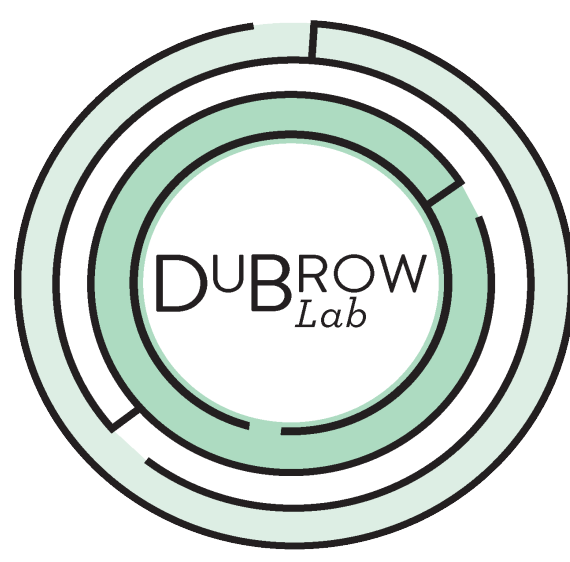


Time-dependent contributions of hippocampus and vmPFC to distributed learning

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Introduction

- Spacing effect: spacing learning out over time improves memory¹.

What is the neural mechanism underlying the spacing effect?

- In rodents, spaced learning is associated with greater ensemble activity pattern similarity in medial PFC².
- However, previous studies have focused only on short timescales, with repetitions occurring within a single experimental session (and day).

What happens in the brain that allows spaced learning over long timescales to improve memory?

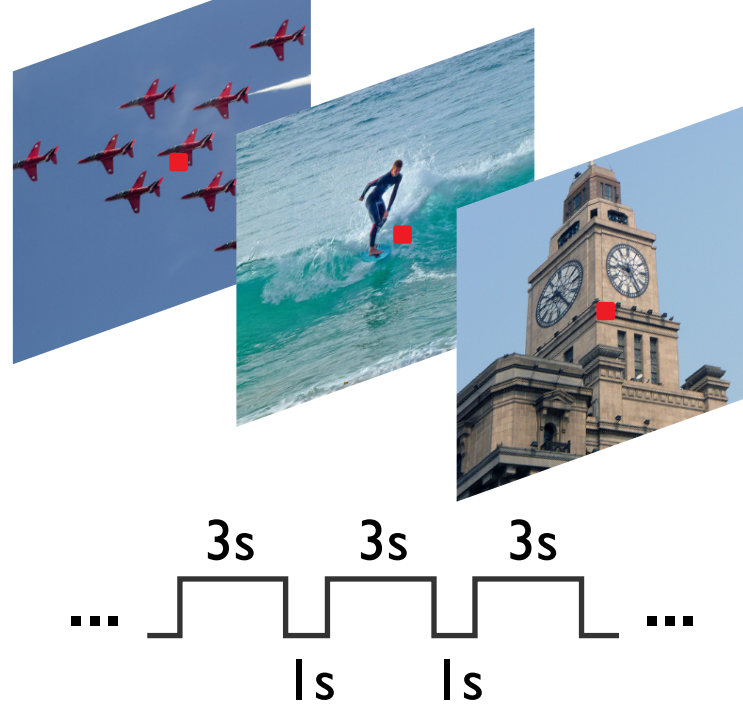
Experimental design

Natural Scenes Dataset³



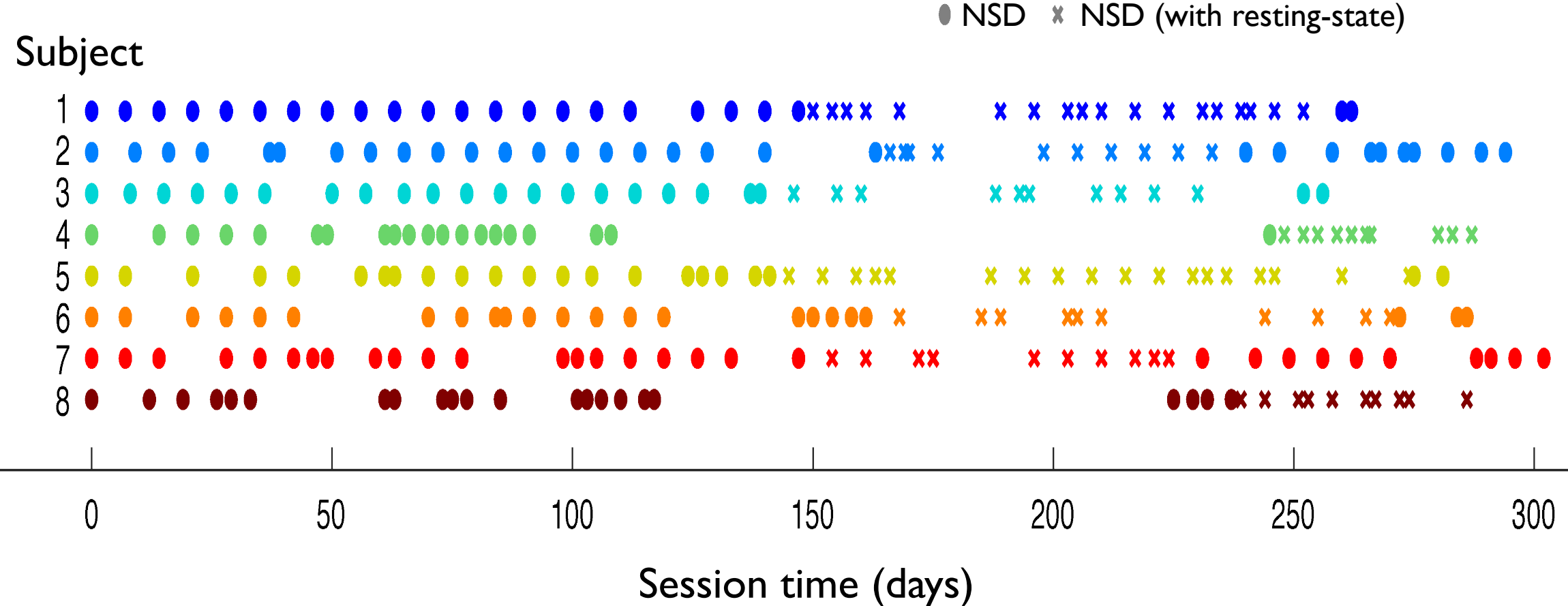
Continuous recognition task

“Have you seen this image before?”
“Yes” or “No”



- 8 human subjects
- 30-40 sessions of 7T fMRI distributed over a year per subject
- ~30,000 trials per subject
- ~10,000 images presented up to three times

Timeline of fMRI scans



Behavioral results

Timeline of an example image



Memory outcome ~ Spacing + E1 onset + E2-E3 lag + FA rates + (1|Subject)

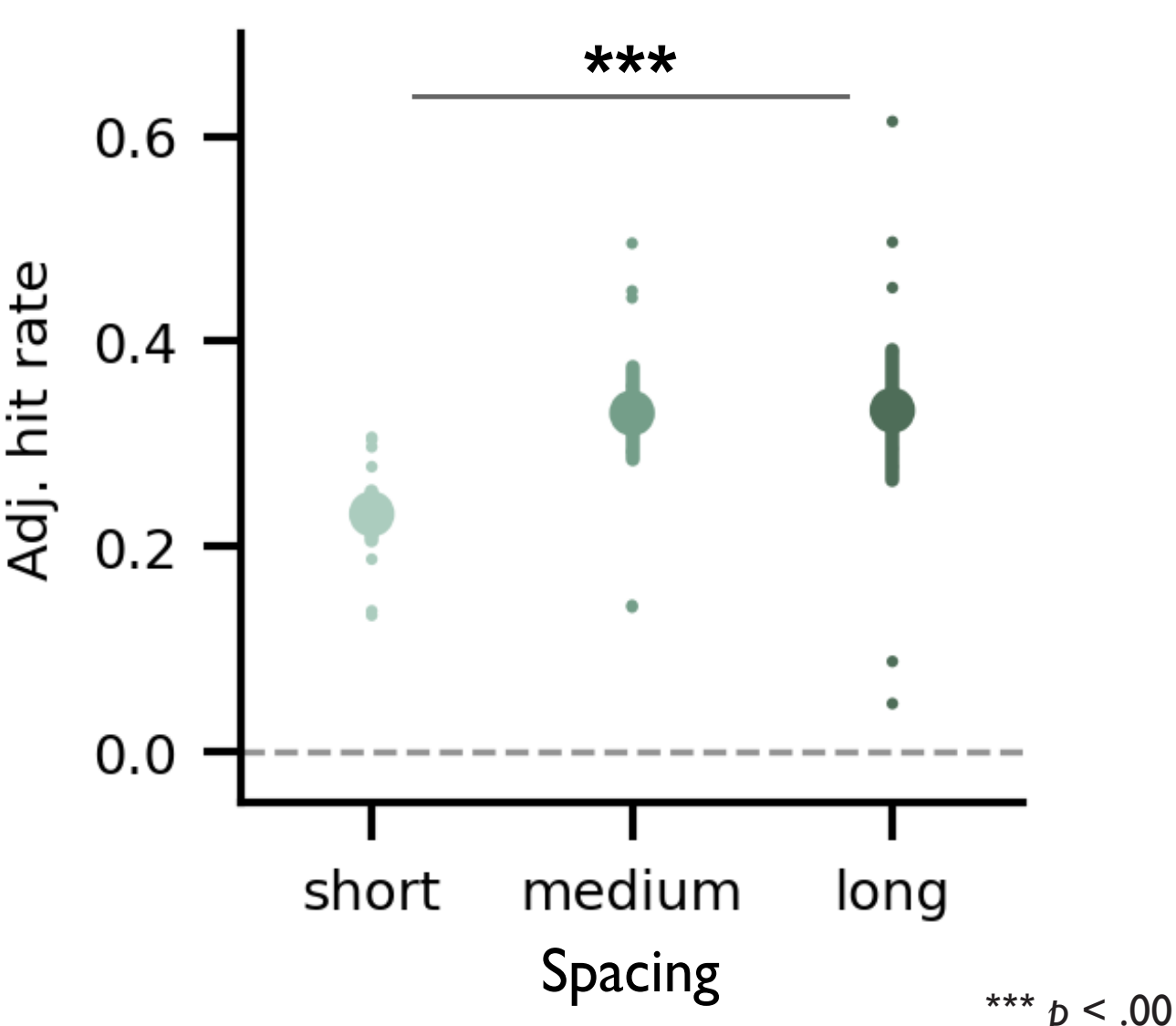
Spacing:

- Short: within-day
- Medium: across-day short (mean: 18 d)
- Long: across-day long (mean: 122 d)

(Items with E2-E3 lag > 1 day)

Spaced learning across long timescales enhances subsequent recognition, with a spacing ranging from 4s to 288 days.

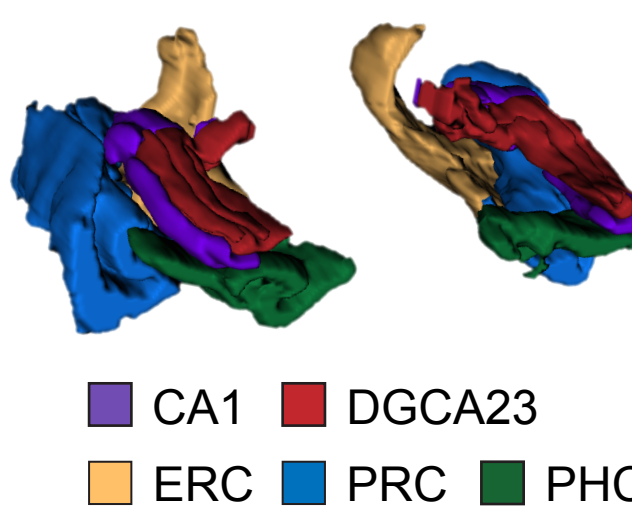
Memory increases as a function of spacing



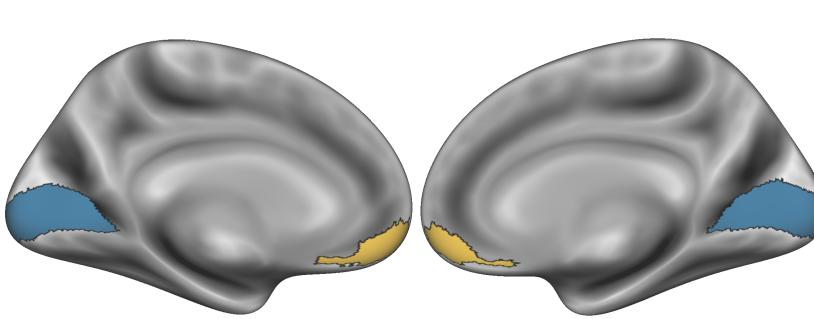
fMRI analysis

Regions of interest

Medial temporal lobe



vmPFC and V1 (control region)



Target item - E1

Target item - E2

Same session Different run

Same session Different run

Foil items - E1'

Foil items - E2'

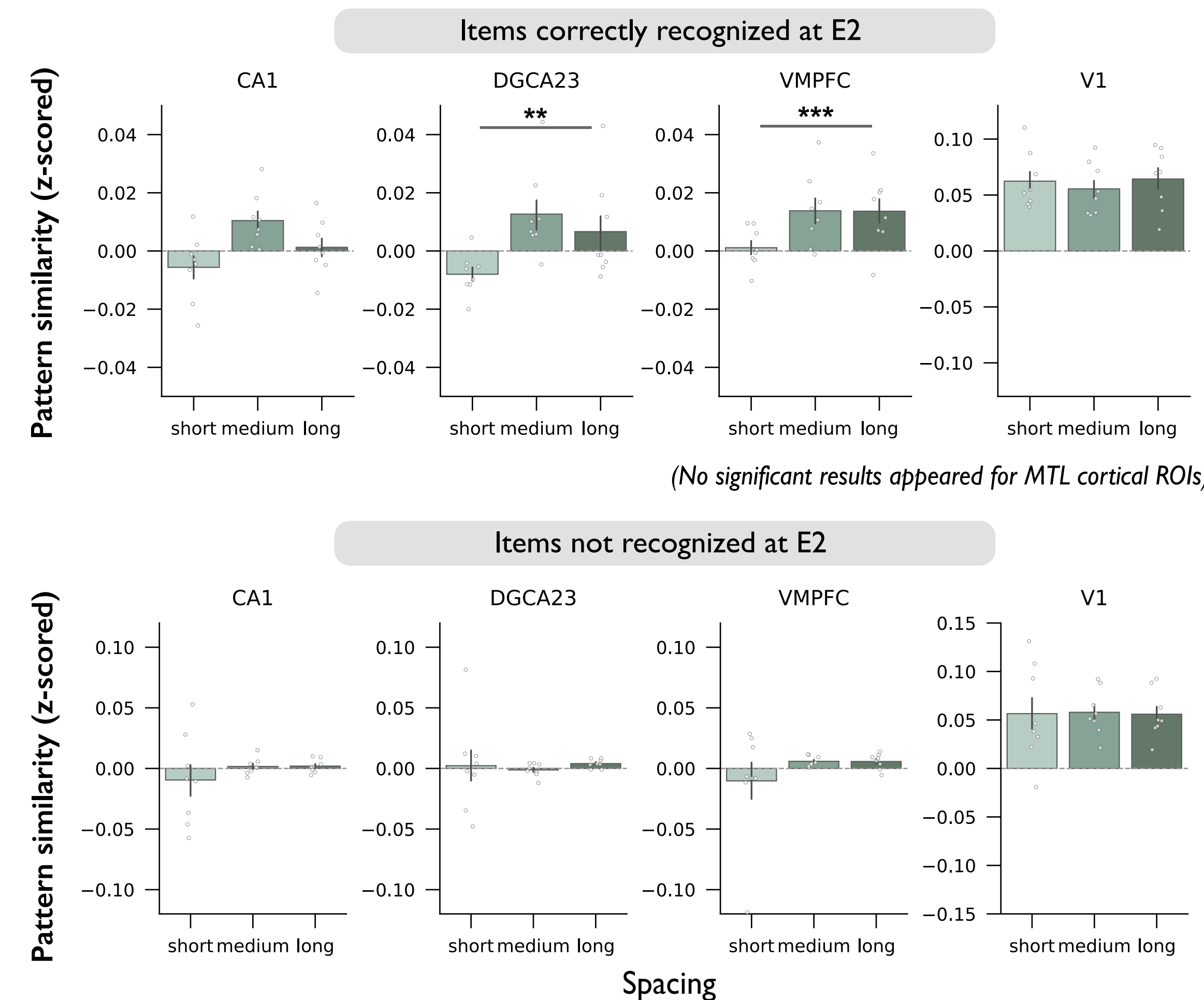
Item-specific similarity = $r(E1E2) - \text{median}(r(E1E2'))$

References and Acknowledgments:

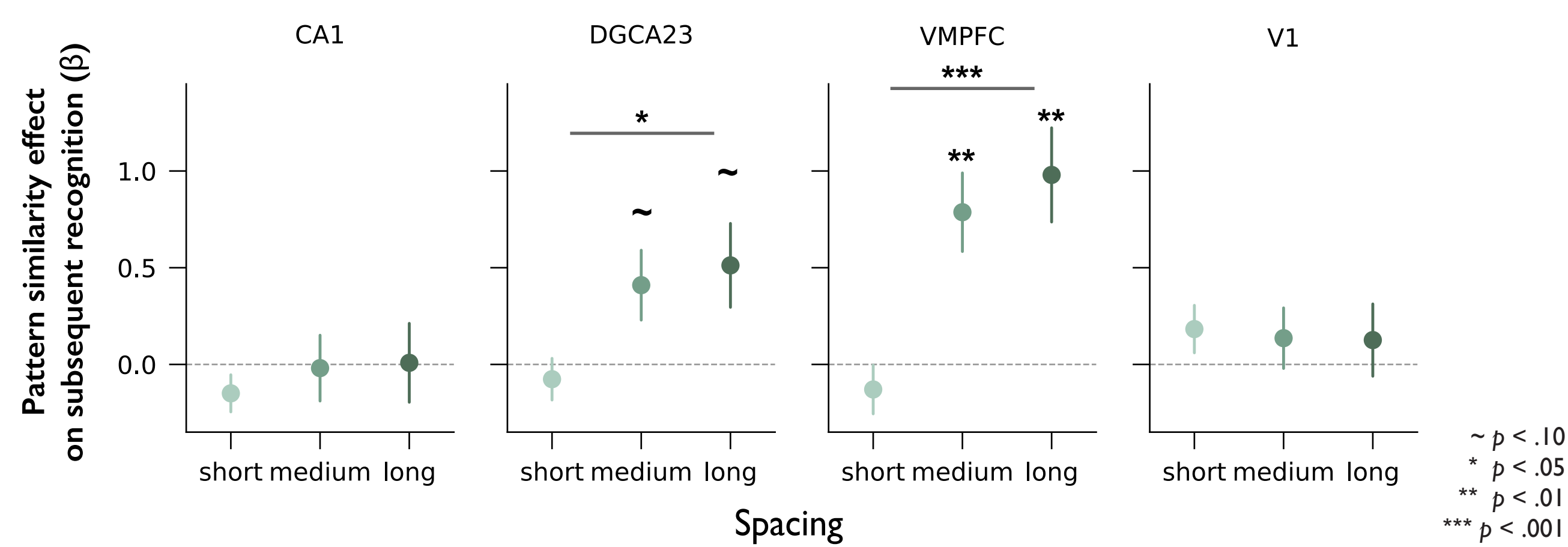
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2. Glas A, Hübener M, Bonhoeffer T, Goltstein PM (2021). Spaced training enhances memory and prefrontal ensemble stability in mice. Current Biology.
3. Allen EJ, St-Yves G, Wu Y, Breedlove JL, Prince JS, Dowdle LT, Nau M, Caron B, Pestilli F, Charest I, Hutchinson JB, Naselaris T, Kay K (2021). A massive 7T fMRI dataset to bridge cognitive neuroscience and artificial intelligence. Nature Neuroscience.
4. Collection of the NSD dataset was supported by NSF CRCNS grants IIS-1822683 (to K.K.) and IIS-1822929 (to T.N.).

fMRI results

Spaced learning enhances item-specific pattern similarity in DGCA23 and vmPFC but is dependent on memory states during second exposure.



Pattern similarity in DGCA23 and vmPFC predicts subsequent recognition for medium/long spacing (across-day), but not short spacing (within-day).



Summary

- Spacing effect operates over long timescales, from seconds to months.
 - Spaced learning is associated with greater item-specific pattern similarity in DGCA23 and vmPFC.
 - Subsequent recognition is predicted by pattern similarity in DGCA23 and vmPFC, but only for spaced (across-day) learning.
- Spaced learning enhances neural pattern similarity in DGCA23 and vmPFC, thus strengthening memory and increasing the probability of subsequent recognition.**