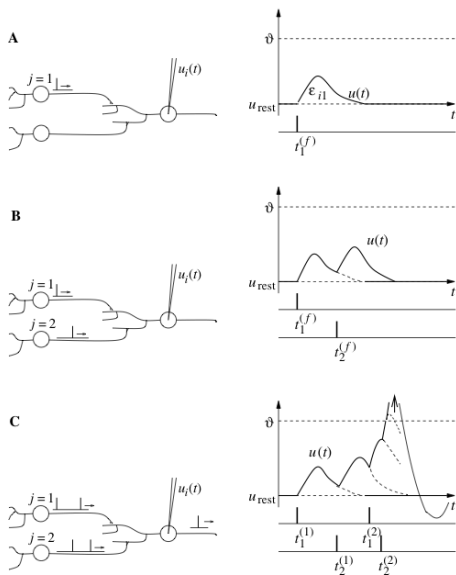


Neuroinformatics 2013

March 7, 2013

Lecture 4: Associators and synaptic plasticity

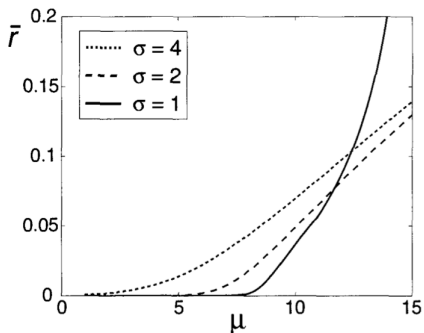
PSP and AP - Supplement of lec4:LIF-neuron noise simulation II



Activation function

- ▶ IF neuron driven by I_{ext} with normal pdf \rightarrow stochastic differential equation
- ▶ t^f is random variable
- ▶ average firing rate

$$\bar{r} = \left(t^{ref} + \tau_m \int_{v_{res} - RI_{ext}/\sigma}^{v^{\vartheta} - RI_{ext}/\sigma} \right) \sqrt{(\pi)} e^{v^2} [1 + erf(v) dv]^{-1}$$

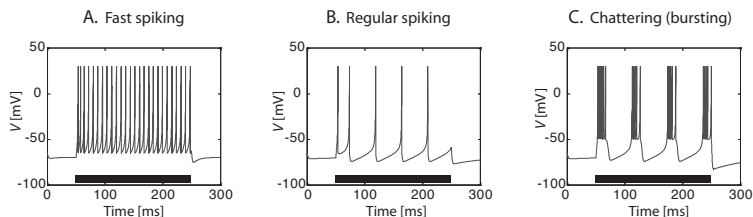


The Izhikevich neuron I

$$\frac{dv(t)}{dt} = 0.04v^2 + 5v + 140 - u + I(t)$$

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

$$v(v > 30) = c \text{ and } u(v > 30) = u + d$$

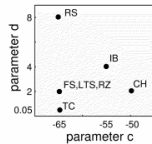
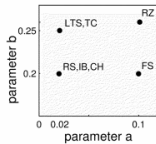
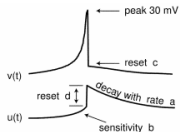


The Izhikevich neuron II

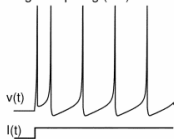
$$v' = 0.04v^2 + 5v + 140 - u + I$$

$$u' = a(bv - u)$$

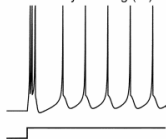
**if $v = 30$ mV,
then $v \leftarrow c$, $u \leftarrow u + d$**



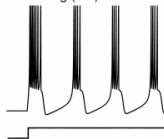
regular spiking (RS)



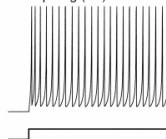
intrinsically bursting (IB)



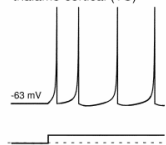
chattering (CH)



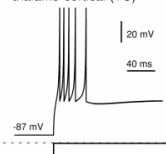
fast spiking (FS)



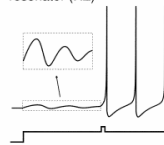
thalamo-cortical (TC)



thalamo-cortical (TC)



resonator (RZ)



low-threshold spiking (LTS)



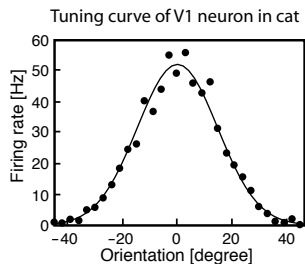
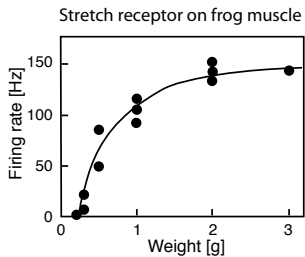
McCulloch-Pitts neuron

- ▶ first model of neuron, A logical calculus of the ideas immanent in nervous activity, Bulletin of Mathematical Biophysics 5:115-133, 1943
- ▶ 1958 - perceptron by Rosenblatt
- ▶ heavy side function as transfer (activation) function, simple logical OR, AND

$$h = \sum_i x_i^{\text{in}}$$

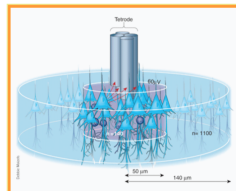
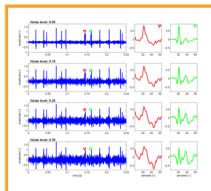
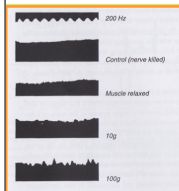
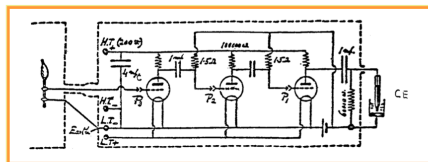
$$x^{\text{out}} = \begin{cases} 1 & \text{if } h > \Theta \\ 0 & \text{otherwise} \end{cases}$$

The firing rate hypothesis



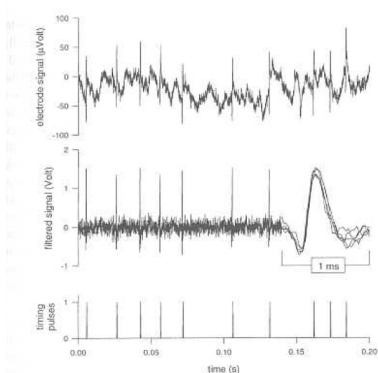
History: Adrian

- ▶ Sir Edgarda Adriana (Nobel price for medicine - 1932)
http://nobelprize.org/nobel_prizes/medicine/laureates/1932/adrian-bio.html

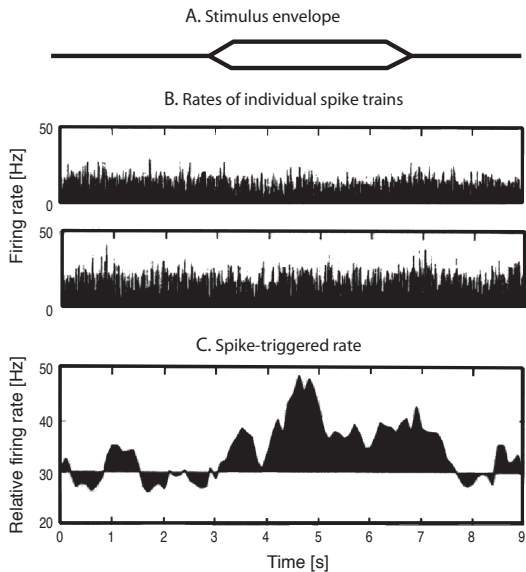


All or none coding

- ▶ top: tungsten electrode in fly's brain, middle: low frequency removal, bottom: spikes

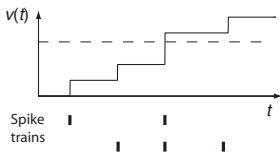


Correlation coding

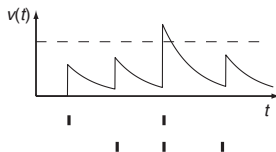


Integrator or coincidence detector?

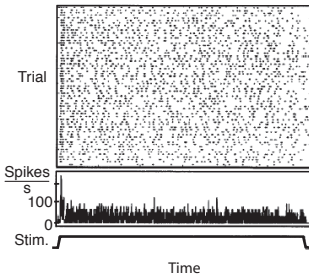
A. Perfect integrator



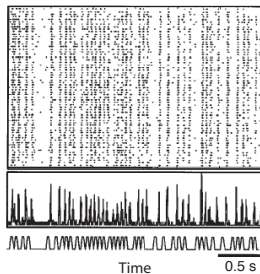
B. Coincidence detector



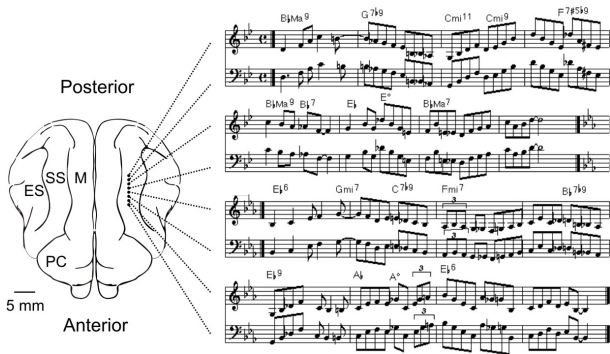
A. Constant stimulus



B. Rapidly changing stimulus



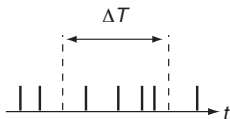
Neuronal music



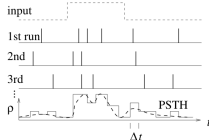
http://cns.iaf.cnrs-gif.fr/alain_music.html

Rate codes

- ▶ Spike count - average over time
- ▶ Spike density - average over several trial:
peri-stimulus-time histogram (PSTH)
- ▶ Population activity - organization of many neurons in columns
cells. Idealized situations - neurons with the same properties.



rate = average over several runs
(single neuron, repeated runs)



spike density
in PSTH

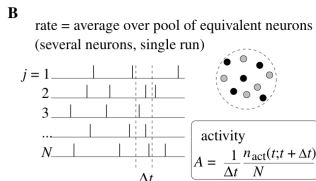
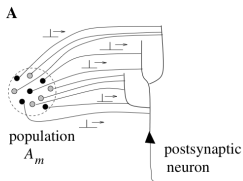
$$\rho = \frac{1}{\Delta t} \frac{1}{K} n_K(t; t + \Delta t)$$

Population model

$$a(t) = \frac{\text{number of spikes in } \Delta T}{\Delta T} = \frac{1}{\Delta T} \int_{t-\Delta T/2}^{t+\Delta T/2} \delta(t' - t^f) dt'$$

$$A(t) = \lim_{\Delta T \rightarrow 0} \frac{\text{number of spikes in population of size } N}{N}$$

$$= \lim_{\Delta T \rightarrow 0} \frac{1}{\Delta T} \int_{t-\Delta T/2}^{t+\Delta T/2} \frac{1}{N} \sum_{i=1}^N \delta(t' - t^f) dt'$$



Population dynamics

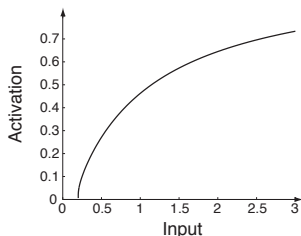
For slow varying input (adiabatic limit), when all nodes do practically the same, same input, etc (Wilson and Cowan, 1972):

$$\tau \frac{dA(t)}{dt} = -A(t) + g(RI^{\text{ext}}(t)). \quad (1)$$

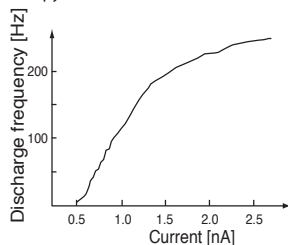
Gain function:

$$g(x) = \frac{1}{t^{\text{ref}} - \tau \log\left(1 - \frac{1}{\tau x}\right)}, \quad (2)$$





A. Activation function for population average in adiabatic limit



B. Activation function of hippocampal pyramidal neuron

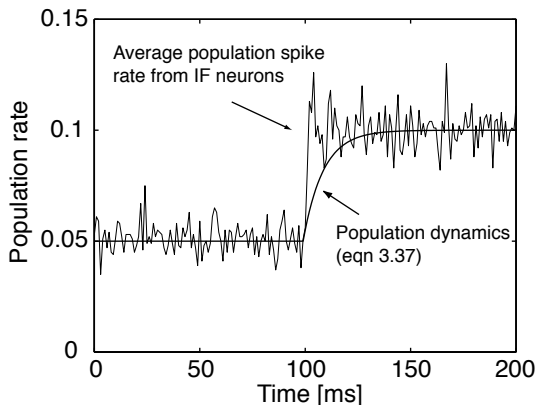


Other gain functions

Type of function	Graphical represent.	Mathematical formula	MATLAB implementation
Linear		$g^{\text{lin}}(x) = x$	<code>x</code>
Step		$g^{\text{step}}(x) = \begin{cases} 1 & \text{if } x > 0 \\ 0 & \text{elsewhere} \end{cases}$	<code>floor(0.5*(1+sign(x)))</code>
Threshold-linear		$g^{\text{theta}}(x) = x \Theta(x)$	<code>x.*floor(0.5*(1+sign(x)))</code>
Sigmoid		$g^{\text{sig}}(x) = \frac{1}{1+\exp(-x)}$	<code>1./(1+exp(-x))</code>
Radial-basis		$g^{\text{gauss}}(x) = \exp(-x^2)$	<code>exp(-x.^2)</code>

SIMULATION - Fast population response

- ▶ Simulation with 1000 independent (NOT CONNECTED) IF neurons: $I_{ext} = I_{ext} + \eta, \eta \in N(0, 1), \tau_m = 10 \text{ ms}, \vartheta = 10 \text{ mV}$
- ▶ Switching from $RI_{ext} = 11 \text{ mV}$ at $t = 100 \text{ ms}$ to $RI_{ext} = 16 \text{ mV}$



Types of plasticity

- ▶ **Structural plasticity** is the mechanism describing the generation of new connections and thereby redefining the topology of the network.
- ▶ **Functional plasticity** is the mechanism of changing the strength values of existing connections.

Hebbian plasticity

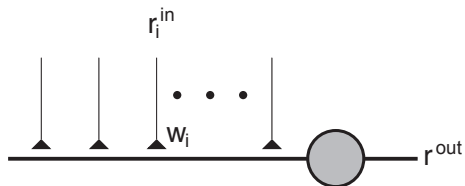
”When an axon of a cell A is near enough to excite cell B or repeatedly or persistently takes part in firing it, some growth or metabolic change takes place in both cells such that A’s efficiency, as one of the cells firing B, is increased.”

Donald O. Hebb, **The organization of behavior**, 1949

See also Sigmund Freud, **Law of association by simultaneity**, 1888

Santiago Ramn y Cajal - memories might instead be formed by strengthening the connections between existing neurons to improve the effectiveness of their communication, 1894

Association



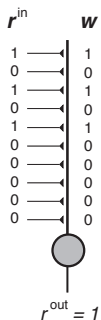
Neuron model: In each time step the model neurons fires if

$$\sum_i w_i r_i^{\text{in}} > 1.5$$

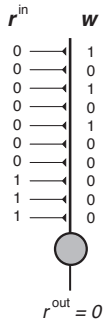
Learning rule: Increase the strength of the synapses by a value $\Delta w = 0.1$ if a presynaptic firing is paired with a postsynaptic firing.

Associative learning

A. *Before learning, only odour cue*



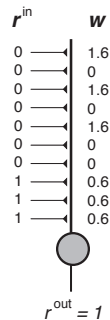
B. *Before learning, only visual cue*



C. *After 1 learning step, both cues*



D. *After 6 learning steps, only visual cue*

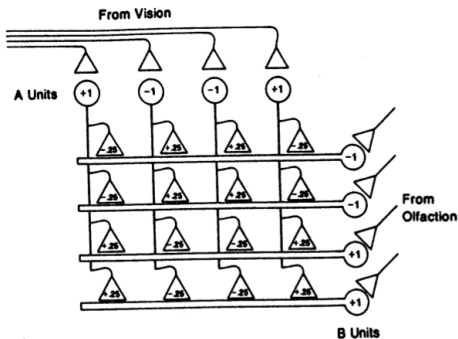


Features of associators and Hebbian learning

- ▶ Pattern completion and generalization, recall from partial input, overlap between input and trained pattern (recognition of noisy numbers)
- ▶ Prototypes and extraction of central tendencies, training on many similar but not equivalent examples (individual face, many common features in all faces)
- ▶ Graceful degradation and fault tolerance (loss of synapses or whole neurons)

How memory is stored?

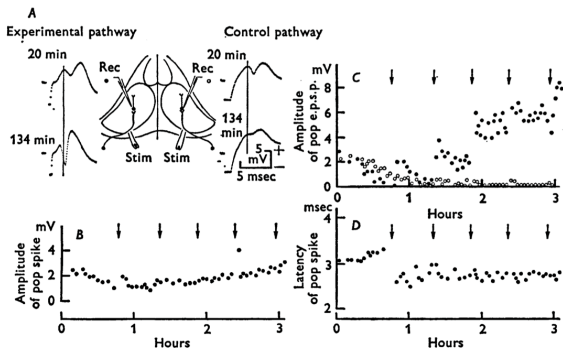
- ▶ Connectionist modeling of memory, one neural network (associator) for visual and olfactory system!
- ▶ machine learning theory of neural networks -e.g. back propagation principle



$$\begin{bmatrix} - & + & + & - \\ - & + & + & - \\ + & - & - & + \\ + & - & - & + \end{bmatrix} + \begin{bmatrix} + & - & + & - \\ - & + & - & + \\ - & + & - & + \\ + & - & + & - \end{bmatrix} = \begin{bmatrix} & ++ & -- \\ -- & ++ & \\ ++ & -- & \end{bmatrix}$$

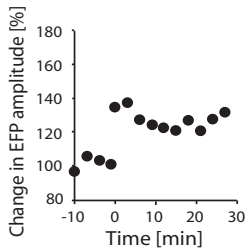
Original LTP by Bliss and Lomo, 1973

- ▶ Long-lasting changes of synaptic response characteristics
- ▶ High frequency-stimulus is applied (plasticity-induced tetanus) → long-term potentiation (to strengthen, make more potent) (LTP) average amplitude of EPSP increased
- ▶ Long frequency stimulus → long-term depression (LTD)

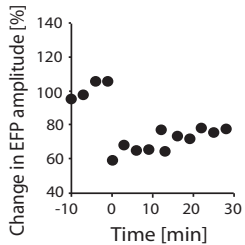


Classical LTP and LTD

A. Long term potentiation

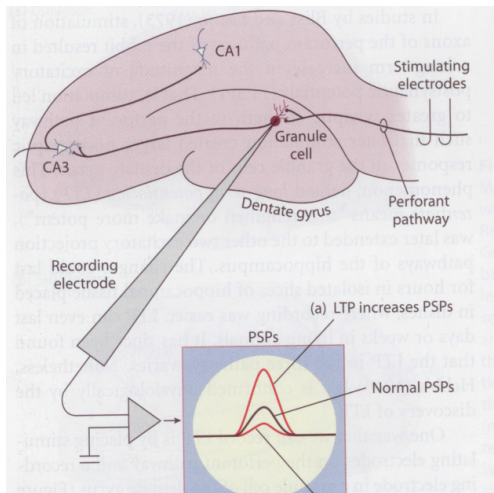


B. Long term depression



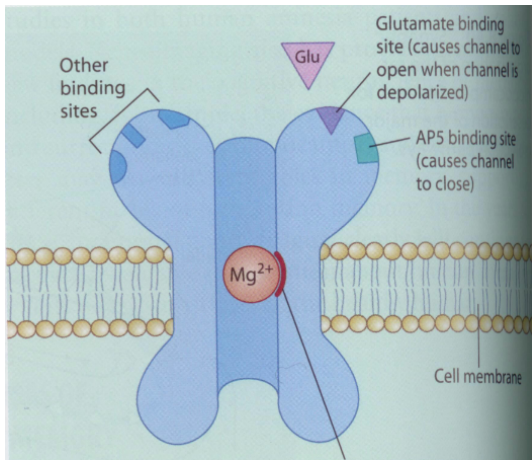
LTP experiment

- ▶ EXPERIMENTAL confirmation of Hebb's rule (1949)
- ▶ i) single pulse is presented ii) stimulation with burst of pulses: 100 pulses/sec ii) After LTP induced, single pulse stimulation
- ▶ Postsynaptic cells must be depolarized to LTP be produced AND receiving excitatory input - see Associative learning slide.



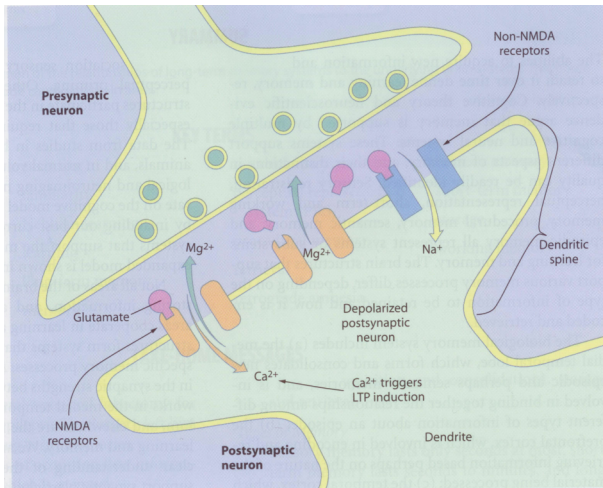
NMDA receptors

- ▶ N-methyl-D-aspartate receptor located on dendritic spines of postsynaptic neurons showing LTP
- ▶ i) NMDA receptors are blocked by Mg^{2+} ii) Channel unblocking after glutamate binding (glutamate is major excitatory transmitter in hippocampus) AND membrane depolarized (NMDA are voltage gated) → Mg^{2+} ejection, Ca^{2+} influx



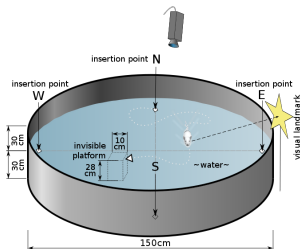
Ca²⁺ role

- ▶ Ca²⁺ changes enzyme activities that influence synaptic strength
- ▶ LTP raises sensitivity of non-NMDA glutamate receptors prompting release of more glutamate



Morris Water Maze - spatial memory

- ▶ i) mice training ii) Chemical blocking of LTP by AP5 impair spatial learning, keep control group iii) AP5-treated mice significantly impaired
- ▶ i) slices of the hippocampus were taken from both groups ii) LTP was easily induced in controls, but could not be induced in the brains of APV-treated rats
- ▶ Alzheimer's disease → cognitive decline seen in individuals with AD may result from impaired LTP ??



Further Readings

Laurence F. Abbott and Sacha B. Nelson (2000), **Synaptic plasticity: taming the beast**, in **Nature Neurosci. (suppl.)**, 3: 1178–83.

Alain Artola and Wolf Singer (1993), **Long-term depression of excitatory synaptic transmission and its relationship to long-term potentiation**, in **Trends in Neuroscience** 16: 480–487.

Mark C. W. van Rossum, Guo-chiang Bi, and Gina G. Turrigiano (2000) **Stable Hebbian learning from spike timing-dependent plasticity**, in **J. Neuroscience** 20(23): 8812–21