

Biological Modeling of Neural Networks

Wulfram Gerstner
EPFL, Lausanne, Switzerland

Week 1: A first simple neuron model/
neurons and mathematics
Week 2: Hodgkin-Huxley models and
biophysical modeling
Week 3: Two-dimensional models and
phase plane analysis
Week 4: Two-dimensional models,
type I and type II models
Week 5,6: Associative Memory,
Hebb rule, Hopfield
Week 7,8: Noise models, noisy neurons
and coding
Week 9: Estimating neuron models for
coding and decoding: GLM
Week 10-13: Networks, cognition, learning
Week x: Online video: Dendrites/Biophysics

LEARNING OUTCOMES

- Solve linear one-dimensional differential equations
- Analyze two-dimensional models in the phase plane
- Develop a simplified model by separation of time scales
- Analyze connected networks in the mean-field limit
- Formulate stochastic models of biological phenomena
- Formalize biological facts into mathematical models
- Prove stability and convergence
- Apply model concepts in simulations
- Predict outcome of dynamics
- Describe neuronal phenomena

Transversal skills

- Plan and carry out activities in a way which makes optimal use of available time and other resources.
- Collect data.
- Write a scientific or technical report.

Look at samples of
past exams

Use a textbook,
(Use video lectures)
don't use slides (only)

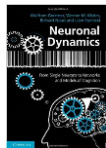
miniproject

Biological Modeling of Neural Networks

Written Exam (2/3)
+ miniproject (1/3)

<http://neurondynamics.epfl.ch/>

Textbook:



Videos (for half the material):

<http://lcn.epfl.ch/~gerstner/NeuronalDynamics-MOOC1.html>

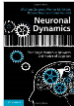
Biological Modeling of Neural Networks



Week 1 – neurons and mathematics: a first simple neuron model

Wulfram Gerstner
EPFL, Lausanne, Switzerland

Reading for week 1:
NEURONAL DYNAMICS
- Ch. 1 (without 1.3.6 and 1.4)
- Ch. 5 (without 5.3.1)



Cambridge Univ. Press

- 1.1 Neurons and Synapses:
 - Overview
- 1.2 The Passive Membrane
 - Linear circuit
 - Dirac delta-function
- 1.3 Leaky Integrate-and-Fire Model
- 1.4 Generalized Integrate-and-Fire Model
- 1.5. Quality of Integrate-and-Fire Models

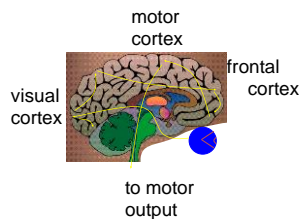
Biological Modeling of Neural Networks



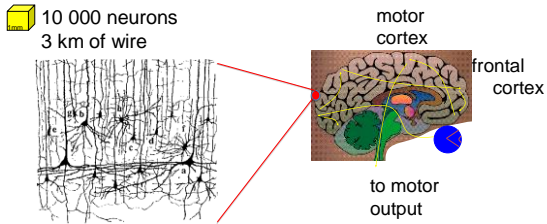
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Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

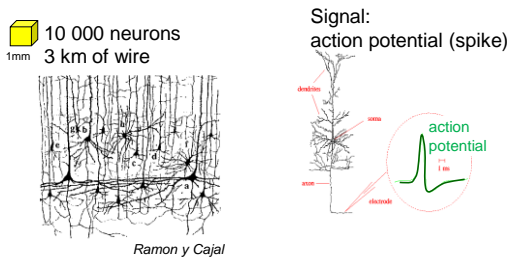
How do we recognize things?
Models of cognition
Weeks 10-14



Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

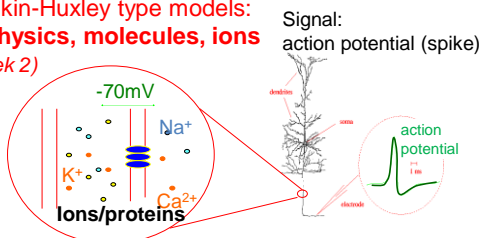


Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

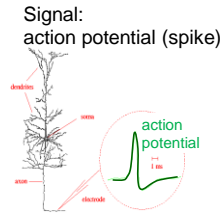


Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

Hodgkin-Huxley type models:
Biophysics, molecules, ions
(week 2)

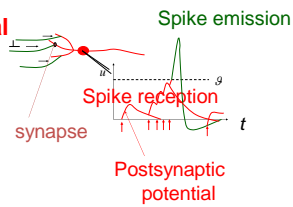


Neuronal Dynamics – 1.1. Neurons and Synapses/Overview



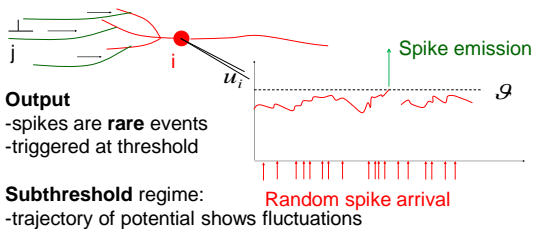
Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

Integrate-and-fire models:
Formal/phenomenological
(week 1 and week 7-9)



- spikes are events
- triggered at threshold
- spike/reset/refractoriness

Noise and variability in integrate-and-fire models



Neuronal Dynamics – membrane potential fluctuations

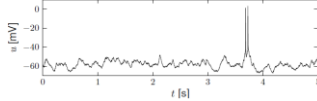
Spontaneous activity *in vivo*

What is noise?

What is the neural code?

(week 7-9)

awake mouse, cortex, freely whisking,



Lab of Prof. C. Petersen, EPFL Crochet et al., 2011

electrode



Brain

Biological Modeling of Neural Networks – Quiz 1.1

A cortical neuron sends out signals which are called:

- ☐ action potentials
- ☐ spikes
- ☐ postsynaptic potential

The dendrite is a part of the neuron

- ☐ where synapses are located
- ☐ which collects signals from other neurons
- ☐ along which spikes are sent to other neurons

In an integrate-and-fire model, when the voltage hits the threshold:

- ☐ the neuron fires a spike
- ☐ the neuron can enter a state of refractoriness
- ☐ the voltage is reset
- ☐ the neuron explodes

In vivo, a typical cortical neuron exhibits

- ☐ rare output spikes
- ☐ regular firing activity
- ☐ a fluctuating membrane potential

Multiple answers possible!

Biological Modeling of Neural Networks

Wulfram Gerstner

EPFL, Lausanne, Switzerland

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Week 5,6: Associative Memory, Hebb rule, Hopfield

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Biological modeling of Neural Networks

Course: Monday : 9:15-13:00

A typical Monday:

1st lecture 9:15-9:50

1st exercise 9:50-10:00

2nd lecture 10:15-10:35

2nd exercise 10:35-11:00

3rd lecture 11:15 – 11:40

3rd exercise 11:45-12:40

Course of 4 credits = 6 hours of work per week
4 'contact' + 2 homework

have your laptop
with you

paper and pencil

paper and pencil

paper and pencil
OR interactive toy
examples
on computer

<http://lcn.epfl.ch/~gerstner/>

moodle.epfl.ch

Week 1 – part 2: The Passive Membrane



Biological Modeling of Neural Networks

Week 1 – neurons and mathematics:
a first simple neuron model

Wulfram Gerstner
EPFL, Lausanne, Switzerland

1.1 Neurons and Synapses:

Overview

1.2 The Passive Membrane

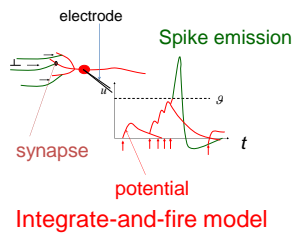
- Linear circuit
- Dirac delta-function

1.3 Leaky Integrate-and-Fire Model

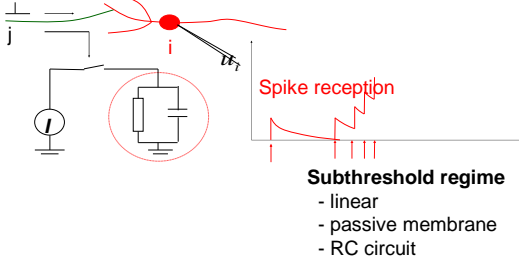
1.4 Generalized Integrate-and-Fire Model

1.5 Quality of Integrate-and-Fire Models

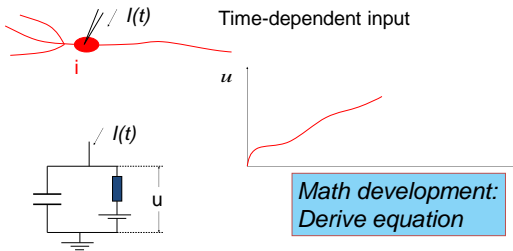
Neuronal Dynamics – 1.2. The passive membrane



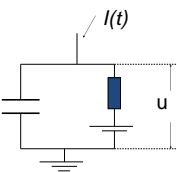
Neuronal Dynamics – 1.2. The passive membrane

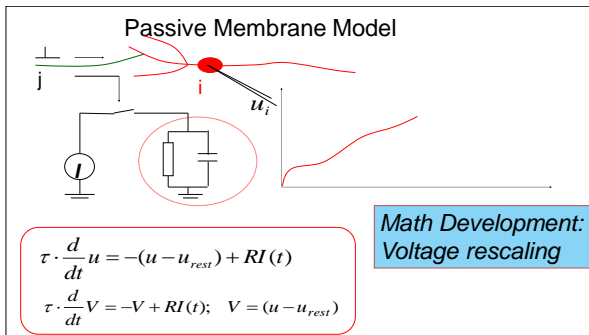


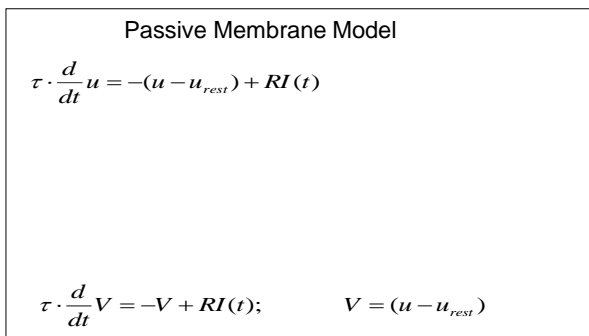
Neuronal Dynamics – 1.2. The passive membrane

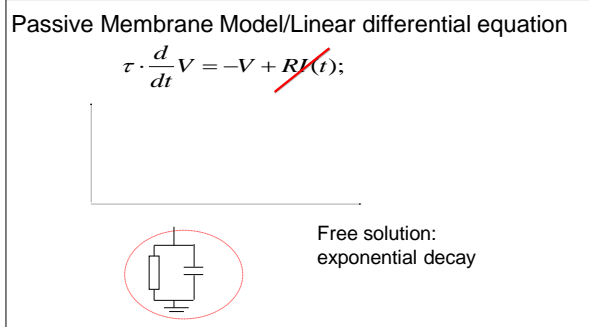


Passive Membrane Model









Neuronal Dynamics – Exercises NOW

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

$$\tau \cdot \frac{d}{dt} V = -V + RI(t); \quad V = (u - u_{rest})$$

**Start Exerc. at 9:47.
Next lecture at 10:15**

Step current input: _____

Pulse current input: _____

arbitrary current input: _____

Calculate the voltage,
for the
3 input currents

Passive Membrane Model – exercise 1 now

TA's:
Marco Lehmann

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Step current input: _____

impulse reception: _____

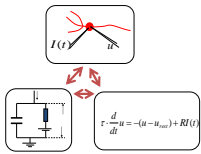
impulse response function: _____

**Start Exerc. at 9:47.
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Triangle: neuron – electricity - math

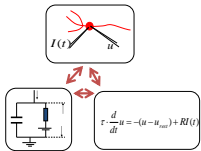
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Pulse input – charge – delta-function



$$I(t) = q \cdot \delta(t - t_0) \quad \text{Pulse current input}$$

Dirac delta-function



$$I(t) = q \cdot \delta(t - t_0)$$

Graph of $I(t)$ versus time t , showing a Dirac delta function at $t = t_0$.

$$1 = \int_{t_0-a}^{t_0+a} \delta(t - t_0) dt$$

$$f(t_0) = \int_{t_0-a}^{t_0+a} f(t) \delta(t - t_0) dt$$

Neuronal Dynamics – Solution of Ex. 1 – arbitrary input

$$\tau \cdot \frac{du}{dt} = -(u - u_{rest}) + RI(t)$$

Arbitrary input

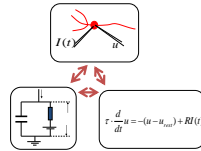
$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{c} e^{-(t-t')/\tau} I(t') dt'$$

Single pulse

$$\Delta u(t) = \frac{q}{c} e^{-(t-t_0)/\tau}$$

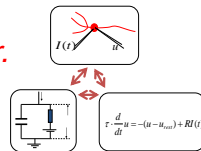
*you need to know the solutions
of linear differential equations!*

Passive membrane, linear differential equation



Passive membrane, linear differential equation

*If you have difficulties,
watch lecture 1.2detour.*

**Three prerequisites:**

- Analysis 1-3
- Probability/Statistics
- Differential Equations or Physics 1-3 or Electrical Circuits

LEARNING OUTCOMES

- Solve linear one-dimensional differential equations
- Analyze two-dimensional models in the phase plane
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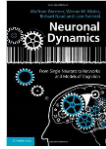
miniproject

Biological Modeling of Neural Networks

Written Exam (2/3)
+ miniproject (1/3)

<http://neurondynamics.epfl.ch/>

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Wulfram Gerstner
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Videos (for half the material):

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Questions?

Week 1 – part 3: Leaky Integrate-and-Fire Model



Biological Modeling of Neural Networks

Week 1 – neurons and mathematics:
a first simple neuron model

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EPFL, Lausanne, Switzerland

1.1 Neurons and Synapses:

Overview

1.2 The Passive Membrane

- Linear circuit
- Dirac delta-function
- Detour: solution of 1-dim linear differential equation

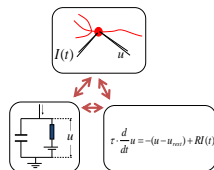
1.3 Leaky Integrate-and-Fire Model

1.4 Generalized Integrate-and-Fire Model

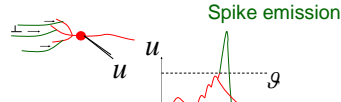
1.5. Quality of Integrate-and-Fire Models

Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



Neuronal Dynamics – Integrate-and-Fire type Models



Input spike causes an EPSP
= excitatory postsynaptic potential

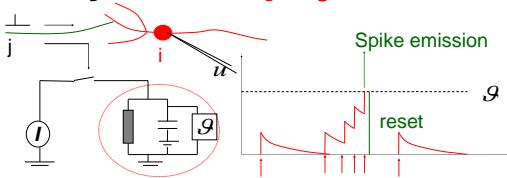
Simple Integrate-and-Fire Model:

passive membrane
+ threshold

- output spikes are events
- generated at threshold
- after spike: reset/refractoriness

Leaky Integrate-and-Fire Model

Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model



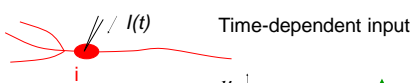
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

linear

$$u(t) = g \Rightarrow \text{Fire+reset } u \rightarrow u_r$$

threshold

Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model



Time-dependent input

Math development:
Response to step current

- spikes are events
- triggered at threshold
- spike/reset/refractoriness

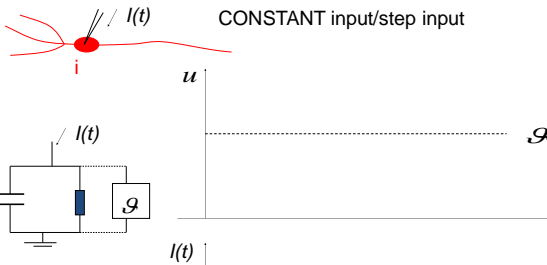
Week1- Quiz 2.**Take 90 seconds:**

Consider the linear differential equation $\tau \cdot \frac{d}{dt}x = -x + x_c$
with initial condition $at t=0 : x=0$

The solution for $t>0$ is

- (i) $x(t) = x_c \exp(t/\tau)$
- (ii) $x(t) = x_c \exp(-t/\tau)$
- (iii) $x(t) = x_c [1 - \exp(-t/\tau)]$
- (iv) $x(t) = 0.5x_c [1 + \exp(-t/\tau)]$

You will have to use the results/response to constant input/step input again and again

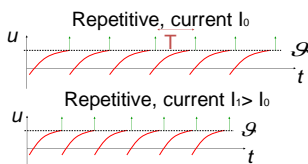
Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model

Leaky Integrate-and-Fire Model (LIF)

$$\tau \cdot \frac{d}{dt}u = -(u - u_{rest}) + RI_0$$

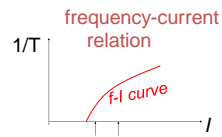
LIF
If $u(t) = g \Rightarrow u \rightarrow u_r$

'Firing'



Neuronal Dynamics – First week, Exercise 2

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



EXERCISE 2 NOW: Leaky Integrate-and-fire Model (LIF)

LIF $\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI_0$ If firing: $u \rightarrow u_r$

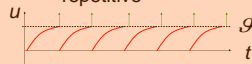
Exercise!

Calculate the interspike interval T for constant input I .
Firing rate is $f=1/T$.
Write f as a function of I .

What is the frequency-current curve $f=g(I)$ of the LIF?

assume: $u_r = u_{rest}$

repetitive



Start Exerc. at 10:53.
Next lecture at 11:15

Week 1 – part 4: Generalized Integrate-and-Fire Model



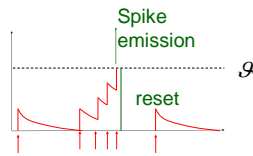
Biological Modeling of Neural Networks

Week 1 – neurons and mathematics:
a first simple neuron model

Wulfram Gerstner
EPFL, Lausanne, Switzerland

- ✓ 1.1 Neurons and Synapses: Overview
- ✓ 1.2 The Passive Membrane
 - Linear circuit
 - Dirac delta-function
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- 1.4 Generalized Integrate-and-Fire Model
- 1.5. Quality of Integrate-and-Fire Models

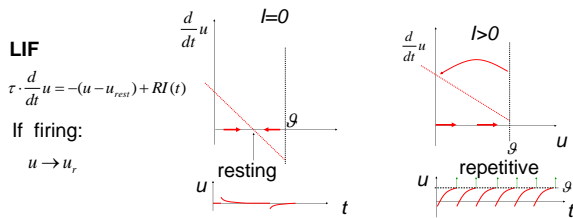
Neuronal Dynamics – 1.4. Generalized Integrate-and-Fire



Integrate-and-fire model

LIF: linear + threshold

Neuronal Dynamics – 1.4. Leaky Integrate-and Fire revisited



Neuronal Dynamics – 1.4. Nonlinear Integrate-and Fire

LIF

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

NLIF

$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

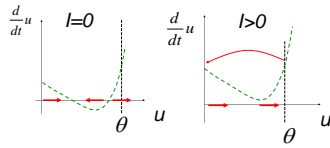
If firing:
 $u \rightarrow u_{reset}$

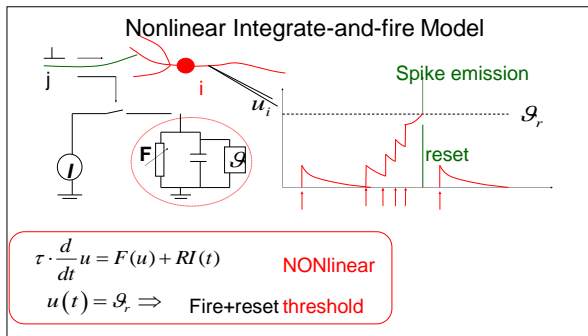
Neuronal Dynamics – 1.4. Nonlinear Integrate-and-Fire

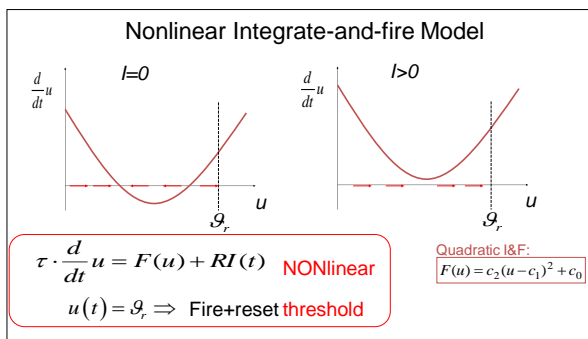
Nonlinear Integrate-and-Fire NLIF

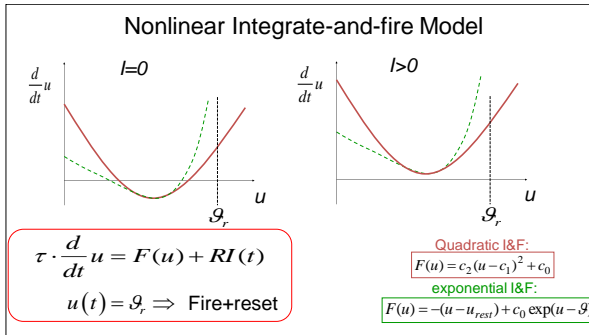
$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

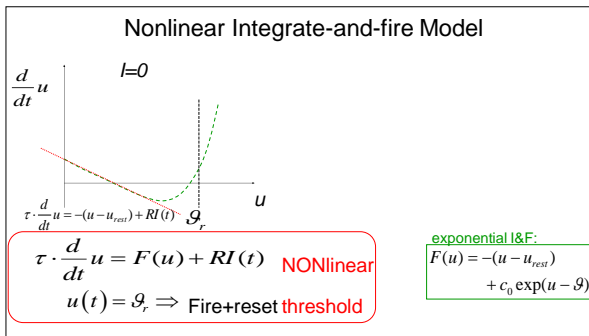
firing: $u(t) = \theta \Rightarrow$
 $u \rightarrow u_r$

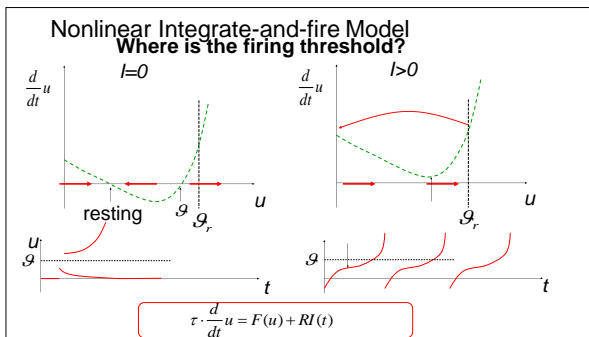












Week 1 – part 5: How good are Integrate-and-Fire Model?



Biological Modeling of Neural Networks

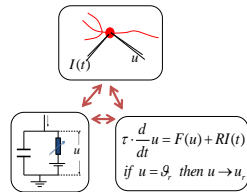
Week 1 – neurons and mathematics:
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- 1.2 The Passive Membrane
 - Linear circuit
 - Dirac delta-function
- ✓ 1.3 Leaky Integrate-and-Fire Model
- ✓ 1.4 Generalized Integrate-and-Fire Model
 - where is the firing threshold?
- 1.5. Quality of Integrate-and-Fire Models
 - Neuron models and experiments

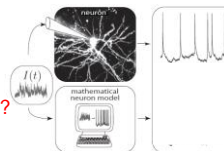
Neuronal Dynamics – 1.5. How good are integrate-and-fire models?

Can we compare neuron models
with experimental data?



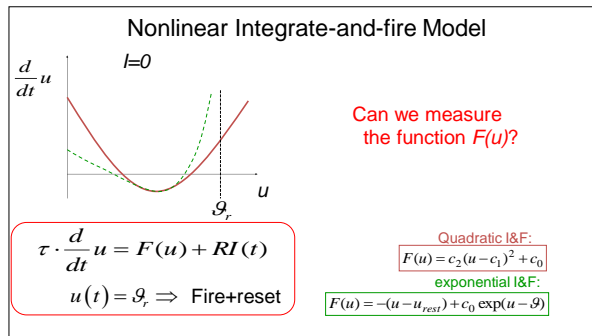
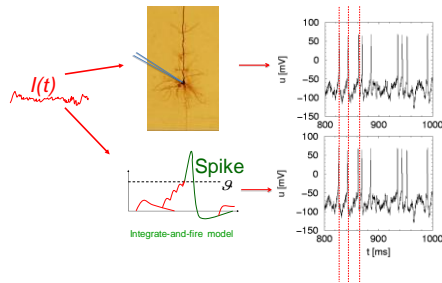
Neuronal Dynamics – 1.5. How good are integrate-and-fire models?

What is a good neuron model?

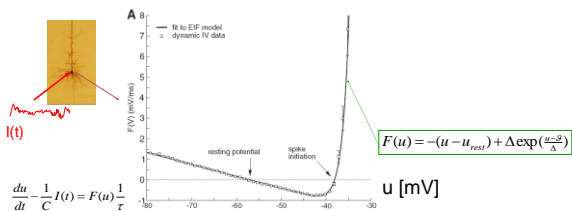


Can we compare neuron models
with experimental data?

Neuronal Dynamics – 1.5. How good are integrate-and-fire models?



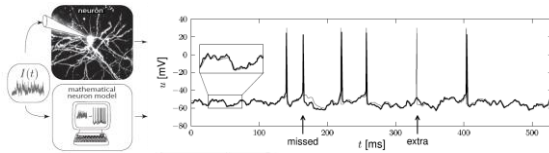
Neuronal Dynamics – 1.5. How good are integrate-and-fire models?



Badel et al., J. Neurophysiology 2008



Neuronal Dynamics – 1.5. How good are integrate-and-fire models?



Neuronal Dynamics – 1.5. How good are integrate-and-fire models?

Nonlinear integrate-and-fire models
are good

Mathematical description → prediction

Computer exercises:
Python

Need to add

- adaptation
- noise
- dendrites/synapses

Biological Modeling of Neural Networks

TA's:

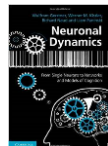
Marco Lehmann
Vasiliki Liakoni
Florian Colombo

<http://neurondynamics.epfl.ch/>

Textbook:

Lecture today:

- Chapter 1
- Chapter 5



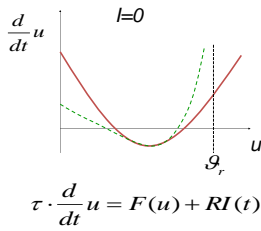
Exercises today:

- Install PYTHON for Computer Exercises
- Exercise 3, on sheet

Videos (for today: 'week 1'):

<http://lcn.epfl.ch/~gerstner/NeuronalDynamics-MOOC1.html>

Biological Modeling of Neural Networks – week1/Exercise 3



Homework!

First week – References and Suggested Reading

Reading: W. Gerstner, W.M. Kistler, R. Naud and L. Paninski, *Neuronal Dynamics: from single neurons to networks and models of cognition*. Chapter 1: Introduction. Cambridge Univ. Press, 2014

Selected references to linear and nonlinear integrate-and-fire models

- Lapique, L. (1907). *Recherches quantitatives sur l'excitation électrique des nerfs traitée comme une polarisation*. J. Physiol. Pathol. Gen., 9:620-635.
- Stein, R. B. (1965). A theoretical analysis of neuronal variability. Biophys. J., 5:173-194.
- Ermentrout, G. B. (1996). *Type I membranes, phase resetting curves, and synchrony*. Neural Computation, 8(5):979-1001.
- Fourcaud-Trocme, N., Hansel, D., van Vreeswijk, C., and Brunel, N. (2003). *How spike generation mechanisms determine the neuronal response to fluctuating input*. J. Neuroscience, 23:11628-11640.
- Badel, L., Lefort, S., Berger, T., Petersen, C., Gerstner, W., and Richardson, M. (2008). Biological Cybernetics, 99(4-5):361-370.
- Latham, P. E., Richmond, B., Nelson, P., and Nirenberg, S. (2000). *Intrinsic dynamics in neuronal networks. I. Theory*. J. Neurophysiology, 83:808-827.

First week

THE END (of main lecture)

MATH DETOUR SLIDES
(for online VIDEO)

Week 1 – part 2: Detour/Linear differential equation



**Neuronal Dynamics:
Computational Neuroscience
of Single Neurons**

Week 1 – neurons and mathematics:
a first simple neuron model

Wulfram Gerstner
EPFL, Lausanne, Switzerland

✓ 1.1 Neurons and Synapses:
Overview

✓ 1.2 The Passive Membrane

- Linear circuit
- Dirac delta-function
- Detour: solution of 1-dim linear differential equation

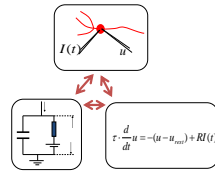
1.3 Leaky Integrate-and-Fire Model

1.4 Generalized Integrate-and-Fire Model

1.5 Quality of Integrate-and-Fire Models

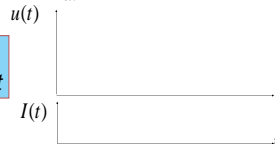
Neuronal Dynamics – 1.2 Detour – Linear Differential Eq.

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

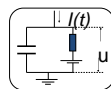


Neuronal Dynamics – 1.2 Detour – Linear Differential Eq.

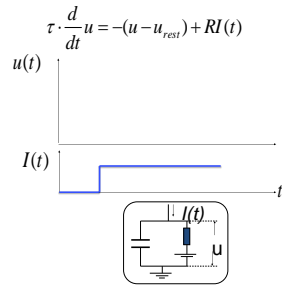
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



Math development:
Response to step current



Neuronal Dynamics – 1.2Detour – Step current input

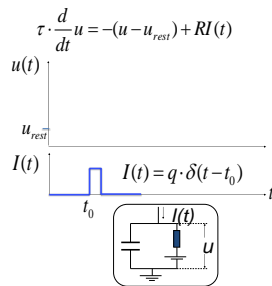


Neuronal Dynamics – 1.2Detour – Short pulse input

$$u(t) = u_{rest} + RI_0 \left[1 - e^{-(t-t_0)/\tau} \right]$$

short pulse: $(t - t_0) \ll \tau$

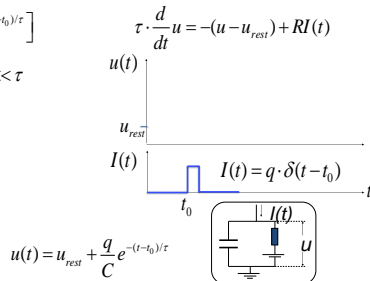
*Math development:
Response to short
current pulse*



Neuronal Dynamics – 1.2Detour – Short pulse input

$$u(t) = u_{rest} + RI_0 \left[1 - e^{-(t-t_0)/\tau} \right]$$

short pulse: $(t - t_0) \ll \tau$



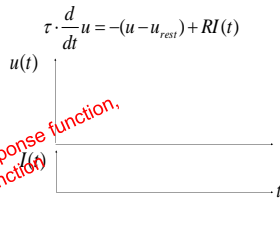
Neuronal Dynamics – 1.2Detour – arbitrary input

Single pulse

$$u(t) = u_{rest} + \frac{q}{C} e^{-(t-t_0)/\tau}$$

Multiple pulses:

$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{C} e^{-(t-t')/\tau} I(t') dt'$$



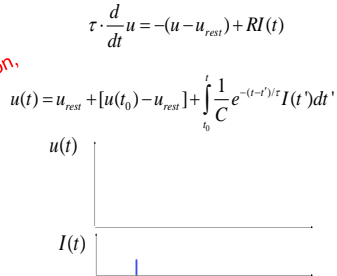
Neuronal Dynamics – 1.2Detour – Greens function

Single pulse

$$\Delta u(t) = q \frac{1}{C} e^{-(t-t_0)/\tau}$$

Multiple pulses:

$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{C} e^{-(t-t')/\tau} I(t') dt'$$



Neuronal Dynamics – 1.2Detour – arbitrary input

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

*If you don't feel at ease yet,
spend **10 minutes** on these
mathematical exercise
And quiz 2 in week 1.*

Arbitrary input

$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{C} e^{-(t-t')/\tau} I(t') dt'$$

Single pulse

$$\Delta u(t) = \frac{q}{C} e^{-(t-t_0)/\tau}$$

*you need to know the solutions
of linear differential equations!*
