

# NEUROIMAGING

(...at the NIH)

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Functional MRI Facility  
Unit on Functional Imaging Methods  
Laboratory of Brain and Cognition

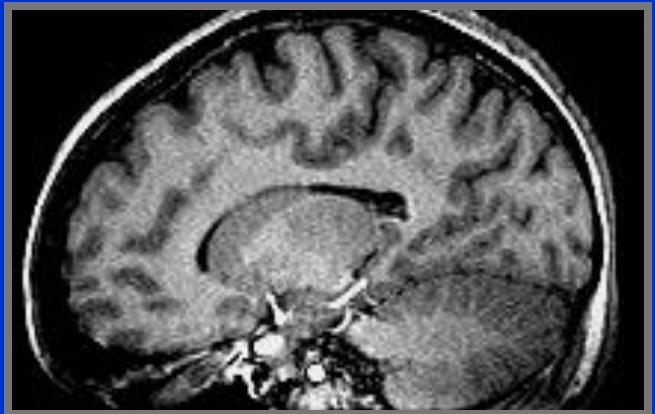


# Two Types of Neuroimaging

- Structural/Anatomical Imaging
- Functional Imaging

# Structural Brain Imaging

- Reveals the anatomy of the brain and the physical structure of brain pathology.
- Primarily associated with Computerized Axial Tomography (CAT or CT) and Magnetic Resonance Imaging (MRI).



- Structural/Anatomical Imaging
  - X-ray
  - Computerized Tomography (CT)
  - Magnetic Resonance Imaging (MRI)
    - Angiography
    - Venography
    - Perfusion
    - Diffusion Tensor Imaging

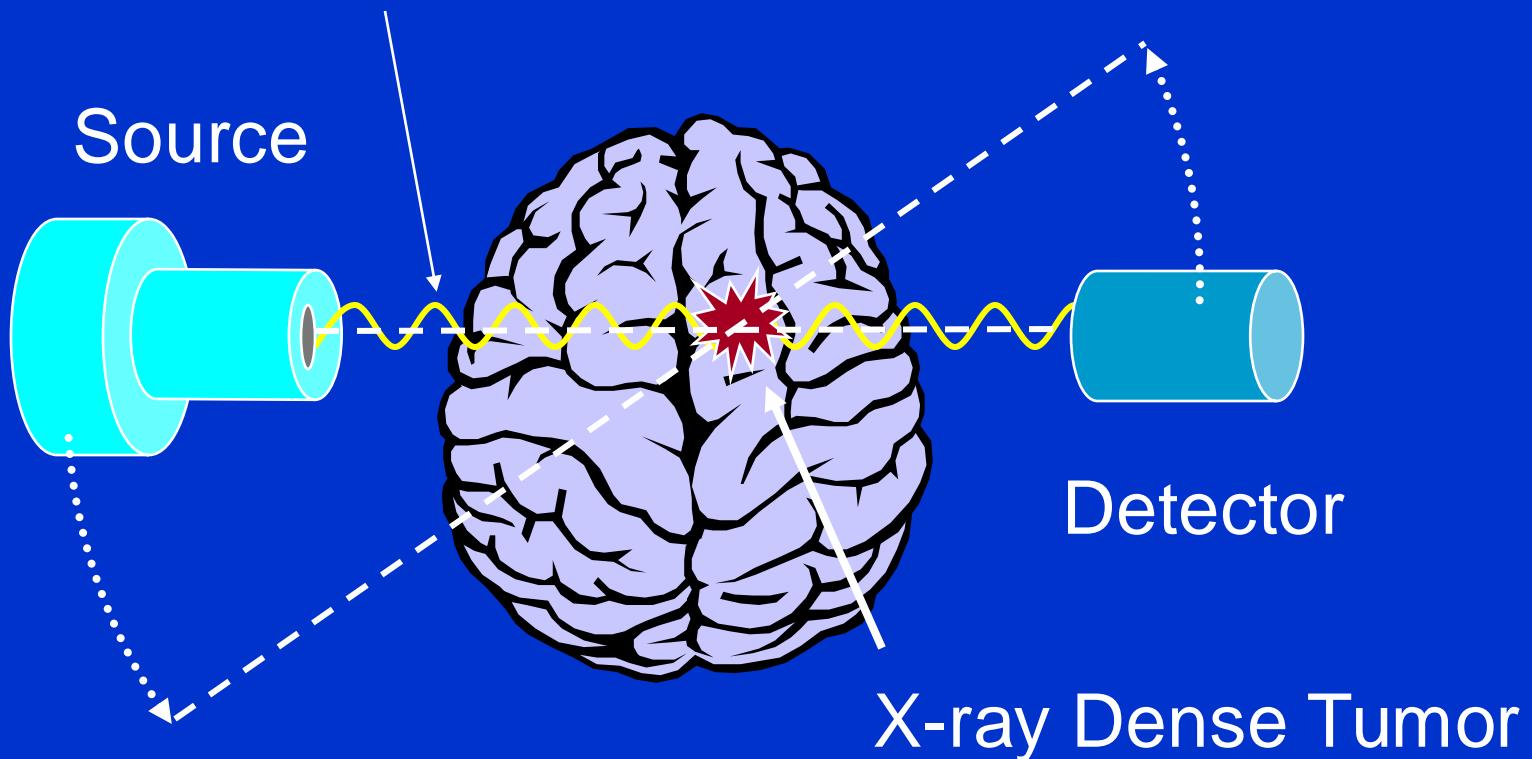
# •Functional Imaging

- 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)

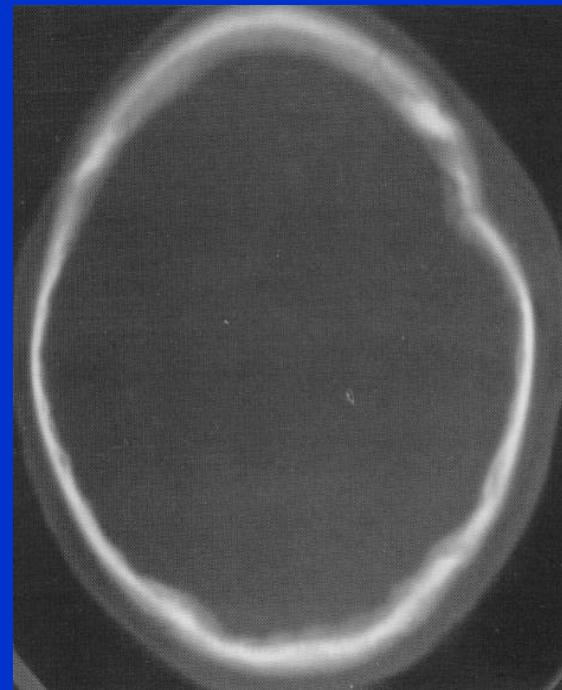
# Computerized Tomography (CT)

Creation of images in slices or sections.

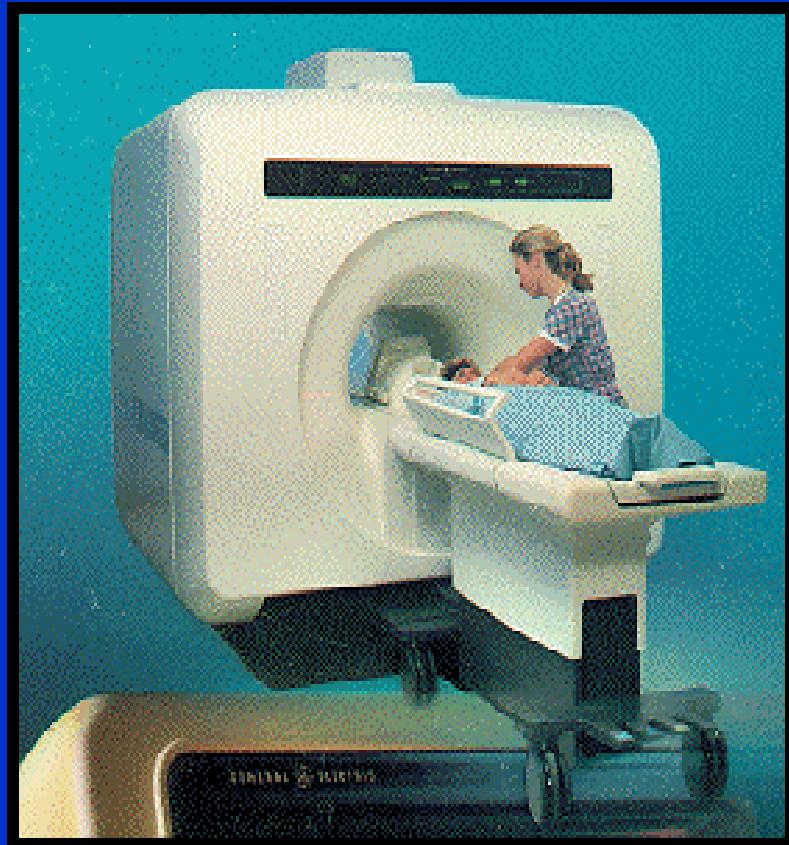
Narrow X-ray beam



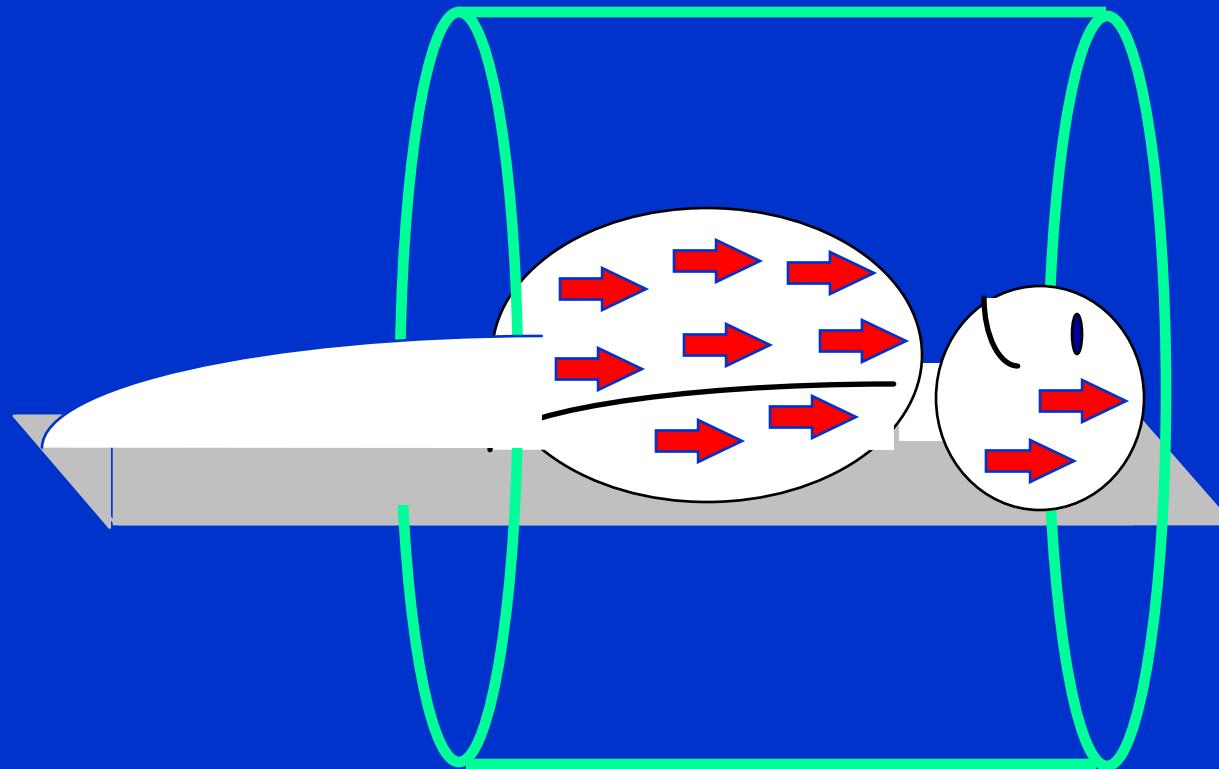
# CT Images



# Magnetic Resonance Imaging

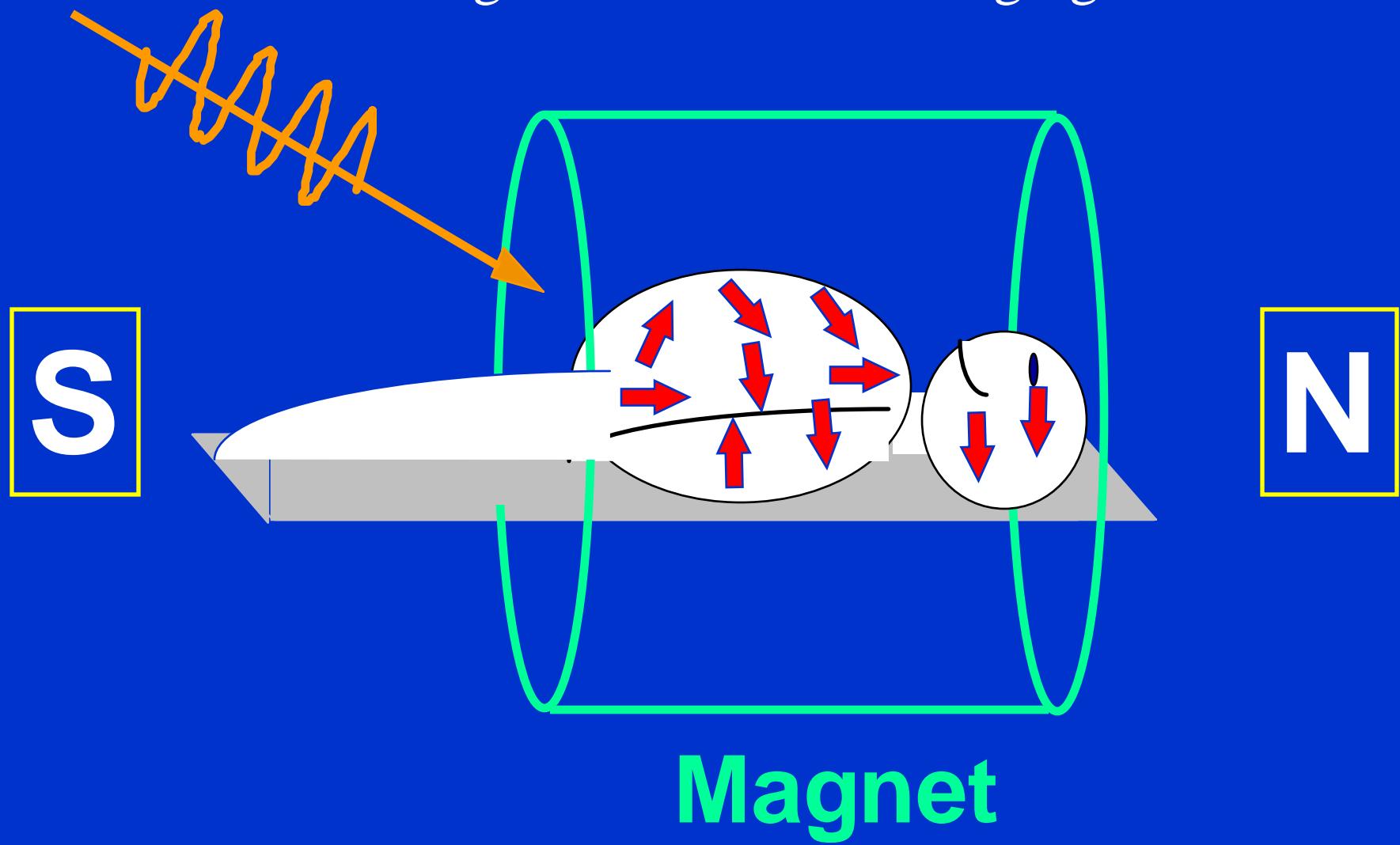


# Magnetic Resonance Imaging

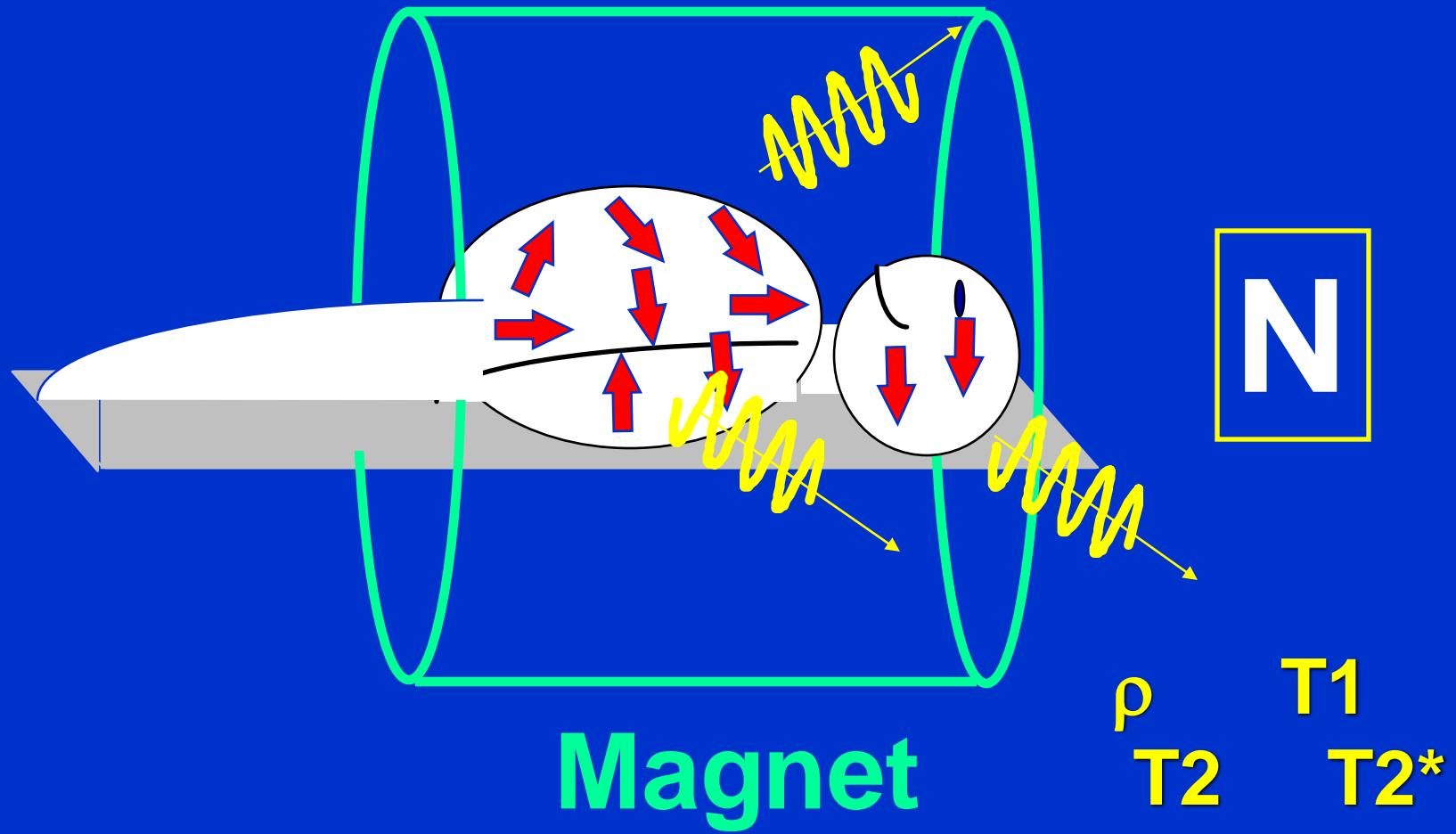


Magnet

# Magnetic Resonance Imaging

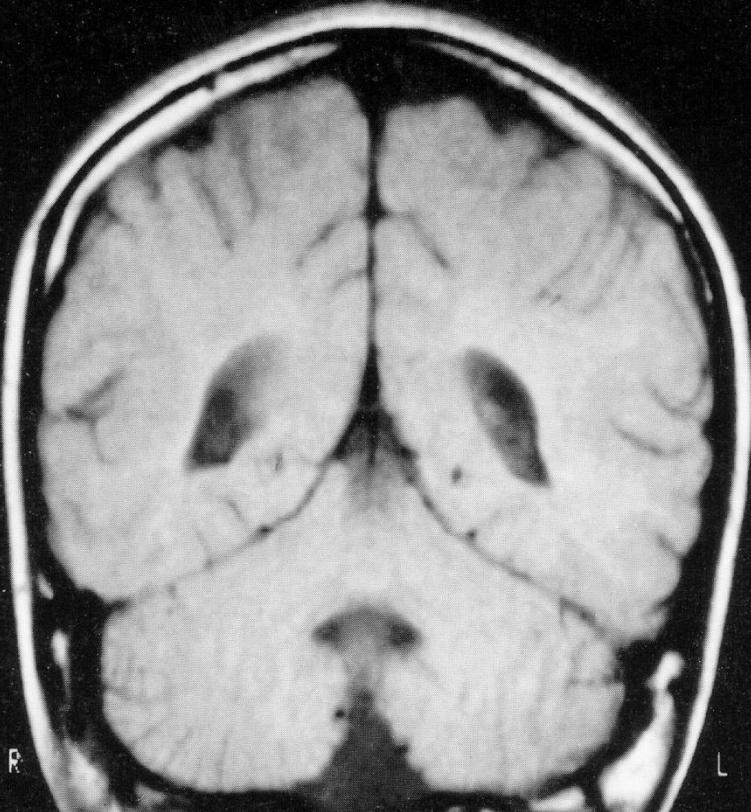


# Magnetic Resonance Imaging

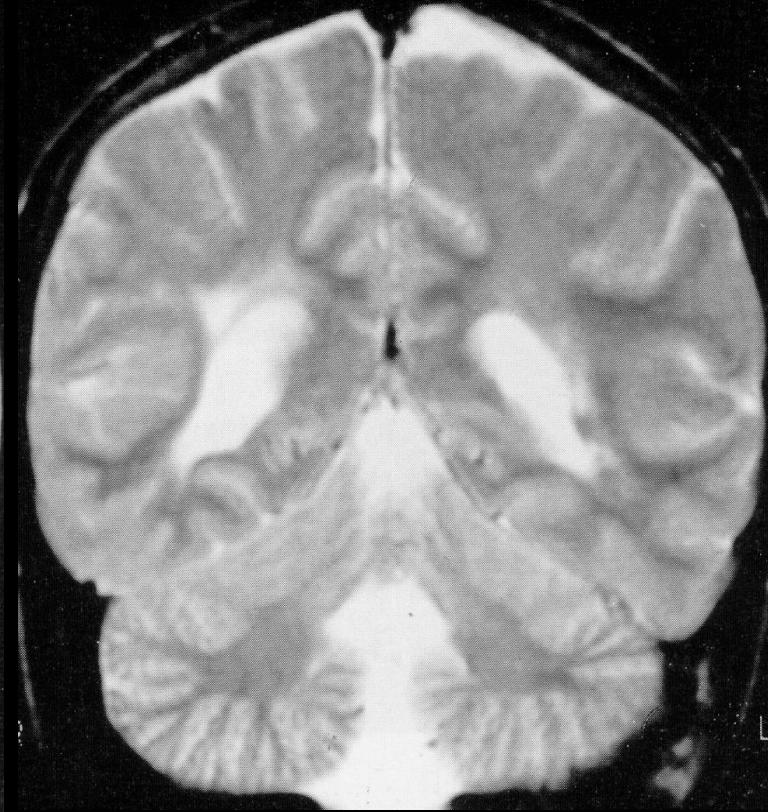


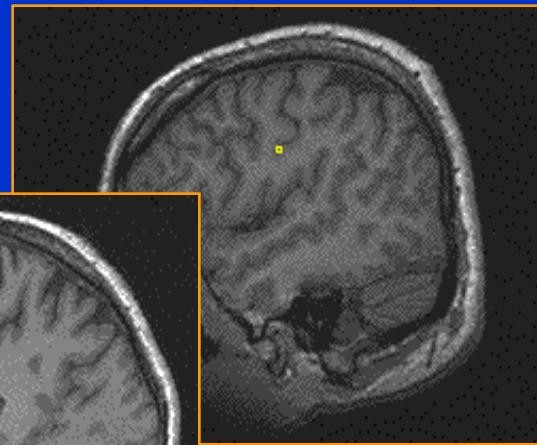
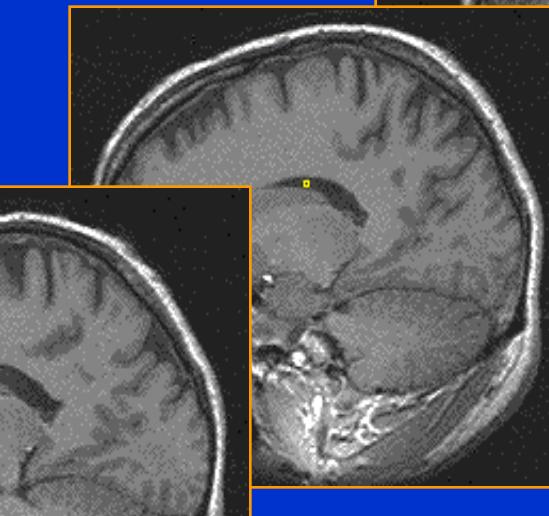
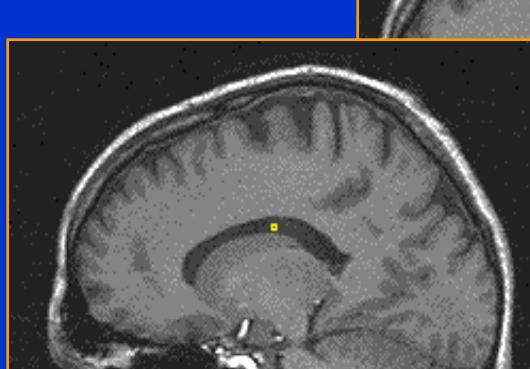
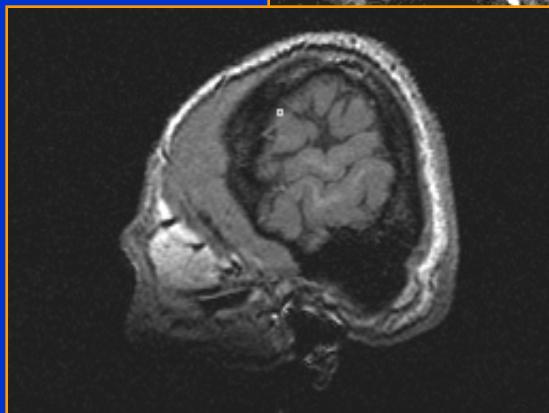
# MRI Images with Different Contrast Weighting

T1 Weighted

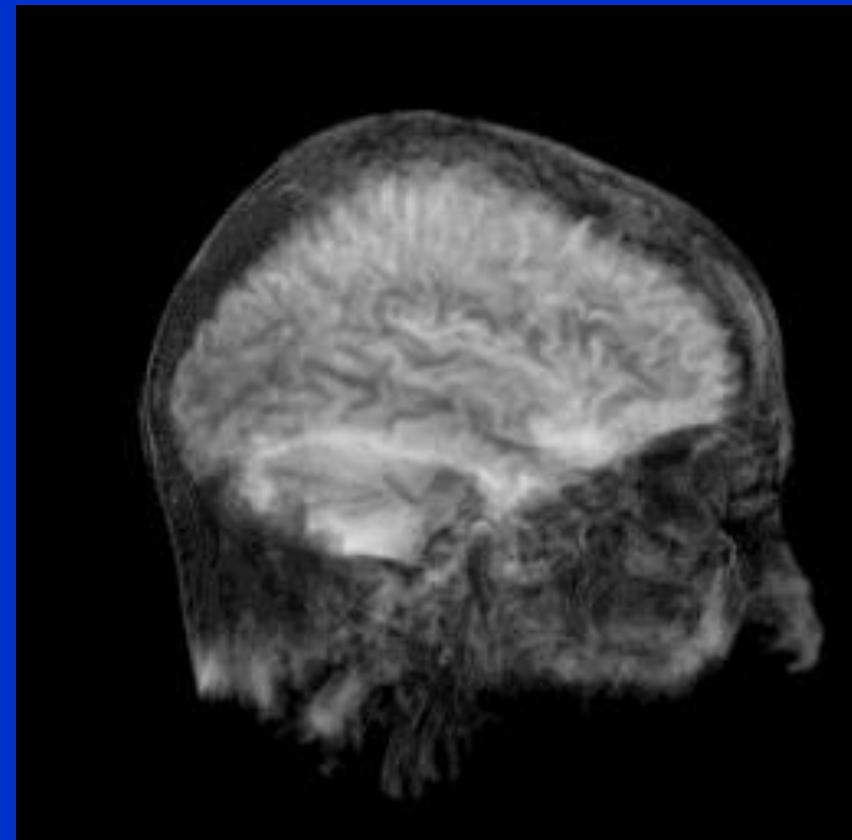
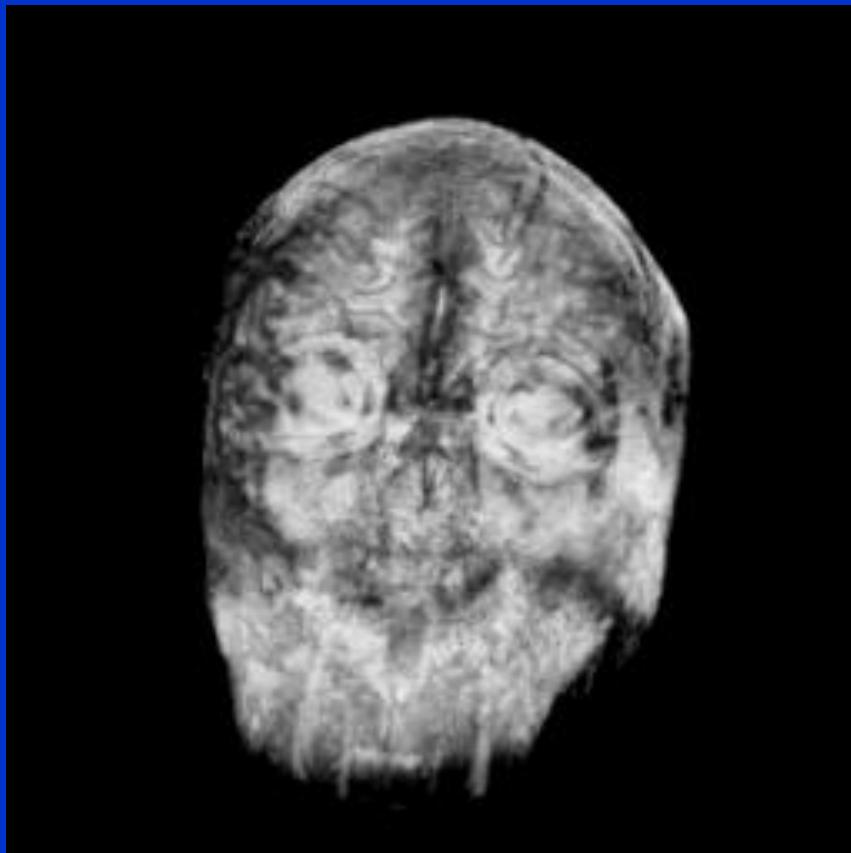


T2 Weighted



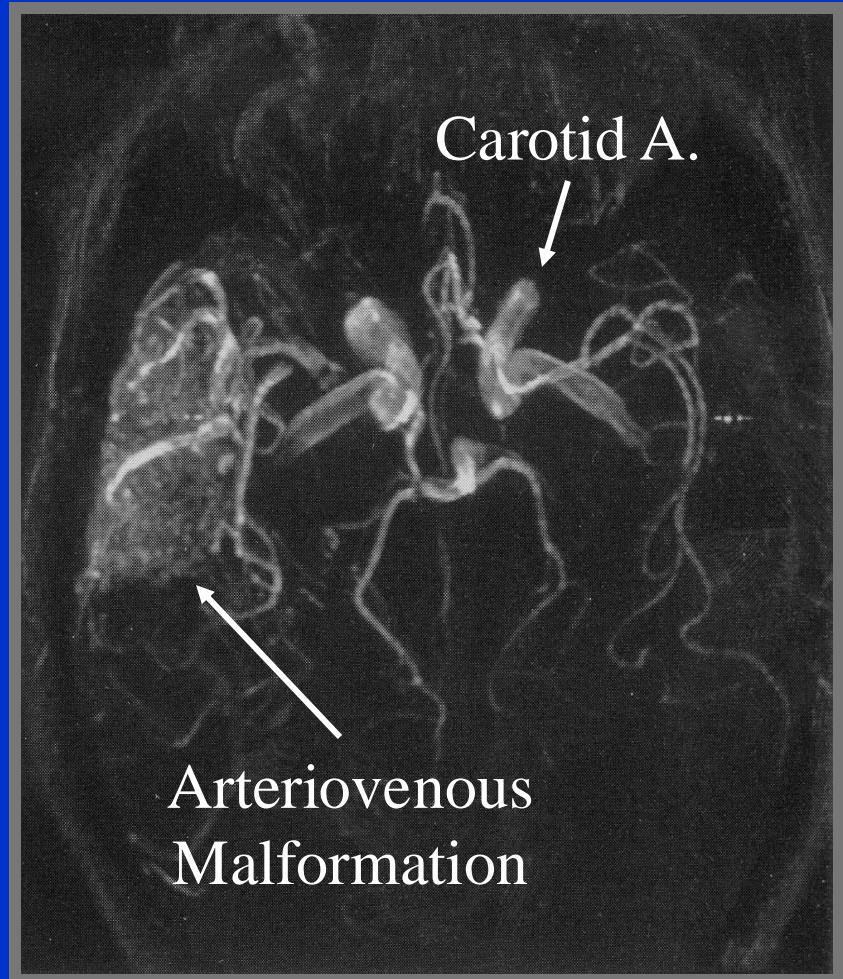
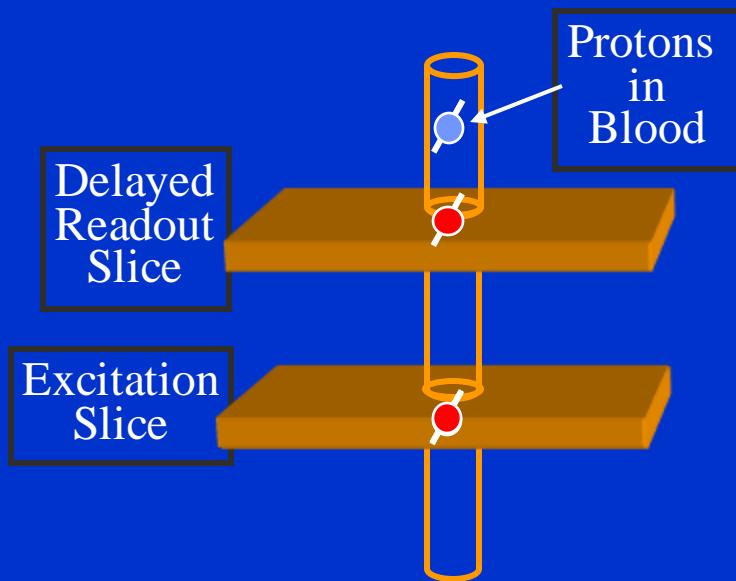


# 3D Rendered MRI

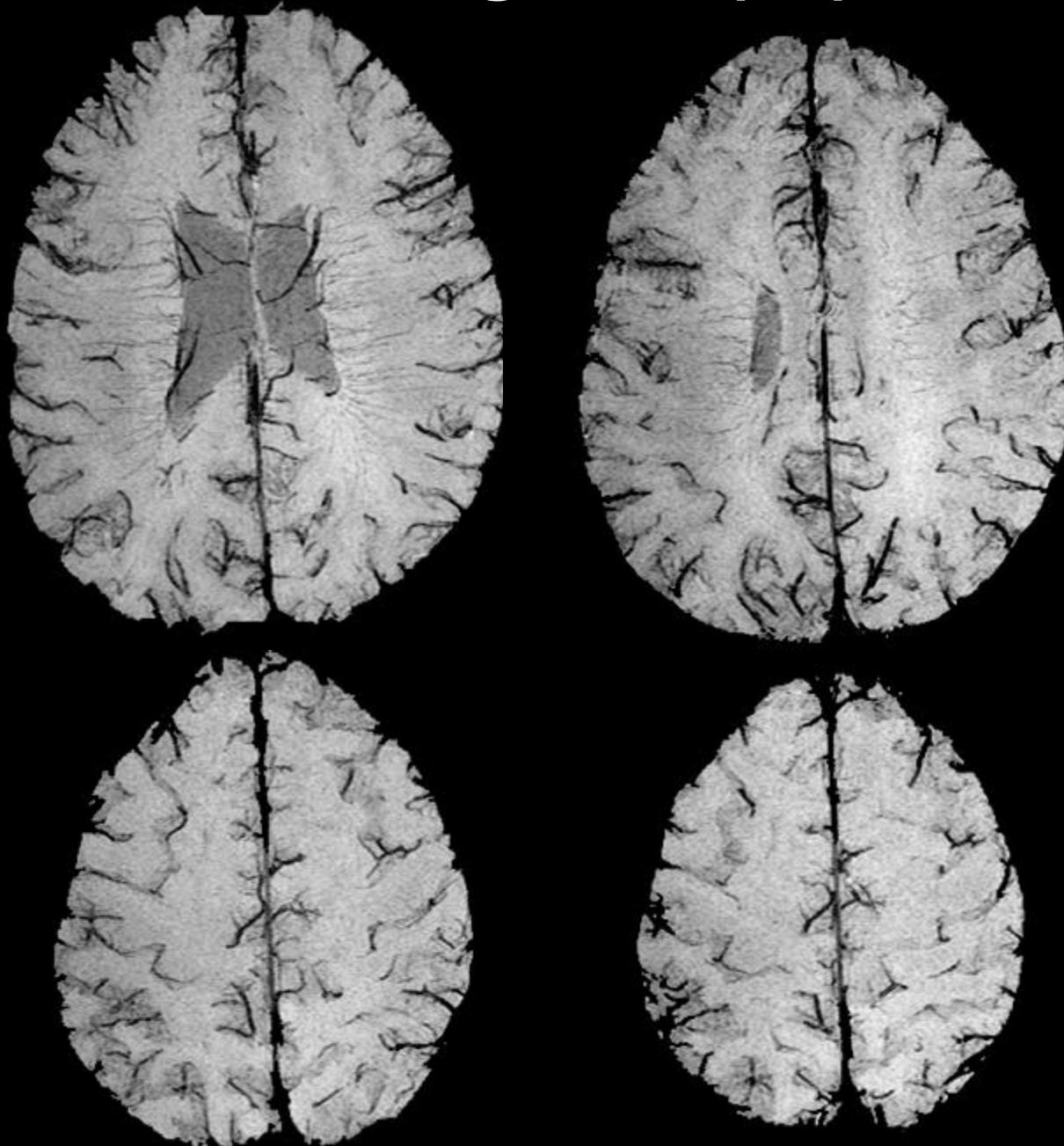


# MR Angiography Shows Blood Vessel Structure

- Blood vessel structure can be visualized by injection of MR tracers or by “spin tagging” techniques.



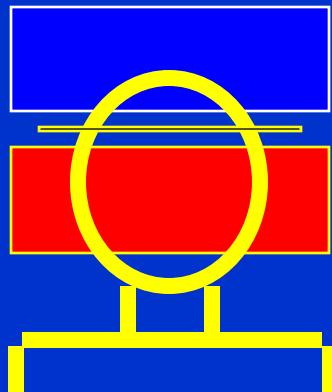
# Venograms (3T)



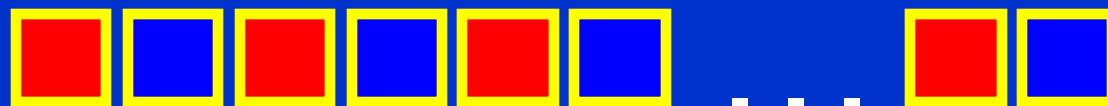
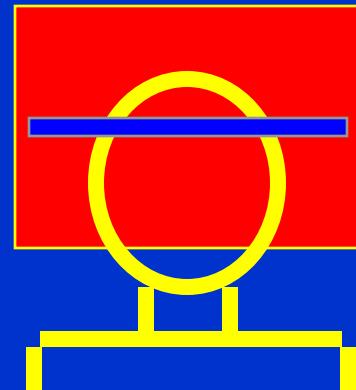


# Perfusion / Flow Imaging

EPISTAR

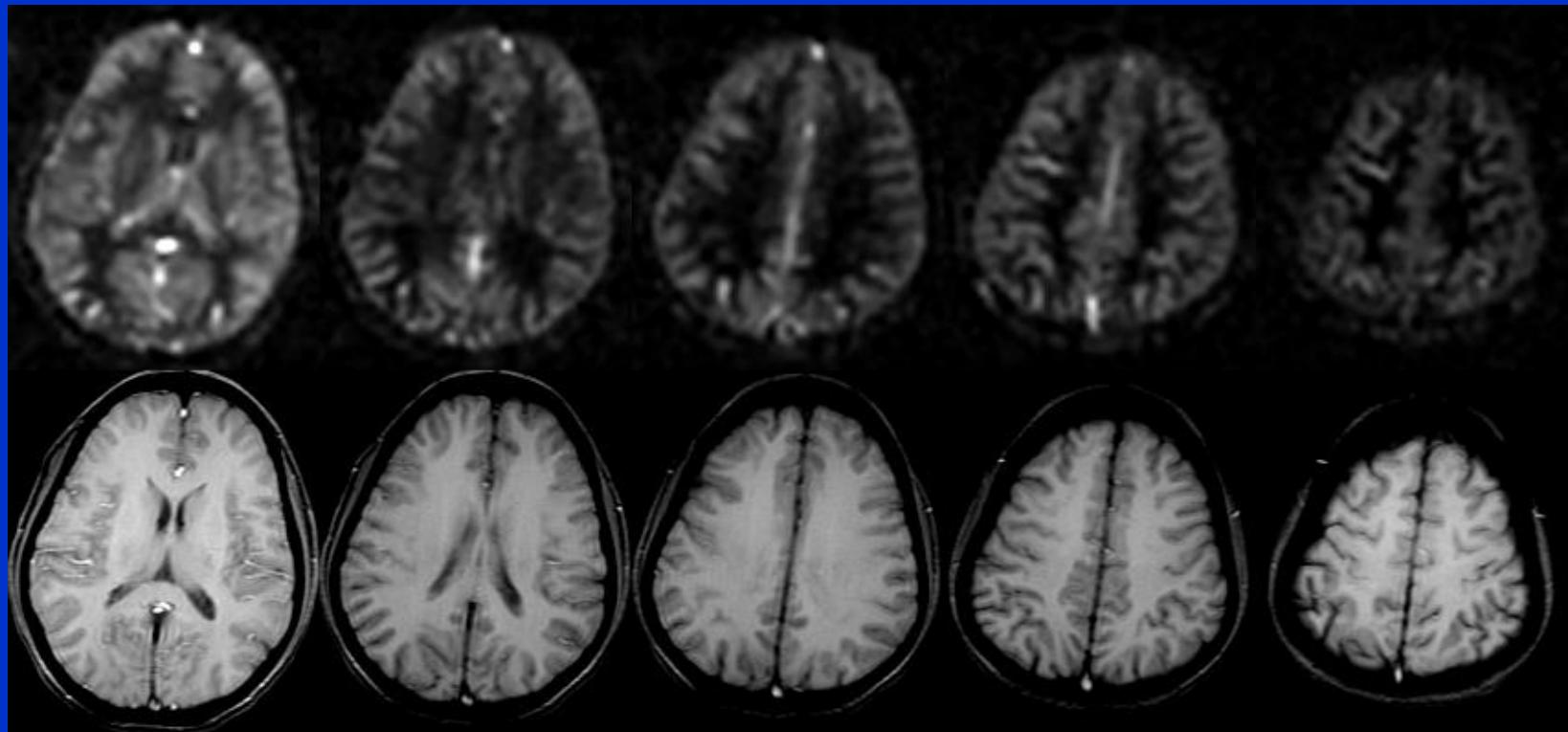


FAIR

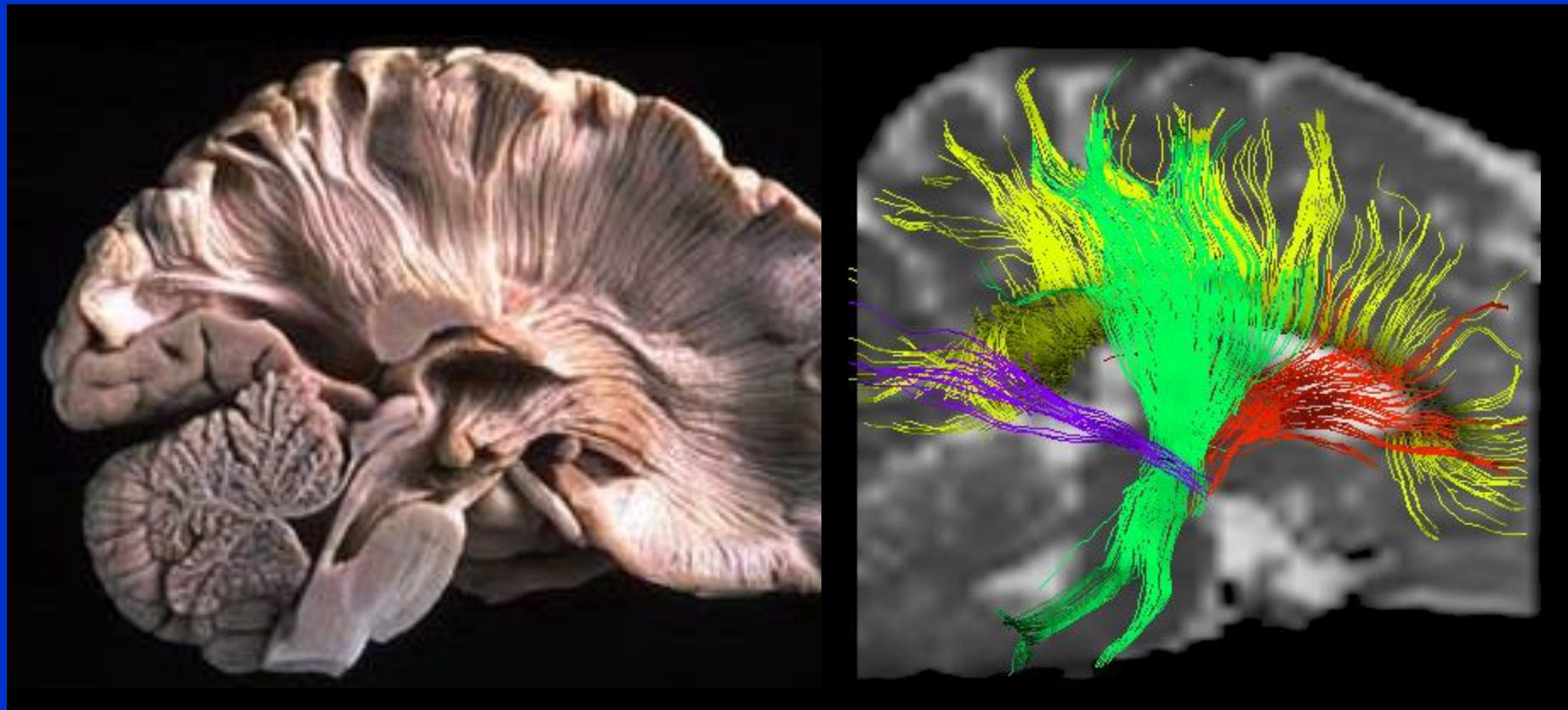


Perfusion  
Time Series

# Perfusion Imaging with MRI

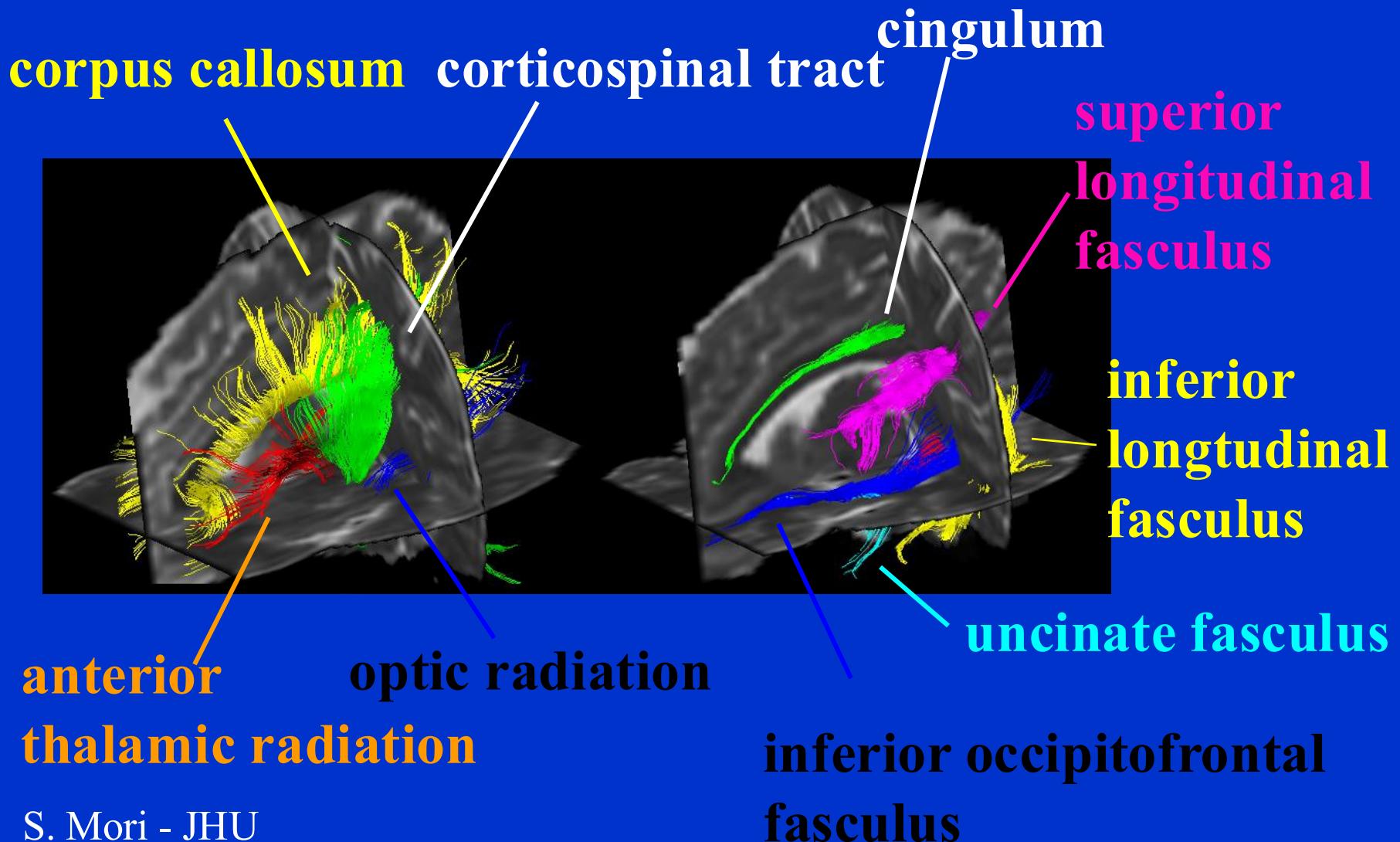


# Diffusion Tensor Imaging



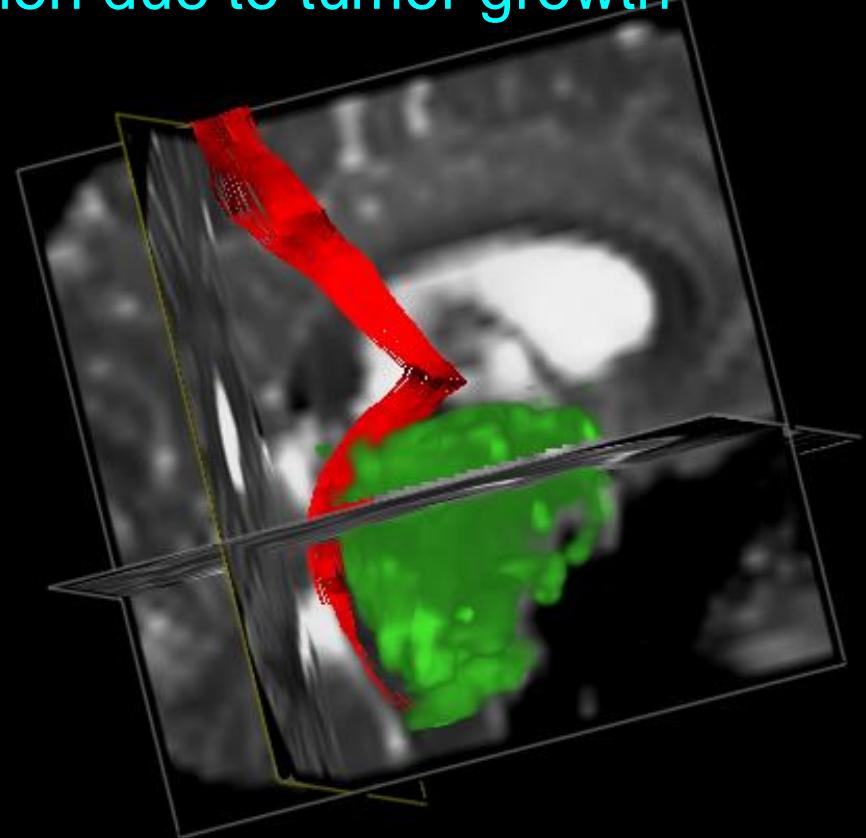
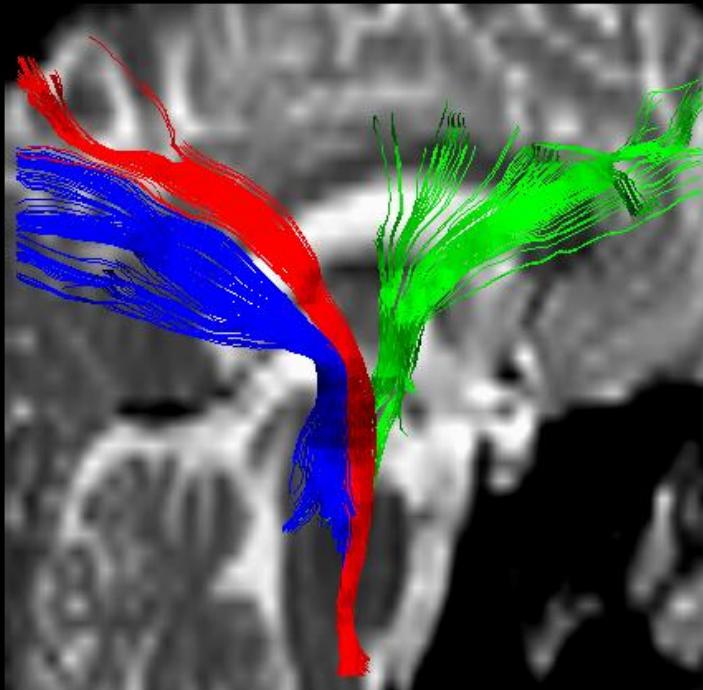
S. Mori - JHU

# Diffusion Tensor Imaging

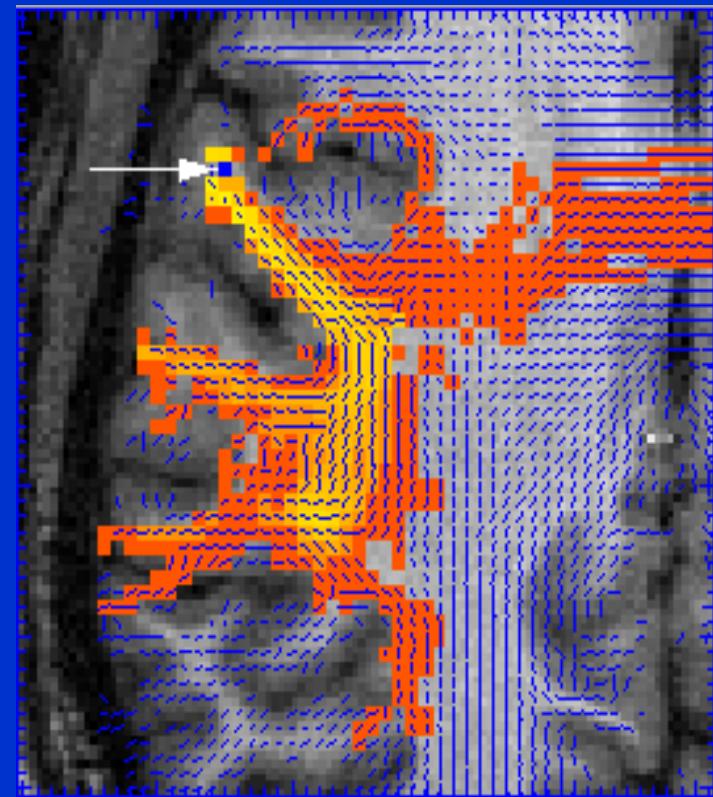
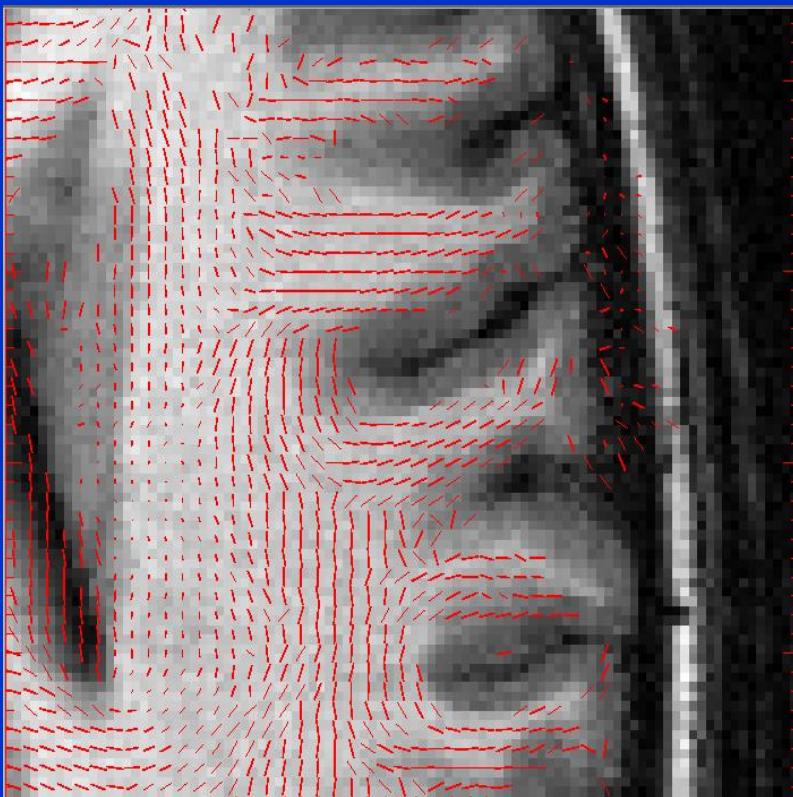


# Anatomical guidance with DTI:

Example: Anatomical deformation due to tumor growth

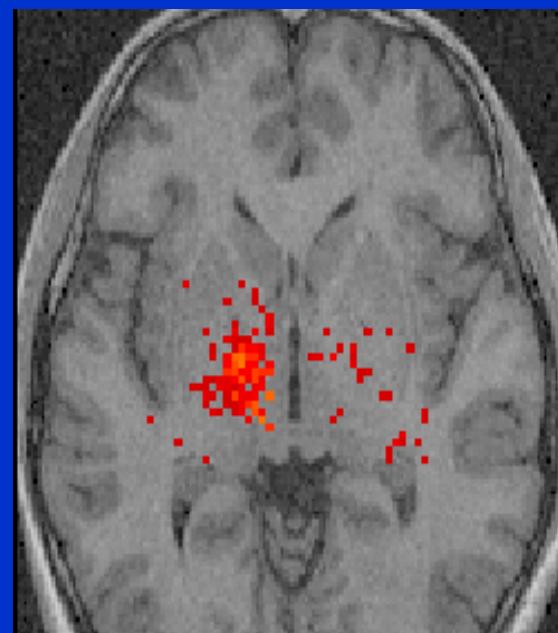
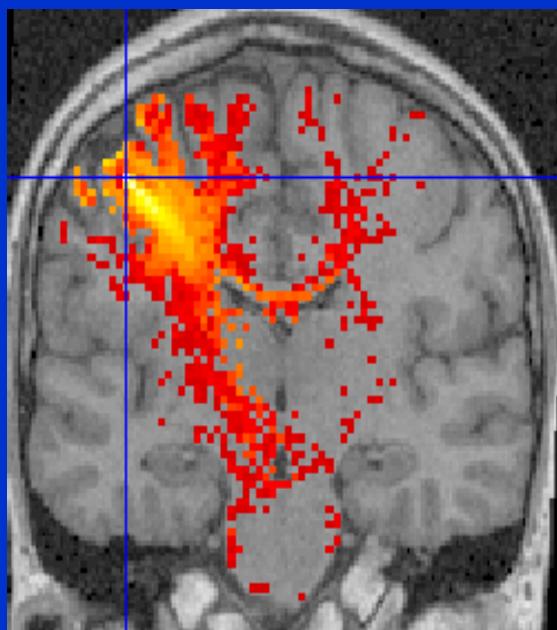


# DTI Connectivity



Koch and Norris, Max Planck, Leipzig

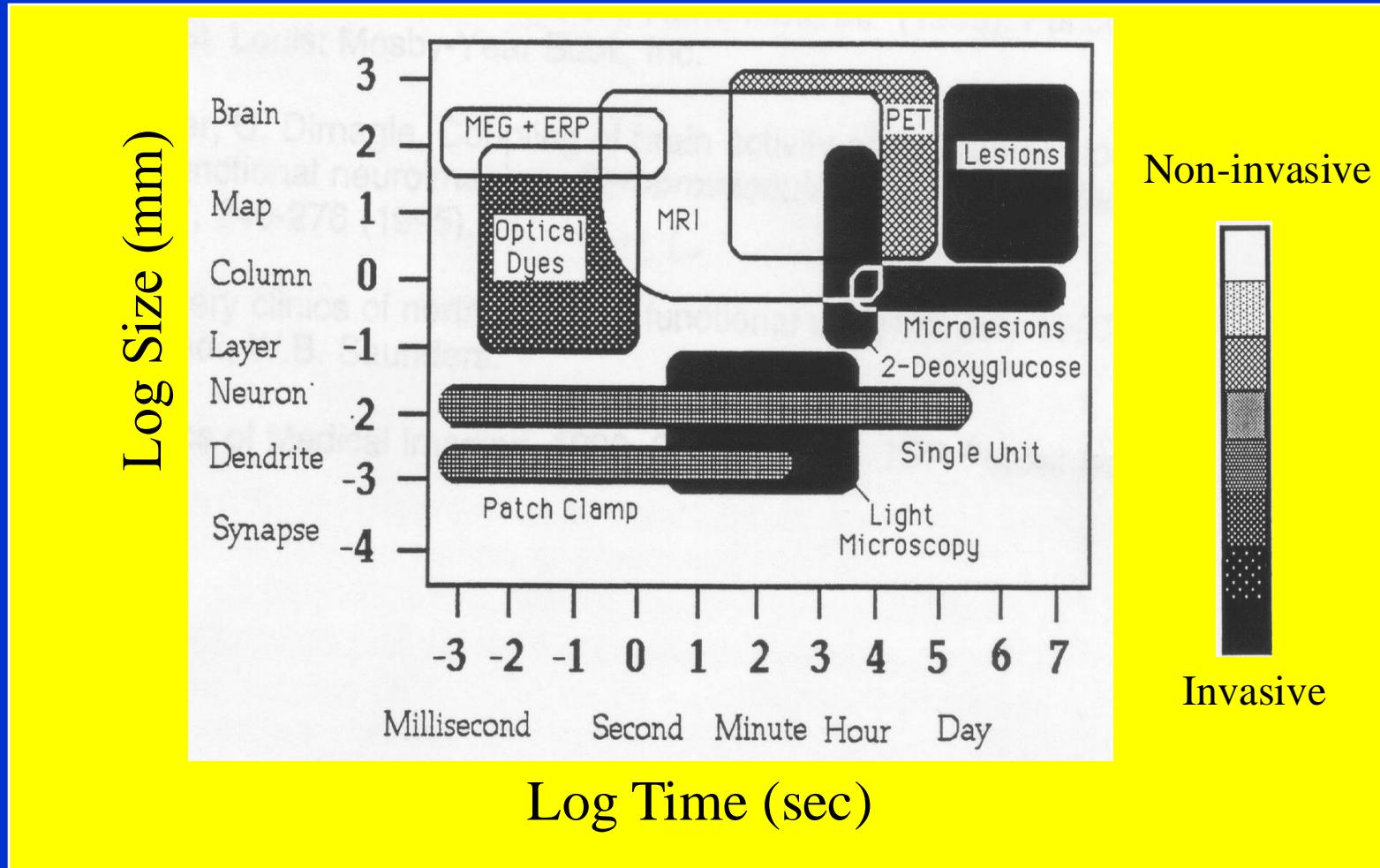
# DTI Connectivity



# Functional Imaging

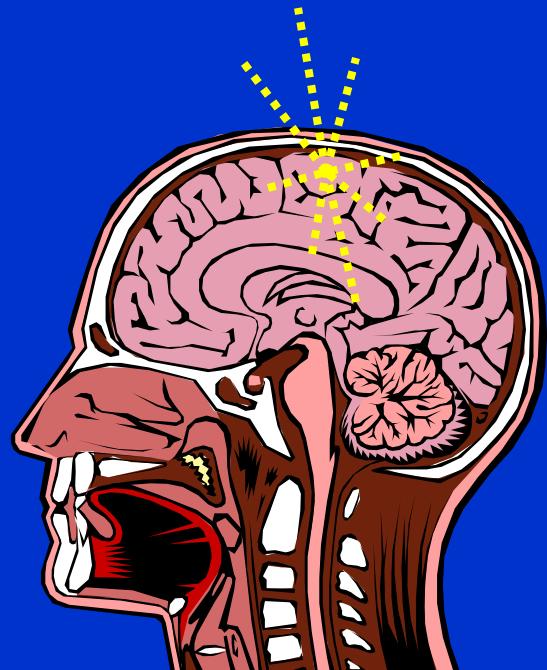
- Unlike structural imaging, functional imaging provides pictures of brain physiology or chemistry.
- By targeting factors that are related to brain activity (eg. blood flow and oxygenation), images of brain activation can be obtained.
- Functional imaging has been used for pre-surgical mapping of function and, eventually, may replace or augment more traditional tests.
- Functional imaging is now a major new research paradigm in neuroscience.

# Functional Neuroimaging Techniques

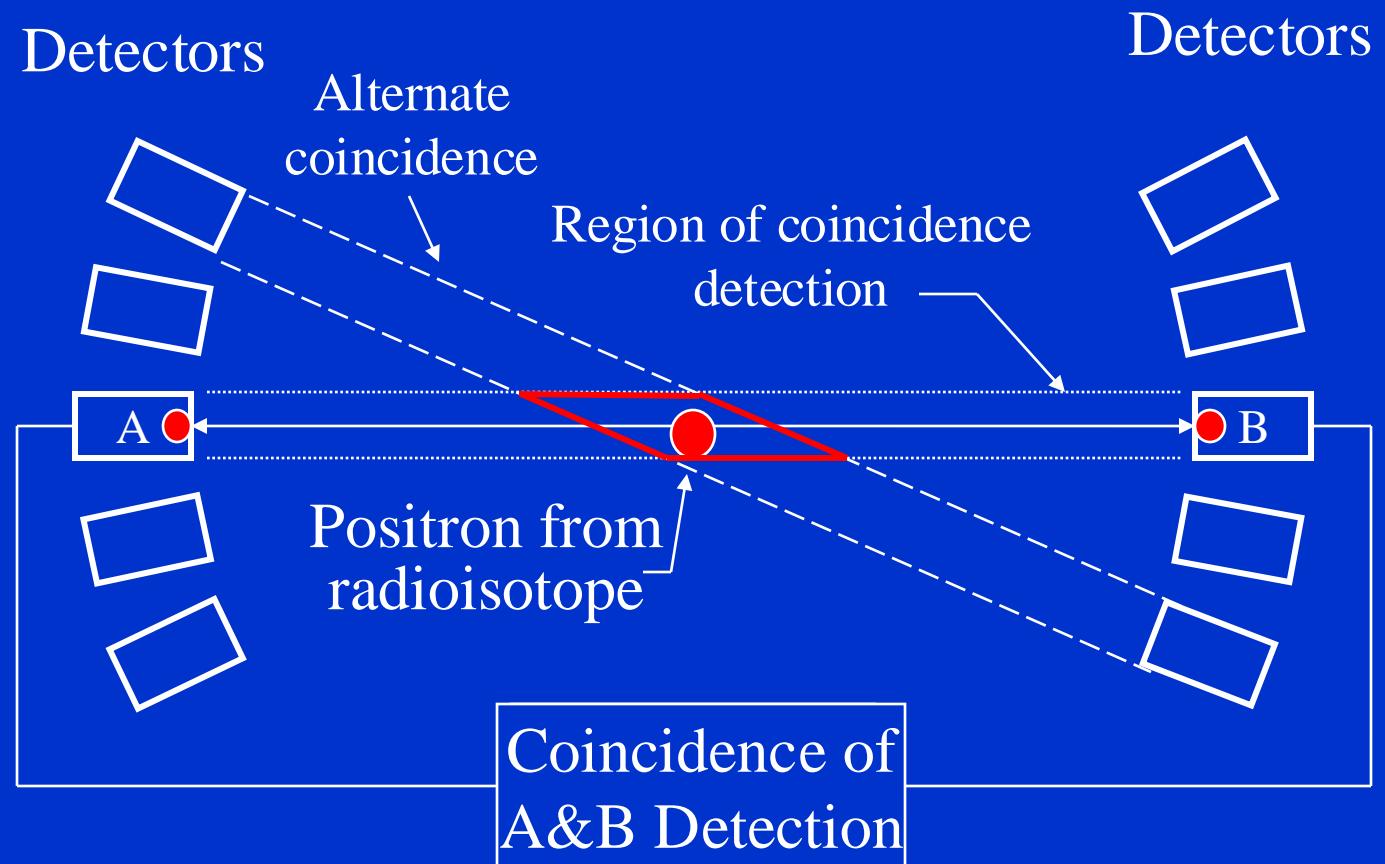


# Positron Emission Tomography (PET)

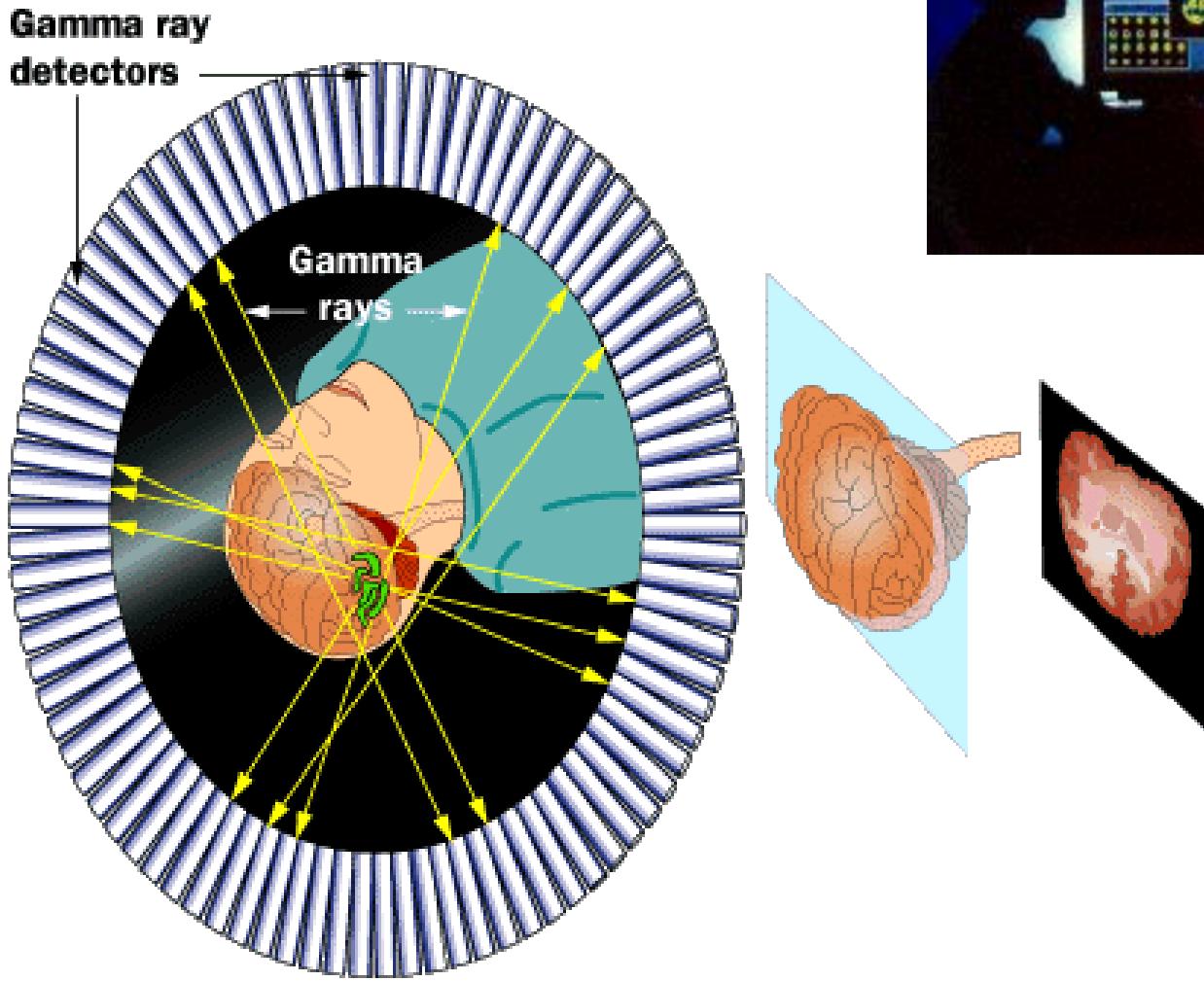
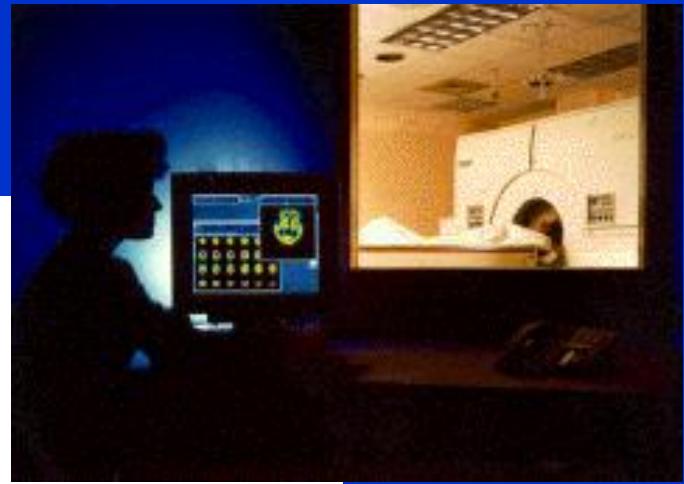
- Positron emission tomography (PET) is a technique for studying functional processes *in vivo* by measuring the concentrations of positron-emitting radioisotopes within the subject.
- PET is primarily used to study biochemical and physiological processes within living organs with 3-dimensional spatial resolution.



# PET mechanism



# Positron Emission Tomography

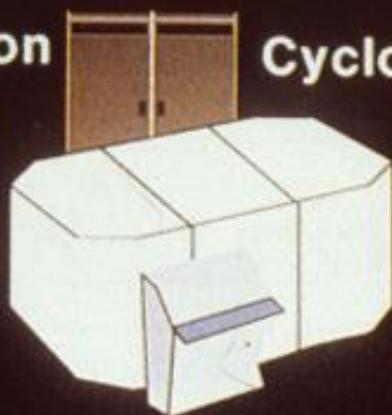




# Imaging of neuroreceptors by PET

Isotope production

[ $^{11}\text{C}$   $^{18}\text{F}$   $^{13}\text{N}$   $^{15}\text{O}$ ]



Cyclotron

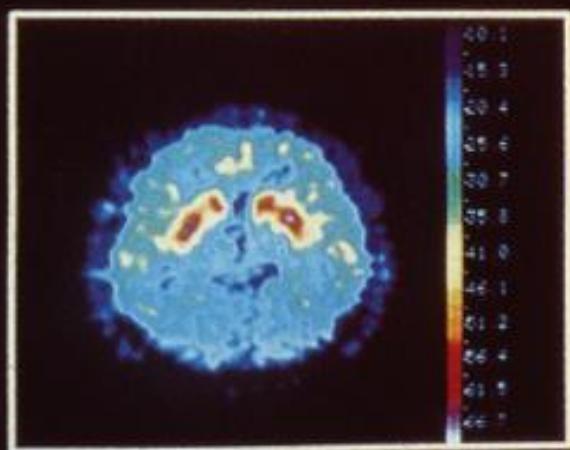
$^{11}\text{CO}_2$

Radio chemistry

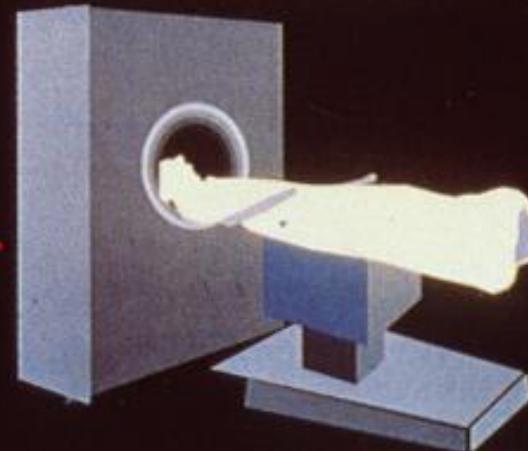
Precursor



Image of  
ligand distribution  
in brain

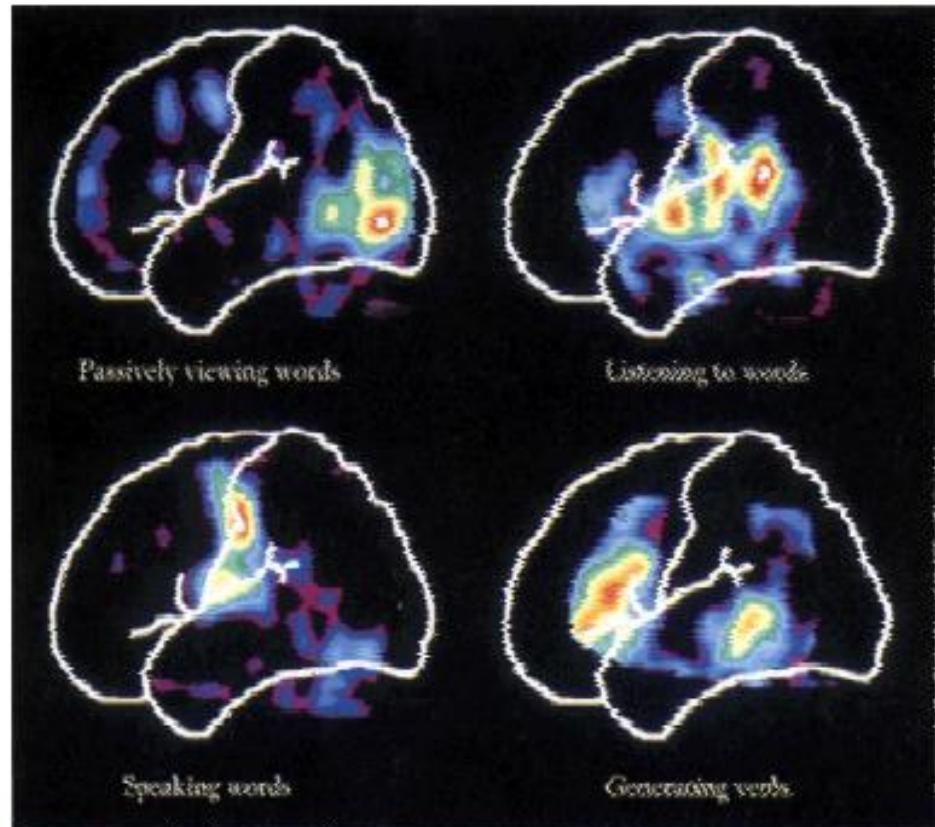
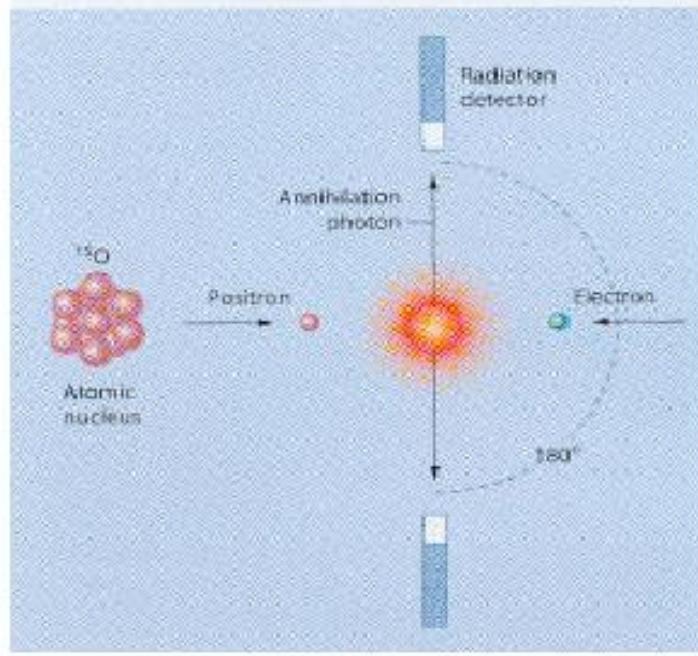


Positron camera

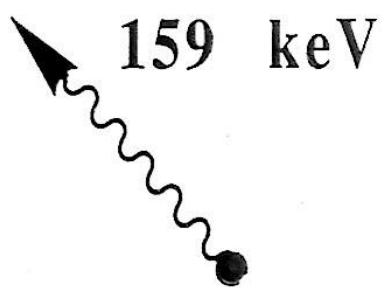


$^{11}\text{C}$ -ligand



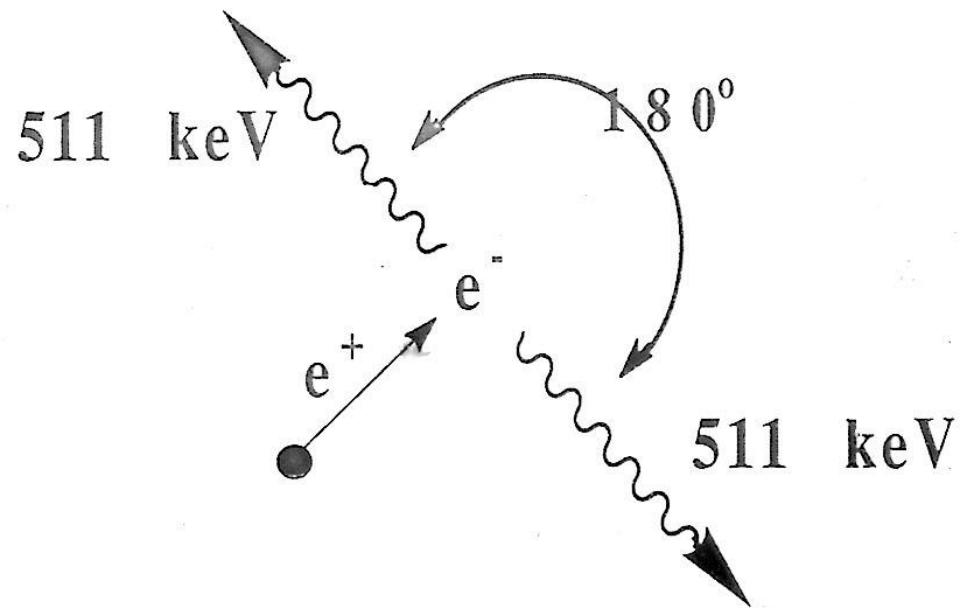


# SPECT



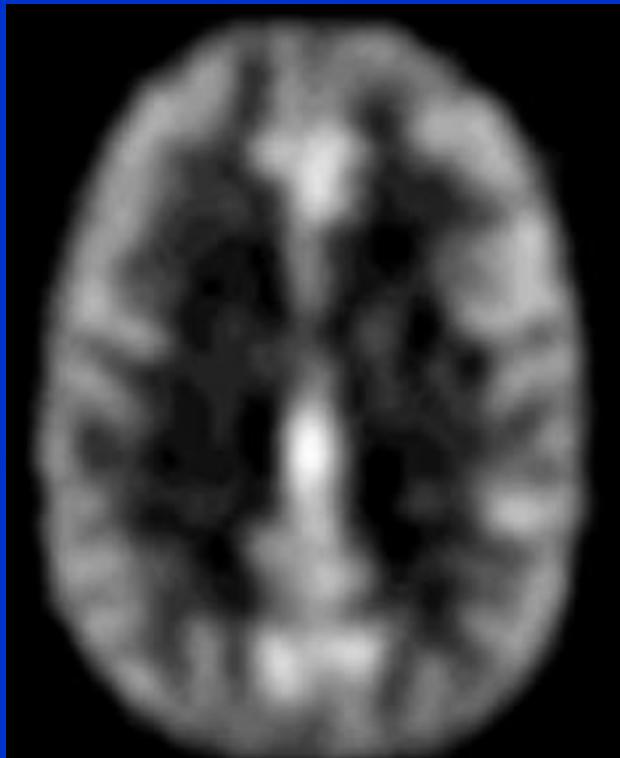
Single Photon  
Source  $^{123}\text{I}$

# PET

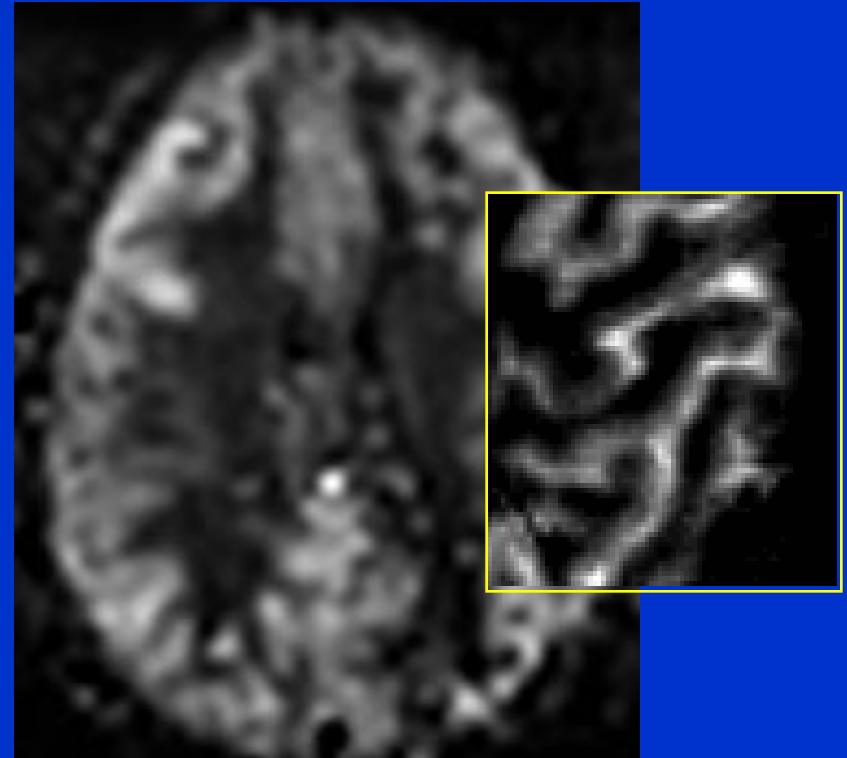


Positron Source

# Comparison with Positron Emission Tomography

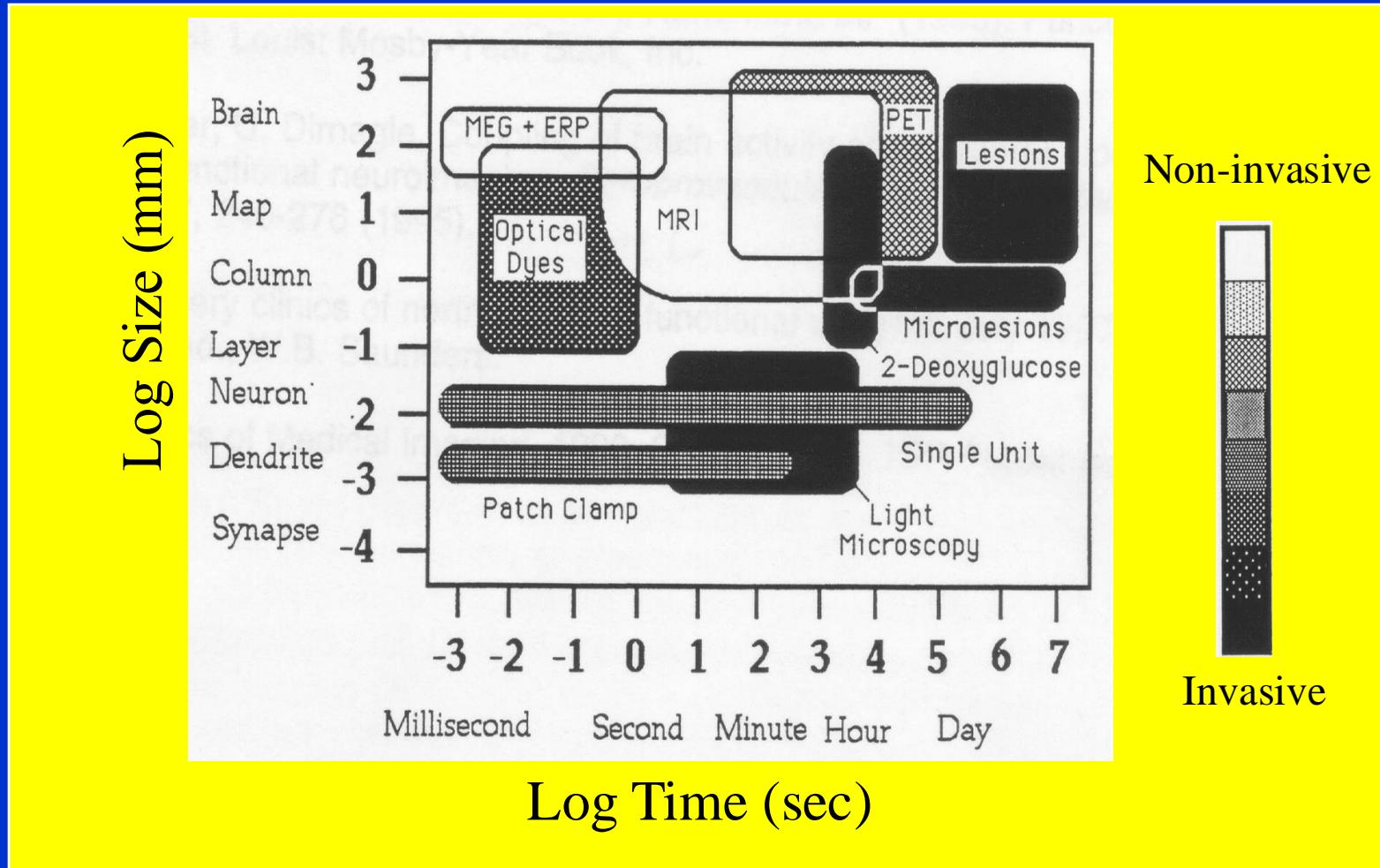


PET:  $\text{H}_2^{15}\text{O}$



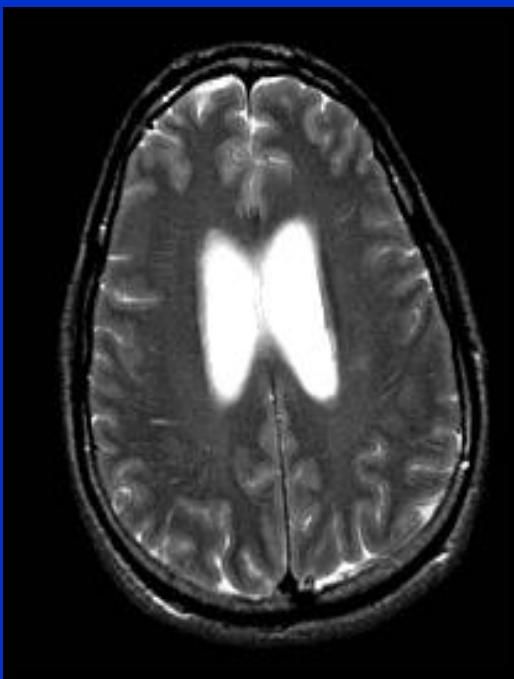
MRI: ASL

# Functional Neuroimaging Techniques

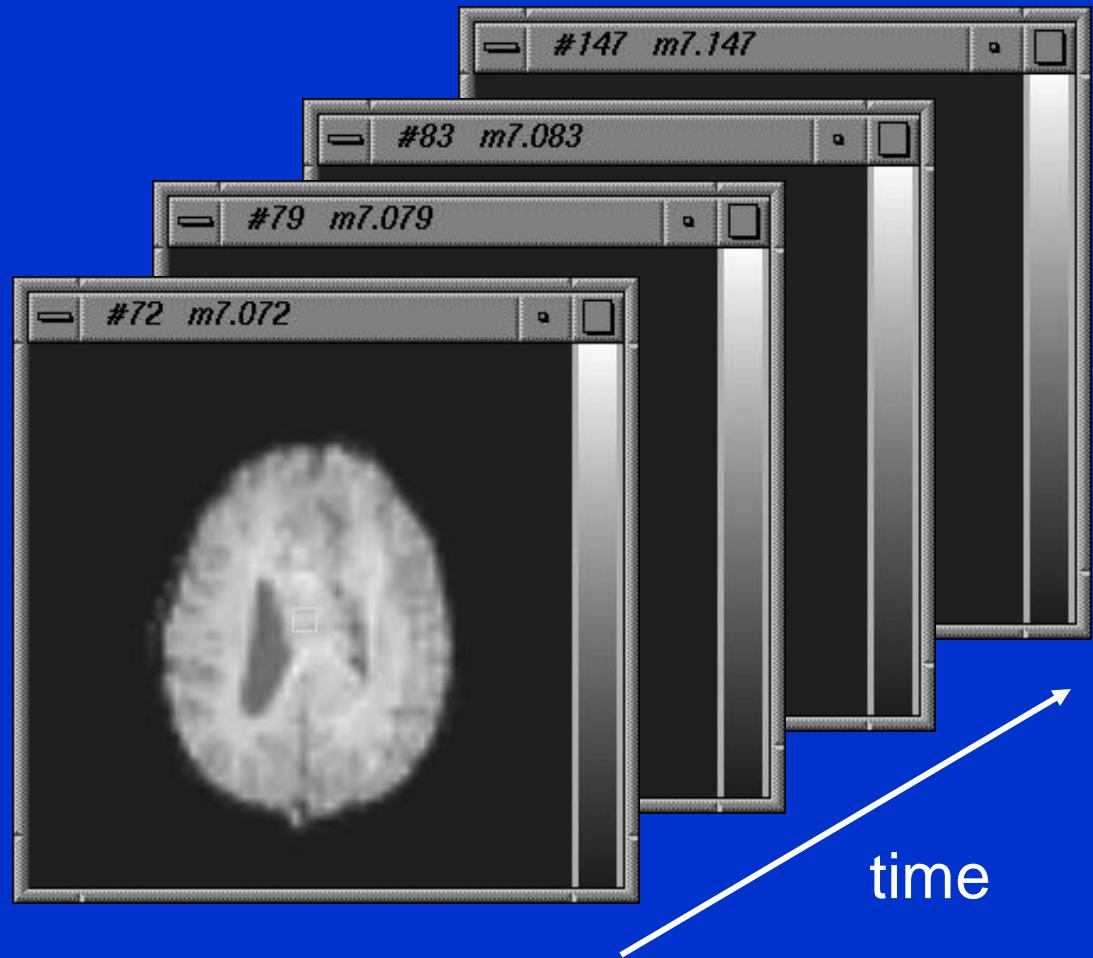


# Contrast in Functional MRI

- Blood Volume
- **BOLD** (Blood Oxygenation Level Dependent Contrast)
- Perfusion
- CMRO<sub>2</sub>

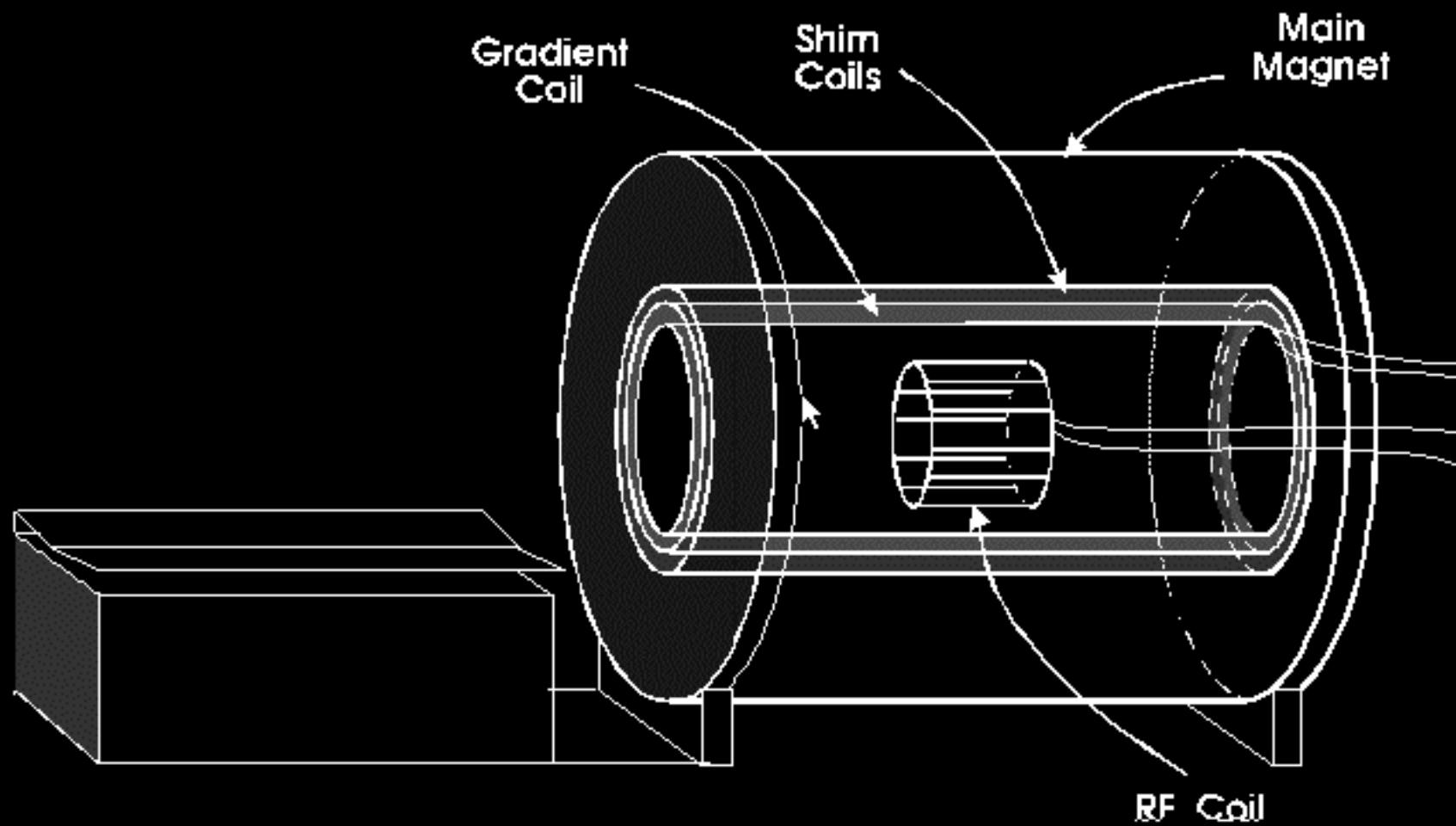


Anatomic



Functional

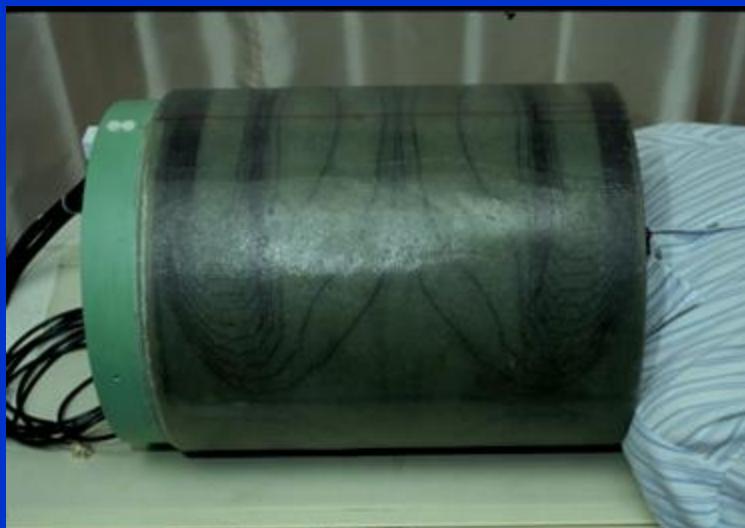
time



**1991-1992**



**1992-1999**



# General Electric 3 Tesla Scanner





# MRI Instrumentation

Computer  
Graphics



Video Image  
Source

Baseline Task

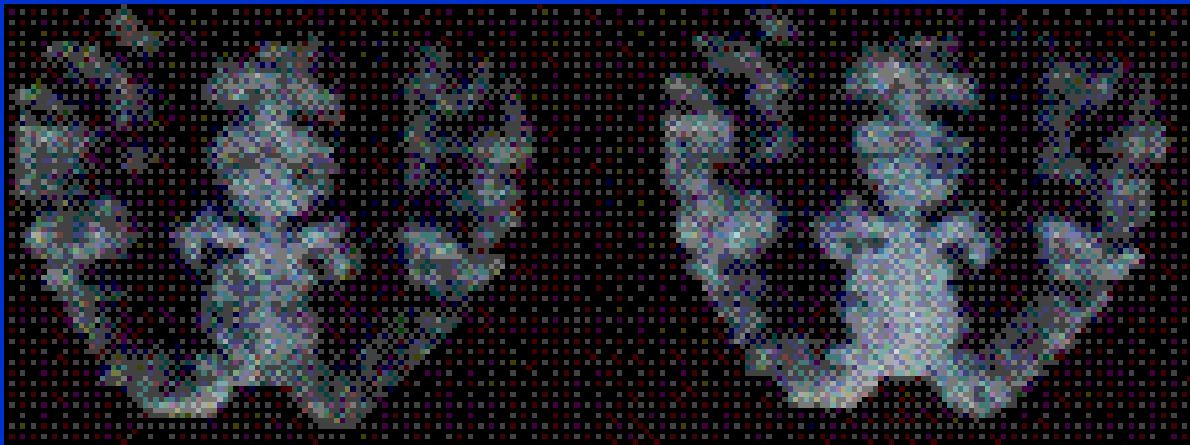
Experimental Task



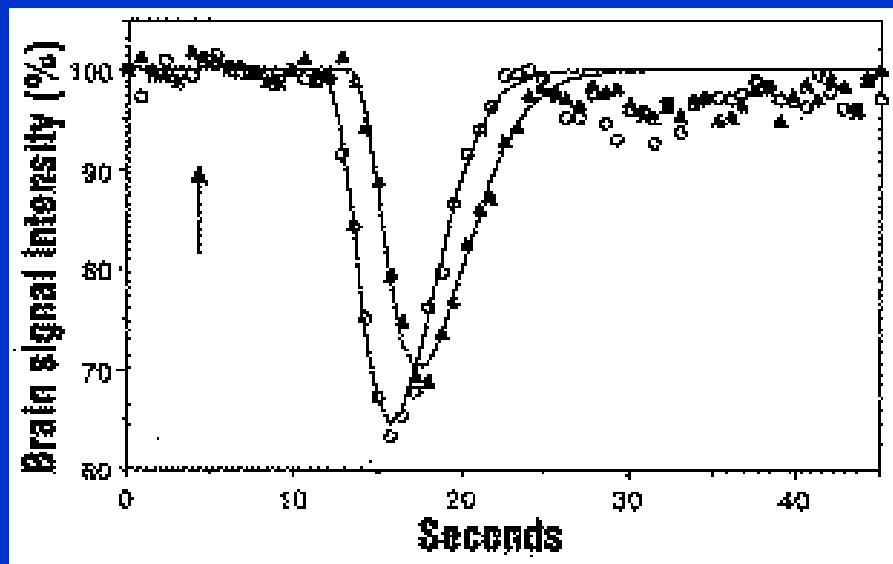
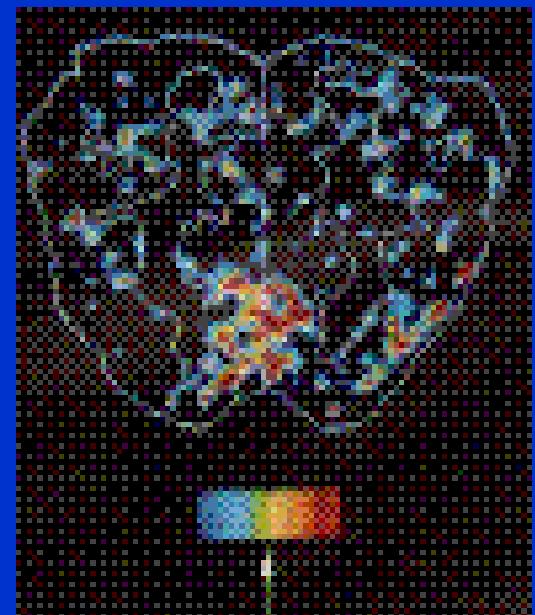


# Blood Volume Changes with Brain Activation

**Resting**



**Active**



# Photic Stimulation

MRI Image showing  
activation of the  
Visual Cortex

From Belliveau, et al.  
Science Nov 1991

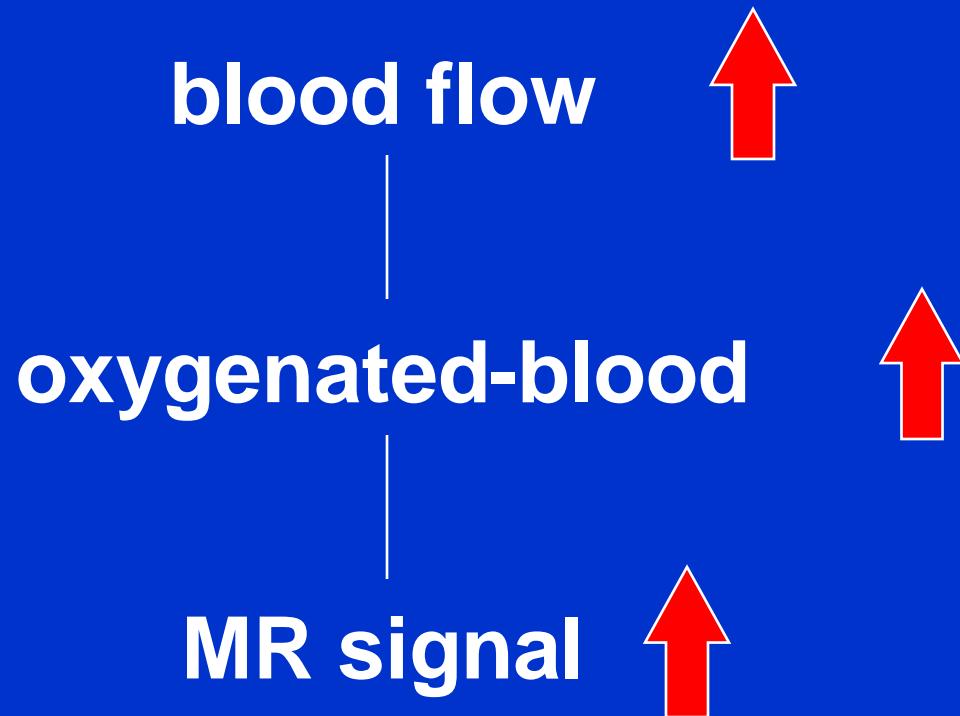
MSC - perfusion



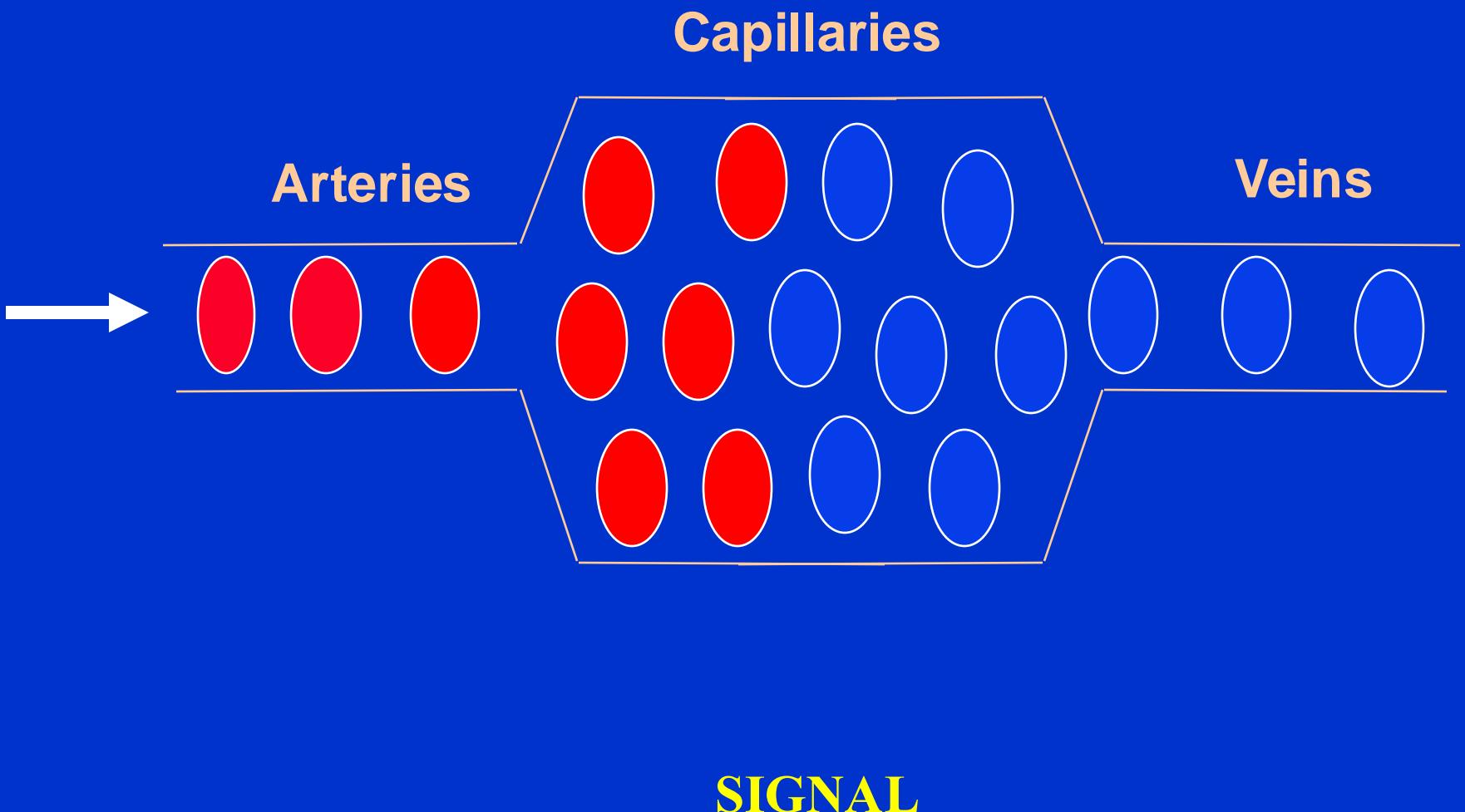
# BOLD

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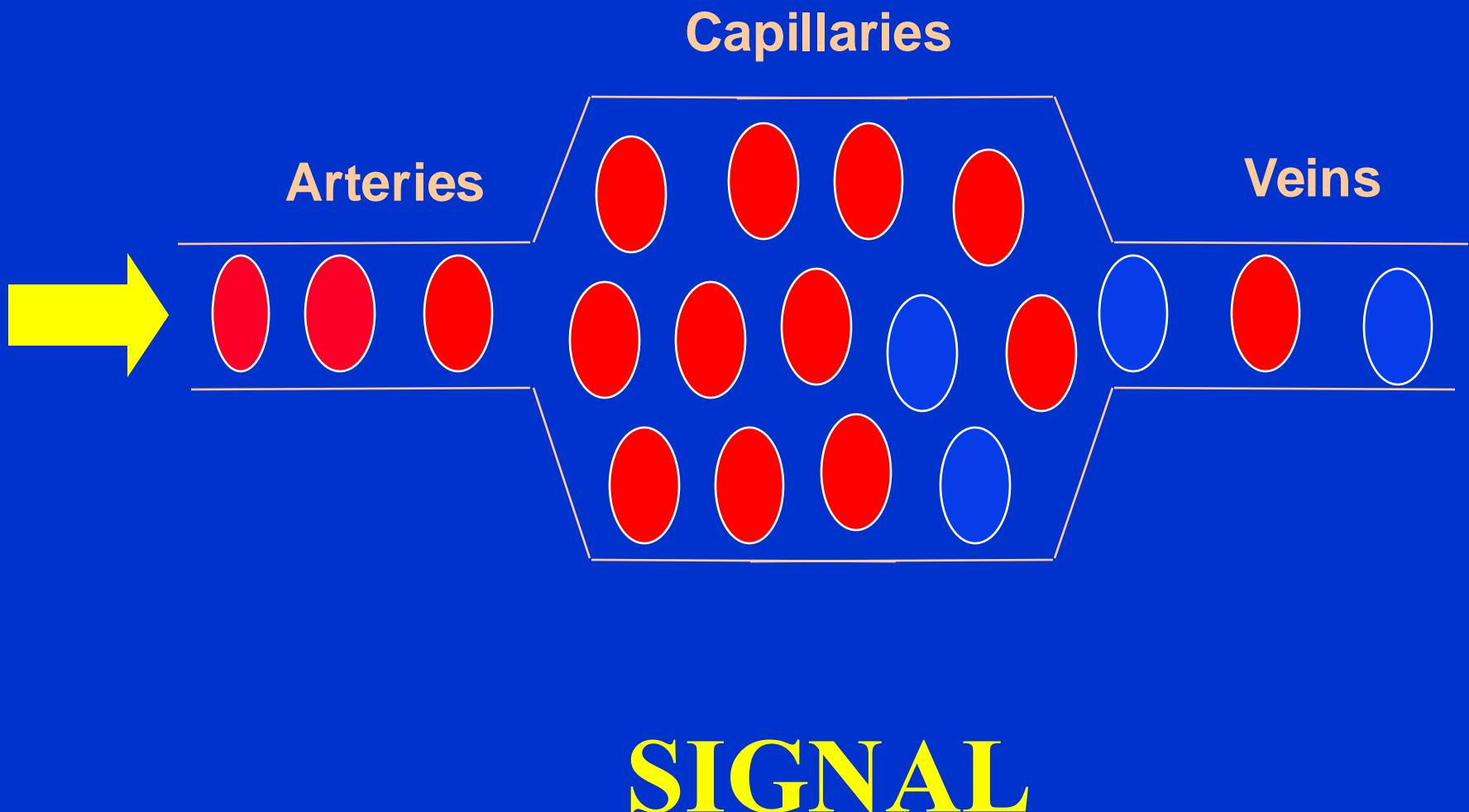
(**blood** oxygenation level dependence)



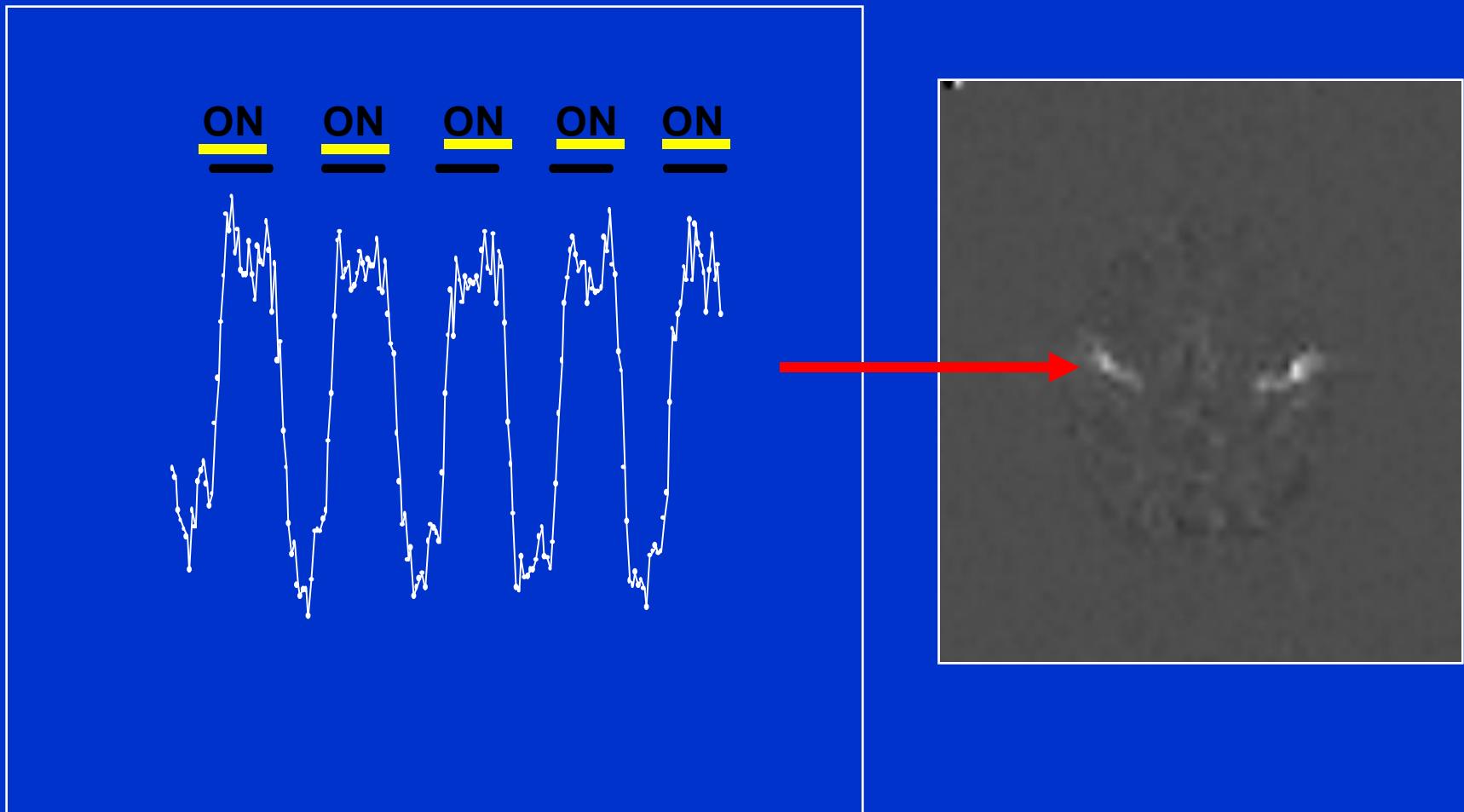
# **BOLD:** Resting Perfusion



# BOLD: Activated Perfusion

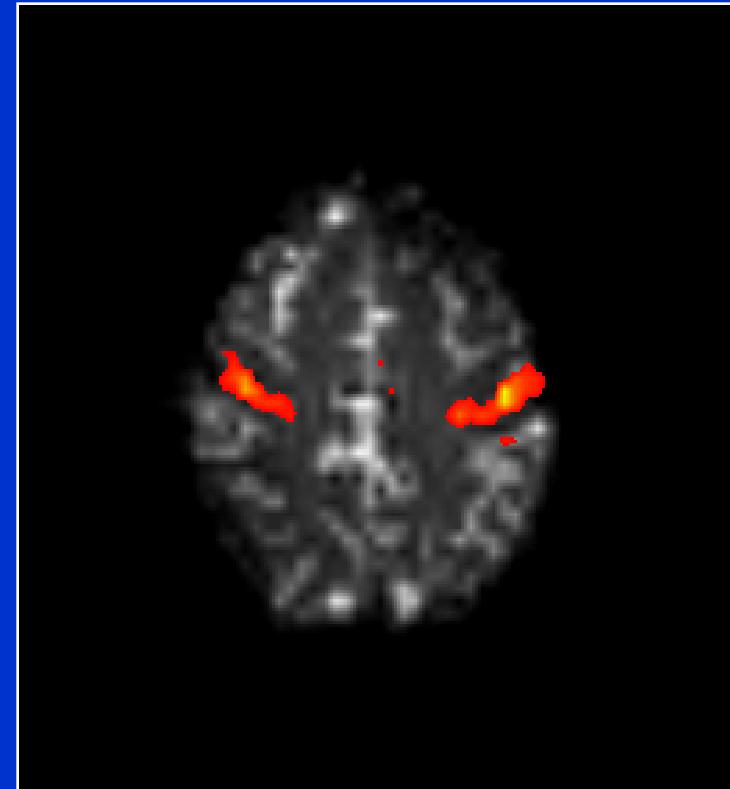


# BOLD: Motor Cortex Activation





Cross Correlation Image

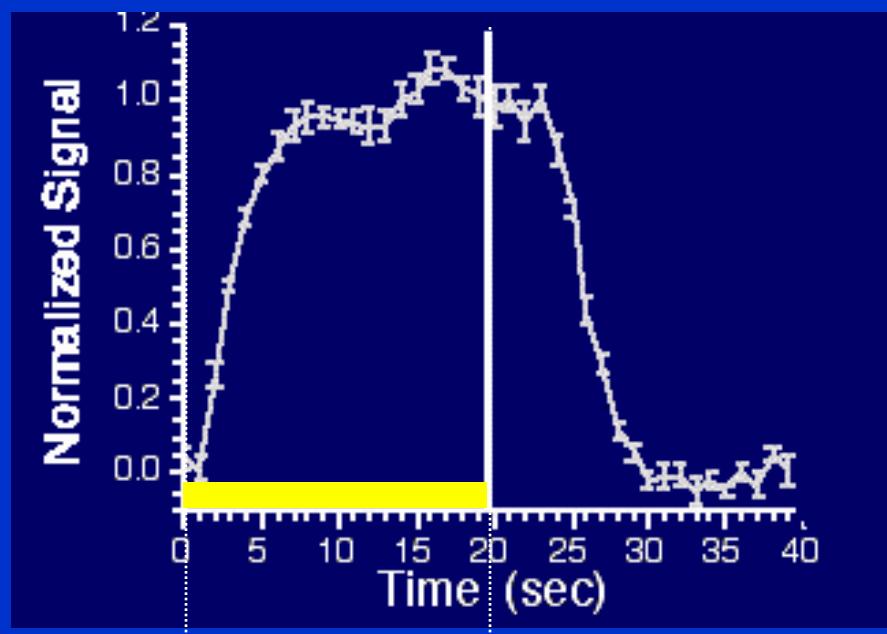


Cross Correlation Image  
Anatomical Image

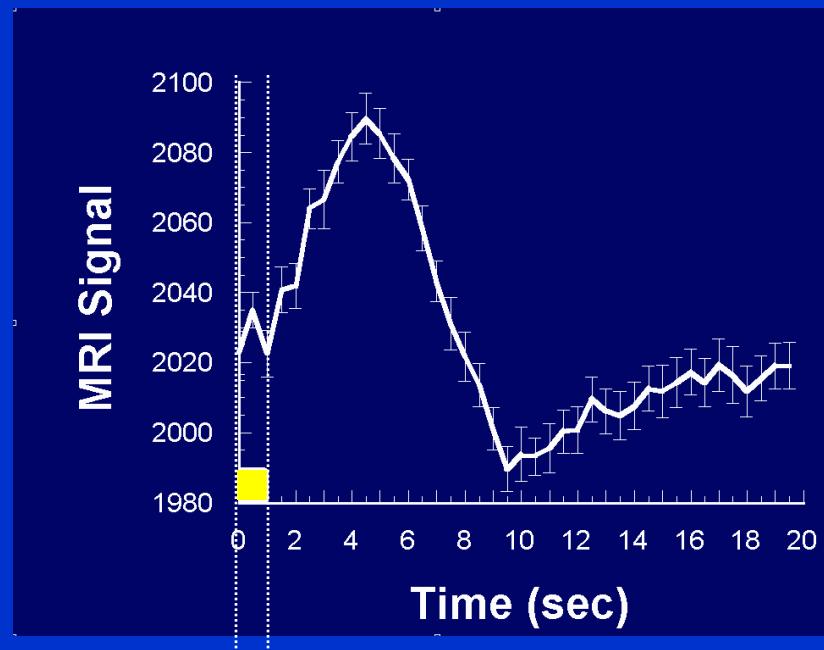


# The BOLD Signal

Blood Oxygenation Level Dependent (BOLD) signal changes



*task*

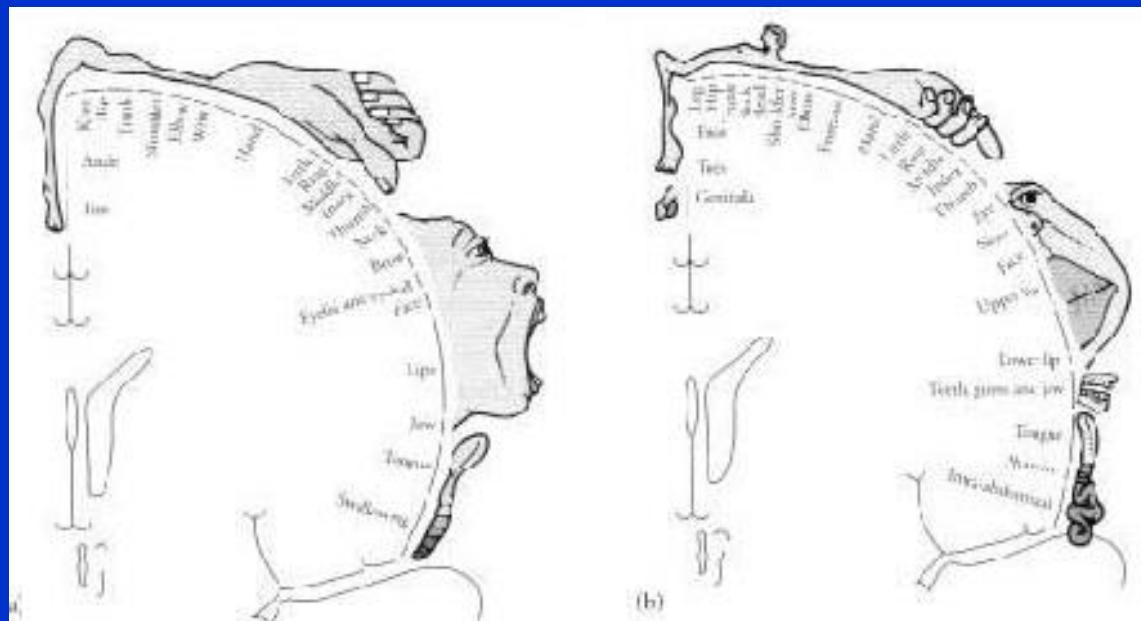
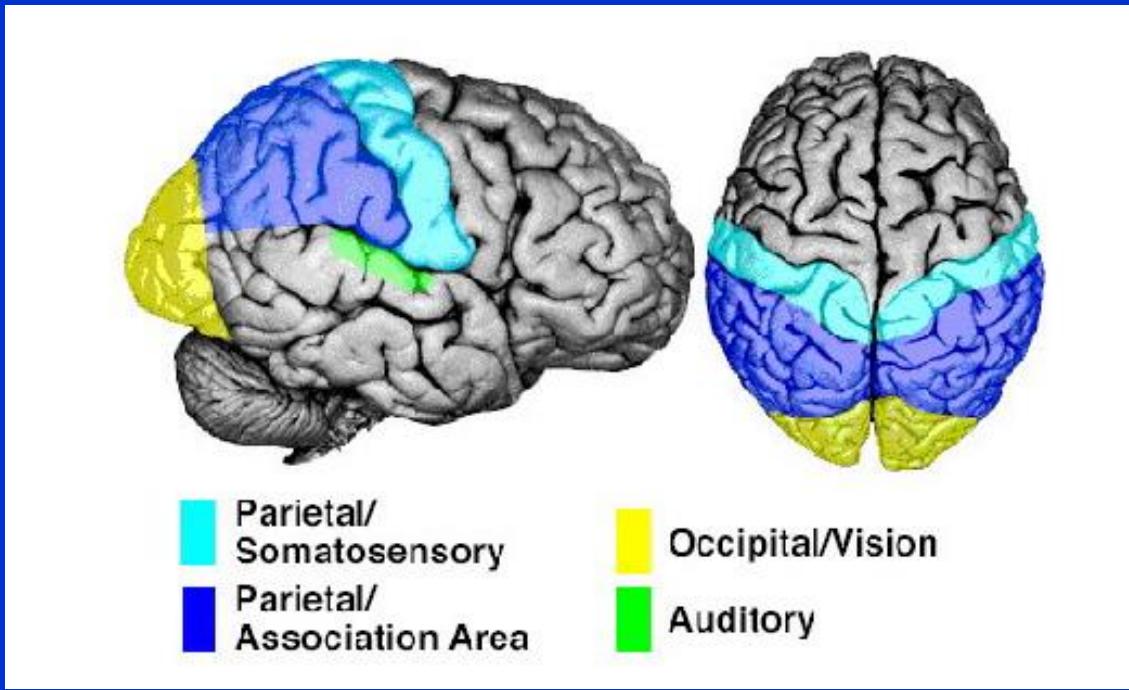


*task*

# Alternating Left and Right Finger Tapping



~ 1992



# Finger Movement

Left

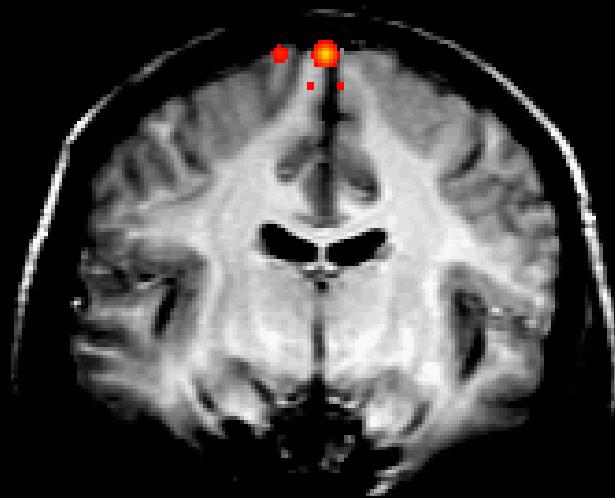


Right

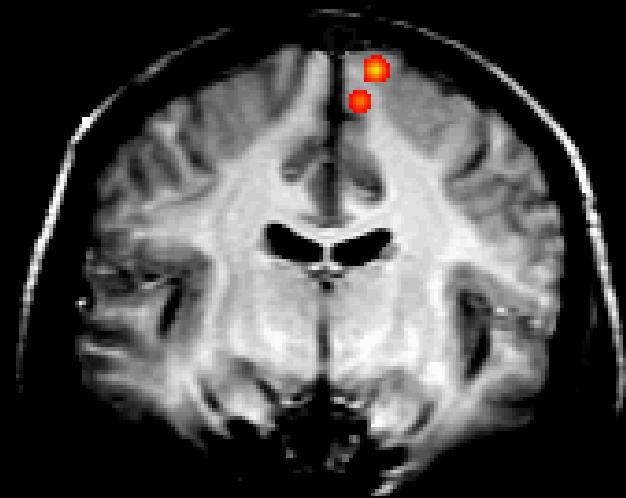


# Toe Movement

Left



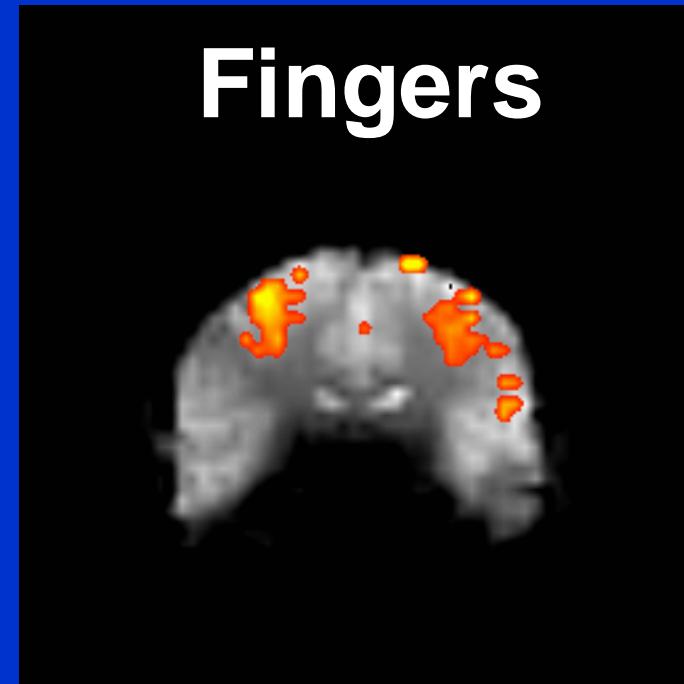
Right



**Wrist**

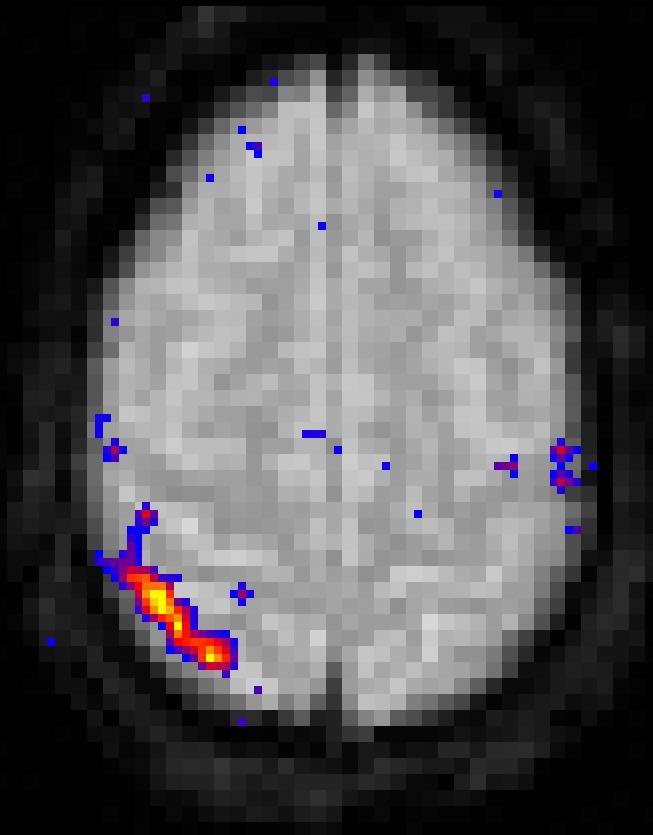
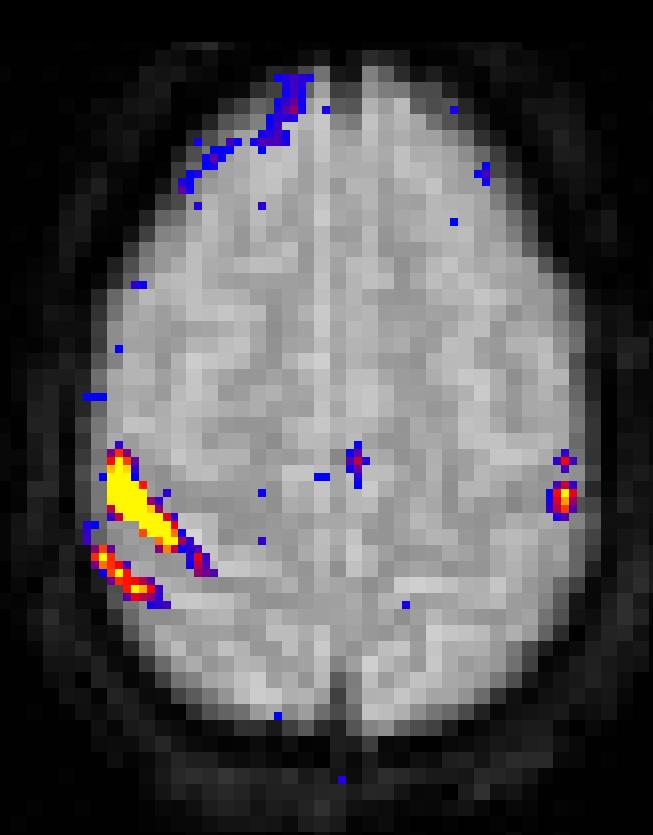


**Fingers**

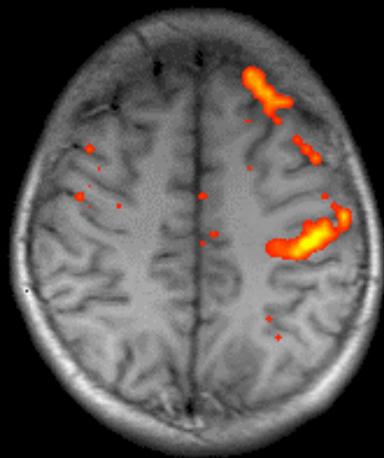


# Finger Movement

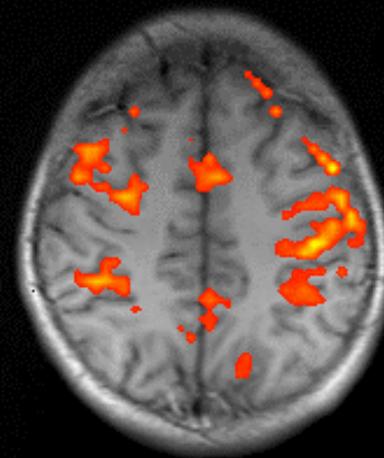
# Tactile Stimulation



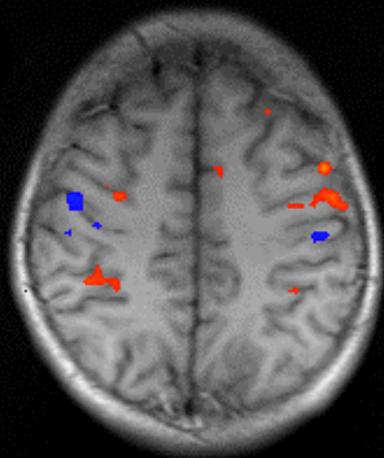
Simple Right



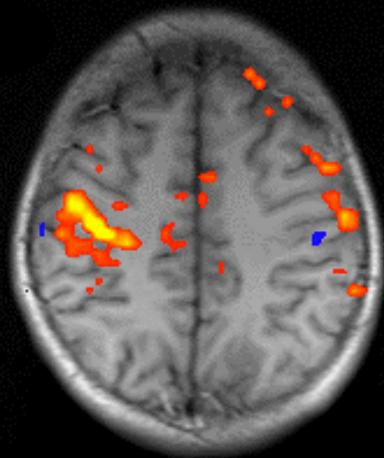
Complex Right



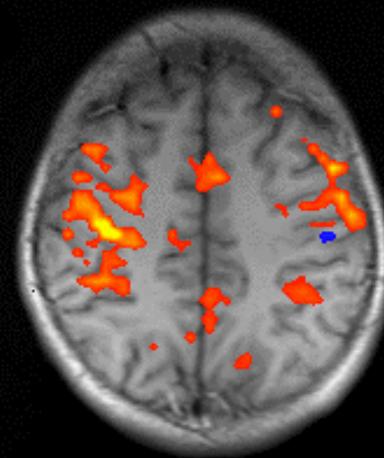
Imagined  
Complex Right



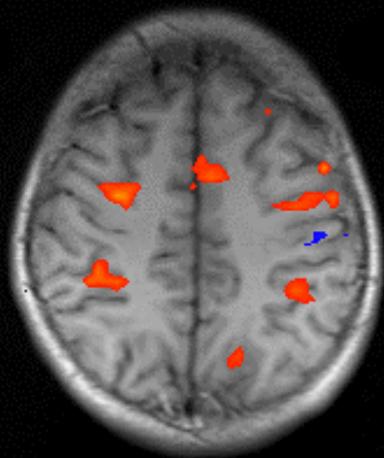
Simple Left



Complex Left



Imagined  
Complex Left



*Left*

## Simple Finger Movement on the Left Hand

1

2

3

4

5

6

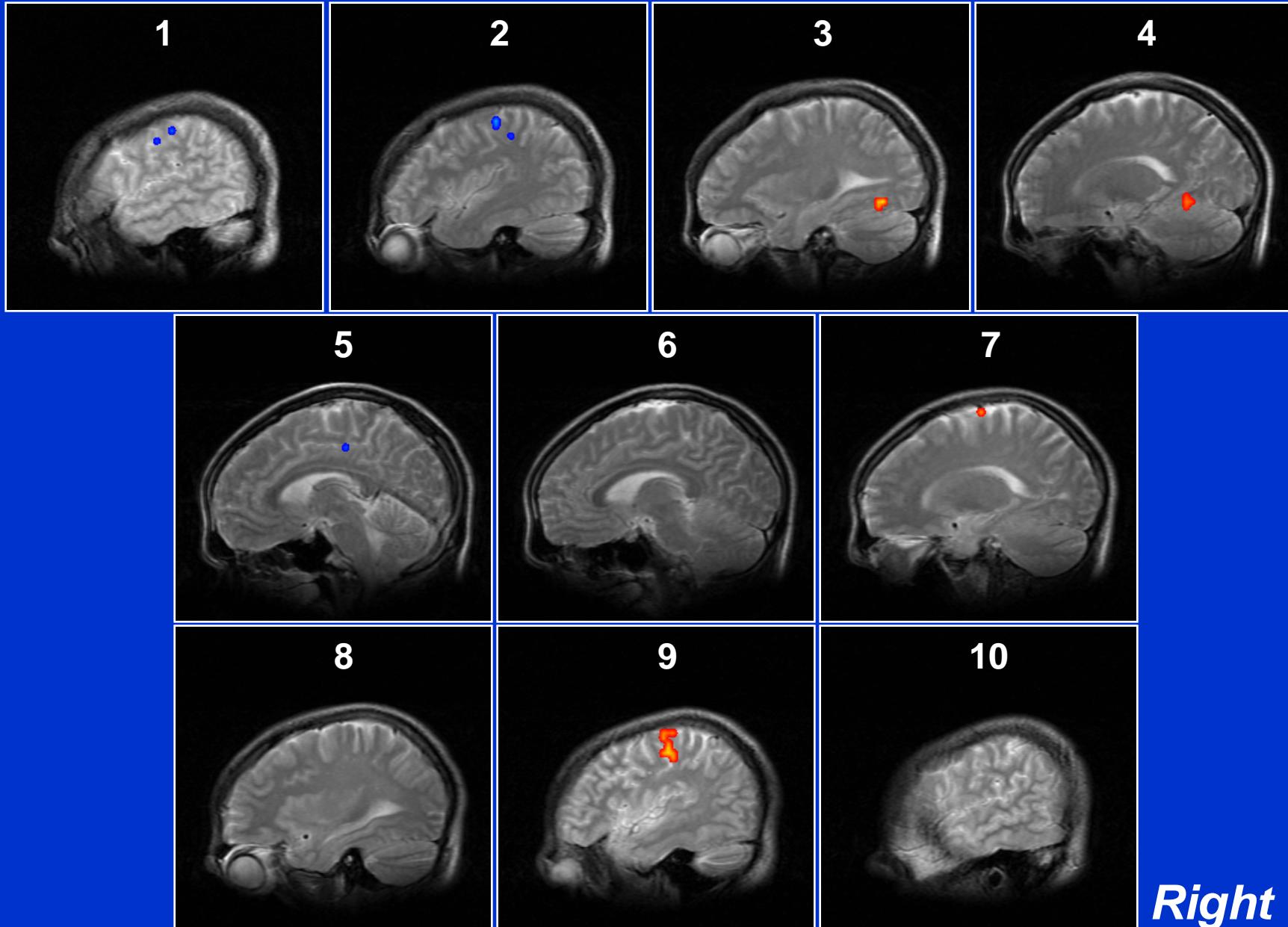
7

8

9

10

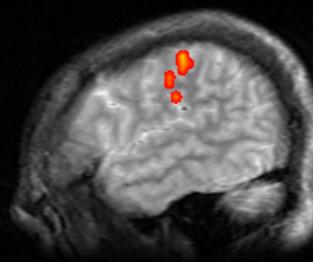
*Right*



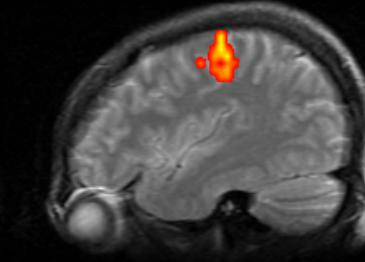
*Left*

## Simple Finger Movement on the Right Hand

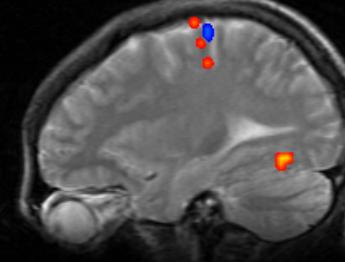
1



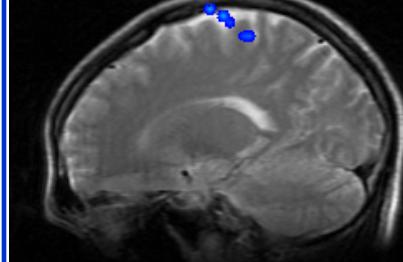
2



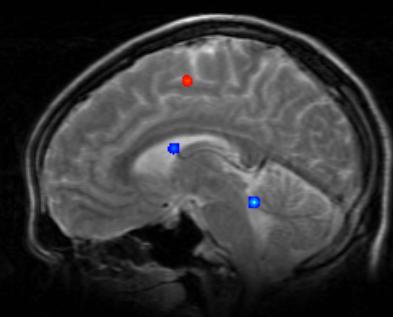
3



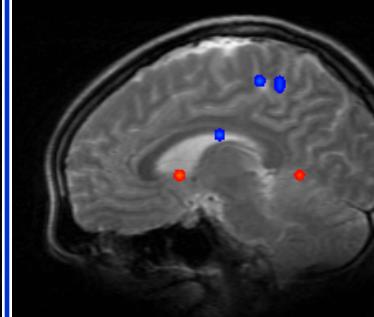
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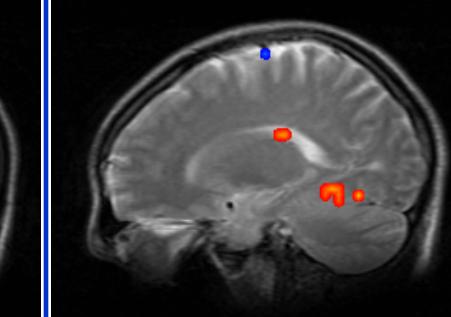
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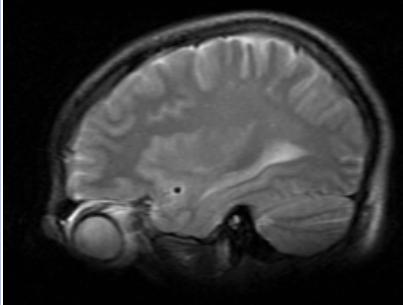
6



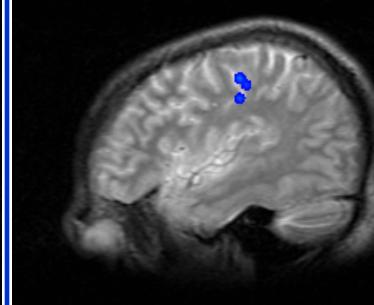
7



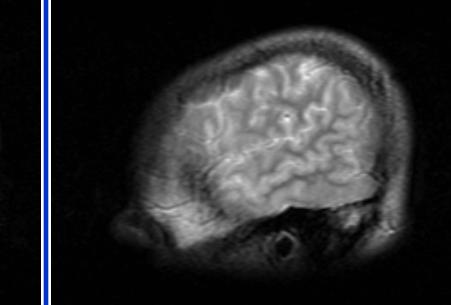
8



9



10



*Right*

*Left*

## Complex Finger Movement on the Right Hand

1

2

3

4

5

6

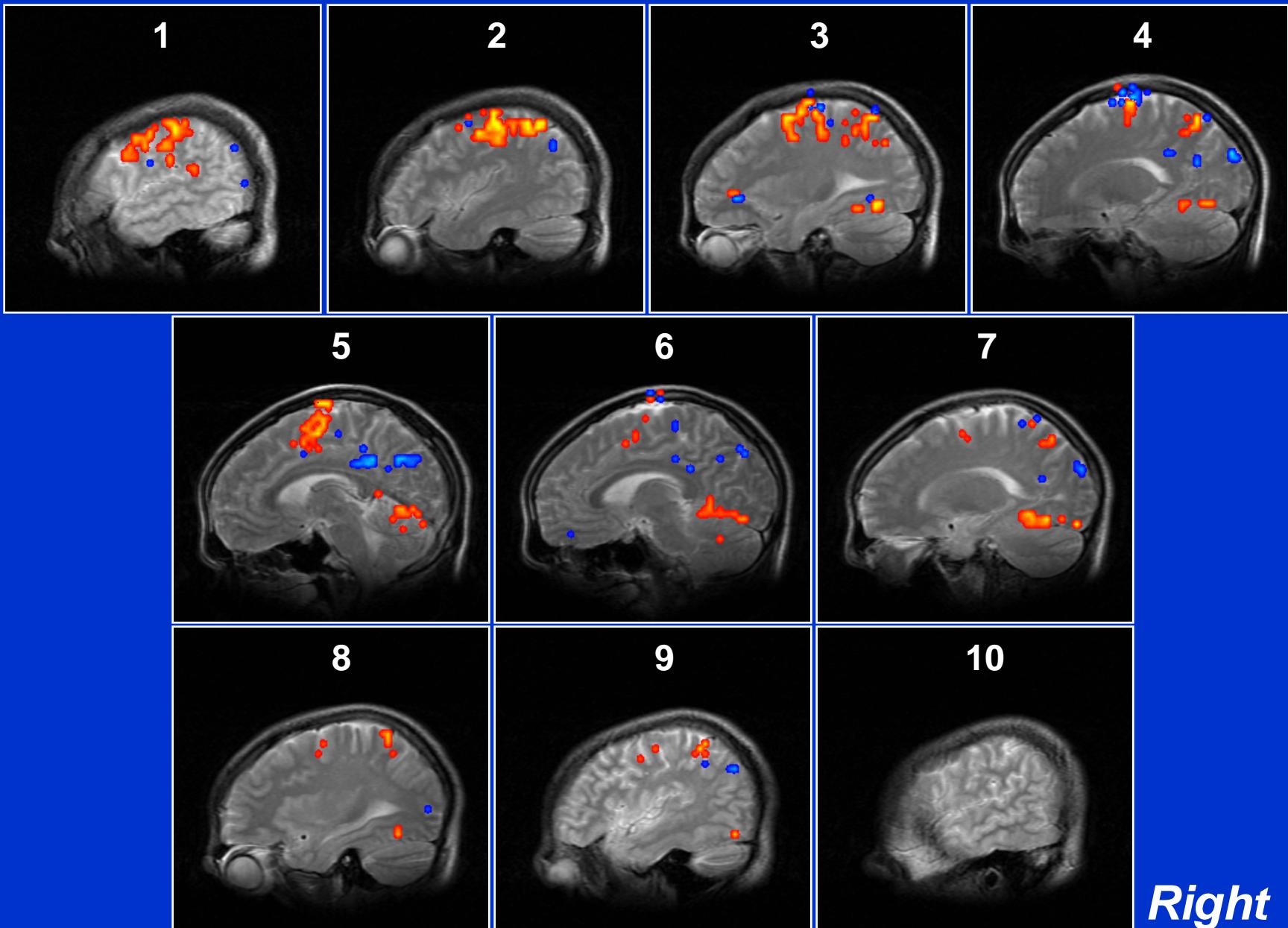
7

8

9

10

*Right*



*Left*

## Complex Finger Movement on the Left Hand

1

2

3

4

5

6

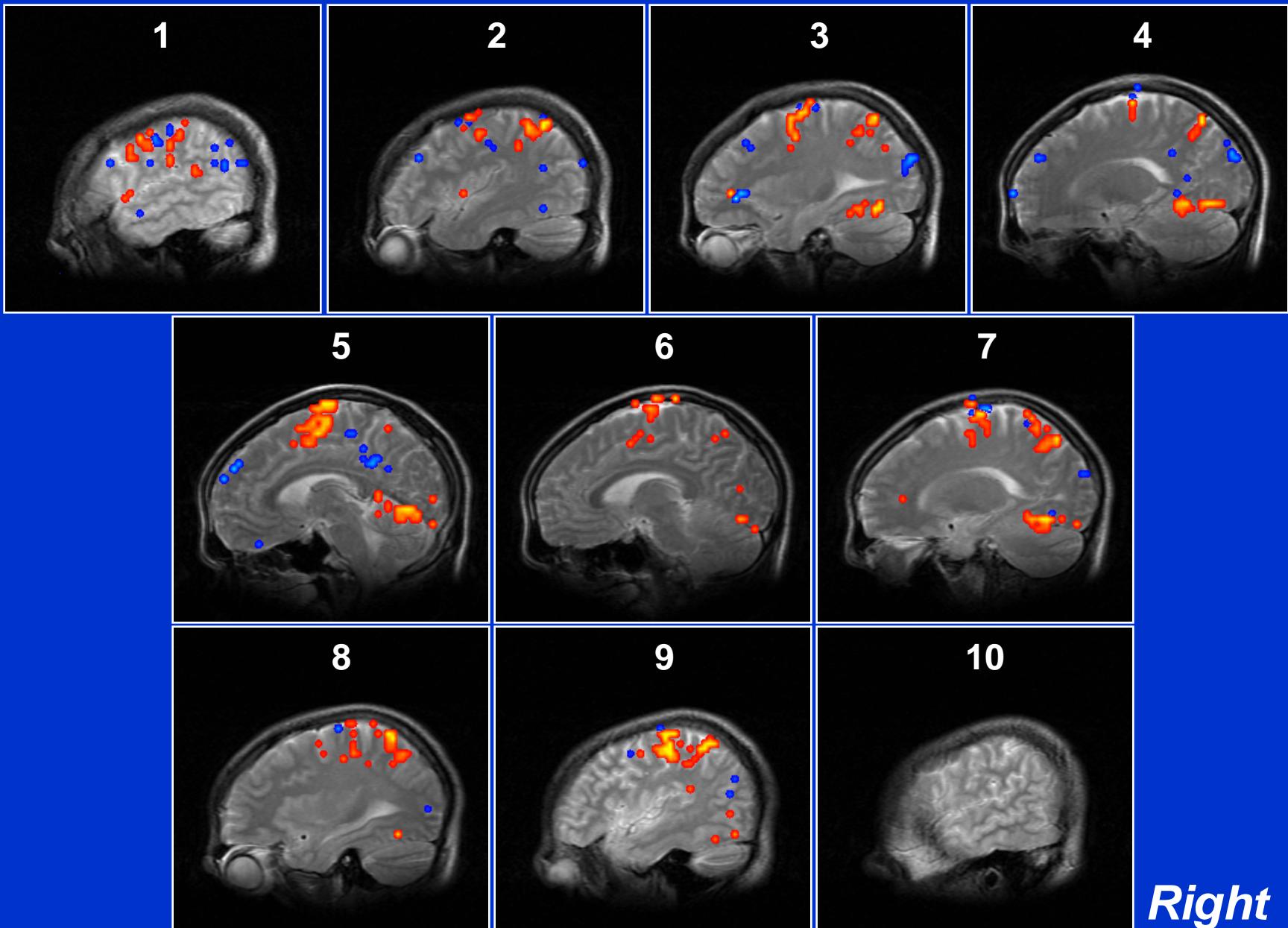
7

8

9

10

*Right*



*Left*

## Imagined Complex Finger Movement on the Left Hand

1

2

3

4

5

6

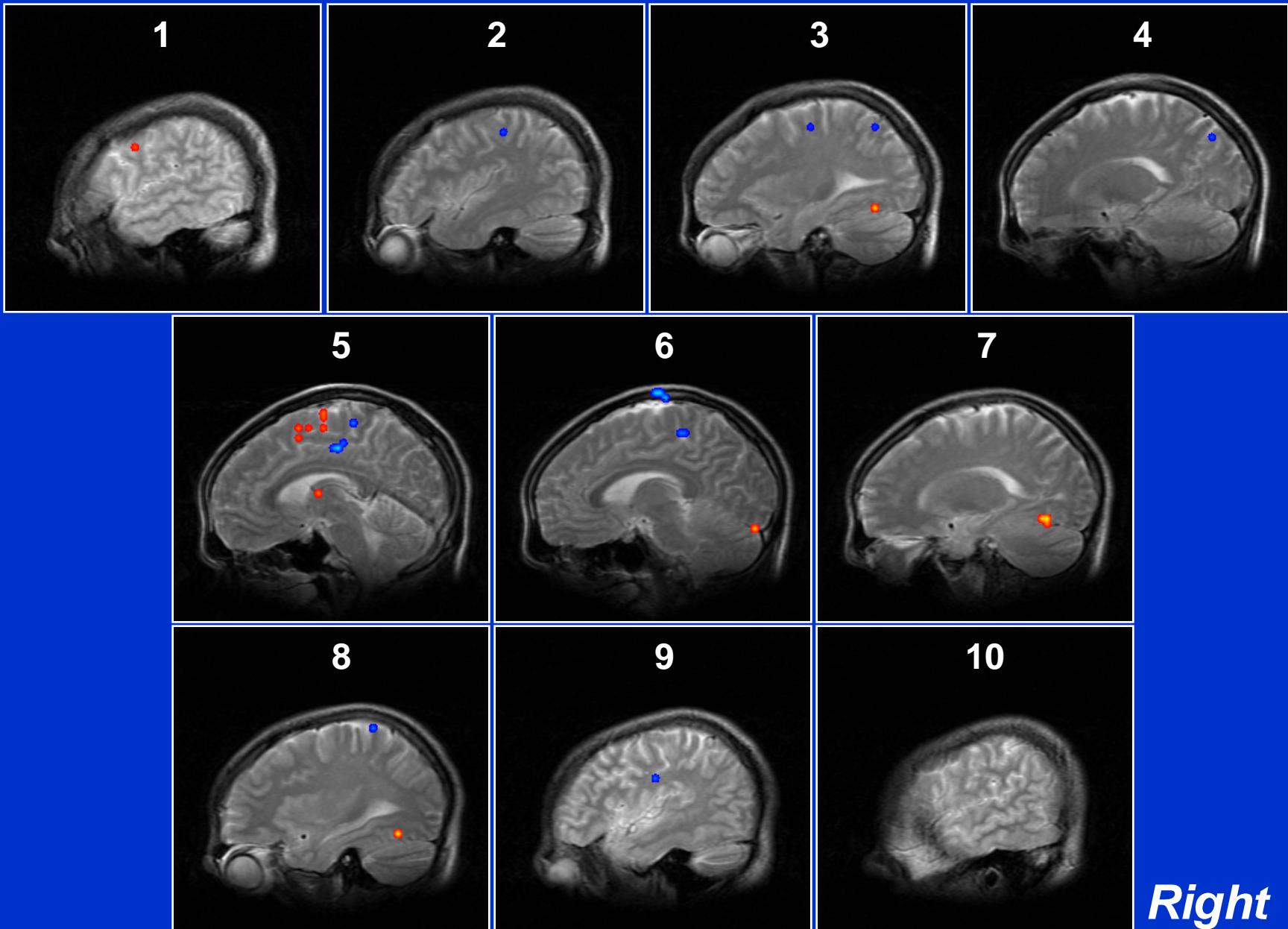
7

8

9

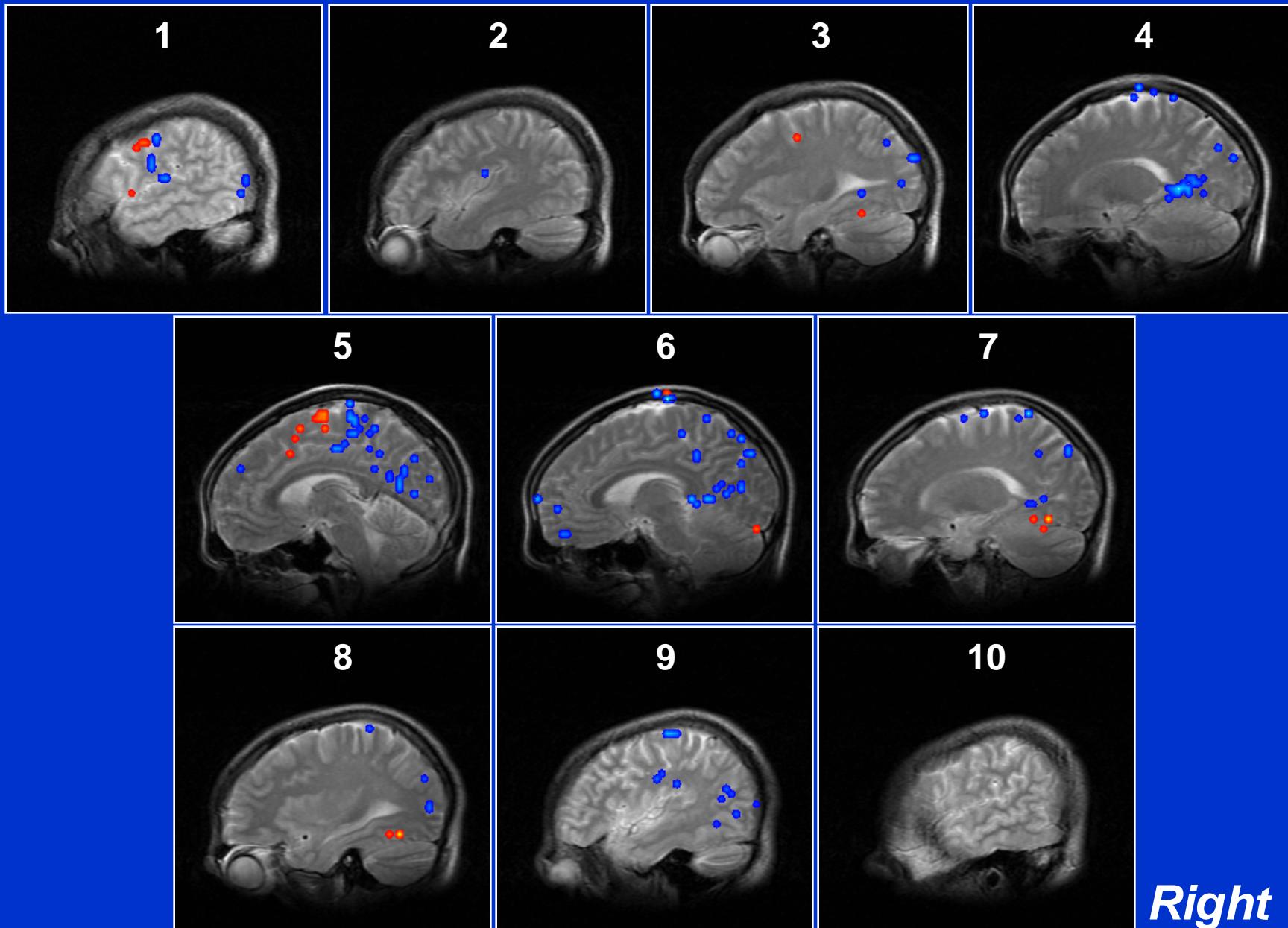
10

*Right*



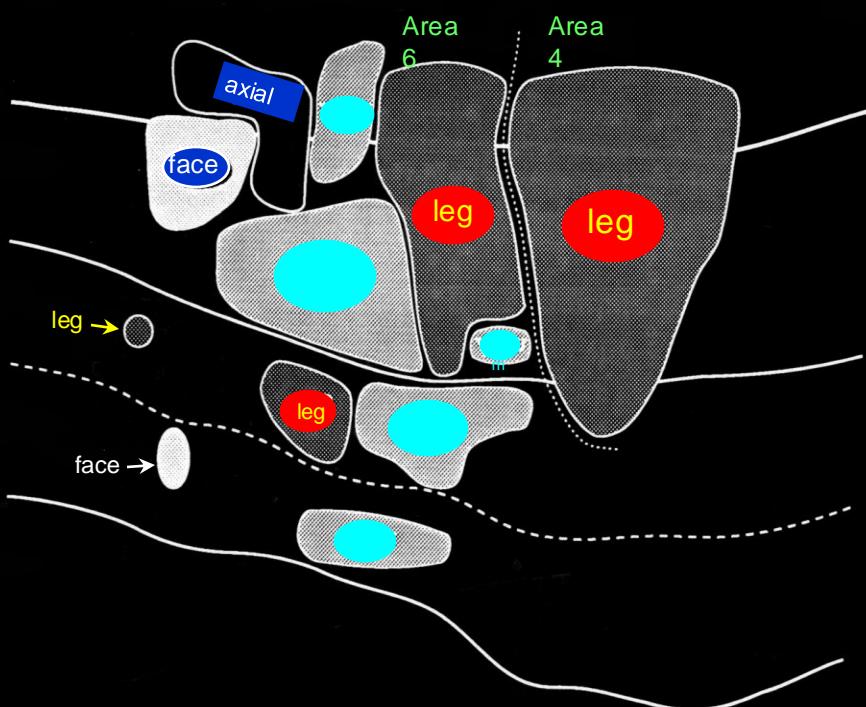
*Left*

## Imagined Complex Finger Movement on the Right Hand

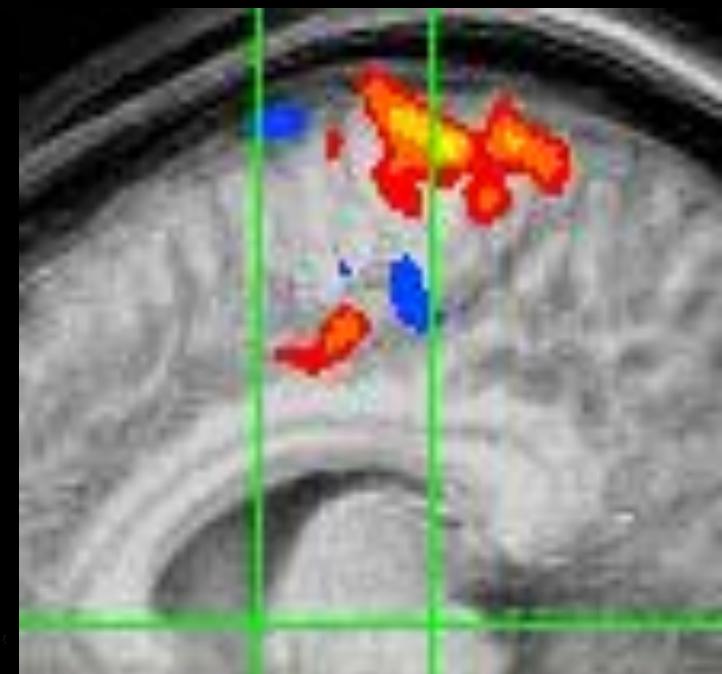


*Right*

# FMRI of Leg Motor Topography - Medial Wall



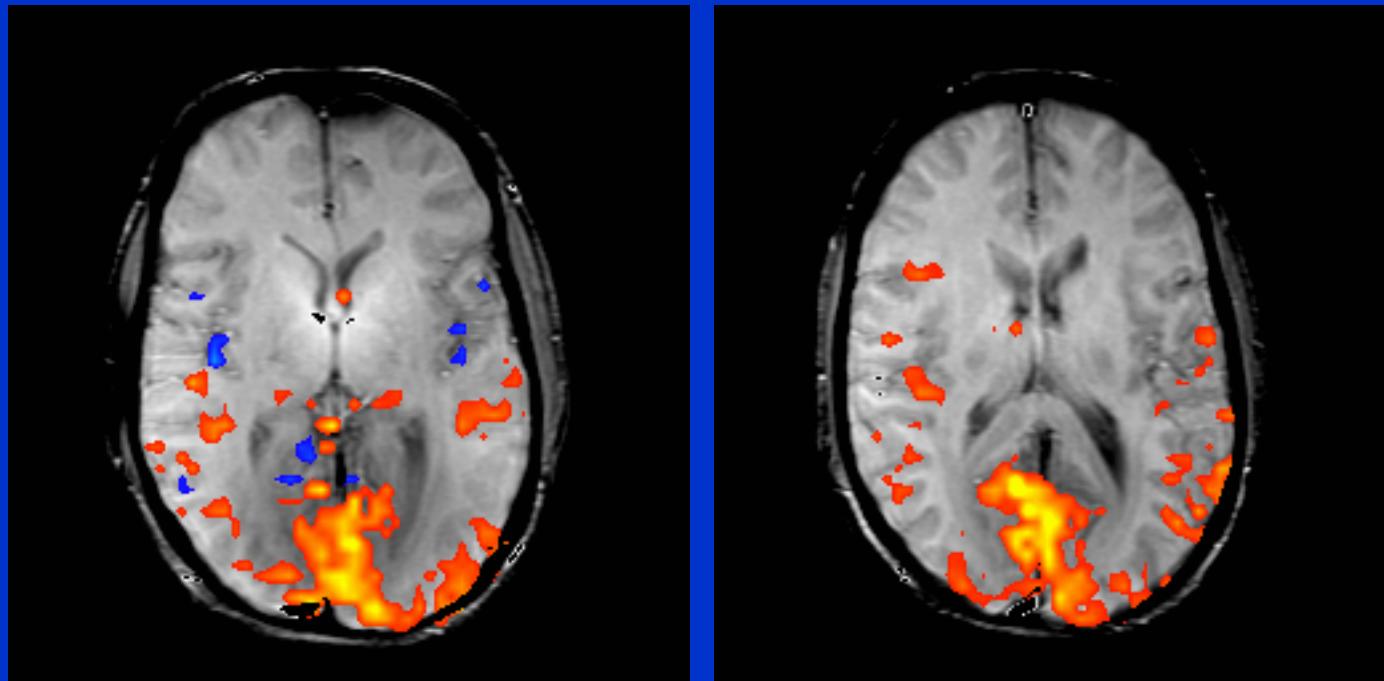
*Macaque*



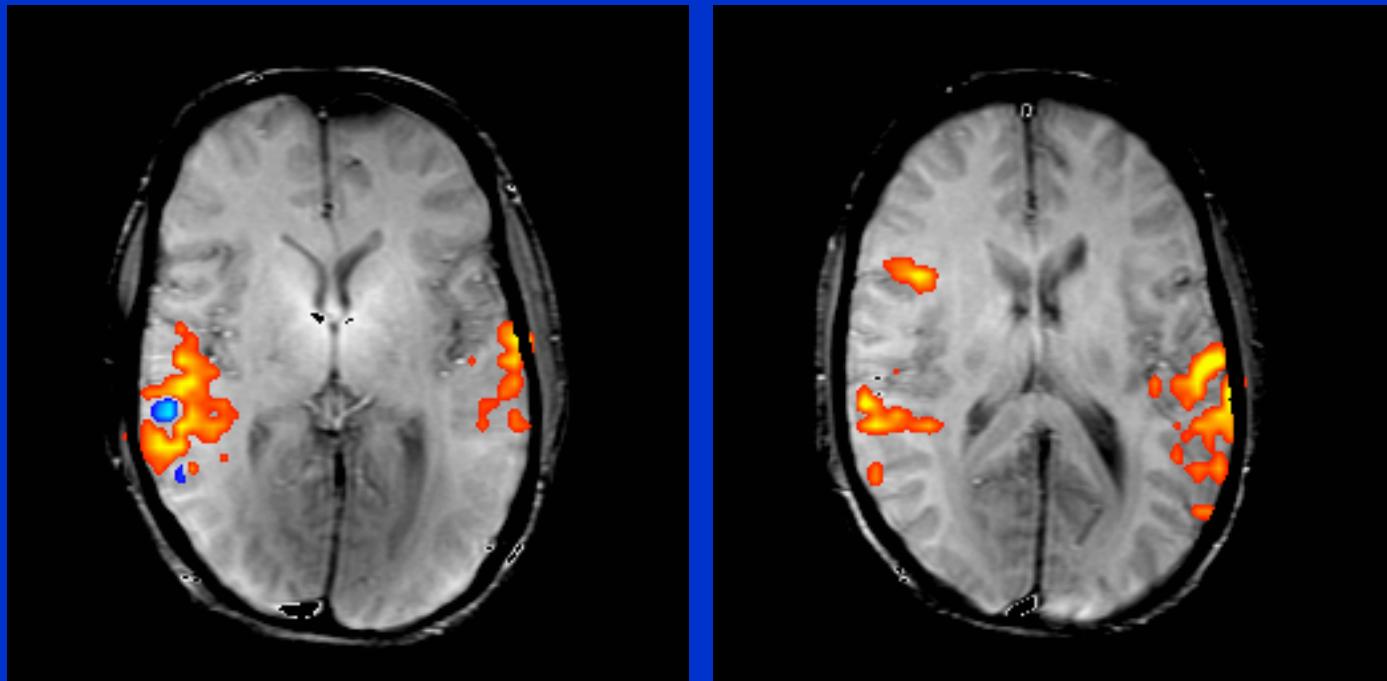
*Human*

Thanks to S. Rao

# Reading

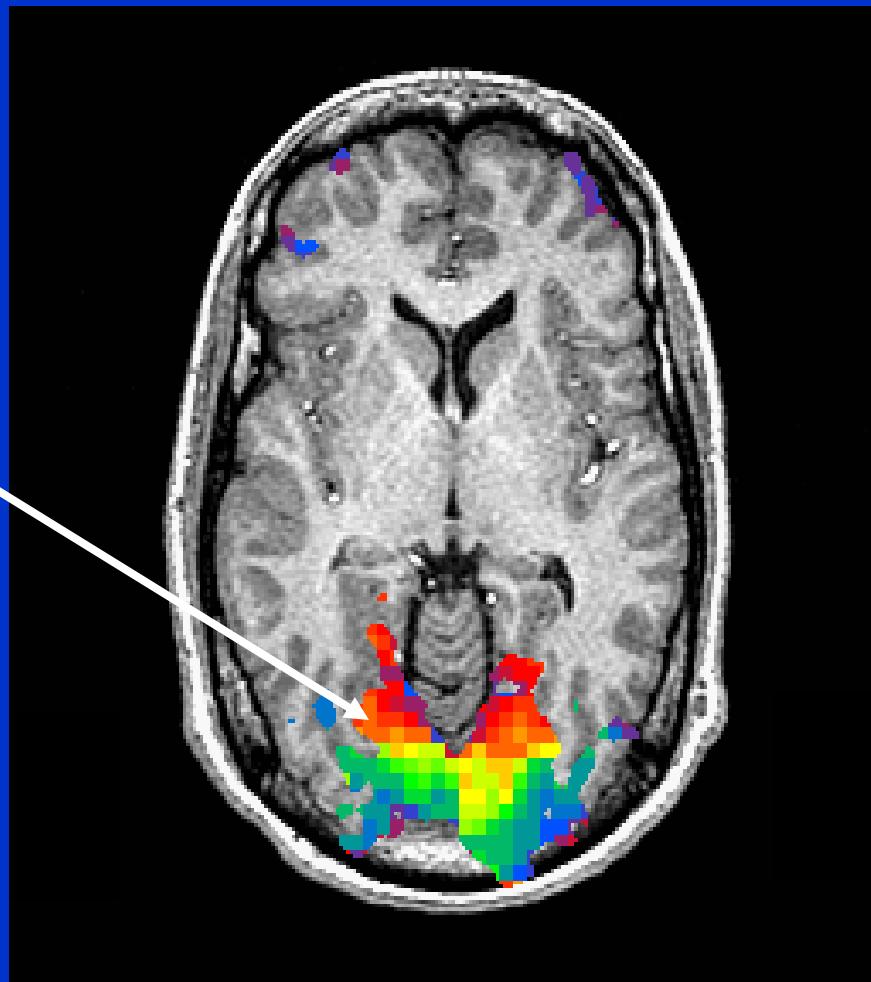


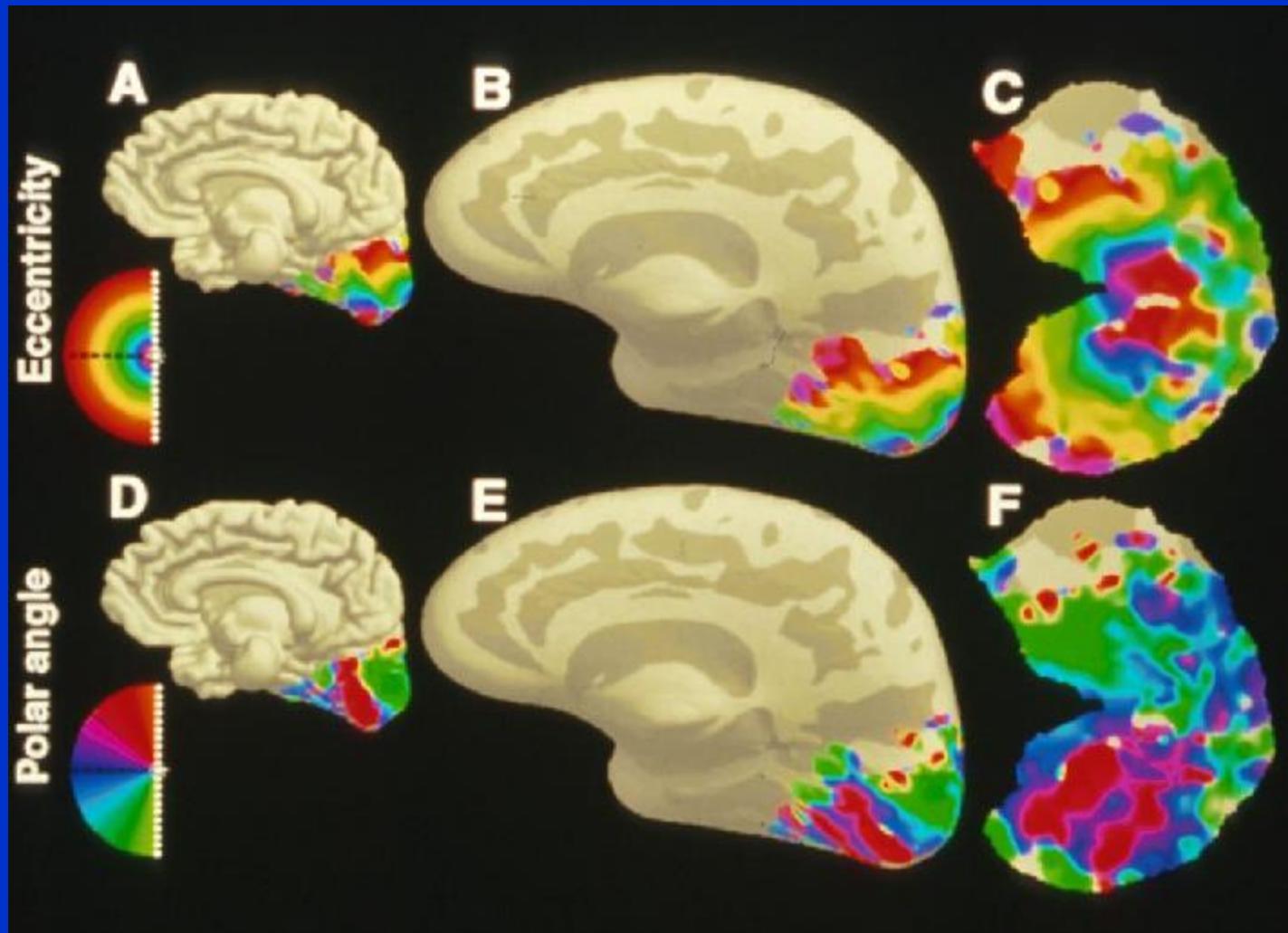
# Listening to Spoken Words



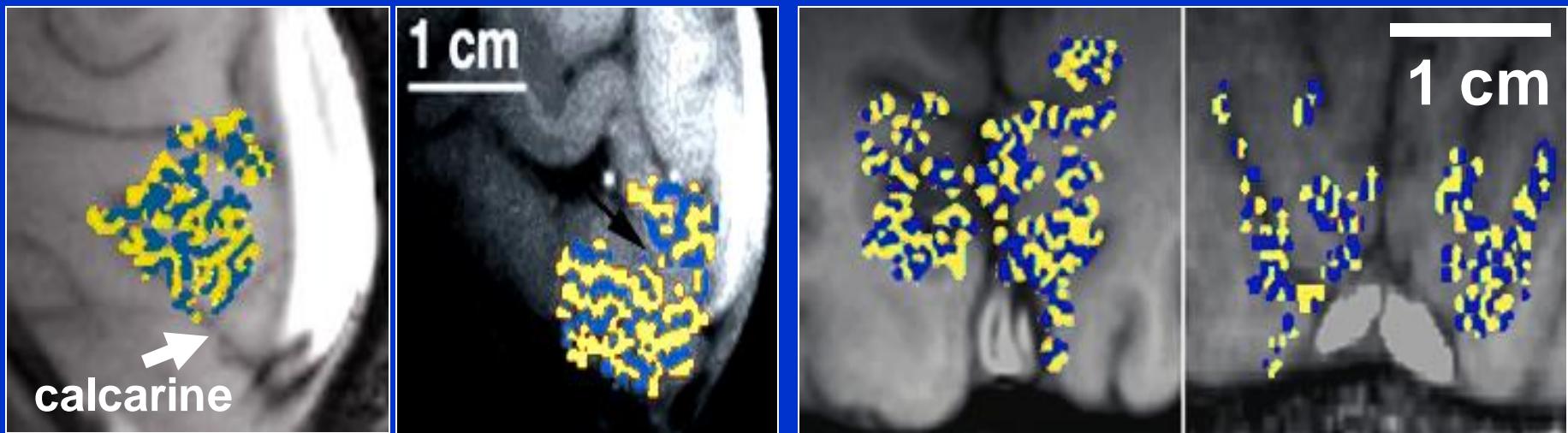
# fMRI Mapping of Visual Cortex

Functional Activation  
(Color codes visual field eccentricity)





# ODC Maps using fMRI



- Identical in size, orientation, and appearance to those obtained by optical imaging<sup>1</sup> and histology<sup>3,4</sup>.

Menon et al.

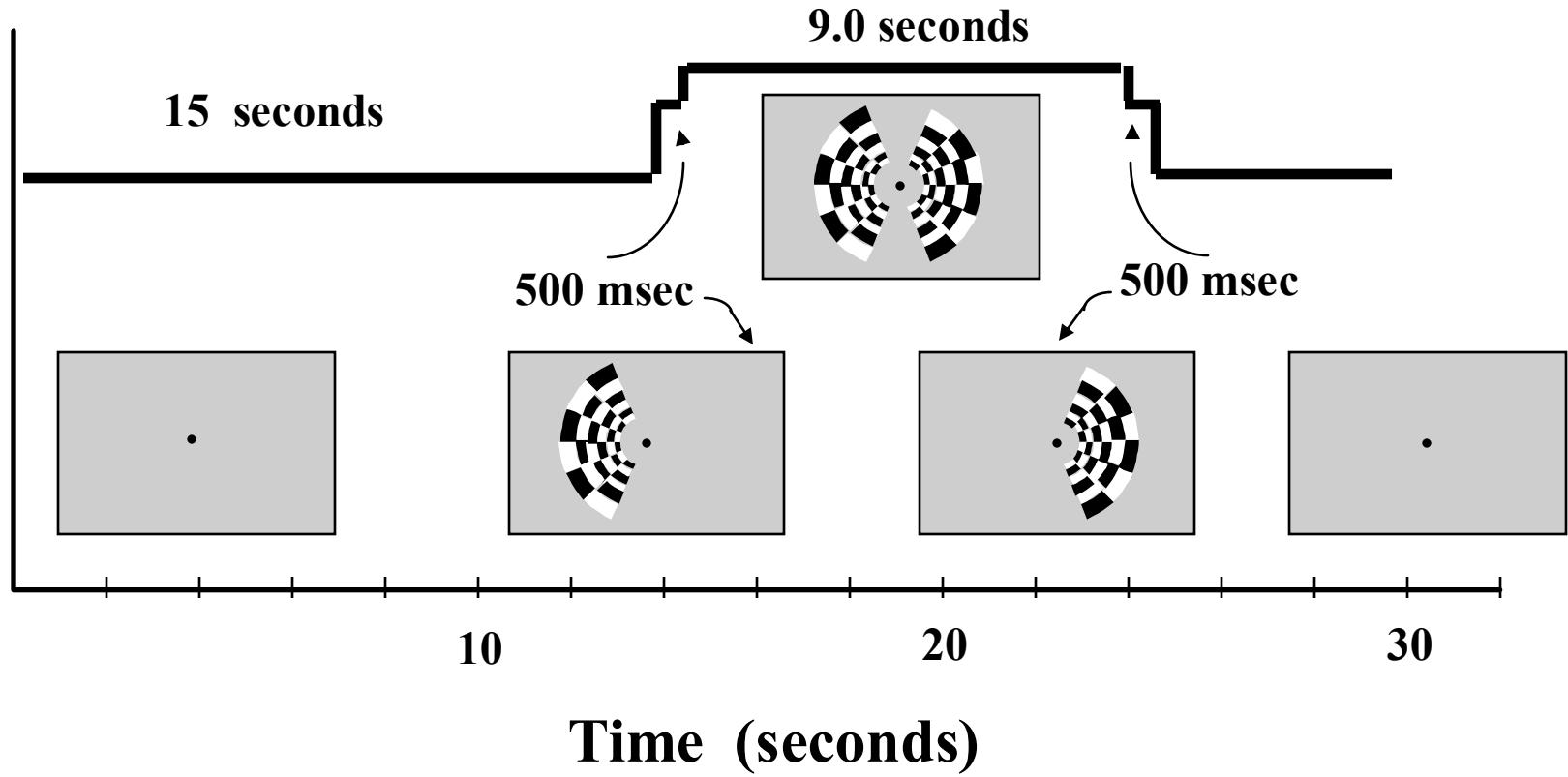
<sup>1</sup>Malonek D, Grinvald A. *Science* 272, 551-4 (1996).

<sup>3</sup>Horton JC, Hocking DR. *J Neurosci* 16, 7228-39 (1996).

<sup>4</sup>Horton JC, et al. *Arch Ophthalmol* 108, 1025-31 (1990).

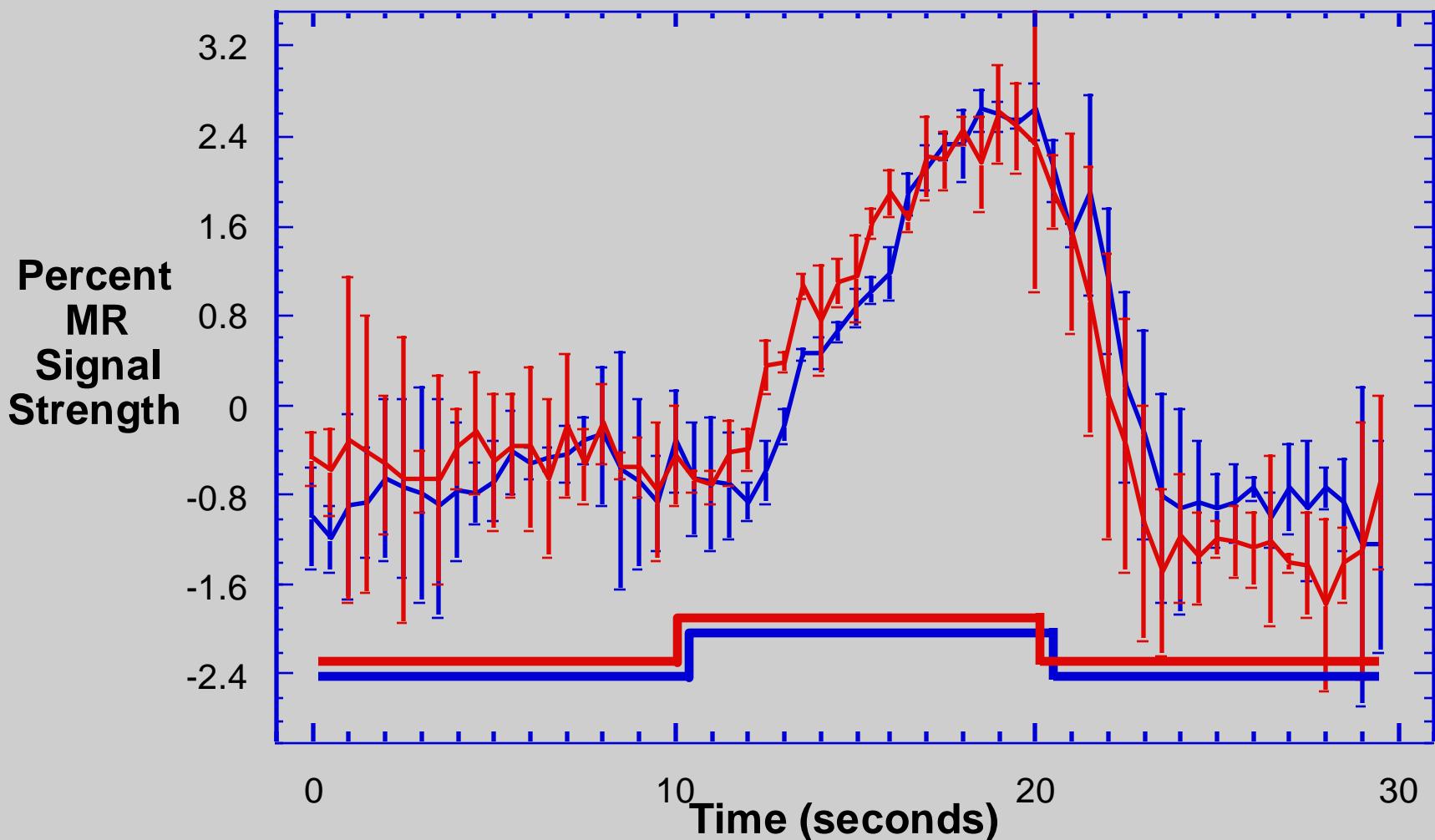
# Regions of Interest Used for Hemi-Field Experiment

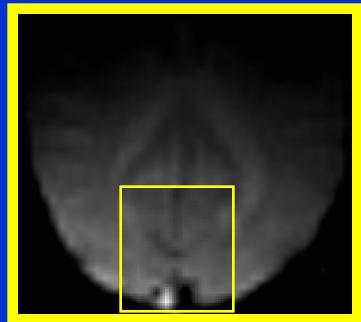




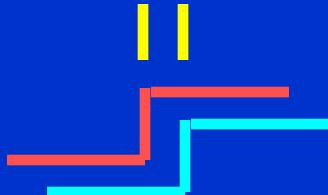
# Hemi-field with 500 msec asynchrony

Average of 6 runs    Standard Deviations Shown

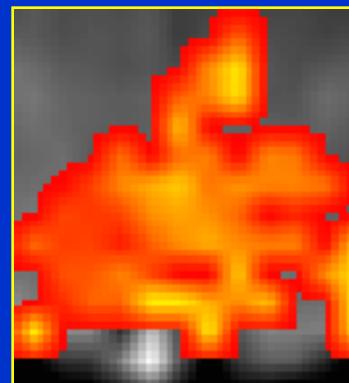




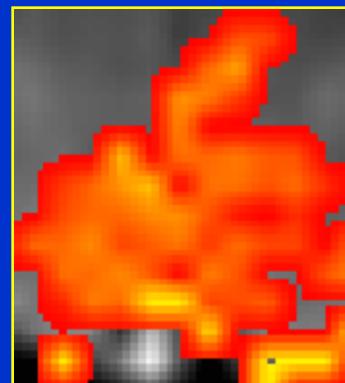
**500 ms**



+ 2.5 s



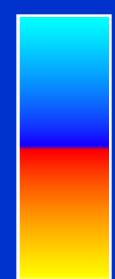
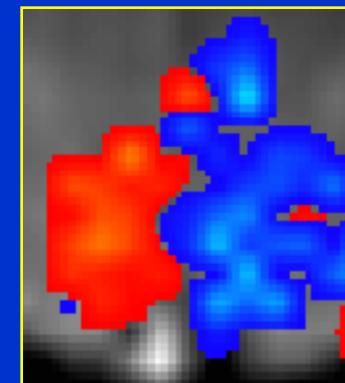
0 s



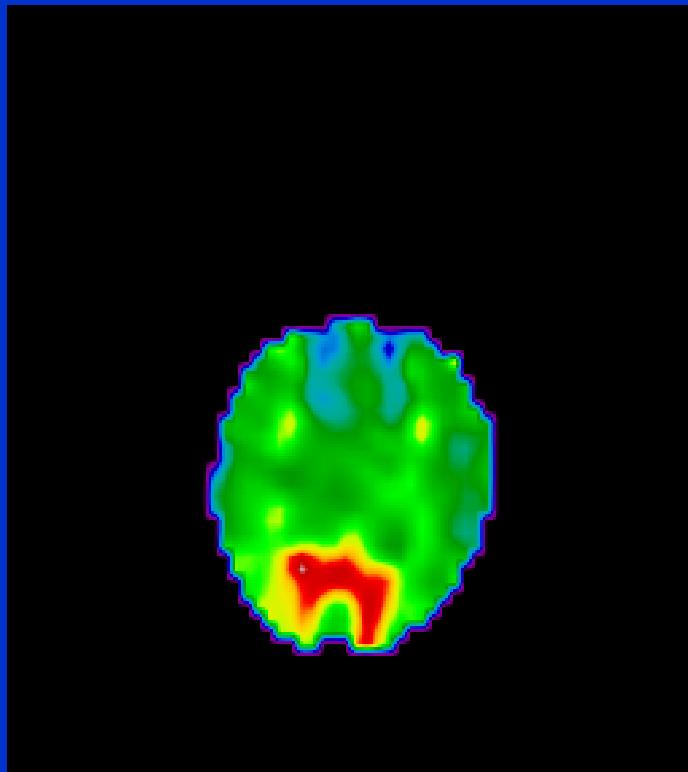
- 2.5 s

# Right Hemifield

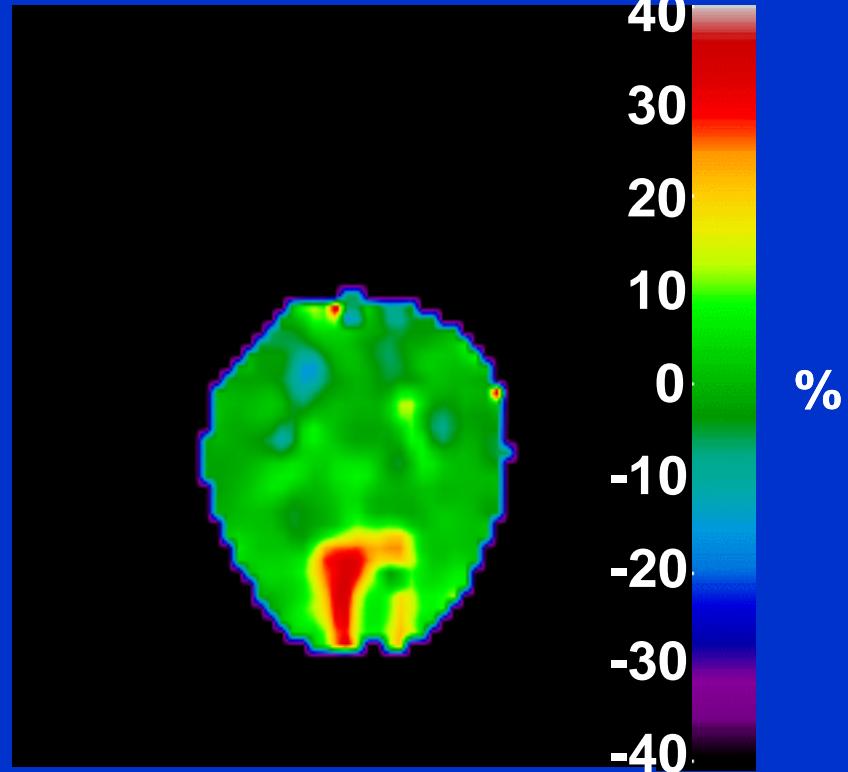
# **Left Hemifield**



# Computed CMRO<sub>2</sub> changes

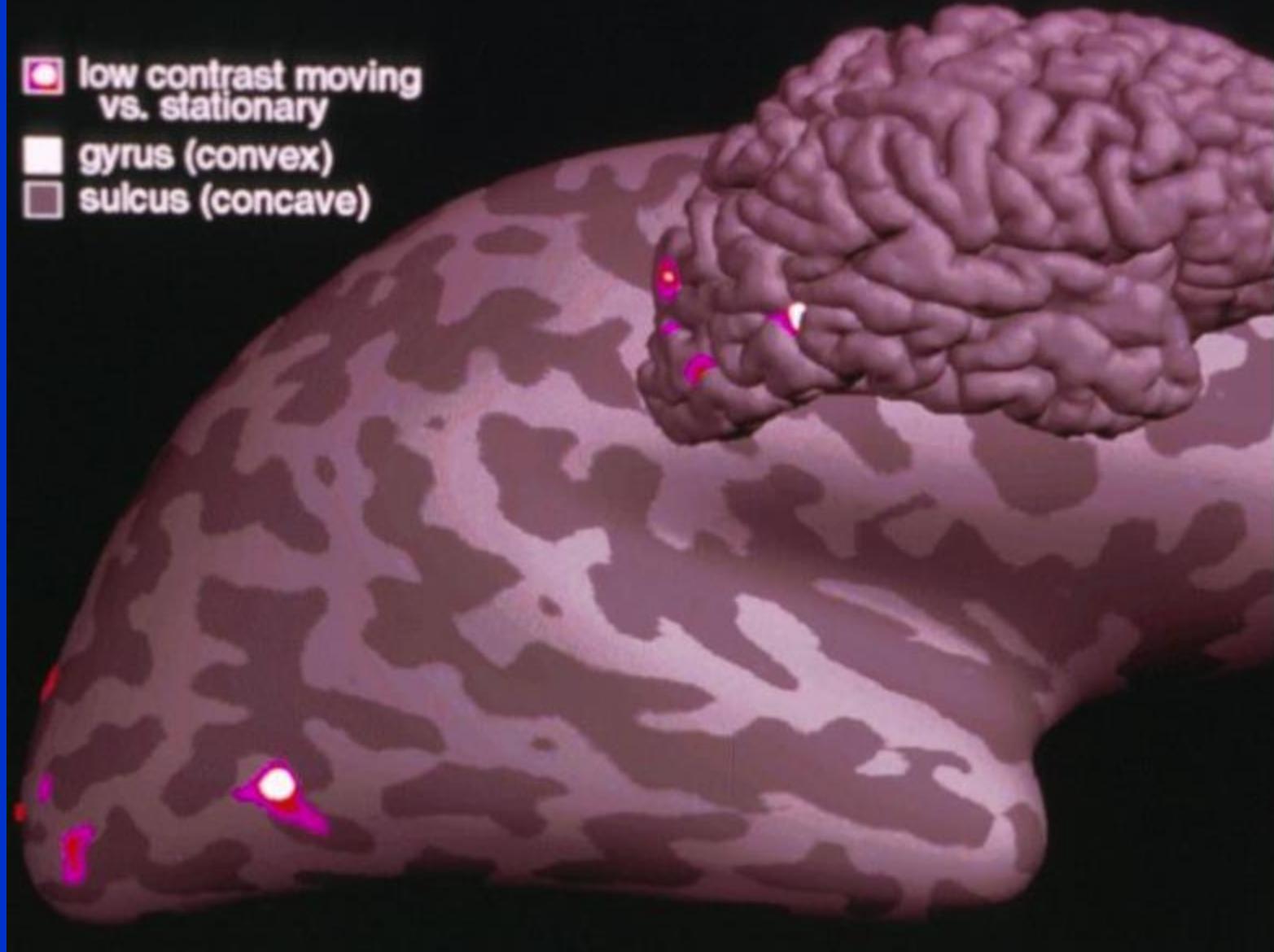


Subject 1

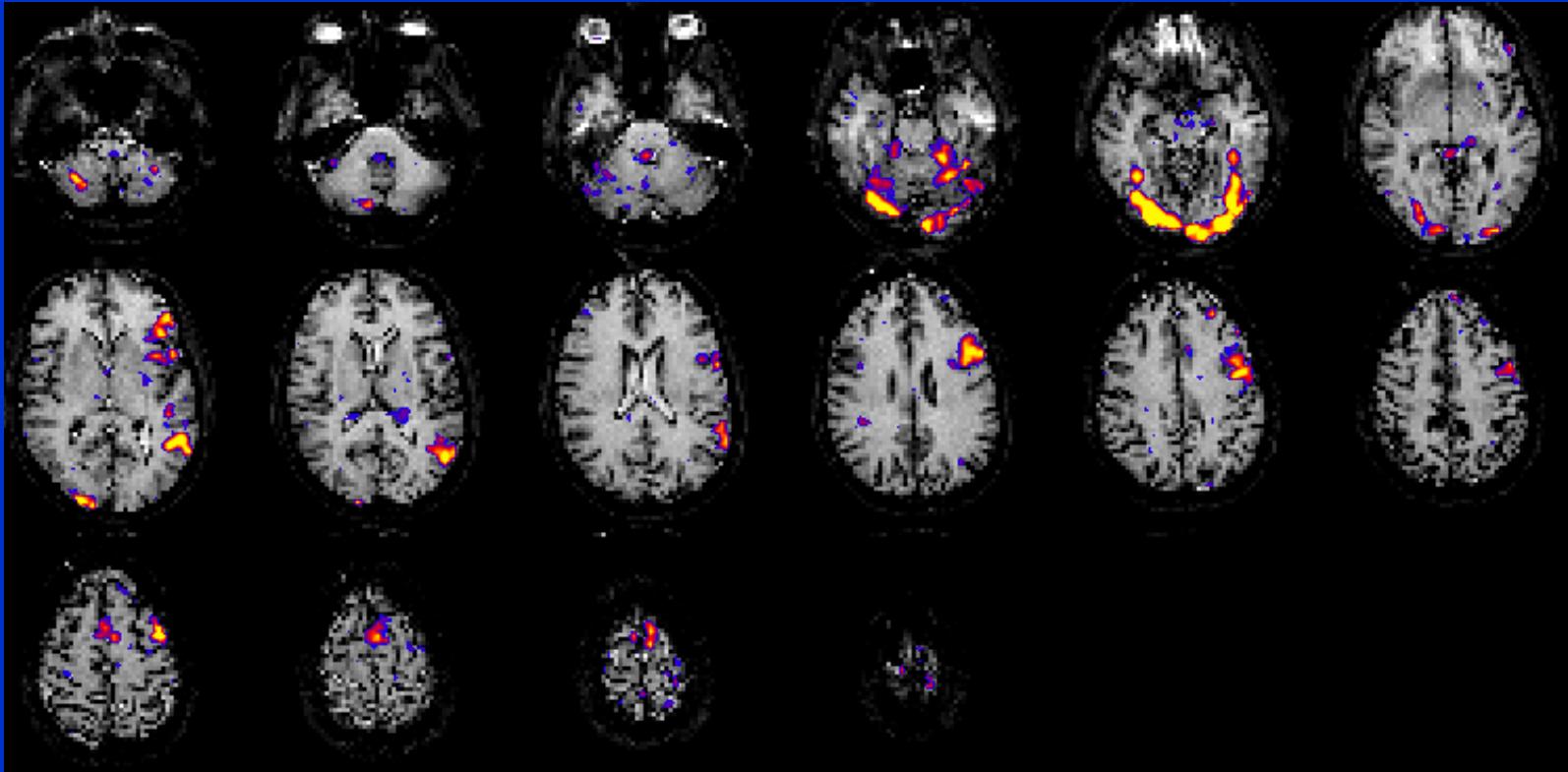


Subject 2

- low contrast moving vs. stationary
- gyrus (convex)
- sulcus (concave)

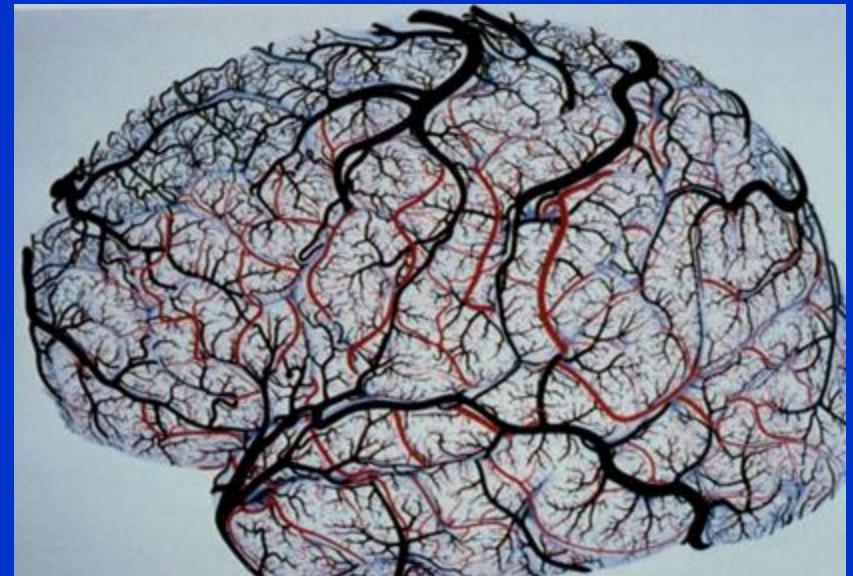
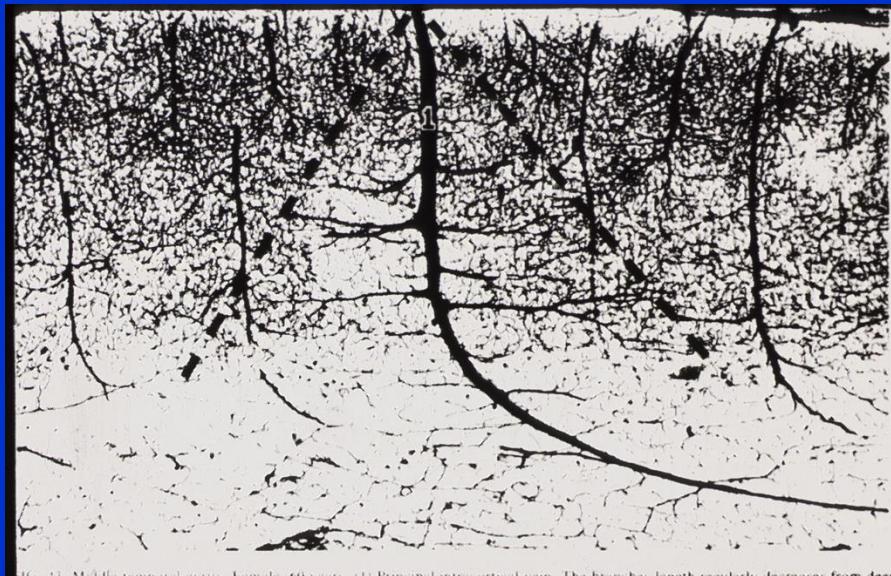


# Word stem completion



# A Primary Challenge:

**...to make progressively more precise inferences using fMRI without making too many assumptions about non-neuronal physiologic factors.**



**Anatomy**



**BOLD**



**Perfusion**



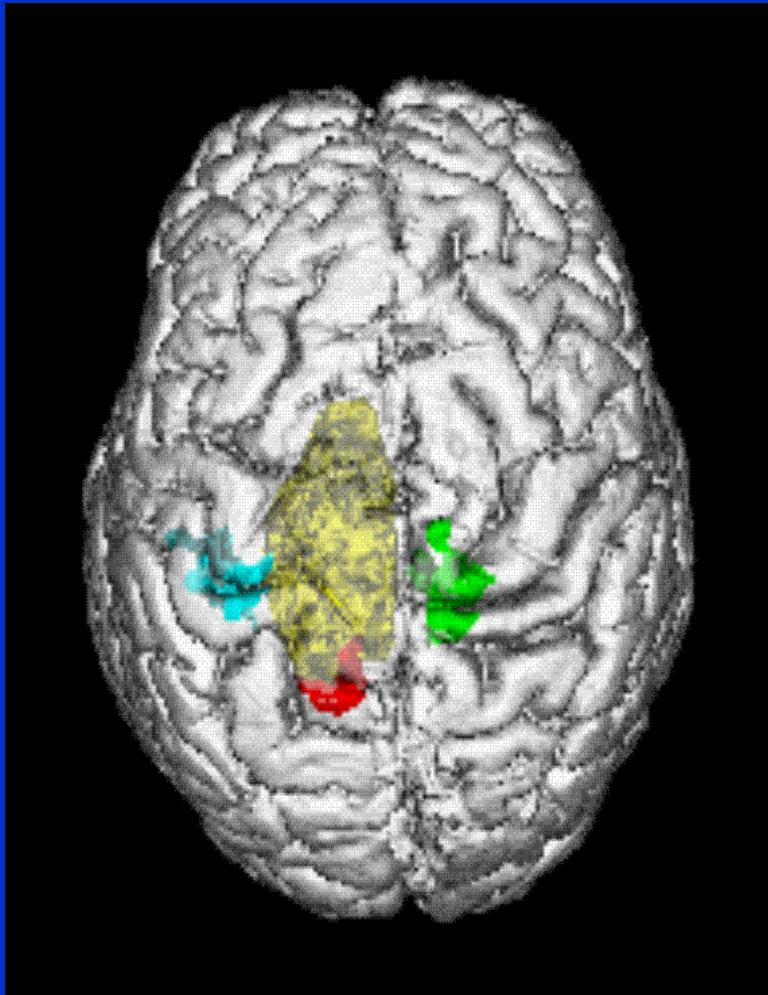
# Presurgical Mapping

Left Foot

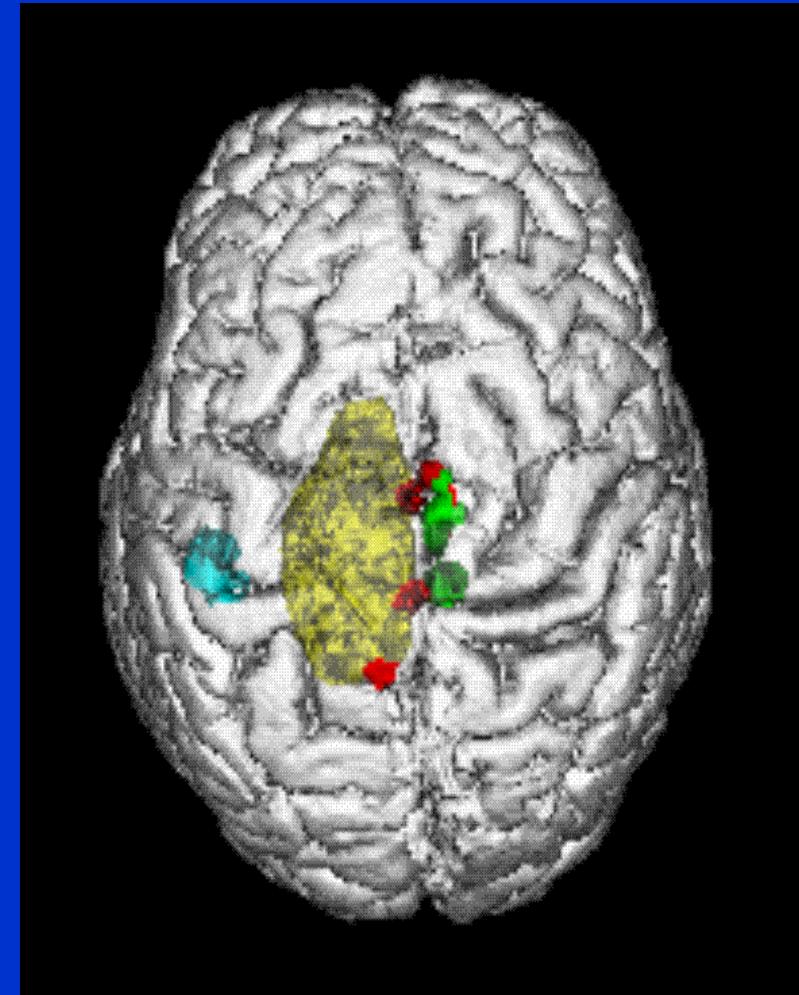
Tumor

Right Foot

Right Hand

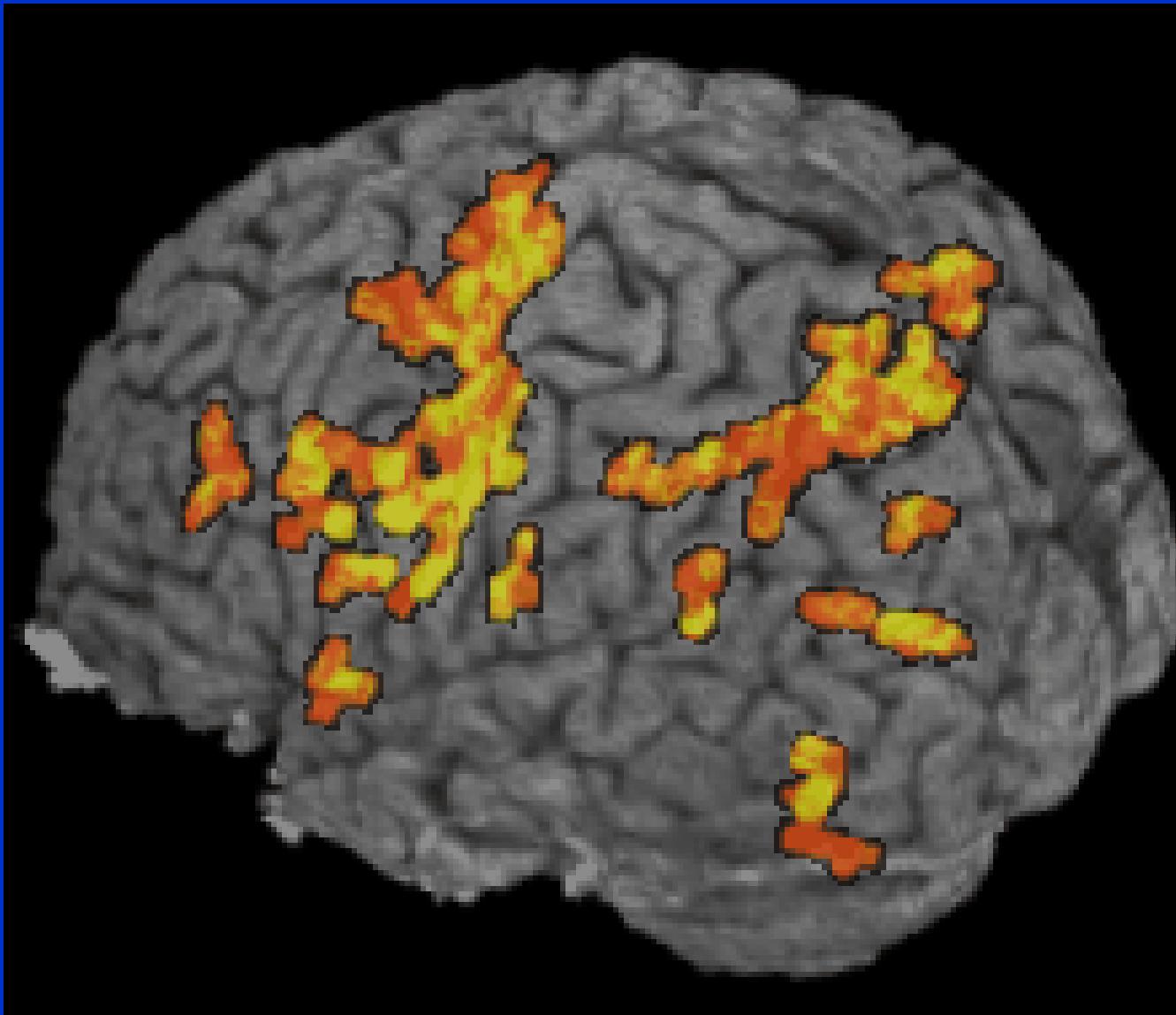


fMRI



O-15 PET

# End of Acquisition



< 1 s to render

Blocked trials:  
20 s on/20 s off  
8 blocks

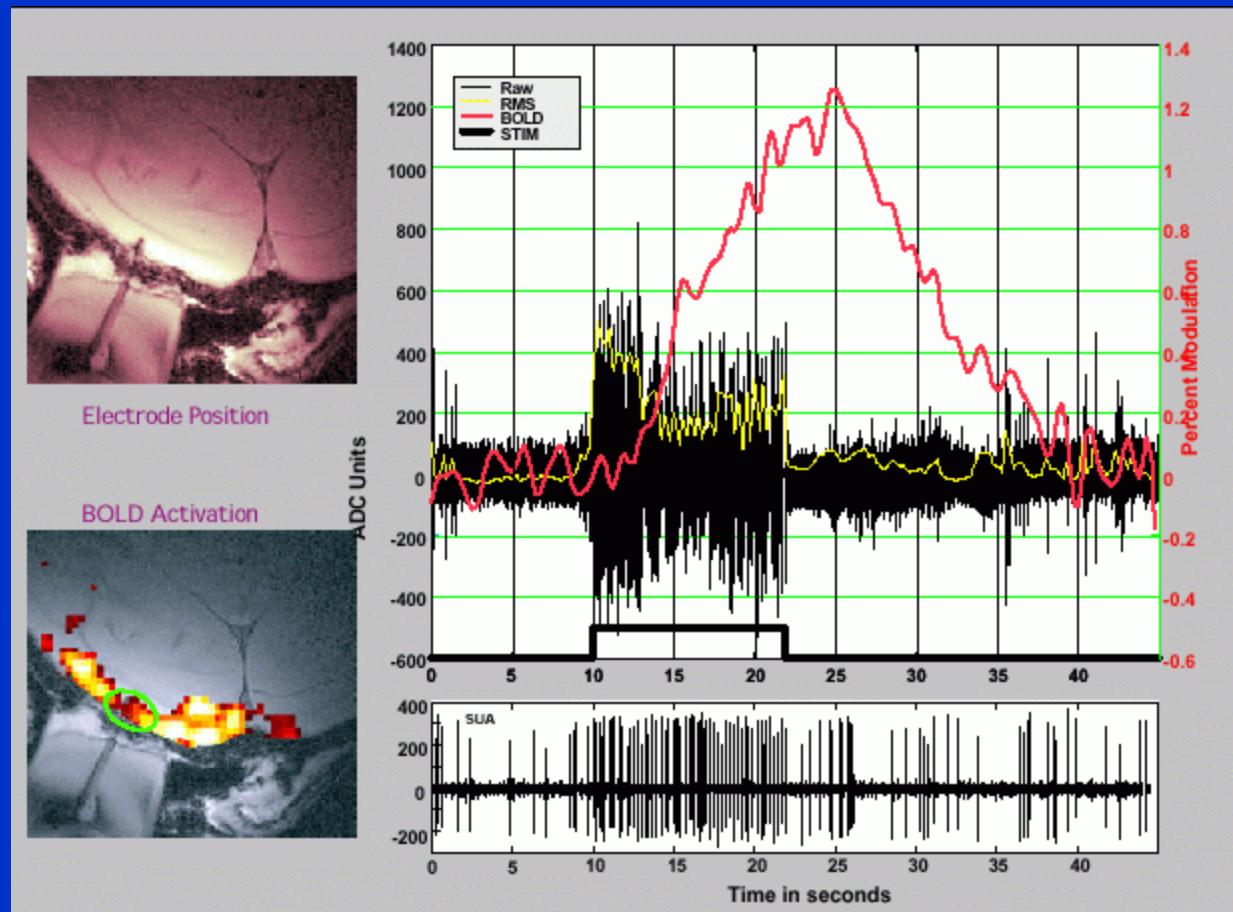
Blocks: 12345678

Color shows  
through brain

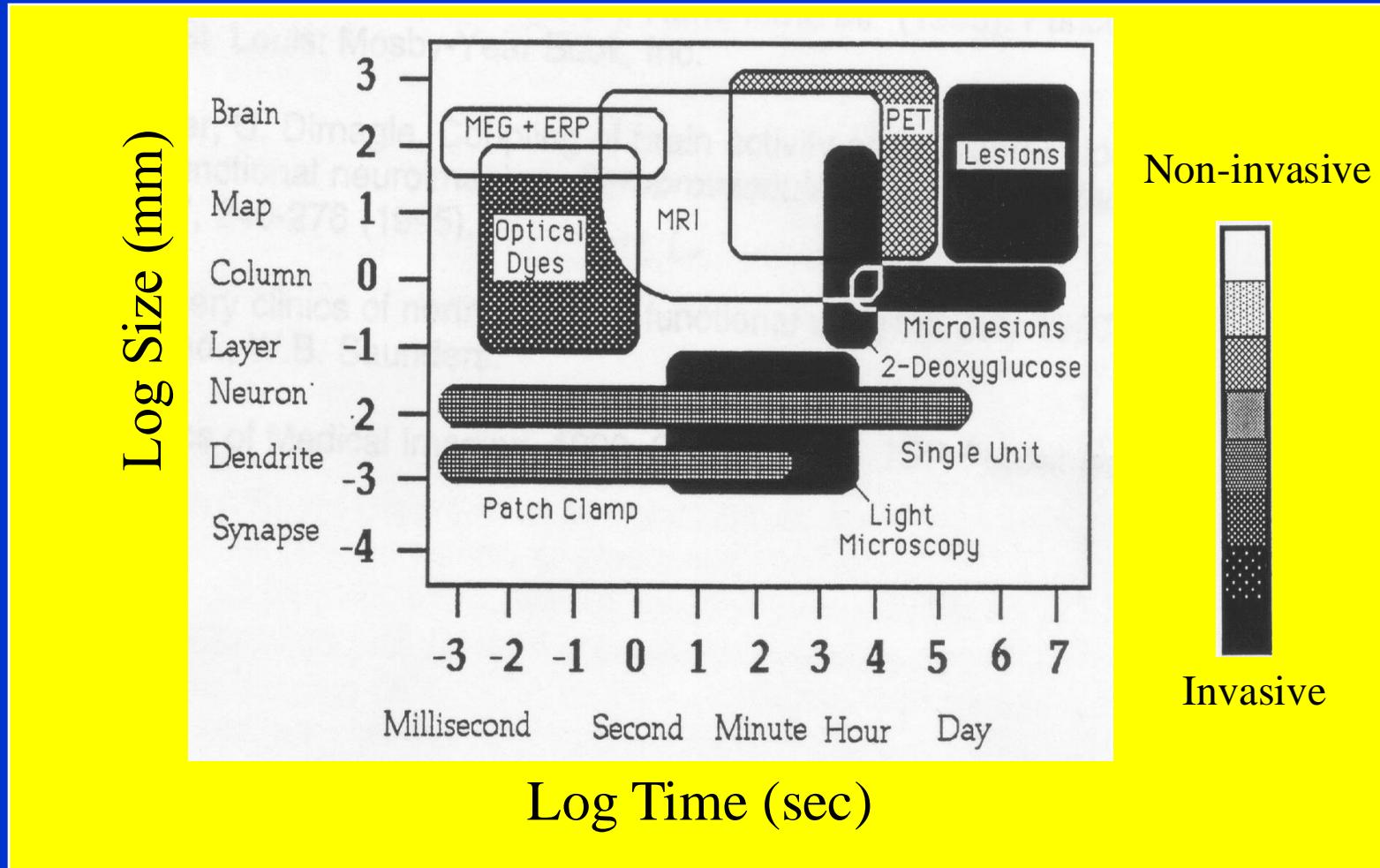
Correlation > 0.45



# Combined Electrophysiological Measurement and fMRI



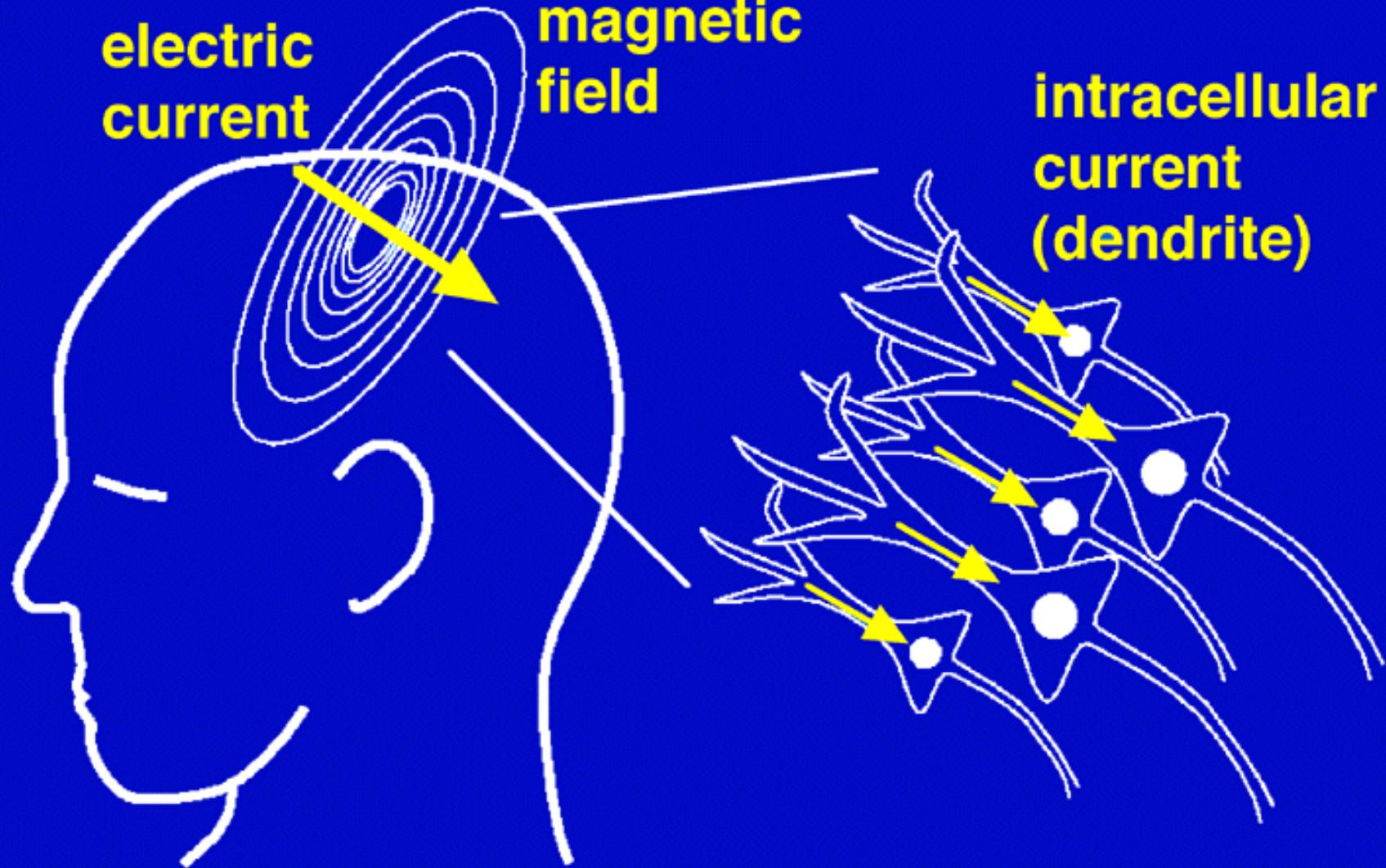
# Functional Neuroimaging Techniques



**electric  
current**

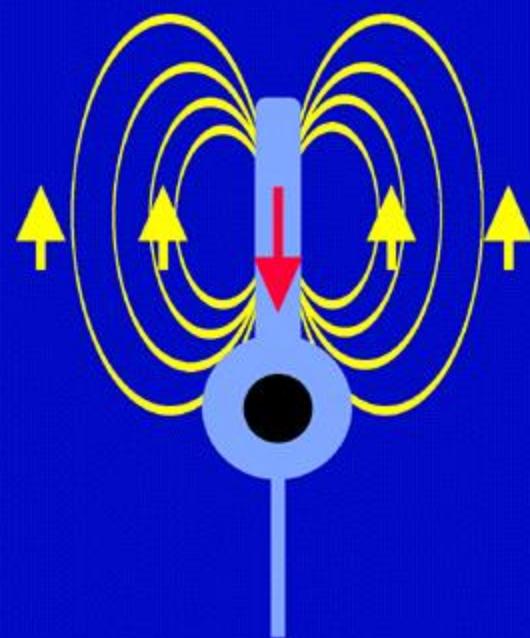
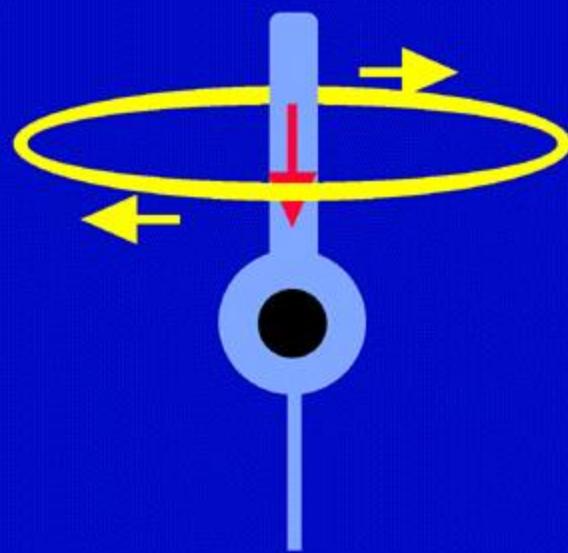
**magnetic  
field**

**intracellular  
current  
(dendrite)**

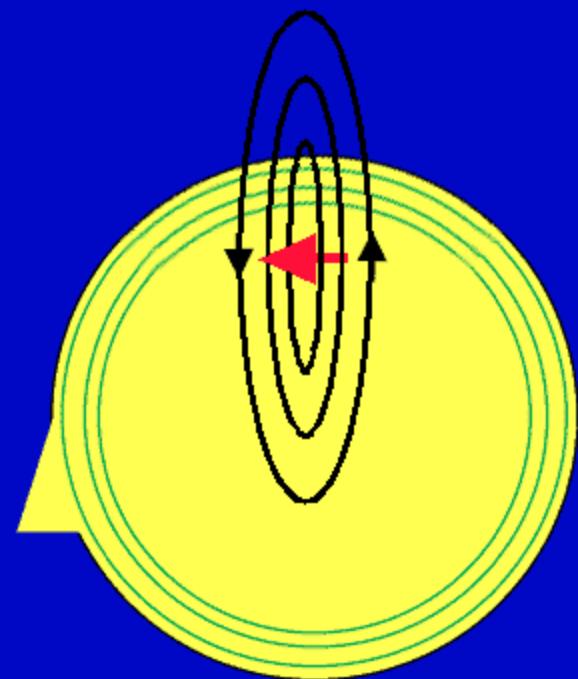


**MEG:**  
intracellular  
current

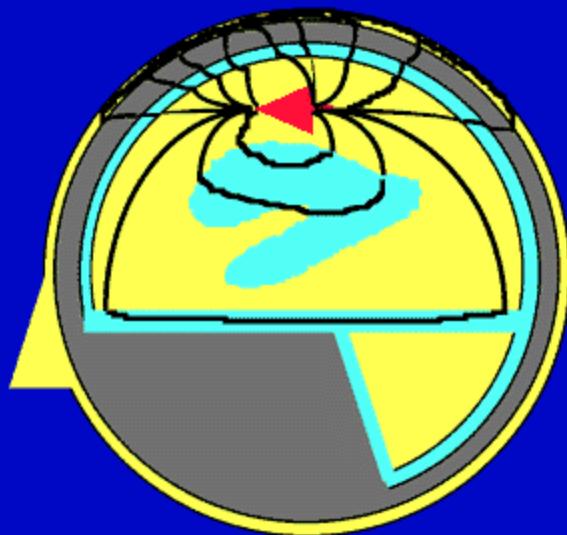
**EEG:**  
extracellular  
current

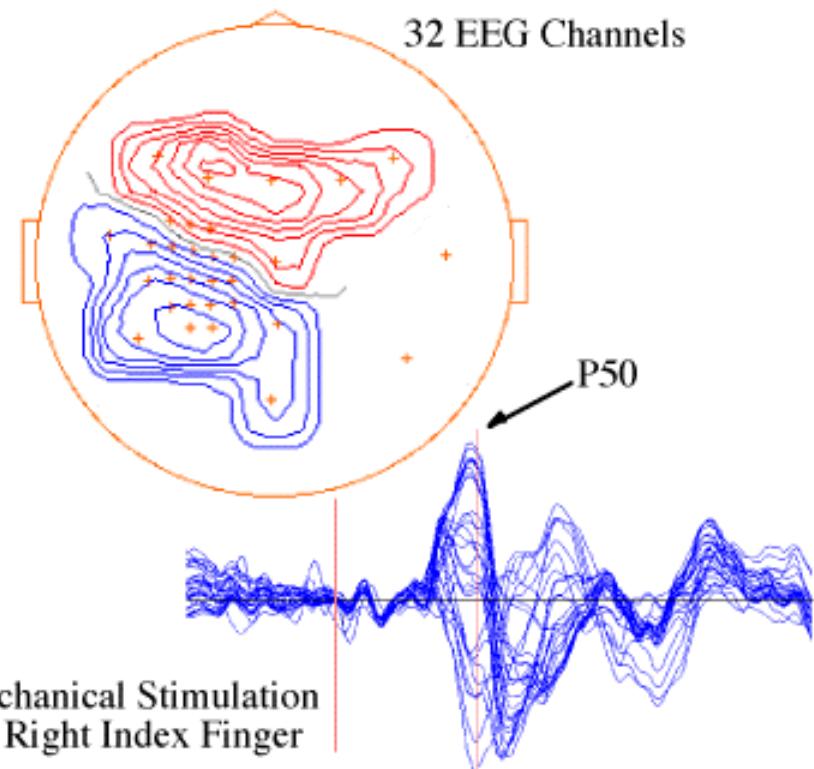
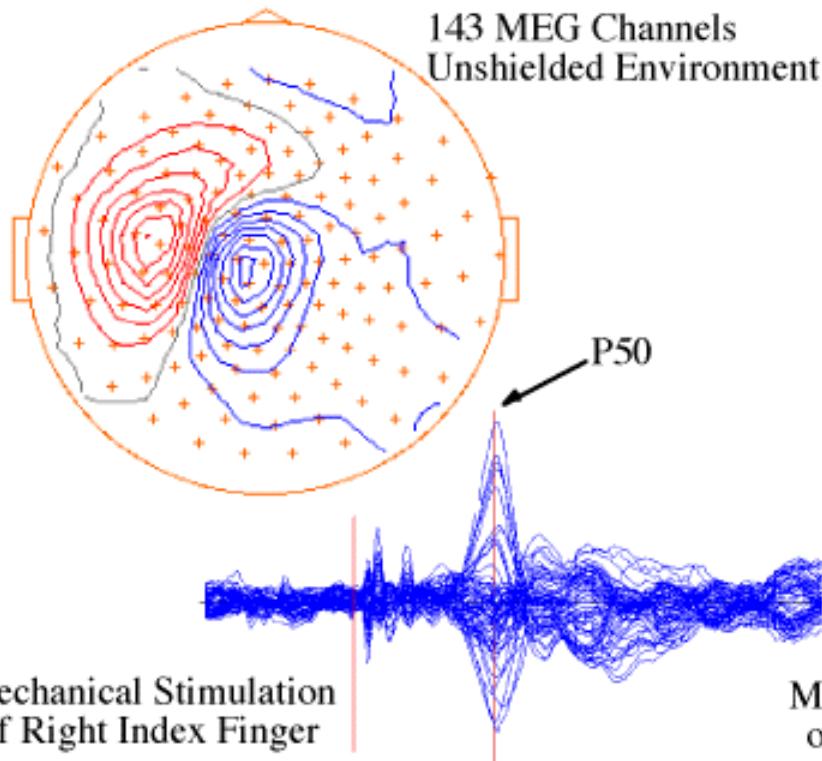


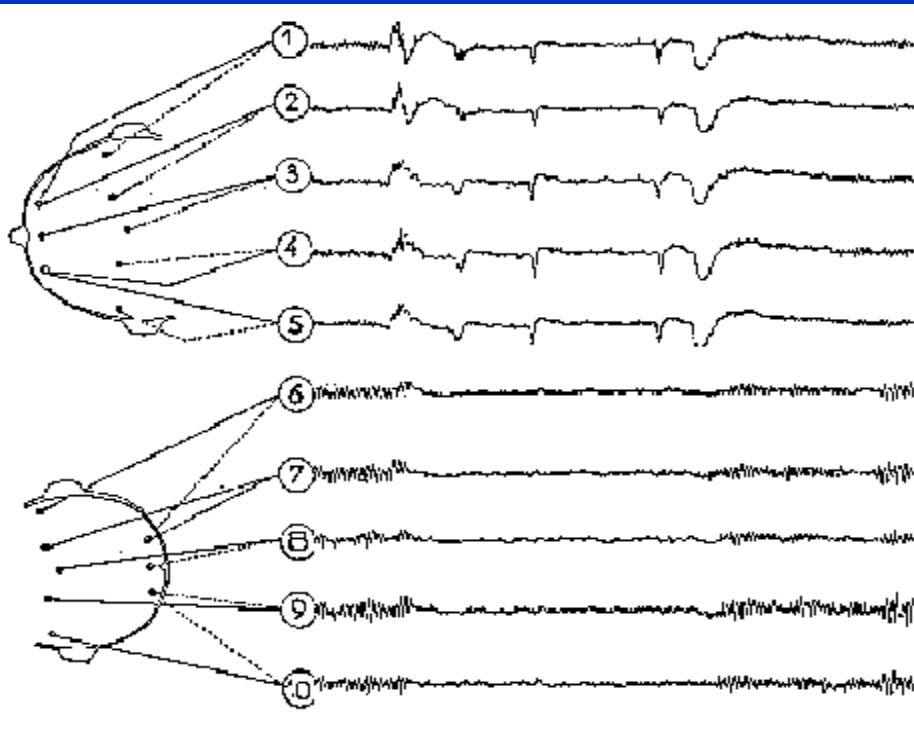
**MEG**



**EEG**



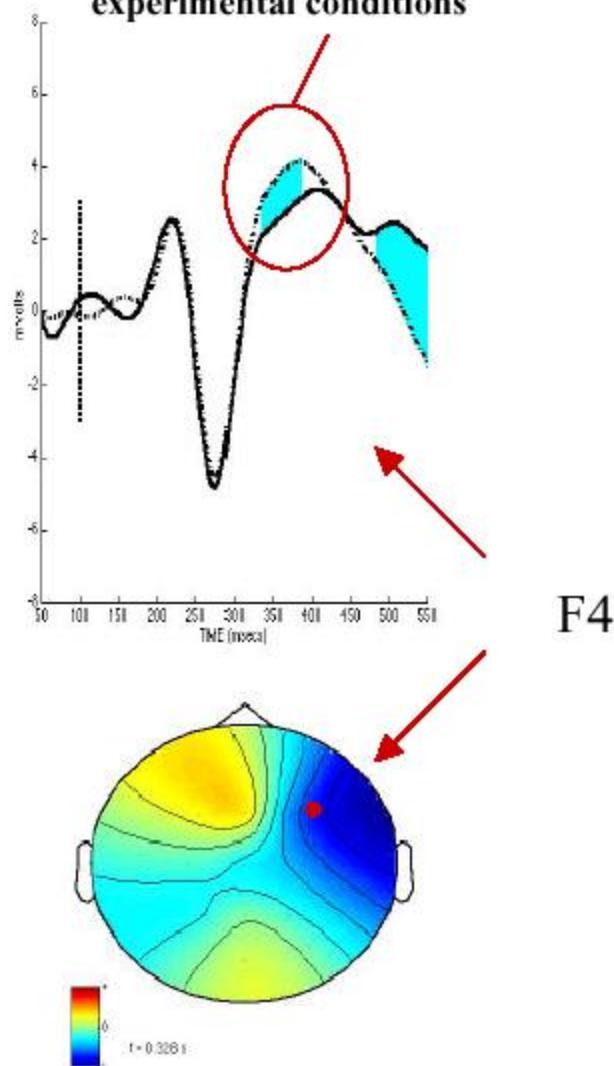




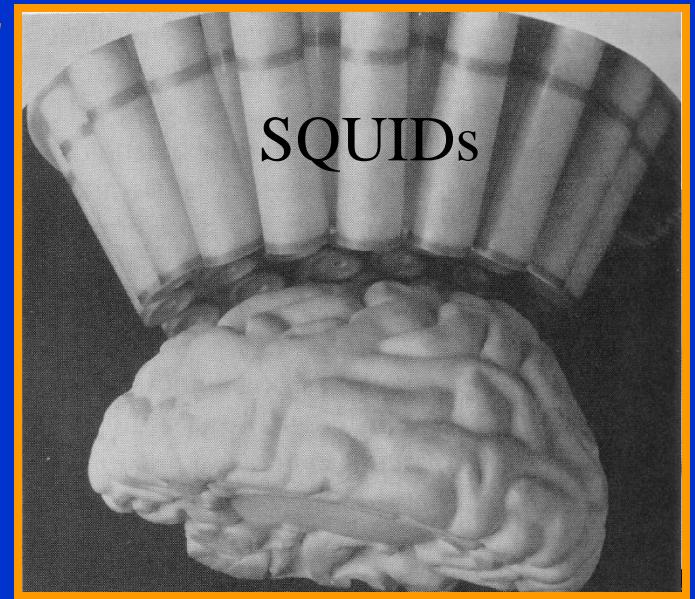
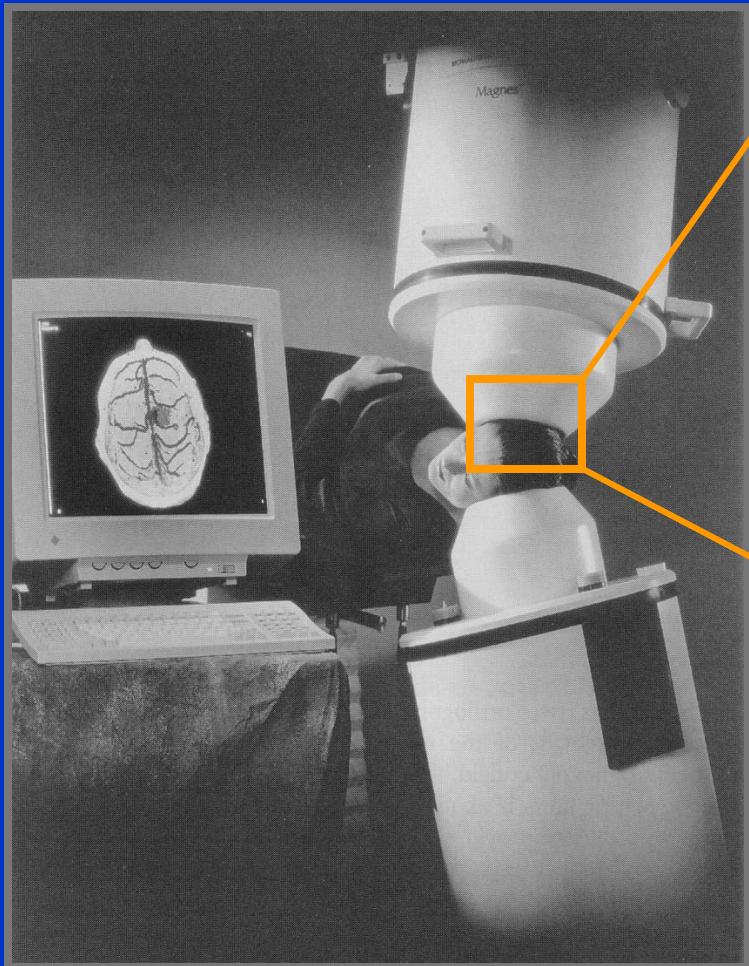
## Electroencephalography (EEG) recording



Statistical (T test) between two experimental conditions



# Magnetoencephalography (MEG)

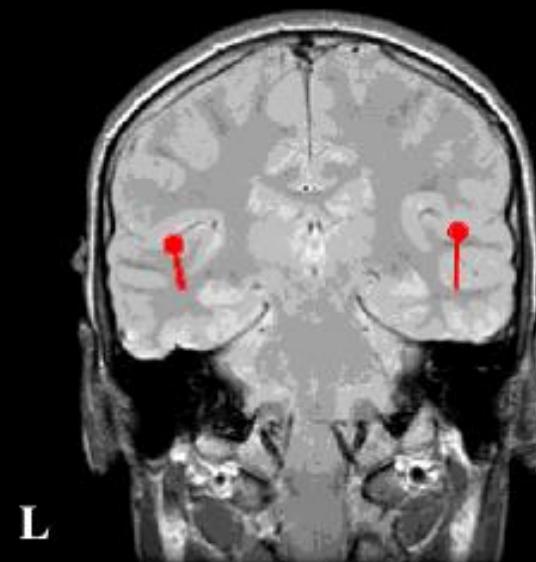
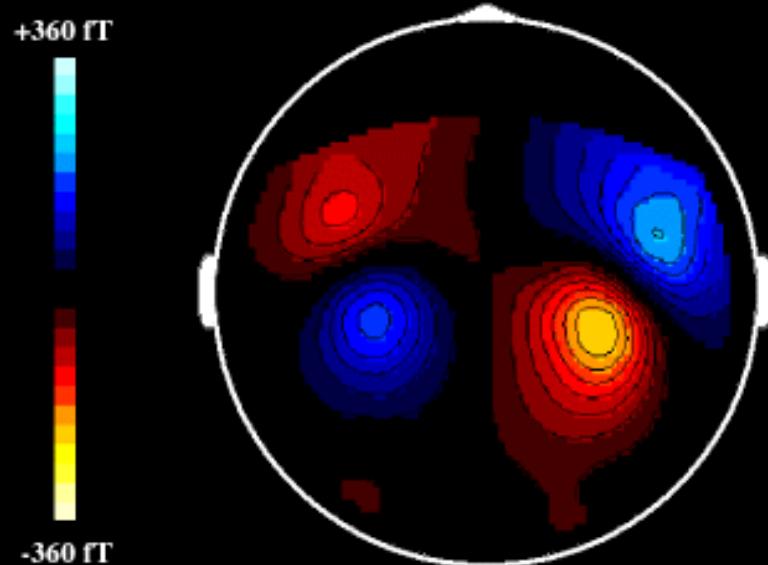
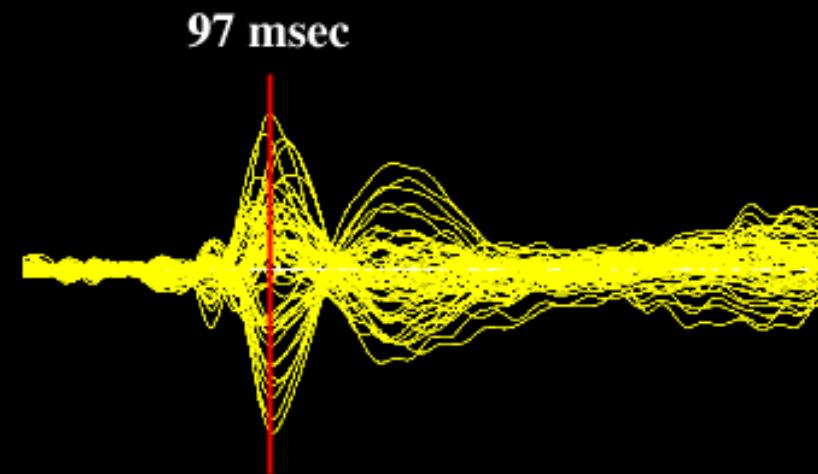
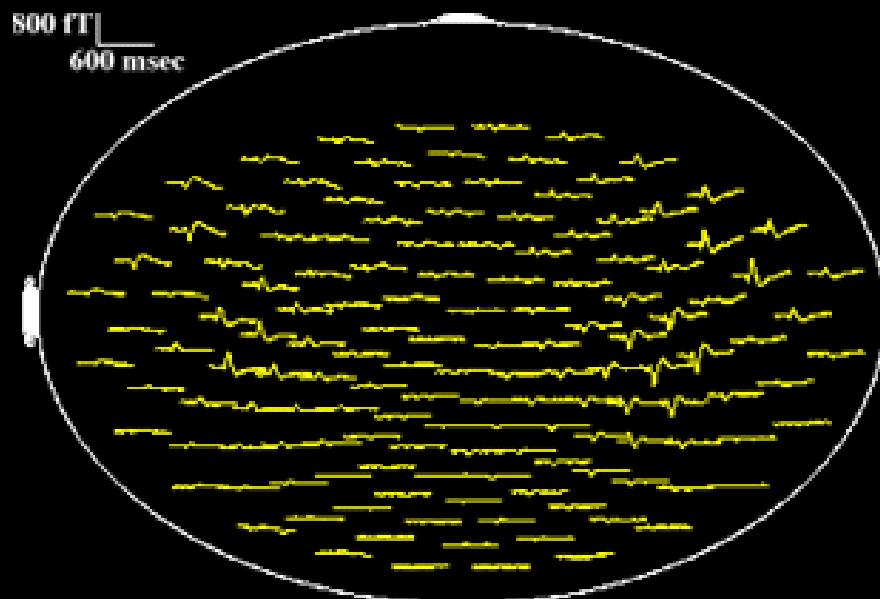


SQUID:  
Superconducting Quantum  
Interference Device



Combined MEG and EEG

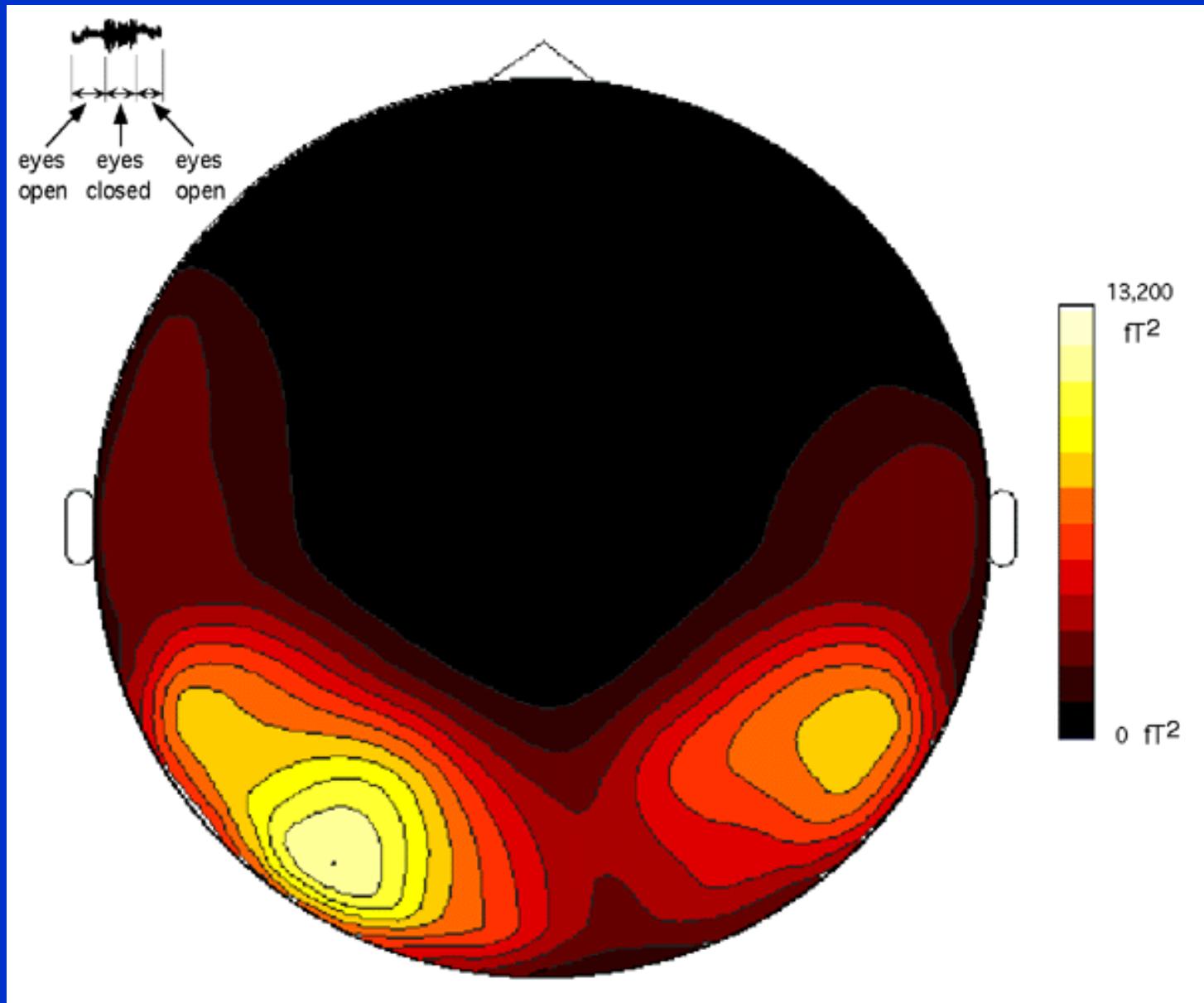
# MEG Mapping



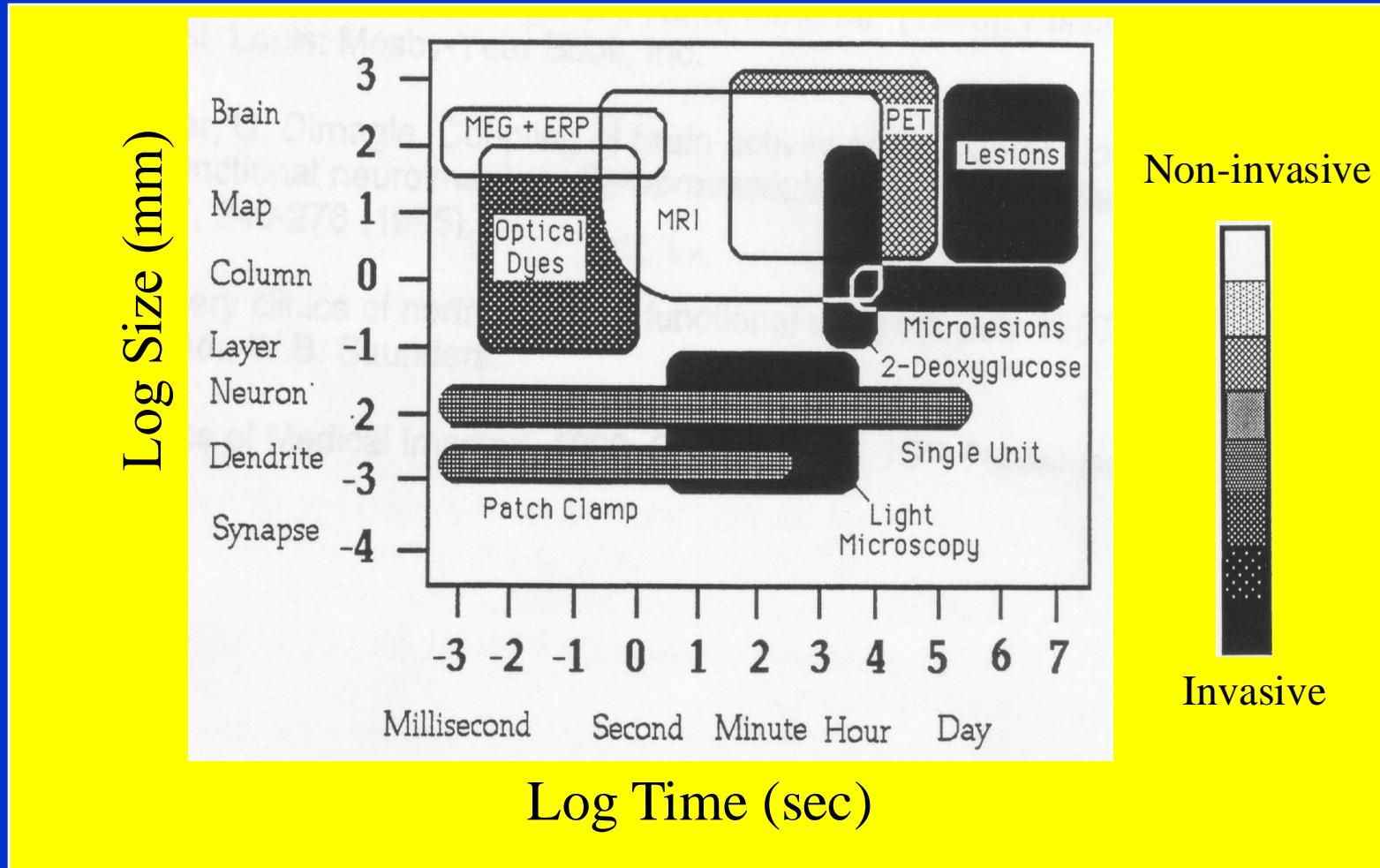
R

L

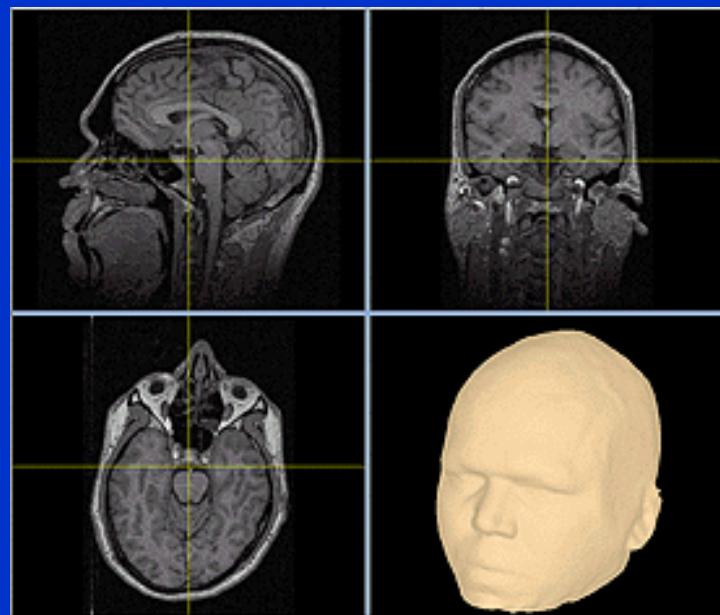
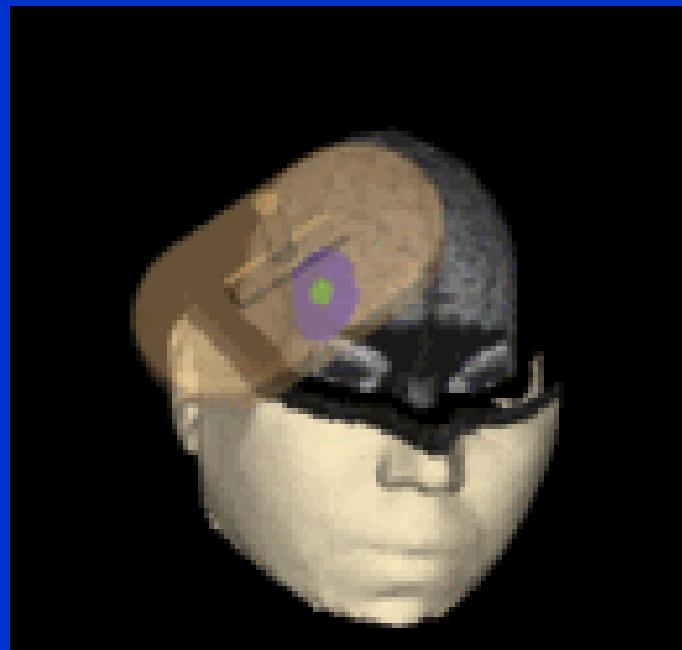
# Alpha Wave Activity Mapped with MEG



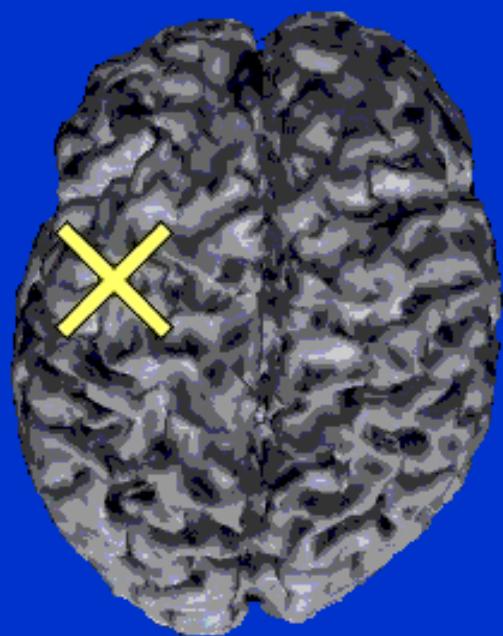
# Functional Neuroimaging Techniques



# Transcranial Magnetic Stimulation



# Transcranial Magnetic Stimulation (TMS)



# Acknowledgements

Ted Deyoe, Medical College of Wisconsin  
Kathleen Schmainda, Medical College of Wisconsin  
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Richard Coppola, National Institute of Mental Health  
Sosumu Mori, Johns Hopkins University  
Robert Cox, National Institute of Mental Health  
Ziad Saad, National Institute of Mental Health  
Eric Wong, University of California, San Diego  
Ravi Menon, University of Western Ontario  
Nikos Logothetis, Max Plank Institute, Germany

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<b>Jerzy Bodurka</b>	<b>August Tuan</b>	<b>Carla Wettig</b>
<b>Frank Ye</b>	<b>Dan Kelley</b>	<b>Kang-Xing Jin</b>
<b>Wen-Ming Luh</b>	<b>Visiting Fellows:</b>	<b>Program Assistant:</b>
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<b>Adam Thomas</b>	<b>Marta Maierov</b>	<b>Scanning Technologists:</b>
<b>Post Docs:</b>	<b>Guosheng Ding</b>	<b>Karen Bove-Bettis</b>
<b>Rasmus Birn</b>	<b>Clinical Fellow:</b>	<b>Paula Rowser</b>
<b>Hauke Heekeren</b>	<b>James Patterson</b>	
<b>David Knight</b>	<b>Psychologist:</b>	
<b>Patrick Bellgowan</b>	<b>Julie Frost</b>	
<b>Ziad Saad</b>		