# Chapter 2: Studies of Human Learning and Memory

From *Mechanisms of Memory*, second edition By J. David Sweatt, Ph.D.

## Definitions

Learning: The acquisition of an altered behavioral response due to an environmental stimulus.

Memory: The processes through which learned information is stored.

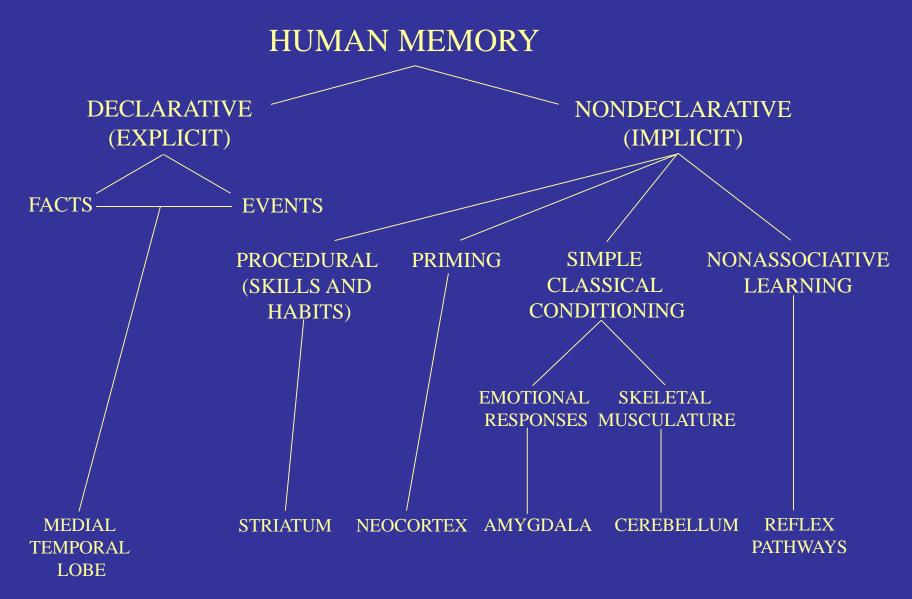
Recall:The conscious or unconscious retrieval<br/>process through which this<br/>alteredalteredbehavior is manifest.

### **Movies Using Amnesia as a Plot Device**

Movie Anterograde Amnesia: <i>Memento</i>	Director (plus Notable Actors)	
	Christopher Nolan (Guy Pearce)	2001
Finding Nemo	Andrew Stanton (Animated - Ellen DeGeneris as Dory)	2003
50 First Dates	Peter Segal (Adam Sandler)	2004
Retrograde Amnesia: The Bourne Identity	Doug Liman (Matt Damon)	2002
Total Recall	Paul Verhoeven (Arnold Schwarzenegger, Sharon Stone)	1990
Spellbound	Alfred Hitchcock (Ingrid Bergman, Gregory Peck)	1945
Eternal Sunshine of the Spotless Mind	Michel Gondry (Jim Carrey, Kate Winslet)	2004
The Forgotten	Joseph Ruben (Julianne Moore)	2004
Men in Black	Barry Sonnenfeld (Tommy Lee Jones, Will Smith)	1997
Not Sure: Mulholland Drive	David Lynch (Michael J. Anderson, Diane Baker)	2001



### **Subdivisions of Human Memory**



Milner B, Squire LR, Kandel ER: "Cognitive neuroscience and the study of memory". Neuron 1998, 20:445-468.

### **Multiple Memory Systems in the Human CNS**

Memory Subtype Working Memory	<b>Corresponding CNS Subregion</b> Prefrontal Cortex, contributions from Caudate Nucleus
Declarative, Episodic, and Spatial Memory	Medial Temporal Lobe system including the Hippocampus, Dentate Gyrus, Ento-rhinal and Peri- rhinal cortices
Habits, Motor Skills, Procedural Memory	Striatum (Caudate Nucleus and Putamen), Globus Pallidus*, and Cerebellum
Priming	Occipital Cortex, Neocortex in general
Aversive Associative Conditioning	Amygdala
Motoric Associative Conditioning	Cerebellum
Olfactory and Taste Conditioning	Olfactory Bulb, Insular Cortex, Nucleus Tractus Solitarius (NTS), positive and negative reinforcement systems (Amygdala or Nucleus Accumbens)
Sensitization, Habituation	Spinal Cord, Brainstem Nuclei, Amygdala
Circadian Rhythm	Hypothalamus – Suprachiasmatic Nucleus (SCN)
Reward, Positive Reinforcement, Addiction	Nucleus Accumbens, Ventral Tegmental Area (VTA)
to-lleader balance Devi	and Olahus Dallidus are referred to as the Decal

\*Collectively the Caudate Nucleus, Putamen, and Globus Pallidus are referred to as the Basal Ganglia.

#### Table 2

### Memory-associated Cognitive Dysfunction in Humans

#### **Disease or Syndrome**

Schizophrenia

**Major Depression** 

**Aging-related Dementias** 

Korsakoff's Syndrome (Alcoholism, Malnutrition)

Huntington's Disease and Parkinson's Disease

**Attention Deficit** 

Hyperactivity Disorder (ADHD)

Post-traumatic Stress Disorder\* (PTSD) Type of Memory Affected or Involved\*

Working Memory

**Declarative Memory** 

Long-term Declarative and Episodic Memory

Long-term Declarative and Episodic Memory

Motor Learning, Habit Learning

Generalized or Specific Learning

Disabilities, Dyslexia

Aversive Associative Conditioning, Lack of Extinction Likely Anatomical Locus

**Prefrontal Cortex** 

?

Temporal Lobe Hippocampal System

Mamillary Bodies, Limbic System, Thalamus

Nigro-striatal System

Frontal Lobes?

**Basal Ganglia?** 

Amygdala, Hippocampus

Table 3

#### Short-term Memory and the Multi-store Memory Model

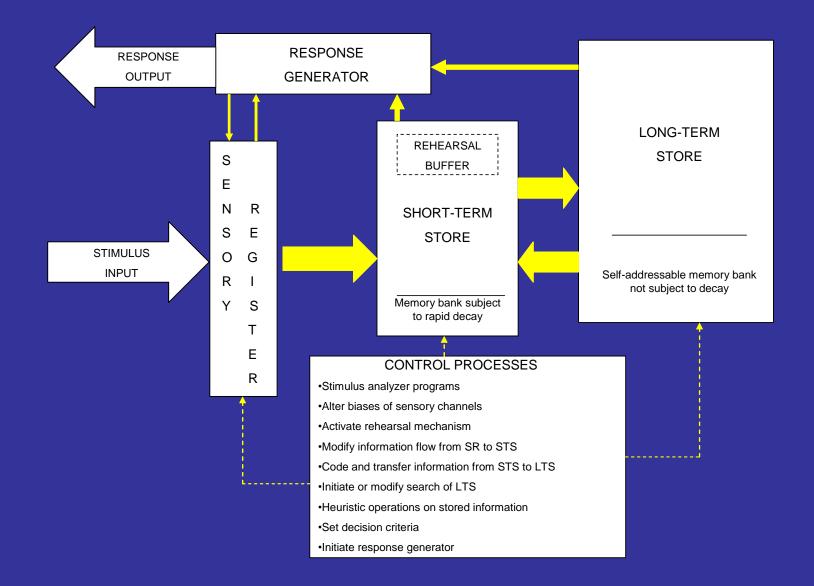


Figure 6A









#### Short-term Memory and the Multi-store Memory Model

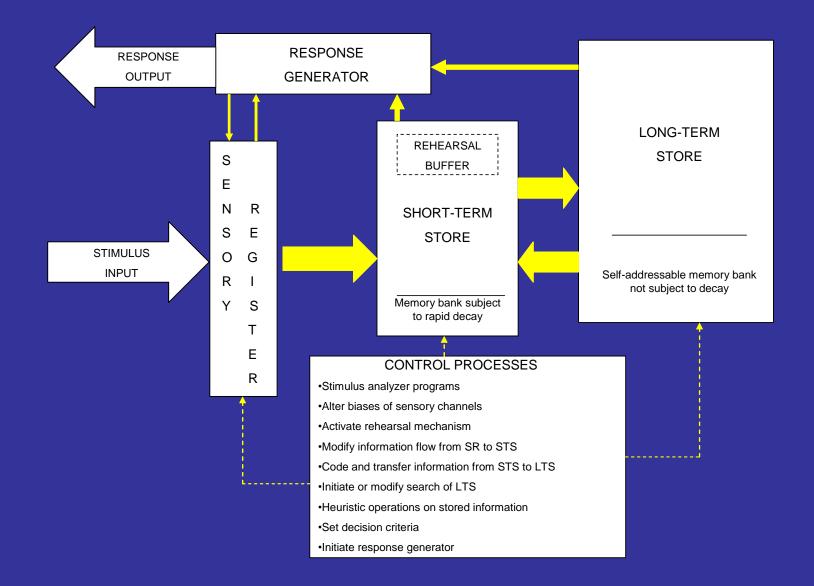
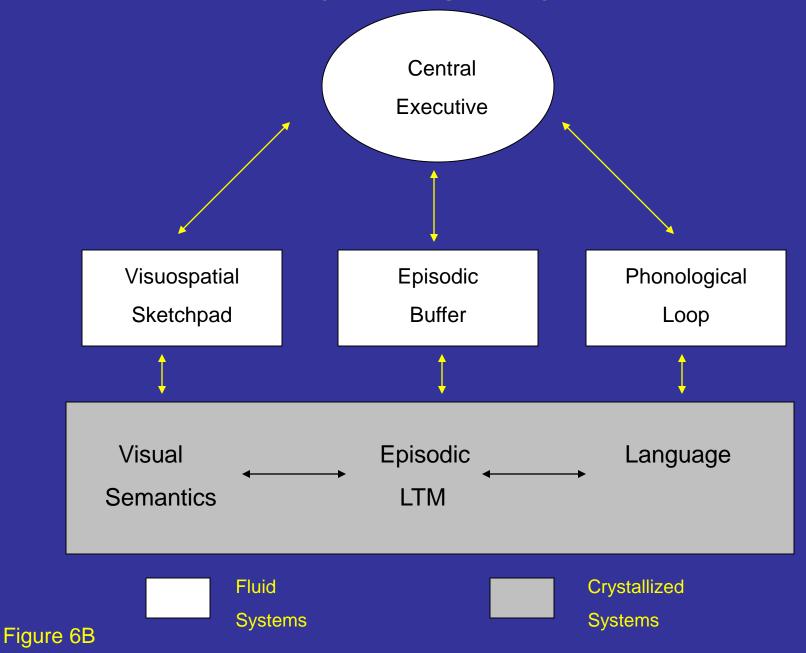


Figure 6A

#### **Baddeley's Working Memory Module**



### Anatomical Subdomains of Working Memory

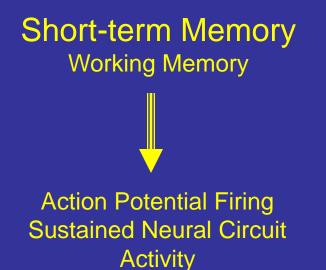
Prefrontal Cortex Working Memory

Ventrolateral PFC Non-spatial memory

(color, shape, etc)

Dorsolateral PFC Spatial memory

#### Figure 7

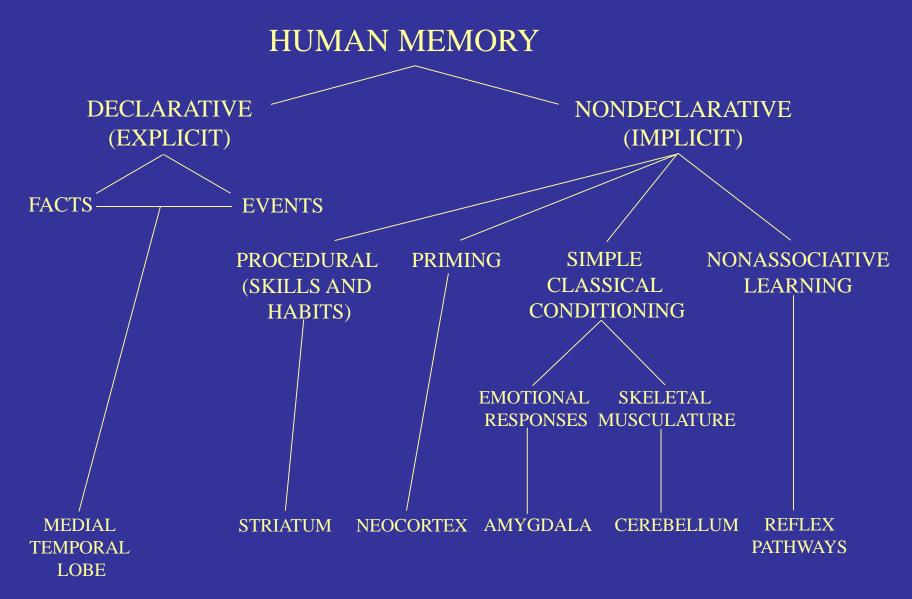


Long-term Memory

Persisting Molecular and Cellular Changes Changes in Synaptic Structure Anatomical Circuit Alterations

#### Figure 8

### **Subdivisions of Human Memory**

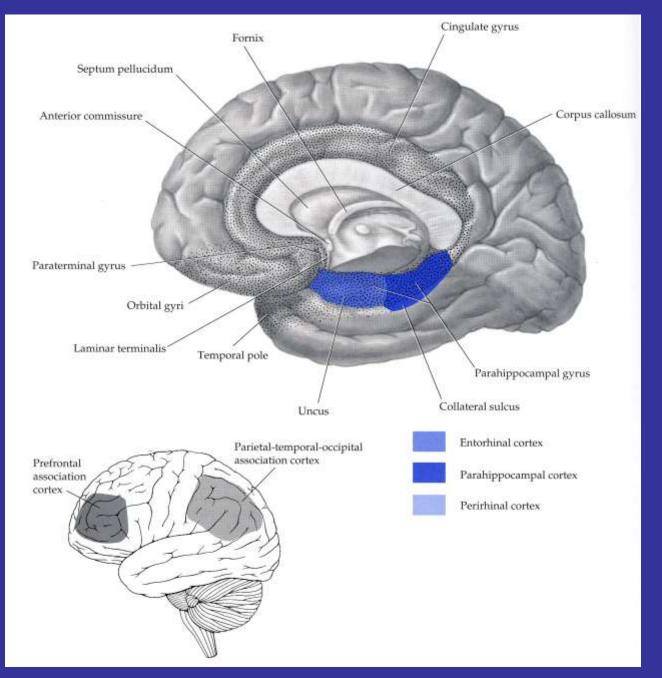


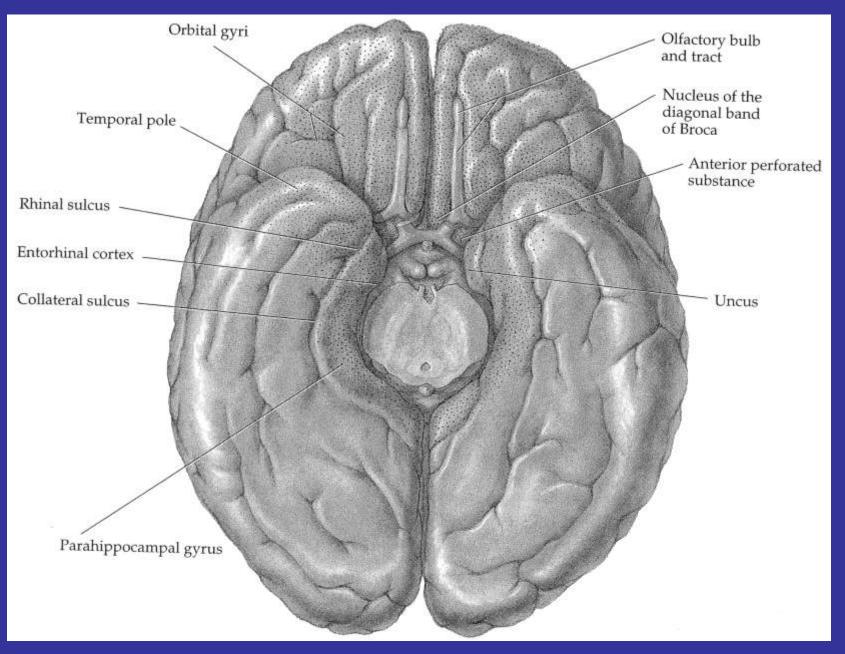
Milner B, Squire LR, Kandel ER: "Cognitive neuroscience and the study of memory". Neuron 1998, 20:445-468.

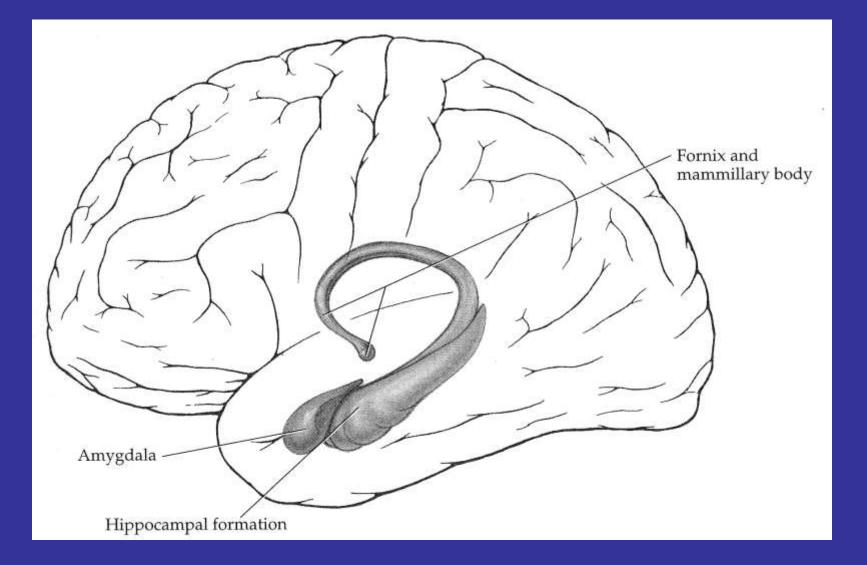
### **Functions of the Hippocampus**

 Cognitive Processing: Space Time Relationships

Memory Consolidation



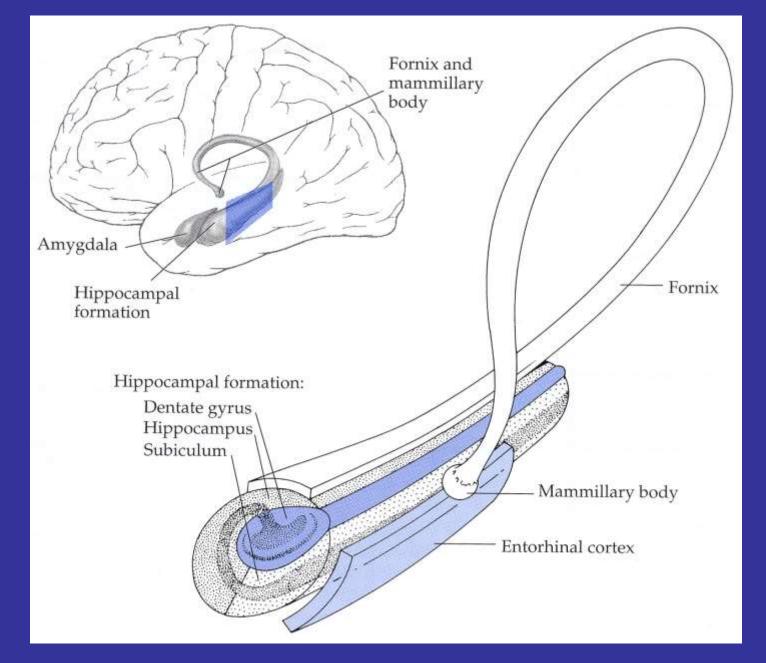




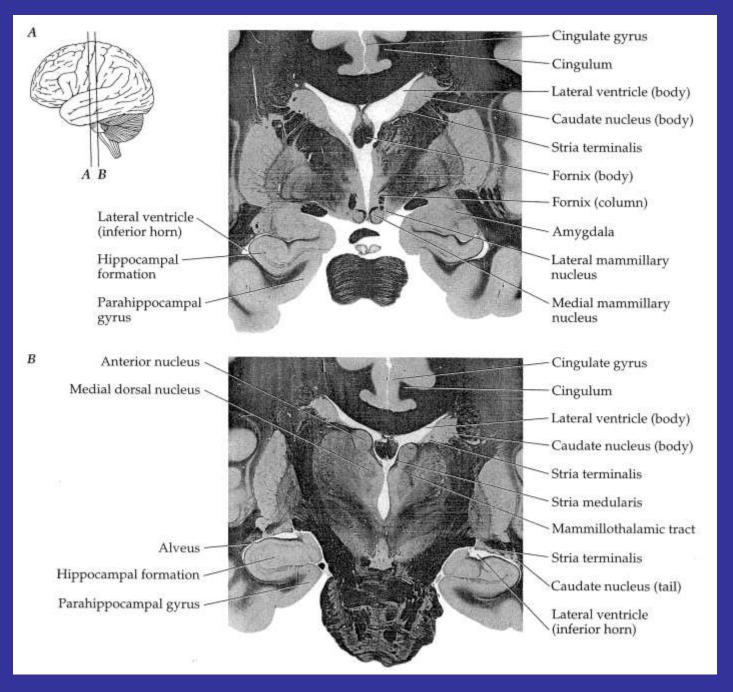


### Lined Seahorse, *Hippocampus erectus*

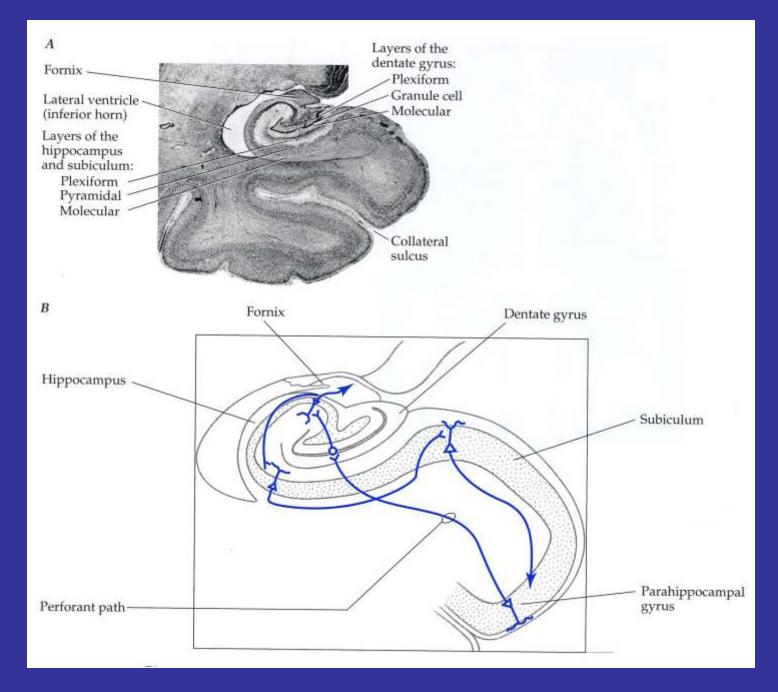
Vertebrate Field Zoology Photo #148, Courtesy Samford University



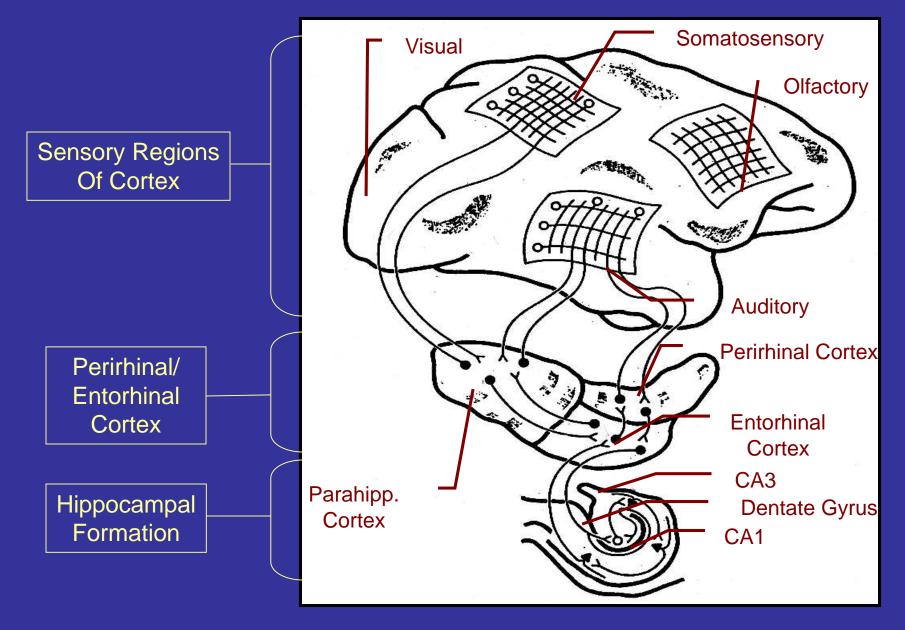
From JH Martin, *Neuroanatomy* 



From JH Martin, Neuroanatomy



### **Hippocampal Connectivity in the CNS**



### **Hippocampal Output Pathway & Intrinsic Circuit**

B Α CA1 **Entorhinal** Cortex CA3 Perforant Pathway Dentate **Entorhinal Cortex** Gyrus CA1 Outputs Mossy **Fibers** С CA3 CA1 Axon Schaffer ANDAD **Collaterals** GABAergic Interneuron CA1 Norepinephrine Acetylcholine Serotonin Amygdala, Subiculum/ Cortex **Entorhinal Cortex** Lateral Septum Schaffer Collaterals

Figure 5

### **Hippocampal Pyramidal Neuron**

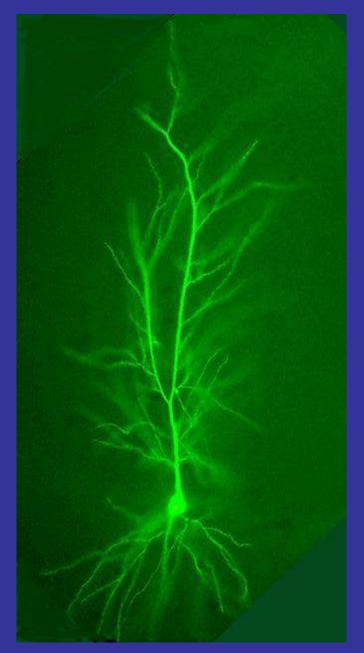


Figure 4

### Major Neurotransmitters in the Hippocampus

Neurotransmitter	Abbreviation, Synonym	General role
Serotonin	5HT, 5-hydroxytryptamine	neuromodulation
Dopamine	DA	neuromodulation
Acetylcholine	ACh	neuromodulation
Norepinephrine	NE	neuromodulation
Gamma-amino Butyric Acid	GABA	inhibitory transmission
Glutamate	Glu	excitatory transmission



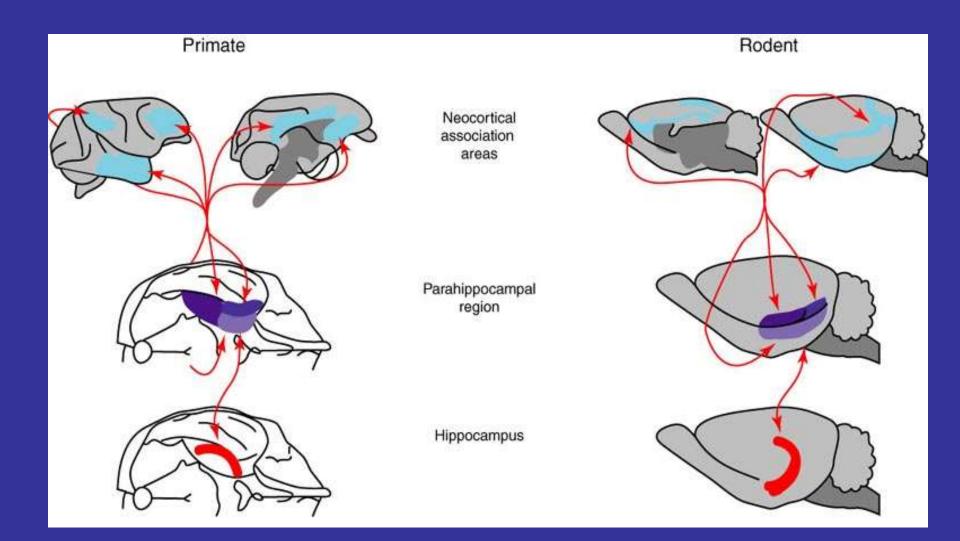


FIGURE 3 The anatomy of the hippocampal memory system in monkeys and rats. Multiple association areas in the cerebral cortex send outputs that converge on subdivisions of the cortex surrounding the hippocampus, which in turn sends its outputs to the hippocampus. The output path involves return projections from the hippocampus to the surrounding cortical region, which in turn projects back to the same cortical association areas. From Eichenbaum (2001).

### **Functions of the Hippocampus**

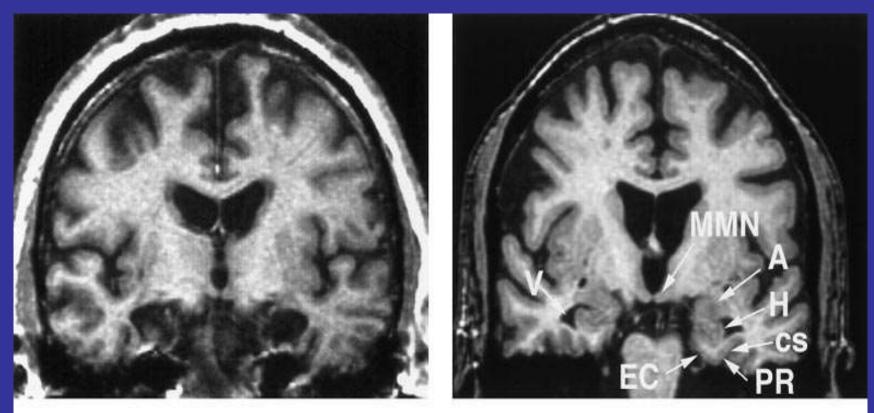
 Cognitive Processing: Space Time Relationships

Memory Consolidation

### MRI scan of H.M. and control brain

### H.M.

### Control

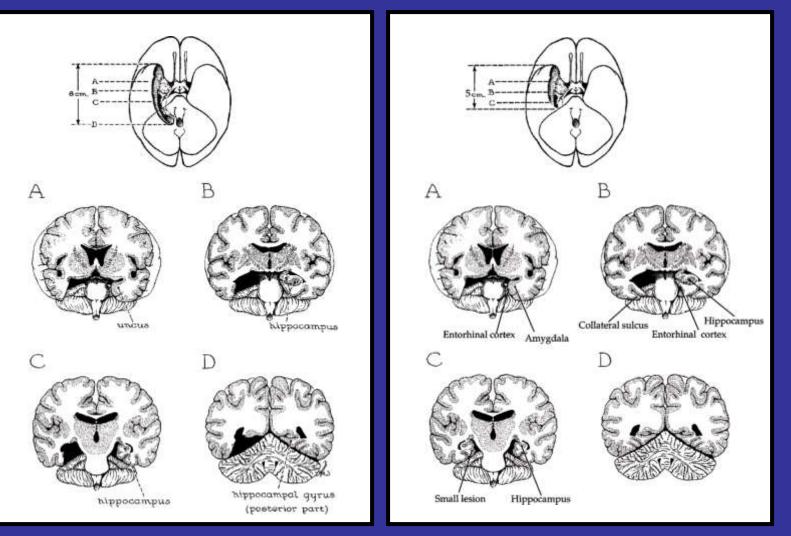


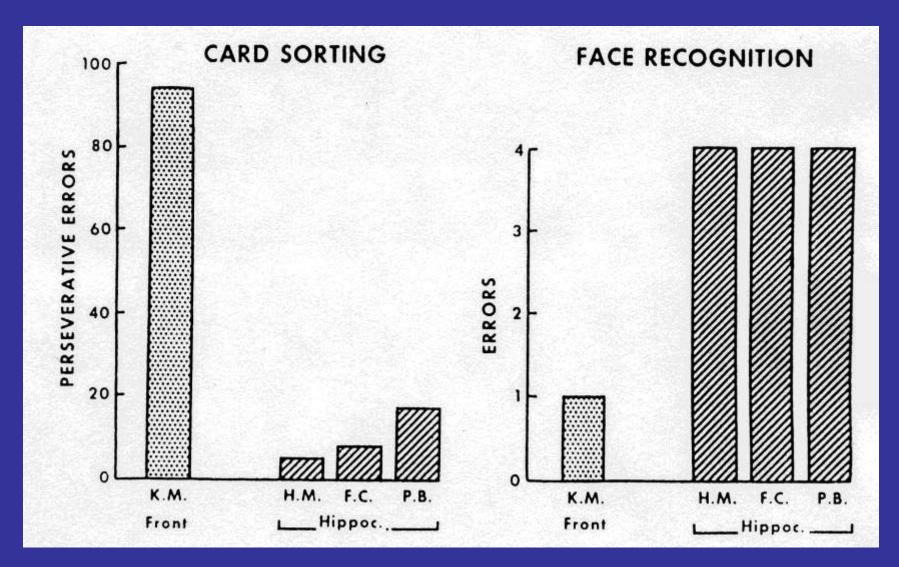
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### MRI of HM's Brain lesions

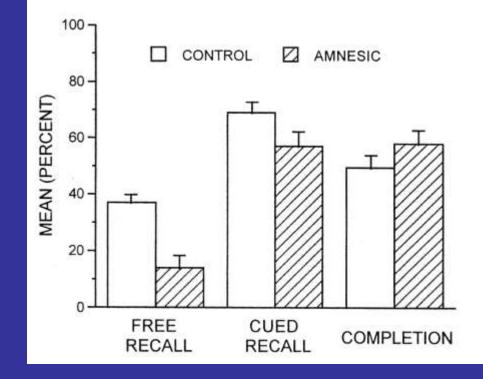
### Surgeon's Estimate of H.M.'s lesions

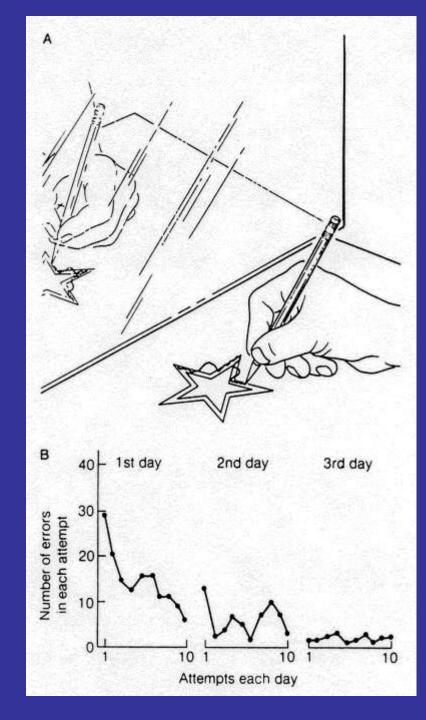
#### Revised Estimate based on MRI





ABSENT	ABS
INCOME	INC
FILLY	FIL
DISCUSS	DIS
CHEESE	CHE
ELEMENT	ELE





## **Four Types of Experiments**

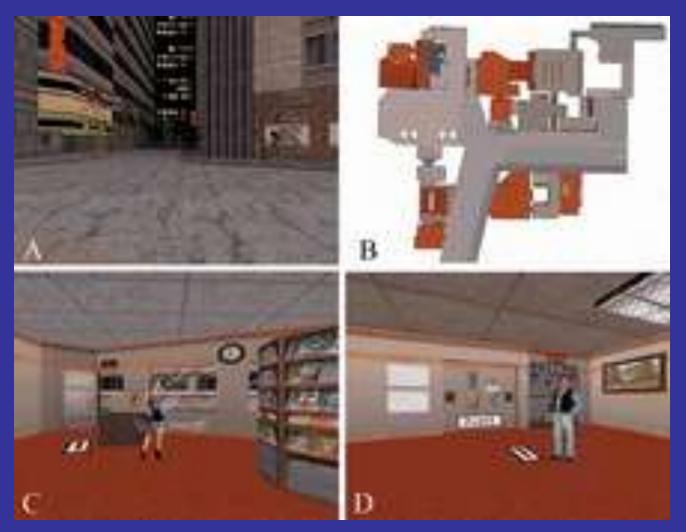
### Hypothesis: $A \rightarrow B \rightarrow C$

Experiment

**Prediction** 

Determine	<ul> <li>None (A makes C happen)</li> </ul>
Block	<ul> <li>Blocking B should block A causing C</li> </ul>
Mimic	<ul> <li>Activating B should cause C</li> </ul>
Measure	A makes B happen

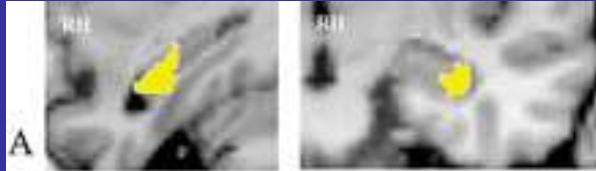
### Virtual Maze



Burgess N, Maguire EA, O'Keefe J. The human hippocampus and spatial and episodic memory. Neuron. 2002 35:625-41.

# **MRI of Hippocampus**

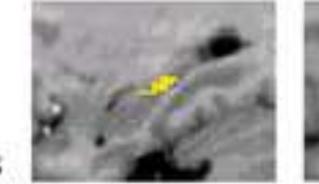
fMRI in Virtual Maze Navigation

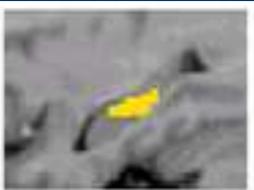


Morphometry of London Taxi Drivers

B

Right Posterior Hippocampal Volume Correlates with Time Spent Driving

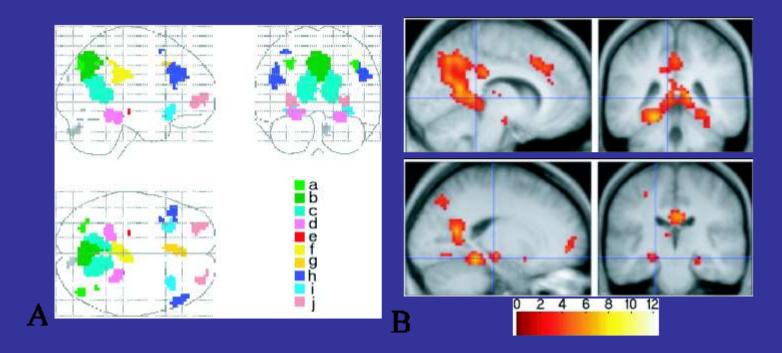


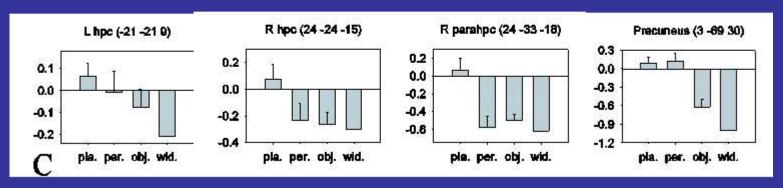




Burgess N, Maguire EA, O'Keefe J. The human hippocampus and spatial and episodic memory. Neuron. 2002 35:625-41.

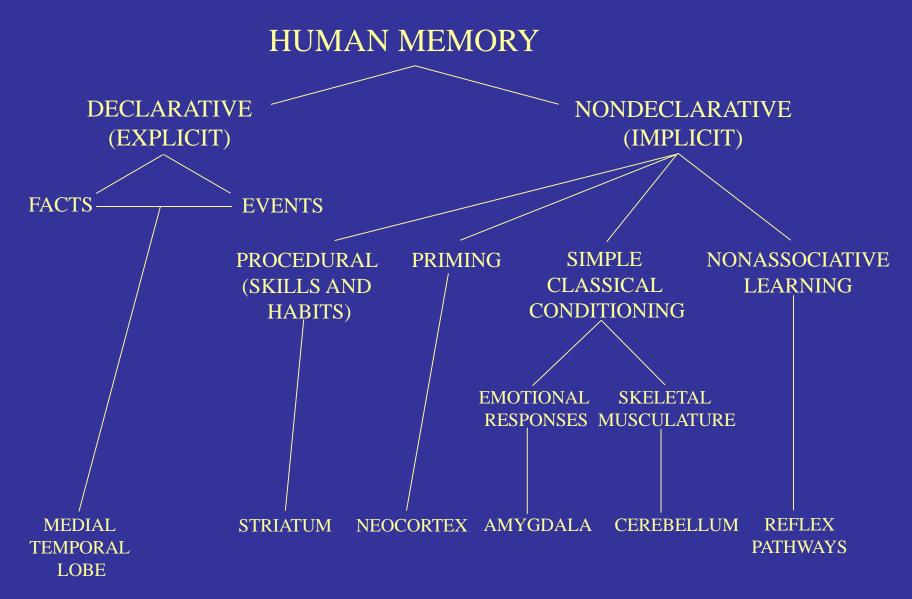
#### Neuroimaging of the retrieval of the Spatial Context of an Event





Blue Box 1

### **Subdivisions of Human Memory**

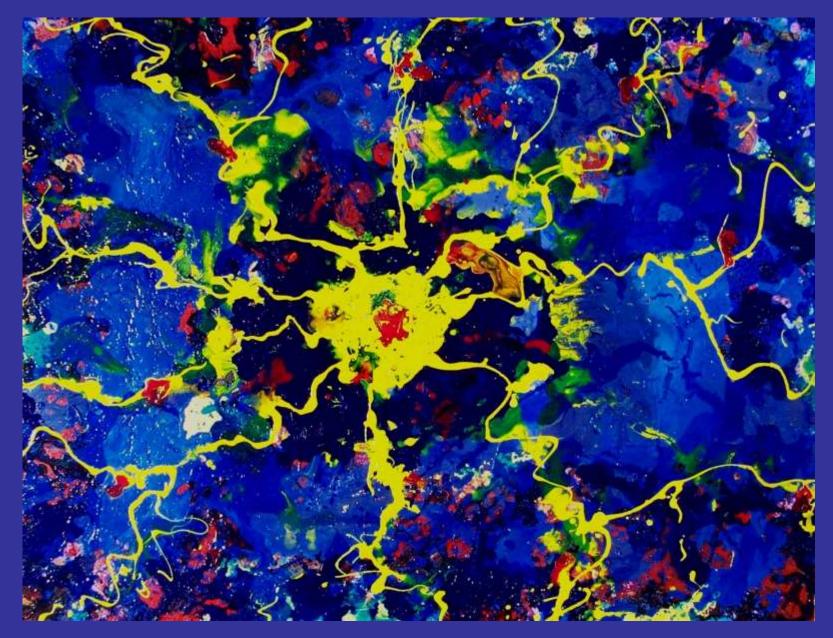


Milner B, Squire LR, Kandel ER: "Cognitive neuroscience and the study of memory". Neuron 1998, 20:445-468.

#### Human Excellence in Motor Learning and Memory







### **Medium Spiny Neuron**

#### **Summary Scheme of Interaction Between Motor Centers**

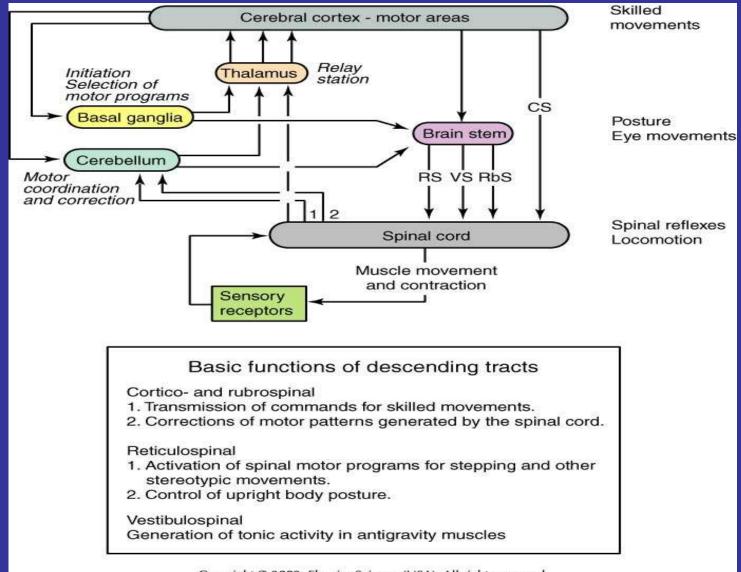
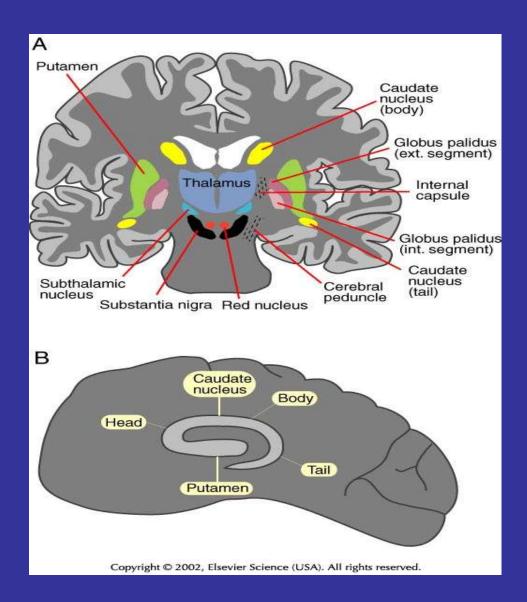


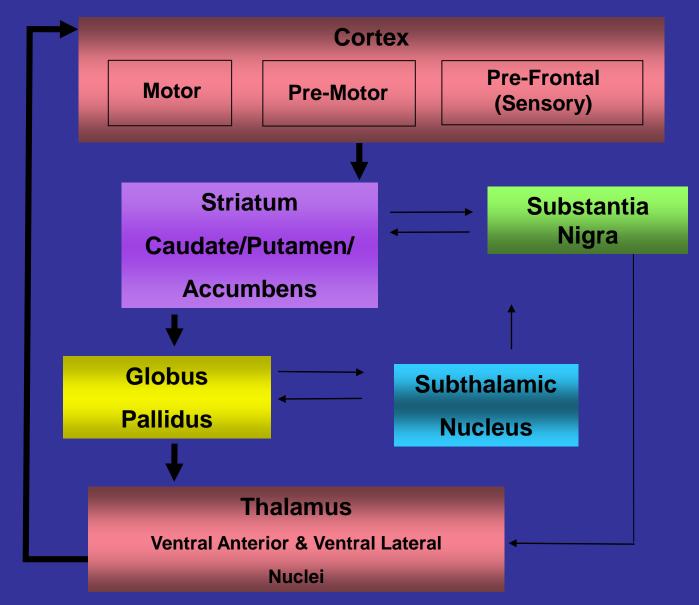
Figure 11

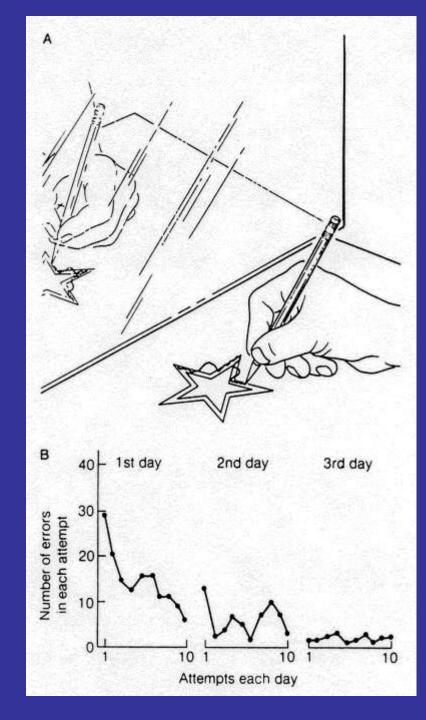
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#### **Basal Ganglia in the Human Brain**



#### **The Striatal Motor Learning System**





#### Memory-associated Cognitive Dysfunction in Humans

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**Major Depression** 

**Aging-related Dementias** 

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Table 3

### Human Excellence in Declarative Learning and Memory

### Rajan Srinavasen Mahadevan

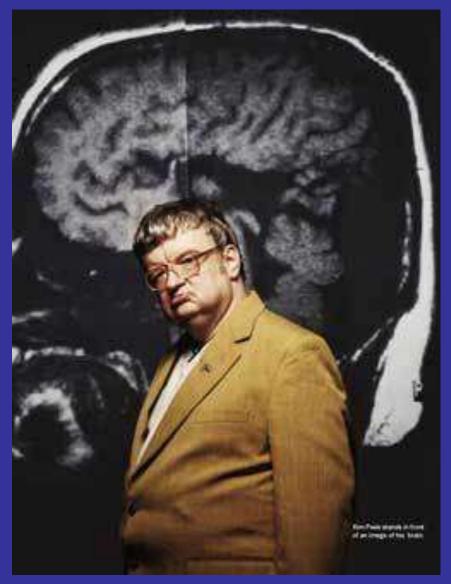


Figure 14

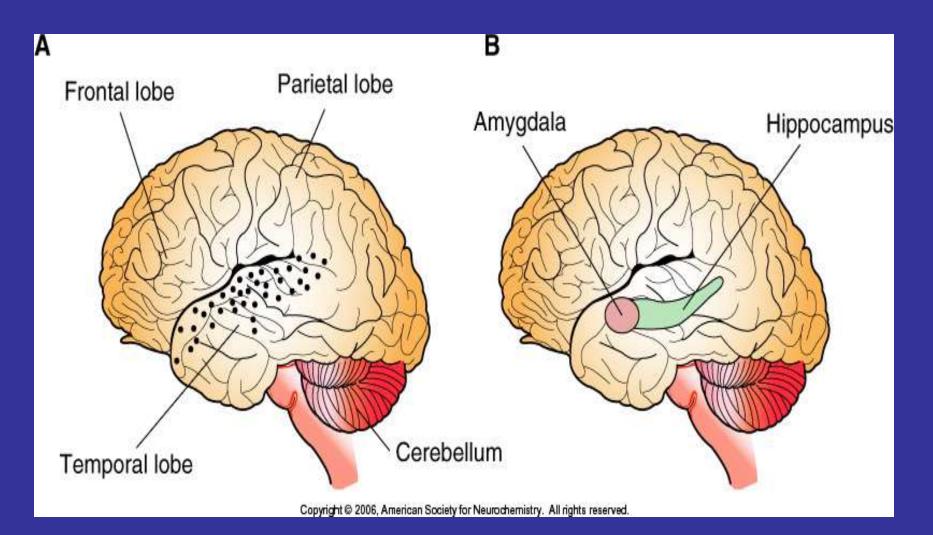
#### 1000 Decimal places of pi

3.14159265358979323846264338327950288419716939937510582097494459 577805321712268066130019278766111959092164201989....

### Kim Peek



### **The Penfield Stimulation Experiments**



# The Mimic Octopus





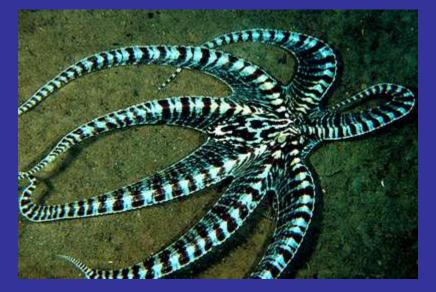






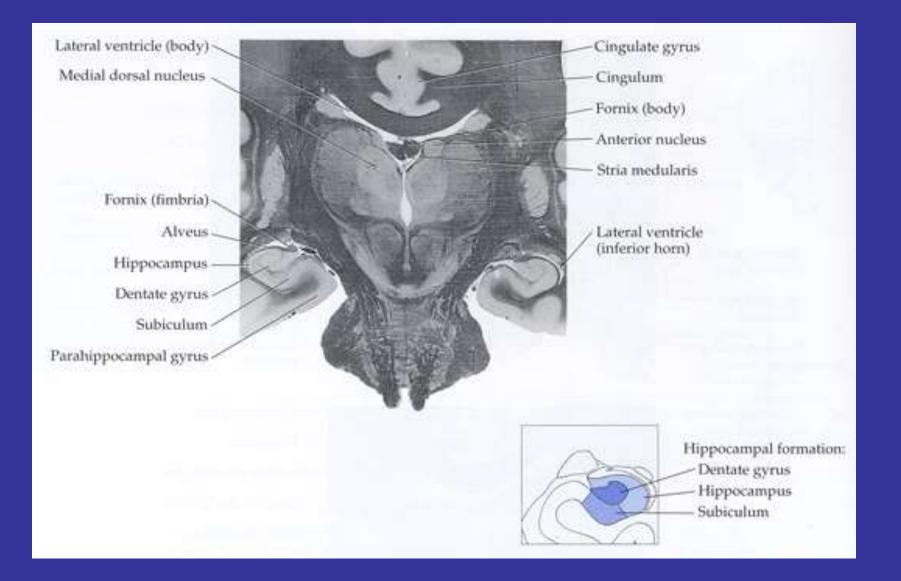




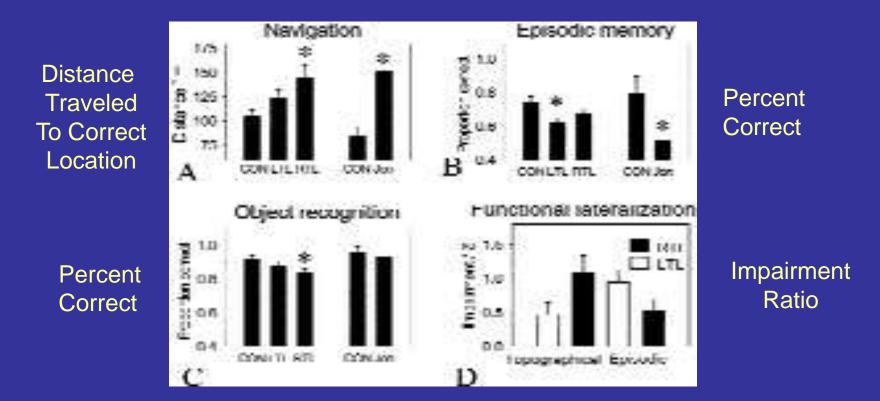


# The Mimic Octopus





## Lateralization of Hippocampal Function



#### Studies of hippocampectomy patients

#### Burgess N, Maguire EA, O'Keefe J.

The human hippocampus and spatial and episodic memory. Neuron. 2002 35:625-41.