

# What fMRI Can, Can't, and Might Do

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Unit on Functional Imaging Methods  
&  
3T Neuroimaging Core Facility

Laboratory of Brain and Cognition  
National Institute of Mental Health

- Functional Imaging

- Xenon Computerized Tomography (Xe CT)

- Positron Emission Tomography (PET)

- Single Photon Computed Tomography (SPECT)

- Functional MRI (fMRI)

- Electroencephalography (EEG)

- Magnetoencphalography (MEG)

- Transcranial Magnetic Stimulation (TMS)

# MRI vs. fMRI

high resolution  
(1 mm)

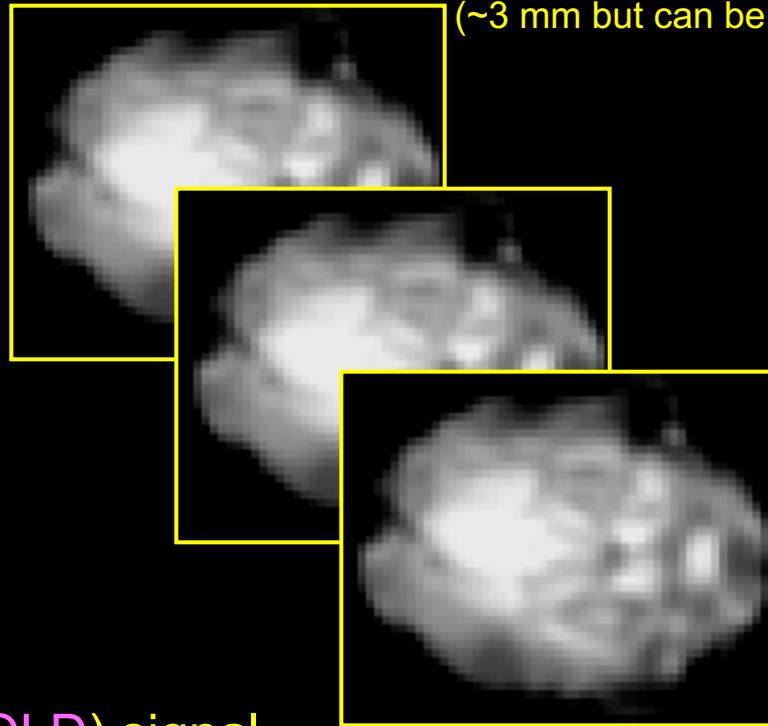
MRI



one image

fMRI

low resolution  
(~3 mm but can be better)



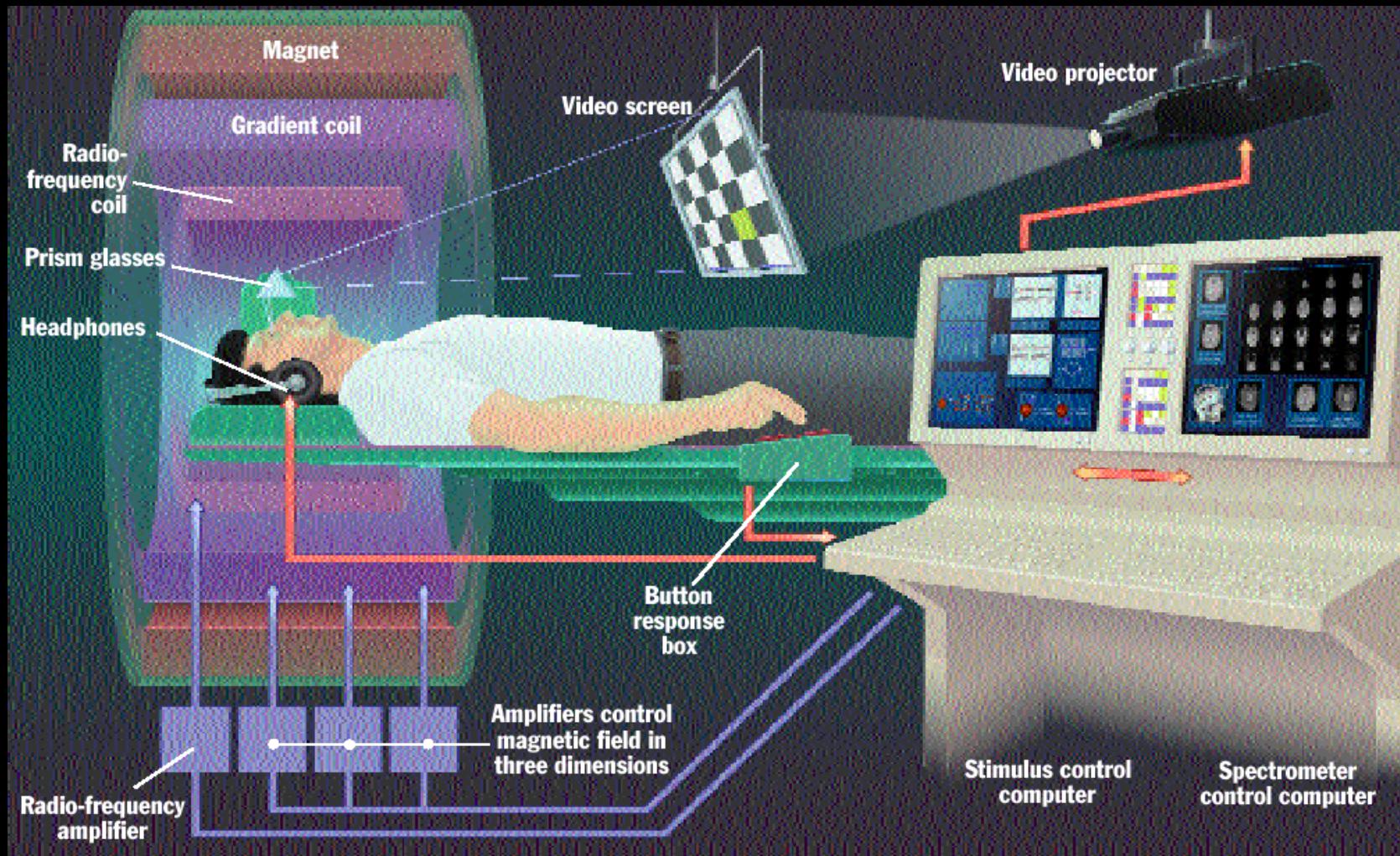
fMRI

Blood Oxygenation Level Dependent (BOLD) signal  
indirect measure of neural activity

many images  
(e.g., every 2 sec for 5 mins)

↑ neural activity → ↑ blood oxygen → ↑ fMRI signal

# fMRI Setup

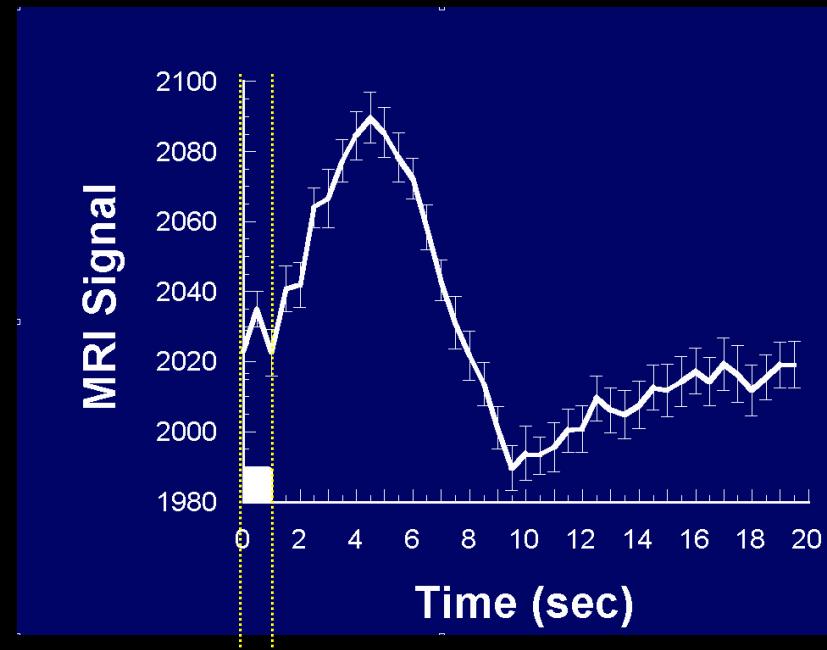
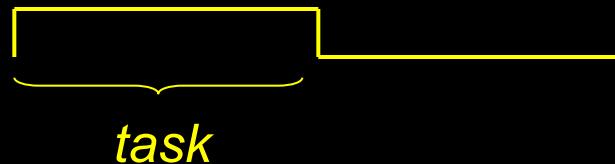
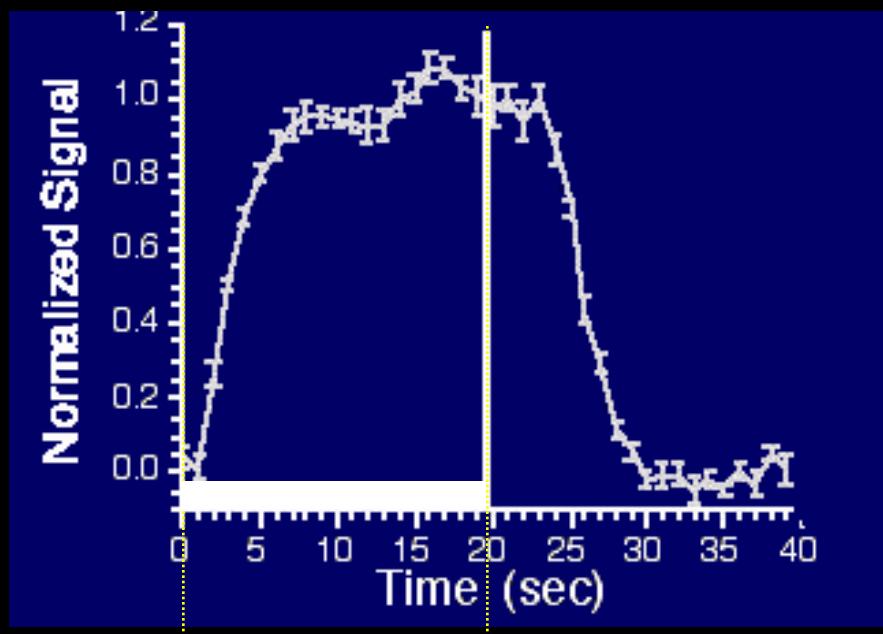


Courtesy, Robert Cox,  
Scientific and Statistical  
Computing Core Facility,  
NIMH



# The FMRI Signal

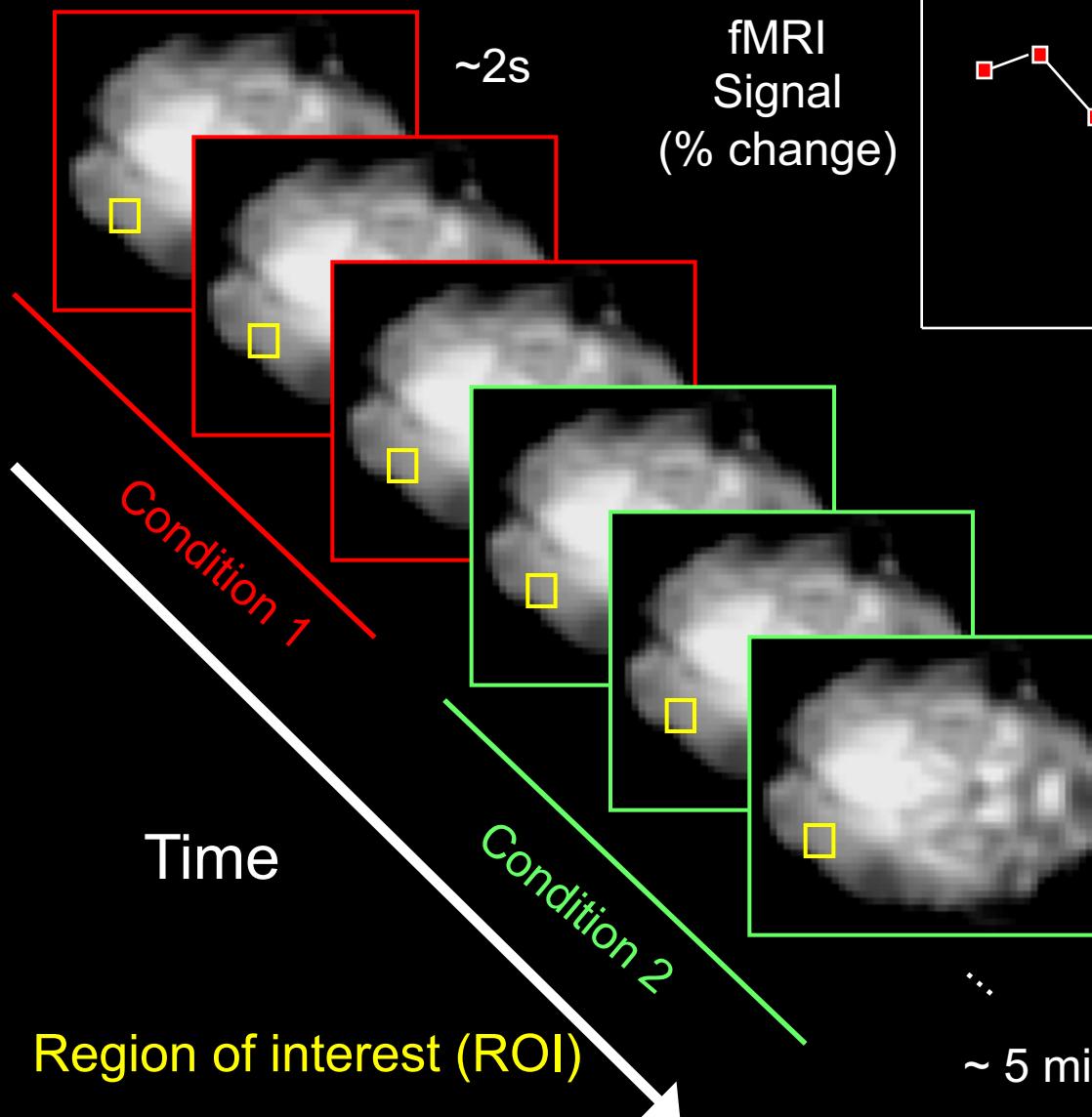
Based on Local Blood Flow Response in the Brain



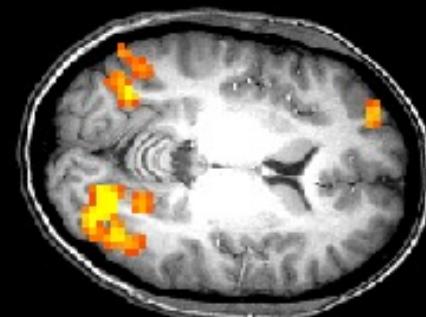


# Activation Statistics

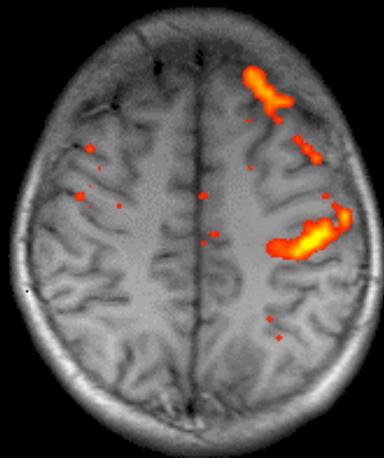
Functional images



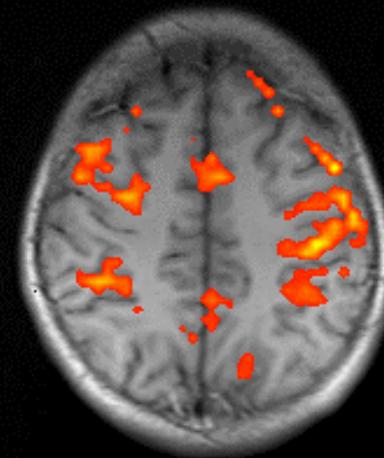
Statistical Map  
superimposed on  
anatomical MRI image



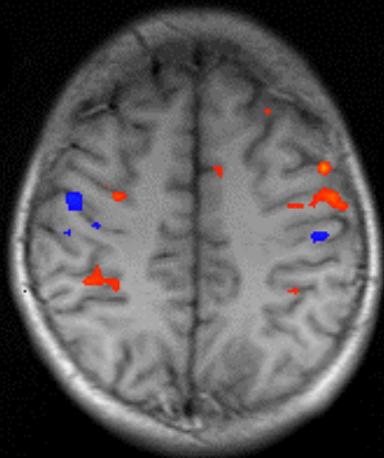
Simple Right



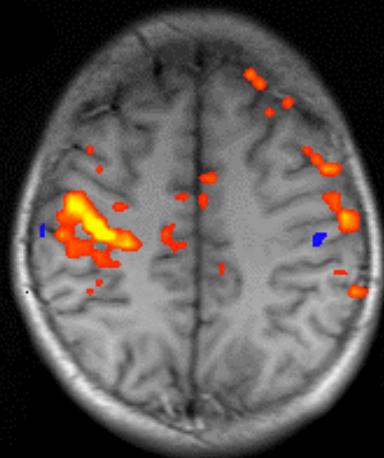
Complex Right



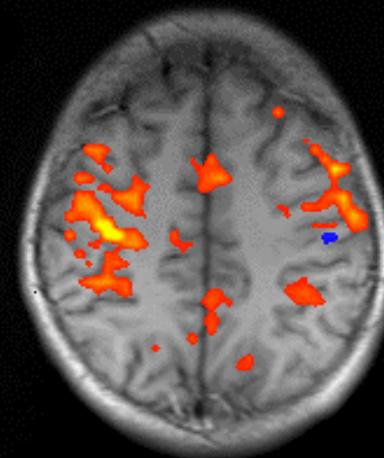
Imagined  
Complex Right



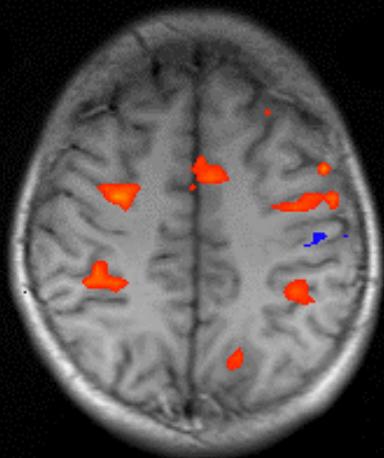
Simple Left



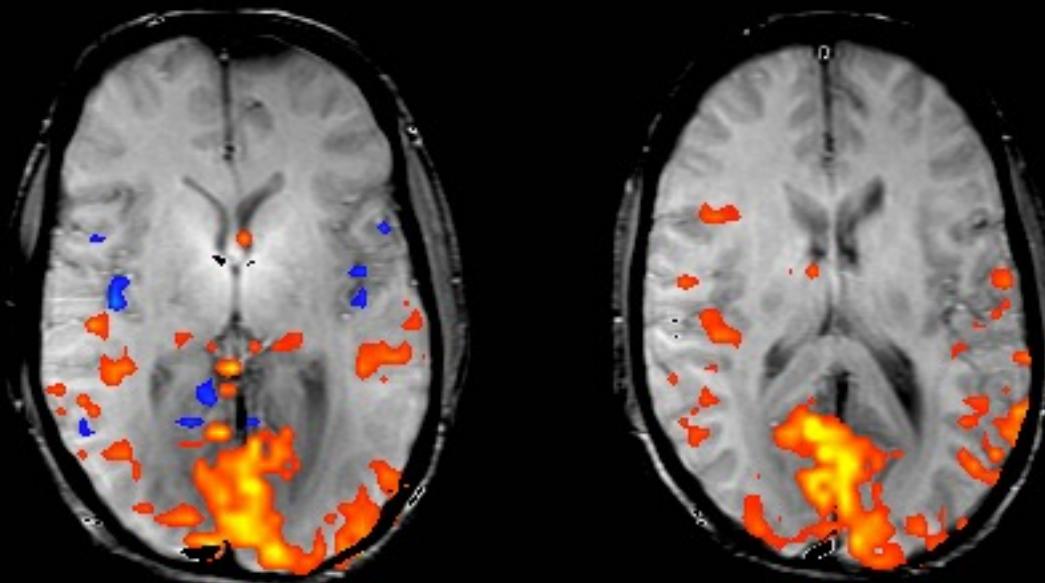
Complex Left



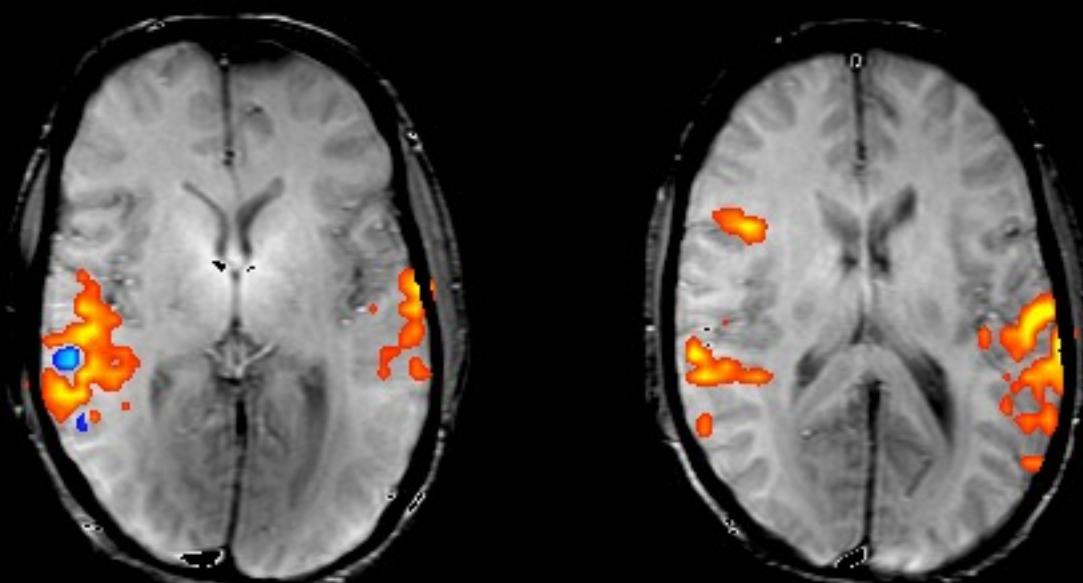
Imagined  
Complex Left



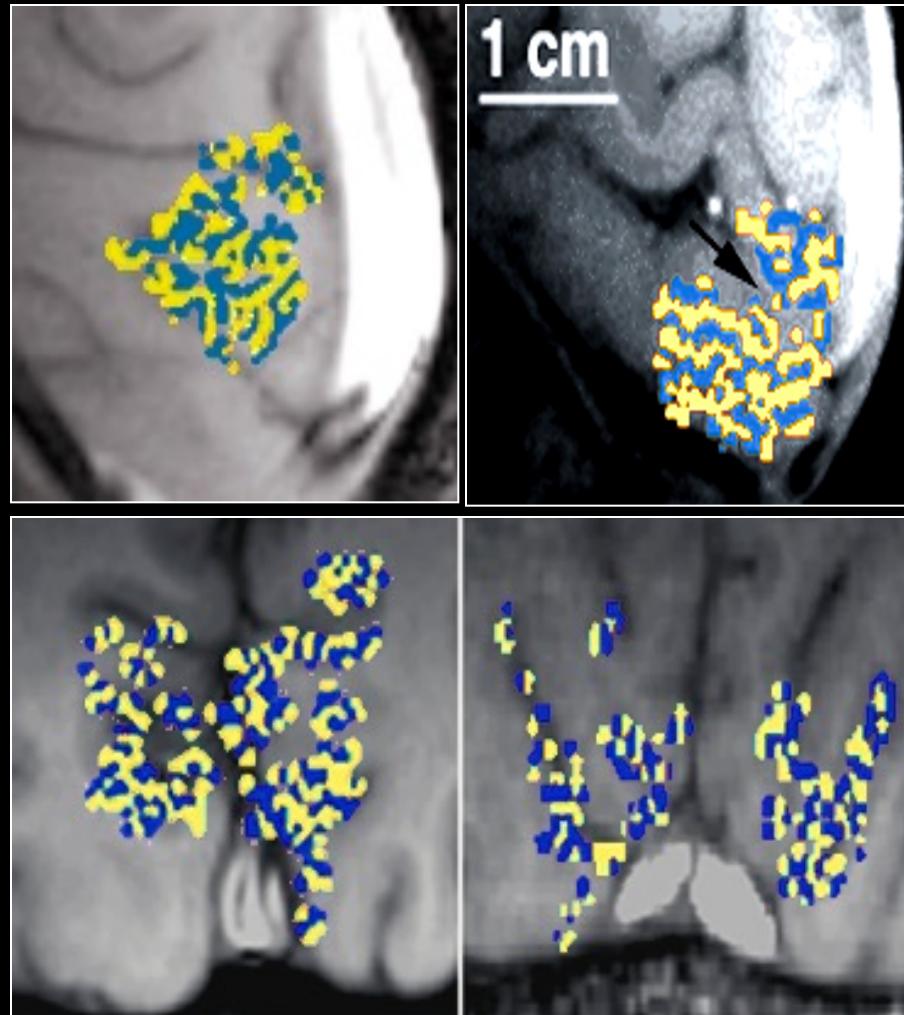
Reading



Listening

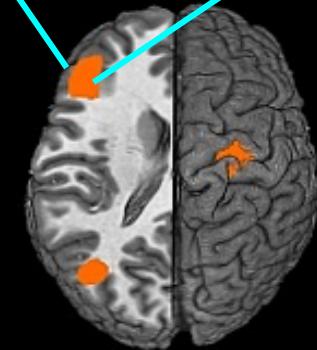
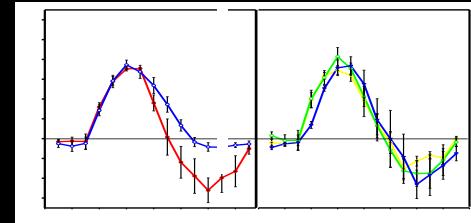


## Spatial Resolution: Ocular Dominance Column Mapping



## Temporal Resolution: Word processing

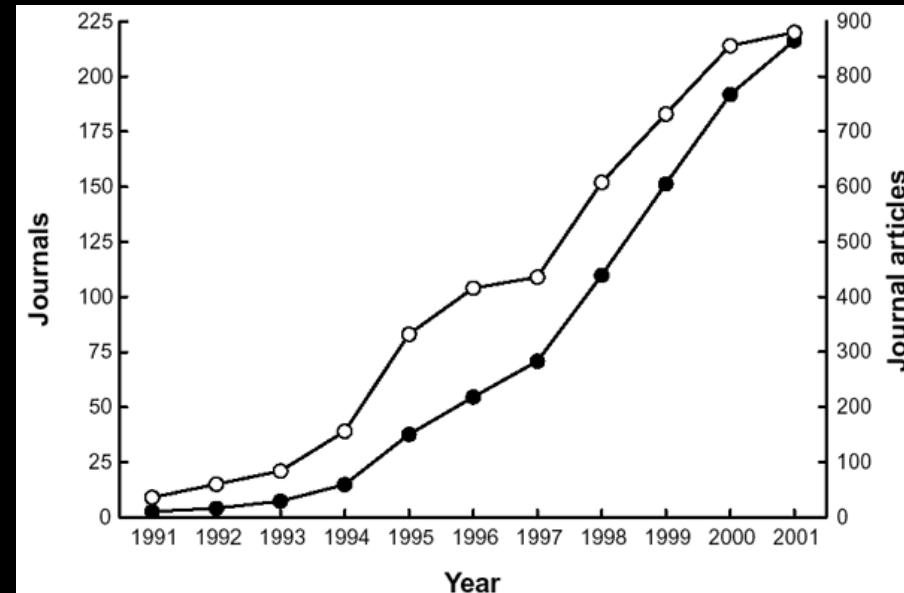
**Word vs. Non-word**  
**0°, 60°, 120° Rotation**



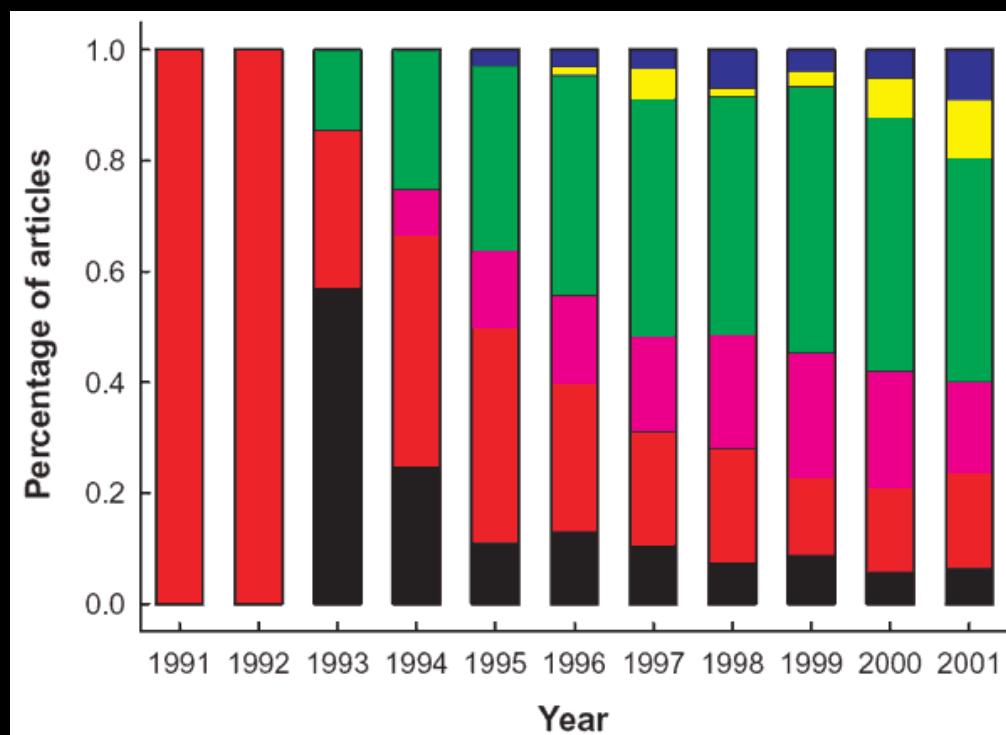
Bellgowan, et al (2003), PNAS 100, 15820–15283

Menon, et al. (1997). J Neurophysiol 77(5): 2780-7.

J. Illes, M. P. Kirschchen,  
J. D. E. Gabrielli,  
Nature Neuroscience,  
6 (3)m p.205

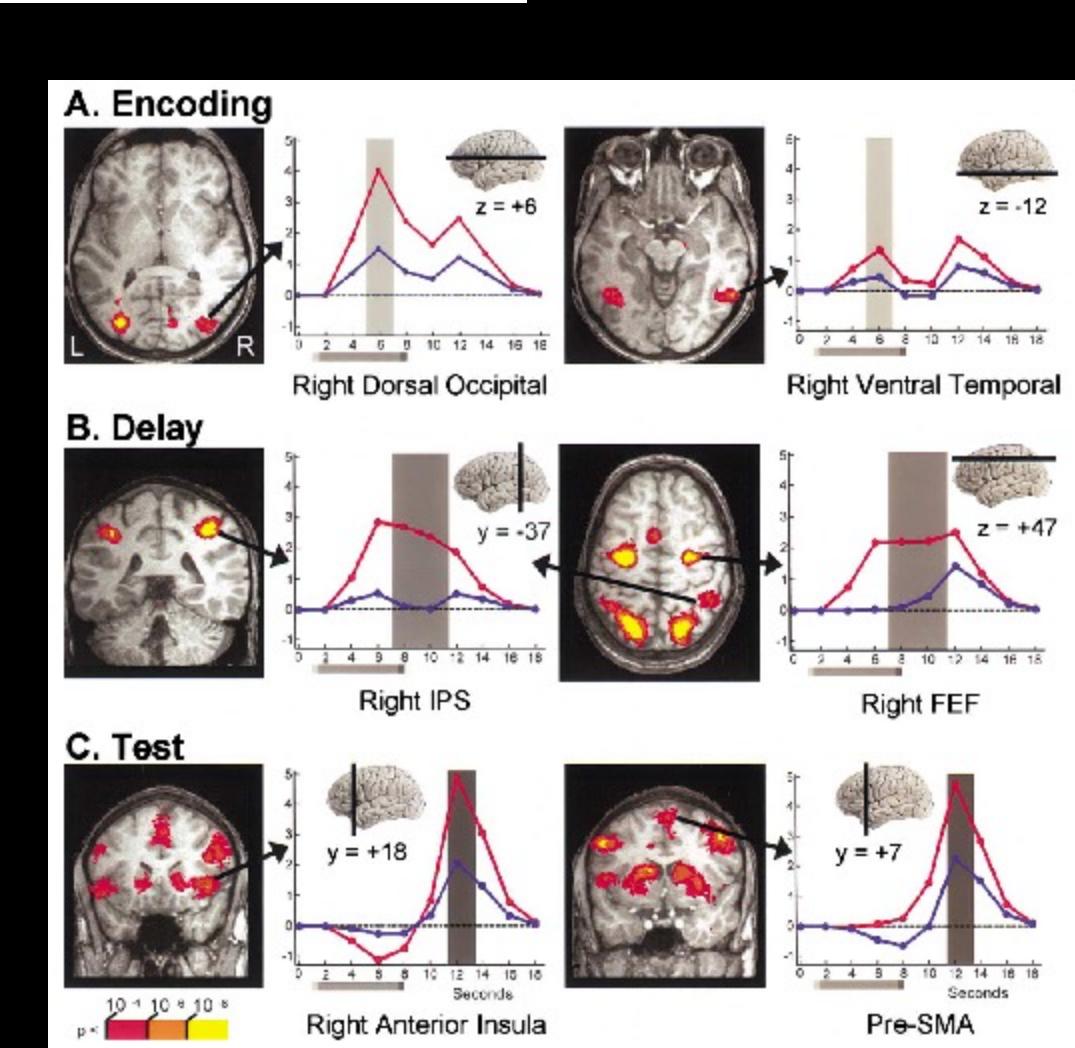
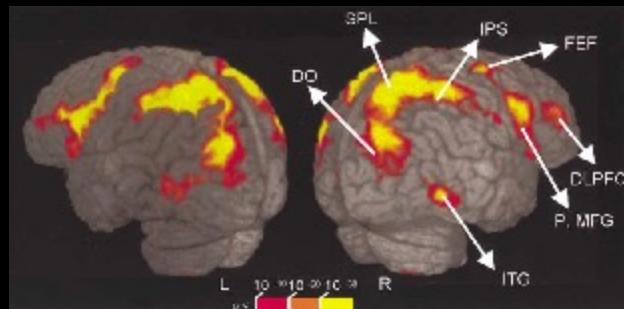


**Motor (black)**  
**Primary Sensory (red)**  
**Integrative Sensory (violet)**  
**Basic Cognition (green)**  
**High-Order Cognition (yellow)**  
**Emotion (blue)**



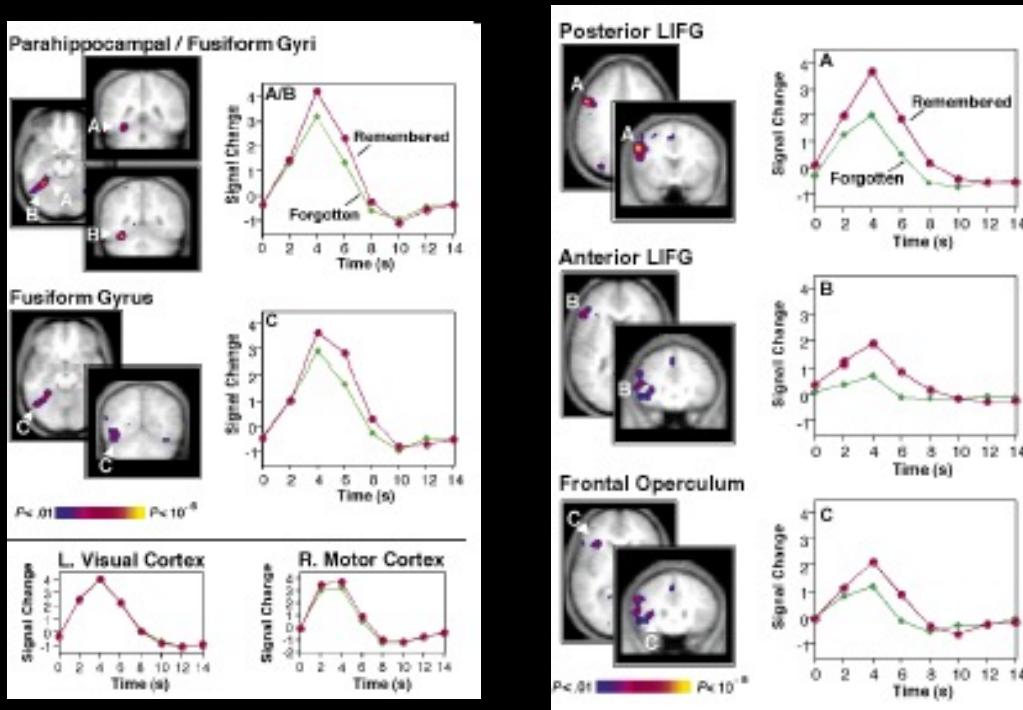
# Neural Correlates of Visual Working Memory: fMRI Amplitude Predicts Task Performance

Luiz Pessoa,<sup>1</sup> Eva Gutierrez, Peter A. Bandettini,  
and Leslie G. Ungerleider  
Laboratory of Brain and Cognition  
National Institute of Mental Health  
National Institutes of Health  
Bethesda, Maryland 20892



# Building Memories: Remembering and Forgetting of Verbal Experiences as Predicted by Brain Activity

Anthony D. Wagner,\* Daniel L. Schacter, Michael Rotte,†  
Wilma Koutstaal, Anat Maril, Anders M. Dale, Bruce R. Rosen,  
Randy L. Buckner



# Classification of spatial patterns

Functional magnetic resonance imaging (fMRI) "brain reading":  
detecting and classifying distributed patterns of fMRI activity  
in human visual cortex

David D. Cox<sup>a,b,\*</sup> and Robert L. Savoy<sup>a,b,c</sup>

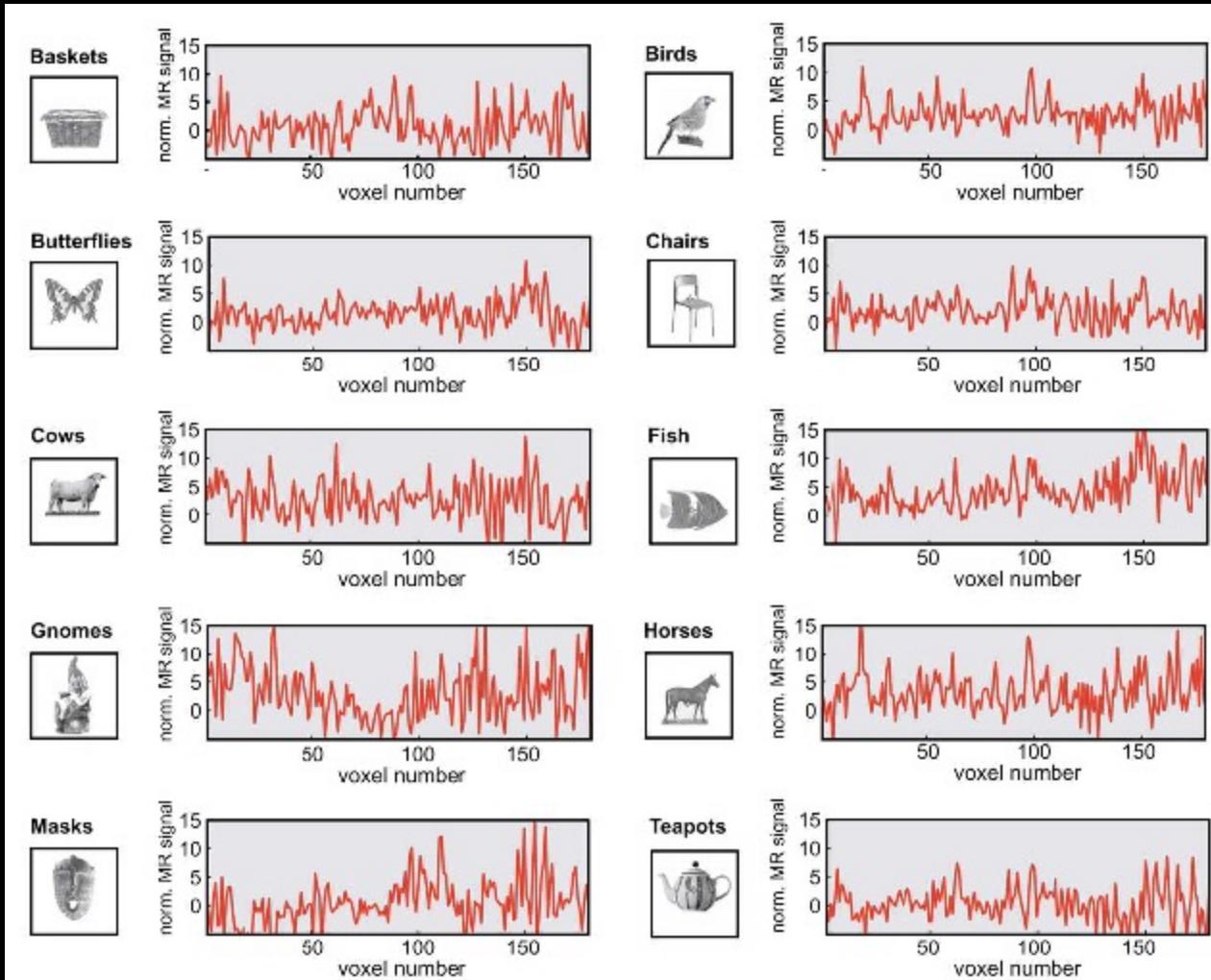
<sup>a</sup> Rowland Institute for Science, Cambridge, MA 02142, USA

<sup>b</sup> Athinoula A. Martinos Center for Structural and Functional Biomedical Imaging, Charlestown, MA 02129, USA

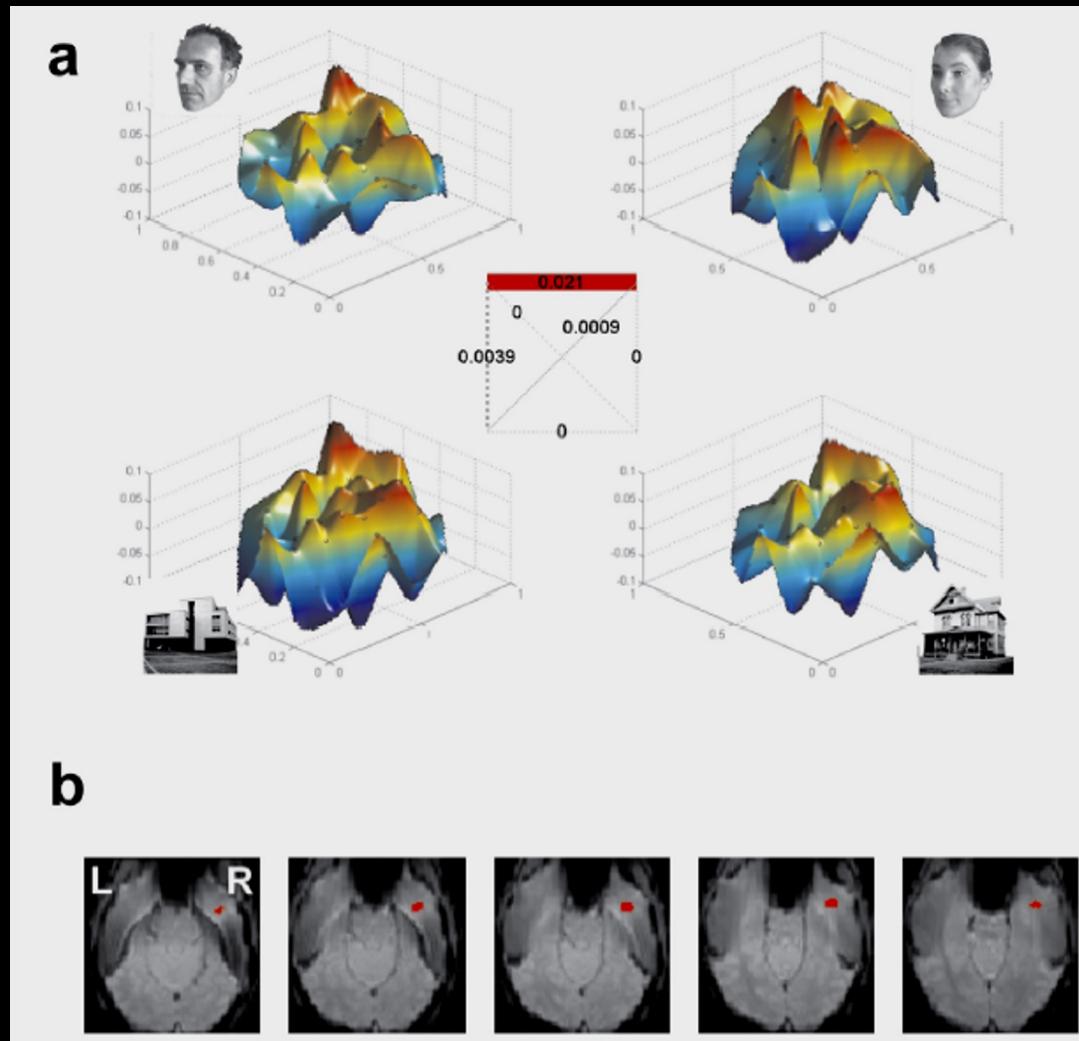
<sup>c</sup> HyperVision, Inc., P.O. Box 158, Lexington, MA 02420, USA

Received 15 July 2002; accepted 10 December 2002

NEUROIMAGE 19 (2): 261-270 Part 1 JUN 2003

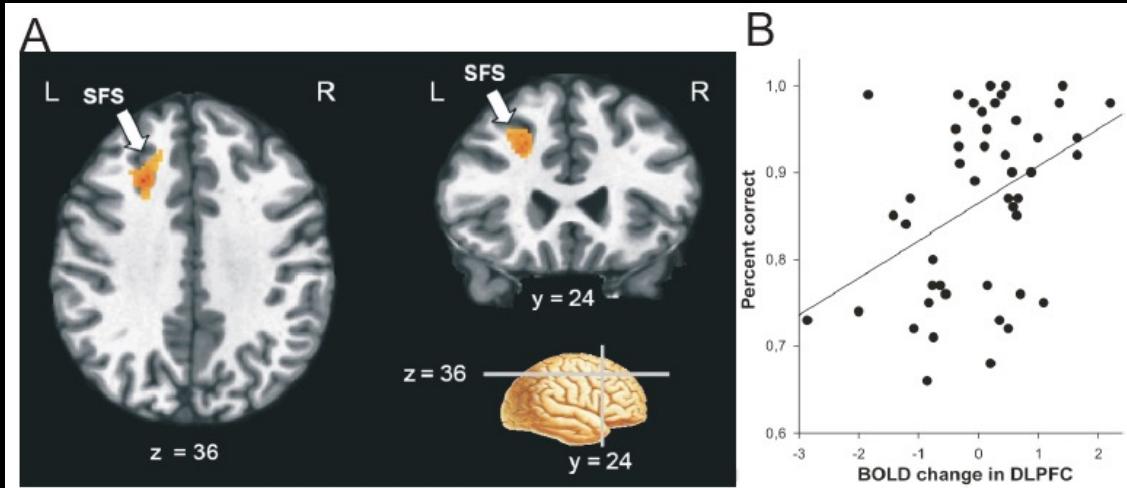
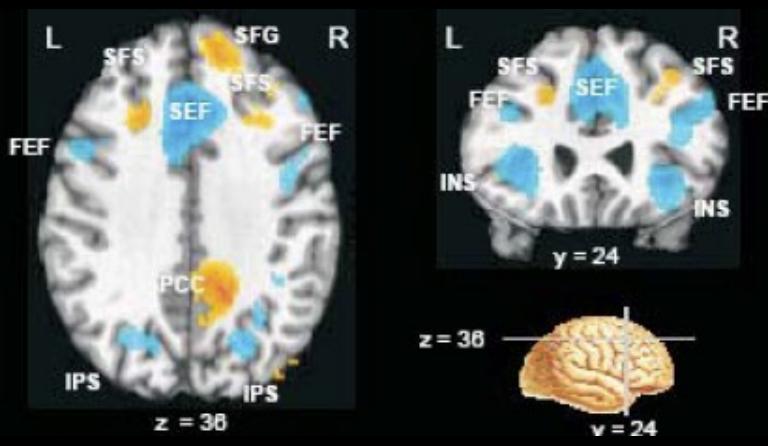
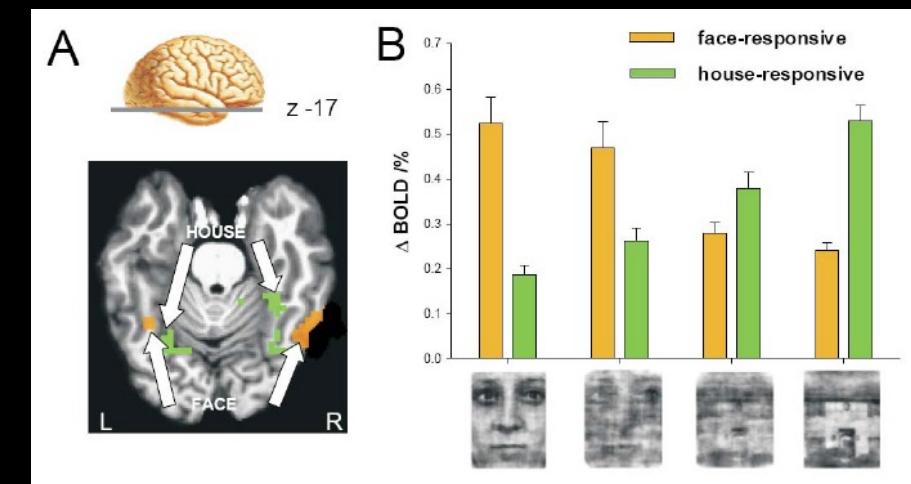
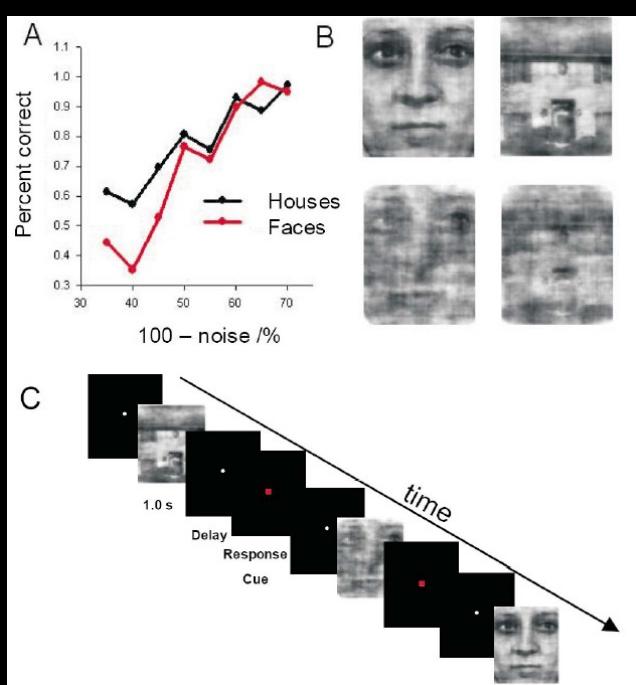


# Voxel-Wise Mapping of Pattern Differences



Kriegeskorte et al. HBM 2004

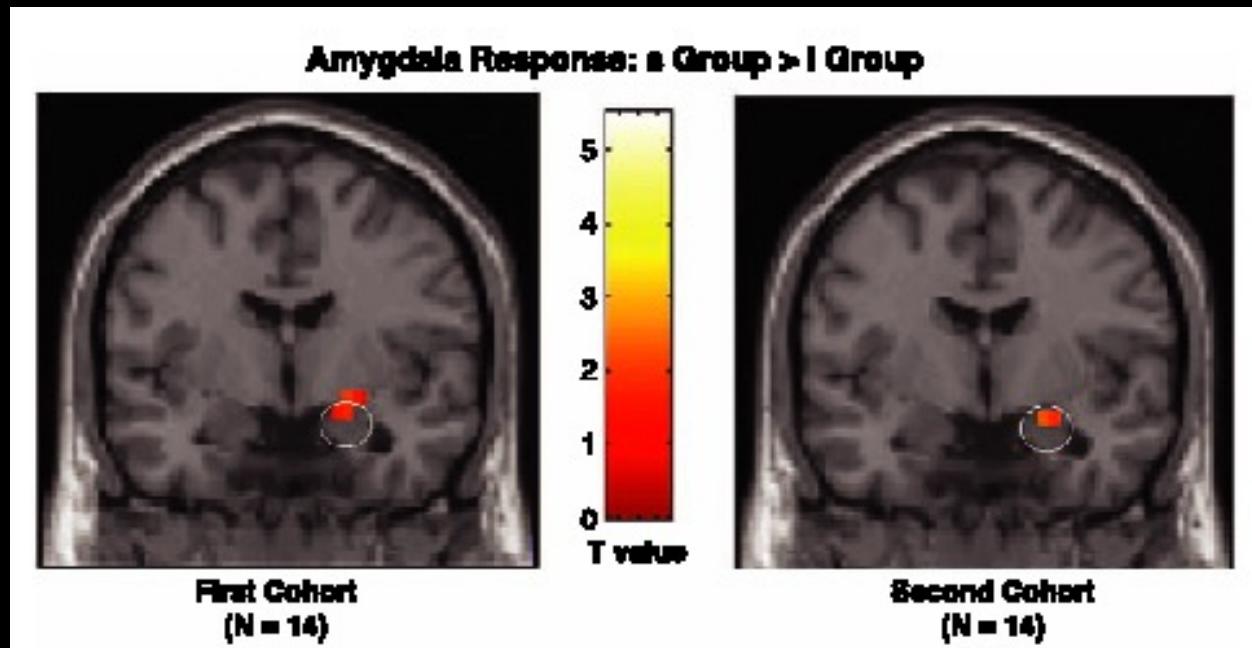
# Perceptual decision making



Comparison of two groups of *normal* individuals with differences in the Serotonin Transporter Gene

# Serotonin Transporter Genetic Variation and the Response of the Human Amygdala

Ahmad R. Hariri,<sup>1</sup> Venkata S. Mattay,<sup>1</sup> Alessandro Tessitore,<sup>1</sup>  
Bhaskar Kolachana,<sup>1</sup> Francesco Fera,<sup>1</sup> David Goldman,<sup>2</sup>  
Michael F. Egan,<sup>1</sup> Daniel R. Weinberger<sup>1\*</sup>

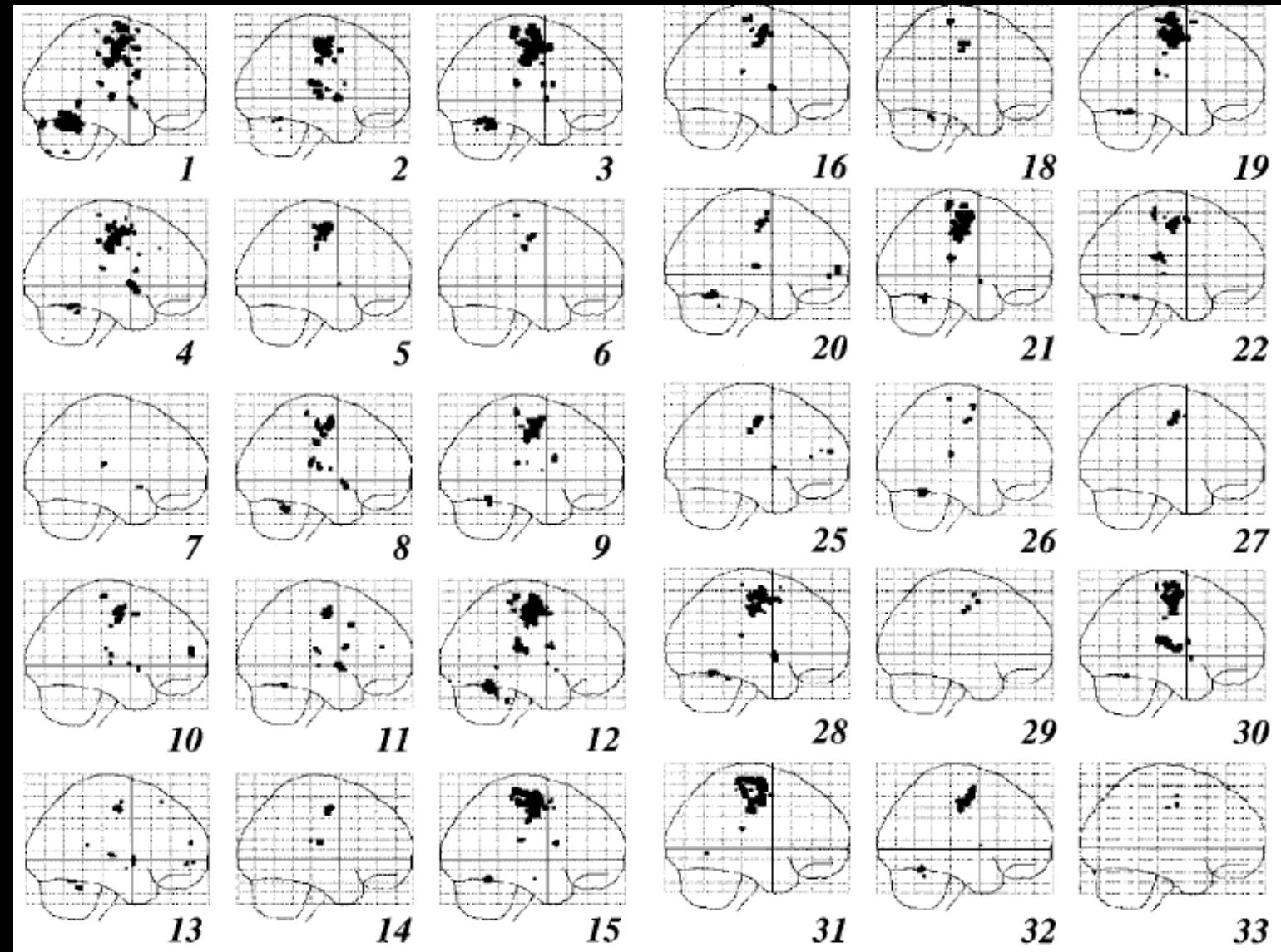


# The Biggest Unknowns in Functional MRI

1. Relationship between neuronal activity and BOLD contrast?
2. Source of BOLD dynamic characteristics?
3. Sources of variability?
4. What's really in the noise?
5. What's "resting" state?
6. Other sources of functional contrast?
7. Ultimate temporal resolution?
8. Ultimate spatial resolution?
9. Ultimate clinical utility?
10. Best display methods?
11. Best processing methods?
12. Optimal Field Strength?

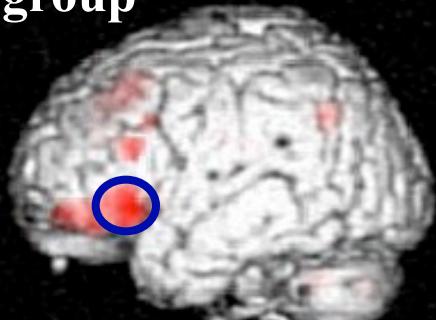
# Sources of Variability in fMRI

- Signal to noise (about 2/1)
- Scanner variability
- Motion
- Hemodynamic response variability
- Differences in structural anatomy
- Differences in functional anatomy
- Differences in strategy for task
- Learning, performance,  
habituation, attention, cognitive  
state



McGonigle, et al (2000), NeuroImage 11, 708-734

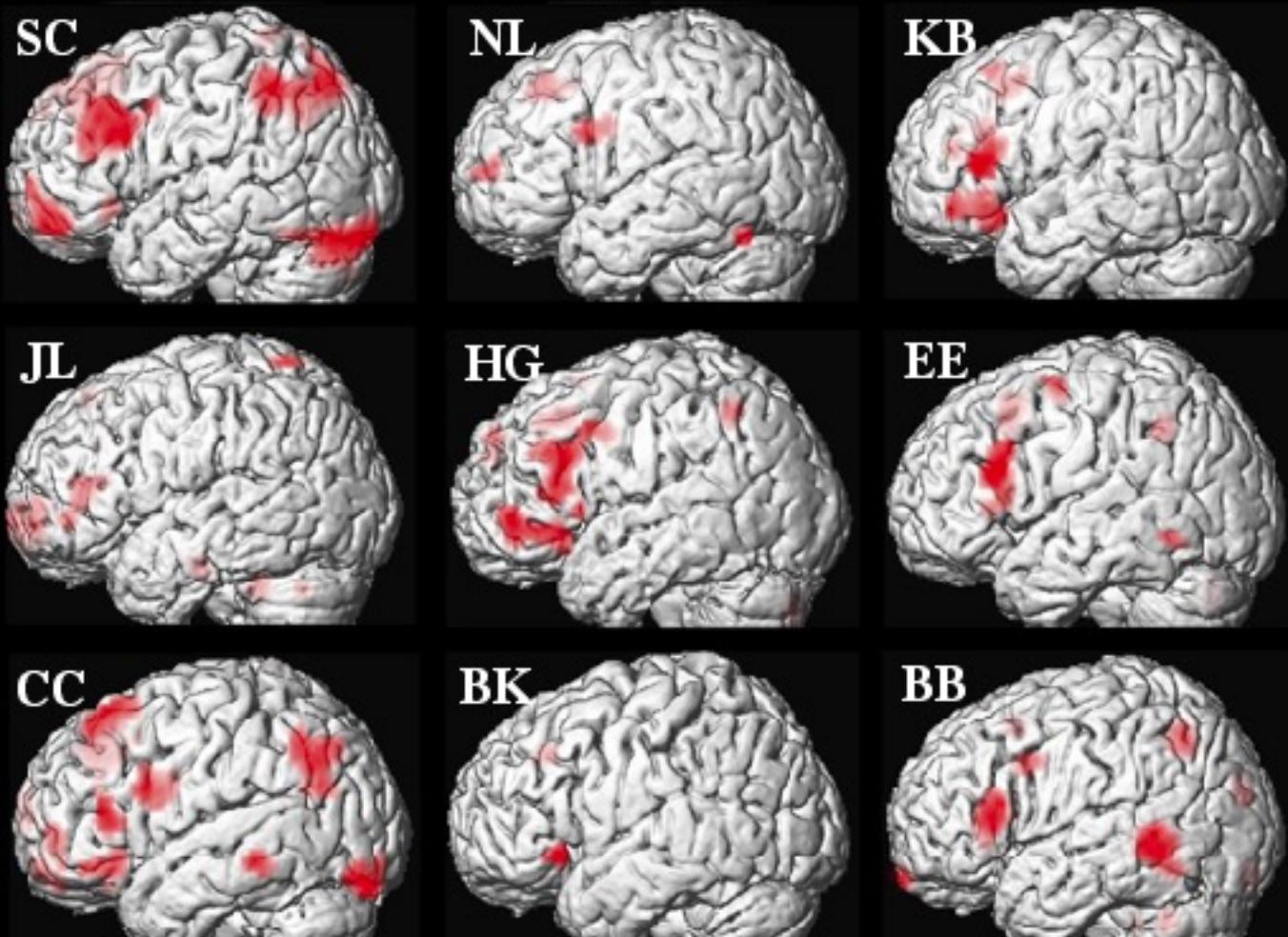
group



# Extensive Individual Differences in Brain Activations During Episodic Retrieval

Miller et al., 2002

Individual activations from the left hemisphere of the 9 subjects



Courtesy, Mike Miller, UC  
Santa Barbara and Jack Van  
Horn, fMRI Data Center,  
Dartmouth University

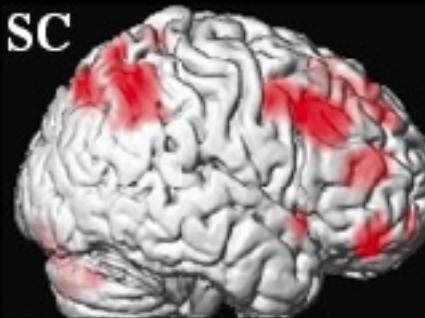
group

# Extensive Individual Differences in Brain Activations During Episodic Retrieval

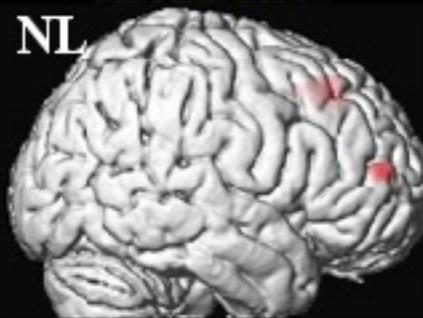
Miller et al., 2002

Individual activations from the right hemisphere of the 9 subjects

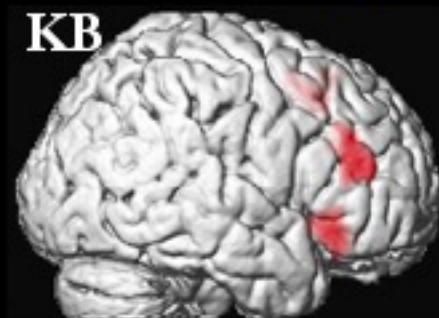
SC



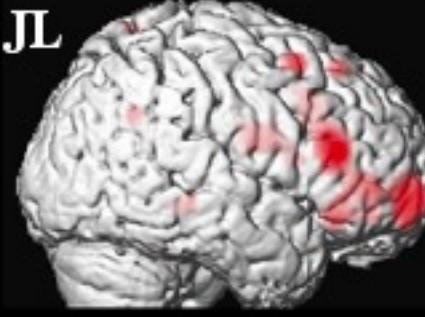
NL



KB



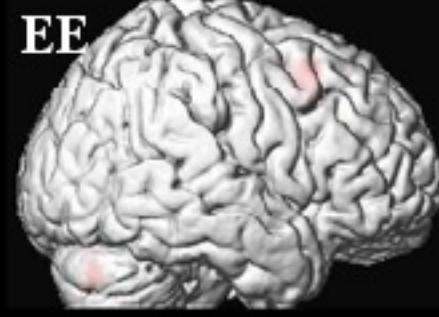
JL



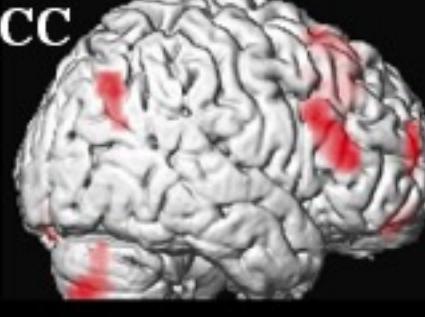
HG



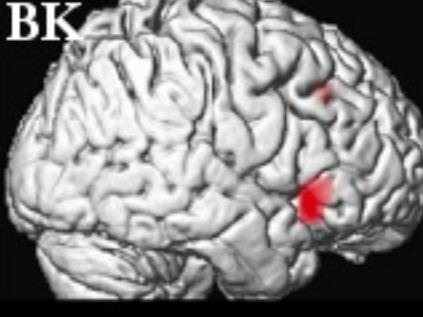
EE



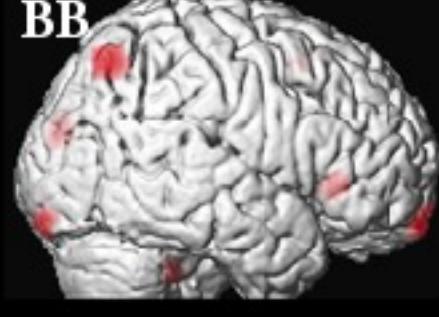
CC



BK

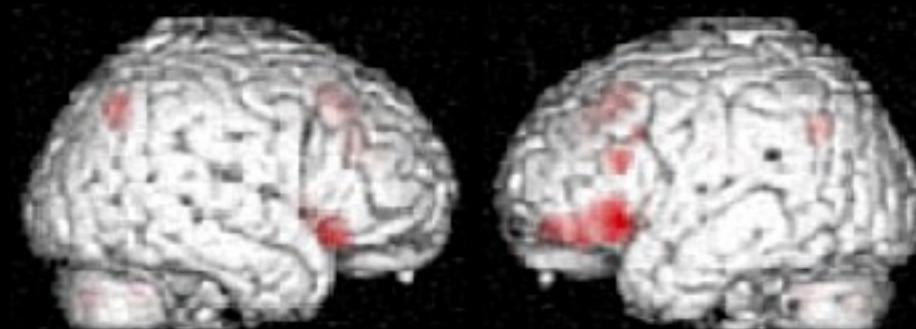


BB



Courtesy, Mike Miller, UC  
Santa Barbara and Jack Van  
Horn, fMRI Data Center,  
Dartmouth University

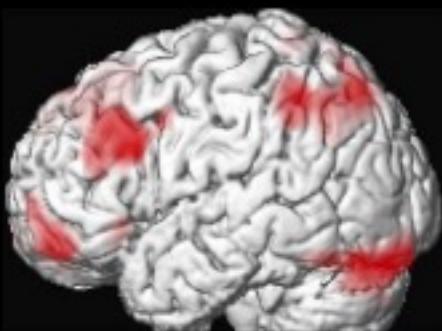
**These individual patterns of activations are stable over time**



**Group Analysis of Episodic Retrieval**



**Subject SC**



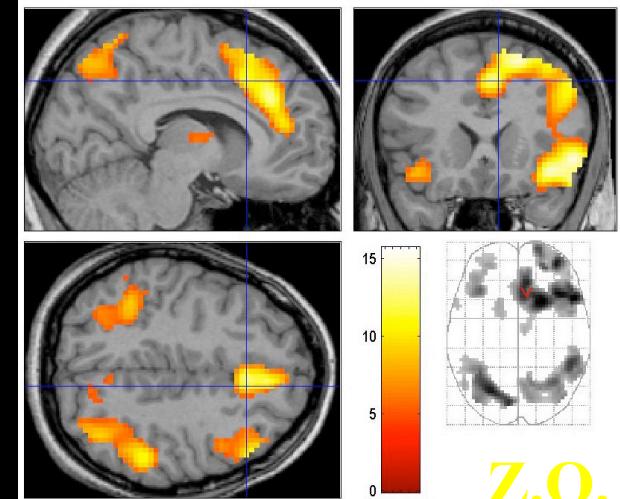
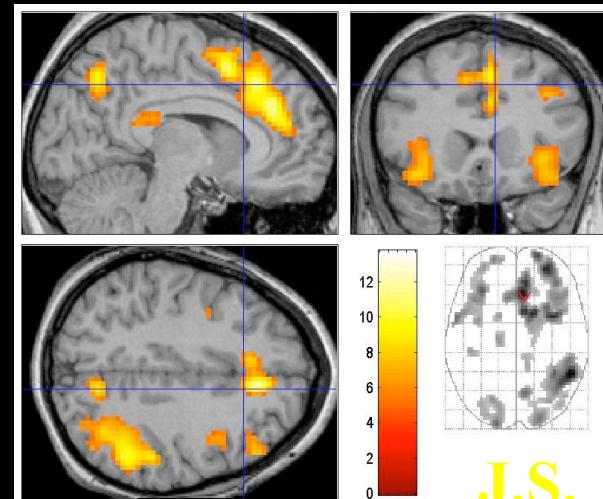
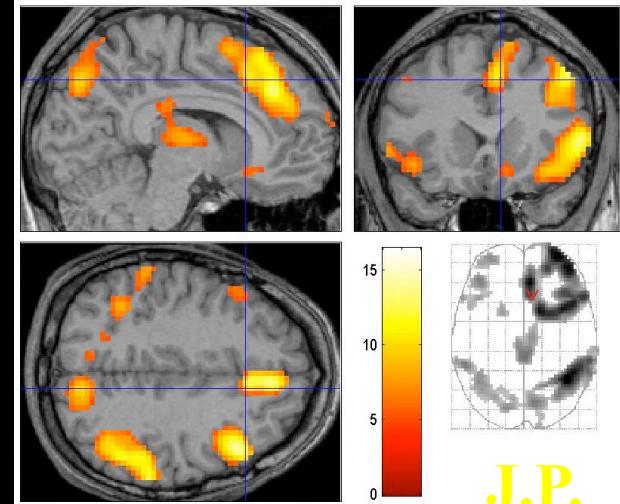
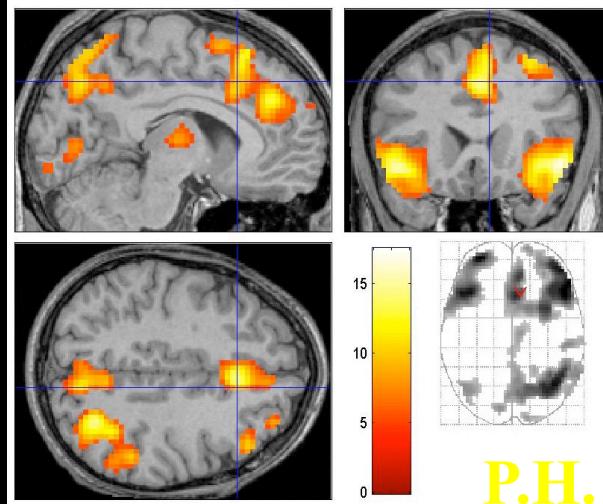
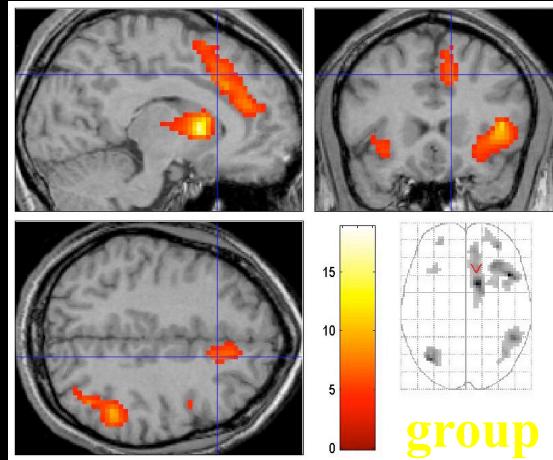
**Subject SC 6 months later**



Courtesy, Mike Miler, UC  
Santa Barbara and Jack Van  
Horn, fMRI Data Center,  
Dartmouth University

**Individual patterns of activity are much more consistent across subjects for other retrieval tasks.**

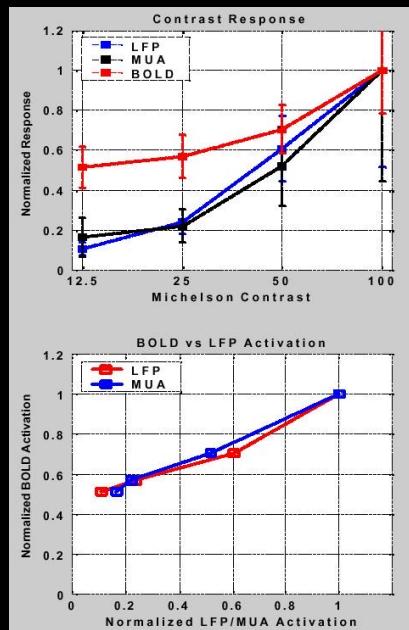
## spatial working memory



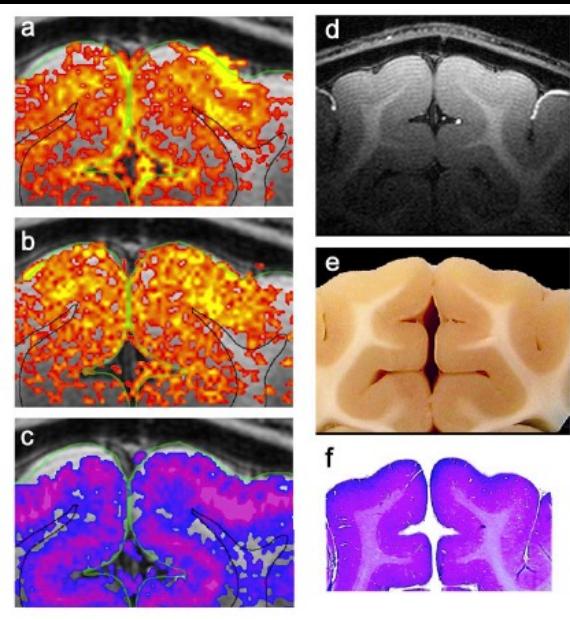
Courtesy, Mike Miler, UC  
Santa Barbara and Jack Van  
Horn, fMRI Data Center,  
Dartmouth University

# Relationship between neuronal activity and BOLD contrast?

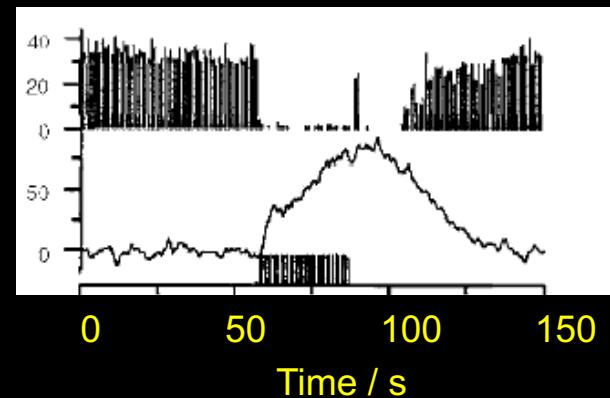
## Magnitude



## Location



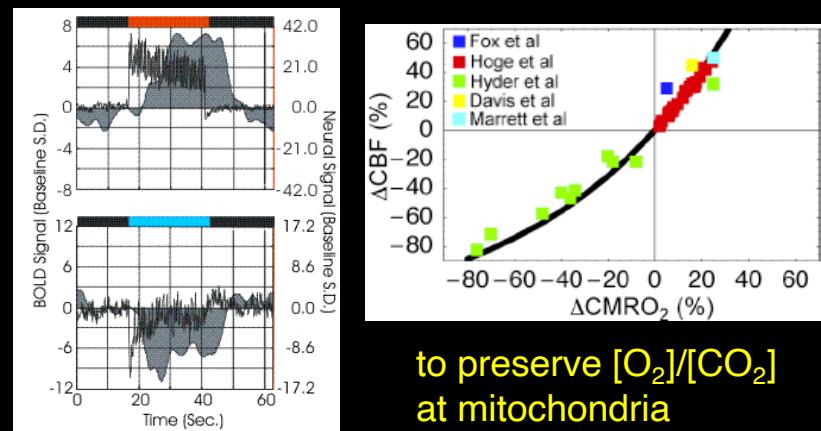
## Inhibition



Mathiesen, et al (1998), J Physiol 512.2:555-566

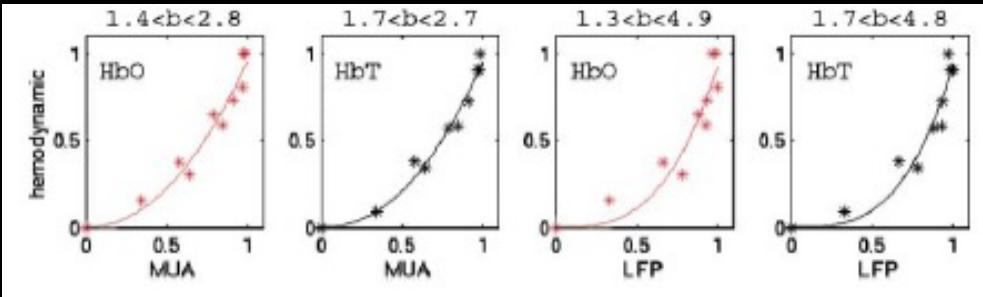
## Neg. BOLD

## Why?



to preserve  $[O_2]/[CO_2]$   
at mitochondria

Logothetis et al. (2001) Nature, 412, 150-157 Harel et al. (2004) ISMRM, 200

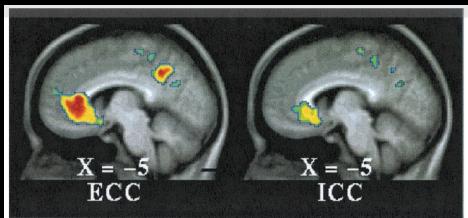


Devor et al. (2001) Neuron, 39, 353-359

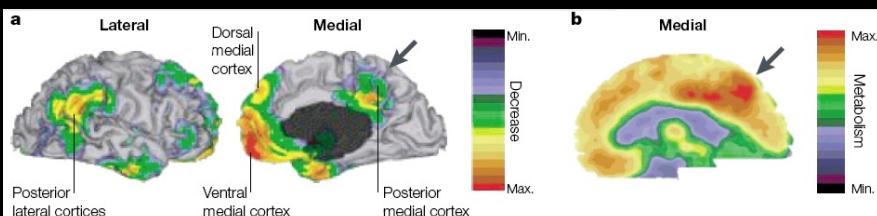
Schmucl et al. (2003) OHBM, 308

Buxton (2004) ISMRM, 273

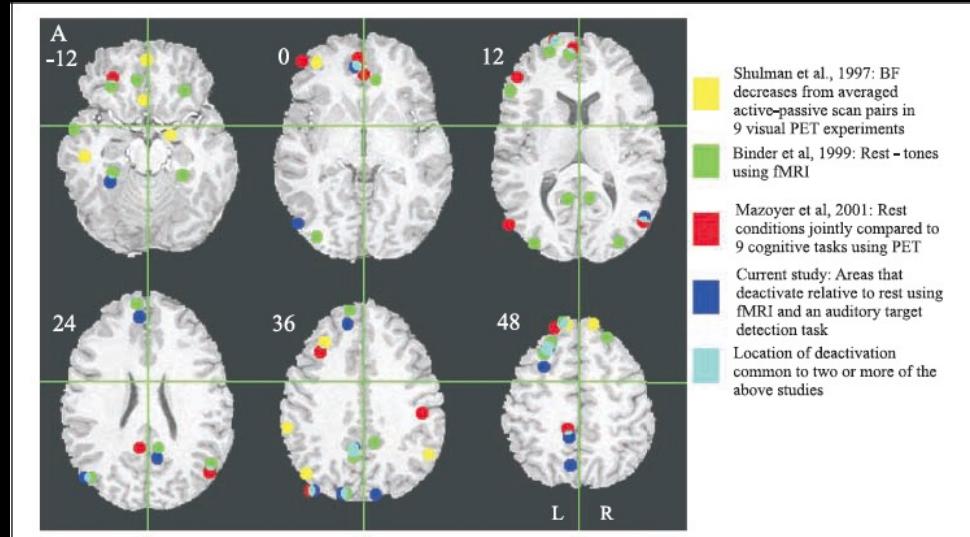
# Negative Signal Changes..



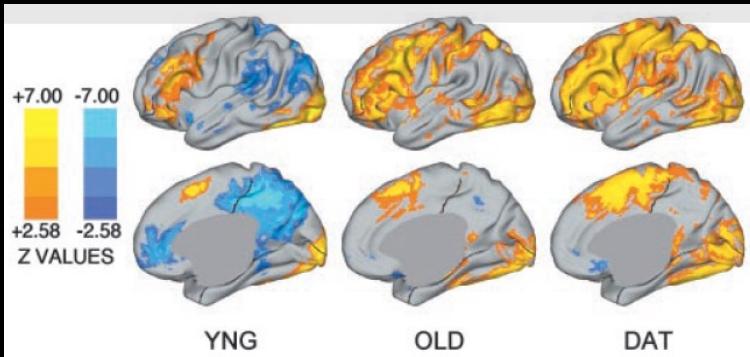
Gusnard, et al (2001), PNAS 98 (7), 4259-4264



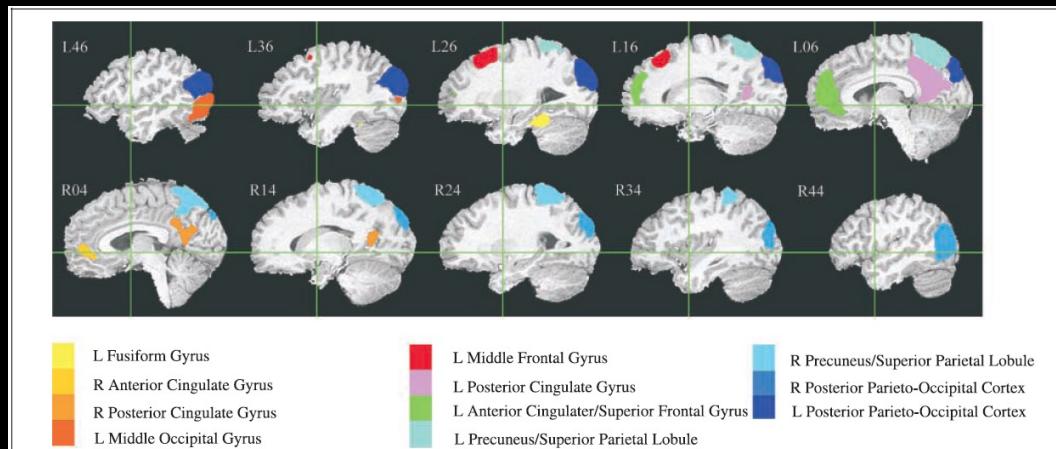
Gusnard, et al (2001), Nature Reviews Neuroscience (2), 685-694



McKiernan, et al (2003), Journ. of Cog. Neurosci. 15 (3), 394-408

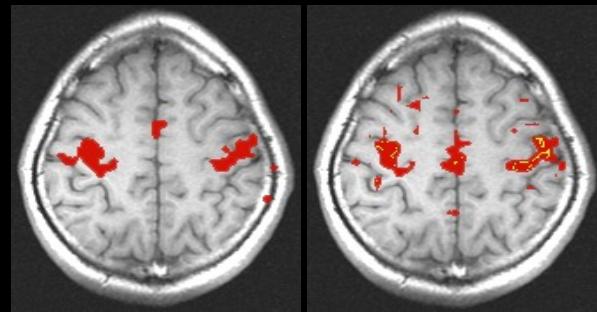
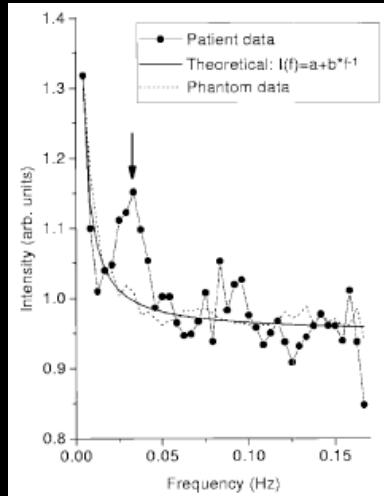


Lustig, et al (2003), PNAS 100 (19), 14504-14509

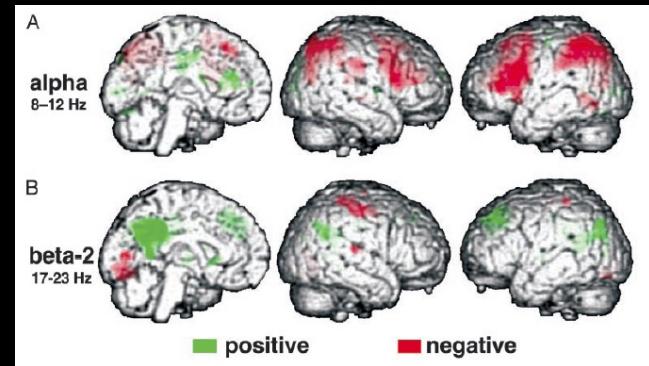


McKiernan, et al (2003), Journ. of Cog. Neurosci. 15 (3), 394-408

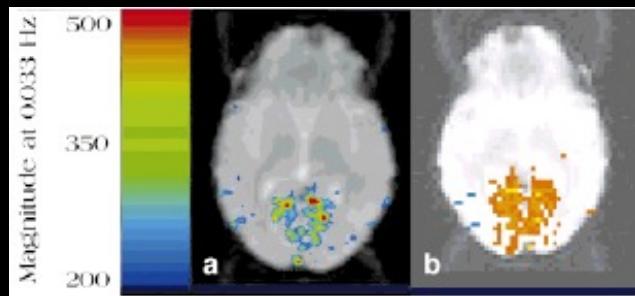
# Baseline fluctuations...



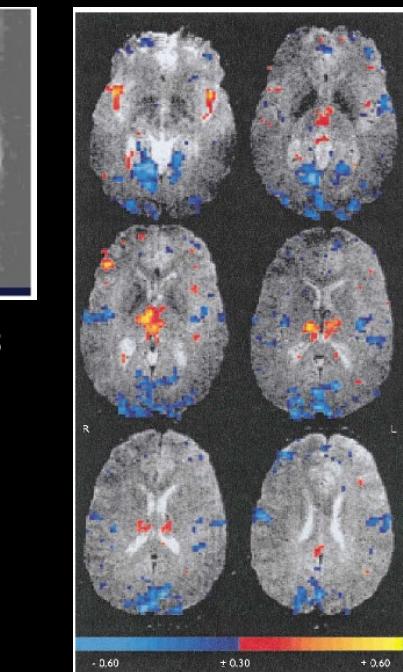
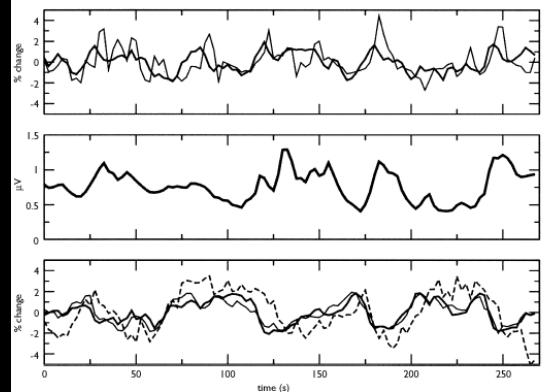
Biswal, et al (1995), MRM 34, 537-541



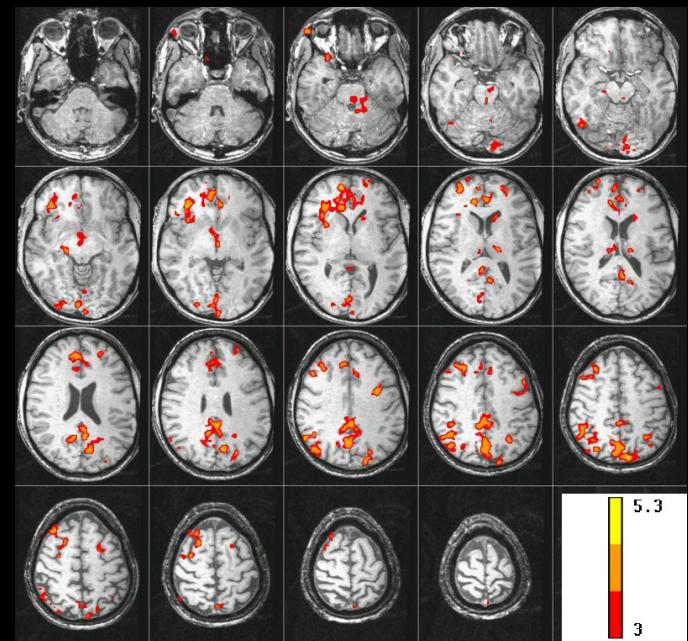
Laufs, et al (1995), PNAS 100 (19), 11053=11058



Kiviniemi, et al (2000), MRM 44, 373-378



Goldman, et al (2002), Neuroreport



Patterson, et al (2002), NeuroImage 17, 1787-1806

## What it can do

- map brain activation in individuals and groups
- spatial resolution: 0.5 mm
- temporal resolution (absolute: 4 sec, relative: 100 ms)
- track changes over time (sec to years)
- predict some behavior

## What it can't do

- differentiate excitation vs. inhibition
- demonstrate causality
- classify individuals
- predict long term behavior

## What it might do

- predict wider range of behavior
- characterize baseline activity
- classify individuals within a narrow context



# Functional Imaging Methods Unit &



## Functional MRI Facility

### Computer Specialist:

Adam Thomas

### Scanning Technologists:

Karen Bove-Bettis

Paula Rowser

Alda Ottley

Ellen Condon

### Staff Scientists:

Sean Marrett

Jerzy Bodurka

Frank Ye

Wen-Ming Luh

Rasmus Birn

### Program Assistant:

Kay Kuhns

### Post Docs:

Hauke Heekeren

David Knight

Anthony Boemio

Niko Kriegeskorte

### Graduate Student:

Natalia Petridou