

Optimal synaptic strategies for different timescales of memory

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What is a synapse?

What is a synapse?

Theorists

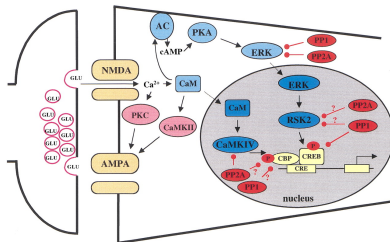
$$W_{ij}$$

What is a synapse?

Theorists

$$W_{ij}$$

Experimenters



[Klann (2002)]

Storage capacity of synaptic memory

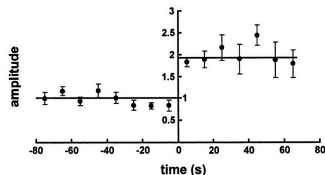
Hopfield, perceptron have capacity $\propto N$, ($\#$ synapses).

Assumes unbounded analog synapses

With discrete, finite synapses:

\implies memory capacity $\sim \mathcal{O}(\log N)$.

[Amit and Fusi (1992), Amit and Fusi (1994)]



[Petersen et al. (1998), O'Connor et al. (2005)]

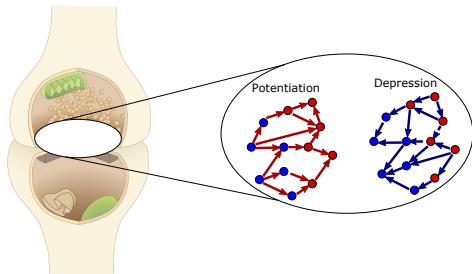
New memories overwrite old \implies stability-plasticity dilemma.

Models of complex synaptic dynamics



Models of complex synaptic dynamics

- Internal functional state of synapse \rightarrow synaptic weight. ● weak
- Candidate plasticity events \rightarrow transitions between states ● strong

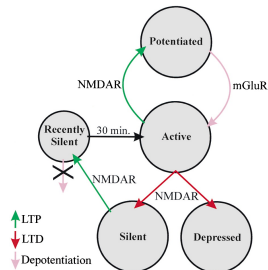
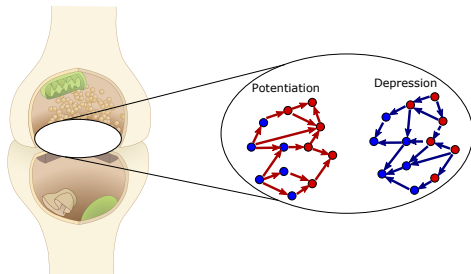


States: #AMPA, #NMDAR, NMDAR subunit composition,
CaMK II autophosphorylation, activating PKC, p38 MAPK,...

[Fusi et al. (2005), Fusi and Abbott (2007), Barrett and van Rossum (2008)]

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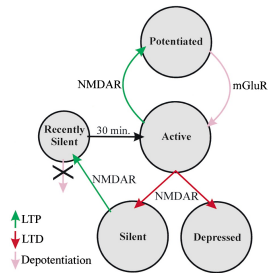
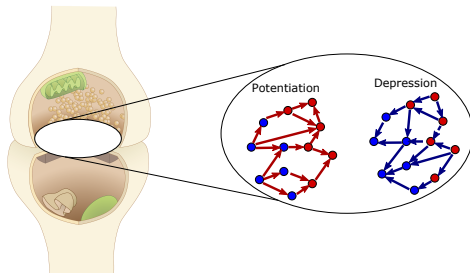
[Montgomery and Madison (2002)]

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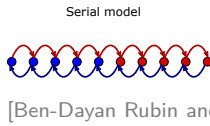
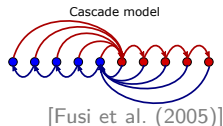
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Models of complex synaptic dynamics

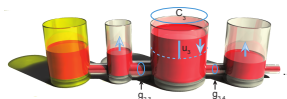
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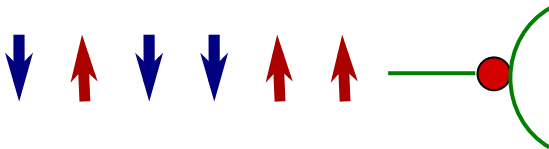
[Montgomery and Madison (2002)]



Leibold and Kempter (2008)]

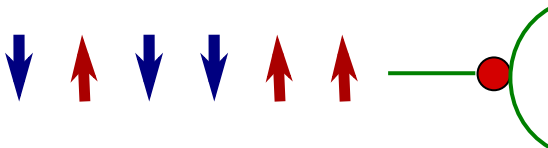


Synaptic memory curves

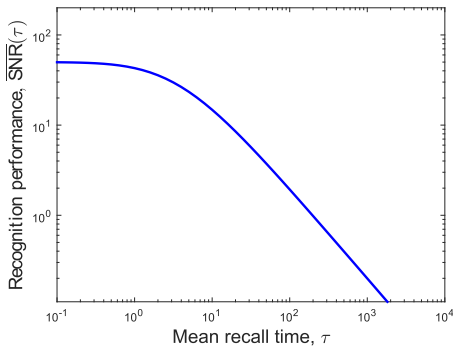


Synapses store a sequence of memories.

Synaptic memory curves



Synapses store a sequence of memories.

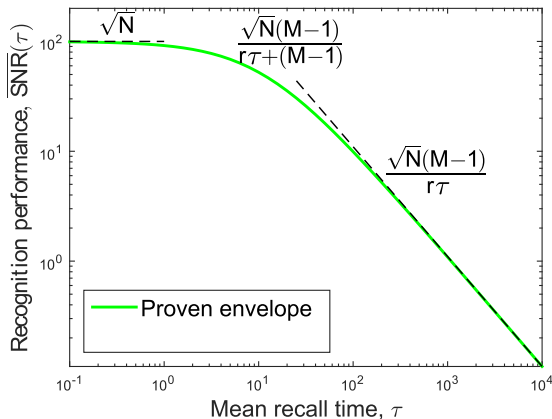


Questions

- What are the upper bounds?
- Which models achieve them?
- What are the theoretical principles behind them?

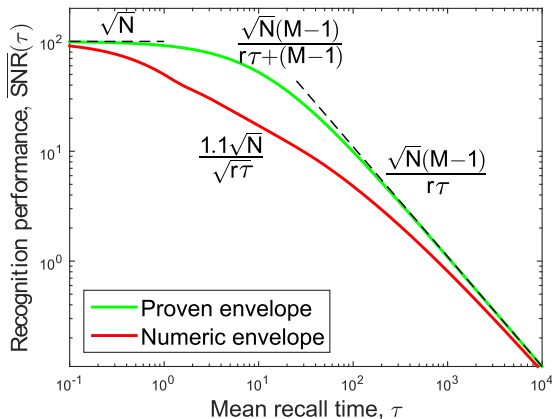
Proven envelope: memory frontier

Upper bound on memory curve at *any* timescale.

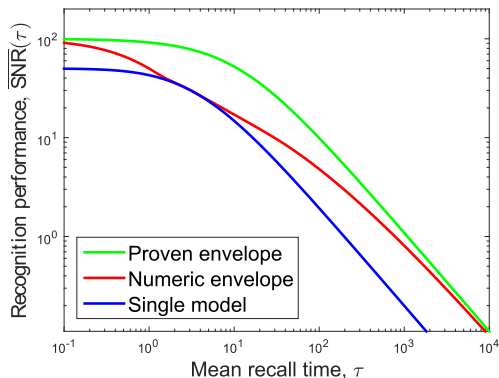


Proven envelope: memory frontier

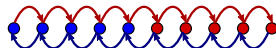
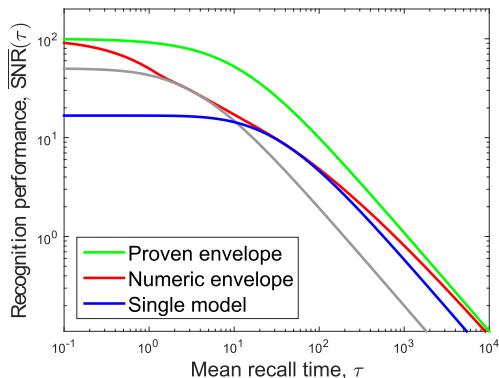
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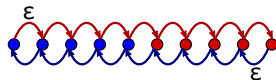
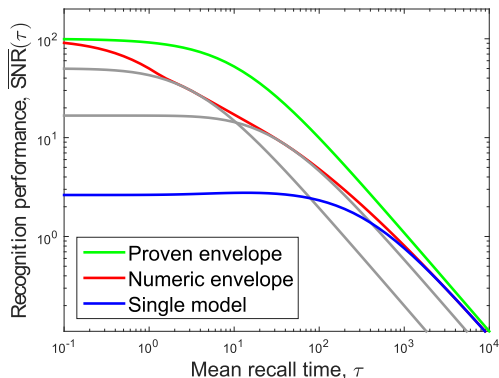
Models that maximize memory for one timescale



Models that maximize memory for one timescale

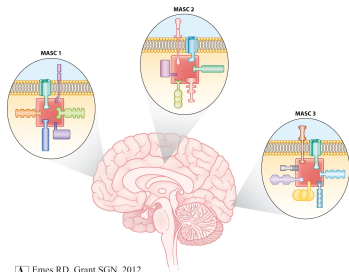


Models that maximize memory for one timescale



Synaptic diversity and timescales of memory

Different synapses have different molecular structures.

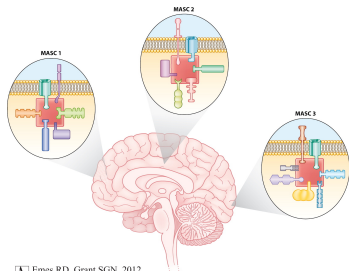


Emes RD, Grant SGN, 2012.
Annu. Rev. Neurosci. 35:111–31

[Emes and Grant (2012)]

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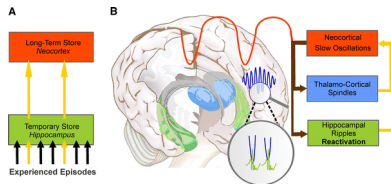
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[Emes and Grant (2012)]

Memories stored in different places for different timescales

[Squire and Alvarez (1995)]

[McClelland et al. (1995)]



[Born and Wilhelm (2012)]

Also: Cerebellar cortex → nuclei.

[Attwell et al. (2002)]

[Cooke et al. (2004)]

Synaptic structures for different timescales of memory

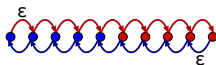
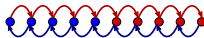
Short timescale



Intermediate timescale

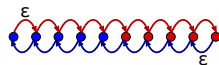


Long timescale



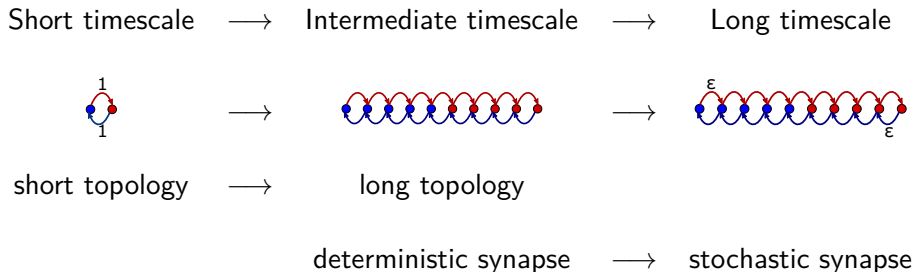
Synaptic structures for different timescales of memory

Short timescale \longrightarrow Intermediate timescale \longrightarrow Long timescale



short topology \longrightarrow long topology

Synaptic structures for different timescales of memory



Experimental tests?

Traditional experiments:



Experimental tests?

Traditional experiments:



Subject a synapse to a sequence of candidate plasticity events.
Observe the changes in synaptic efficacy.



- We have formulated a general theory of learning and memory with complex synapses.
- We find a memory envelope: a single curve that cannot be exceeded by the memory curve of *any* synaptic model.
- We understood which types of synaptic structure are useful for storing memories for different timescales.

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Technical detail: ordering states

Let \mathbf{T}_{ij} = mean first passage time from state i to state j . Then:

$$\eta = \sum_j \mathbf{T}_{ij} \mathbf{p}_j^\infty,$$

is independent of the initial state i (Kemeney's constant).

[Kemeny and Snell (1960)]

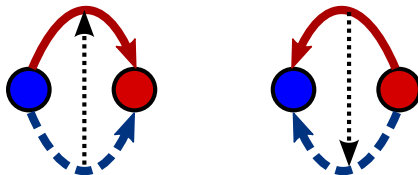
We define:

$$\eta_i^+ = \sum_{j \in \text{strong}} \mathbf{T}_{ij} \mathbf{p}_j^\infty, \quad \eta_i^- = \sum_{j \in \text{weak}} \mathbf{T}_{ij} \mathbf{p}_j^\infty.$$

They can be used to arrange the states in an order (increasing η^- or decreasing η^+). [back](#)

Technical detail: upper/lower triangular

With states in order:



Endpoint: potentiation goes right, depression goes left.

[back](#)