

# The spatial and temporal dynamics of structural plasticity in the memory domain

Yaniv Assaf

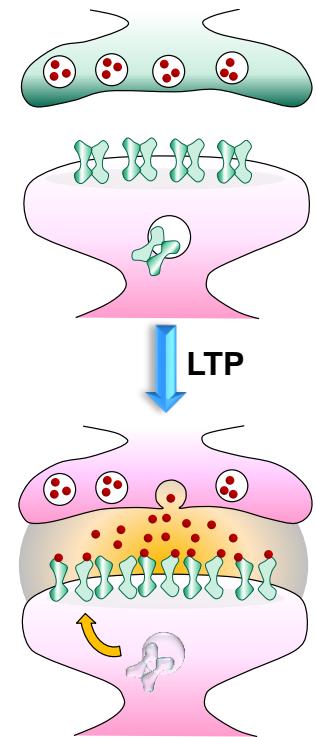
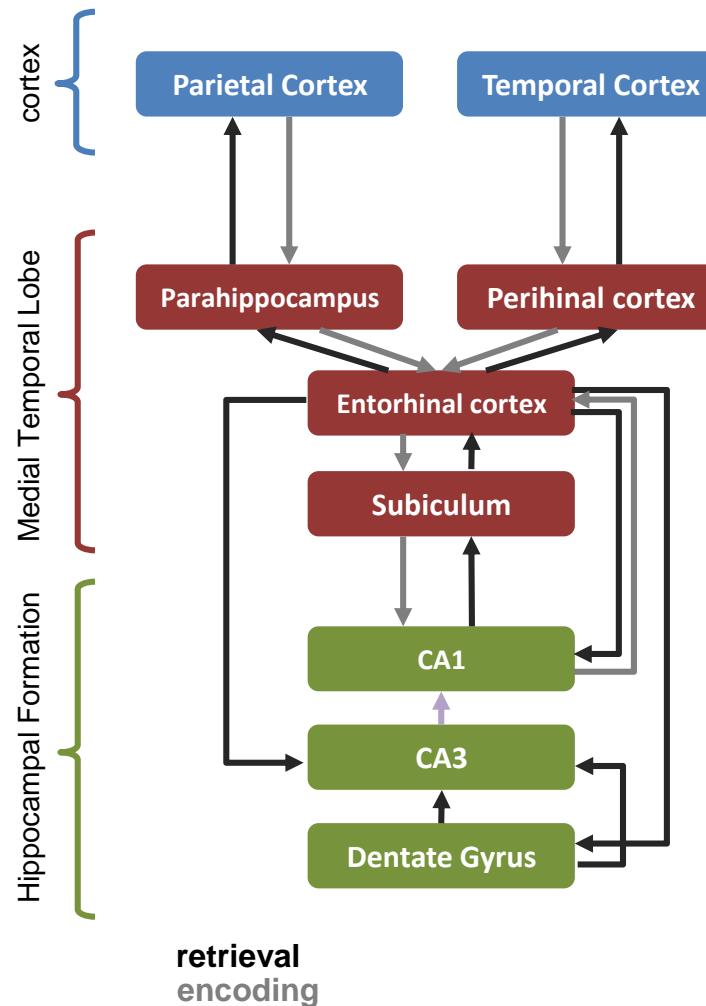


Department of Neurobiology, Faculty of Life Sciences  
Sagol School of Neuroscience, Tel Aviv University  
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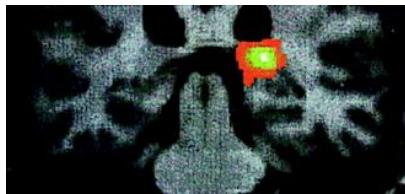


Experimental MRI Centre (EMRIC)  
School of Biosciences, Cardiff University  
Cardiff, The United Kingdom

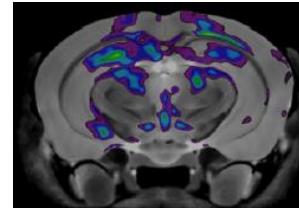
# The memory domain



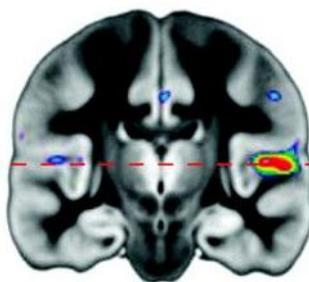
# Neuro-plasticity in Adults



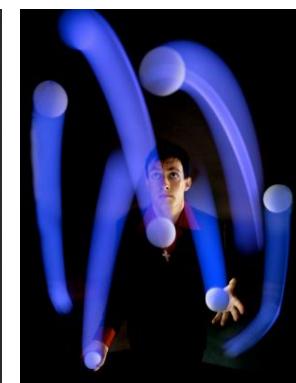
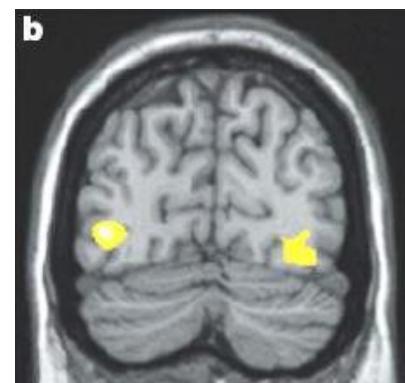
Maguire et-al, PNAS 2000



Lerch et-al,  
Neuroimage,  
2011



Bermudez et-al, Cereb. Cortex 2008

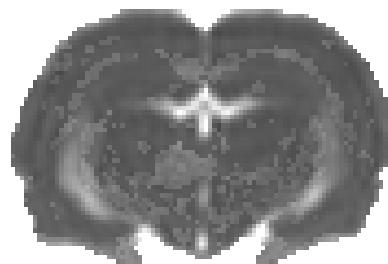


Draganski et-al, Nature 2004

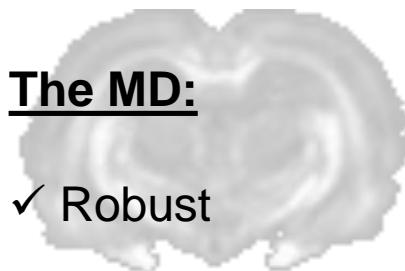
# Diffusion Tensor Imaging



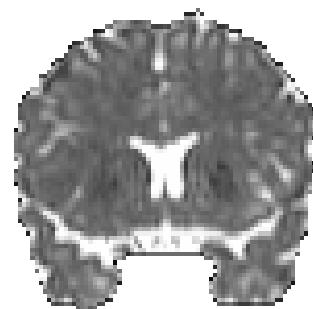
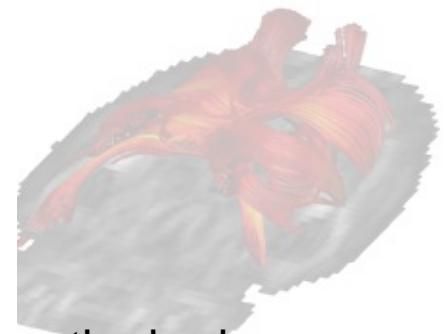
Mean Diffusivity (MD)



Fractional Anisotropy (FA)

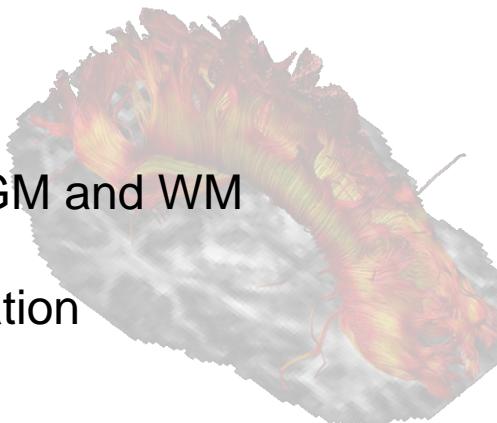


Tractography



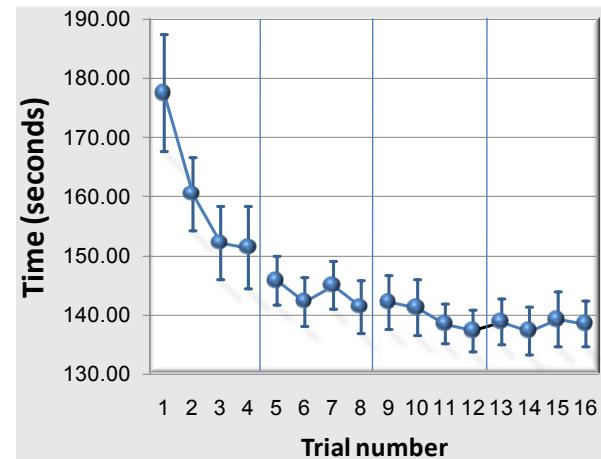
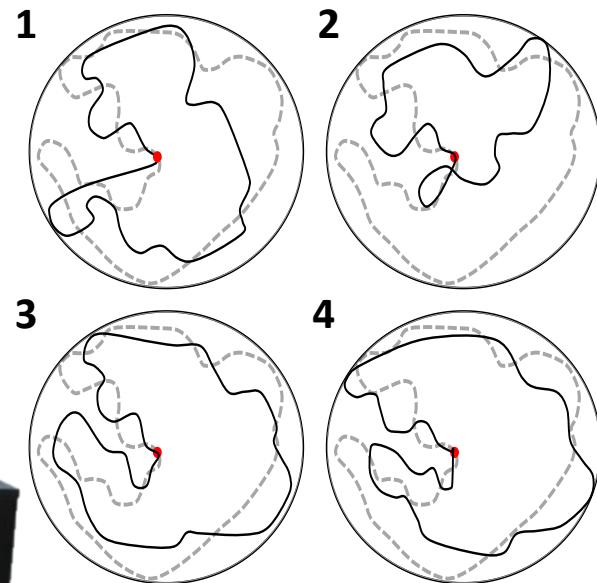
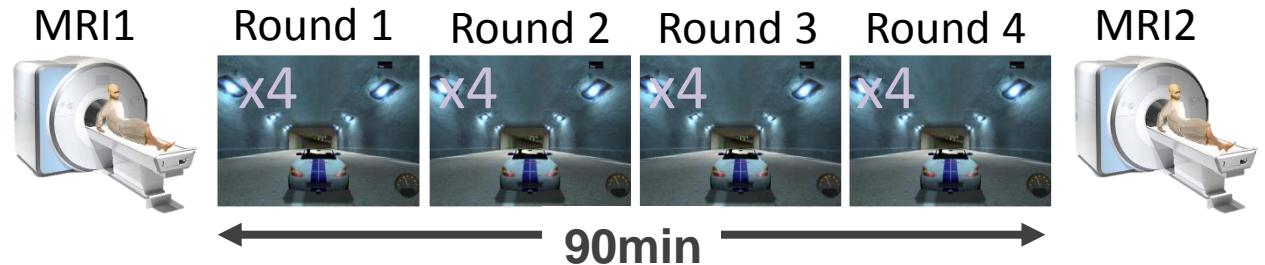
## The MD:

- ✓ Robust
- ✓ Homogeneous Across the brain
- ✓ Fast Acquisition
- ✓ Well defined in GM and WM
- ✓ Simple Interpretation



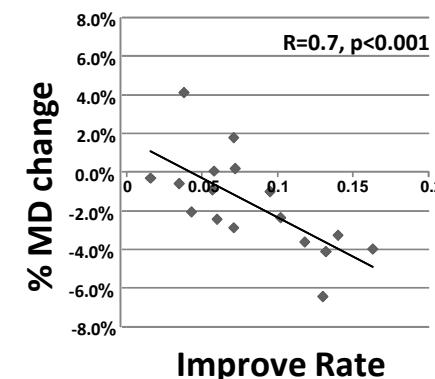
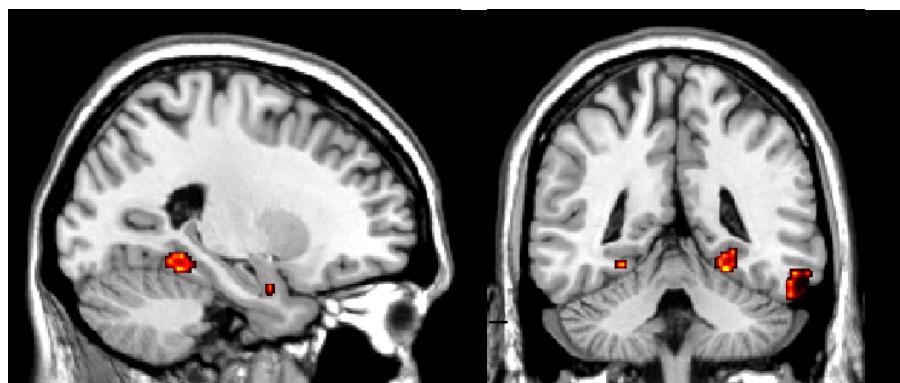
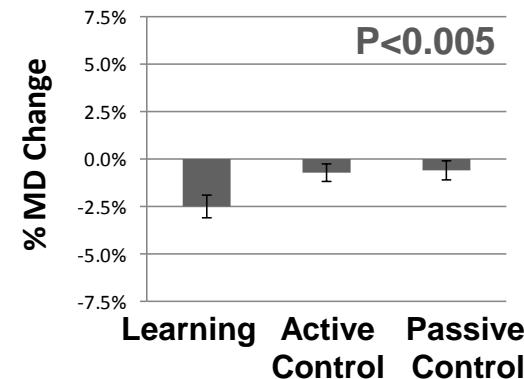
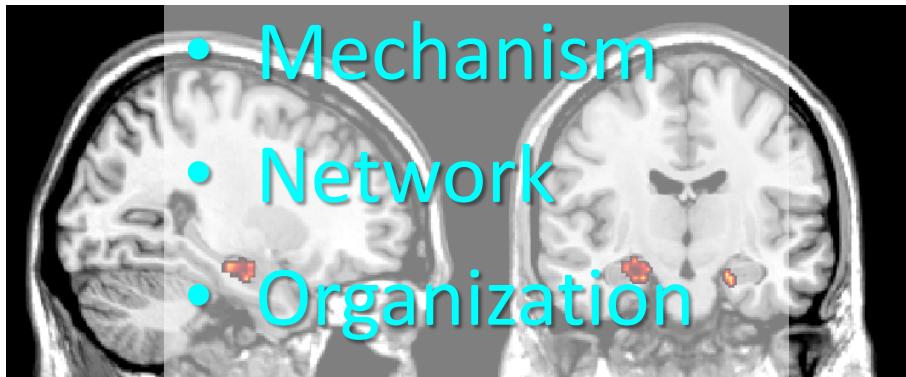
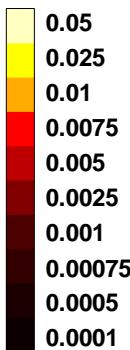
FA → 0.0 0.2 0.4 0.6 0.8 1.0  
MD → 0.0 0.6 1.2 1.8 2.4 3.0 X10<sup>-3</sup> mm<sup>2</sup>/s

# The Need for Speed

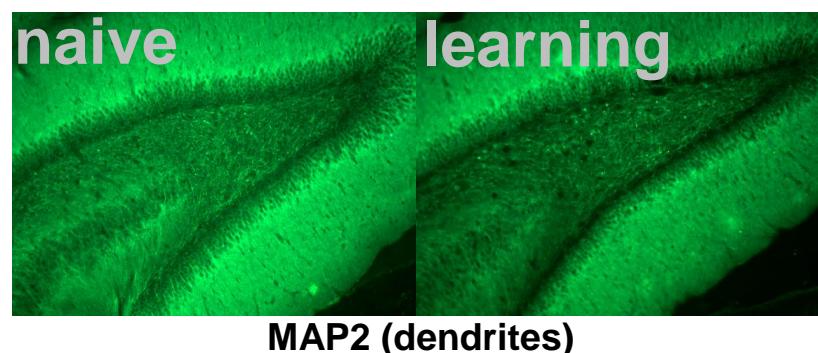
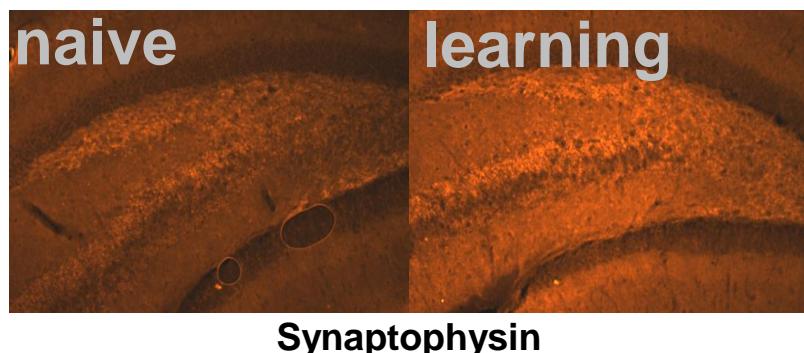
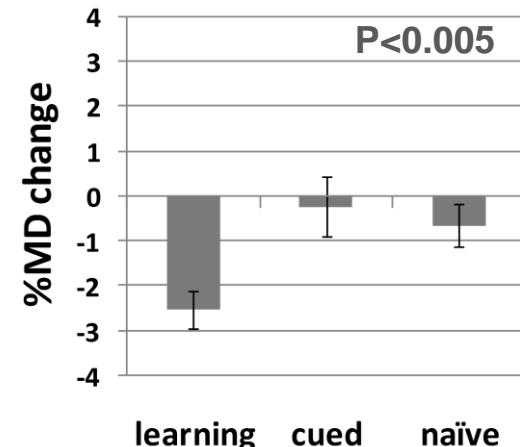
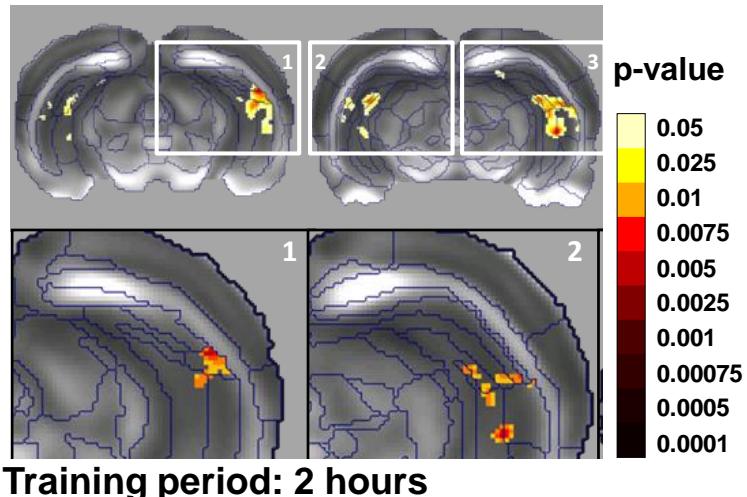
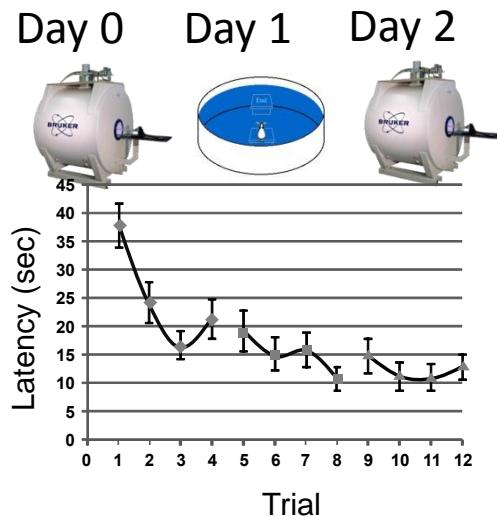


# Effect of Learning and Memory

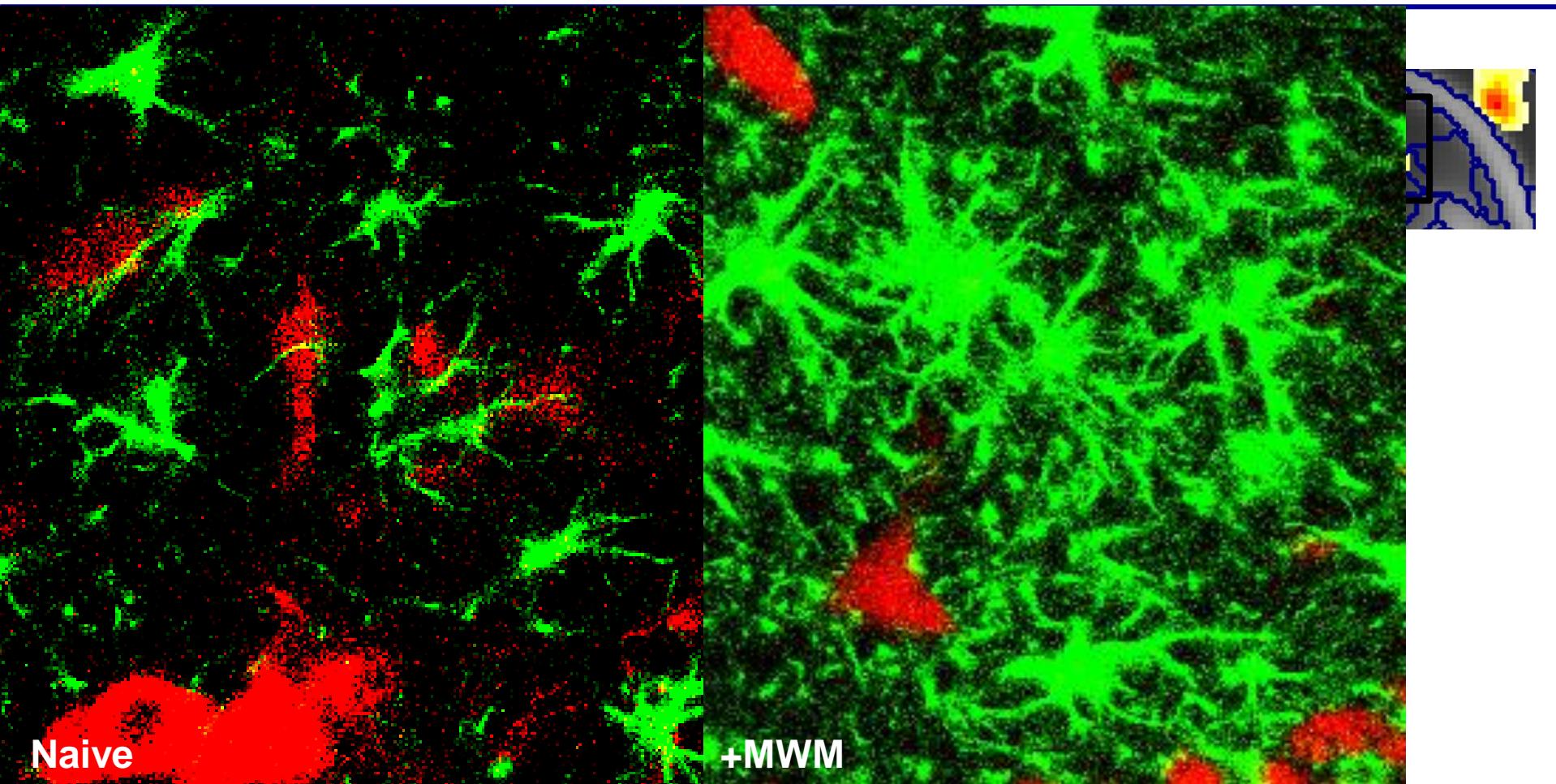
p-value



# Working Memory Structural Plasticity?



# Structural Remodeling



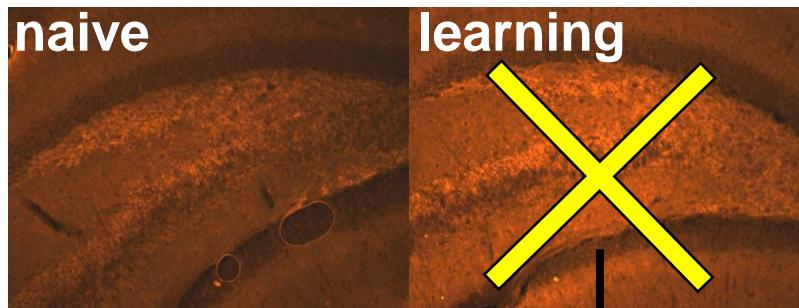
Naive

+MWM

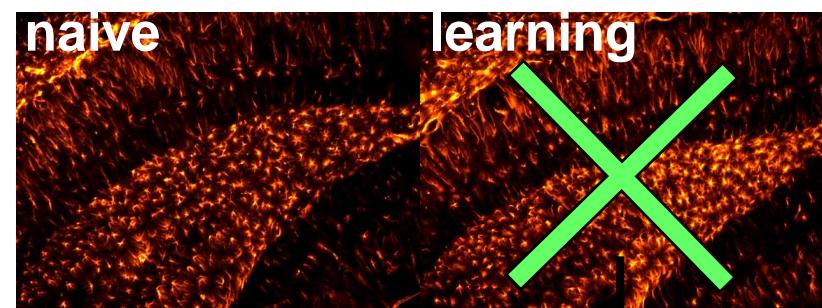


Sagol School  
of Neuroscience  
Tel Aviv University

# Source of MD reduction

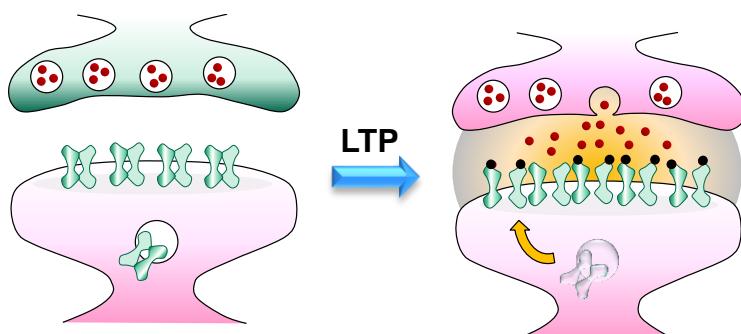


Synaptophysin

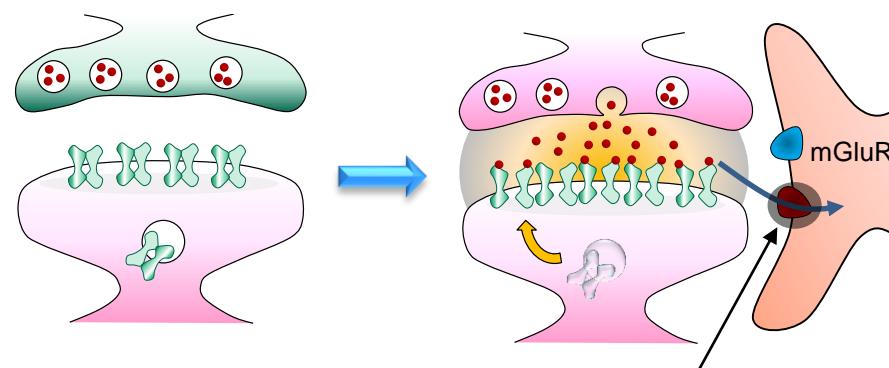


GFAP (astrocytes)

NMDA antagonist



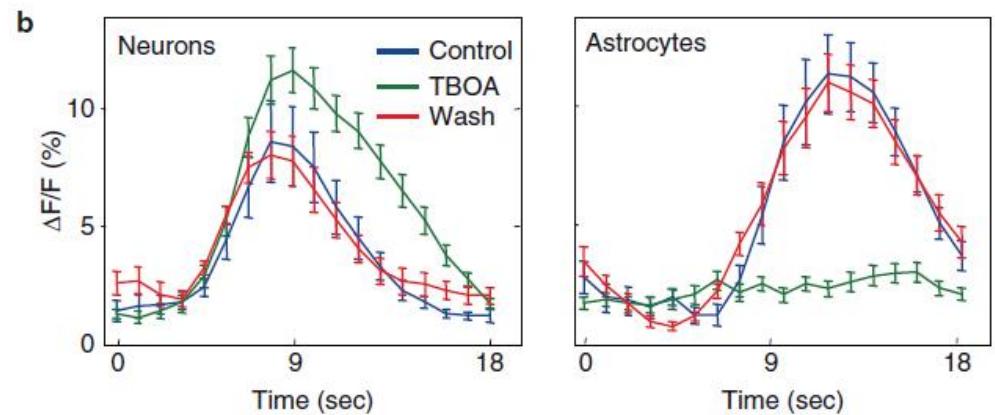
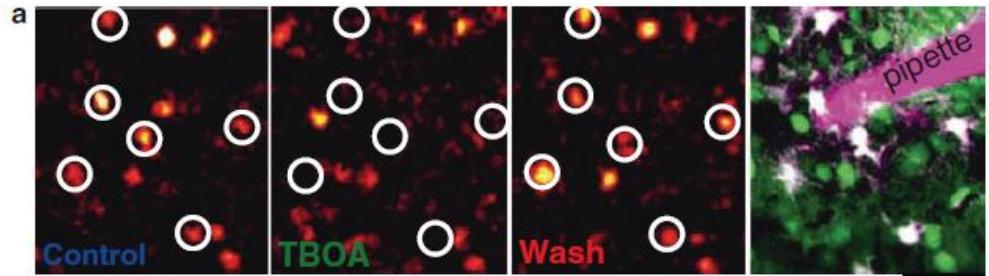
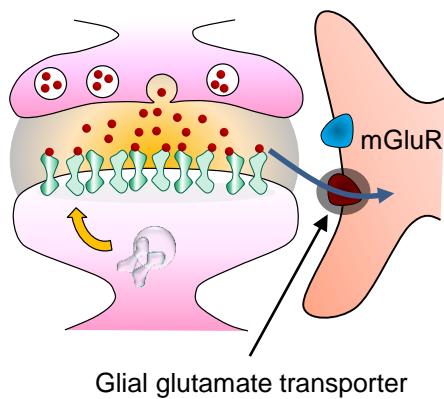
Glutamate transporter antagonist



Glial glutamate transporter



# Blocking astrocyte activation

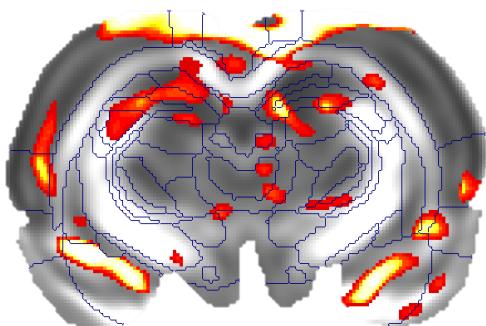
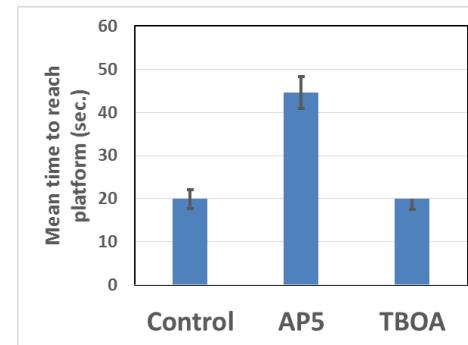


Yu, Schummers, Sur,  
in: "Imaging the brain with optical methods, pp 45-65, 2010

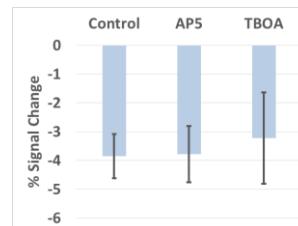
# Pharmacological Interference to learning



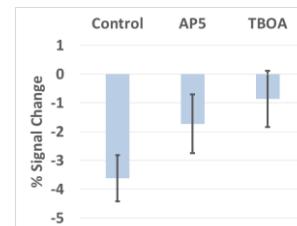
Group1 – aCSF  
Group2 – AP5  
Group3 - TBOA



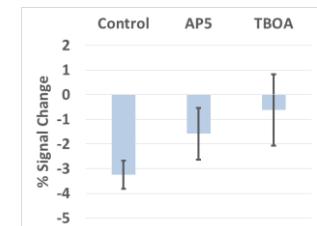
Dentate Gyrus



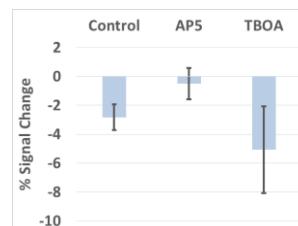
CA1



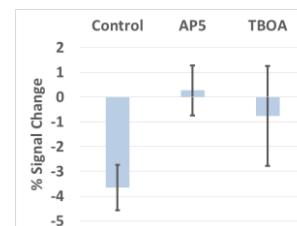
Entorhinal



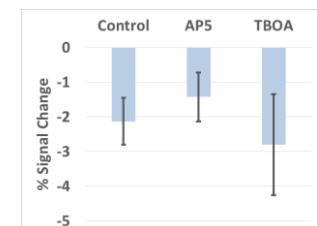
Cingulate Cortex



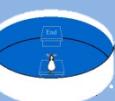
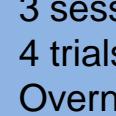
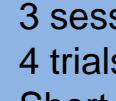
S1



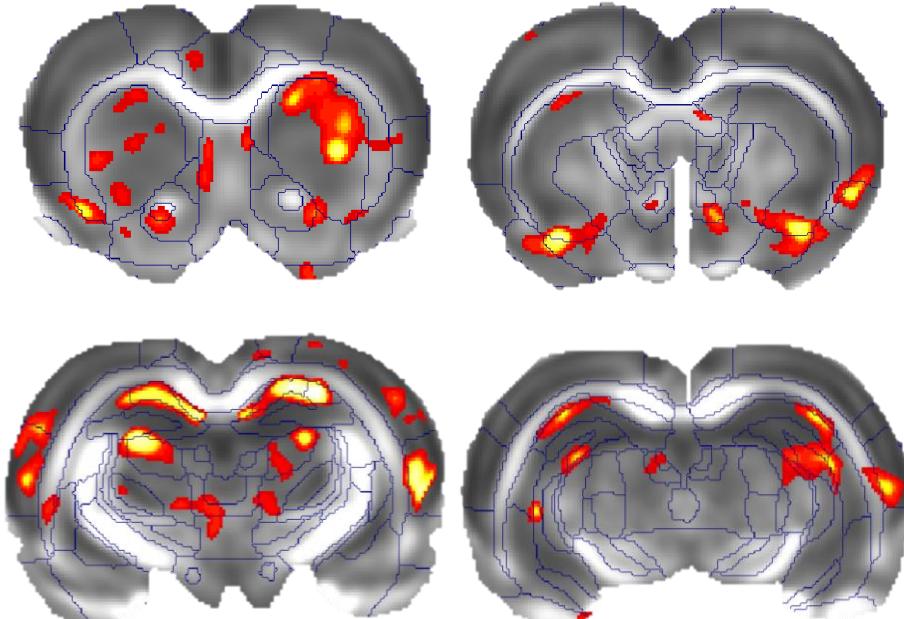
Amygdala



# The temporal dynamics of brain plasticity

Exp. 3	Day 0	Day 1	Day 2	Day 3	Day 4	Day 5	Day 6	<u>Consolidation</u>
								5 session 4 trials in each session Overnight
Exp. 2	Day 0	session1	Session 2	Session 3	Day 1	<u>Long Term Memory</u>		3 session 4 trials in each session Overnight
								
Exp. 1	Day 0	session1	Session 2	Session 3	Day 0	<u>Short Term Memory</u>		3 session 4 trials in each session Short term memory
								

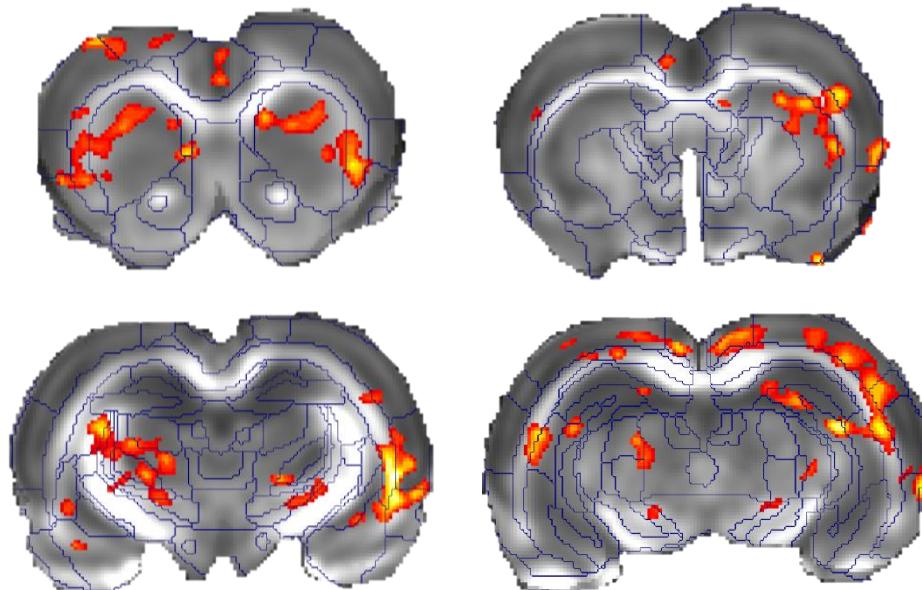
# Short Term Memory



Dorsal Hippocampus (CA1)  
Thalamus (LP)  
Ventral Pallidum  
Dorsal striatum (CPu)  
Entorhinal Cortex  
Auditory Cortex  
Dentate gyrus

	Day 0	session1	Session 2	Session 3	Day 0	
Exp. 1						3 session 4 trials in each session Short term memory

# Long Term Memory



Sensory Cortex (S1/S2)  
Cingulate Cortex  
Entorhinal Cortex  
Dentate Gyrus  
Dorsal Striatum  
Thalamus (ventral nuc.)

Exp. 2

Day 0



session 1



Session 2



Session 3

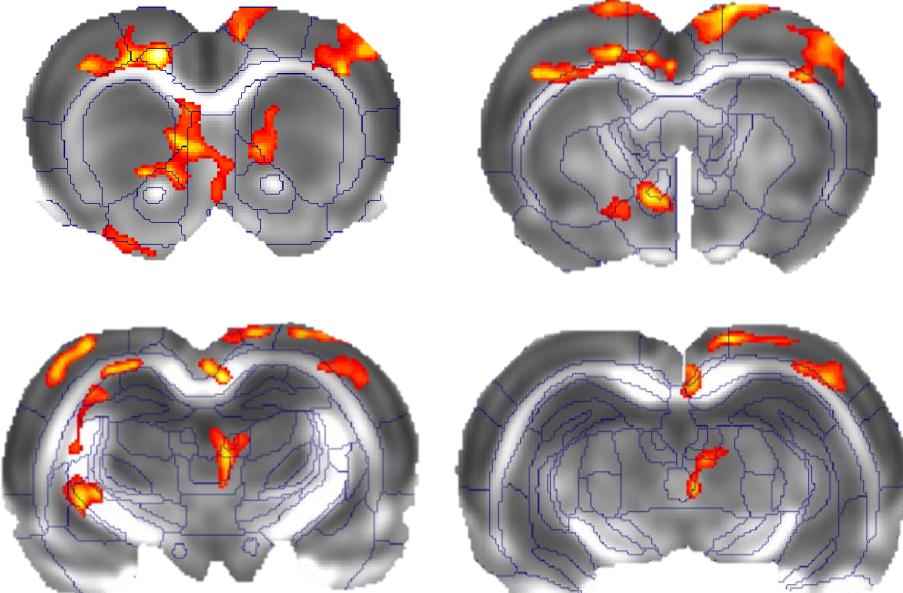


Day 1



3 session  
4 trials in each session  
Overnight

# Retrieval



Motor Cortex (M1/M2)  
Sensory Cortex (S1/S2)  
Cingulate Cortex  
Septal Nuc.  
Visual Cortex  
Dentate gyrus

Corpus Callosum  
Fornix

Exp. 3

Day 0



Day 1



Day 2



Day 3



Day 4



Day 5



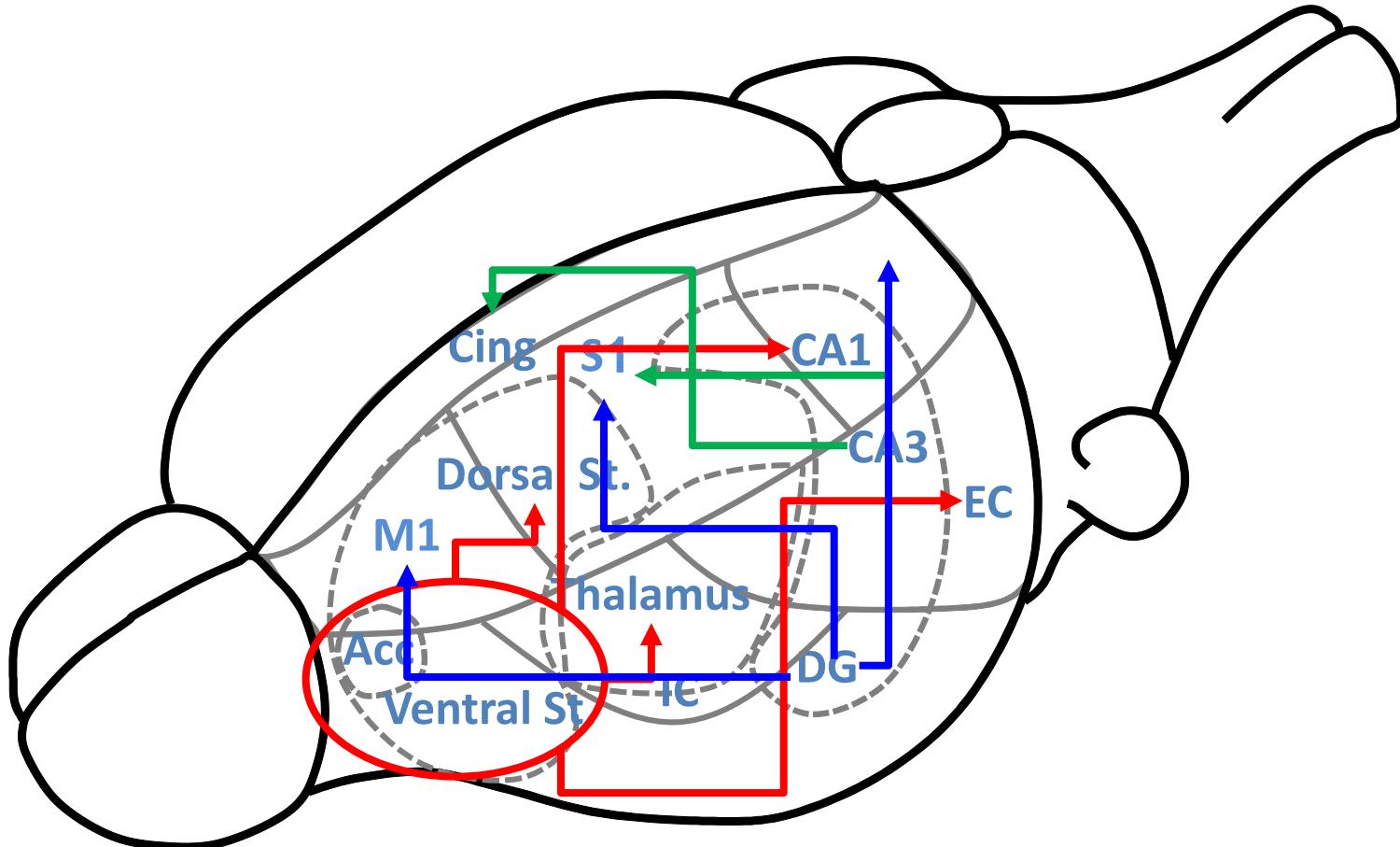
Day 6



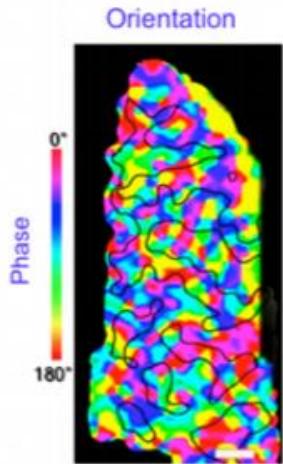
5 session  
4 trials in each session  
Overnight



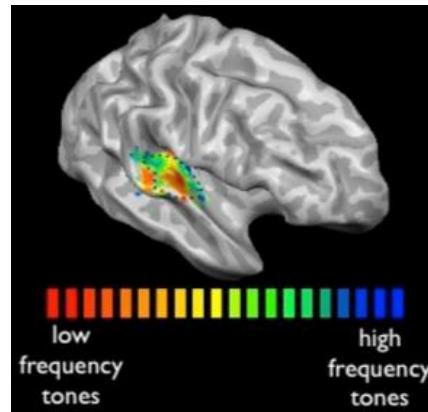
# Dynamics of Plasticity



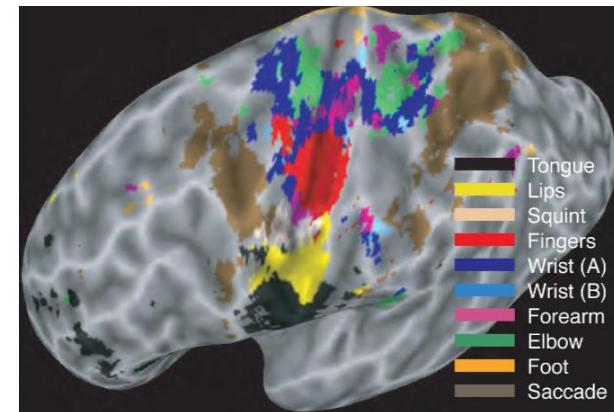
# Spatial Organization?



Yacoub et al.  
PNAS 105, 10607, 2008

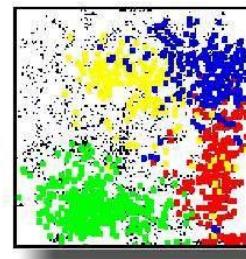
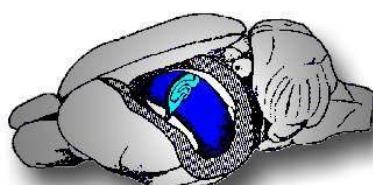


De Costa et al.  
Journal of Neuroscience 31, 14067, 2011



© Michael Graziano

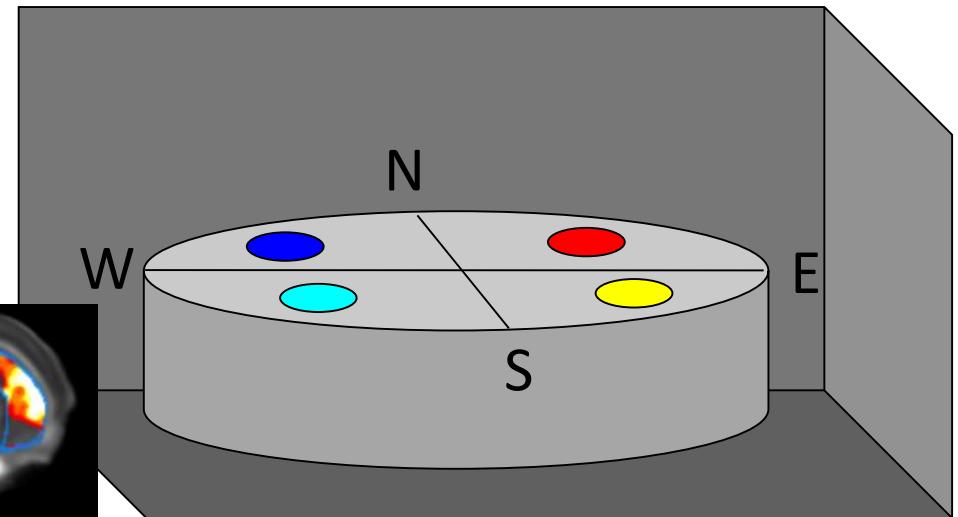
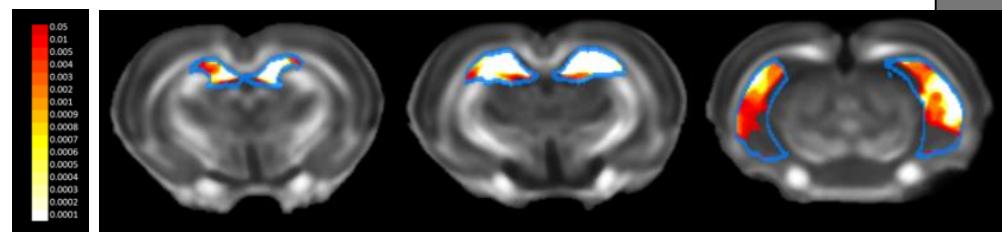
Is there a system level organization within the hippocampus?



No place maps (based on electrophysiology)

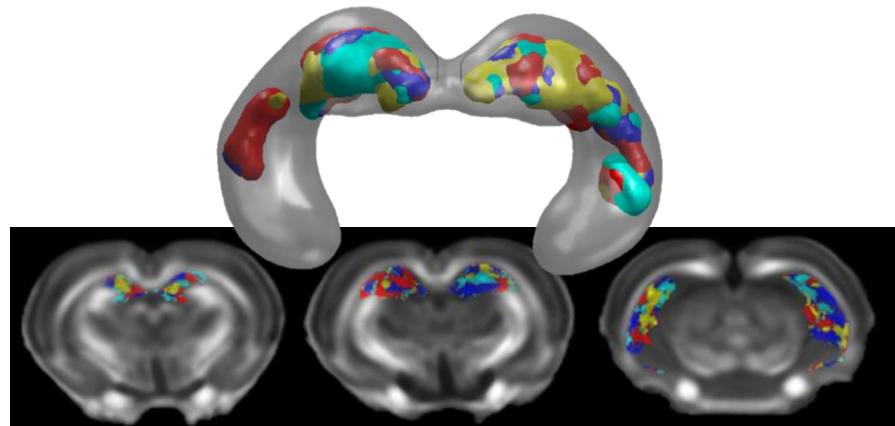
# Spatial Organization in the hippocampus?

4 groups: for each the platform is located at a different quadrant (n=52).

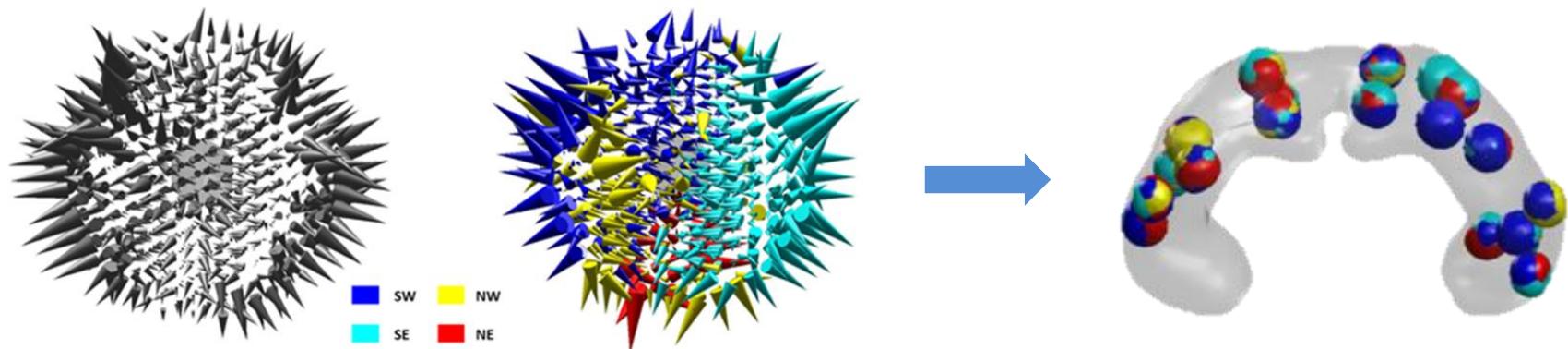
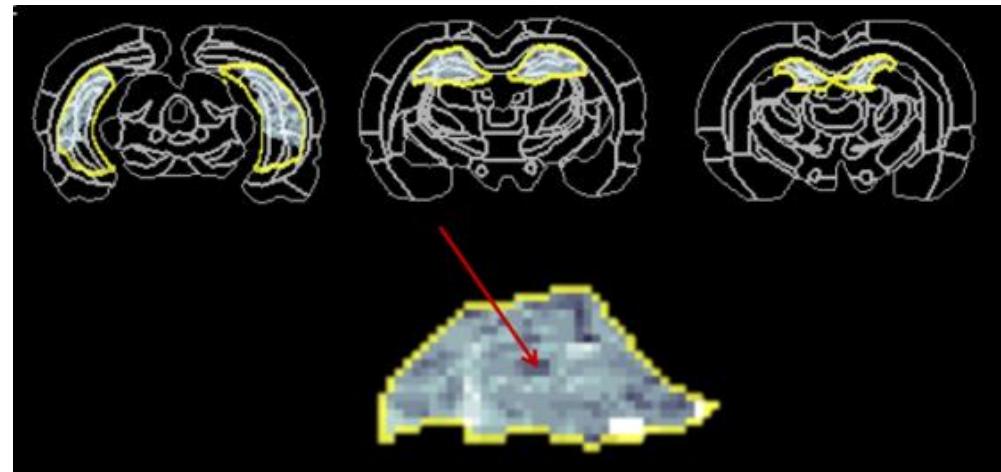
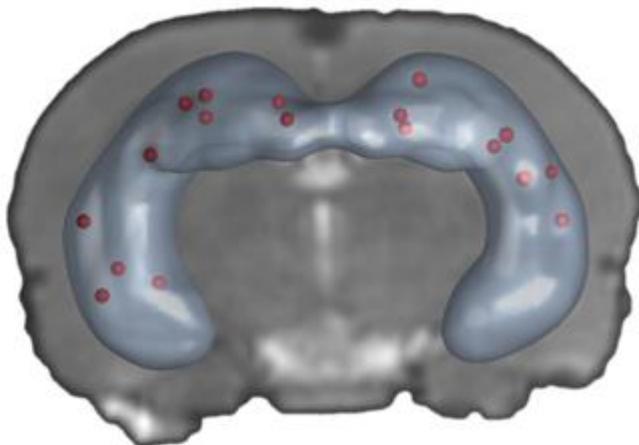


Is there a preference for location?

(i.e. voxels where MD reduction was larger for specific platform location)

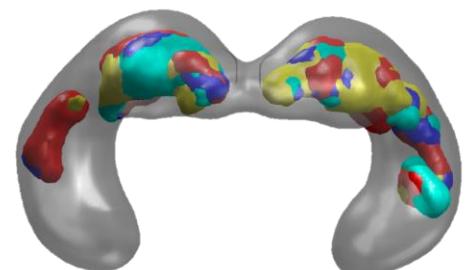


# Preference for Index



# Summary

- Diffusion MRI based characterization of neuroplasticity opens new opportunities to explore the memory domain
- The mechanism of diffusion MRI changes seems to be strongly related to astrocyte activity
- Structural neuroplasticity is a dynamic process that progress in space and time within the brain
- Diffusion MRI enables measurement of neuroplasticity on the system level for the whole brain providing a more holistic and comprehensive characterization of this process



# Acknowledgements

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Prof. Peter Basser, NIH

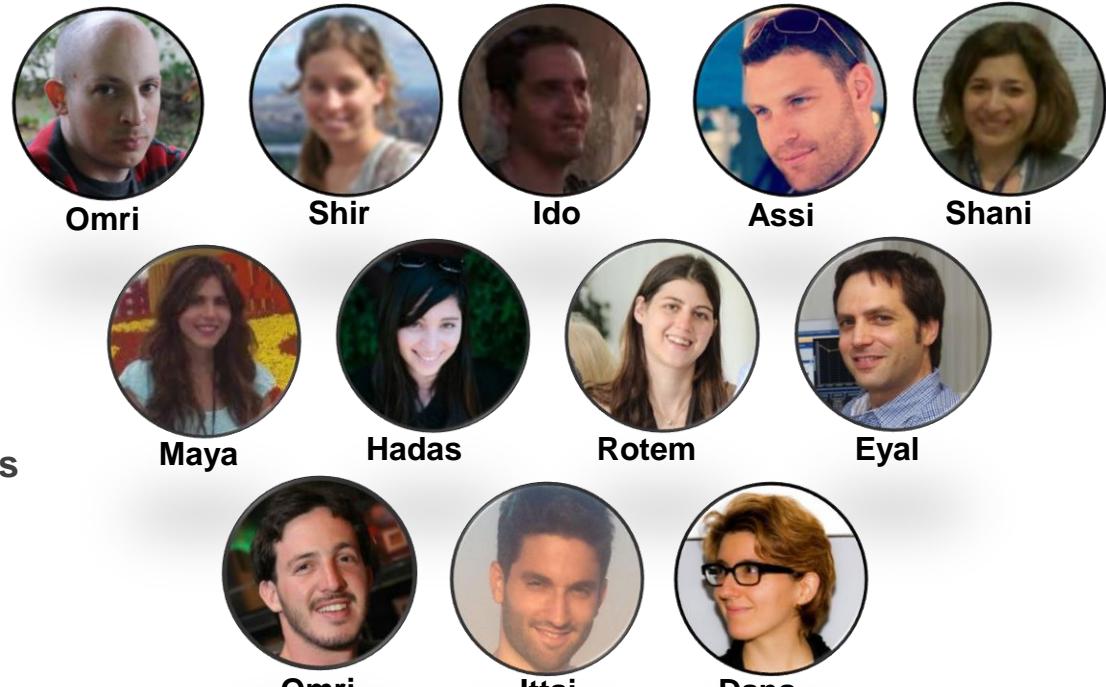
Prof. Derek Jones, UK

Dr. Alexander Leemans, The Netherlands

(ExploreDTI)

Mr. Nadav Mark

Strauss Center for Computational  
Neuroimaging



<http://neuroimaging.tau.ac.il/ya>



Thank You