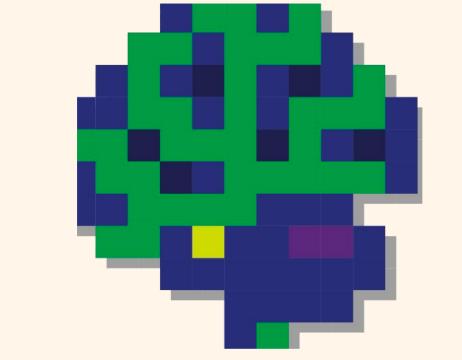
# INVESTIGATING HOW MEMORY REPRESENTATIONS CHANGE AS A FUNCTION OF COMPETITION-DEPENDENT LEARNING AND SLEEP



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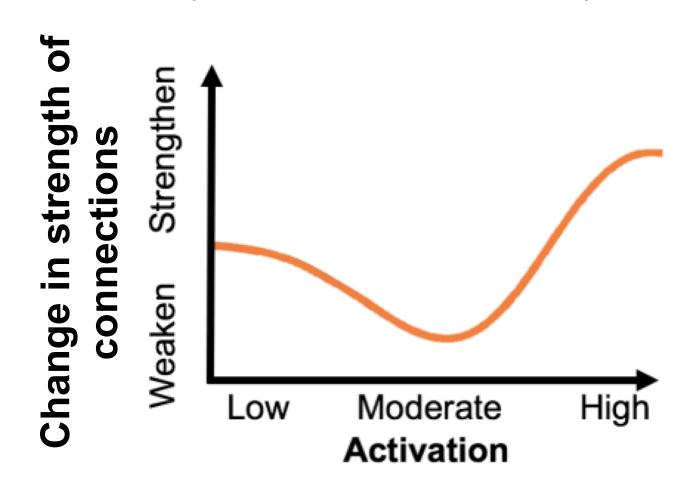


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## Background

When we retrieve a memory, competing memories can come to mind. How does this competition drive plasticity and lead to representational change?

The nonmonotonic plasticity hypothesis (NMPH) posits a U-shaped relationship between coactivation of competing memories and changes to their synaptic connections (Ritvo et al., *TiCS* 2019):



For example, if a competing memory (B) activates moderately when another memory (A) is retrieved, this will trigger weakening of the connections between A and B, leading to neural differentiation (i.e., a decrease in neural overlap).

Sleep is another opportunity to revisit competing memories. REM sleep, in particular, may be a focused period of revisiting competing memories in an interleaved manner (Norman et al., Neural Networks 2005). Can we "mark" memories for representational change during learning, and implement those changes during sleep?

#### Approach:

- Manipulate competition during awake learning
- Manipulate post-learning sleep
- Measure representational change post-sleep

We used a statistical learning paradigm to manipulate pre-sleep competition. One important consequence of statistical learning is the ability to generate predictions based on contextual cues. A predicts B other stimuli

According to the NMPH, if item B is moderately activated due to prediction, but fails to appear, the neural connections between A and B are weakened.

If B is later restudied in a different context, this activates new features not previously shared with A.

Violation Restudy

The result is violation-related neural differentiation: A and B representations are less similar than A-B pairs that did not undergo violation and restudy (Kim et al., *J Neuro* 2017).

### Methods Representational change predictions: post-learning pre-learning **Nonviolation Violation** offline period statistical learning snapshots snapshots podcast listening n = 23Regions of interest: More neural overlap, Less neural overlap. n = 23lower pattern similarity higher pattern similarity n = 23 NREM + REM nap **CA2+3 B** prediction Representational change: similarity of pre-learning A and post-learning B neural patterns Nonviolation initial learning repetitions (3) pre-learning snapshots post-learning snapshots restudy **Violation** B Prediction: similarity of pre-learning B to violation event neural patterns

more neural differentiation?

CA2+3

Correlations are computed within-subject in a pairwise fashion.

B prediction is associated with less representational similarity between

Randomization analysis confirmed this effect is item-specific (p = .02).

A and B (i.e., more differentiation).

DG

WAKE

NREM

### Results

### Relationship between prediction and representational change: Representational change: Do violation pairs become Is stronger activation of B during violation events associated with less similar than nonviolation pairs? 0.04 WAKE NREM 0.02 - REM REM 0.005 0.00 -5 –0.02 **-**-0.04-0.005-0.06-0.08 --0.10CA2+3 DG Plot shows the difference in average representational change between task conditions (violation – nonviolation). Negative values indicate more violation-related neural differentiation. Left CA2+3: The REM group shows the predicted relationship – more

- Right DG: The REM group shows significant violation-related differentiation. Differences between groups are trending (ps < .07).
- Randomization analysis confirmed this effect is item-specific (p = .02).

Error bars are +/- 1 SEM

## Conclusions

### All of the "movement" happens in the REM group:

- Item-specific, violation-related neural differentiation
- Some evidence that B prediction during violation events is related to differentiation
- This suggests a period of REM sleep can promote the plasticity instantiated by competition-dependent learning.

### Next steps:

- Can we improve our neural patterns (e.g., using GLMsingle to derive single-trial estimates)?
- Test for a U-shaped relationship, rather than linear, between prediction strength and representational change (Wammes et al., *eLife* 2022)

Funded by NIH R01-MH06945 and K99-MH126154

