The hippocampus is necessary for binding object identity to location in visual working memory.

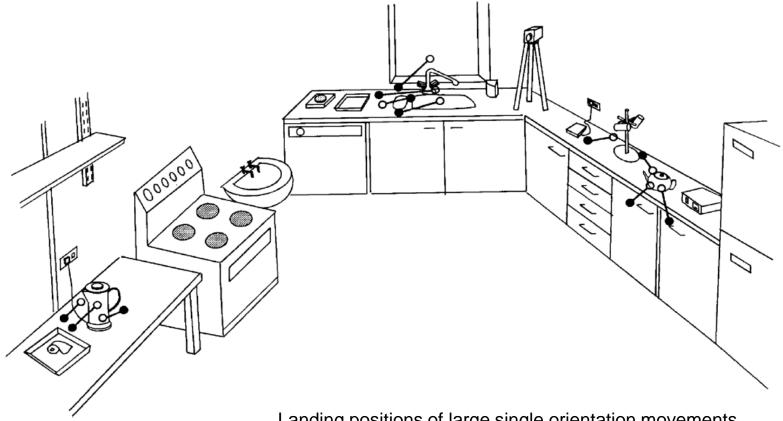
Yoni Pertzov

Department of Psychology The Hebrew University of Jerusalem



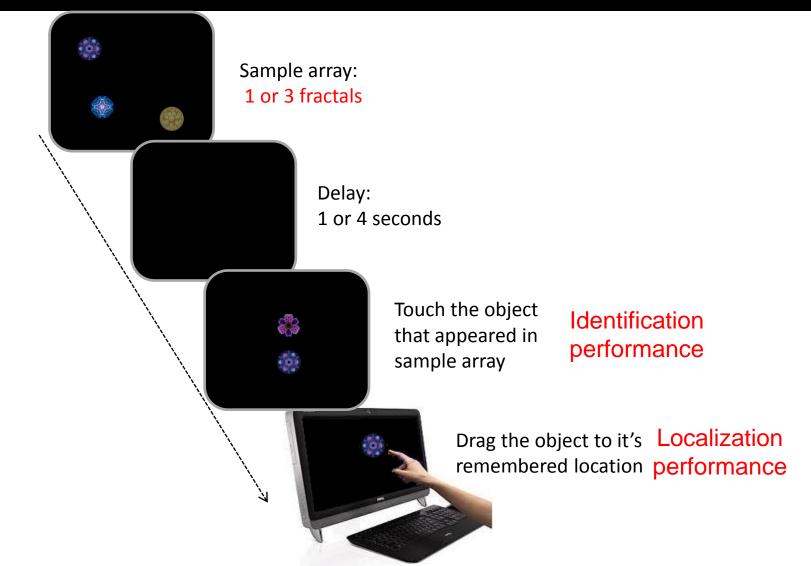
12th International Conference On Cognitive Neuroscience, Brisbane Australia

Visual working memory is often used for localizing objects



Landing positions of large single orientation movements, requires memory (out of the visual field >90°) Tatler & Land. *Phil. Trans.*, B 2011.

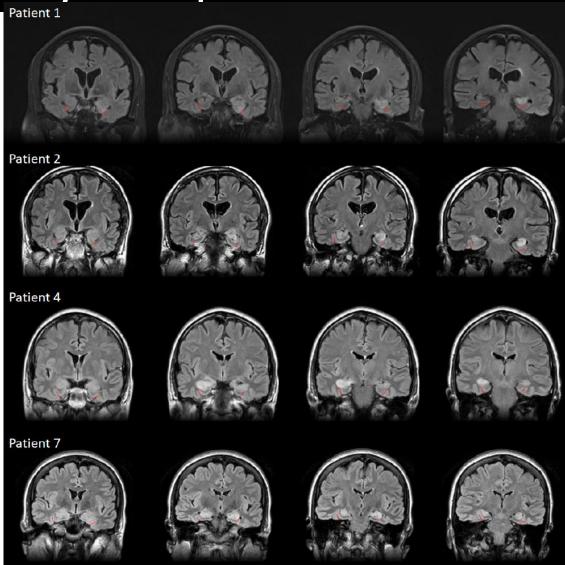
Remembering what was where



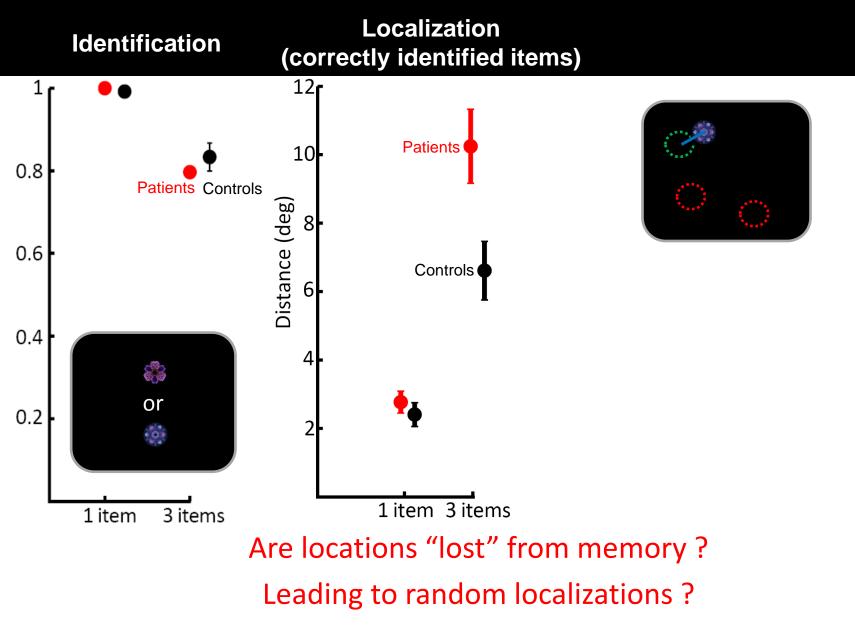
Pertzov et al. Forgetting What Was Where: The Fragility of Object-Location Binding. PLOS ONE 2012

Voltage Gated Potassium Channel Antibody encephalitis

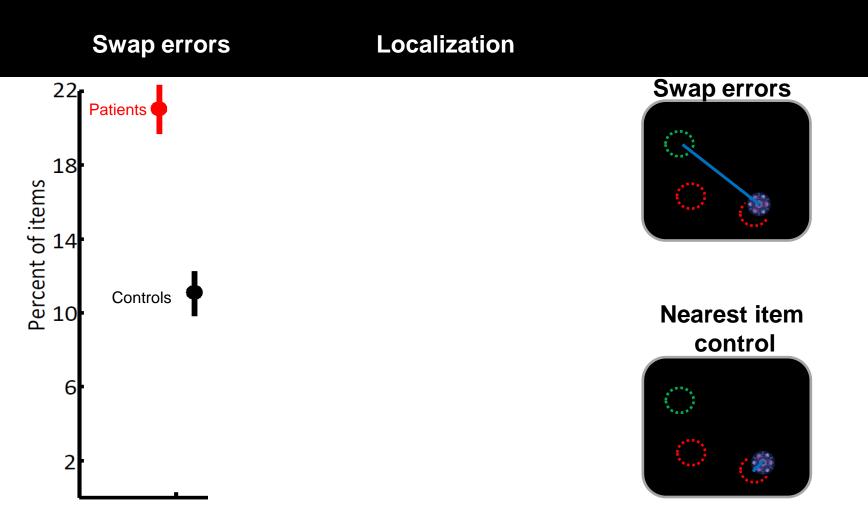
- 7 patients.
- Recently recognized: Vincent et al 2004. Responds to treatment.
- Antibodies target LGI1 protein which is expressed very restrictively in the hippocampus.
- Post mortem study revealed neural loss exclusively in the hippocampus and amygdala.
- Abnormal signal restricted to the MTL.



Pertzov et al. Medial temporal lobe damage impairs binding in visual short-term memory. BRAIN, 2013



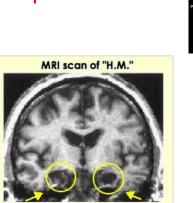
Pertzov et al. Medial temporal lobe damage impairs binding in visual short-term memory. BRAIN 2013.

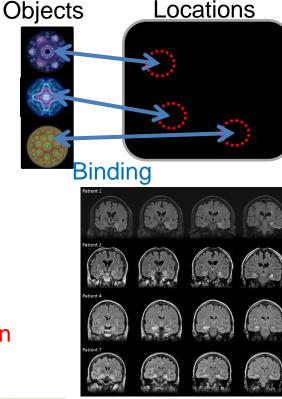


Patients' impairment is associated with swap errors.

Binding in working memory: Neural mechanisms

- When forgotten, items are lost in their entirety ?
- The links that bound objects to their locations could be forgotten in spite of intact memory of item identity and position.
- Medial temporal lobes (MTL) involved in long term memory but not in working memory?
- MTL is involved in binding isolated properties also across brief retention intervals. But not ID or position alone.
- Challenging the 'multi-store hypothesis'.



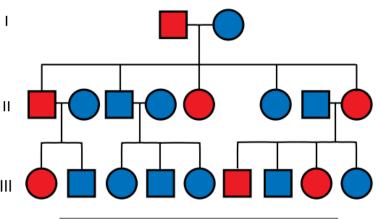


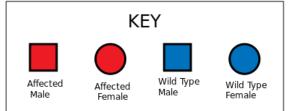
Binding in working memory: Clinical practice ?

- Recent failures of clinical trials imply that we must treat Alzheimer's Disease (AD) prior to its mild to moderate stages.
 - Requires early detection of the disease.
- The MTL is compromised in AD years before onset of behavioral symptoms.
- Standard tests of episodic memory are crude and uncontrolled. Maybe use binding in memory ?
- How can we test patients BEFORE they are diagnosed with AD

Familial Alzheimer's Disease (FAD)

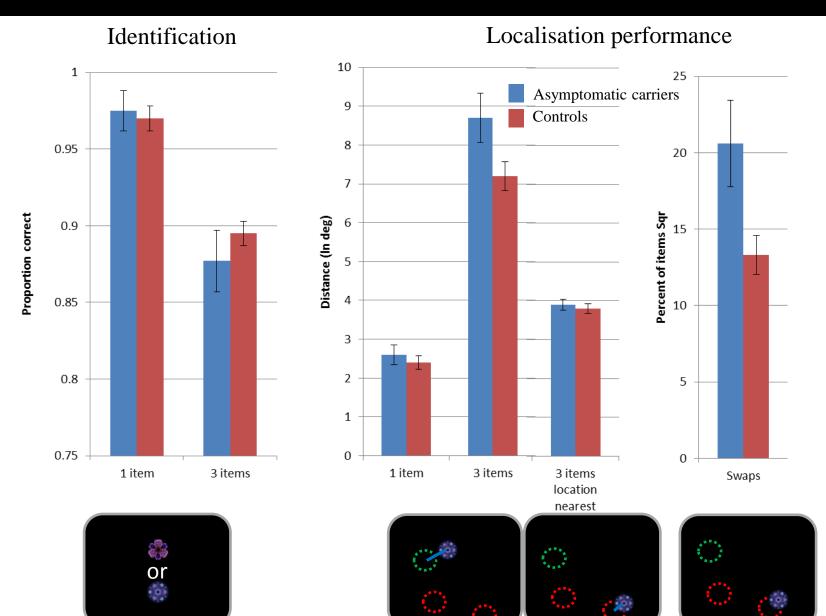
- Autosomal Dominance disease
- Mutation Carriers have 100% chances of developing FAD at the parents age.
- Invaluable for studying presypotomatic stages.
- 12 asymptomatic mutation carriers with no deficits on standard neuropsychological tasks
- 62 healthy controls
- "double blind" design.



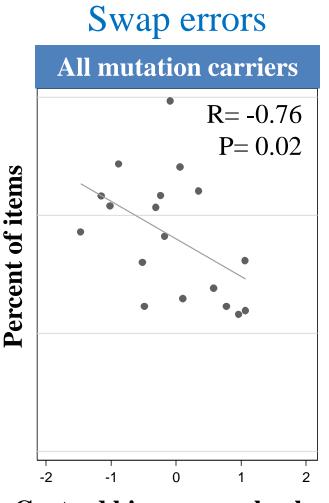


In Autosomal Dominance the chance of receiving and expressing a particular gene is 50% regardless of the sex of parent or child.

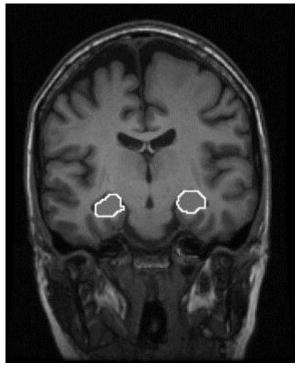
Asymptomatic carriers vs. controls



Correlation with hippocampal volume



Automatic segmentation of Hippocampal volume



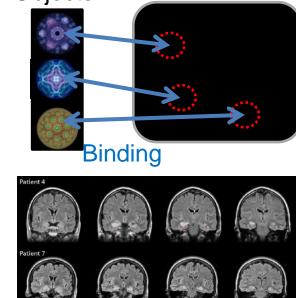
Centred hippocampal volume

Conclusions

Items and their locations are represented independently in visual working memory

Binding in visual working memory related to hippocampal integrity

Binding errors in WM may provide a means for early detection in Alzheimer's disease





Objects

Thanks to:

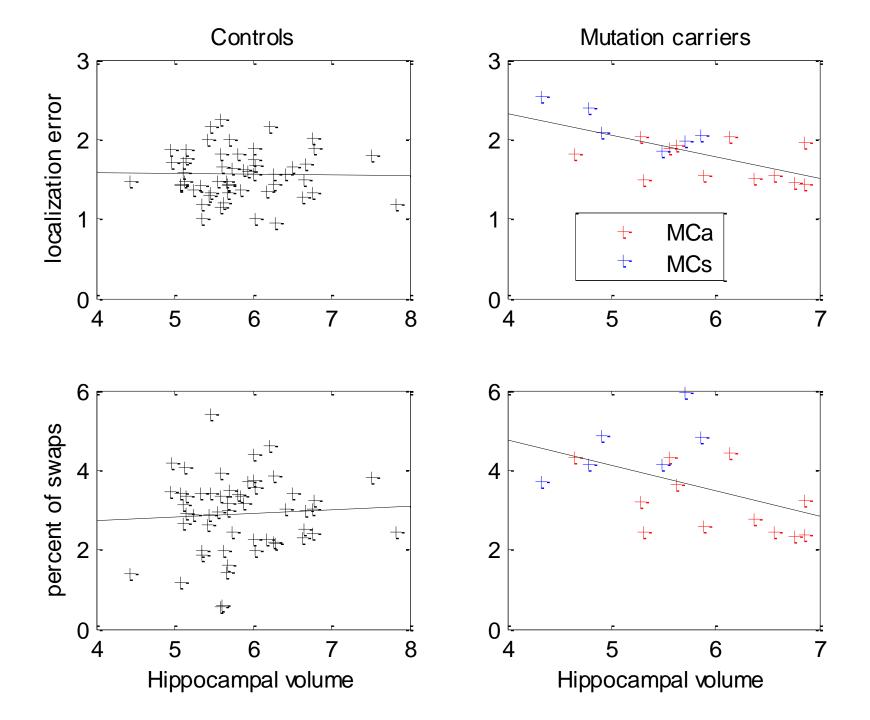
Masud Husain

Cognitive Neurology Research Group Cognitive Neuropsychology Centre University of Oxford

MTL patients: Chris Butler Tom Miller Department of Clinical Neurology University of Oxford

Alzheimer patients: Yuying Liang Sebastian Crutch Dementia Research Centre University College London

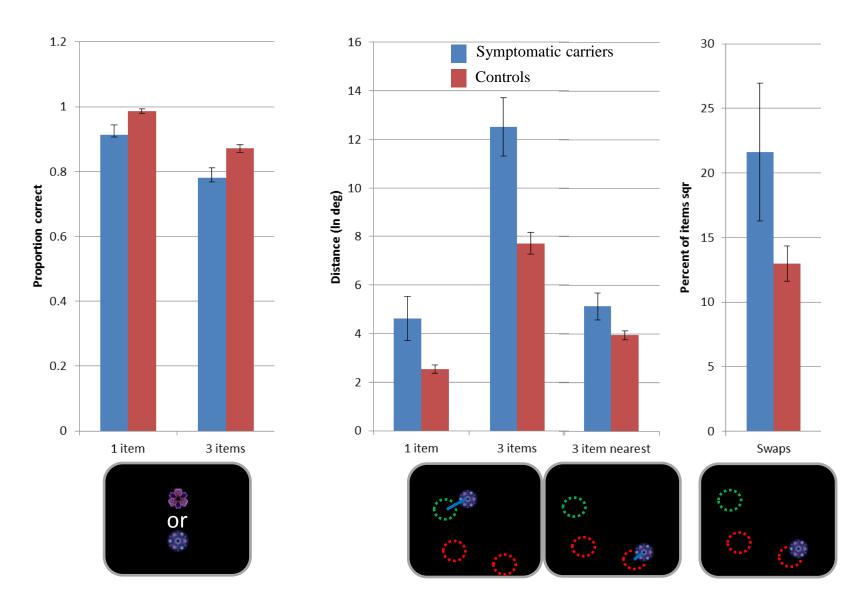
Back-up slides

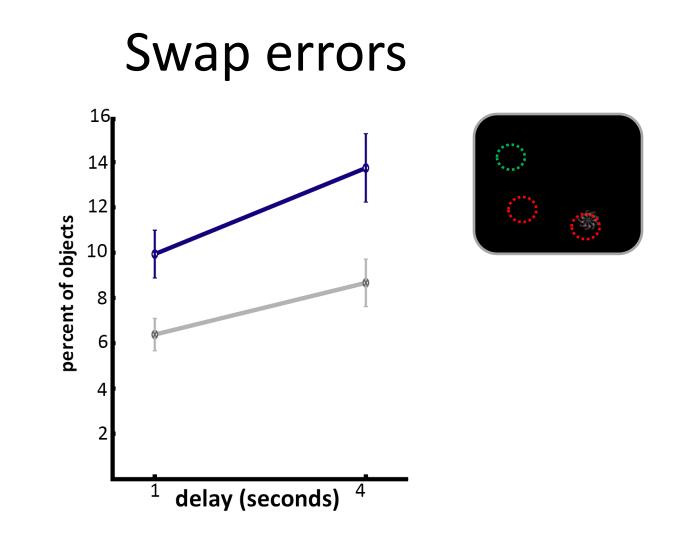


Symptomatic carriers vs. controls

Identification

Localisation performance





Contribution of swap errors to localization ?

Swap across time

