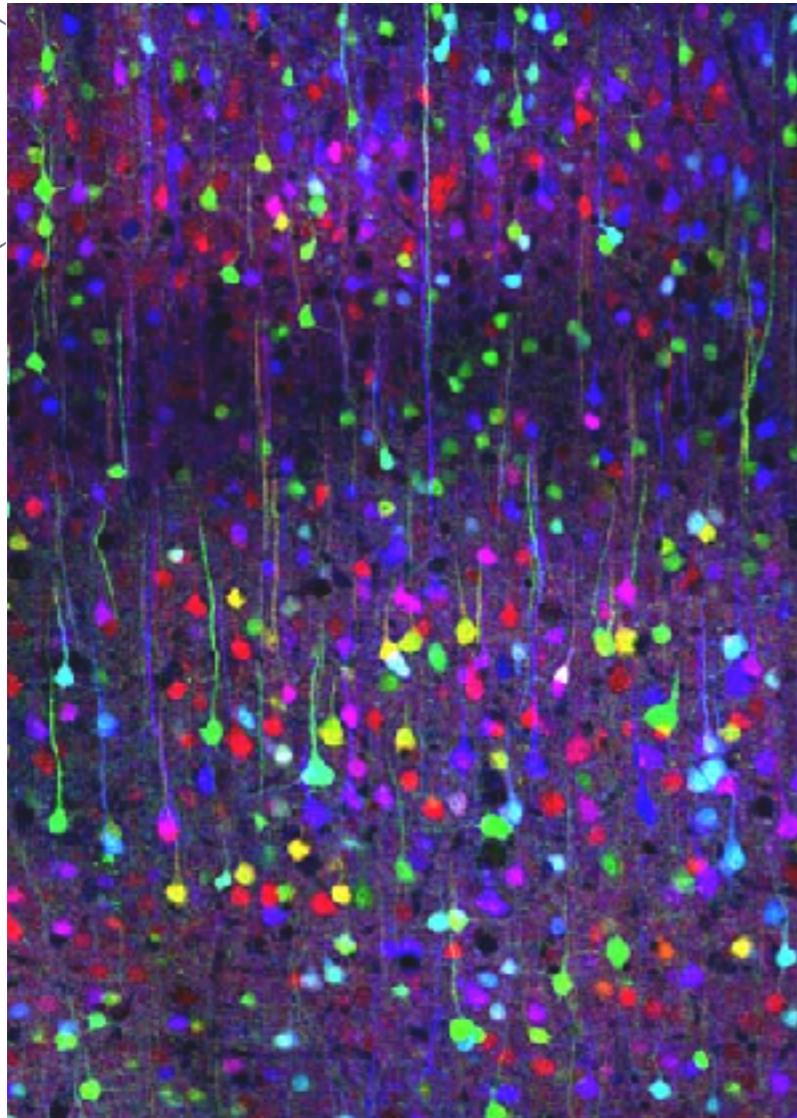


Computational Neuroscience 2018/2019

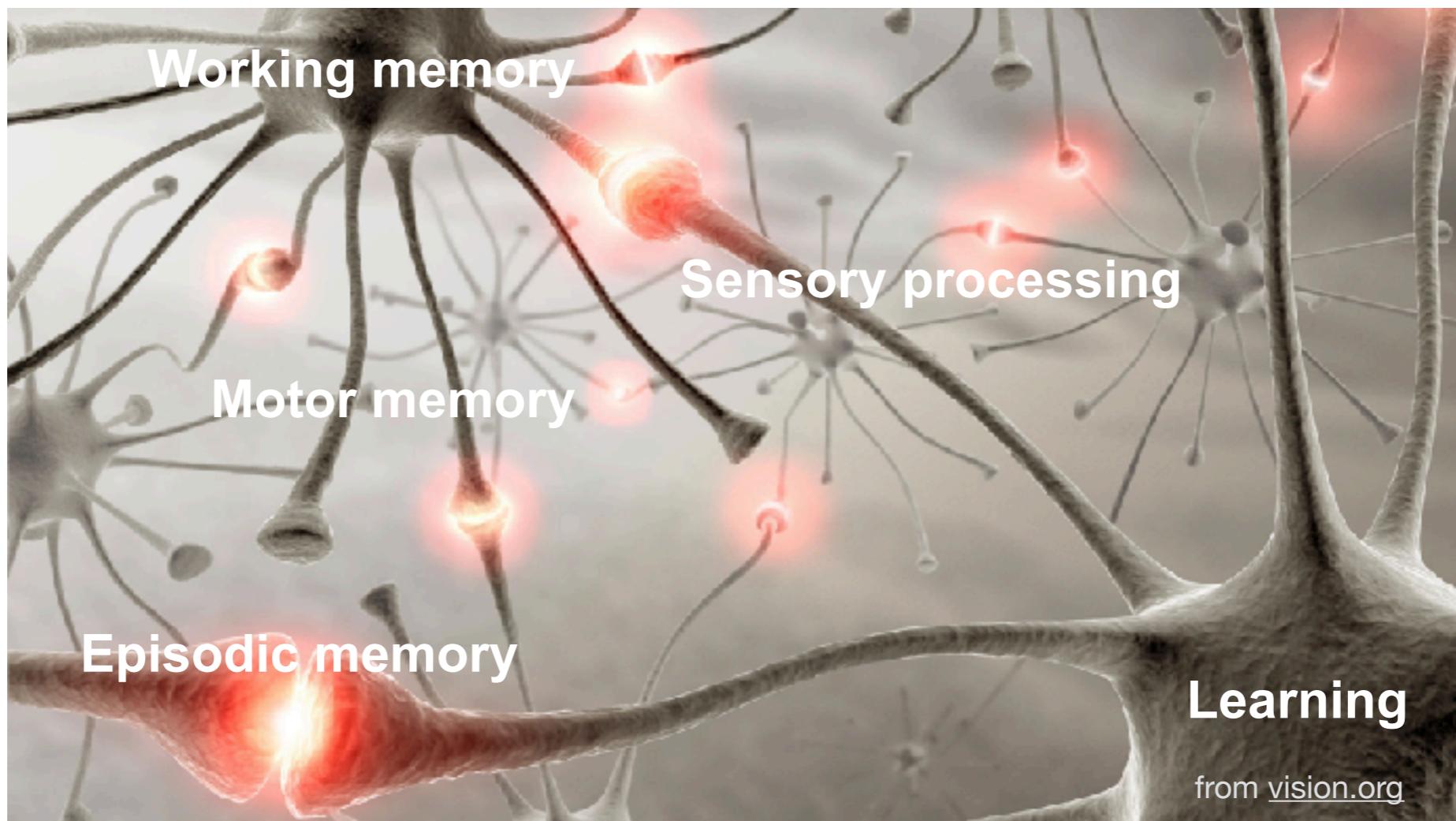


Brainbow (Litchman Lab)

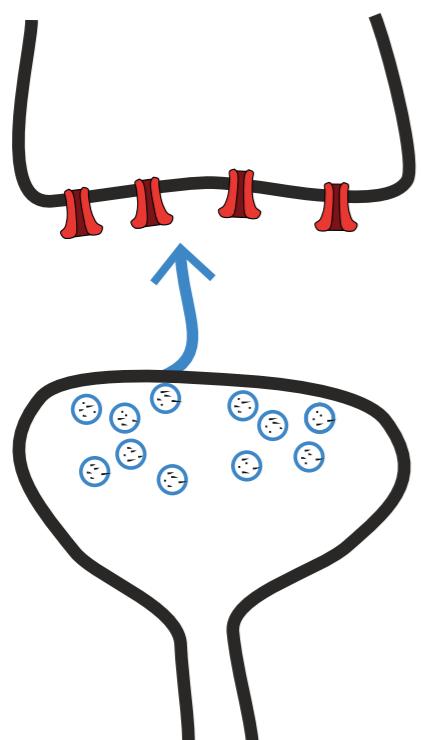


Lecture 16 Synaptic plasticity: Short-term synaptic plasticity

Synaptic plasticity underlies memory and learning



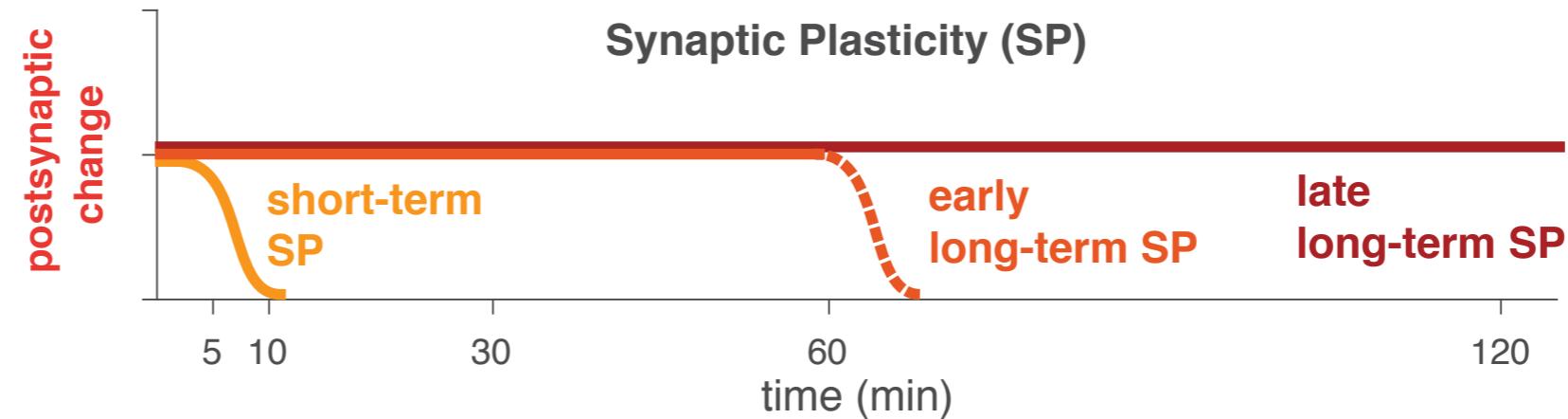
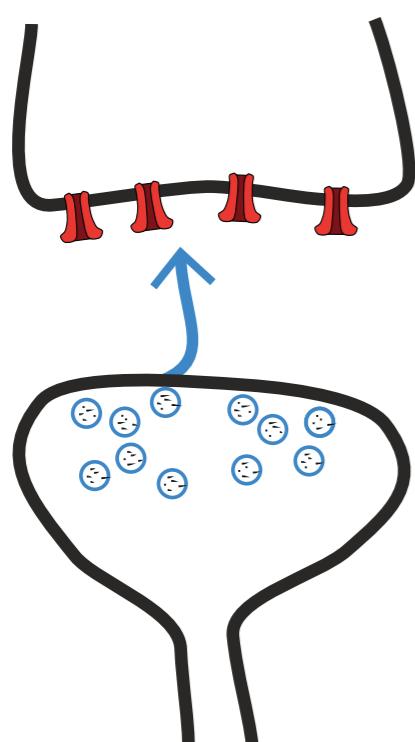
Synaptic plasticity



Synaptic plasticity: Multiple timescales and locations

Synapses can change **postsynaptically**

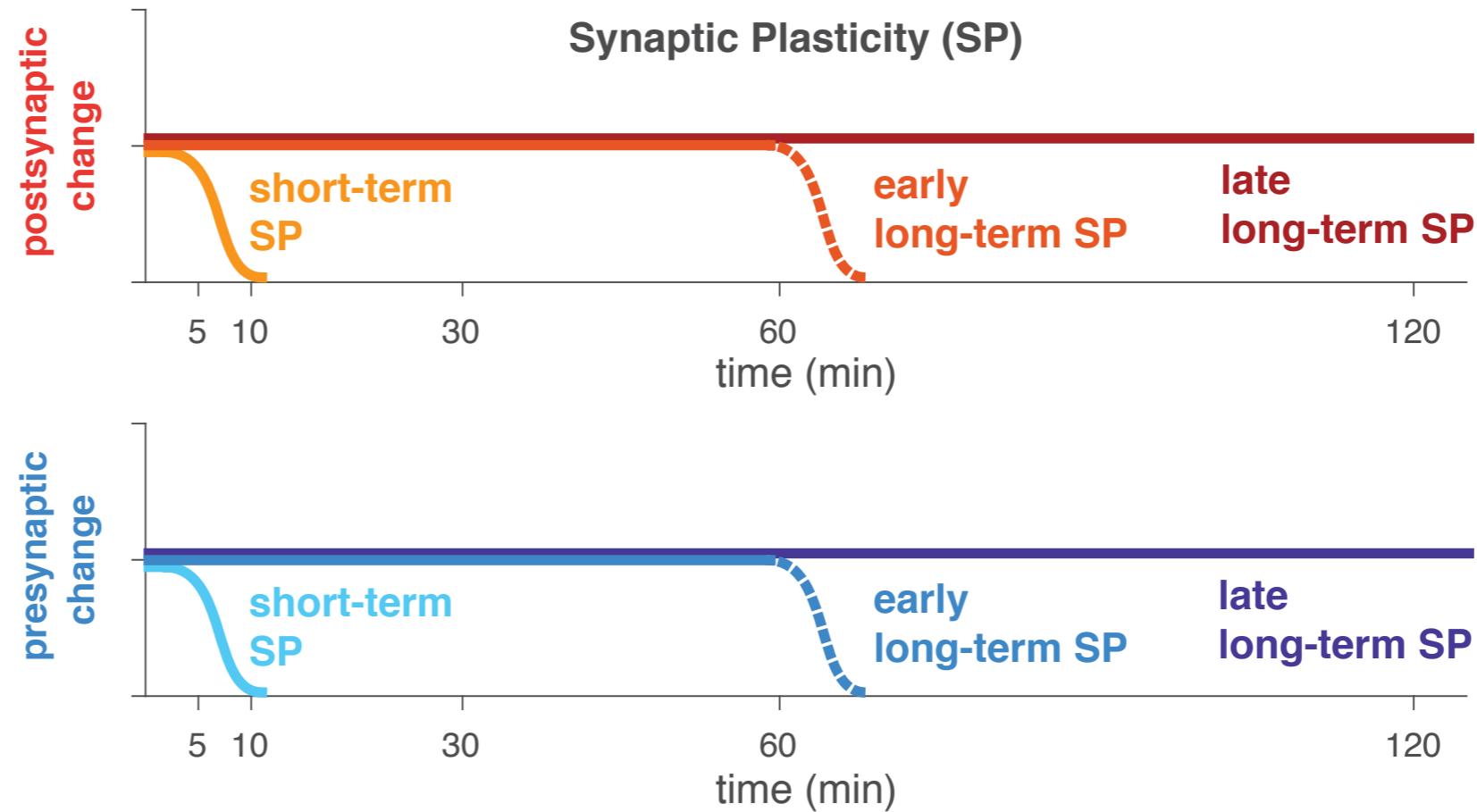
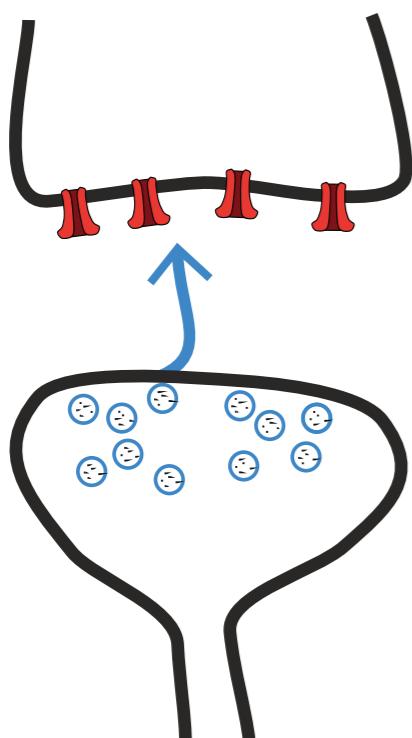
(from a few seconds (**short-term**) to tens of minutes, hours (**long-term**) or even days/years!):



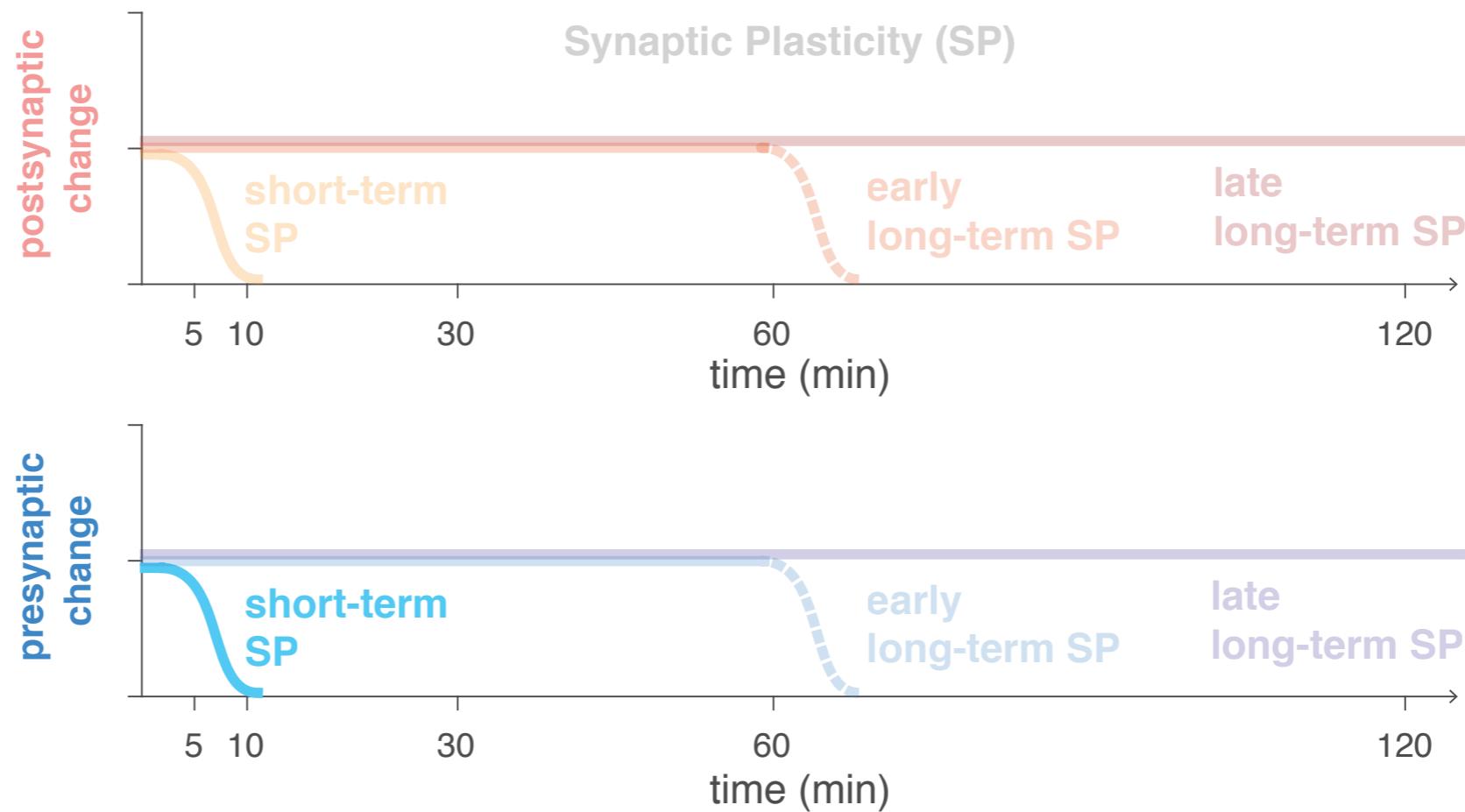
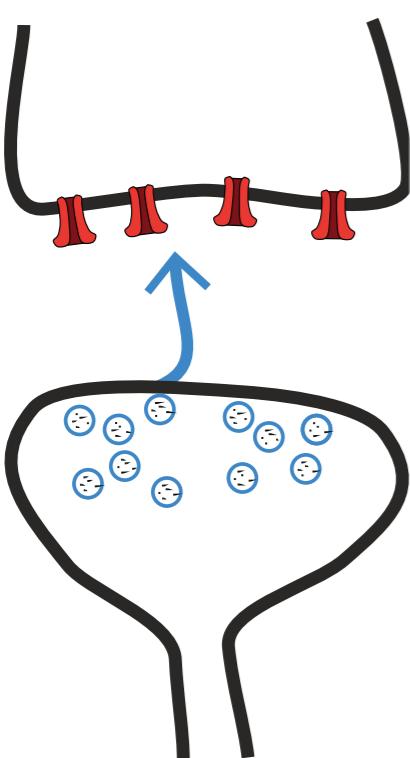
Synaptic plasticity: Multiple timescales and locations

But they can also change **presynaptically**

(from a few seconds (**short-term**) to tens of minutes, hours (**long-term**) or even days/years!):



Presynaptic Short-term synaptic plasticity



Abbott & Regehr Nature 2004

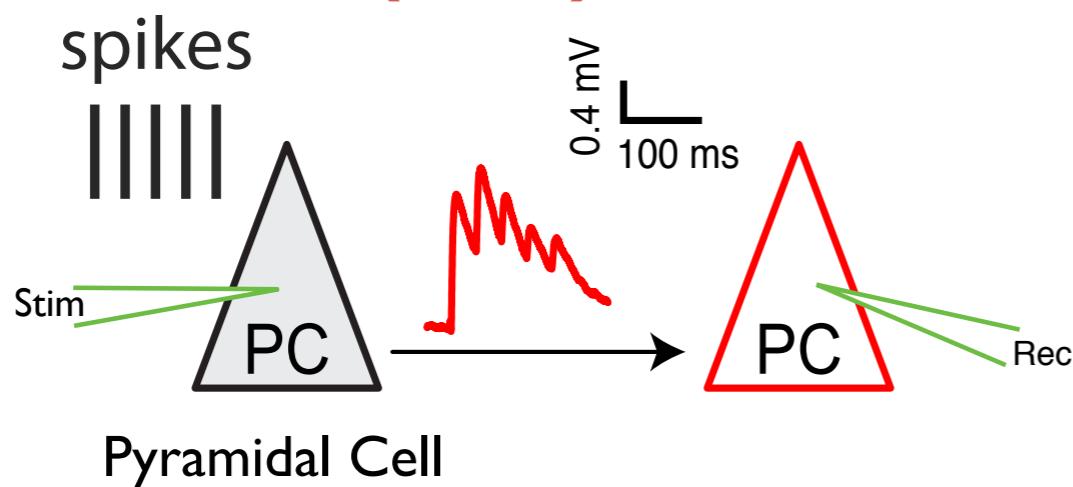
Outline

Short-term synaptic plasticity (STP)

- 1. The two main types**
- 2. Models: Tsodyks-Markram and other models**
- 3. Functions**
 - 3.1. Network dynamics**
 - 3.2. Theory of working memory**

Two main types of short-term synaptic plasticity

Short-term depression (STD)

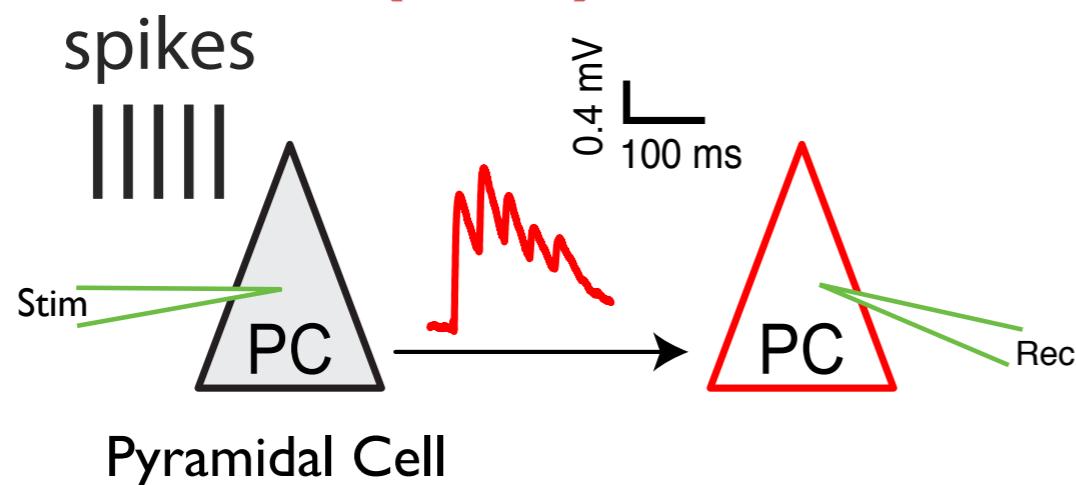


Tsodyks and Markram, PNAS 1997; Costa et al. 2013 Frontiers

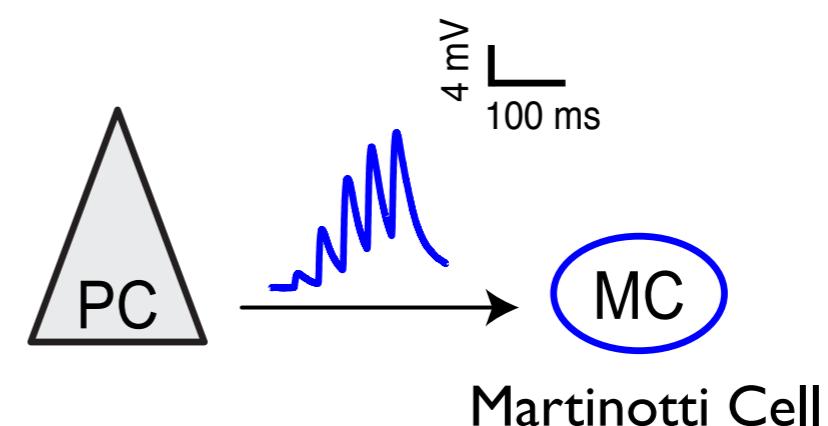
Abbott & Regehr Nature 2004

Two main types of short-term synaptic plasticity

Short-term depression (STD)



Short-term facilitation (STF)



Tsodyks and Markram, PNAS 1997; Costa et al. 2013 Frontiers

Abbott & Regehr Nature 2004

Models of short-term synaptic plasticity

Models of short-term synaptic plasticity (STP)

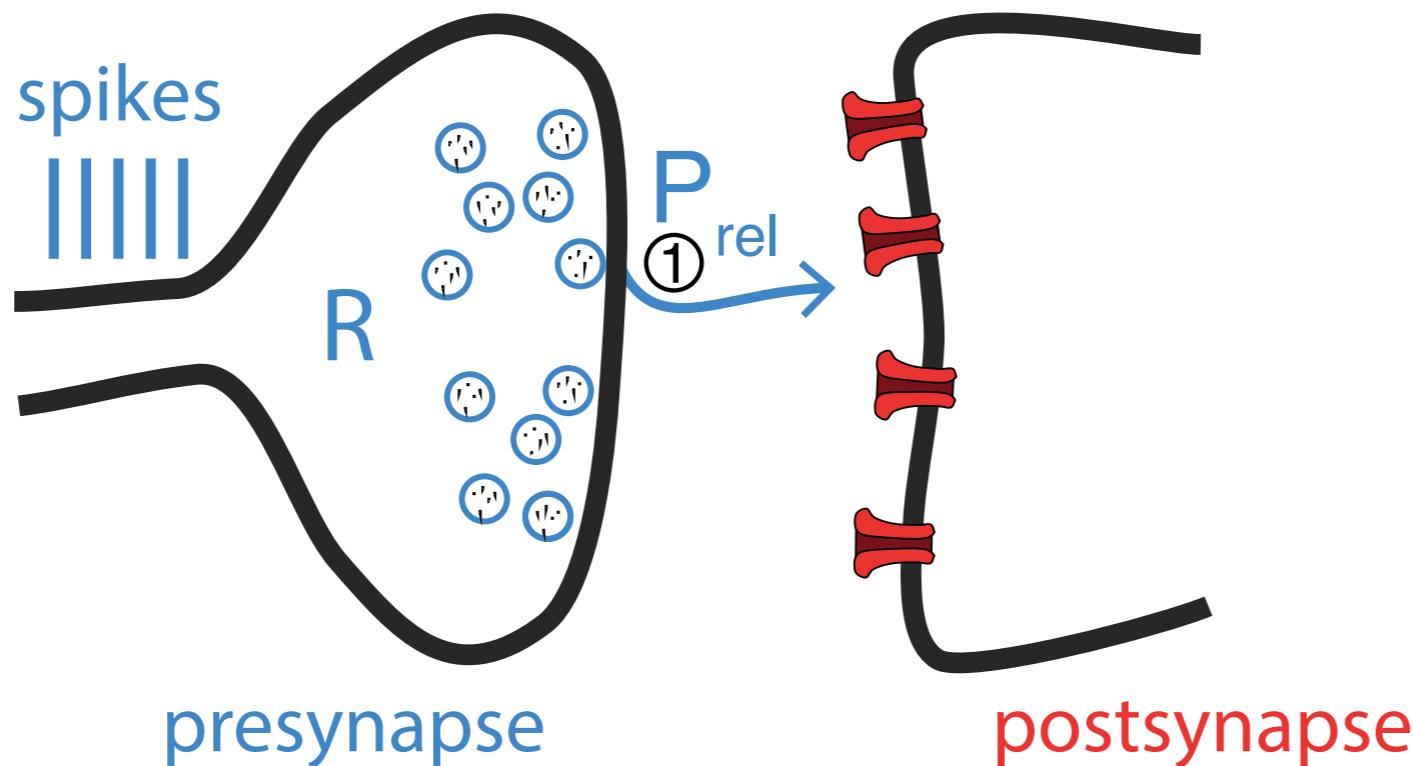
I. Phenomenological models

1. Varela model [more abstract model, Varela et al. JNeurosci 1997]
2. Markram-Tsodyks (STD and STF) [closer to biology]: Tsodyks, Markram et al. PNAS 1997/1998

2. Molecular models

1. Pan and Zucker Neuron 2009

A phenomenological model of STD: Tsodyks-Markram model



Abstracted out of molecular ‘details’!

1: R vesicles are released with probability P_{rel}

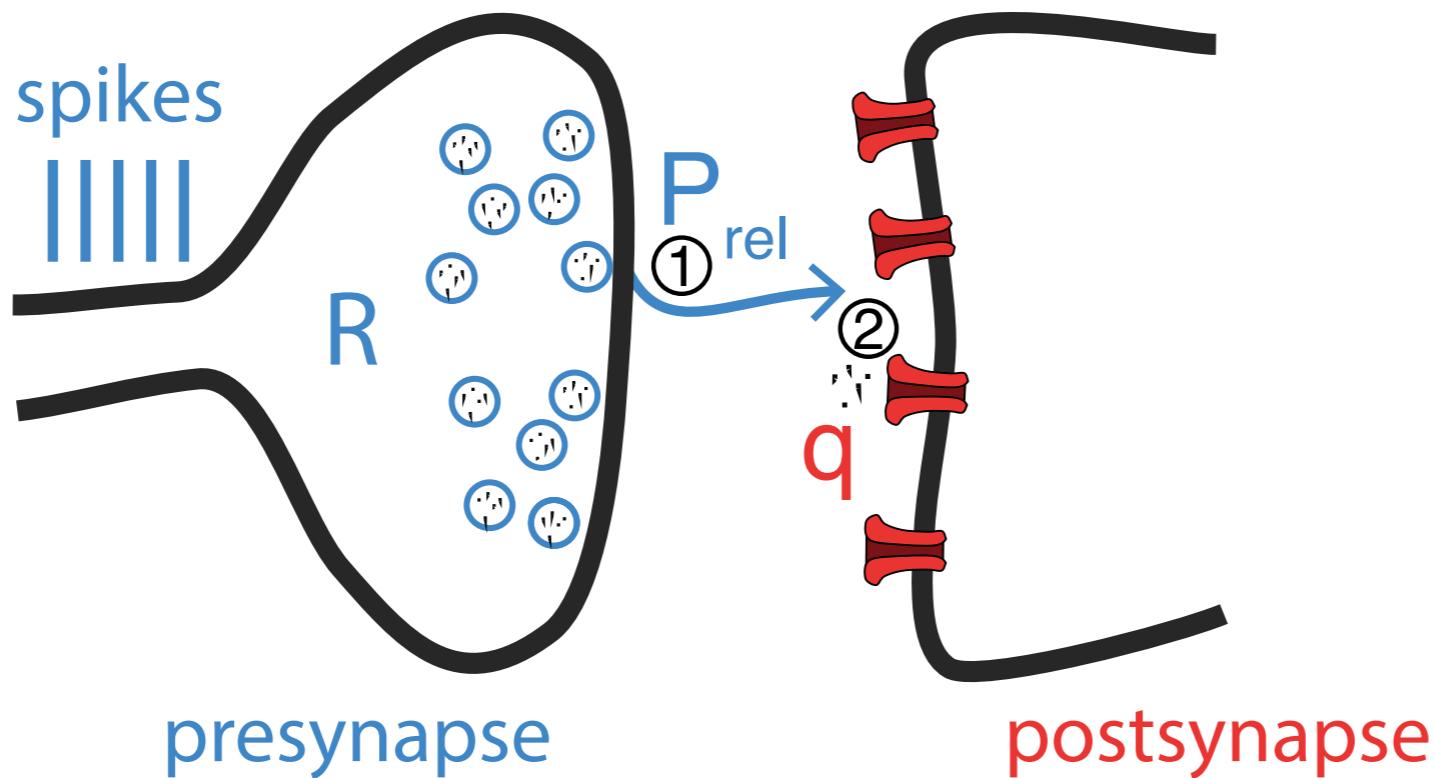
$$\text{Average release} = R \times P_{\text{rel}}$$

Tsodyks, Markram et al. PNAS 1997/1998;
Regehr CSH 2012;
Costa et al. Frontiers 2013

Note:

R is the fraction of vesicles available per release site.
Here we assume the number of release sites to be 1
(i.e. N=1), see previous lecture.

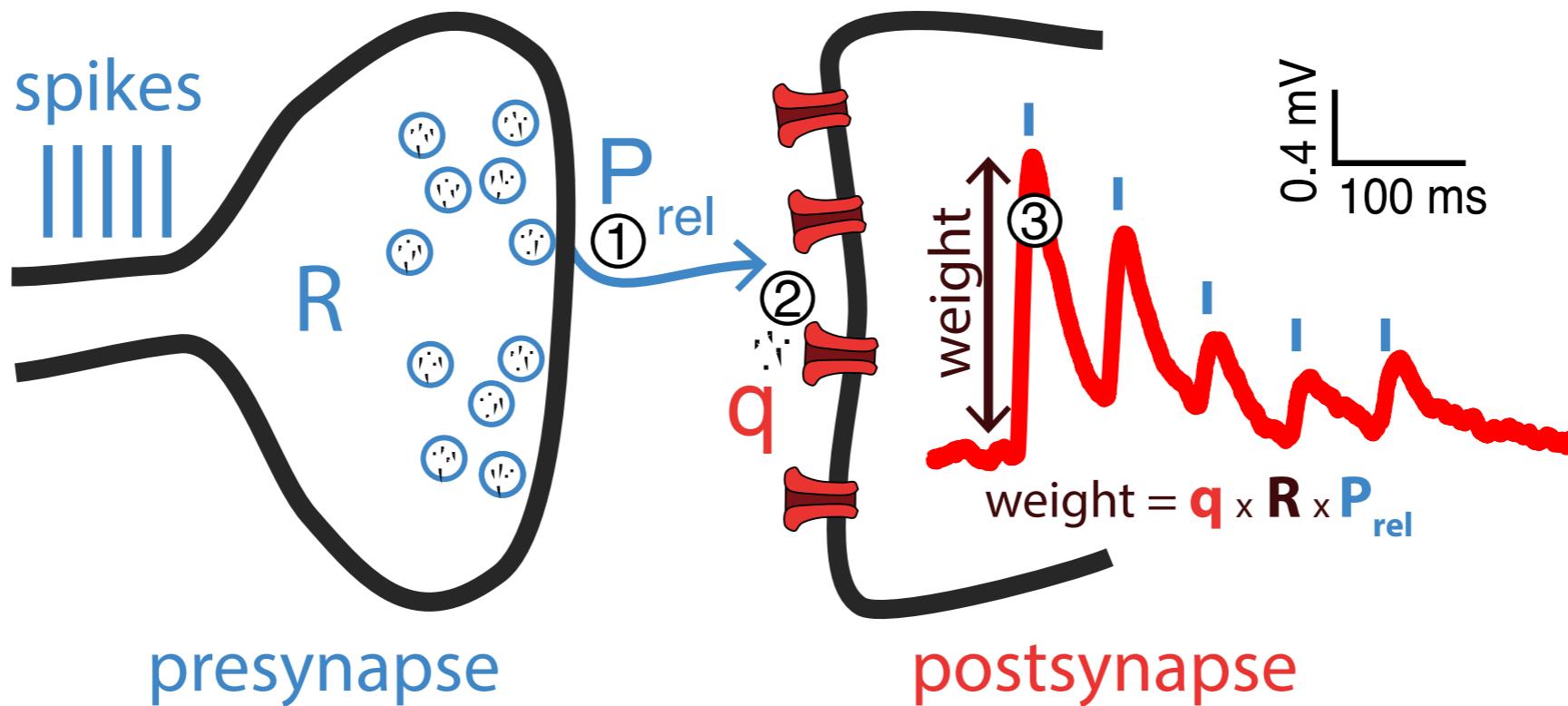
A phenomenological model of STD: Tsodyks-Markram model



2: Released neurotransmitters bind to postsynaptic receptors causing a response with amplitude q

Tsodyks, Markram et al. PNAS 1997/1998;
Regehr CSH 2012;
Costa et al. Frontiers 2013

A phenomenological model of STD: Tsodyks-Markram model

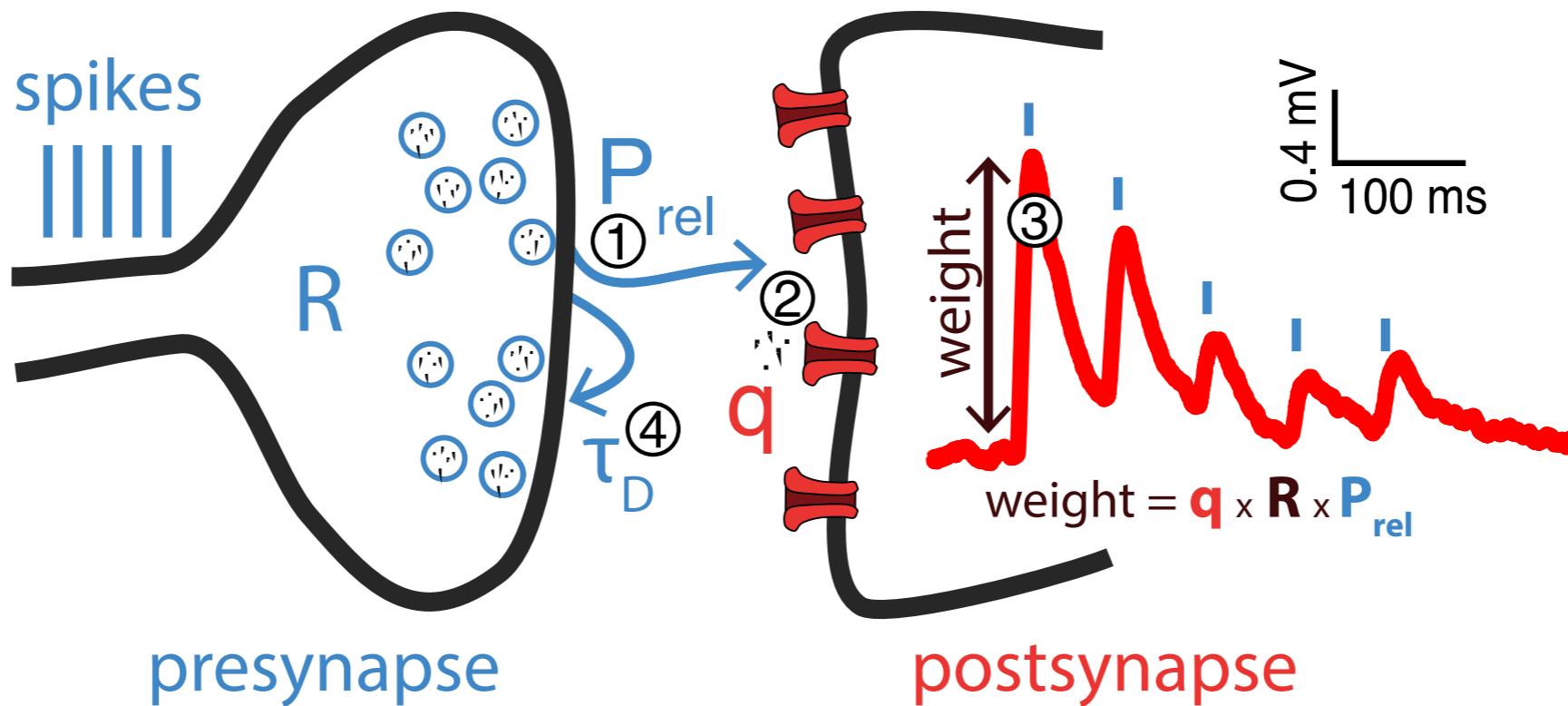


3: The final synaptic response (or weight) is then equal to $q \times R \times P_{rel}$

Tsodyks, Markram et al. PNAS 1997/1998;
Regehr CSH 2012;
Costa et al. Frontiers 2013

Note: This model ignores the stochastic nature of synapses and different release sites (N in the previous lecture).

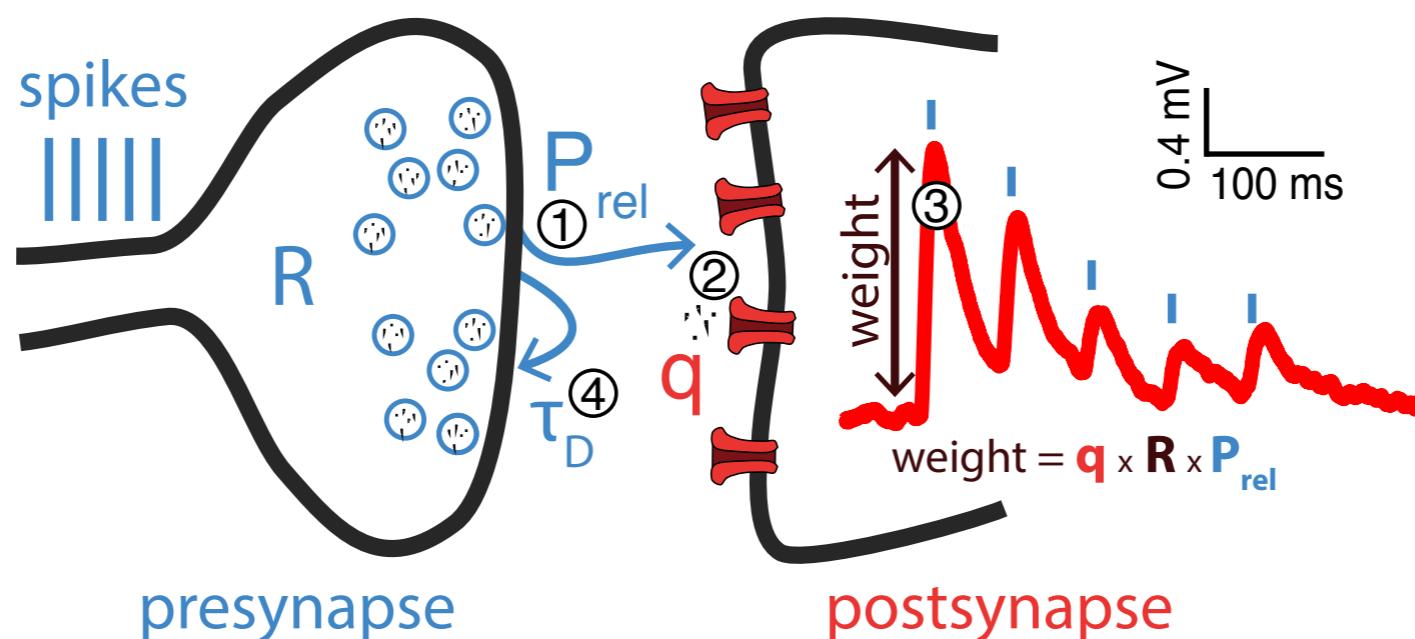
A phenomenological model of STD: Tsodyks-Markram model



4: At the same vesicles are recycled with a time constant **τ_D** , which leads to short-term depression before the vesicles (**R**) are fully recovered

Tsodyks, Markram et al. PNAS 1997/1998;
Regehr CSH 2012;
Costa et al. Frontiers 2013

A phenomenological model of STD: Tsodyks-Markram model

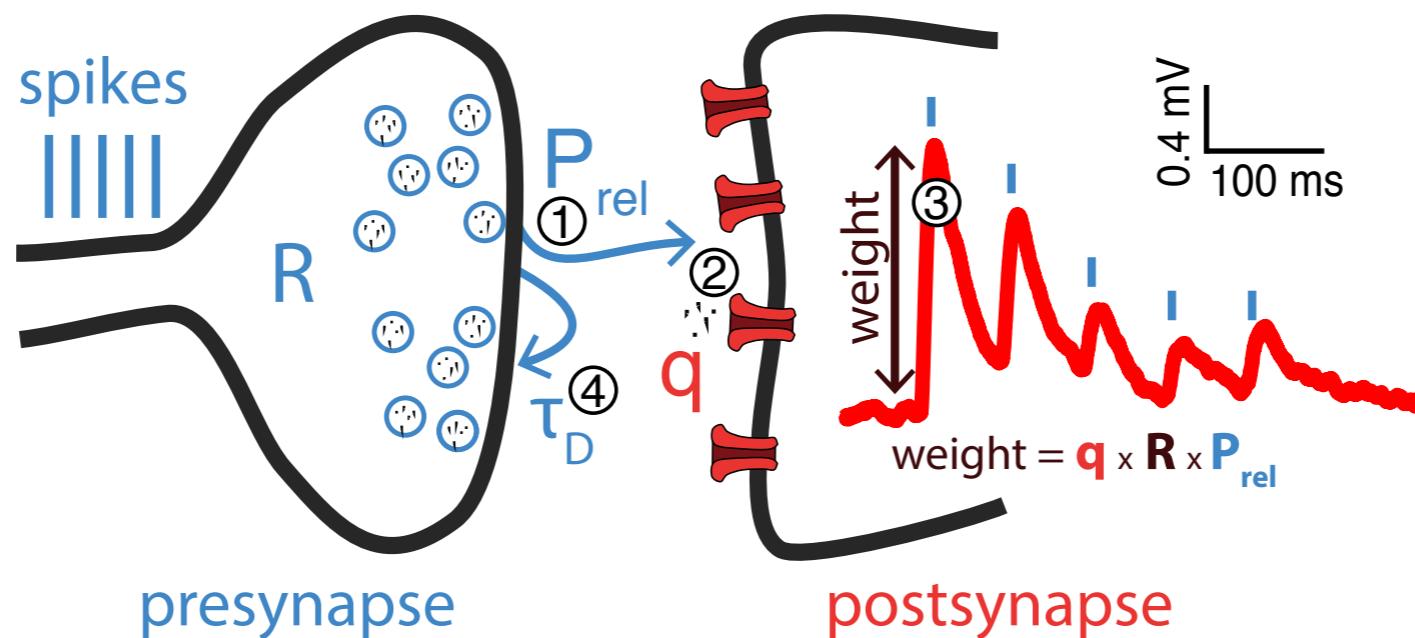


Differential equation that **models the number of vesicles available**:

$$\frac{dR}{dt} = -RP_r\delta_{\text{spike}}$$

Tsodyks, Markram et al. PNAS 1997/1998;

A phenomenological model of STD: Tsodyks-Markram model



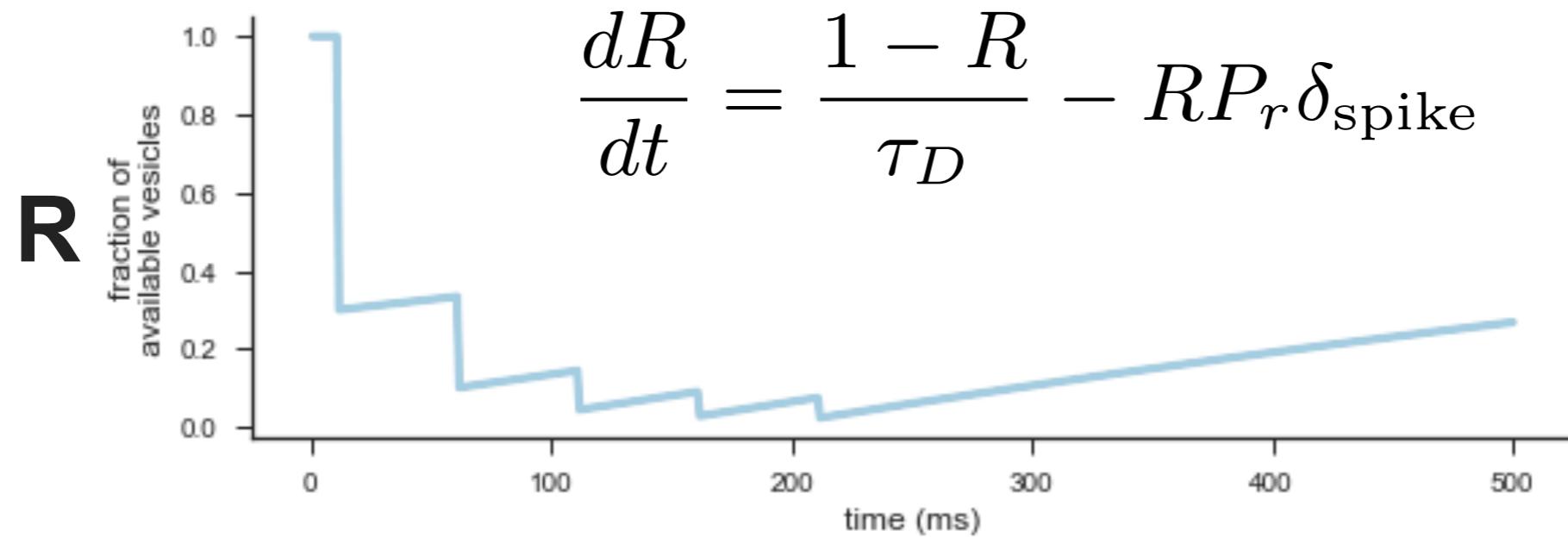
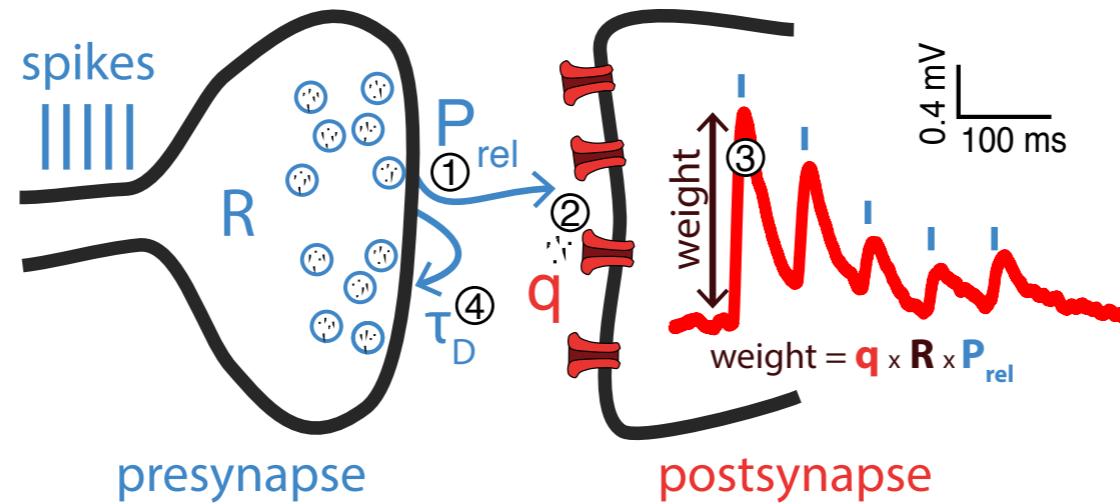
Differential equation that **models the number of vesicles available**:

$$\frac{dR}{dt} = \frac{1 - R}{\tau_D} - RP_r \delta_{\text{spike}}$$

Two parameters: **τ_D** and **P_r**

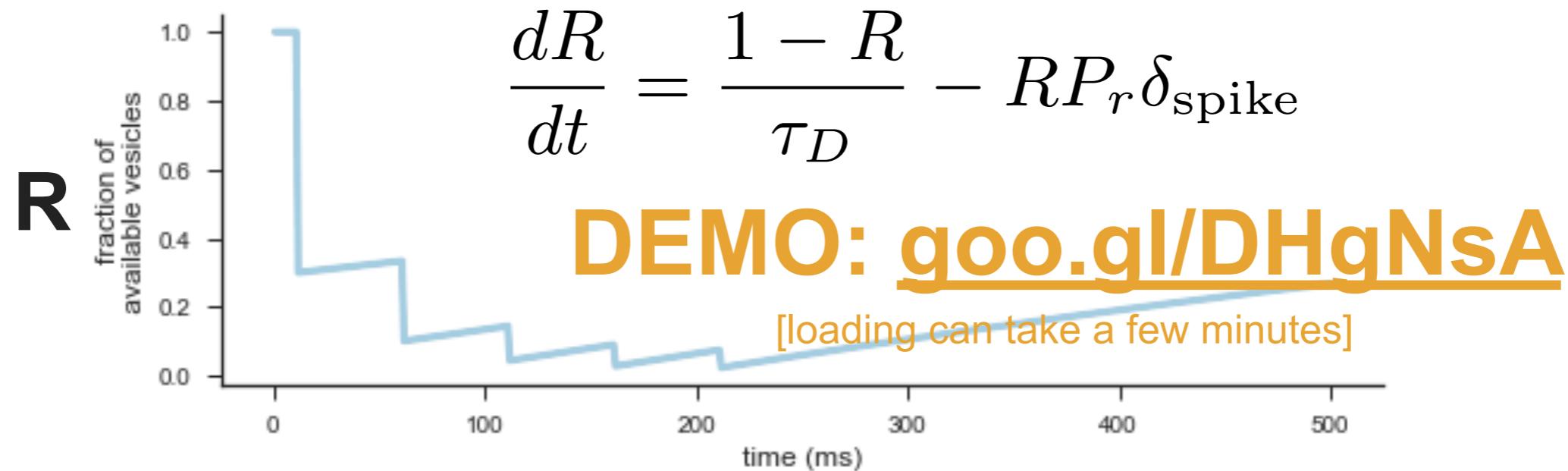
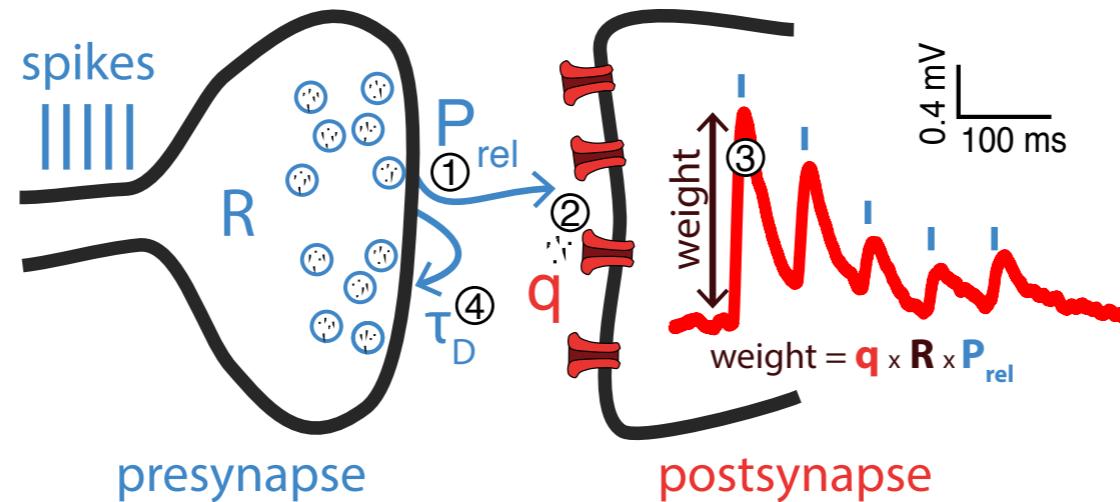
Tsodyks, Markram et al. PNAS 1997/1998;

A phenomenological model of STD: Tsodyks-Markram model



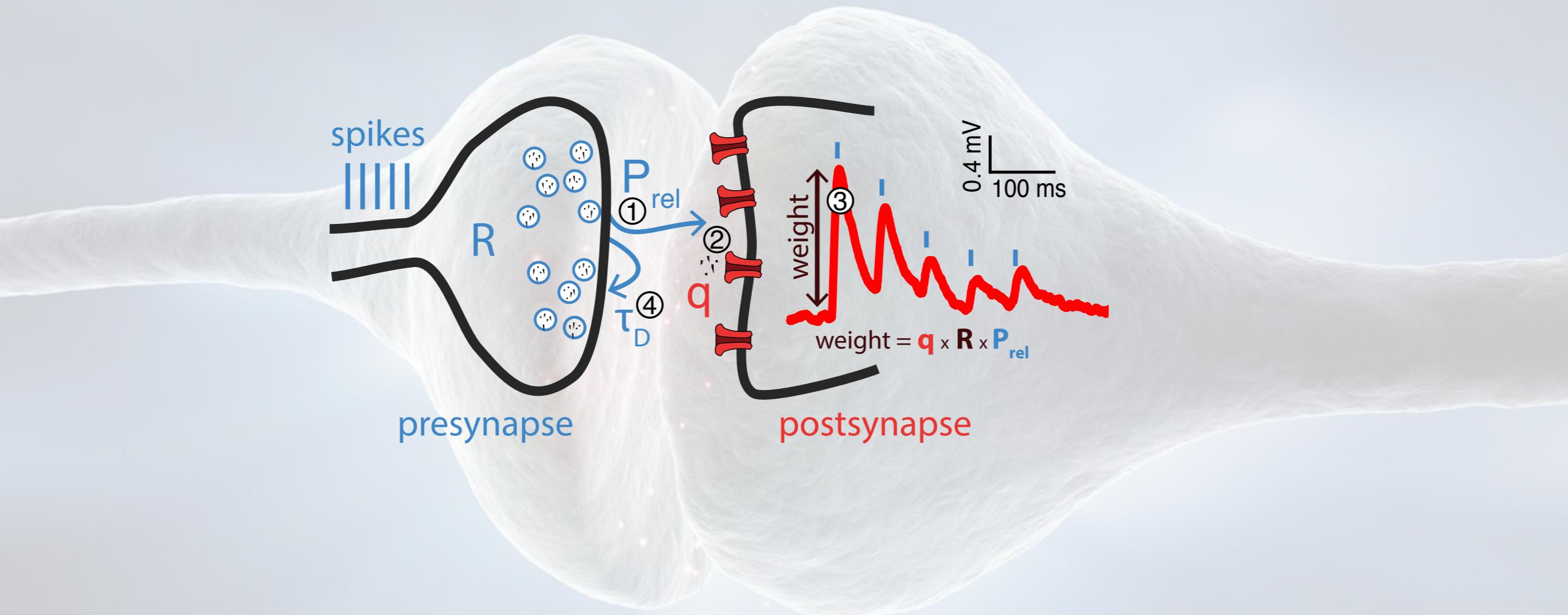
Tsodyks, Markram et al. PNAS 1997/1998;

A phenomenological model of STD: Tsodyks-Markram model

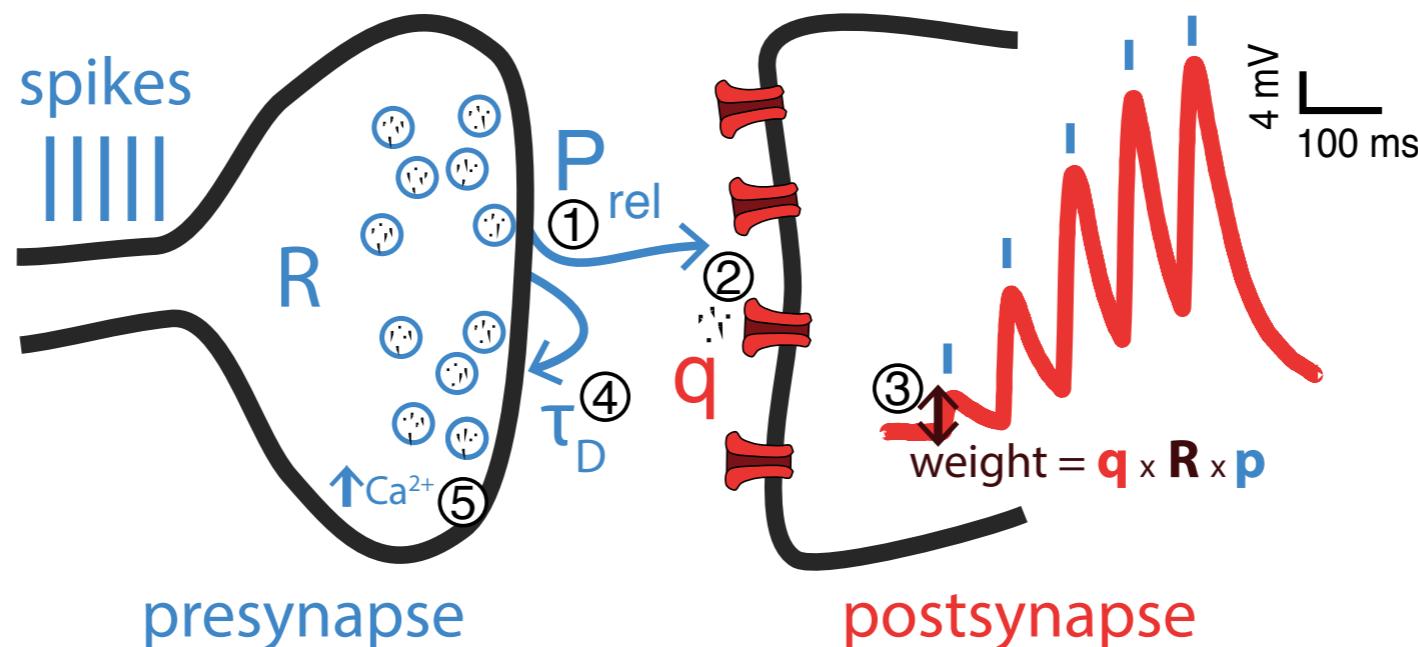


Tsodyks, Markram et al. PNAS 1997/1998;

Questions?



A phenomenological model of STF: Tsodyks-Markram model with facilitation



5: Calcium build up on the presynaptic side, leads to increased release probability p

Number of release vesicles:

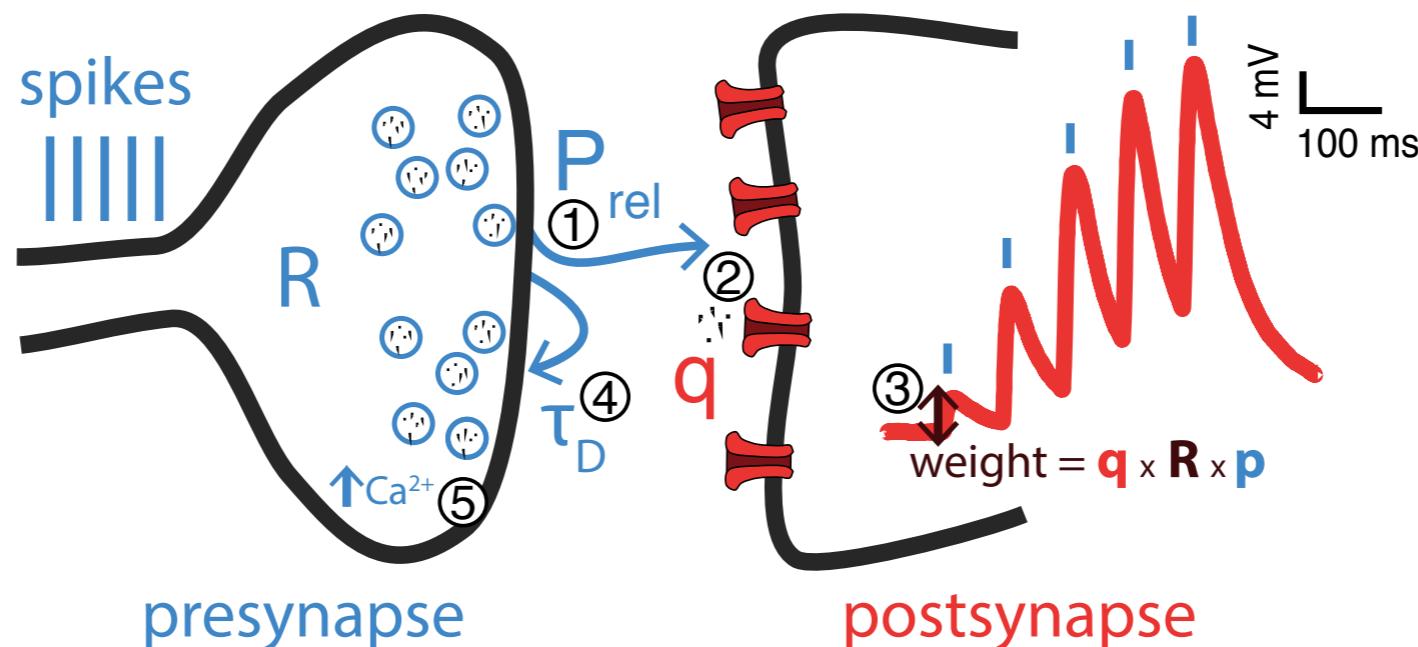
$$\frac{dR}{dt} = \frac{1 - R}{\tau_D} - Rp\delta_{\text{spike}}$$

Dynamic release probability:

$$\frac{dp}{dt} = + (1 - p)\delta_{\text{spike}}$$

Tsodyks, Markram et al. PNAS 1997/1998;

A phenomenological model of STF: Tsodyks-Markram model with facilitation



5: Calcium build up on the presynaptic side, leads to increased release probability p

Number of release vesicles:

$$\frac{dR}{dt} = \frac{1 - R}{\tau_D} - Rp\delta_{\text{spike}}$$

Dynamic release probability:

$$\frac{dp}{dt} = \frac{P_r - p}{\tau_F} + (1 - p)\delta_{\text{spike}}$$

Tsodyks, Markram et al. PNAS 1997/1998;

Group discussion

[groups of 2-3 (5 min)]

- What benefits might *short-term plasticity* have for the brain?

Hint: Remember that the brain is a dynamical system!

Outline

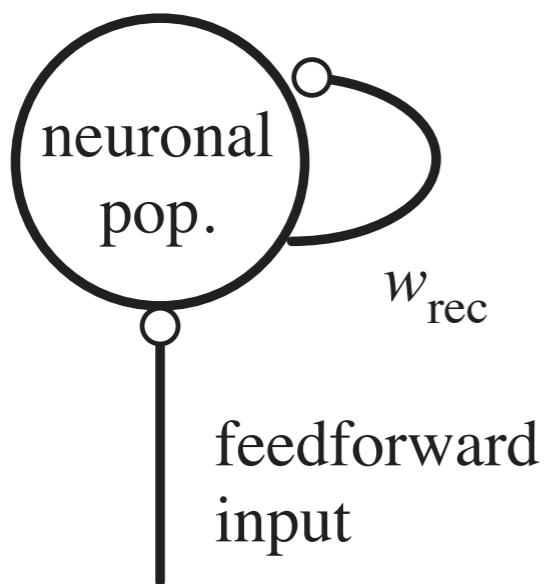
Short-term synaptic plasticity (STP)

1. The two main types
2. Models: Tsodyks-Markram and other models
3. Functions
 - 3.1. Network dynamics
 - 3.2. Theory of working memory

Summary: short-term plasticity (part I)

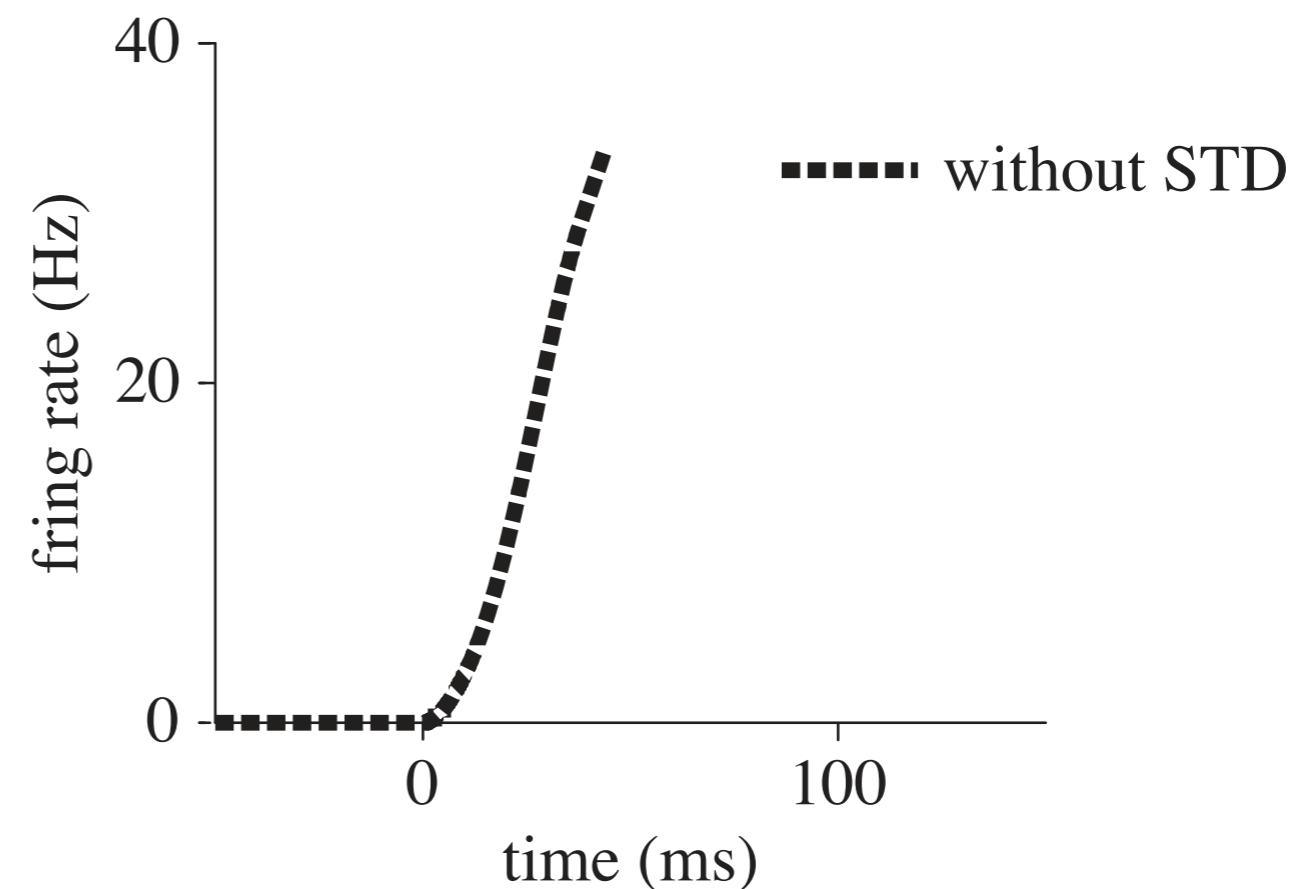
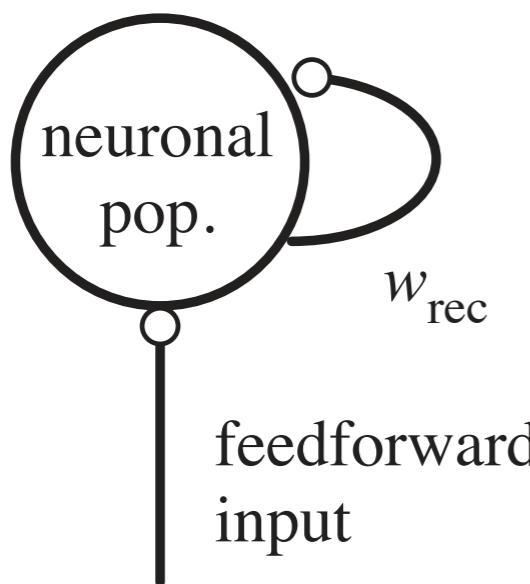
1. Lasts for up to tens of seconds
2. Two types: short-term depression and facilitation
3. Phenomenological and molecular models
4. Markram-Tsodyks model is a popular model with versions that can capture STD and STF

Short-term depression helps prevent seizure-like dynamics



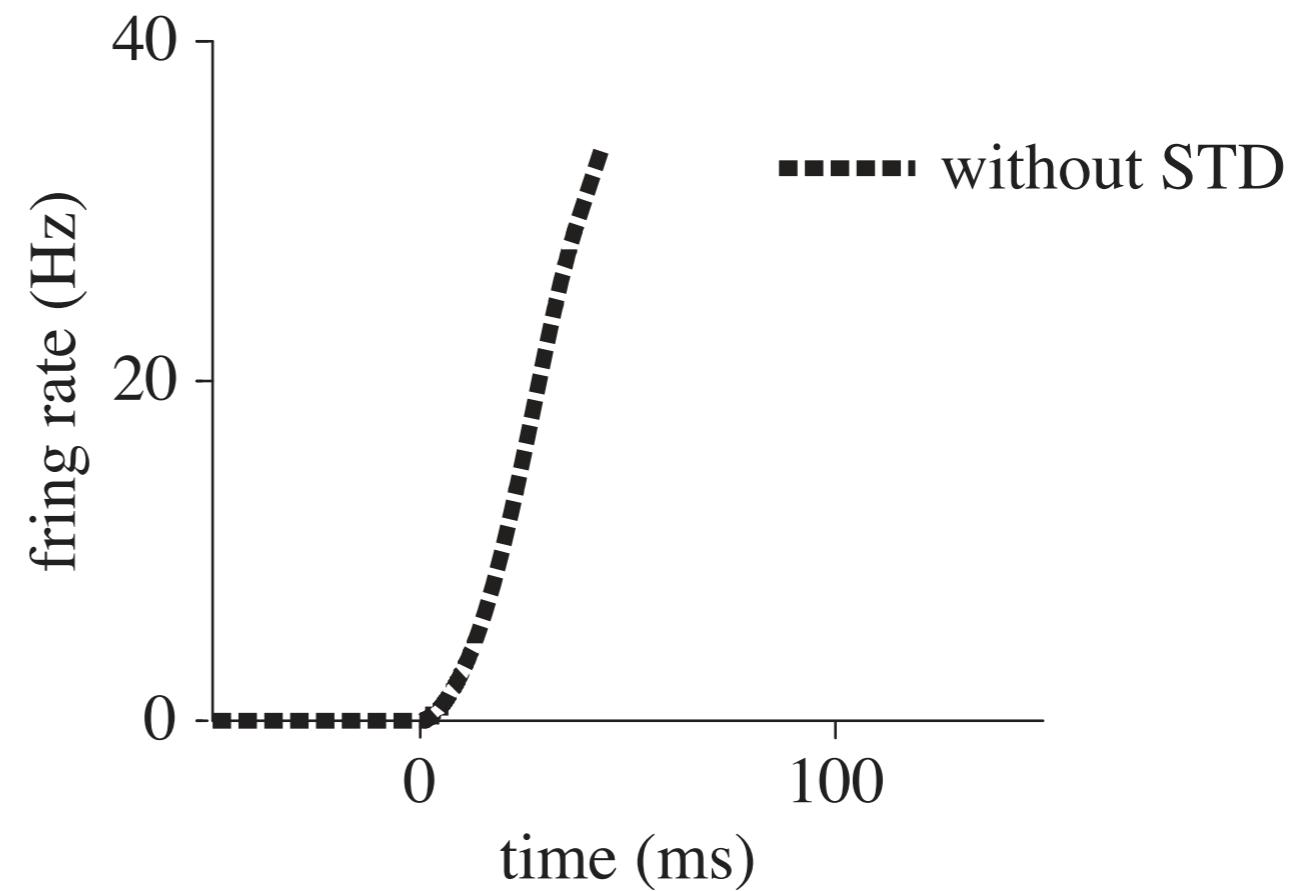
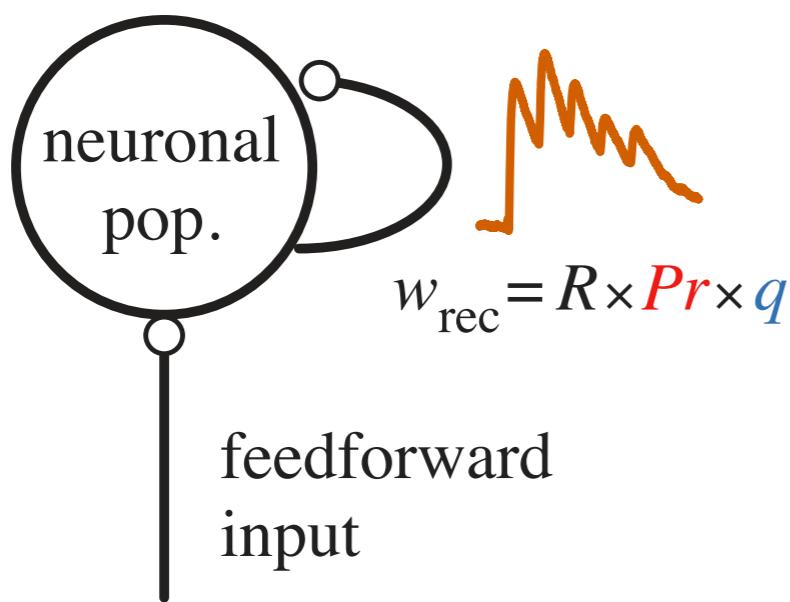
Costa et al. Phil. Trans. R. Soc. 2017
Sussillo et al. JNeurophysiol. 2007
and several others..

Short-term depression helps prevent seizure-like dynamics



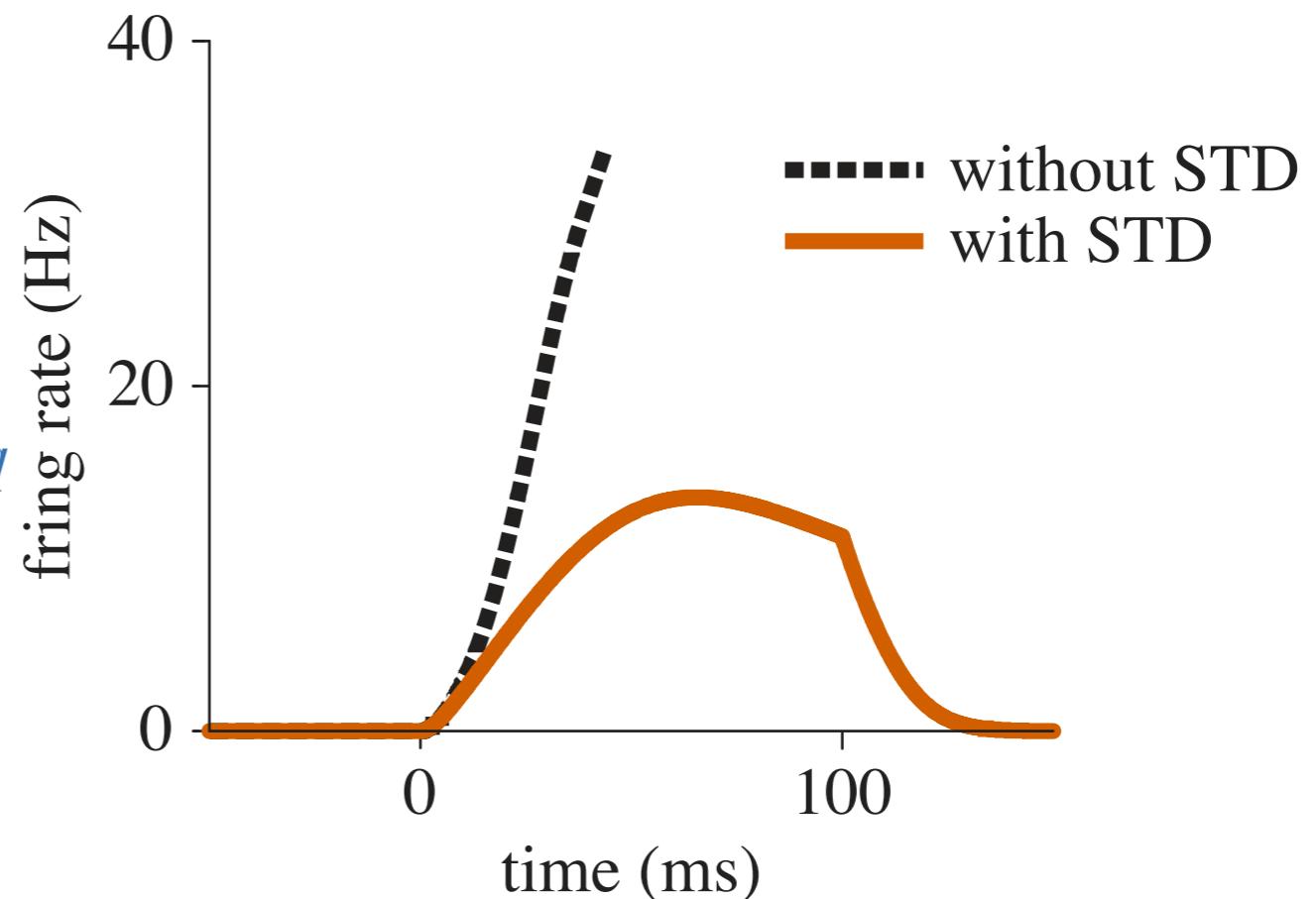
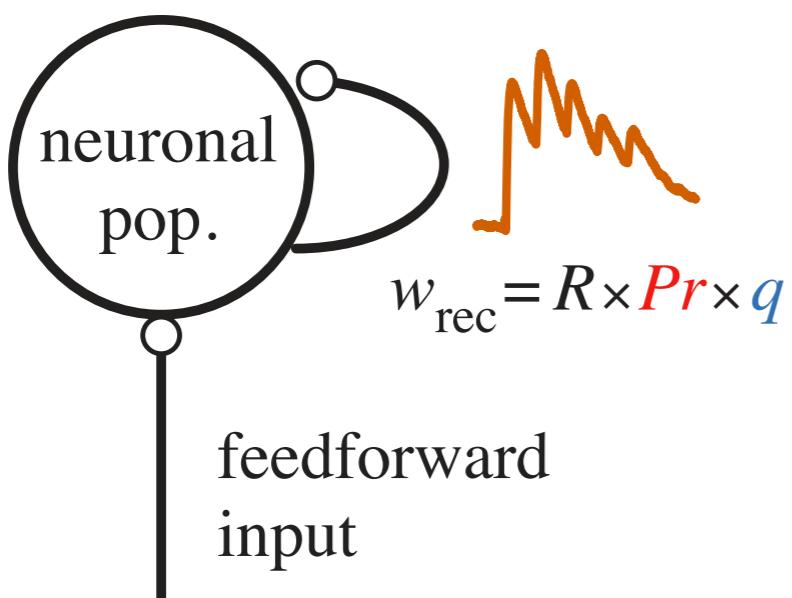
Costa et al. Phil. Trans. R. Soc. 2017
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Short-term depression helps prevent seizure-like dynamics



Costa et al. Phil. Trans. R. Soc. 2017
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Short-term depression helps prevent seizure-like dynamics



Costa et al. Phil. Trans. R. Soc. 2017
Sussillo et al. JNeurophysiol. 2007
and several others..

Short-term facilitation as a theory of working memory

Working memory is our ability to keep an item in memory for a short period of time, with very limited capacity (usually up to 5-10 different items).

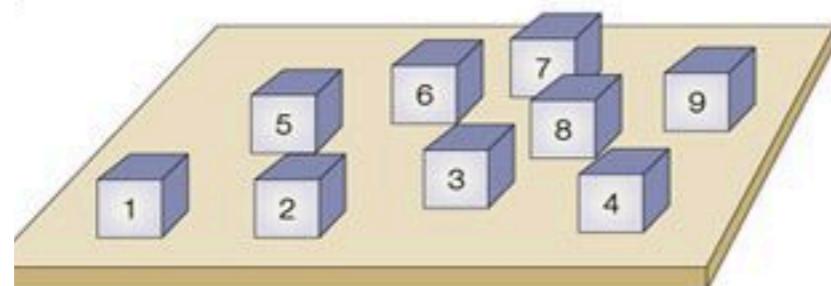
Note: Working memory is key for reasoning: our ability to link different sets of information together.

Examples: Remembering a phone number, while driving.

Where are all your lectures for the day?

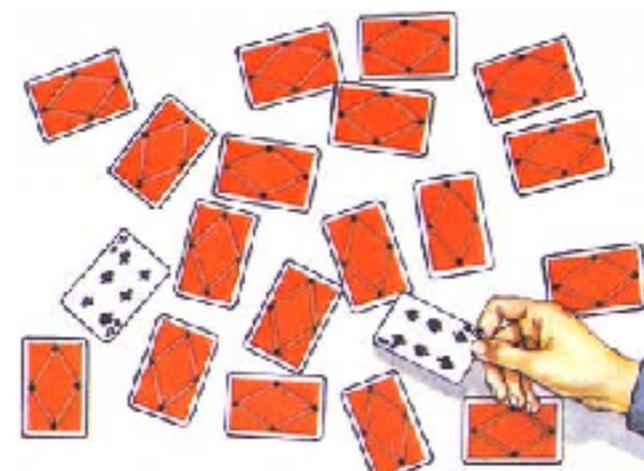
Or these particular tasks:

Corsi block tapping task:
recall the order of tapping on the blocks

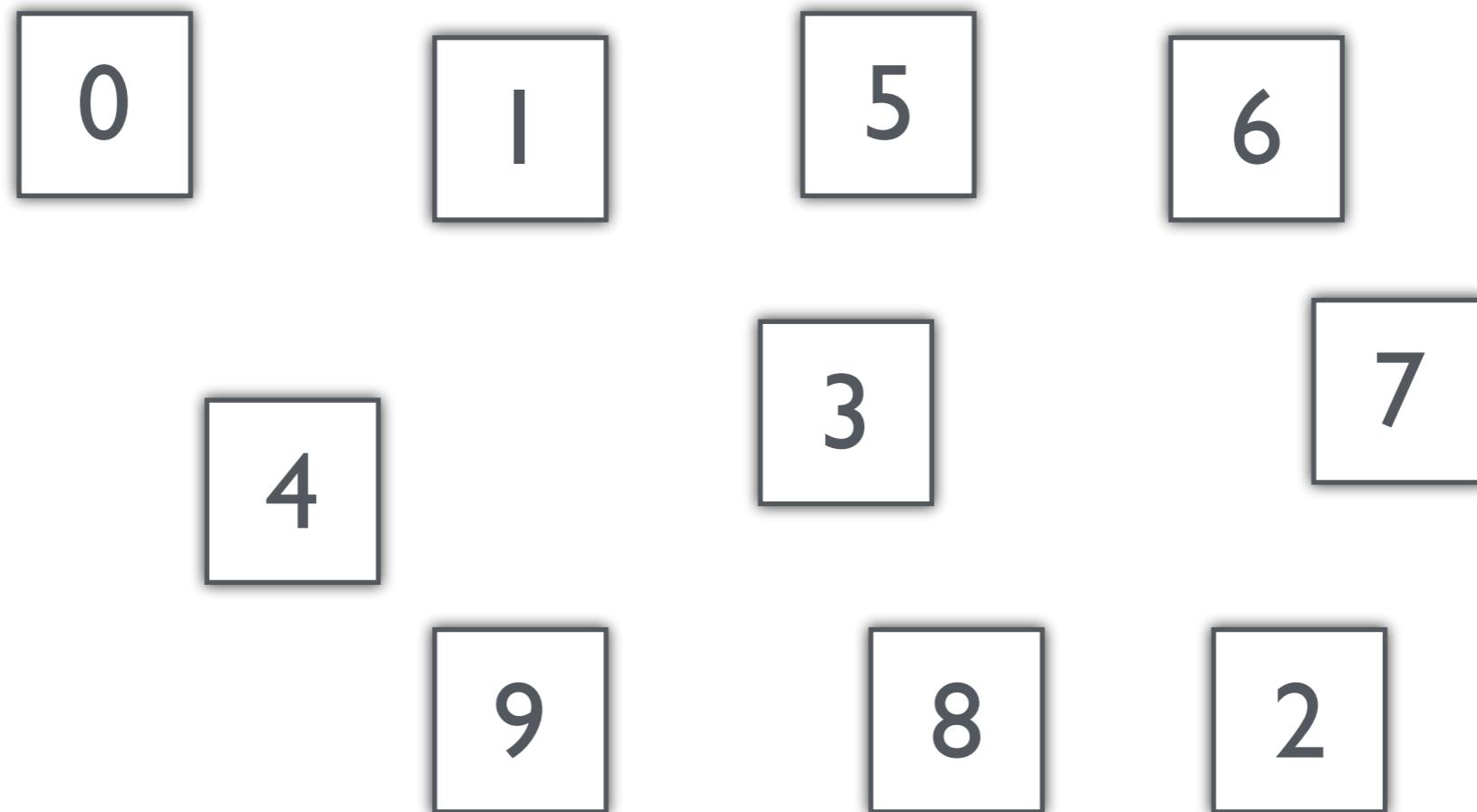


Baddeley, Nature Rev Neurosci 2003

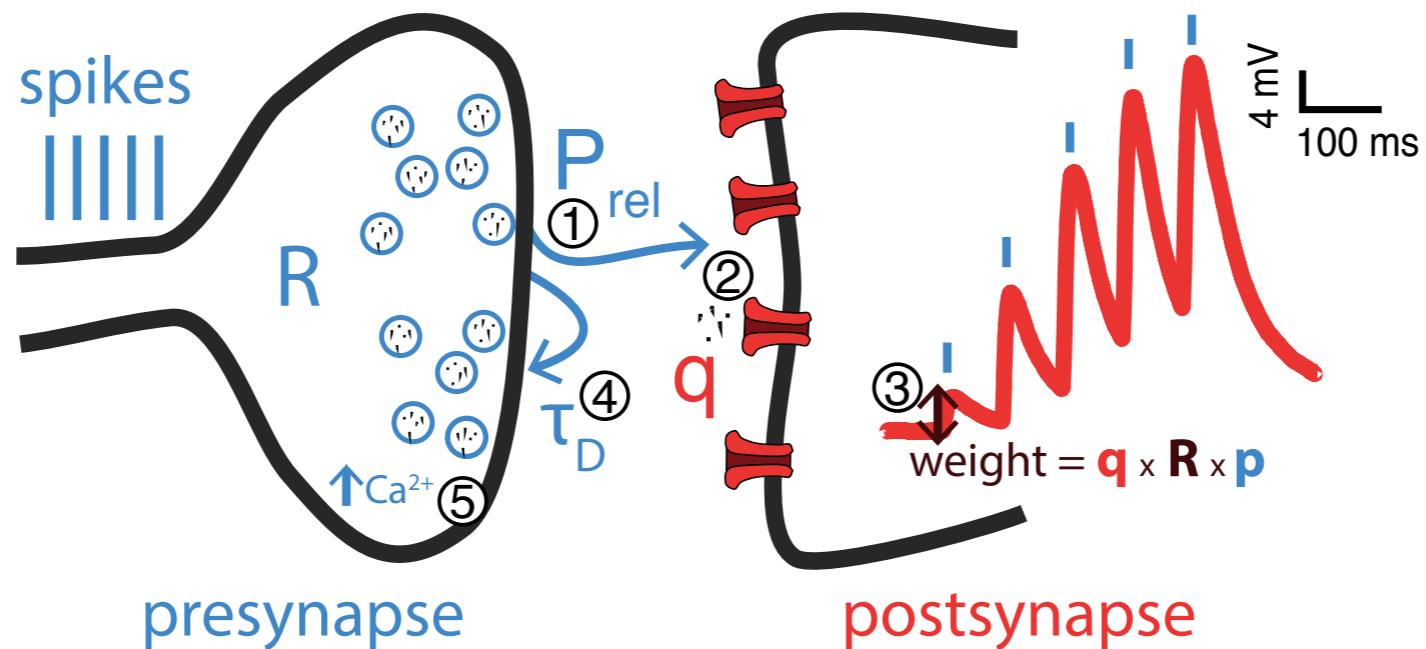
Card matching game:



Lets play block 'tapping'



Short-term facilitation as a theory of working memory



Key idea: Use slow decay in the release probability dynamics p to remember previous inputs.

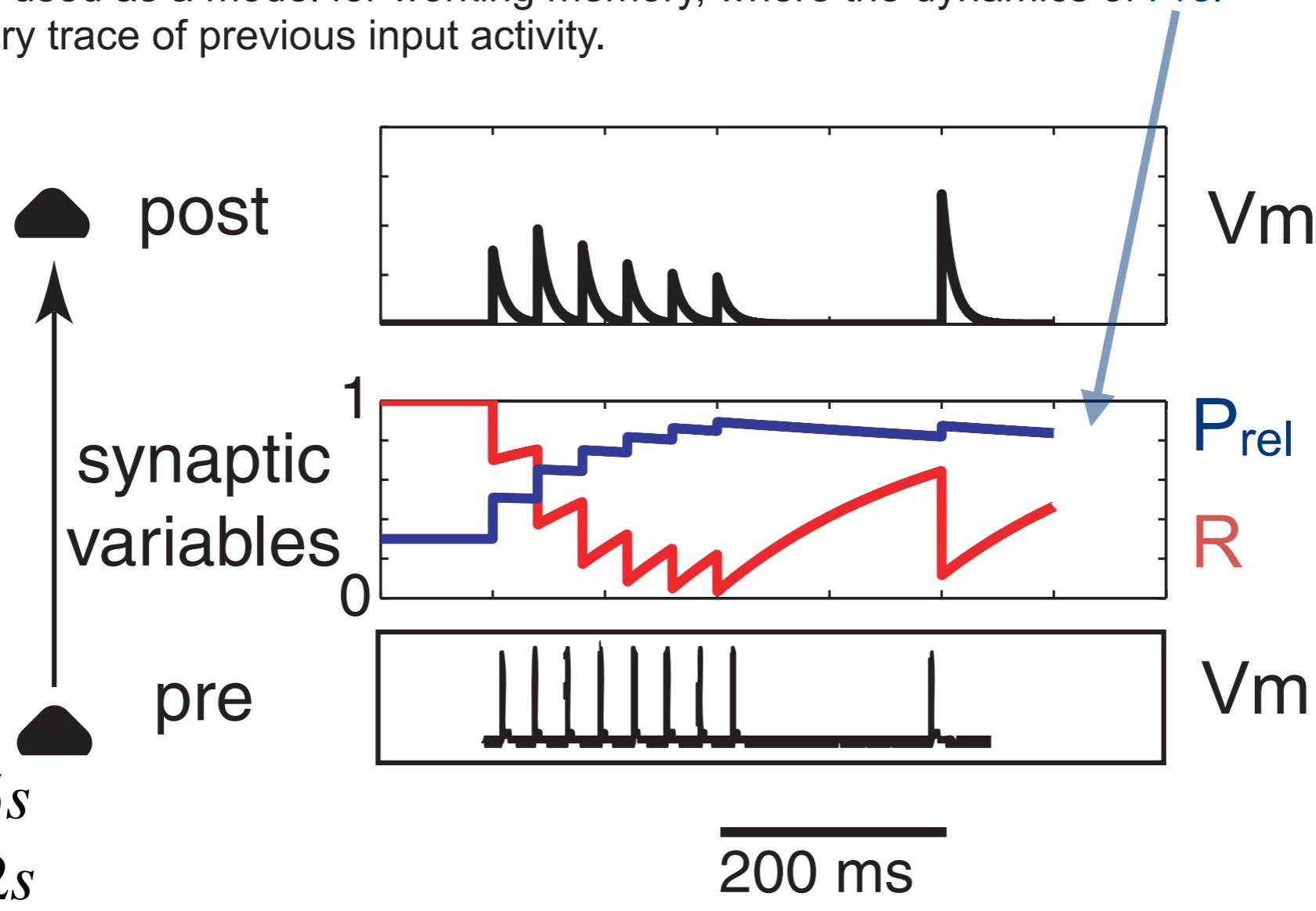
$$\frac{dR}{dt} = \frac{1 - R}{\tau_D} - Rp\delta_{\text{spike}}$$

$$\frac{dp}{dt} = \frac{P_r - p}{\tau_F} + (1 - p)\delta_{\text{spike}}$$

$$\tau_F \gg \tau_D$$

Short-term facilitation as a synaptic theory of working memory

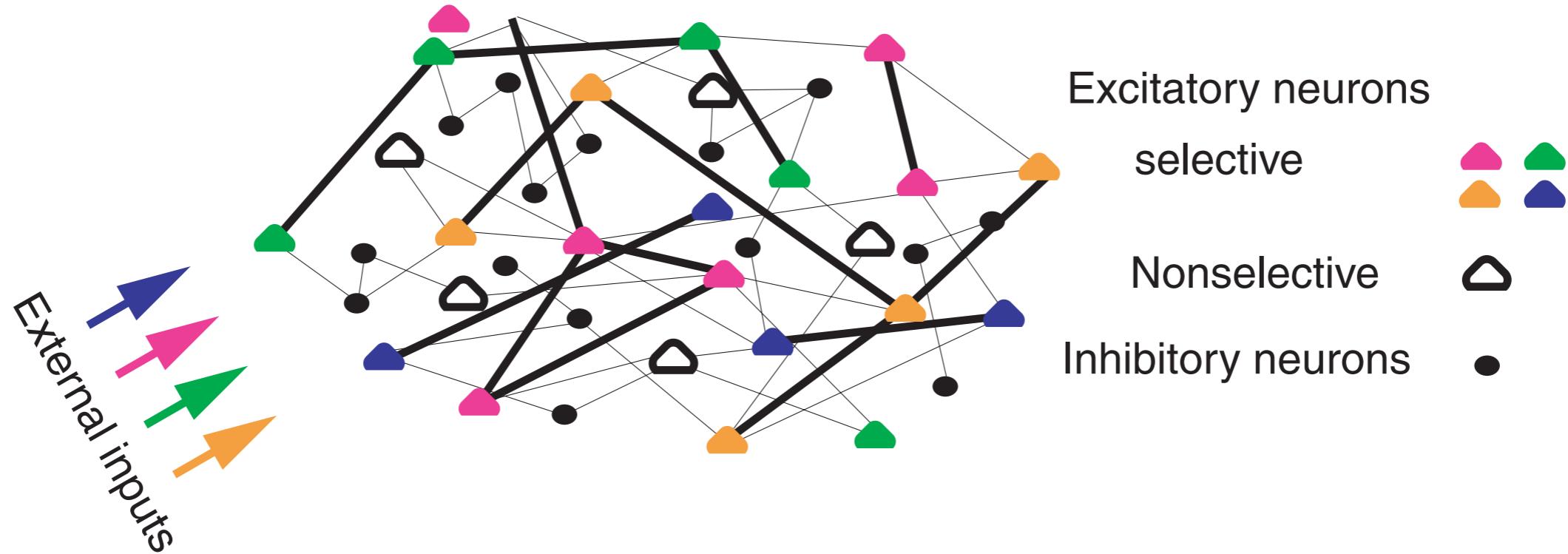
Here STF is used as a model for working memory, where the dynamics of P_{rel} are a memory trace of previous input activity.



Mongillo et al. Science 2008

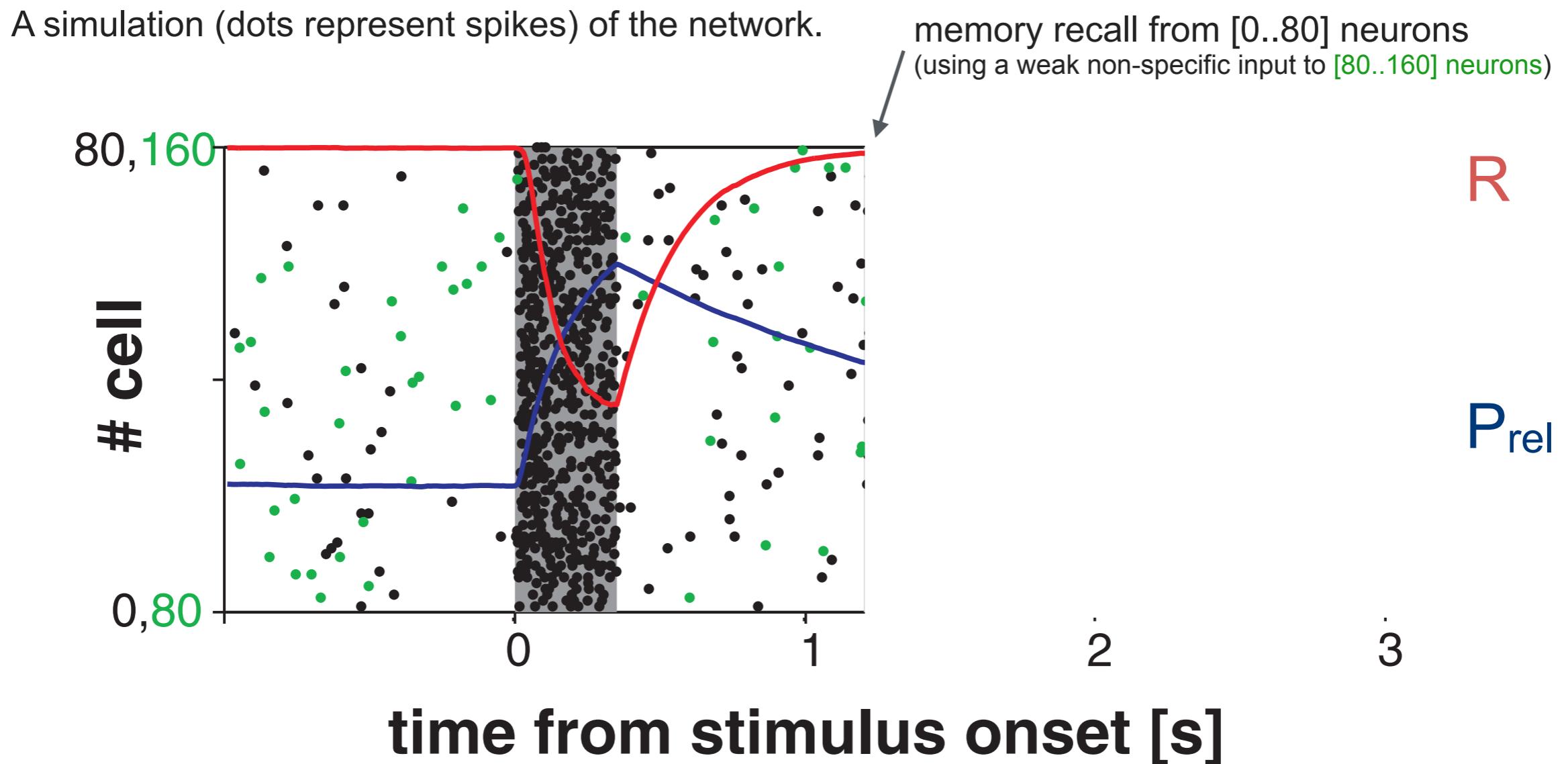
Short-term facilitation as a synaptic theory of working memory

A recurrent neural network of integrate and fire neurons was used to demonstrate the idea.



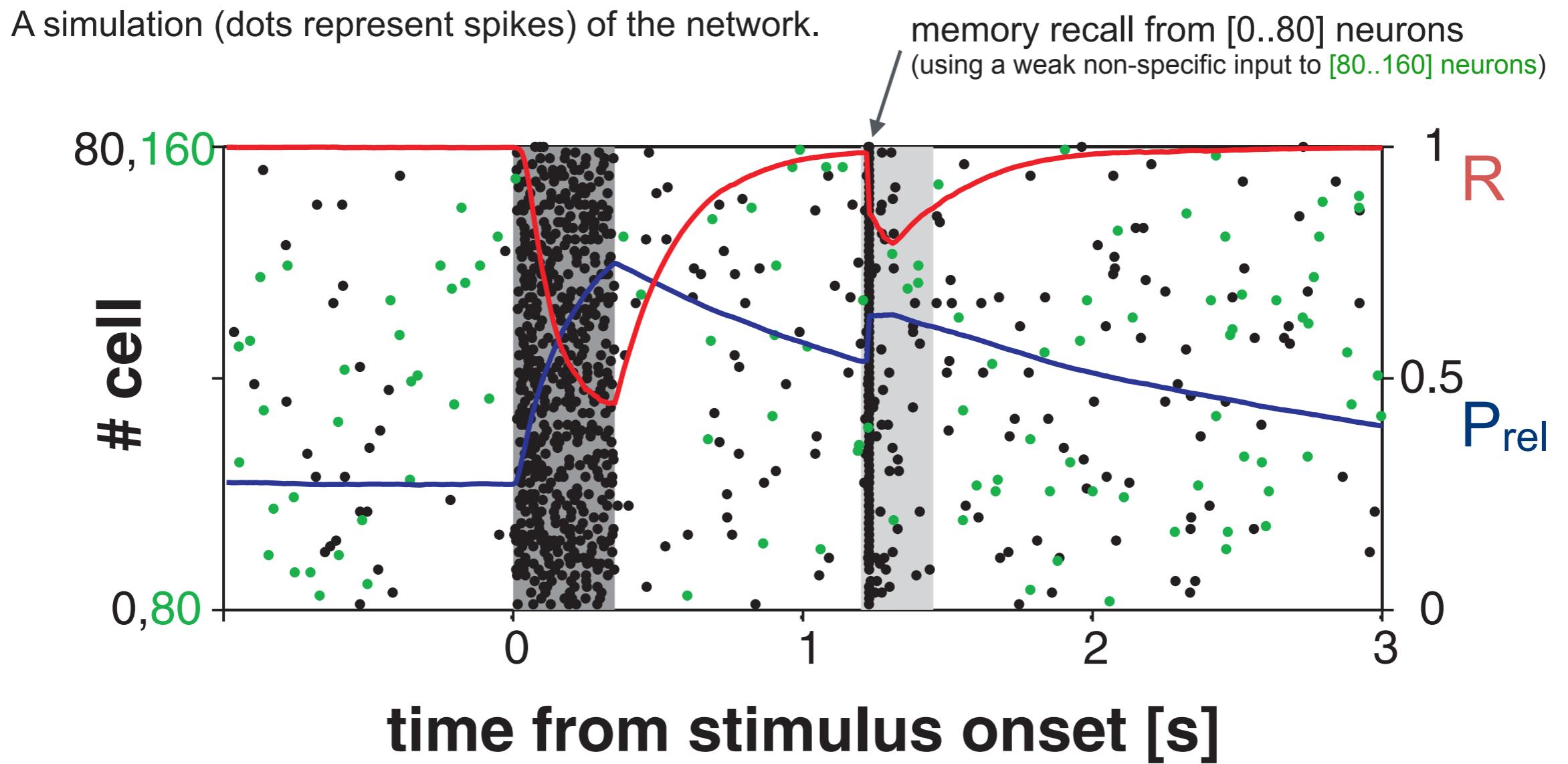
Mongillo et al. Science 2008

Short-term facilitation as a synaptic theory of working memory



Mongillo et al. Science 2008

Short-term facilitation as a synaptic theory of working memory



Note: This suggests that every one of the **1000 trillion synapses** in our brains has some form of working memory, and therefore contribute to reasoning!

Can you recall the number?

0

1

5

6

4

3

7

9

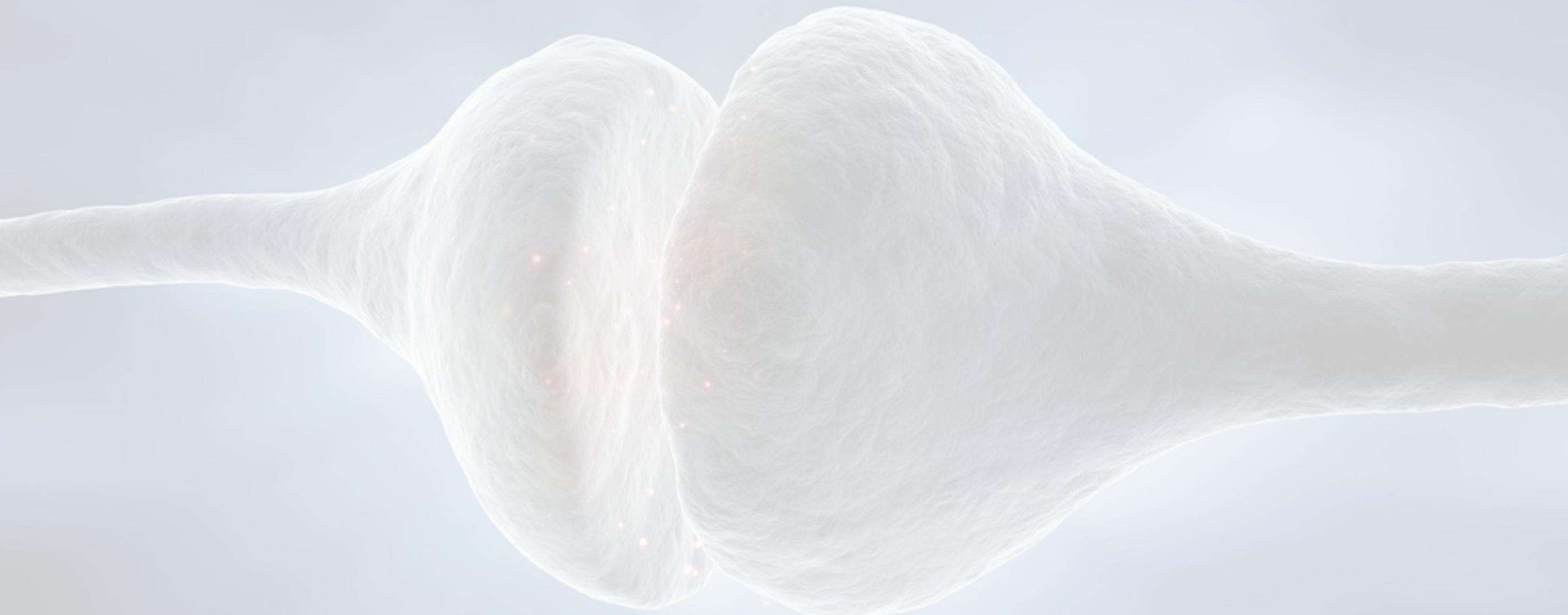
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2

Summary: short-term plasticity (part 2)

1. STD is important to *control network activity*, helping to prevent pathological states
2. STF has been proposed a new (exciting) theory of how we hold items in memory for short periods of time (working memory)

Questions?



References

Text books:

General biology: Wikipedia/Principles of neuroscience [neuroscience bible]

Computational Neuroscience: Neuronal Dynamics by Gerstner, Kistler, Naud and Paninski

Computational Neuroscience: Principles of Computational Modelling in Neuroscience by Sterratt, Gillies, Graham and Willshaw

[Chapter on Synaptic modelling: http://dai.fmph.uniba.sk/courses/comp-neuro/reading/Sterratt_CH7_synapse.pdf]

And the papers referred to during the lecture.