

Advanced Electrophysiology

Lessons 1-2

26 February/ 5 March 2026

Course presentation

Equivalent electrical circuit of a neuron



Synaptic neurophysiology (Prof. Cingolani)

Research Strand: Neurosciences (Biomedicine)



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Biosketch

After graduating in Molecular Biology at the University of Pisa (Italy), I obtained a Ph.D. fellowship to work at the Max-Planck Institute for Experimental Medicine, Göttingen (Germany). Here, I specialized in

- Research Areas
- Biomedicine
- Environmental Biology
- Psychology
- Research Centres
- Alpine Center for Botanical Studies in Pura Pass
- Research Outcomes
- Large International Projects
- Software and databases
- Monthly summary of Q1 publications
- Top cited articles
- ArTS research archive
- Publications of the Department
- Conferences

Building Q - rooms 216 & 309



Lorenzo A. Cingolani



◀ **Postdoc**
University College,
London
Synaptic physiology



◀ **Ph.D., MPI, Göttingen**
Electrophysiology

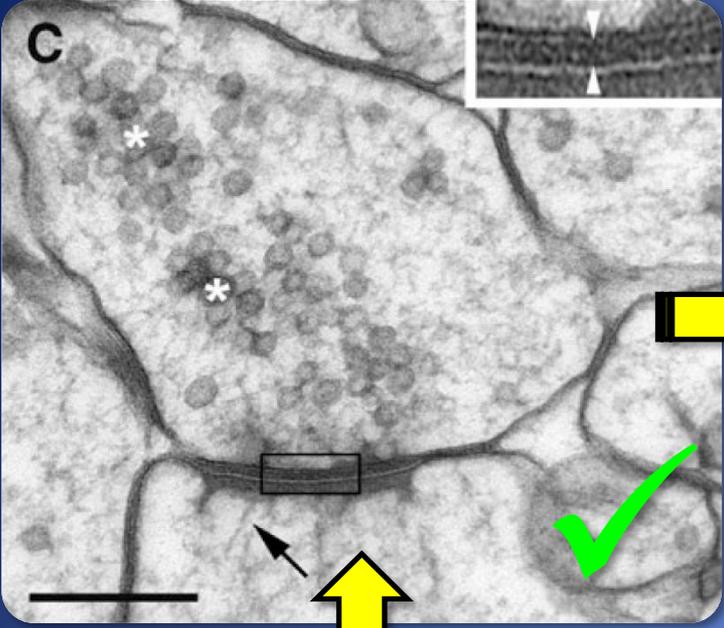


University of Pisa
Molecular Biology

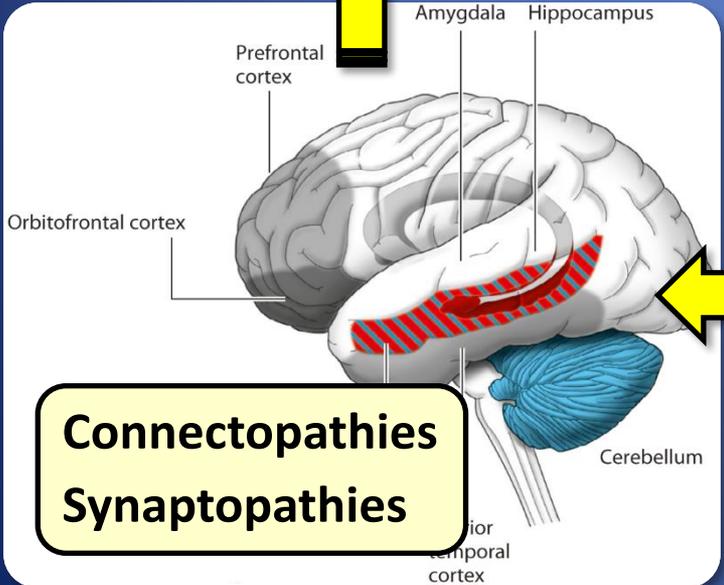


Genoa
IIT

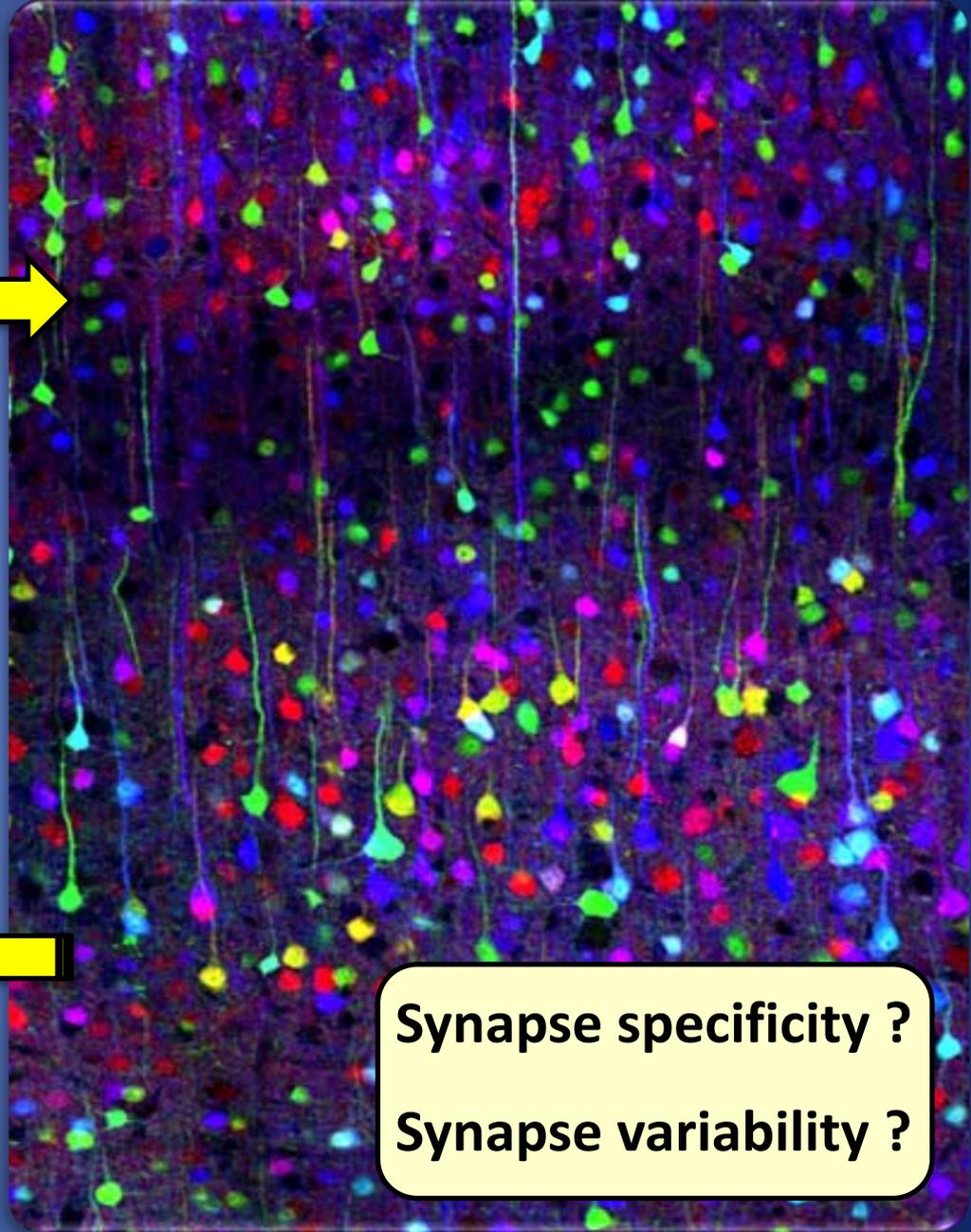
Rostaing et al., 2006



Wang et al., 2014



Connectopathies
Synaptopathies

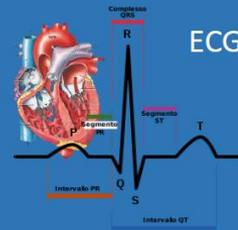
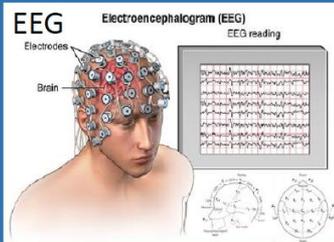


Synapse specificity ?
Synapse variability ?

What is electrophysiology?

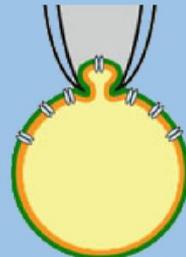
Electrophysiological techniques

ELECTROPHYSIOLOGY



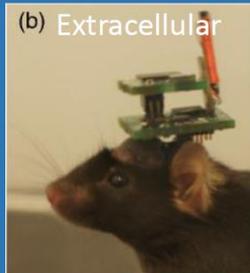
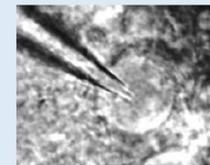
Single cell electrophysiology

Patch-clamp

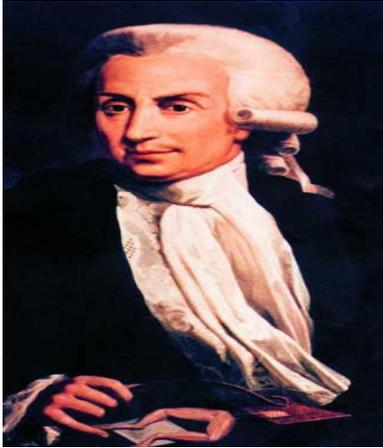


e.g.: cell-attached

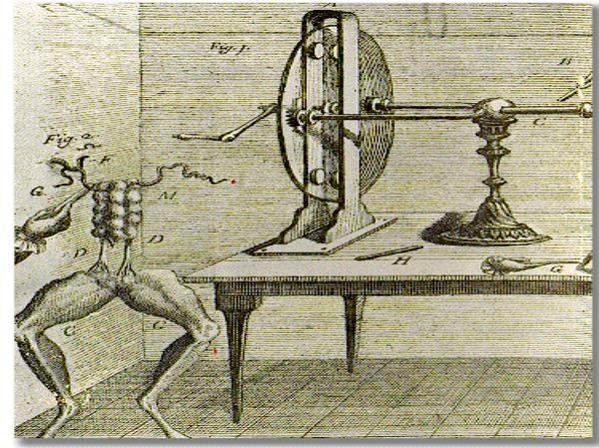
Whole-cell



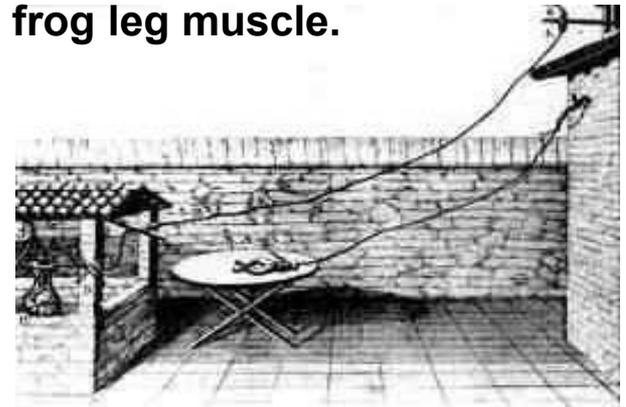
The origins of bioelectricity



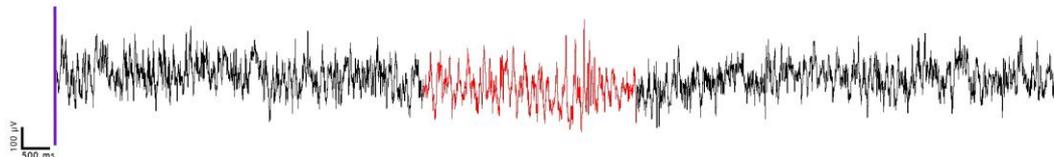
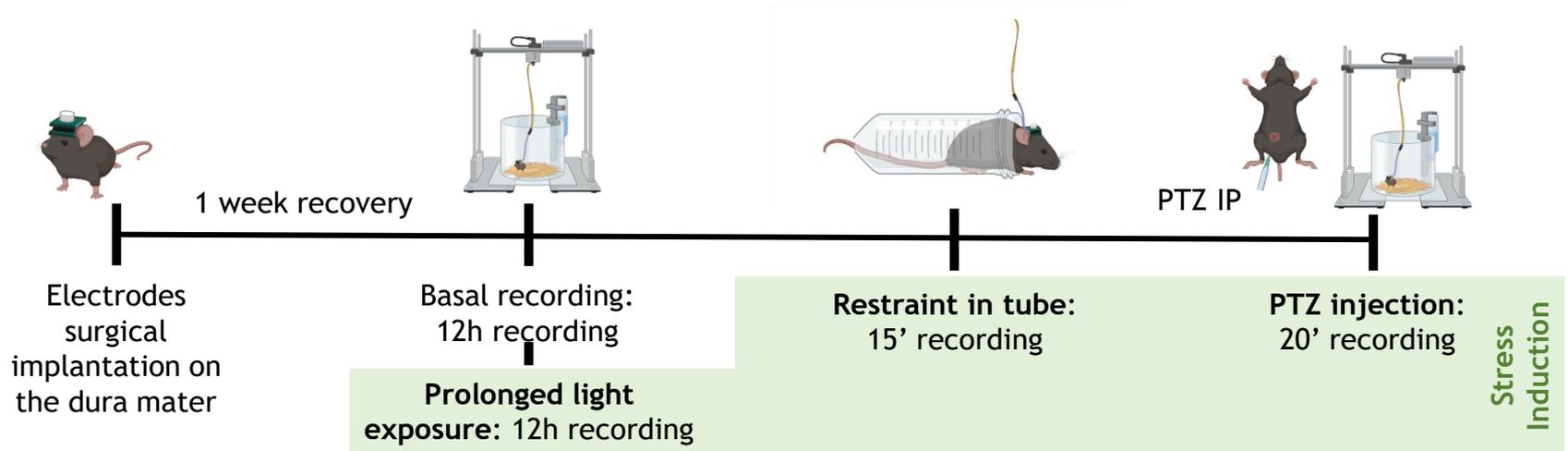
Luigi Galvani (1737-1798), working at the University of Bologna, discovered that nerve conduction and muscle contraction are electrical phenomena. Galvani published his results in 1791 in "De Viribus Electricitatis in Motu Muscolari Commentarius".



Galvani, demonstrated that the application of an electric current caused the contraction of the frog leg muscle.



Electroencephalographic (EEG) recordings in freely moving mice



Course goals

- **Understand the basic biophysics of synapses, neurons and neuronal networks, and the principles and potential of electrophysiological techniques**
- **Become accustomed to performing electrophysiological experiment (patch-clamp and MEA recordings)**
- **Become proficient at data analysis using dedicated software**

Structure of the course

- **Face-to-face course**
- **Work at the computer**
- **Laboratory**

- **Lessons will be recorded and available on MT**
- **Materials (presentations, papers, articles, book chapters) loaded on Moodle before lessons (password: electro)**

Structure of the exam

1. A homework assignment, consisting in the analysis, figure preparation and interpretation of a set of electrophysiological data.

Weight in the final score: **60%**

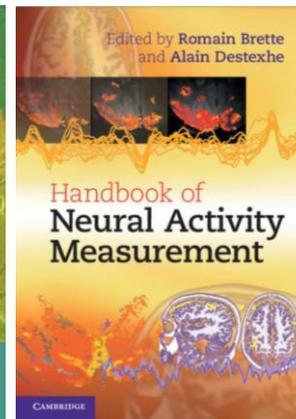
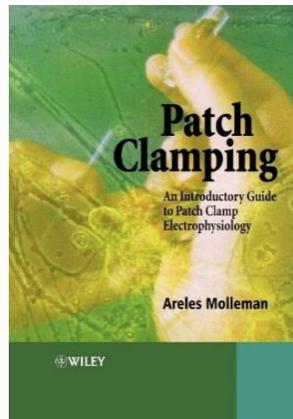
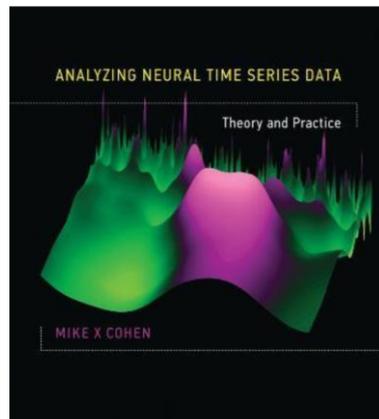
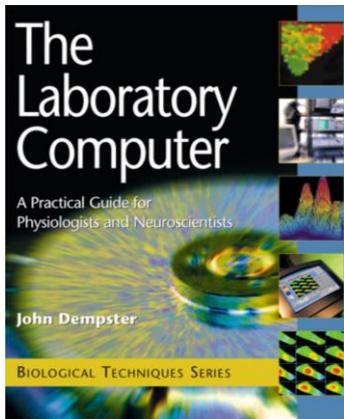
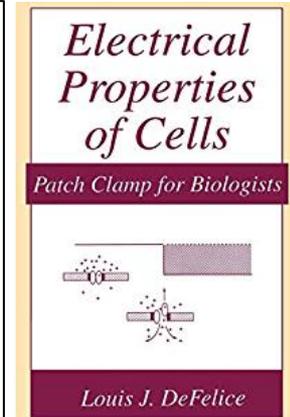
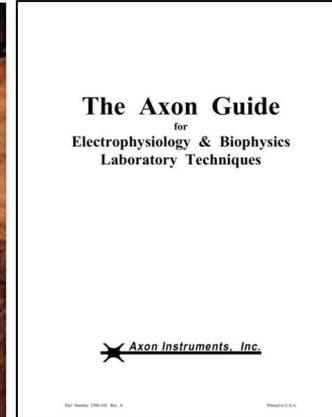
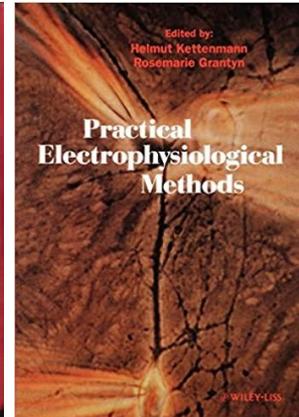
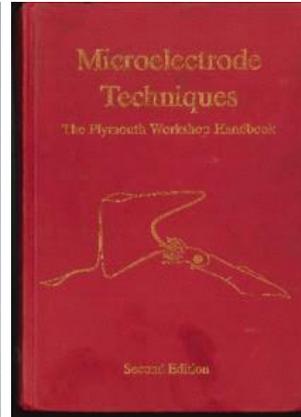
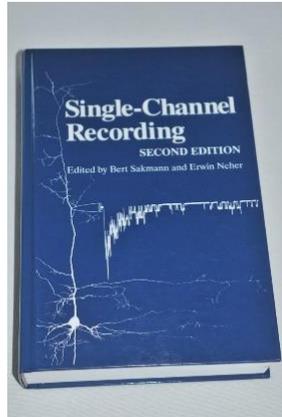
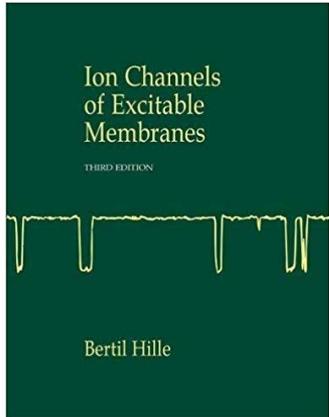
2. Practicum at the electrophysiological setups. Students will be judged for their propensity for experimental work.

Weight in the final score: **10%**

3. Questionnaire on the Moodle platform. 10 multiple choice questions. Each question has only one correct answer, and each correct answer is awarded 1 point.

Weight in the final score: **30%**

Textbooks



... and others

Calendar

- **Th 26 February (14:00-16:00):** Frontal lesson (ex-Cla)
- **Th 26 February (16:00-18:00):** Data analysis project (ex-Cla)
- **F 27 February (14:00-18:00):** Acute brain slice preparation (Room 309, Building Q)
- **Th 5 March (14:00-16:00):** Frontal lesson (ex-Cla)
- **Th 5 March (16:00-18:00):** Data analysis project (ex-Cla; Bring a laptop!)
- **F 6 March (14:00-18:00):** Patch-clamp (group 1; Room 309, Building Q)
- **Th 12 March (14:00-16:00):** Frontal lesson (ex-Cla)
- **Th 12 March (16:00-18:00):** Data analysis project (ex-Cla; Bring a laptop!)
- **F 13 March (14:00-18:00):** Patch-clamp (group 2; Room 309, Building Q)
- **F 20 March (14:00-18:00):** Patch-clamp (group 3; Room 309, Building Q)

Bioelectricity

Learning objective:

To understand the equivalent electrical circuit of a neuron

The Axon Guide
for
Electrophysiology & Biophysics
Laboratory Techniques

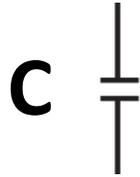
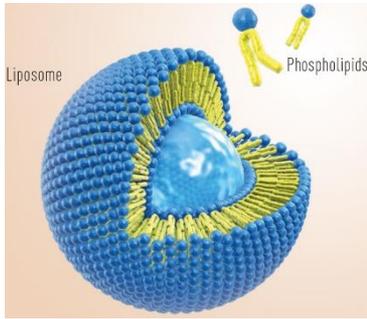
Chapter 1

 **Axon Instruments, Inc.**

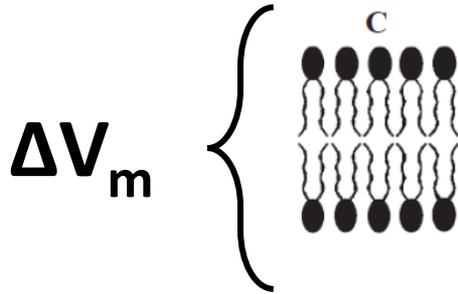
Foolish assumptions:

We know what the membrane potential (ΔV_m) and Nernst equation are

Principles of electrical circuits - Capacitance



$$C = \frac{\Delta Q}{\Delta V_m}$$



$$\Delta Q_{\text{total}} = \Delta Q_1 + \Delta Q_2 = C_1 \Delta V_m + C_2 \Delta V_m = \Delta V_m (C_1 + C_2)$$

$$C_{\text{total}} = C_1 + C_2$$

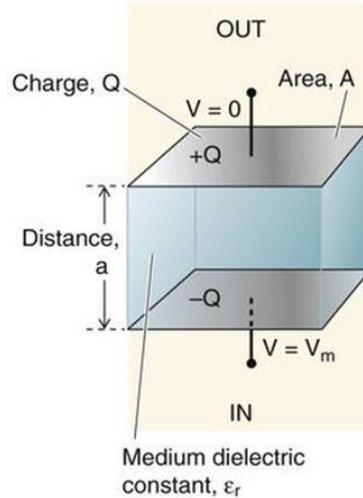
$$C_{\text{total}} = \sum_i^n C_i$$

ΔQ is charge imbalance (Coulombs, C)
C is capacitance (Farads, F)

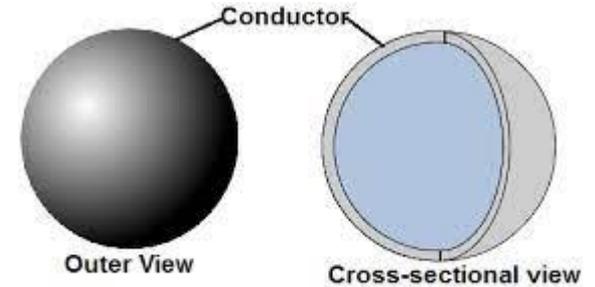
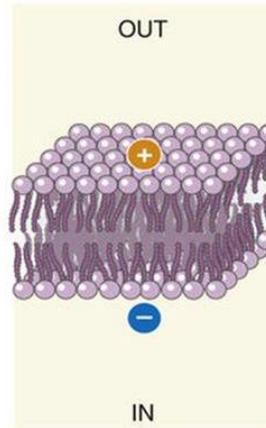
Principles of electrical circuits - Capacitance

$$C = \epsilon_m \frac{A}{d}$$

C PARALLEL-PLATE CAPACITOR



LIPID MEMBRANE



Spherical Capacitor

$$d_m = 10 \text{ nm} = 10^{-8} \text{ m}$$

$$\epsilon_m = 10^2 \text{ pF / m} \quad (\epsilon_0 = 8.85 \text{ pF / m})$$

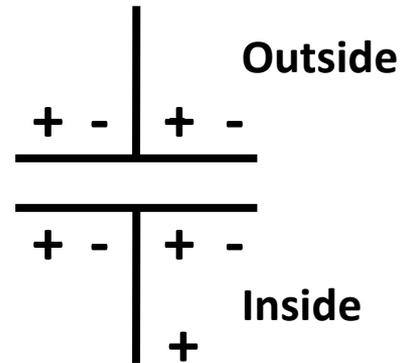
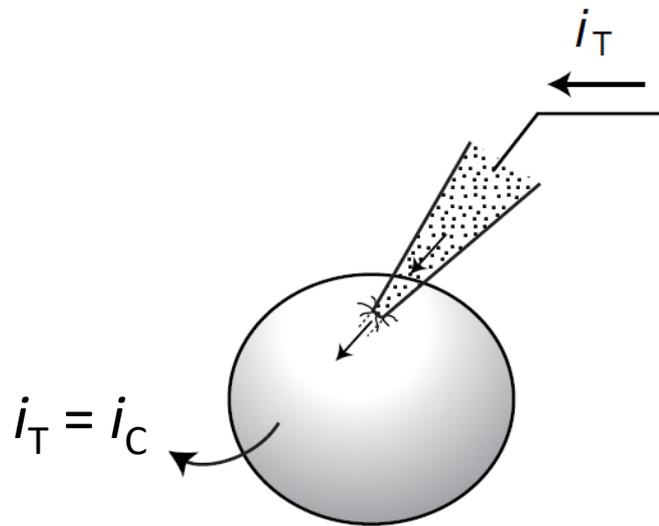
$$C_m / A = \epsilon_m / d = \text{Specific } C_m = 10^{10} \text{ pF / m}^2$$

C_m is proportional to A_m

Specific $C_m = 1 \mu\text{F / cm}^2$

ϵ is dielectric constant of the insulator (Farad / m)
 A is area of the capacitor (m^2)
 d is distance between conducting plates (m)

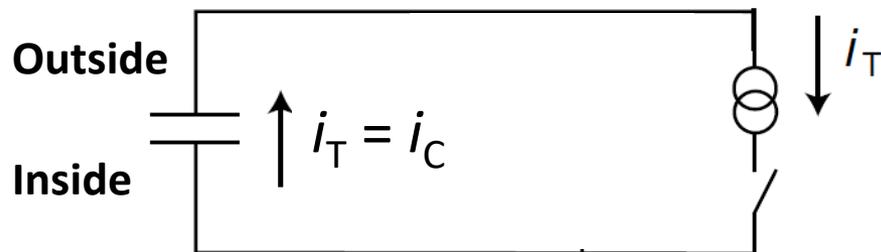
Principles of electrical circuits - Capacitive current



$$\Delta Q = C \Delta V_m$$

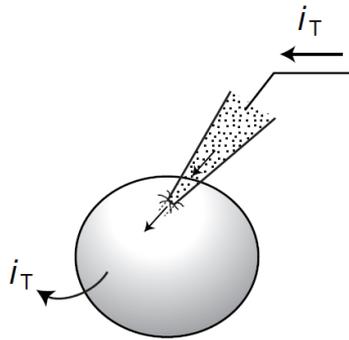
$$i_T = i_C = \frac{dQ}{dt} = C \frac{dV_m}{dt}$$

i_C is C times the time rate of change (d/dt) of V_m



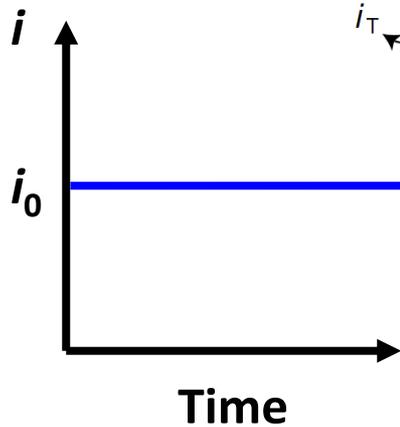
What happens when we inject current into this simple neuron?

Principles of electrical circuits - Capacitive current



$$i_T(t) = C \frac{dV_m}{dt}$$

We can integrate this differential equation over time starting with initial voltage V_0 at time 0



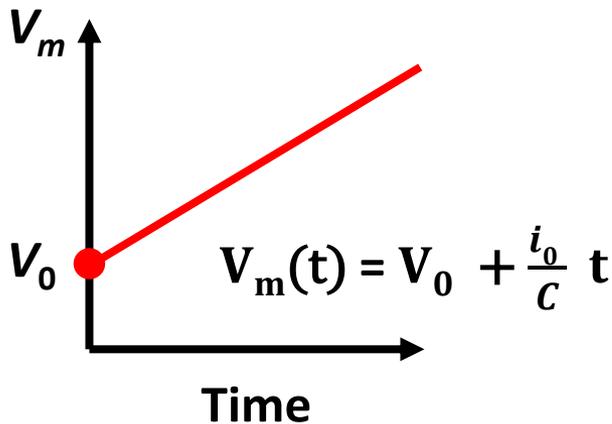
$$dV_m = \frac{1}{C} i_T(t) dt$$

$$\int_{V_0}^{V_t} dV_m = \frac{1}{C} \int_0^t i_T(t) dt$$

$$V_m(t) = V_0 + \frac{1}{C} \int_0^t i_T dt$$

ΔQ

$$V_m(t) = V_0 + \frac{\Delta Q}{C}$$



V_m is the initial V_m at time 0 plus the integral over time of the injected current ($=\Delta Q$) divided by C

Principles of electrical circuits - Resistance

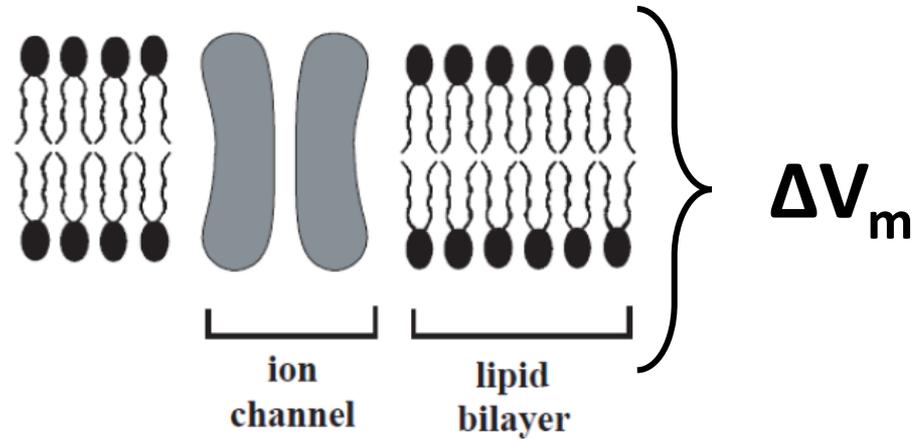
Ohm's (first) law

$$i = \frac{\Delta V_m}{R}$$

$$i = G \Delta V_m$$



$$R = \frac{1}{G}$$



$$i_1 = \frac{\Delta V_m}{R_1}$$

$$i_2 = \frac{\Delta V_m}{R_2}$$

$$i_{\text{total}} = \frac{\Delta V_m}{R_{\text{total}}} = \Delta V_m \left(\frac{1}{R_1} + \frac{1}{R_2} \right)$$

$$\frac{1}{R_{\text{total}}} = \frac{1}{R_1} + \frac{1}{R_2}$$

$$\frac{1}{R_{\text{total}}} = \sum_i^n \frac{1}{R_i}$$

ΔV is voltage (Volts, V)

i is current (Amperes, A)

R is resistance (Ohms, Ω)

G is conductance (Siemens, S)

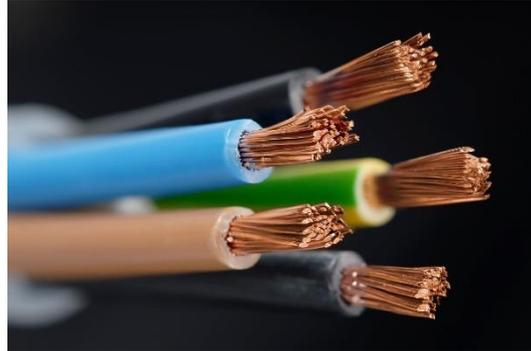
$$G_{\text{total}} = G_1 + G_2$$

$$G_{\text{total}} = \sum_i^n G_i$$

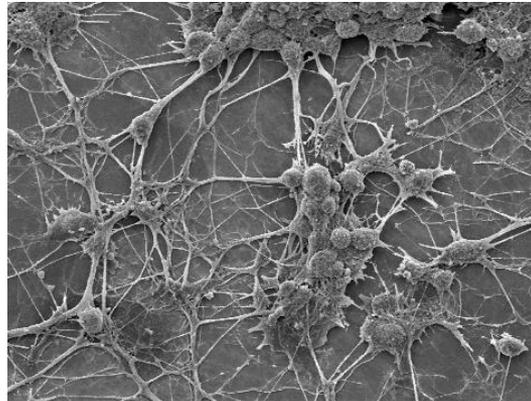
Principles of electrical circuits - Resistivity

Ohm's second law

$$R = \rho \frac{L}{A}$$



$$\rho_{\text{copper}} = 1.7 \mu\Omega \text{ cm}$$



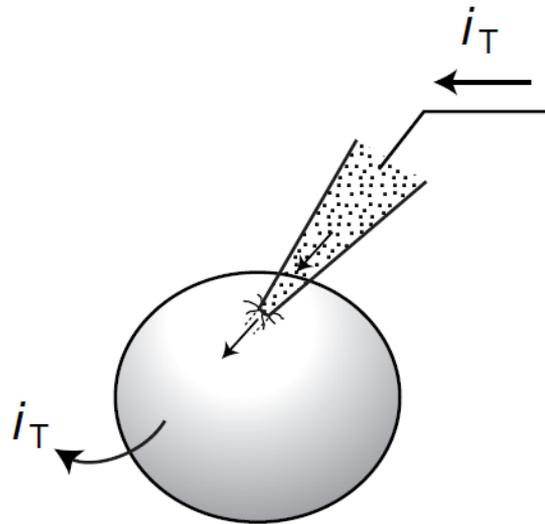
$$\rho_{\text{CSF}} = 60 \Omega \text{ cm}$$

(More than 10 million higher)

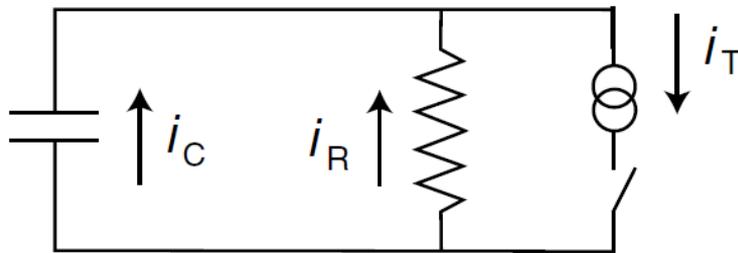
The brain is made of lousy conductors → You need huge ΔV to produce tiny currents ($i = \Delta V/R$), that's why you need special gimmicks like action potentials to send electrical signals

ρ is resistivity of the material ($\Omega \text{ m}$)
L is length of the conductor (m)
A section of the conductor (m^2)

The equivalent electrical circuit of a neuron



Outside



Inside

$$i_C = C \frac{dV_m}{dt}$$

$$i_R = \frac{V_m}{R}$$

$$i_T = \frac{V_m}{R} + C \frac{dV_m}{dt}$$

(first order linear differential equation)

$$Ri_T = V_m + RC \frac{dV_m}{dt}$$

What is the steady-state solution of this equation?

$$\text{Set } \frac{dV_m}{dt} = 0$$

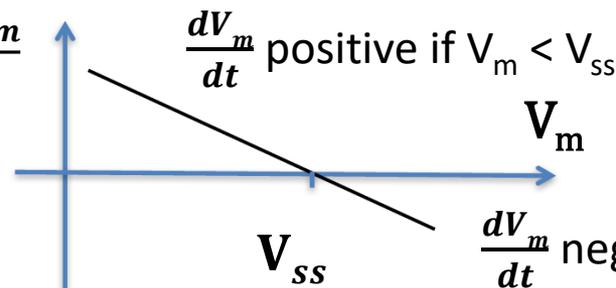
$$V_{ss} = Ri_T$$

$$V_{ss} = V_m + \tau \frac{dV_m}{dt}$$

where $\tau = RC$

$$\frac{dV_m}{dt} = -\frac{1}{\tau} (V_m - V_{ss})$$

$\frac{dV_m}{dt}$

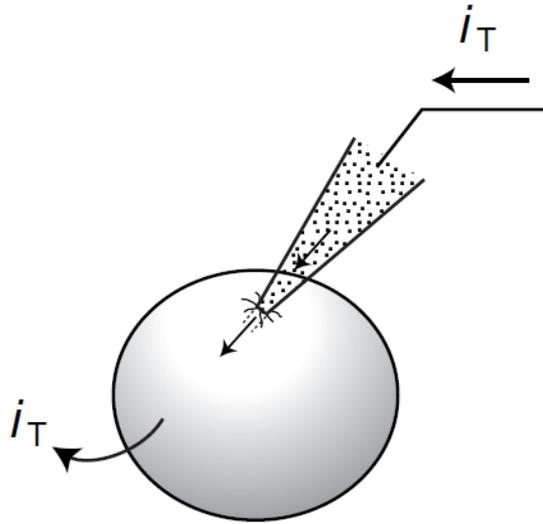


V_{ss}

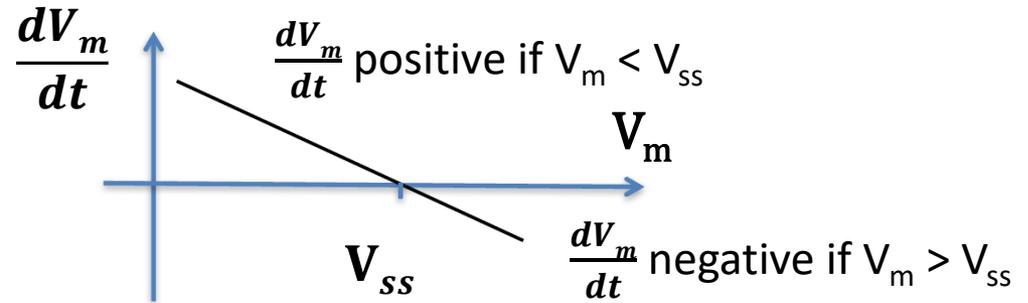
$\frac{dV_m}{dt}$ negative if $V_m > V_{ss}$

The equivalent electrical circuit of a neuron

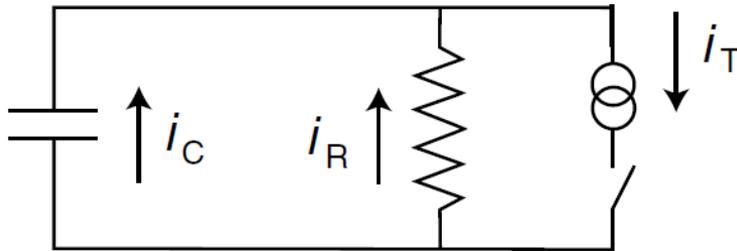
$$\frac{dV_m}{dt} = -\frac{1}{\tau} (V_m - V_{ss})$$



Outside



- V_m is always approaching (relaxing towards) V_{ss}
- V_m approaches V_{ss} at a rate ($\frac{dV_m}{dt}$) proportional to how far away it is from V_{ss}

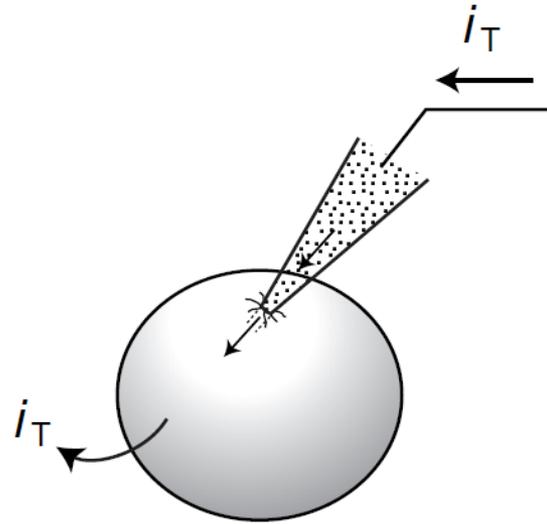


$$i_C = C \frac{dV_m}{dt}$$

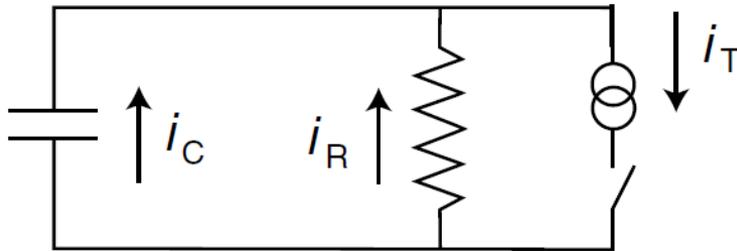
Inside

$$i_R = \frac{V_m}{R}$$

The equivalent electrical circuit of a neuron



Outside



Inside

$$i_C = C \frac{dV_m}{dt}$$

$$i_R = \frac{V_m}{R}$$

$$\frac{dV_m}{dt} = -\frac{1}{\tau} (V_m - V_{ss})$$

$$\frac{dV_m}{V_m - V_{ss}} = -\frac{dt}{\tau}$$

$$\int_{V_0}^{V_t} \frac{dV_m}{V_m - V_{ss}} = -\frac{1}{\tau} \int_0^t dt$$

$$\int \frac{dx}{x+c} = \ln(x+c)$$

$$\ln \frac{V_m - V_{ss}}{V_0 - V_{ss}} = -\frac{t}{\tau}$$

$$\frac{V_m - V_{ss}}{V_0 - V_{ss}} = e^{-\frac{t}{\tau}}$$

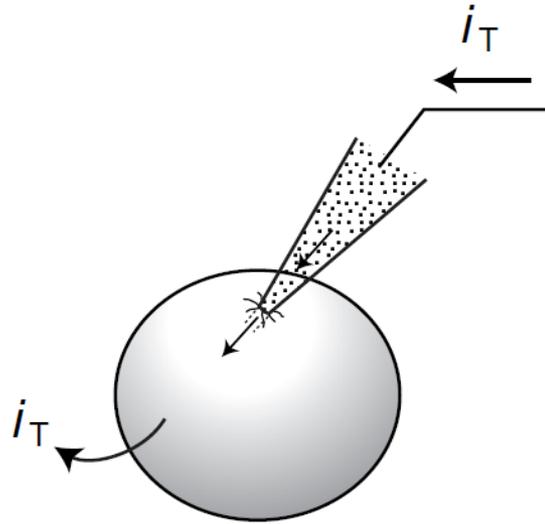
If $V_0 < V_{ss} \rightarrow$ exponential rise
 If $V_0 > V_{ss} \rightarrow$ exponential decay

$$V_m - V_{ss} = (V_0 - V_{ss}) e^{-\frac{t}{\tau}}$$

General solution
 for constant
 current

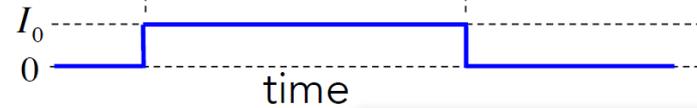
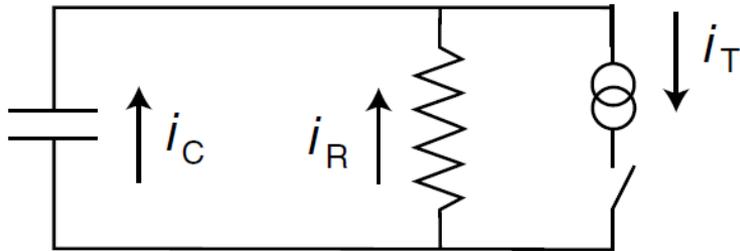
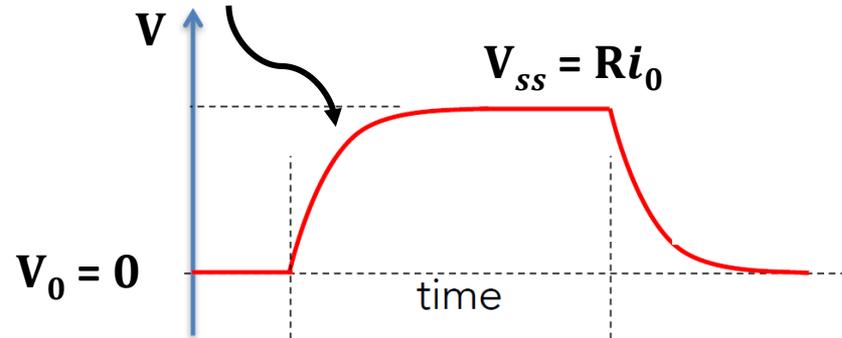
What is relaxing exponentially towards V_{ss} is the difference between V_m and V_{ss}

Response to current injection



Outside

$$V_m = V_{ss} (1 - e^{-\frac{t}{\tau}})$$



$$i_C = C \frac{dV_m}{dt}$$

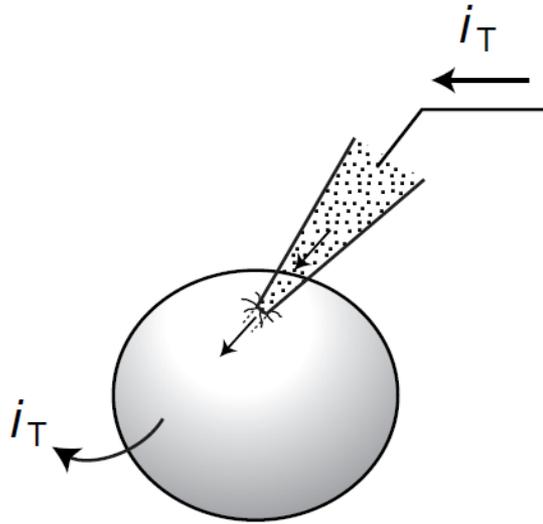
Inside

$$i_R = \frac{V_m}{R}$$

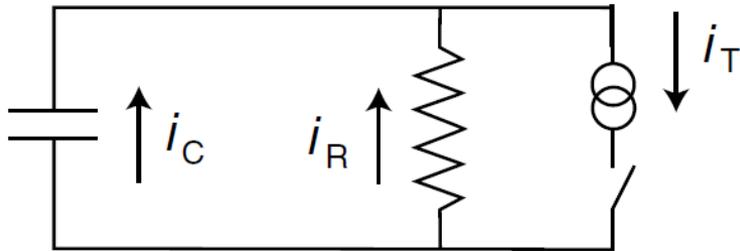
$$V_m - V_{ss} = (V_0 - V_{ss}) e^{-\frac{t}{\tau}}$$

Response to current injection

V_m is always relaxing toward V_{ss} exponentially



Outside

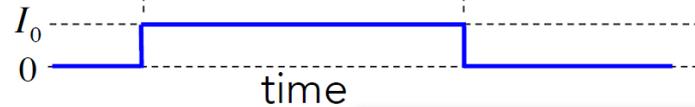
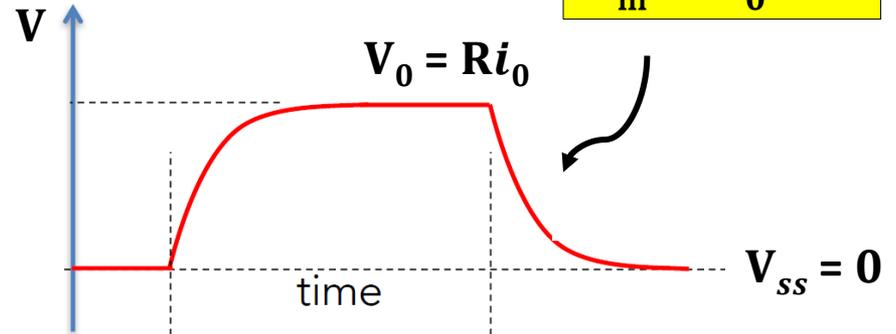


$$i_C = C \frac{dV_m}{dt}$$

Inside

$$i_R = \frac{V_m}{R}$$

$$V_m = V_0 e^{-\frac{t}{\tau}}$$



$$V_m - V_{ss} = (V_0 - V_{ss}) e^{-\frac{t}{\tau}}$$

An RC neuron acts as a low pass filter

(it allows low-frequency signals to pass through while attenuating high-frequency components)

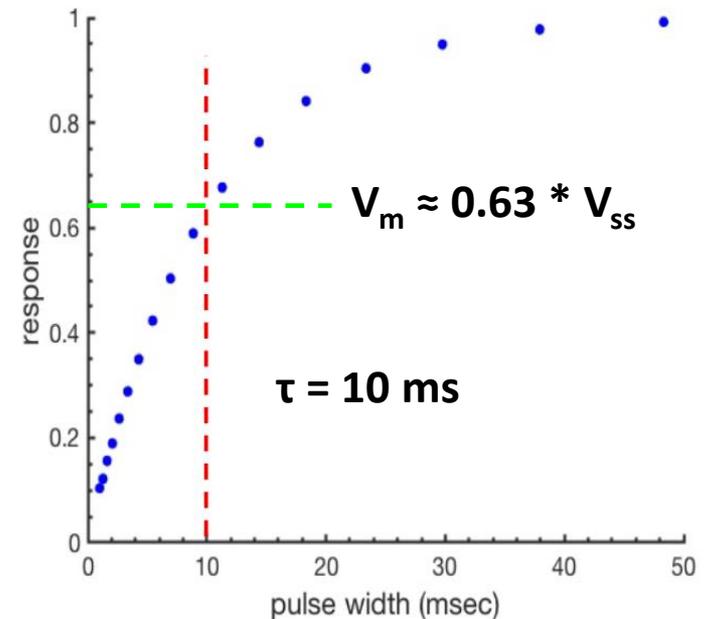
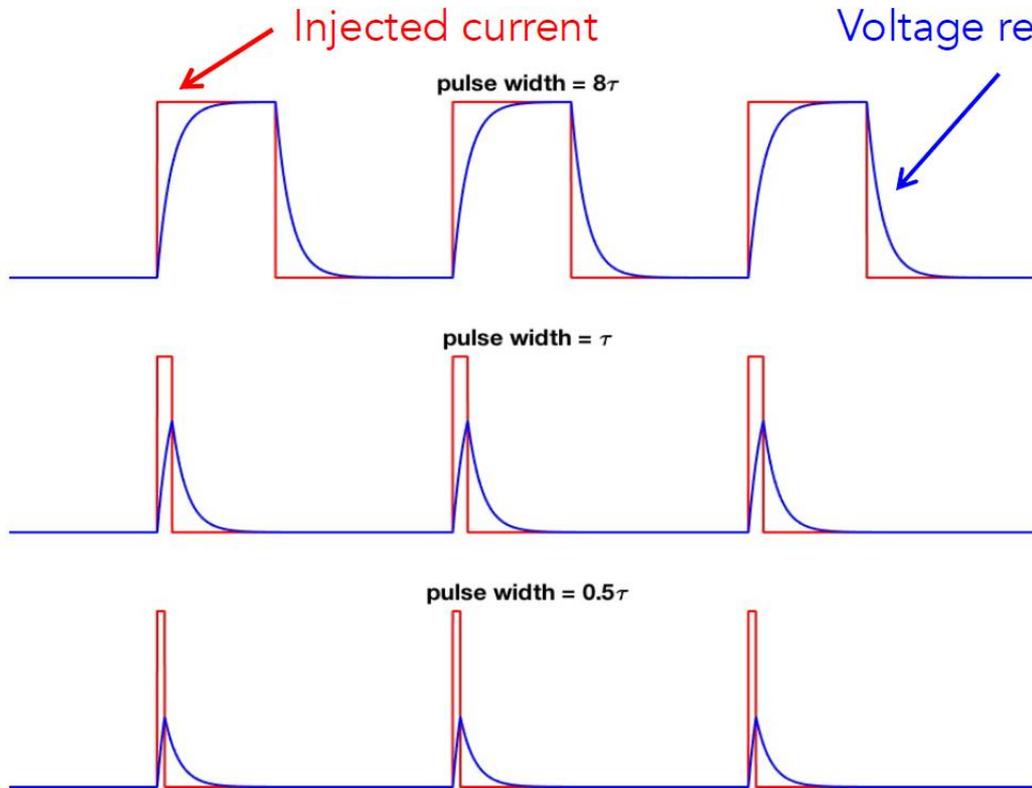
$$V_m = V_{ss} (1 - e^{-\frac{t}{\tau}})$$

$$\tau = RC$$

$$R \approx 10^8 \Omega = 100 \text{ M}\Omega$$

$$C \approx 10^{-10} \text{ F} = 100 \text{ pF}$$

$$\tau = RC \approx 10^{-2} \text{ s} = 10 \text{ ms}$$

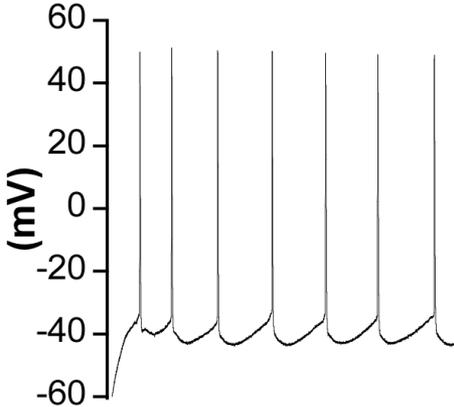


Learning objectives

- 1. To understand what membrane capacitance is from both biological and electrical perspectives**
- 2. To understand what membrane resistance is from both biological and electrical perspectives**
- 3. To understand the equivalent electrical circuit of a neuron**
- 4. To be able to predict the response of a neuron to different current injections**

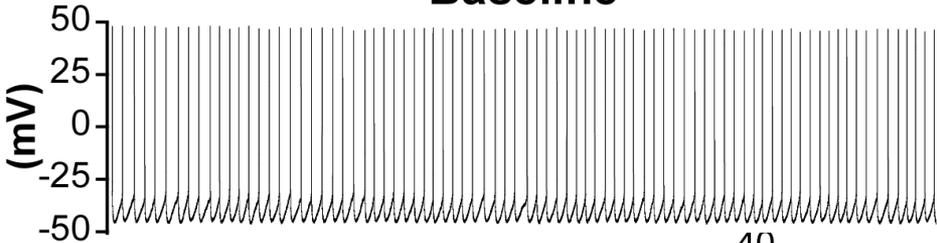
A neuron with active ion channels - action potentials

Baseline

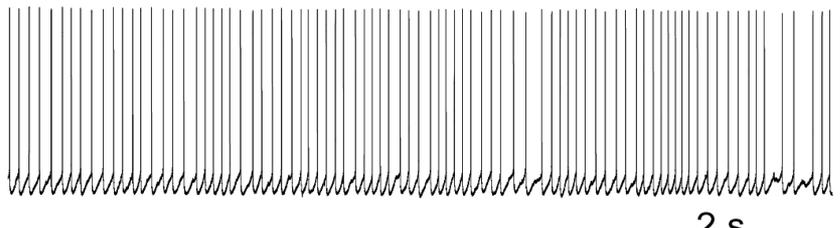


- 1. Frequency**
- 2. Adaptation/acceleration**
- 3. Precision**

Baseline



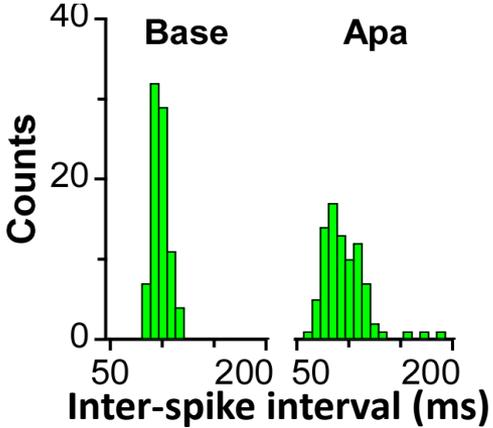
Apamin



2 s

Base

Apa



A neuron with active ion channels beyond action potentials

