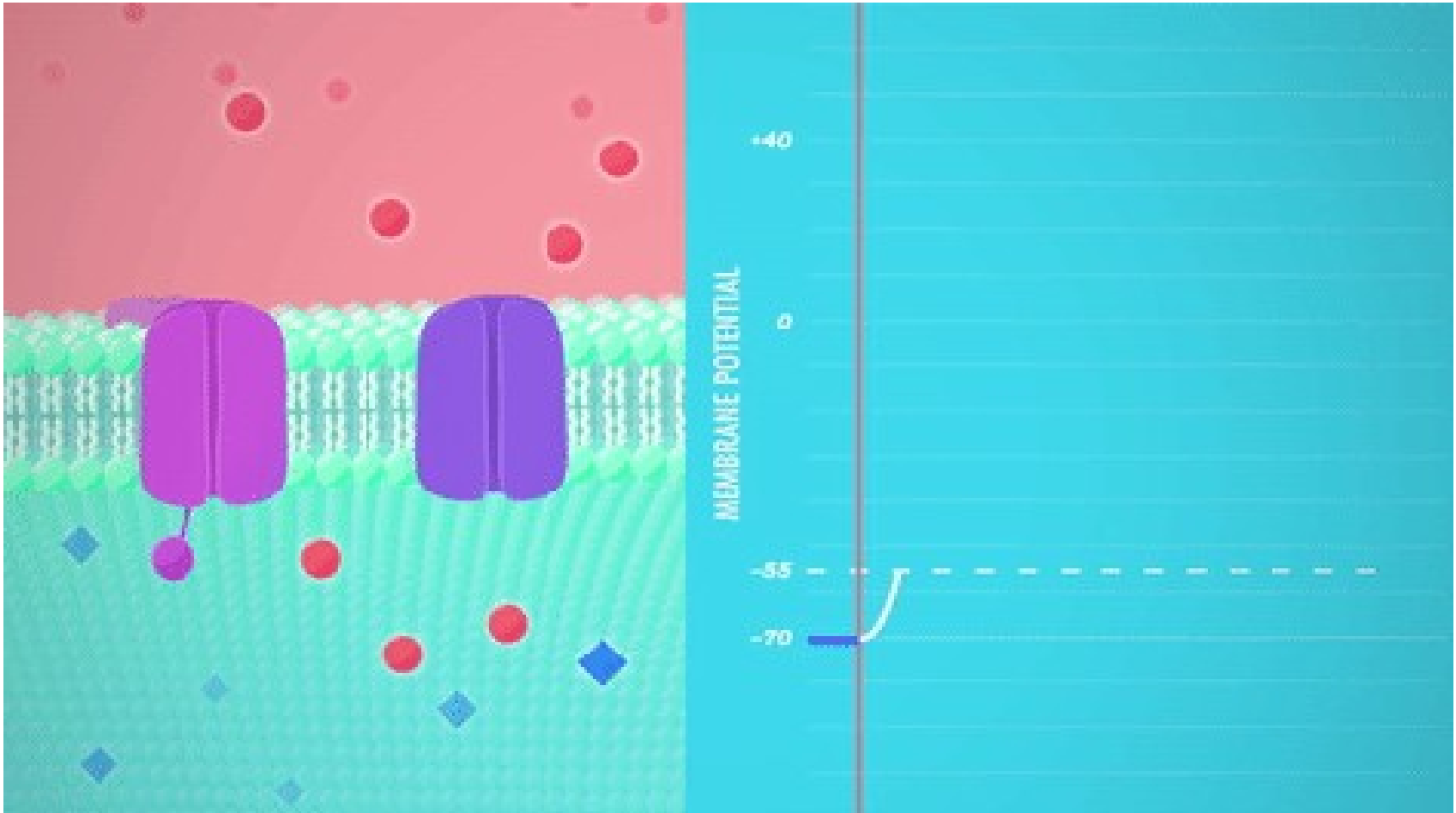
Resting membrane potential



Its all about getting to –55mV



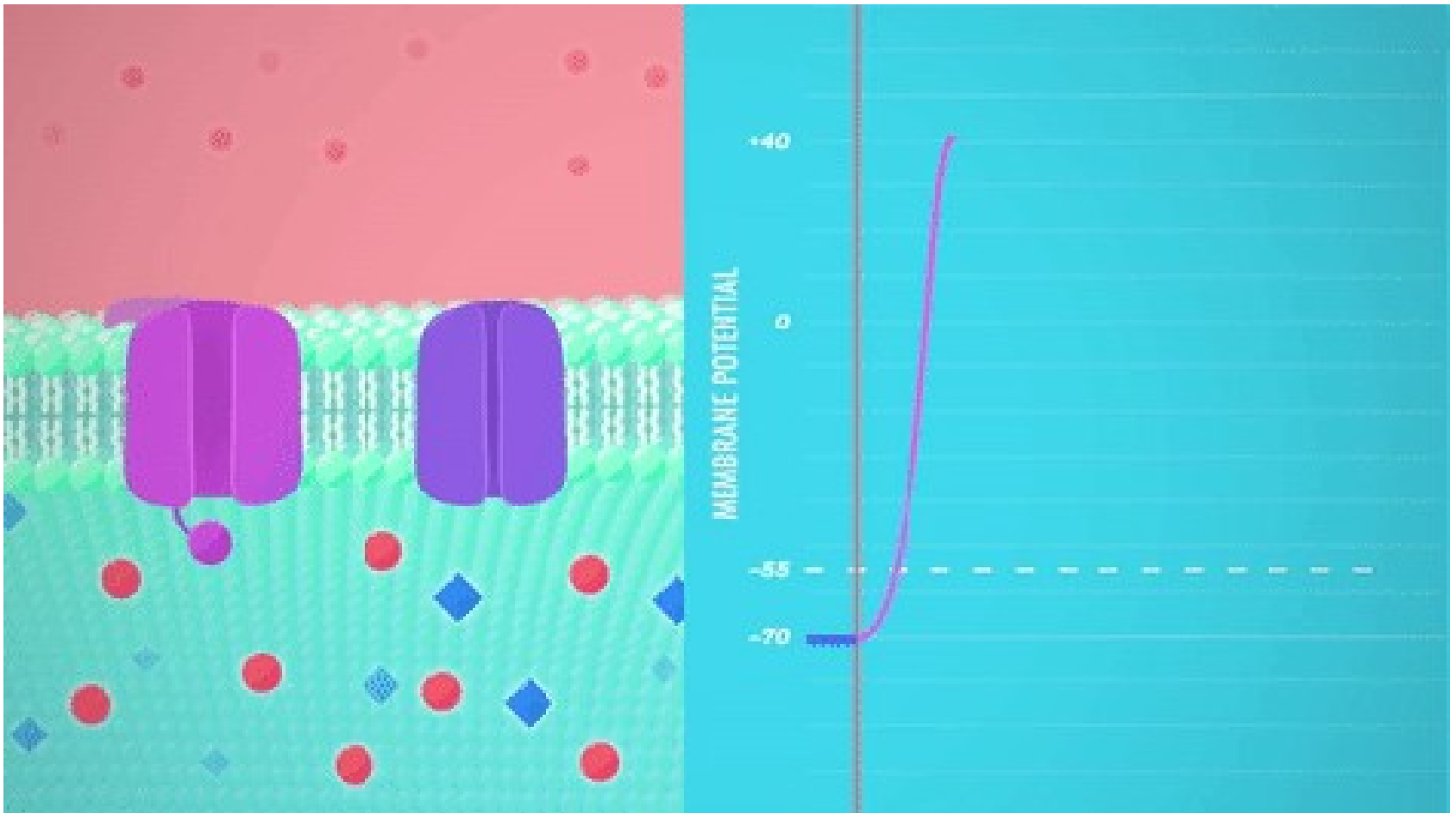
Depolarization



Why does membrane potential go all the way to positive?



Repolarization

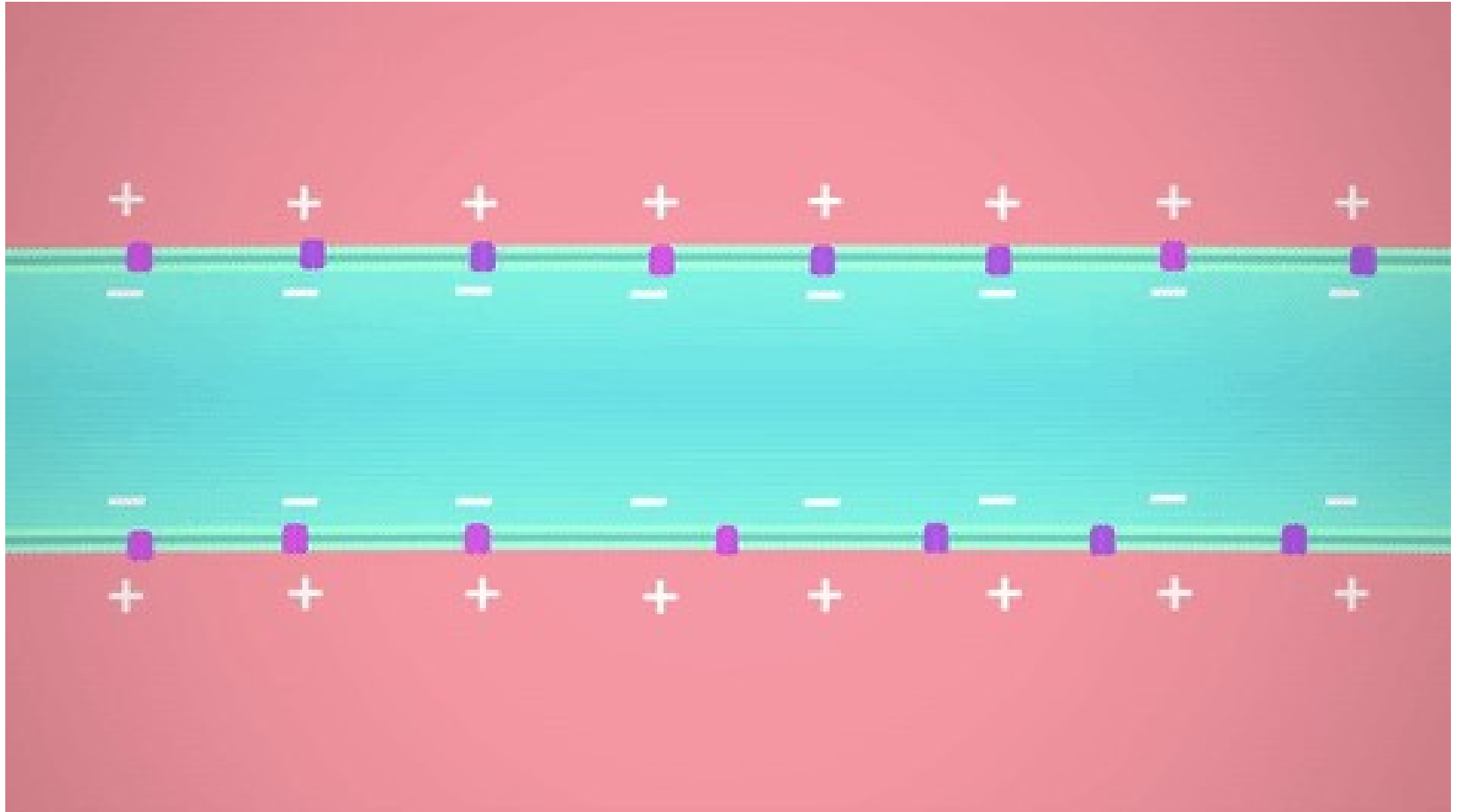


**OK, we have evoked
depolarization in a bit of
membrane.**



**But how does the signal get
further?**

Action Potential

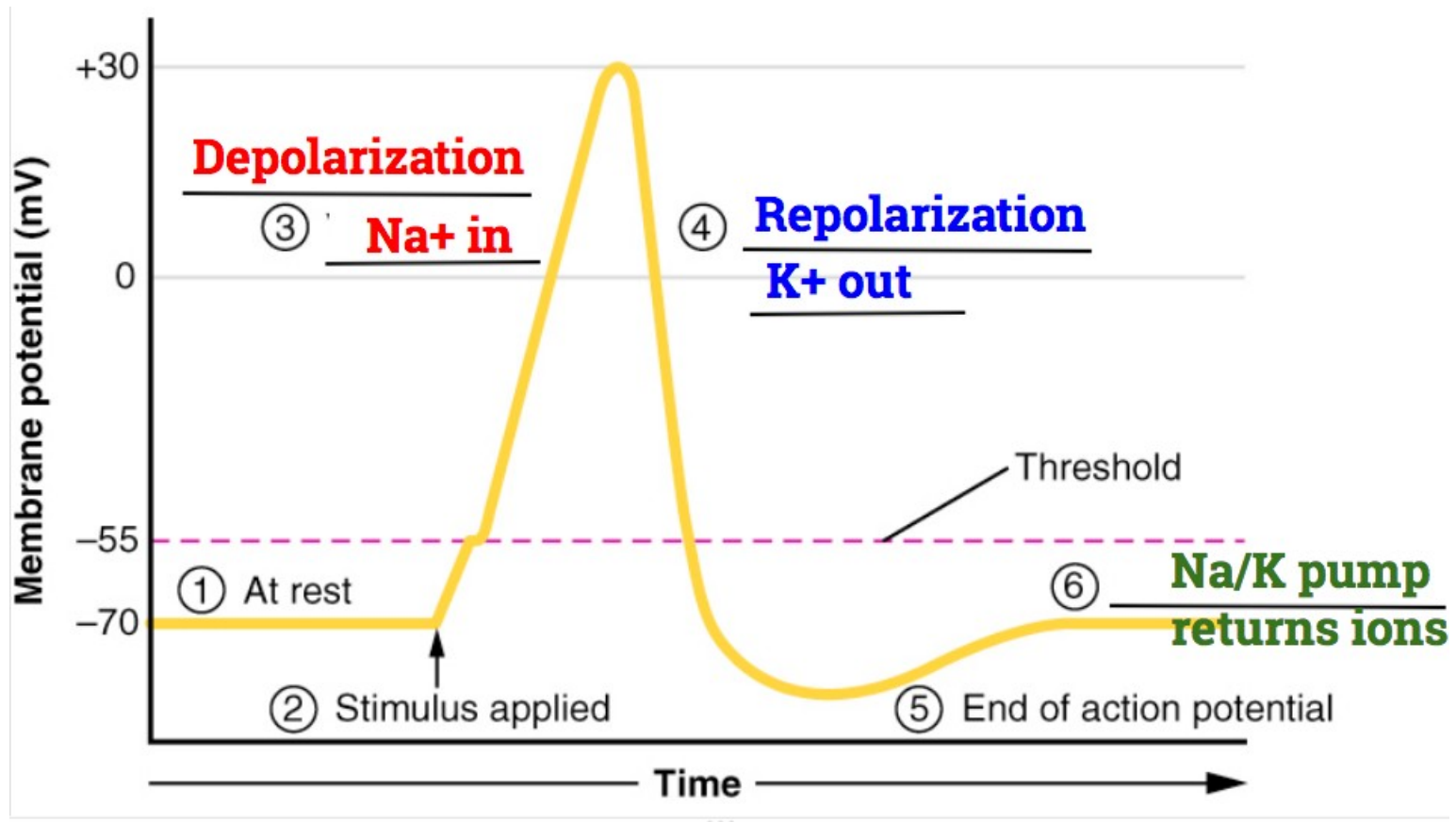


**OK we have Action
Potential wave.**



**Is the mechanism we have
described complete/self-
consistent?**

Action Potential summary



**We are transmitting
signals.**



Are we doing it for free?

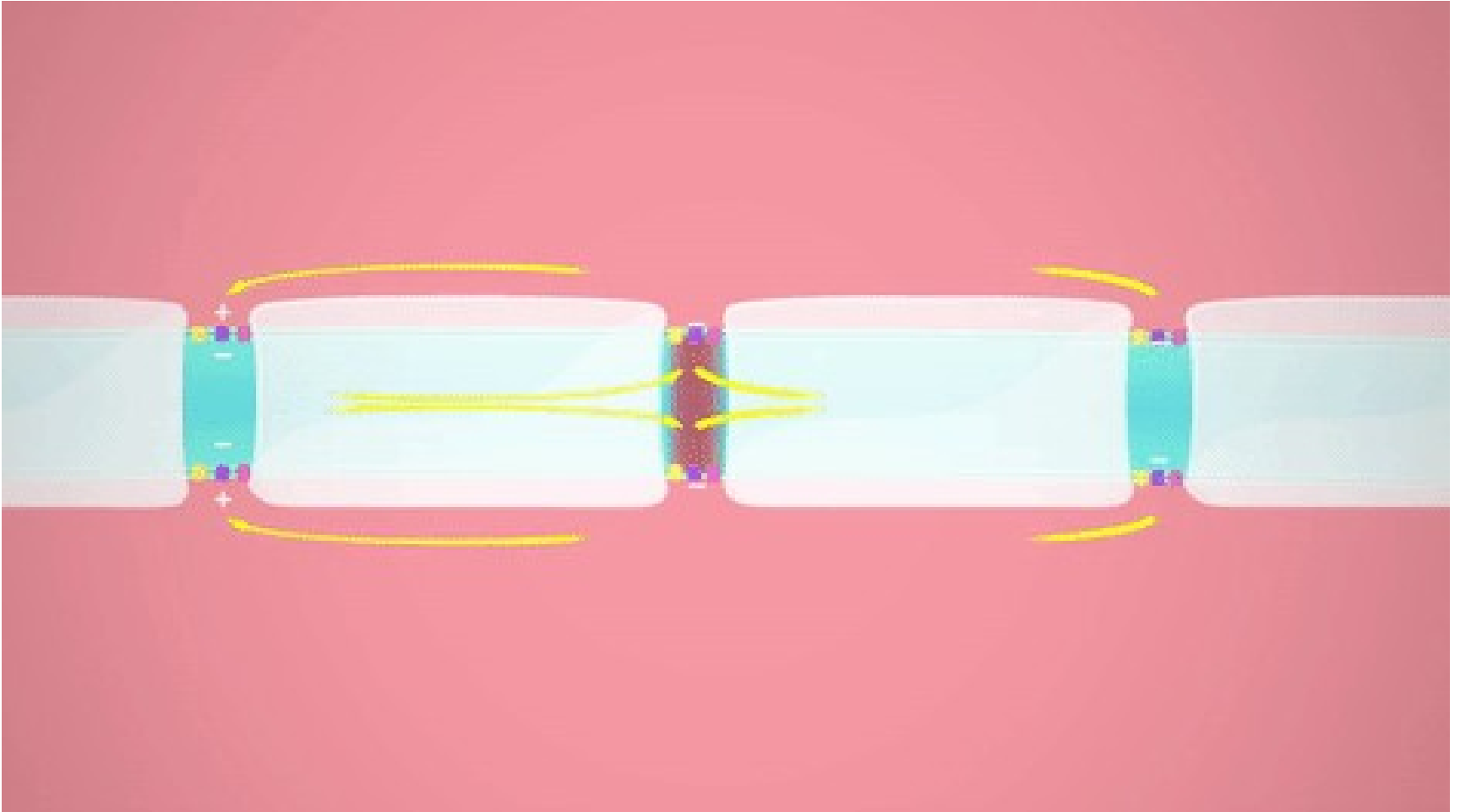
As much as 20–40% of brains' energy expenditure is due to maintenance of membrane potential

**All this ion pumping seems
quite inefficient and slow.**



**Can we do something about
it?**

Myelinated axon



**Now we know how information
comes out.**

How does it come in?

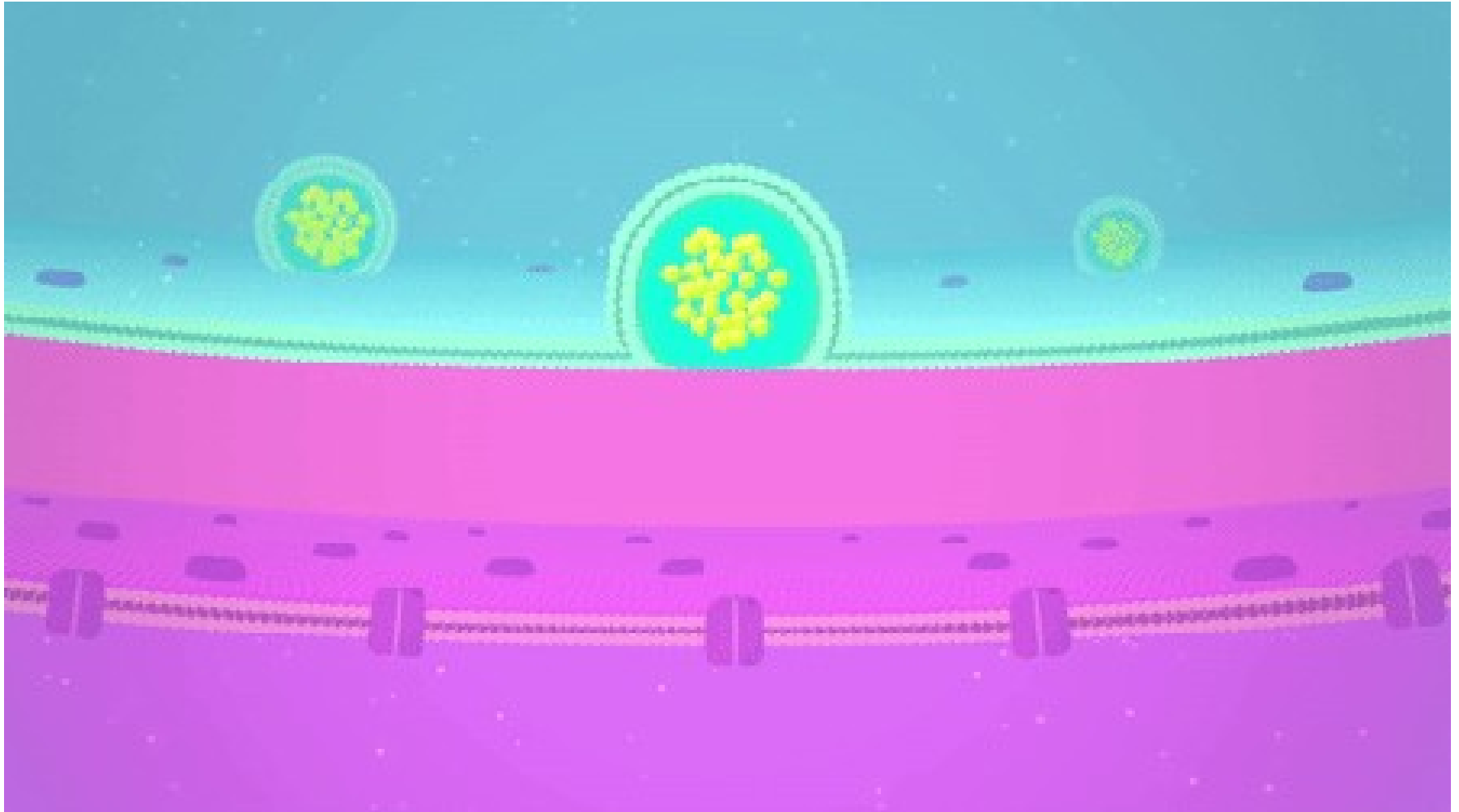
Chemical synapses

- Communication within neurons is electrical
- Communication between neurons is chemical
- **Neurotransmitters** are the molecules that act as chemical signals between neurons

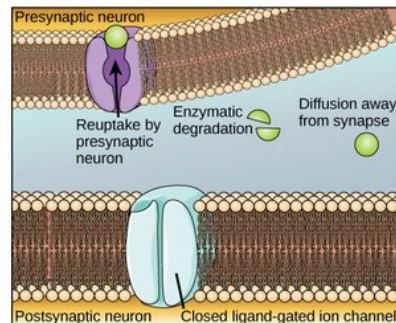
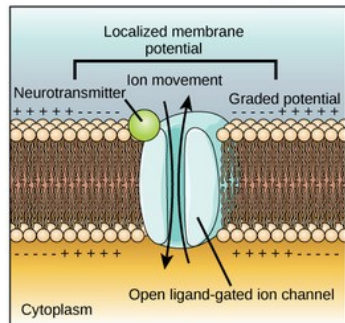
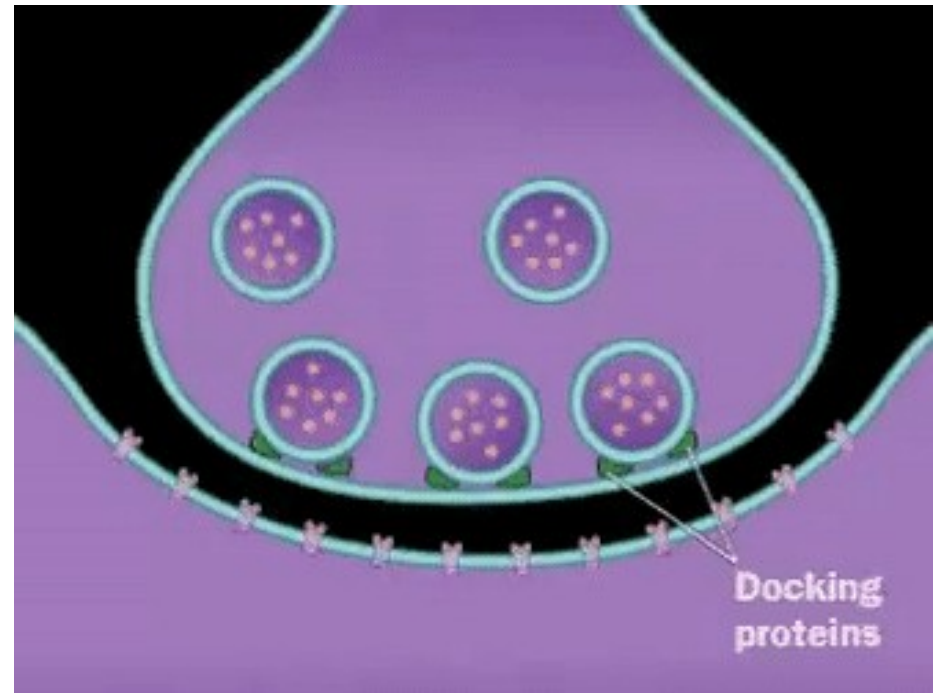
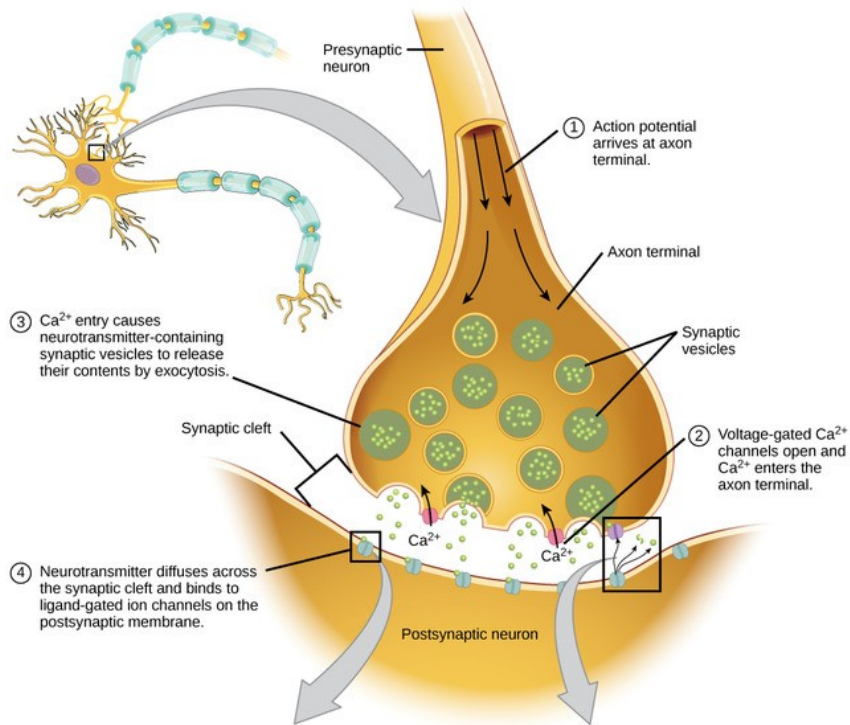
Pre-synaptic neurotransmitter release



Synaptic cleft



Chemical Synapses



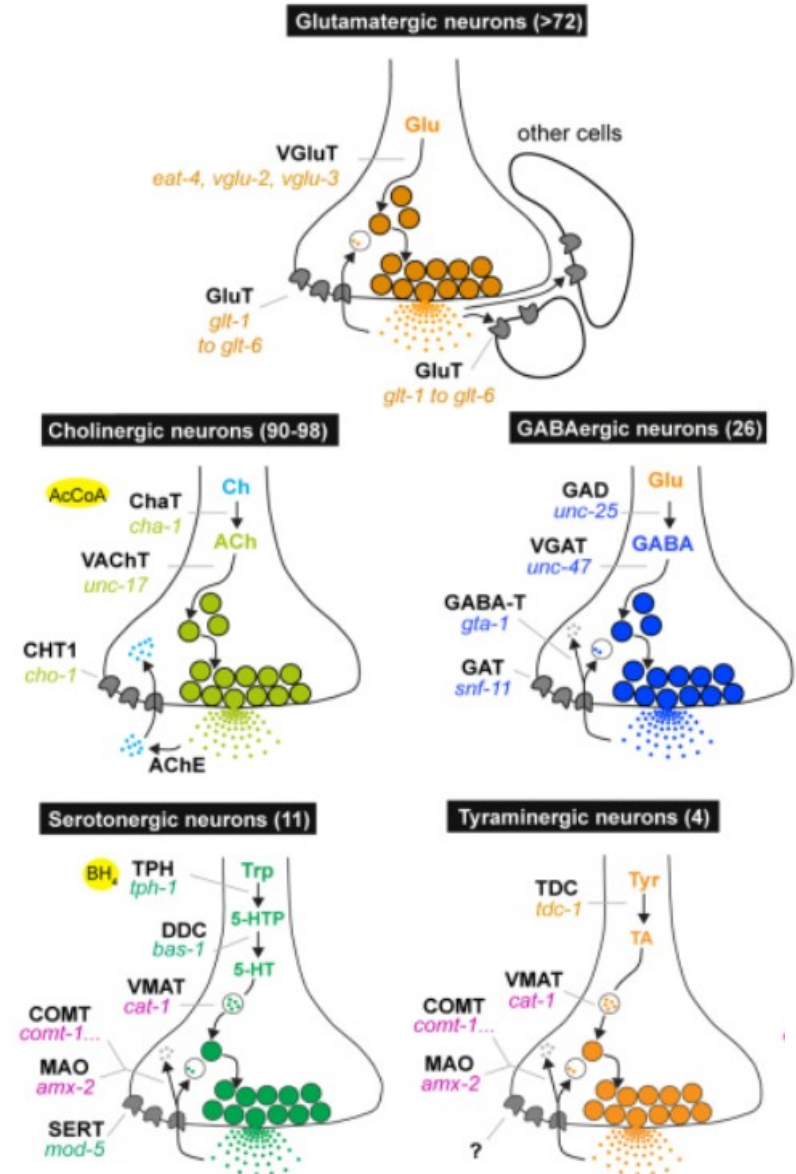
⑤ Binding of neurotransmitter opens ligand-gated ion channels, resulting in graded potentials.

⑥ Reuptake by the presynaptic neuron, enzymatic degradation, and diffusion reduce neurotransmitter levels, terminating the signal.

- Vesicles
- Calcium

Neurotransmitter

- Various types
 - Glutamate
 - GABA
 - Serotonin (5HT)
- Typically one neuron = 1 neurotransmitter
- But neurotransmitters are not exclusive to receptors

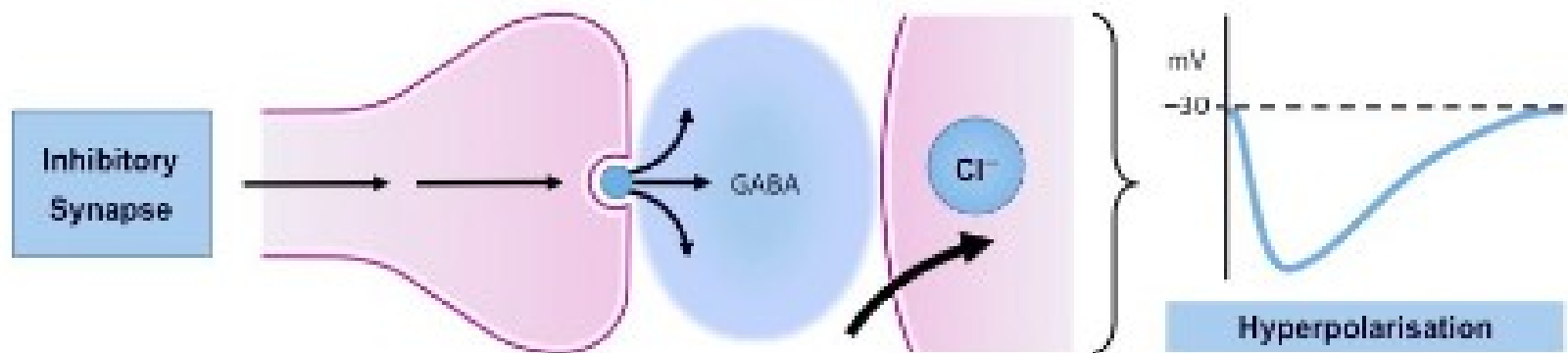
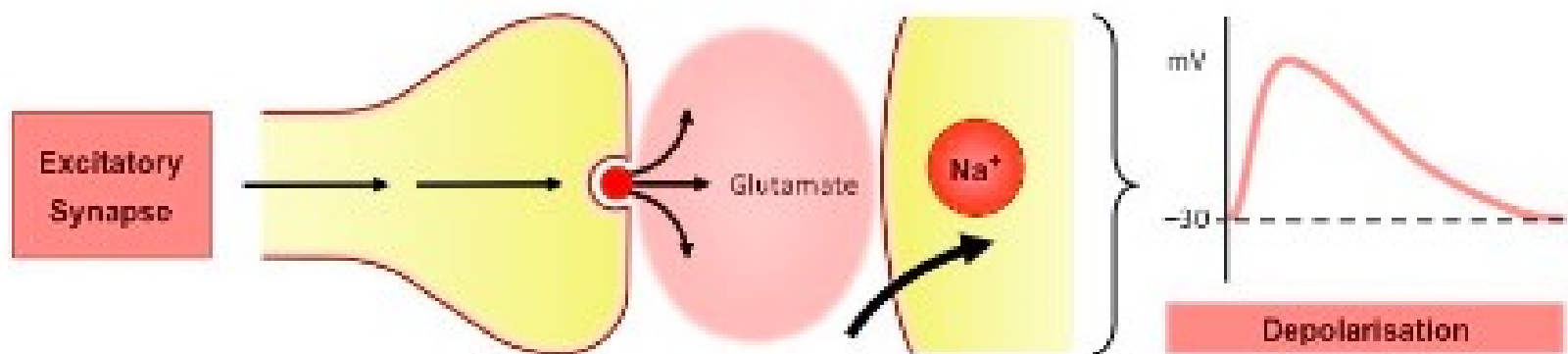


**In ANNs we have positive
and negative connections.**

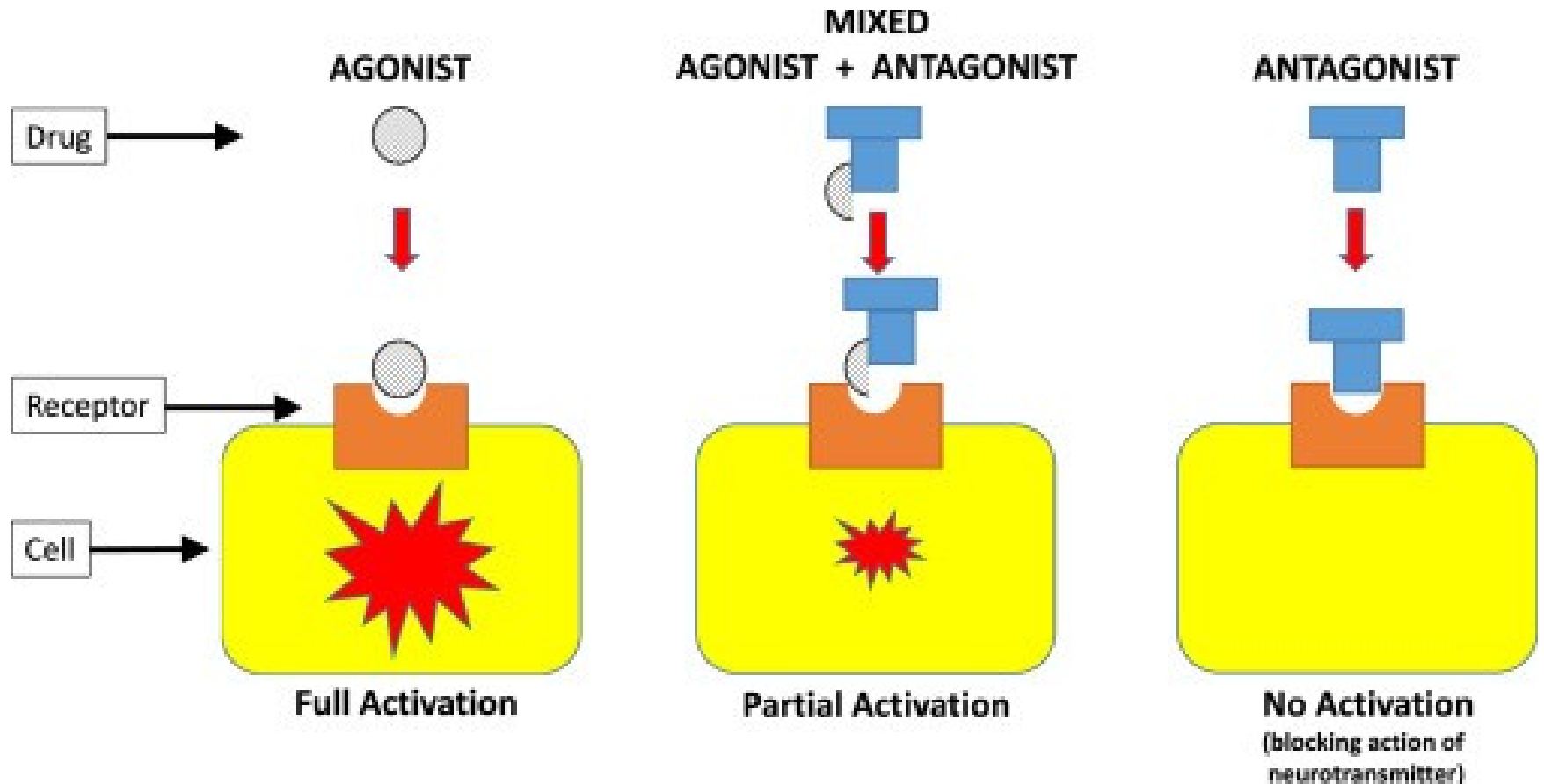


**How would you design
them in the brain?**

EPSP vs. IPSP



Antagonists vs. agonists





**Do you know of real-world
examples where
agonists/antagonists play a
role?**

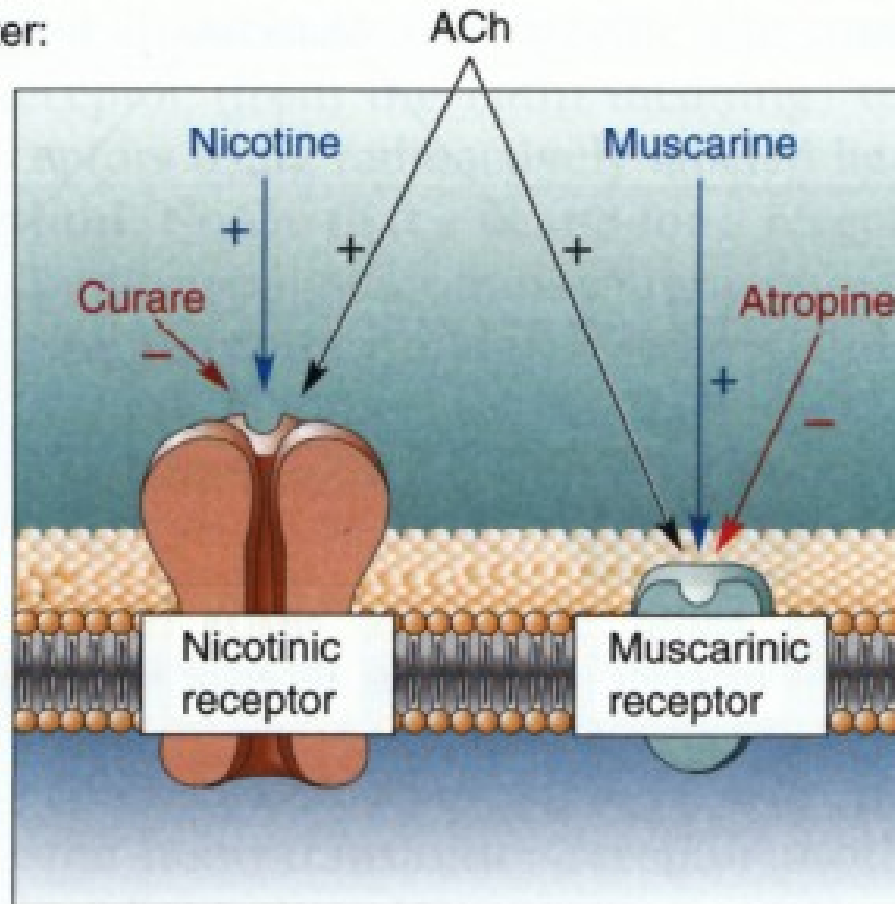
Antagonists vs. agonists

Neurotransmitter:

Agonists:

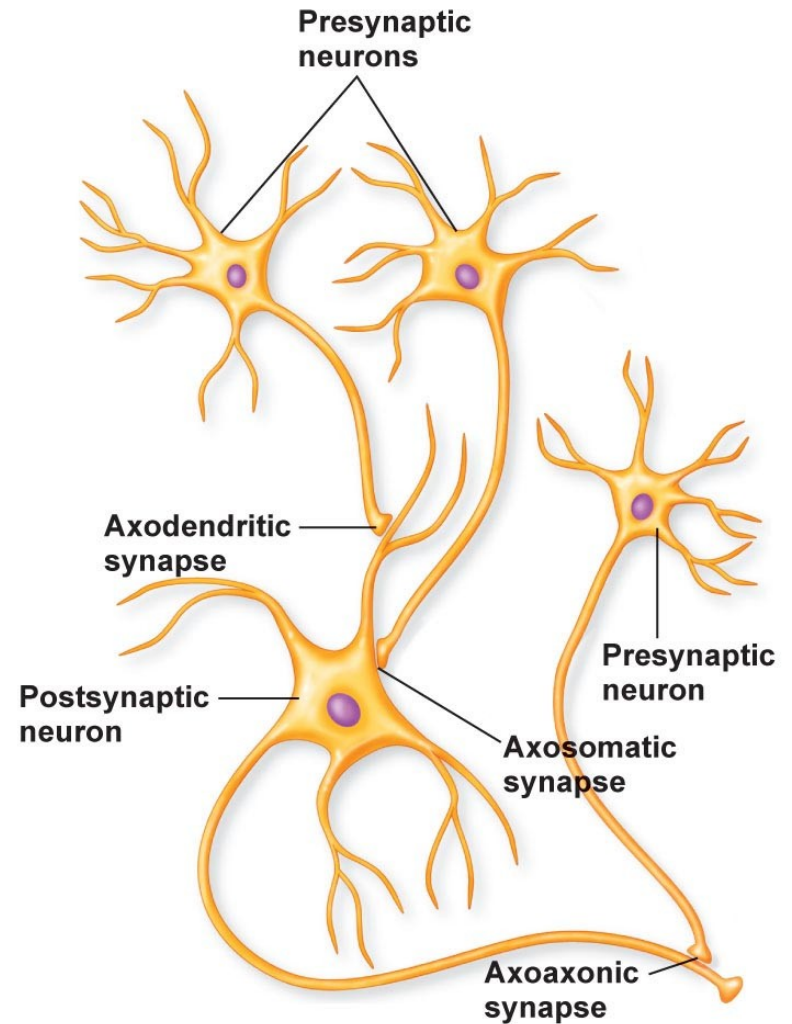
Antagonists:

Receptors:



Types of connections

- Axo-dendritic
- Axo-axonal
- Axo-somatic



The diagram illustrates the integration of excitatory and inhibitory inputs at the axon hillock. It is divided into two parts, (a) and (b).

(a) Excitatory synapse (active), Inhibitory synapse (inactive): This part shows a neuron with an active excitatory synapse on the dendrite and an inactive inhibitory synapse on the soma. The EPSP at the dendrite is large, and the Vm of the soma shows a small depolarization. The axon hillock is shown with a small depolarization, indicating that the excitatory input is sufficient to reach the threshold for an action potential.

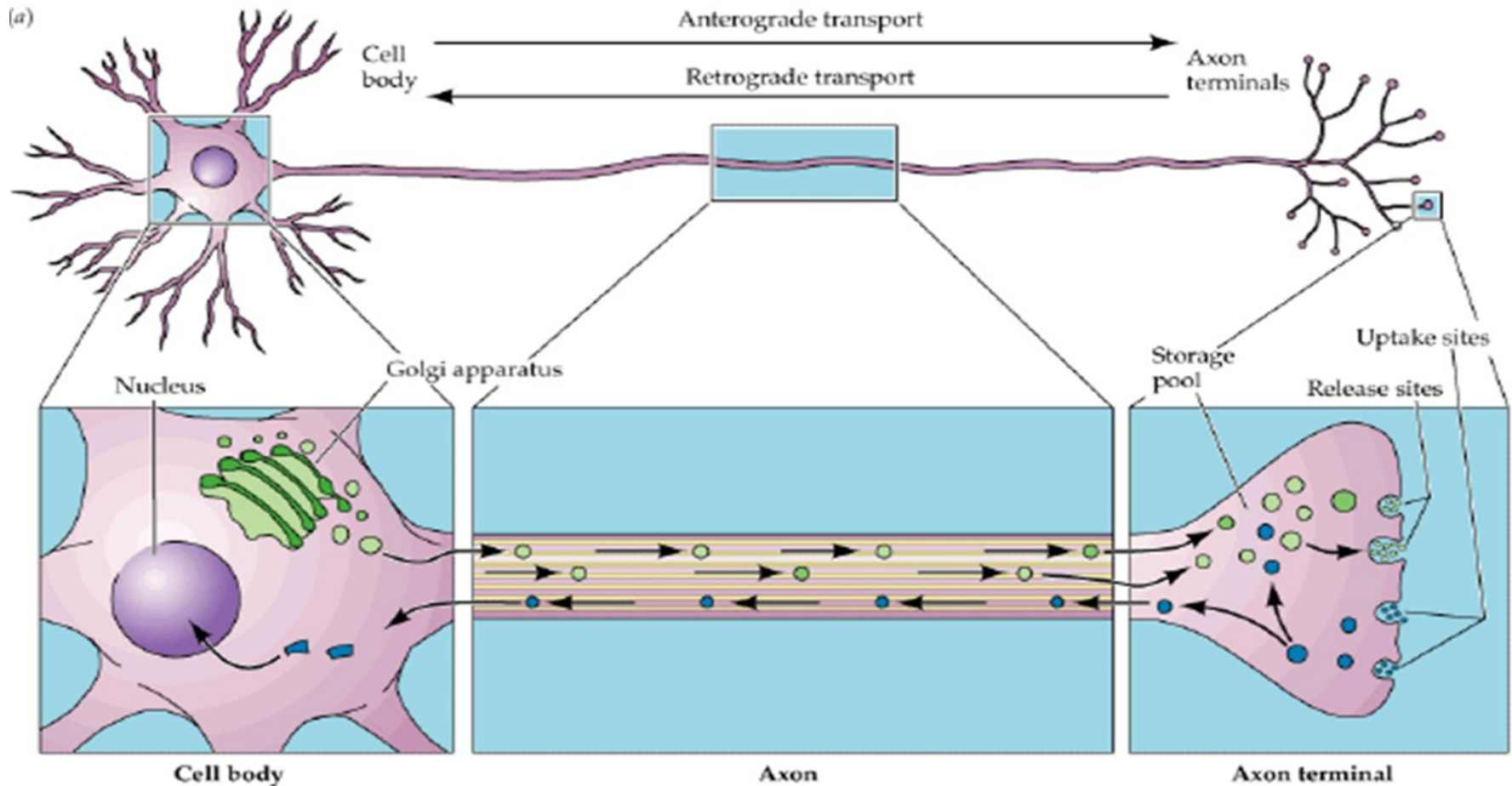
(b) Excitatory synapse (active), Inhibitory synapse (active): This part shows a neuron with both an active excitatory synapse on the dendrite and an active inhibitory synapse on the soma. The EPSP at the dendrite is large, but the Vm of the soma shows a small hyperpolarization due to the inhibitory input. The axon hillock is shown with a small hyperpolarization, indicating that the inhibitory input is sufficient to prevent an action potential.

Shunting inhibition



**Implications for Artificial
Neural Networks?**

Axonal transport



Slow: 1-10mm/day

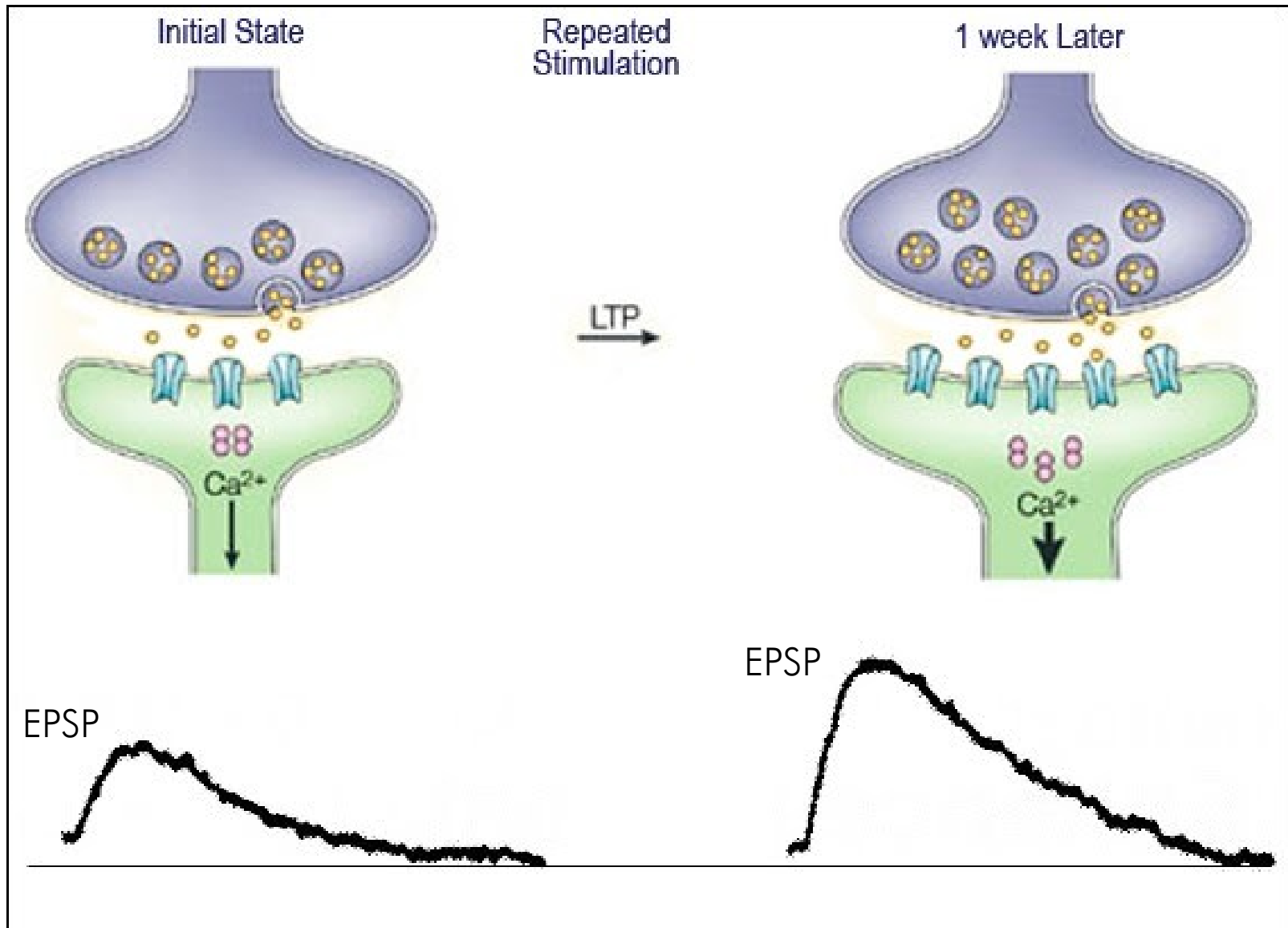
Fast: ~ 1000 mm/day

**The most important
property of ANNs is that
they learn.**

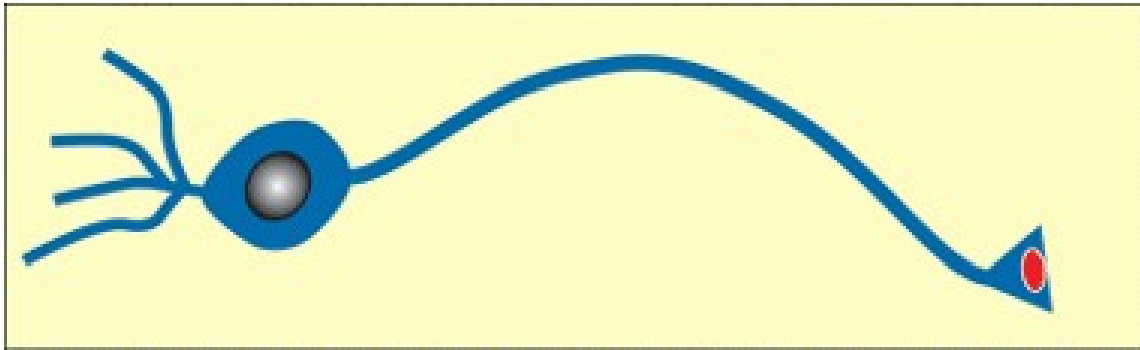


**How would you design that in a
biological neuron?**

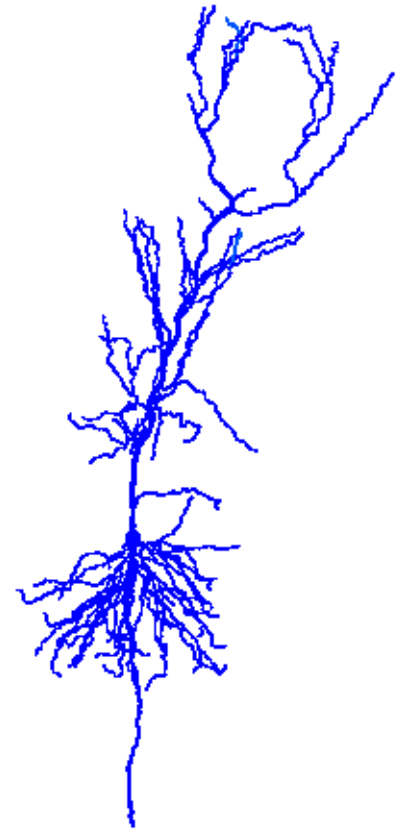
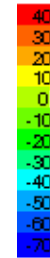
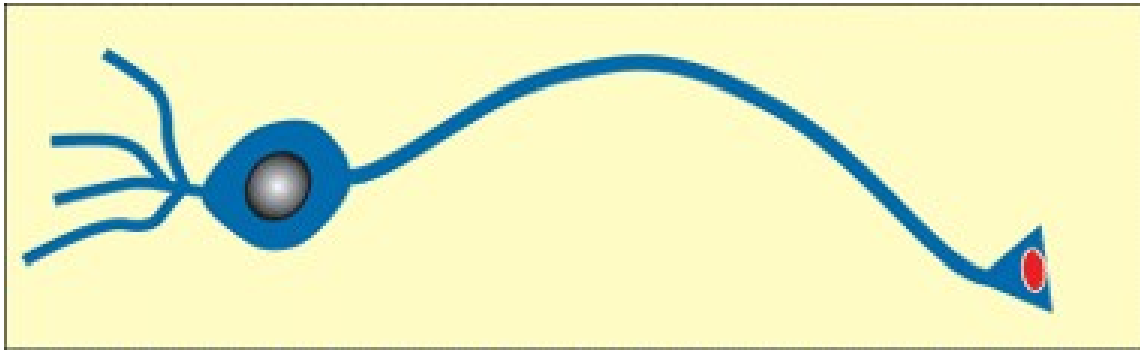
Synapses and learning



Signal integration in a neuron



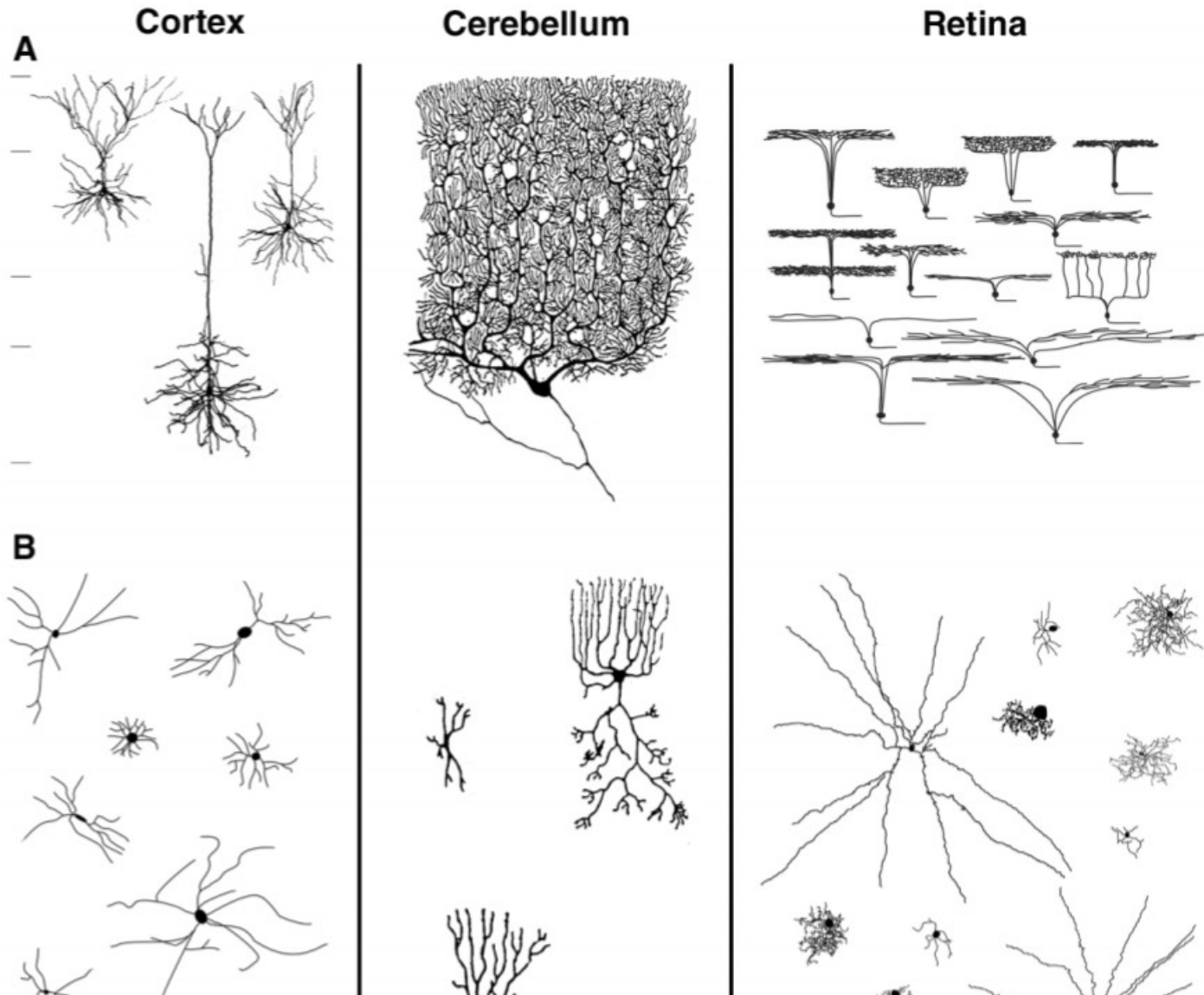
Signal integration in a neuron



Basic parameters

- 10 – 100 μm
- Cerebellum $\sim 10^9$ granule cells
- Cat visual cortex, 1 mm^3
 - $\sim 50\,000$ neurons
 - each $\sim 6\,000$ synapses
 - $\sim 84\%$ Type I, $\sim 16\%$ Type II
- Human (extrapolation from cat)
 - Surface of one hemisphere $\sim 100\,000\text{ mm}^2$
 - $\sim 10^9$ neurons
 - $\sim 60 \cdot 10^{12}$ synapses

Many types of cells

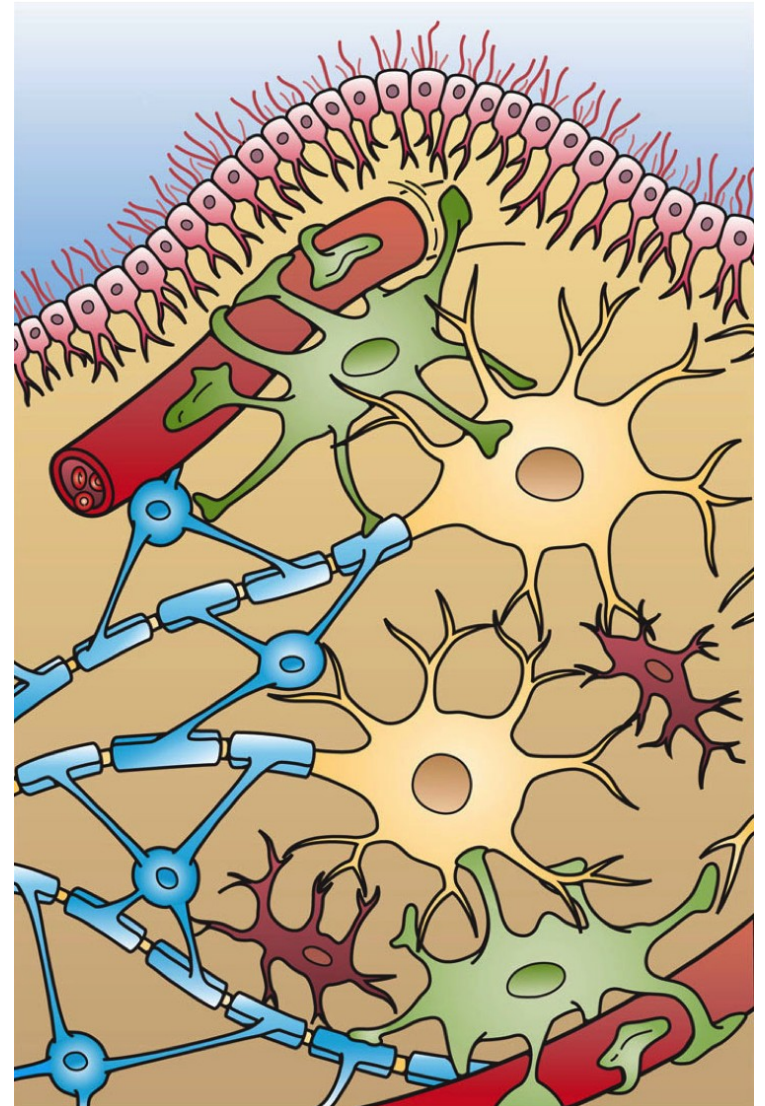
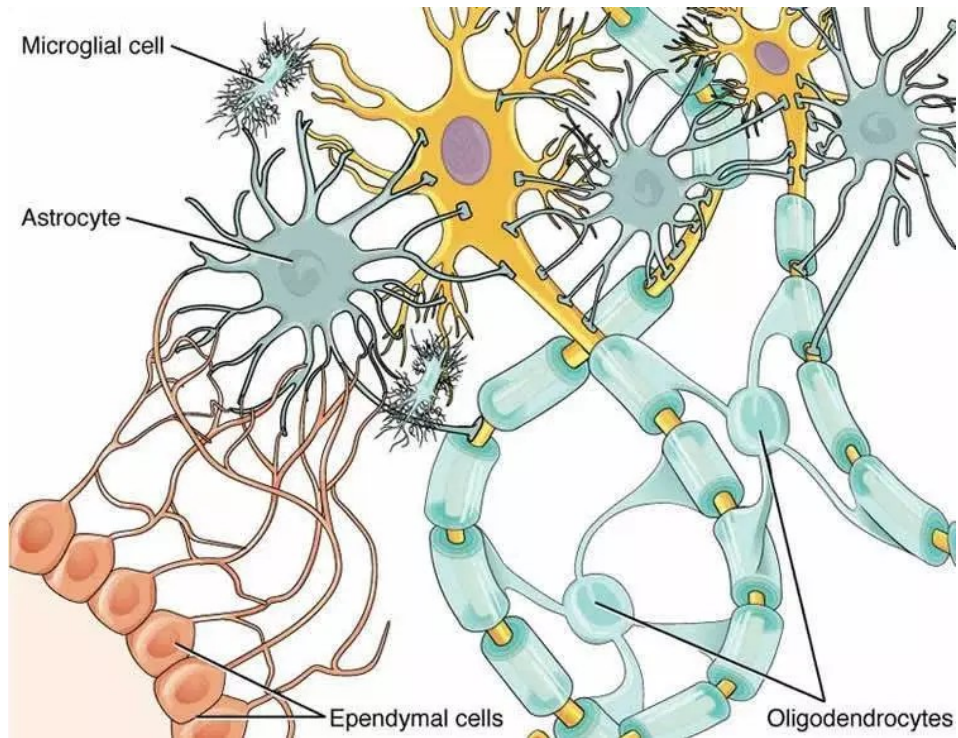


Caveats

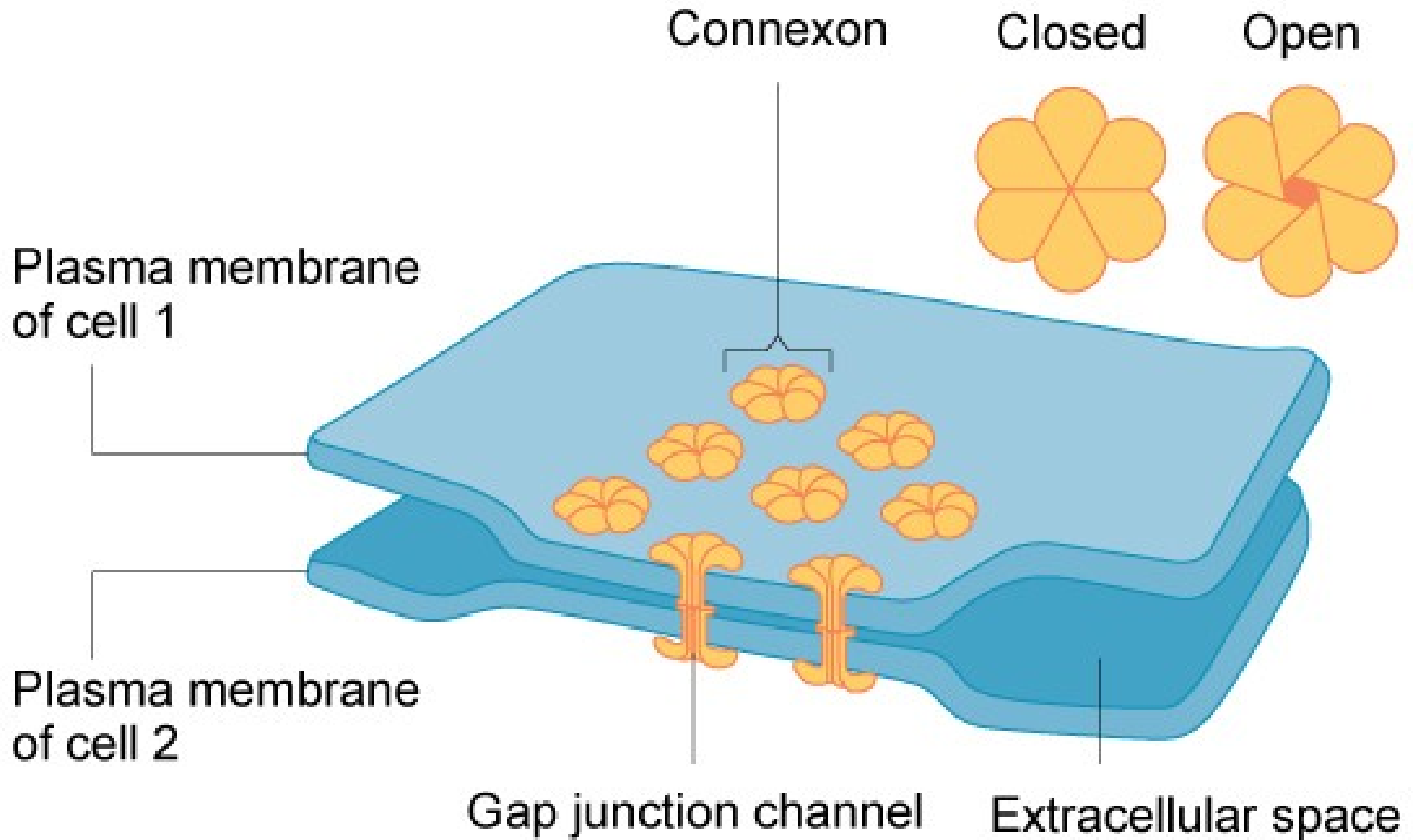
Other cells in CNS

- Neurons: 10%
- Glia: 90%
 - astrocytes: cca 80%
 - ependymal cells: cca 5%
 - oligodendrocytes: cca 5%
 - mikroglia: cca 10%
- Glia as a support system for neural substrate

Glia



Gap junctions



Gap junctions

- Present throughout neural system
- Hypothesized in formation of neural rhythms
- Weak neural-to-glial coupling via gap junctions
- Astrocytes and Oligodendrocytes coupled via gap junctions

Other

- Probabilistic nature of vesicle release – failure to initiate PSP
- Threshold is not fixed
- Dendritic integration