

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

Peter A. Bandettini, Ph.D.

Section on Functional Imaging Methods

<http://fim.nimh.nih.gov>

Laboratory of Brain and Cognition

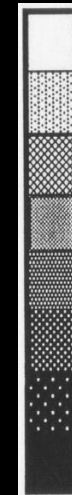
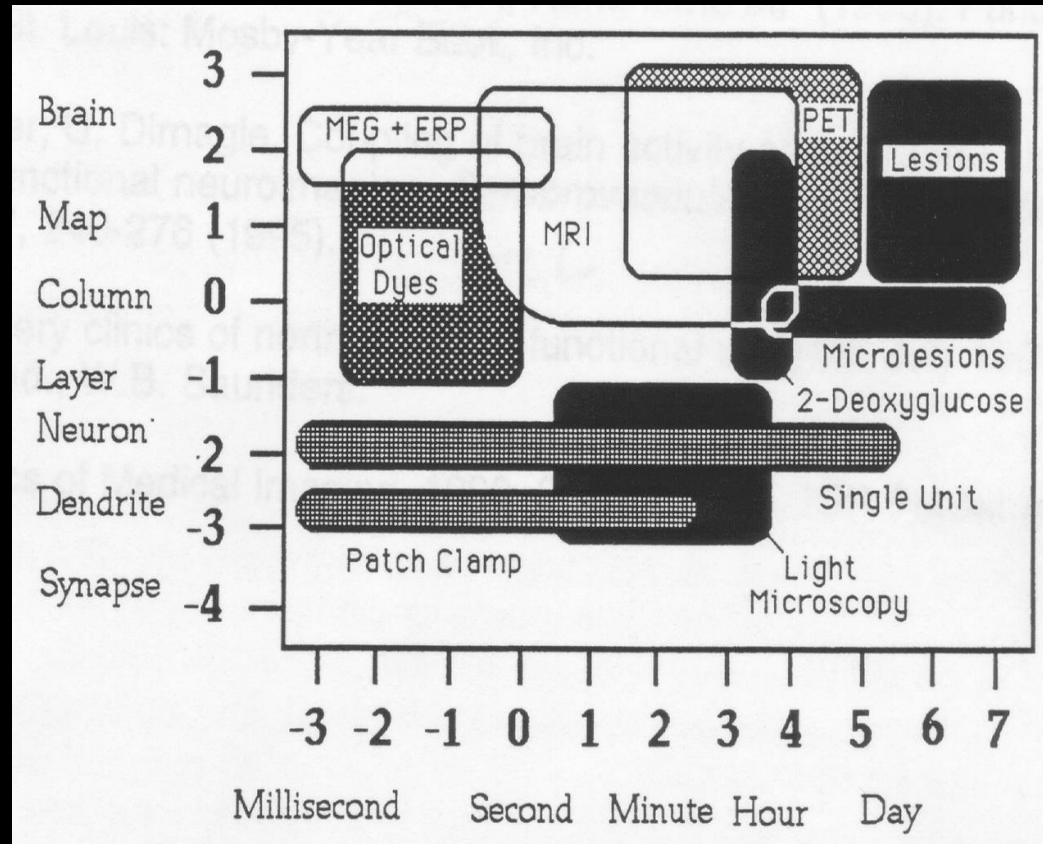
&

Functional MRI Facility

<http://fmrif.nimh.nih.gov>



# Functional Neuroimaging Techniques





**1991**

A brief overview of the three main types of fMRI contrast

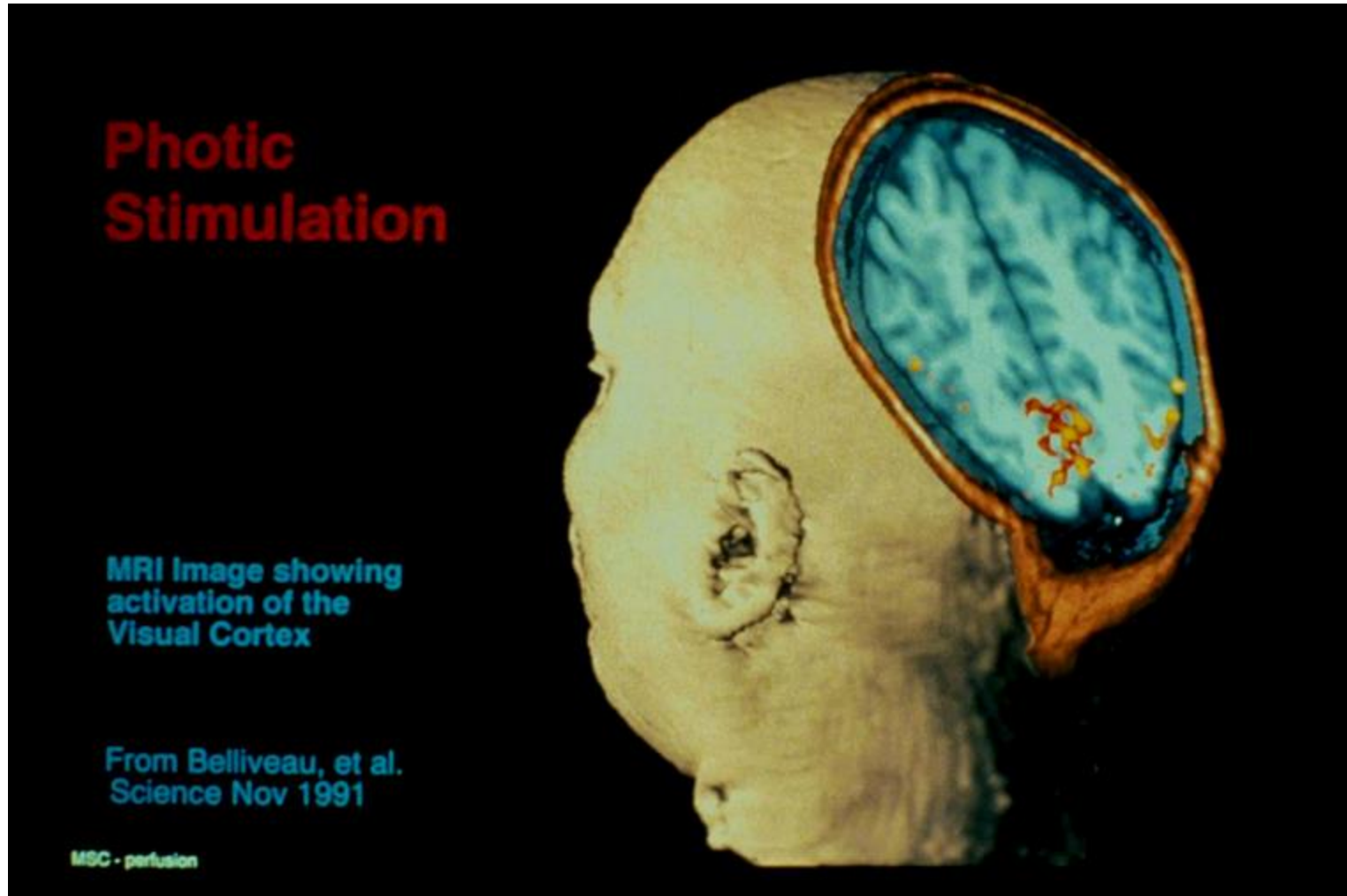
Volume

Flow or Perfusion

Oxygenation

# Blood Volume

What started it all...

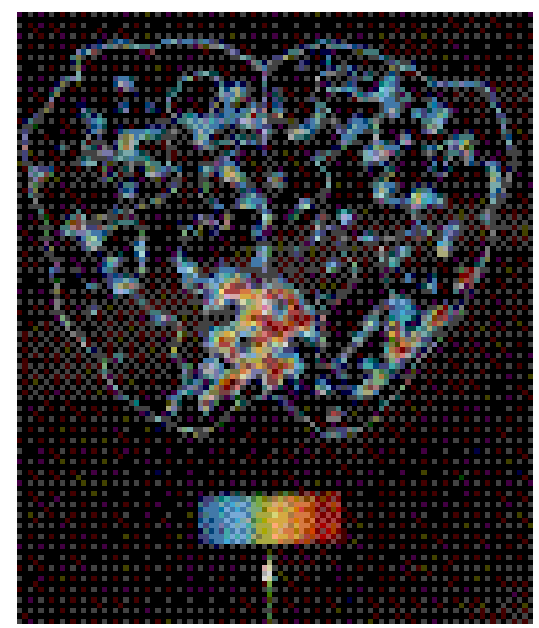
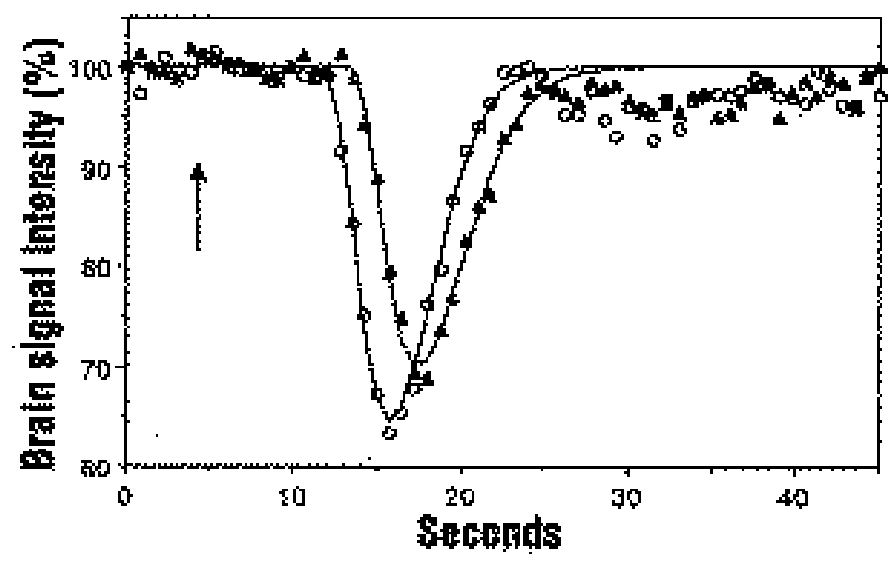
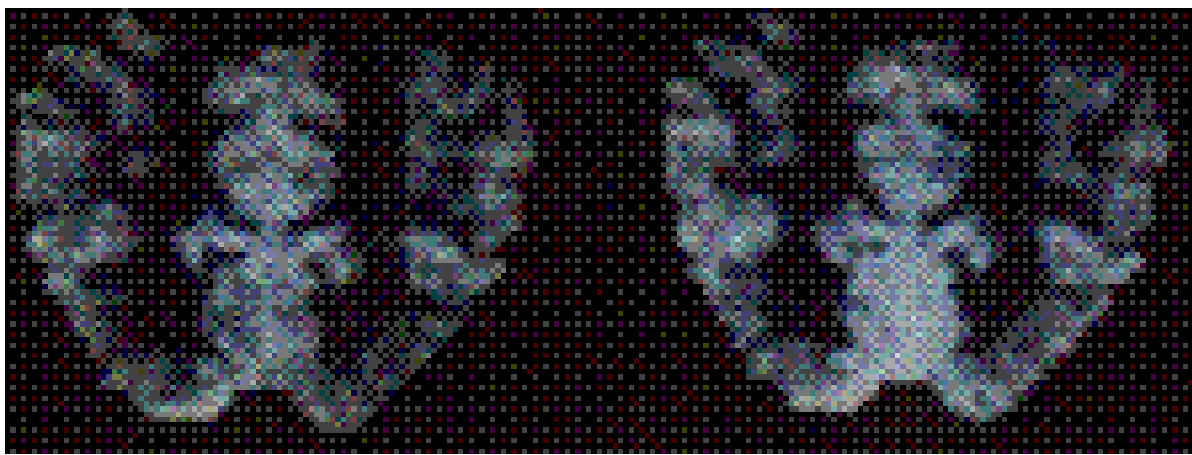




# Blood Volume

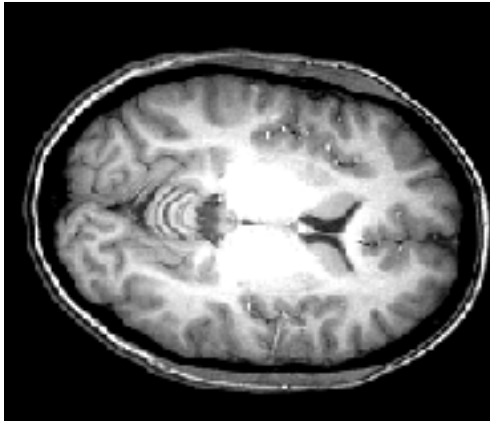
Resting

Active



# MRI vs. fMRI

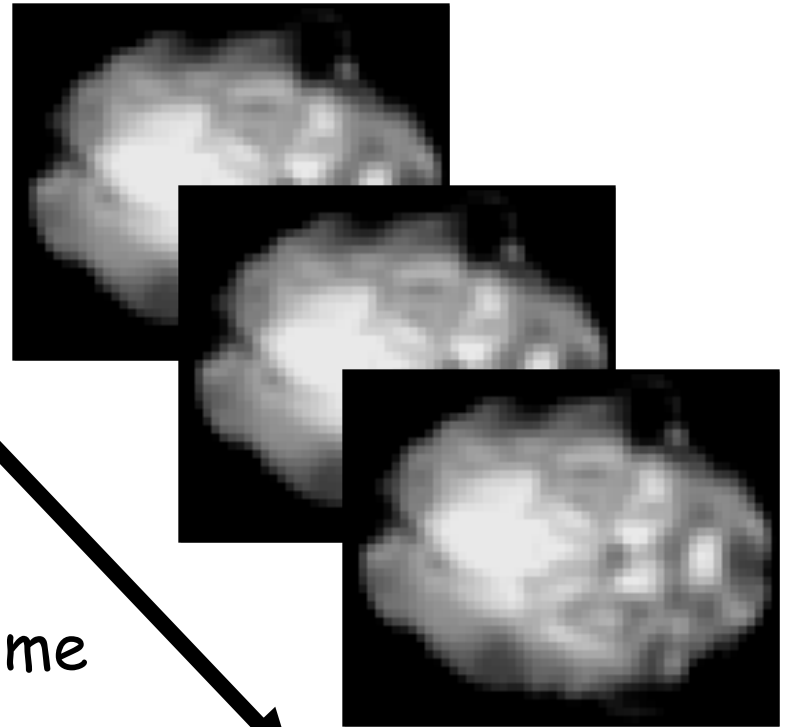
MRI



one image

high resolution  
(1 mm or less)

fMRI



Time

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

low resolution  
(1.5 to 4 mm)



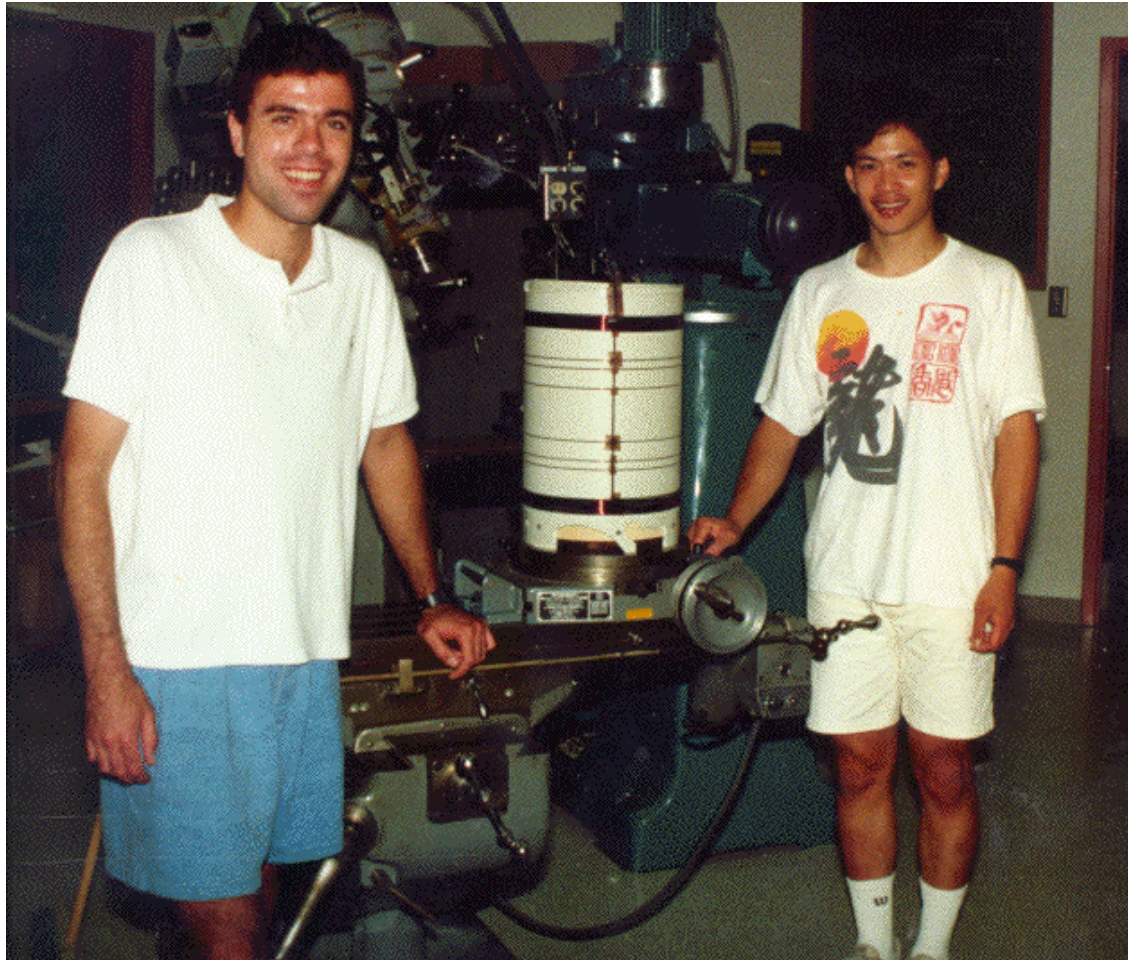


**1991-1992**



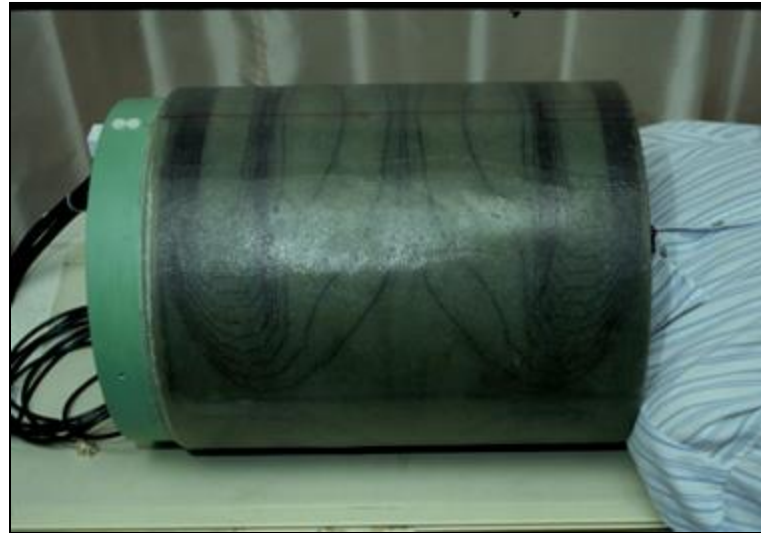
**1992-1999**





**August, 1991**

**Local Gradient Coil  
(low inductance)**



**Whole body gradients  
(more powerful amplifiers)**







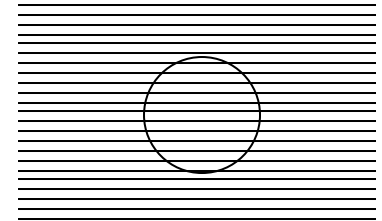
# Blood Oxygenation

*Oxygenated and deoxygenated red blood cells have different magnetic properties*

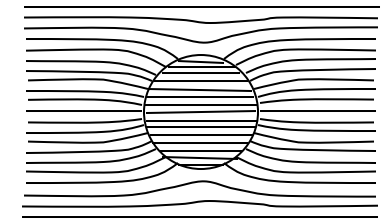


red blood cells

oxygenated



deoxygenated

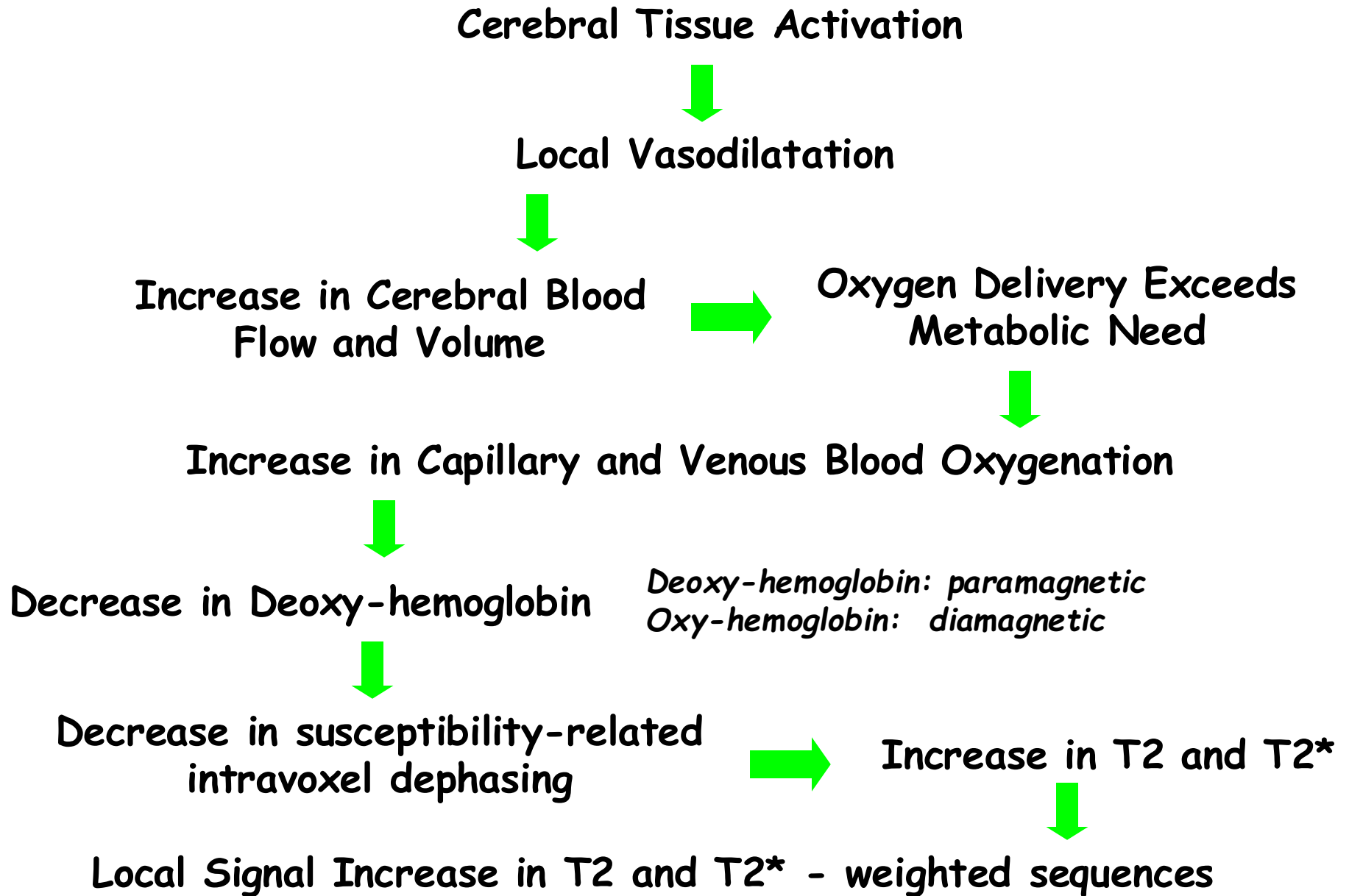


L. Pauling, C. D. Coryell, *Proc. Natl. Acad. Sci. USA* 22, 210-216, **1936**.

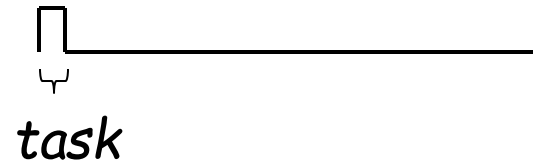
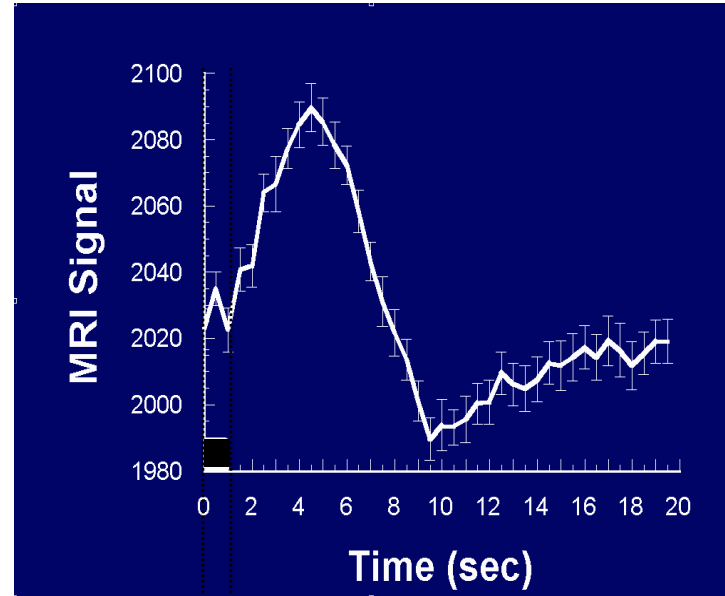
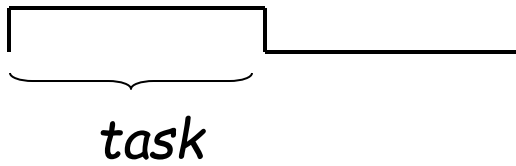
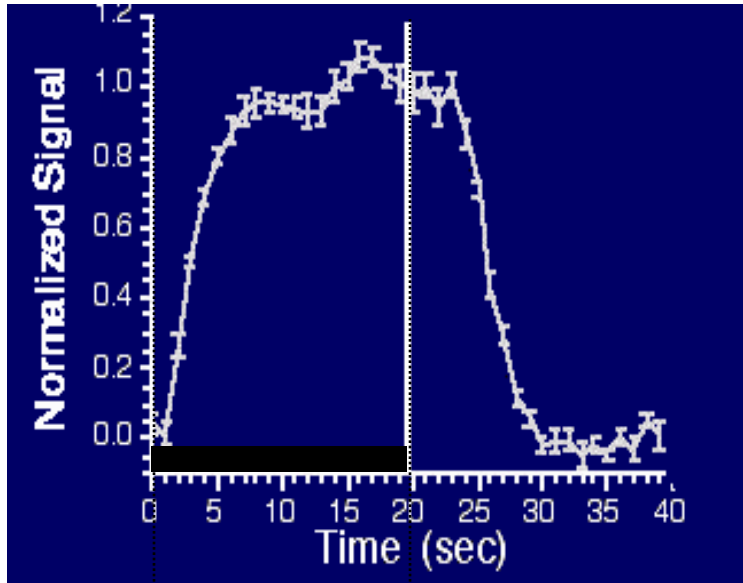
K.R. Thulborn, J. C. Waterton, et al., *Biochim. Biophys. Acta.* 714: 265-270, **1982**.

S. Ogawa, T. M. Lee, A. R. Kay, D. W. Tank, *Proc. Natl. Acad. Sci. USA* 87, 9868-9872, **1990**.

# Blood Oxygenation

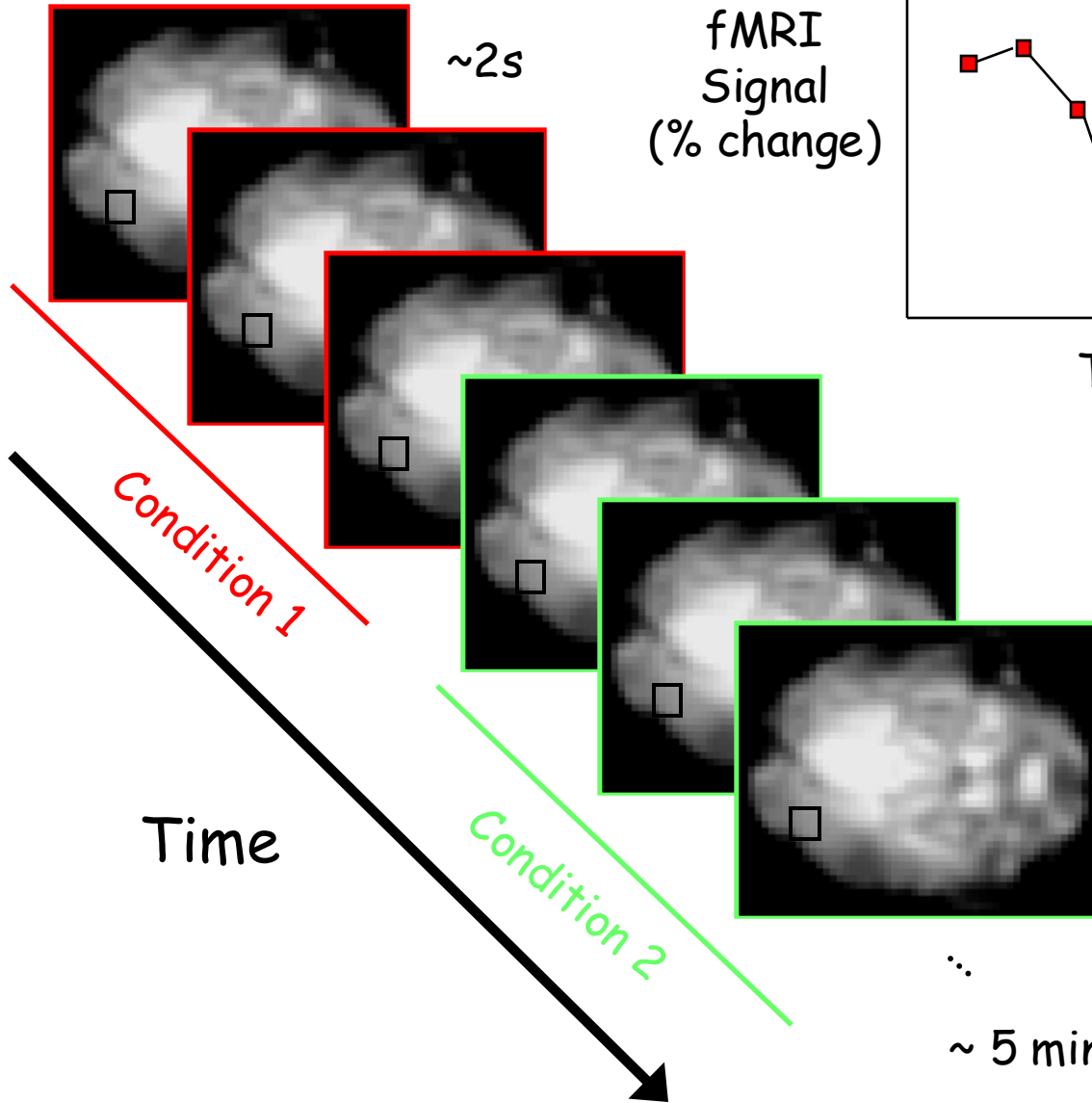


# Blood Oxygenation

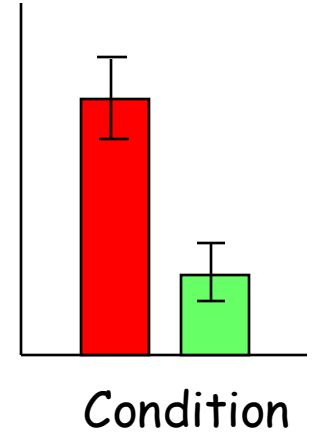
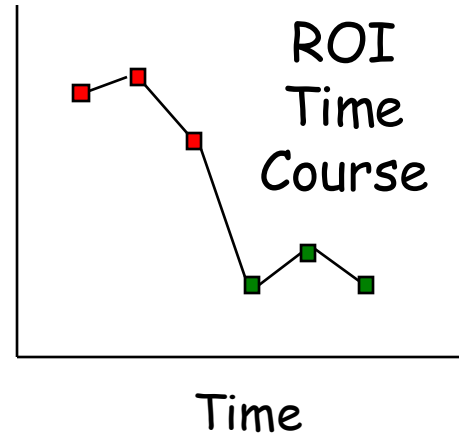


# Activation Statistics

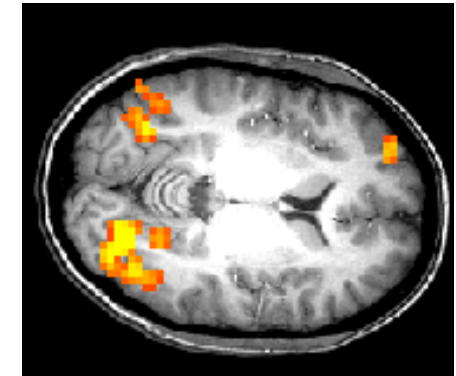
Functional images



fMRI  
Signal  
(% change)

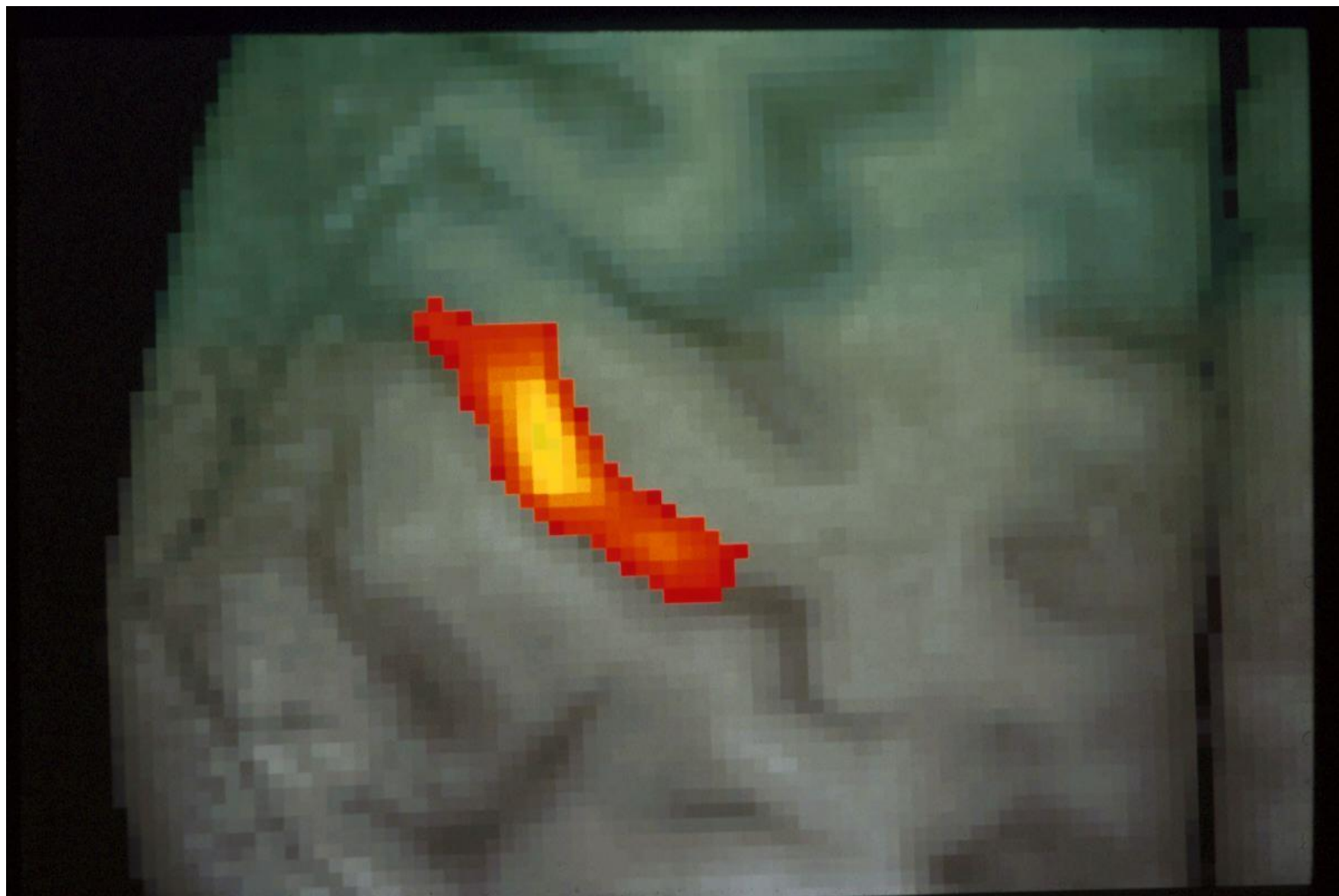


Statistical Map  
superimposed on  
anatomical MRI image



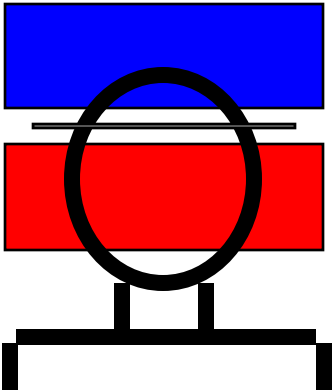
Courtesy, Robert Cox



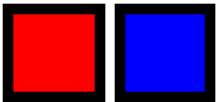
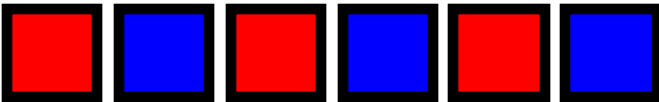
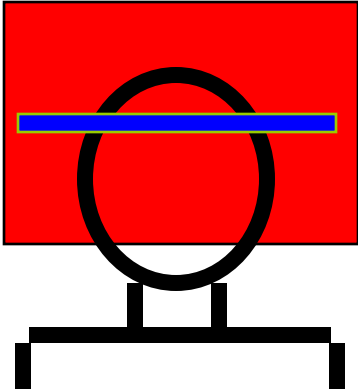


# Perfusion

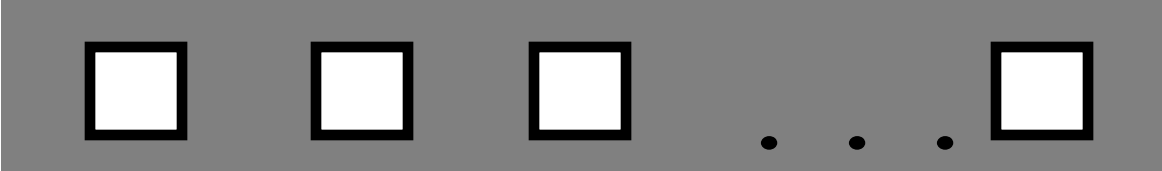
## EPISTAR



## FAIR

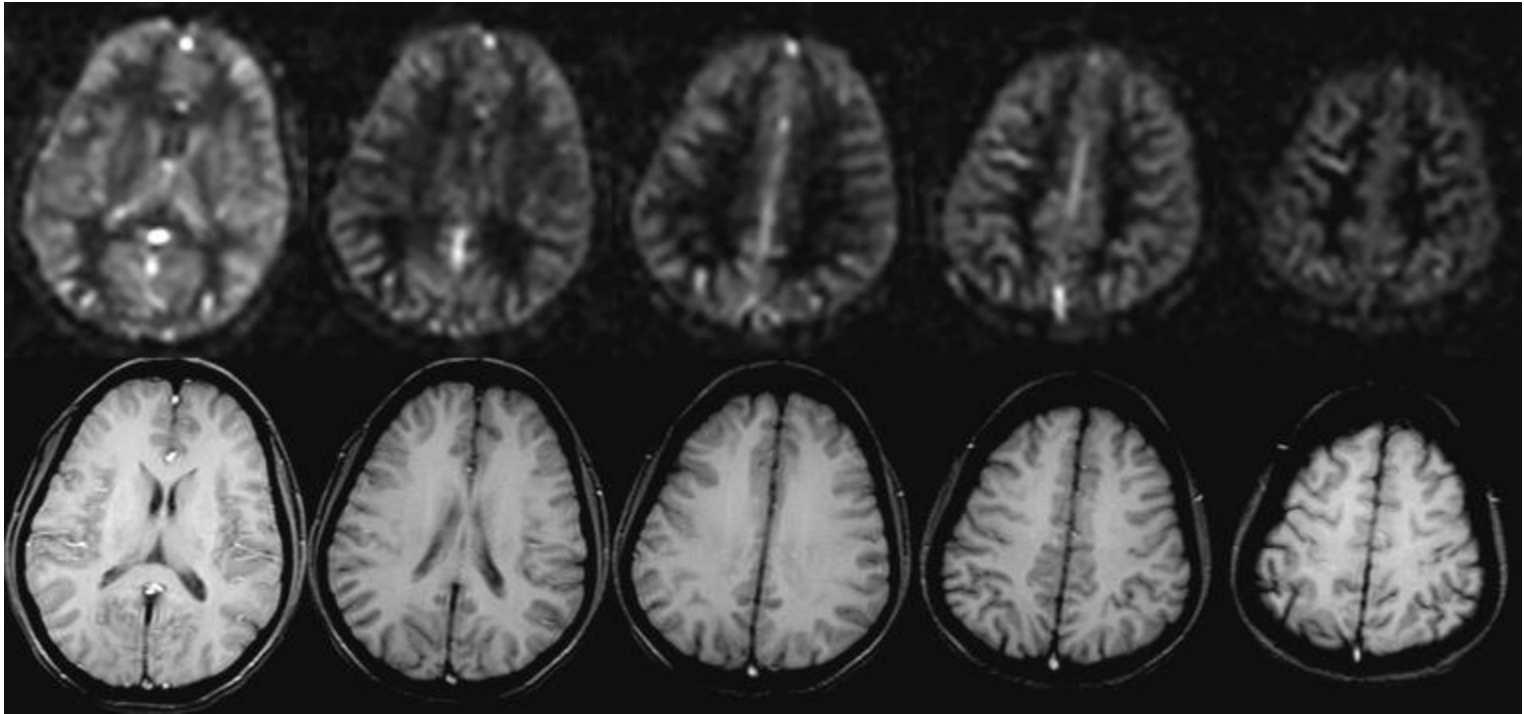


...



Perfusion  
Time Series

# Perfusion



Williams, D. S., Detre, J. A., Leigh, J. S. & Koretsky, A. S. (1992) "Magnetic resonance imaging of perfusion using spin-inversion of arterial water." *Proc. Natl. Acad. Sci. USA* 89, 212-216.

Edelman, R., Siewert, B. & Darby, D. (1994) "Qualitative mapping of cerebral blood flow and functional localization with echo planar MR imaging and signal targeting with alternating radiofrequency (EPISTAR)." *Radiology* 192, 1-8.

Kim, S.-G. (1995) "Quantification of relative cerebral blood flow change by flow-sensitive alternating inversion recovery (FAIR) technique: application to functional mapping." *Magn. Reson. Med.* 34, 293-301.

Kwong, K. K. et al. (1995) "MR perfusion studies with T1-weighted echo planar imaging." *Magn. Reson. Med.* 34, 878-887.

# Perfusion

TI (ms)

FAIR

EPISTAR

200

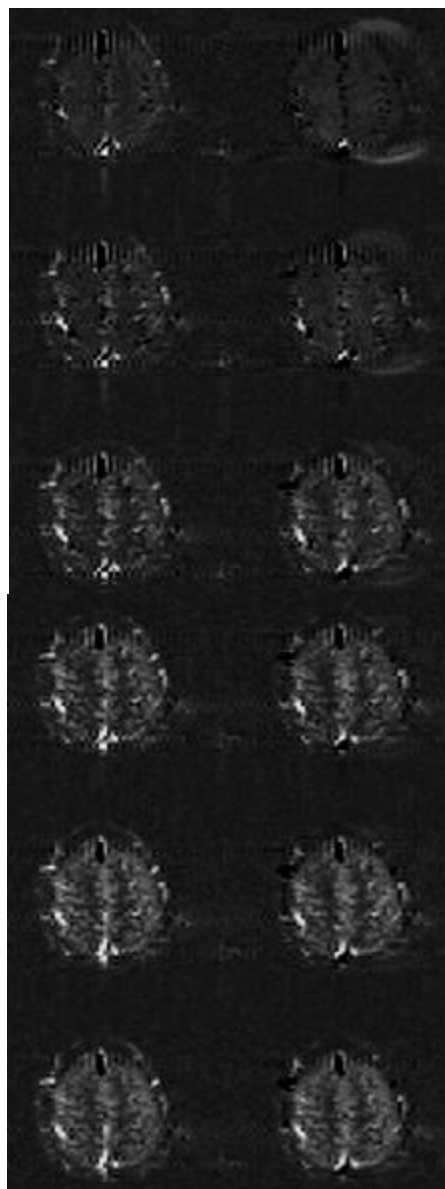
400

600

800

1000

1200

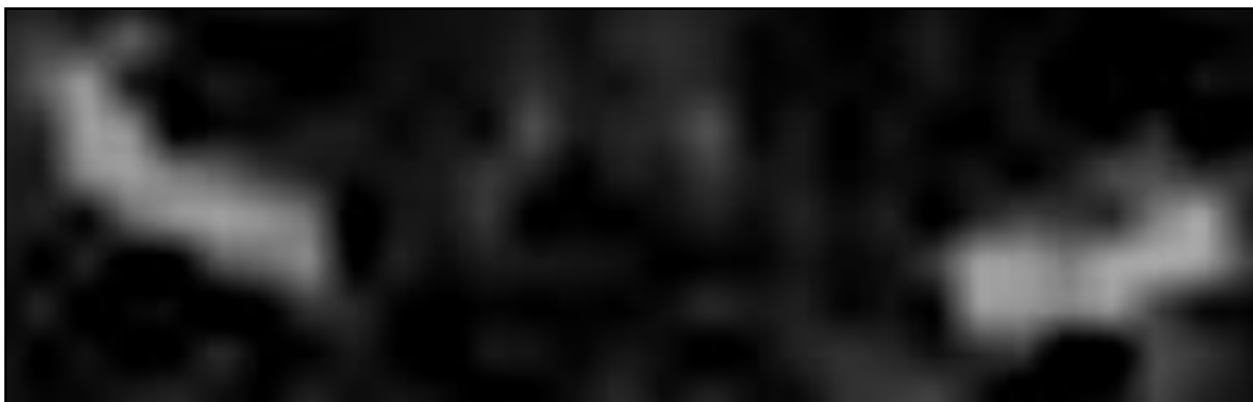


Perfusion

Simultaneous BOLD and Perfusion



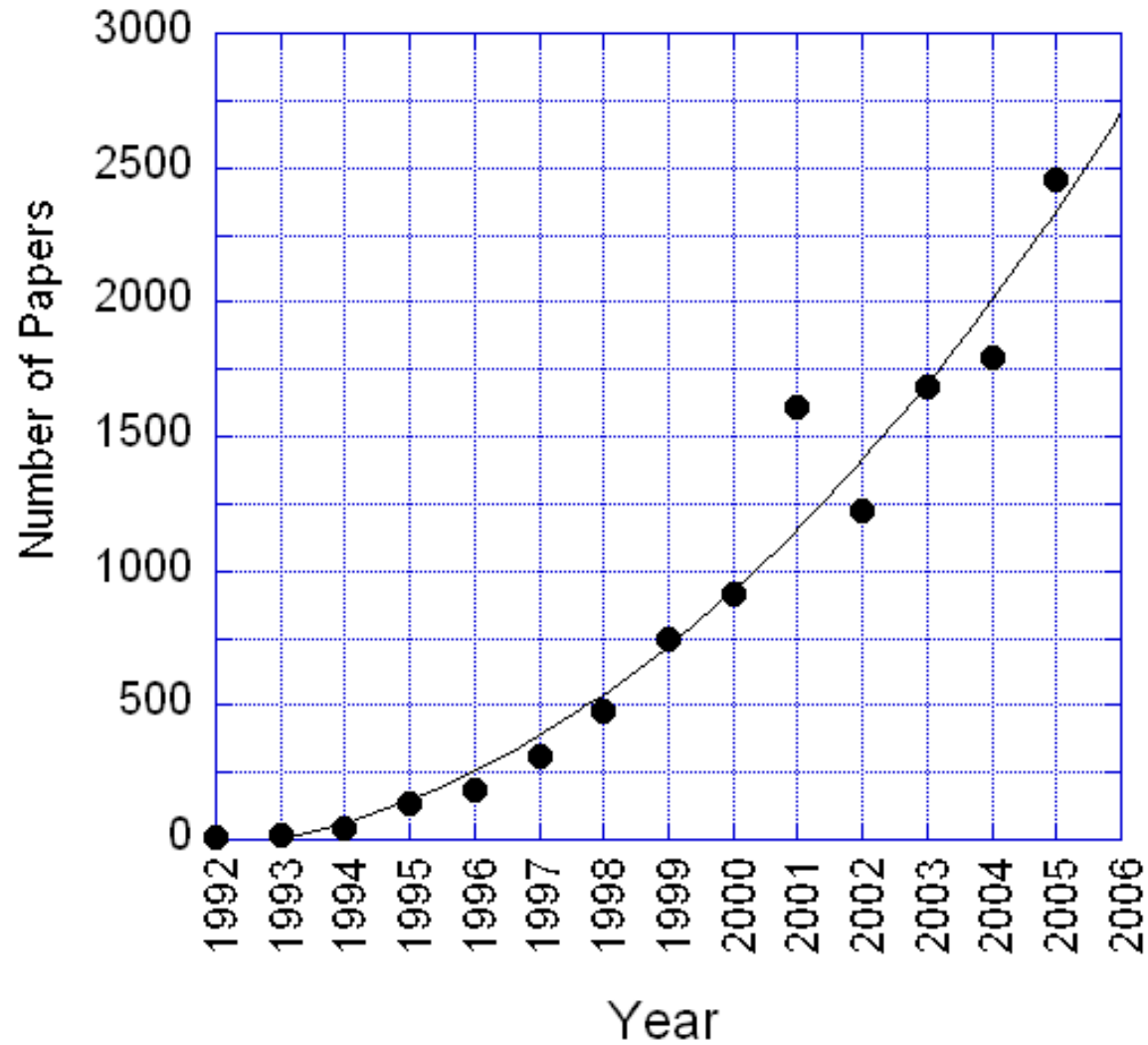
**BOLD**



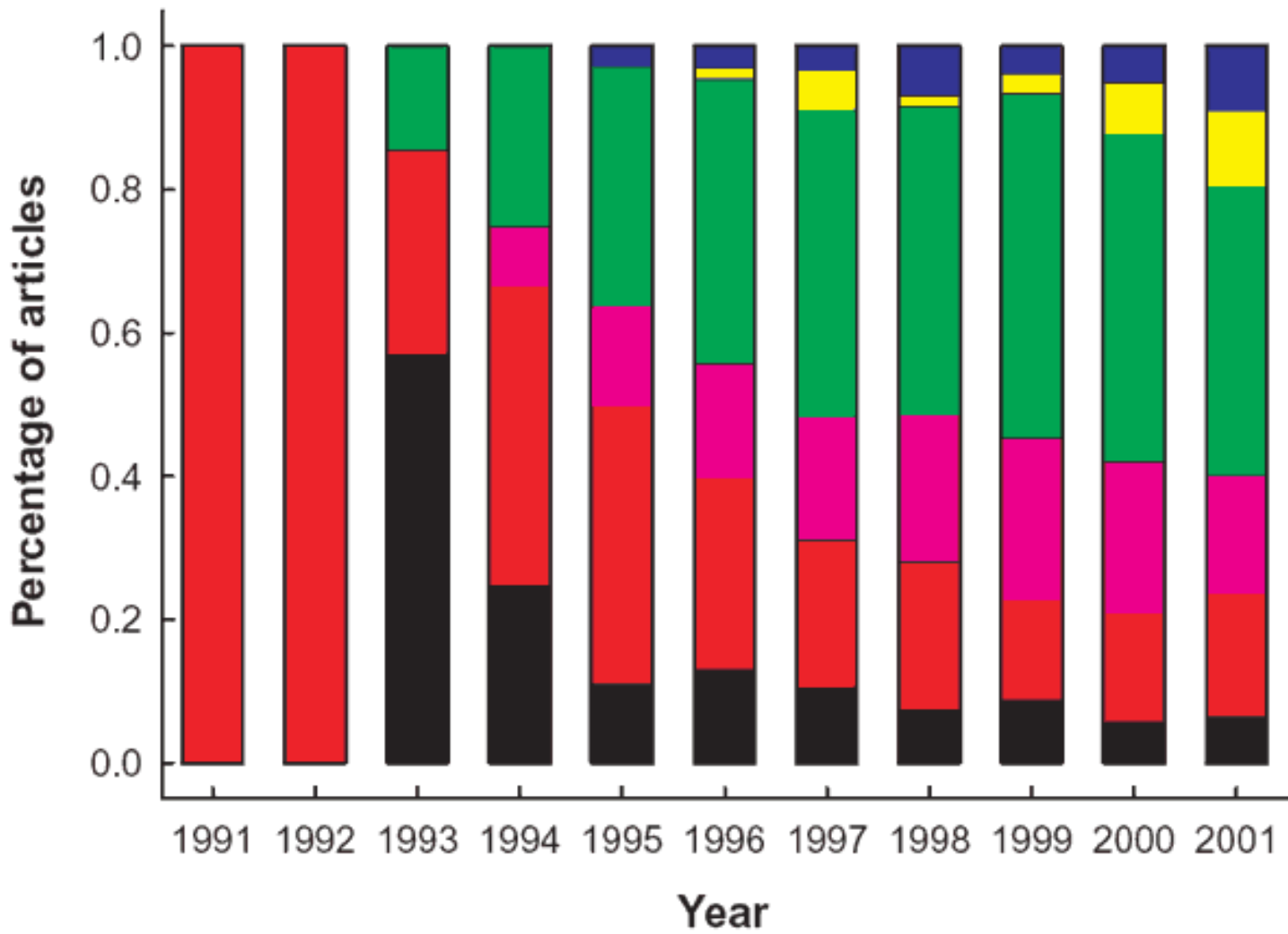
**Perfusion**



# fMRI Papers Published per Year



**"fMRI" or "functional MRI"**



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

J. Illes, M. P. Kirschen, J. D. E. Gabrieli,  
 Nature Neuroscience, 6 (3) p.205

# Technology

Coil arrays  
High field strength  
High resolution  
Novel functional contrast

# Methodology

Connectivity assessment  
Multi-modal integration  
Pattern classification  
Task design

Fluctuations  
Dynamics  
Cross - modal comparison

Basic Neuroscience  
Behavior correlation/prediction  
Pathology correlation

# Interpretation

# Applications



# Technology

Coil arrays  
High field strength  
High resolution  
Novel functional contrast

# Methodology

Connectivity assessment  
Multi-modal integration  
Pattern classification  
Task design

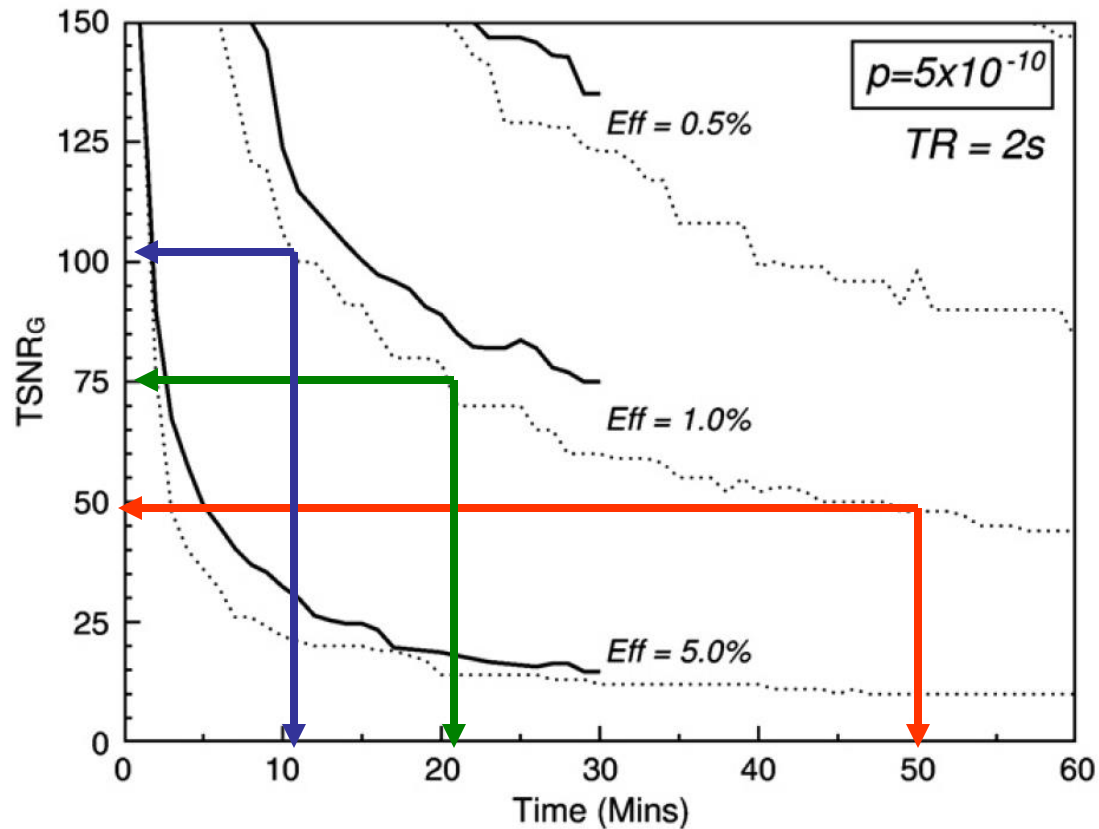
Fluctuations  
Dynamics  
Cross - modal comparison

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# Interpretation

# Applications

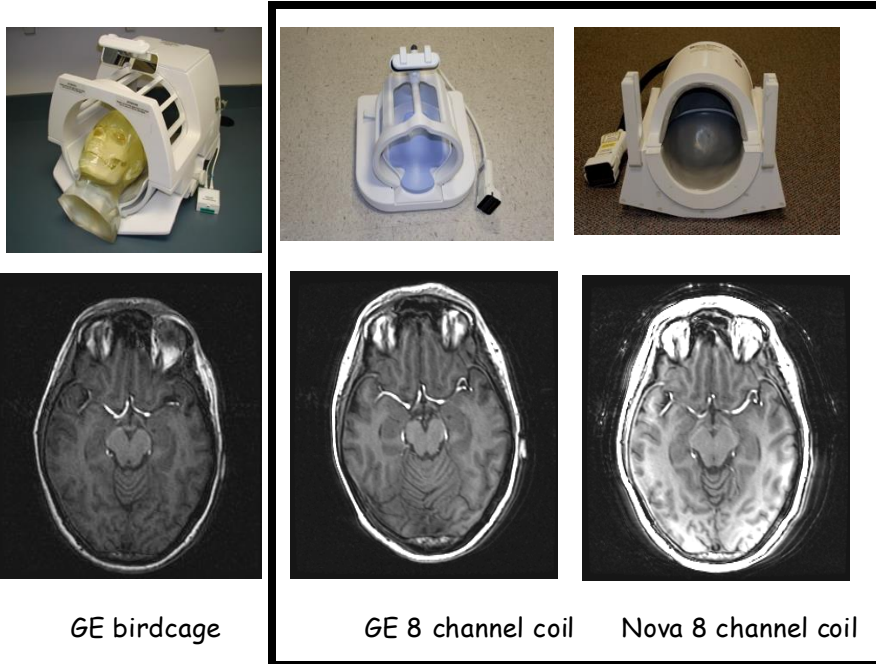
# Technology



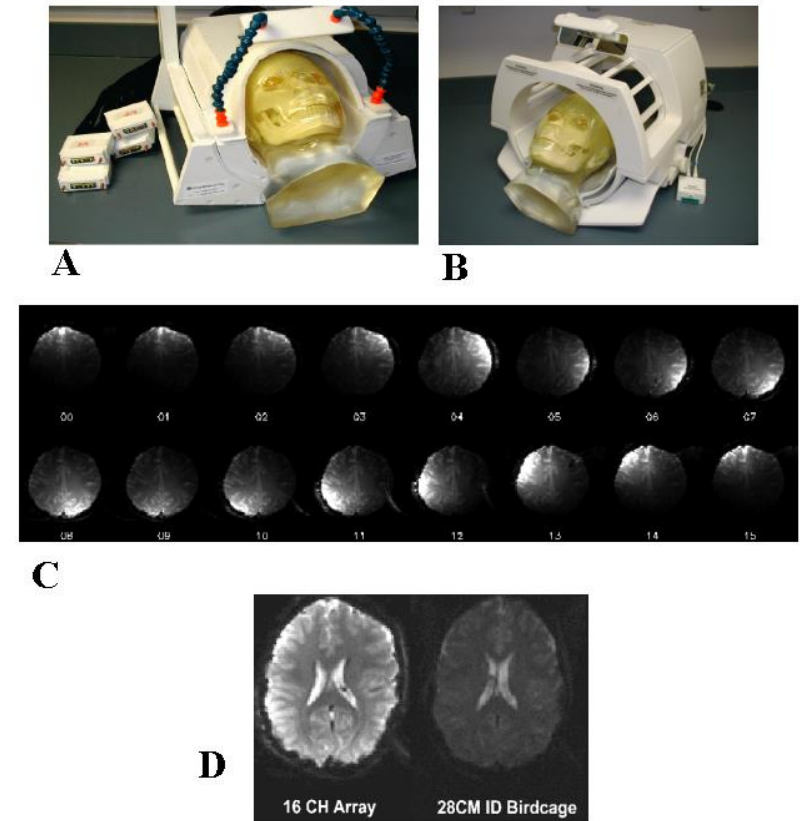
K. Murphy, J. Bodurka, P. A. Bandettini, How long to scan?  
The relationship between fMRI temporal signal to noise and the  
necessary scan duration. *NeuroImage*, 34, 565-574 (2007)

# Technology

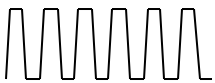
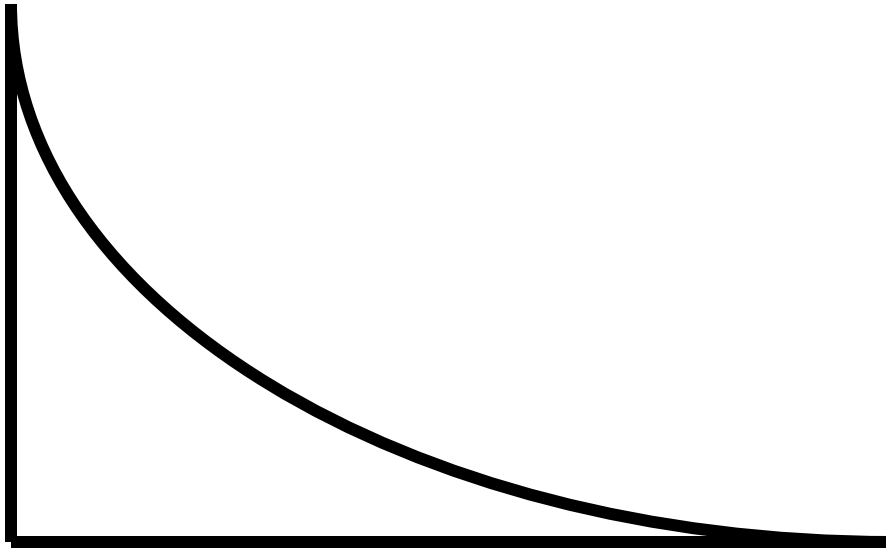
## 8 channel parallel receiver coil



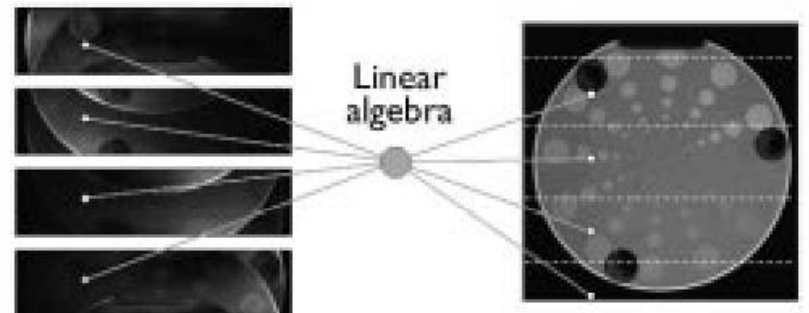
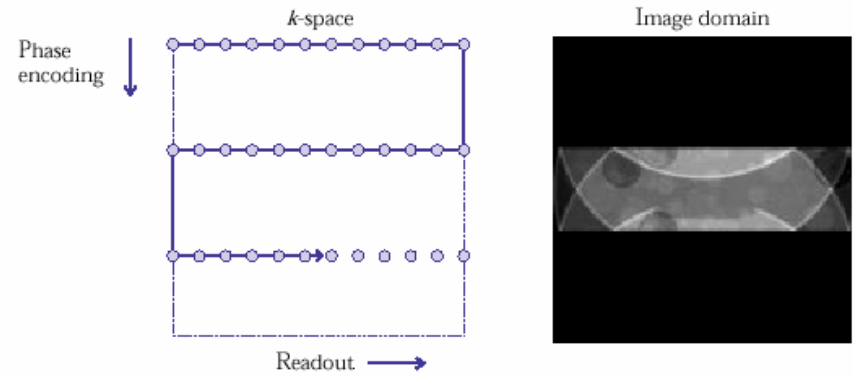
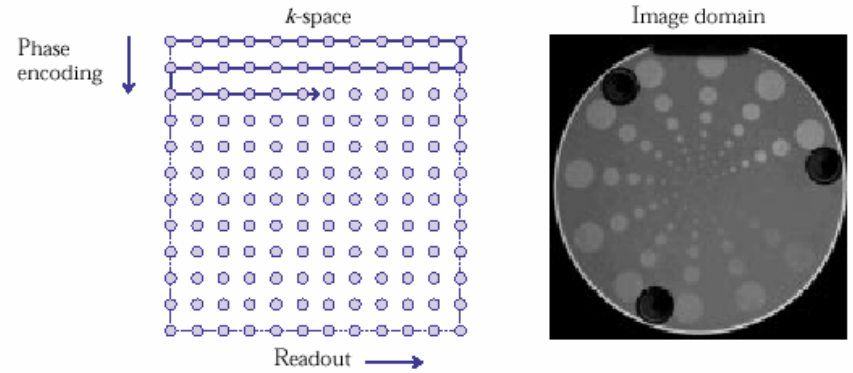
## 16 channel parallel receiver coil



# Technology

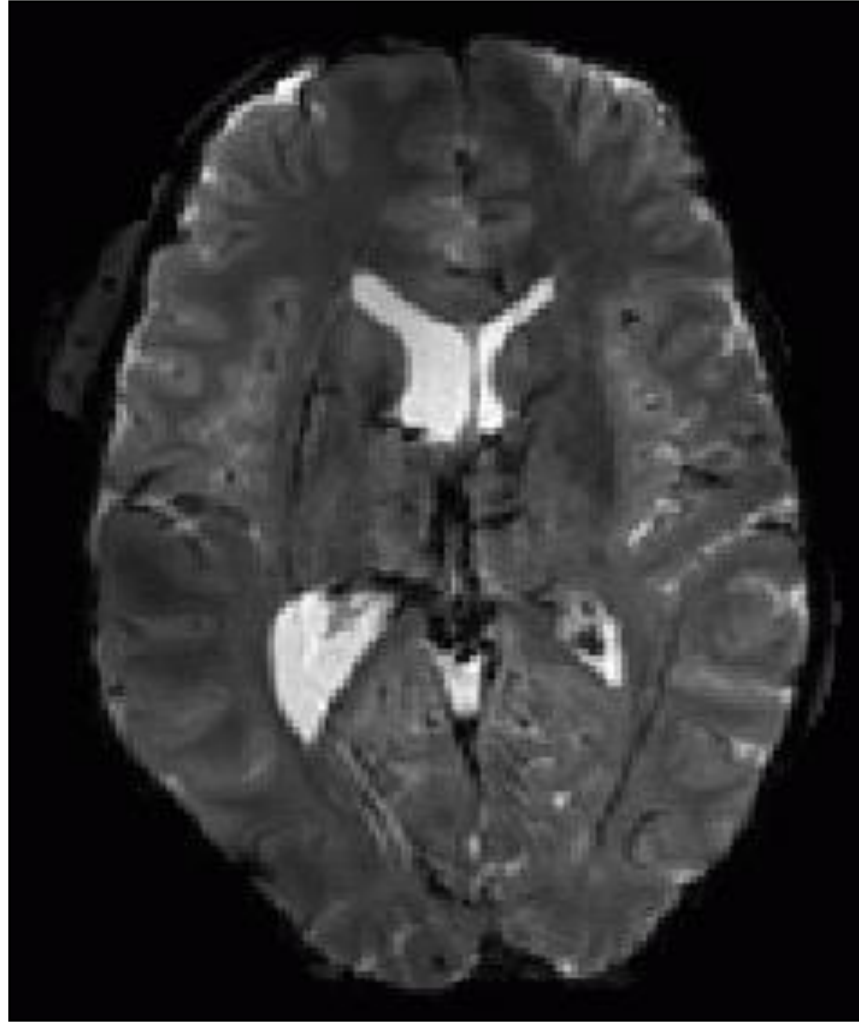


≈ 5 to 30 ms



Pruessmann, et al.

# Technology

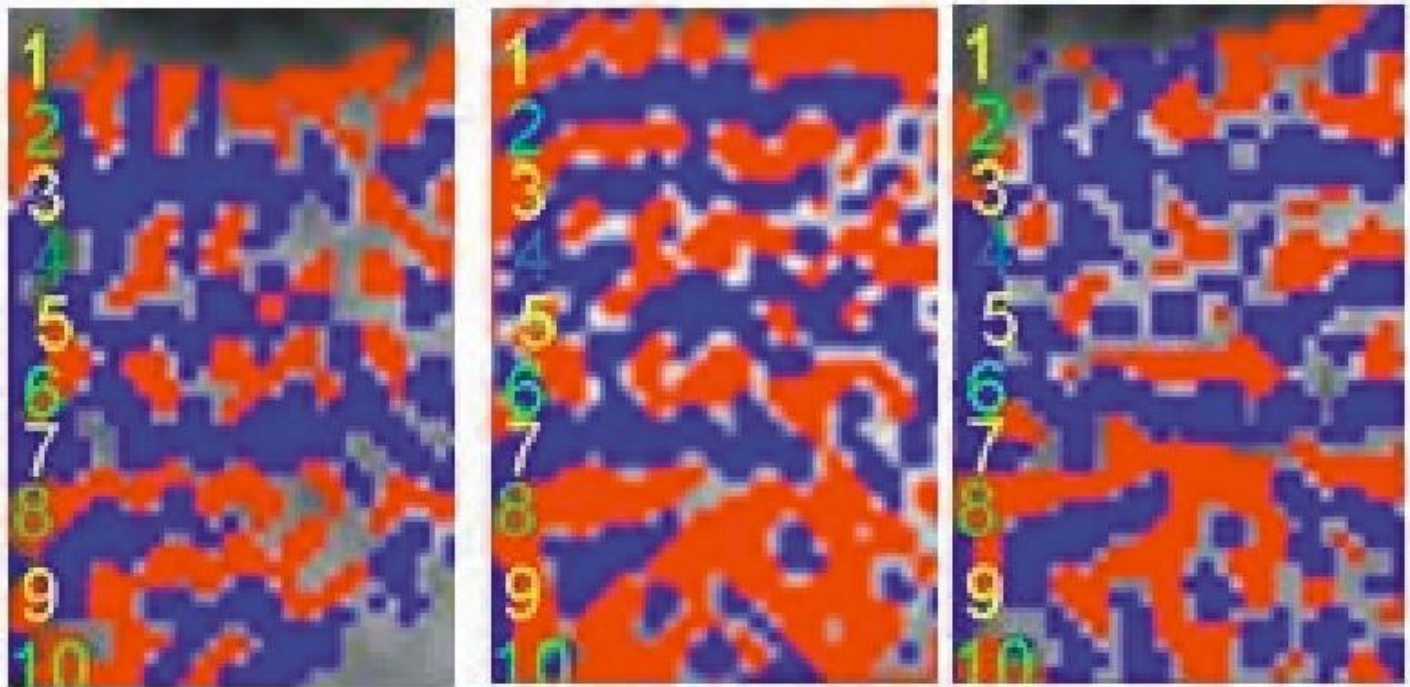


3T single-shot SENSE EPI using 16 channels: 1.25x1.25x2mm

# Technology

## HSE-BOLD demonstration of ocular dominance columns

human, 7T,  $0.5 \times 0.5 \times 3 \text{ mm}^3$



day 1

day 2

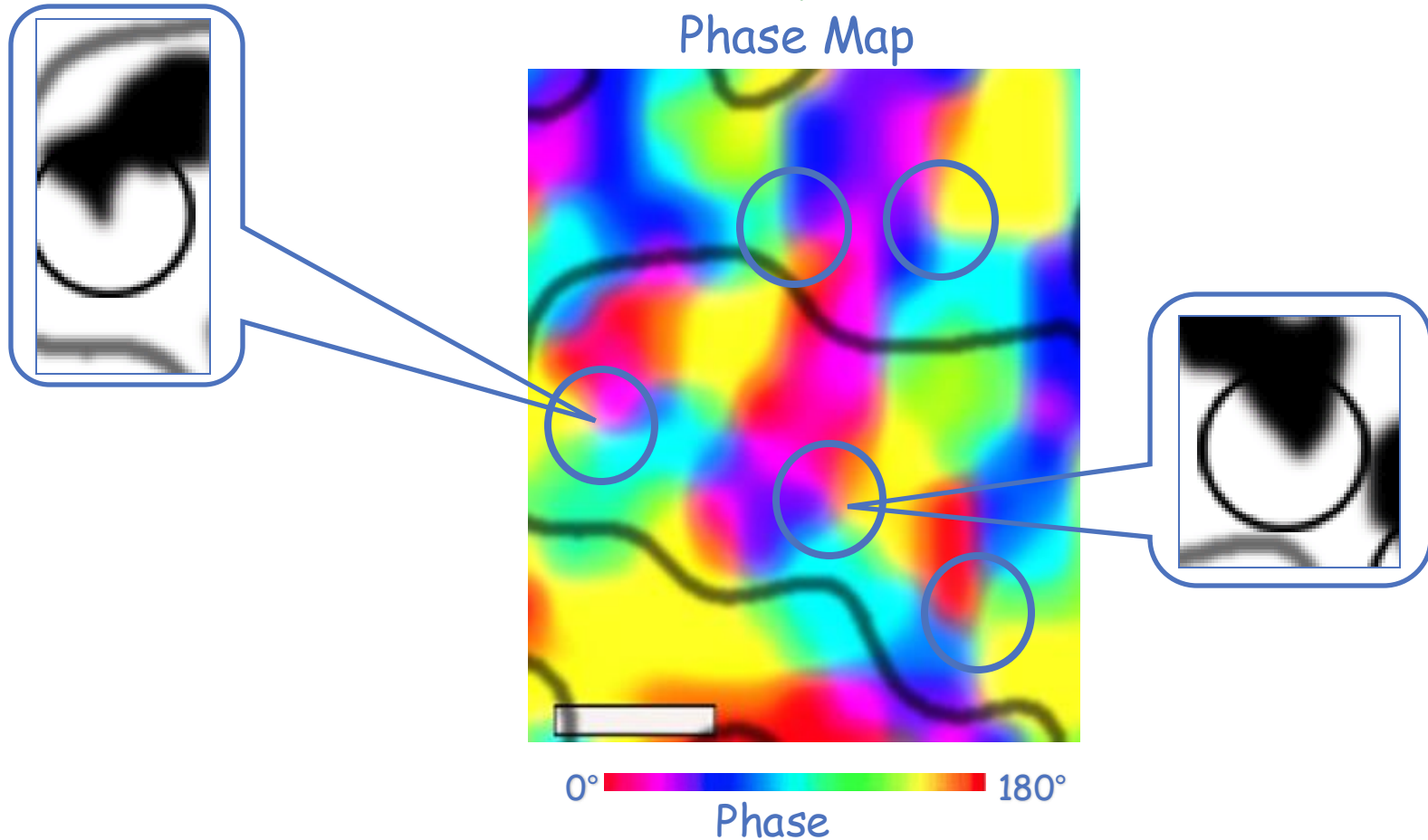
day 3

Yacoub et al: differential maps contrasting stimulation of the left and right eye



# Orientation Columns in Human V1 as Revealed by fMRI at 7T

Phase Map



Yacoub, Ugurbil & Harel  
University of Minnesota / CMRR  
HBM 2006

Scalebar = 0.5 mm

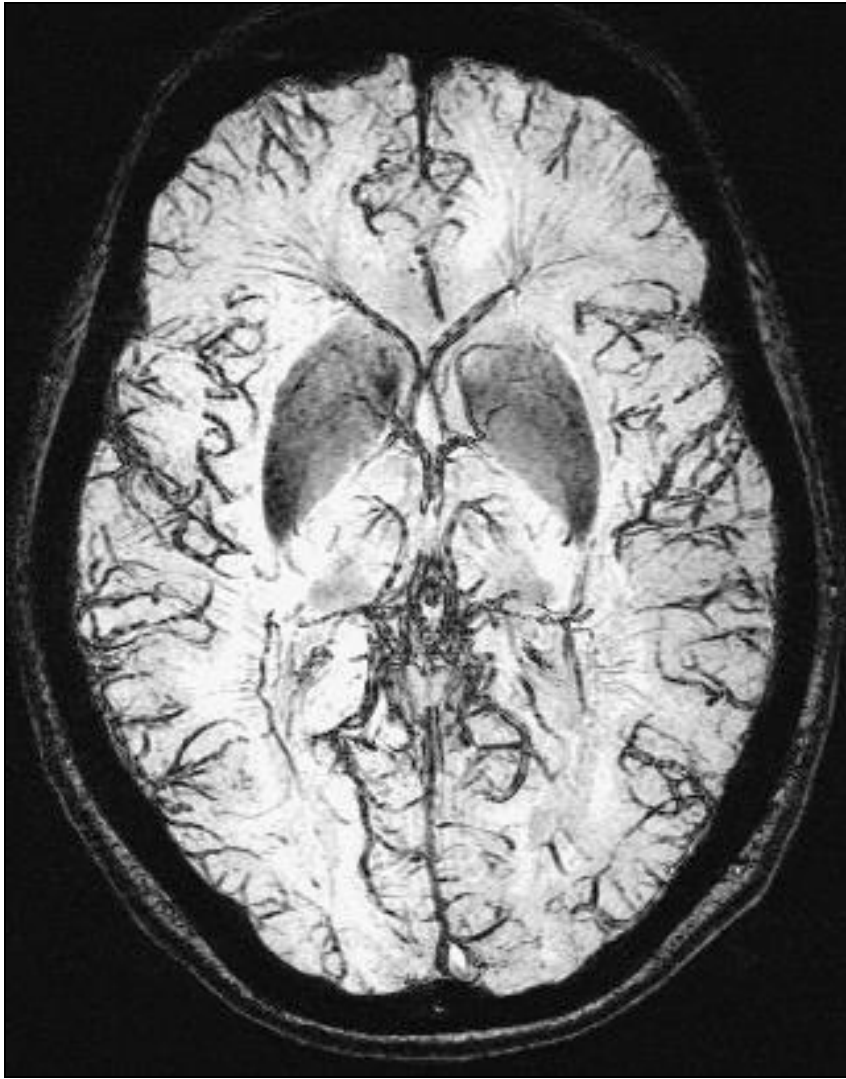
## fMRI Contrast

- Volume (gadolinium)
- BOLD
- Perfusion (ASL)
- $\Delta\text{CMRO}_2$
- $\Delta\text{Volume}$  (VASO)
- Neuronal Currents
- Diffusion coefficient
- Temperature



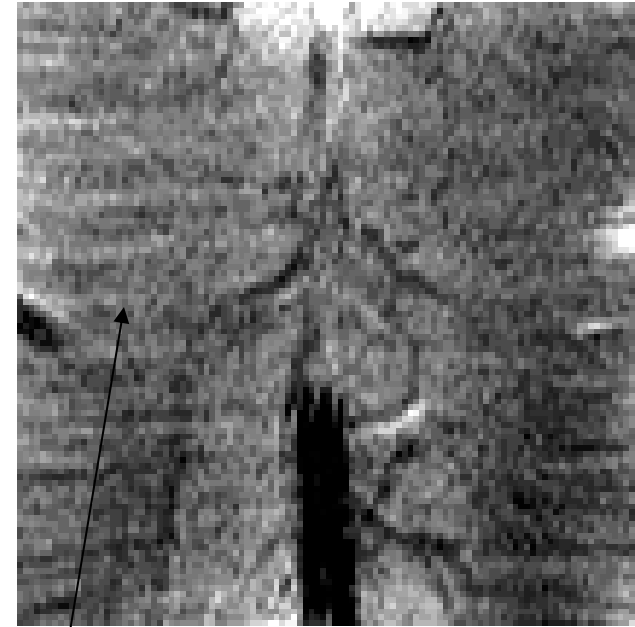
# Technology

BOLD effect to highlight veins: 3 Tesla



**Bove-Bettis, et al (2004), SMRT**

# Technology



fiber bundles?

Courtesy Tie-Qiang Li,  
NINDS

# Technology

Coil arrays  
High field strength  
High resolution  
Novel functional contrast

# Methodology

Connectivity assessment  
Multi-modal integration  
Pattern classification  
Task design

Fluctuations  
Dynamics  
Cross - modal comparison

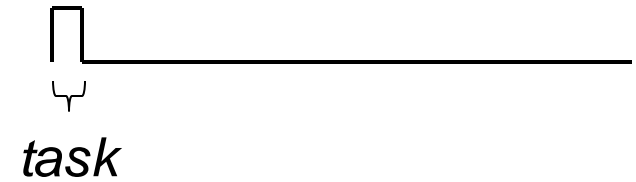
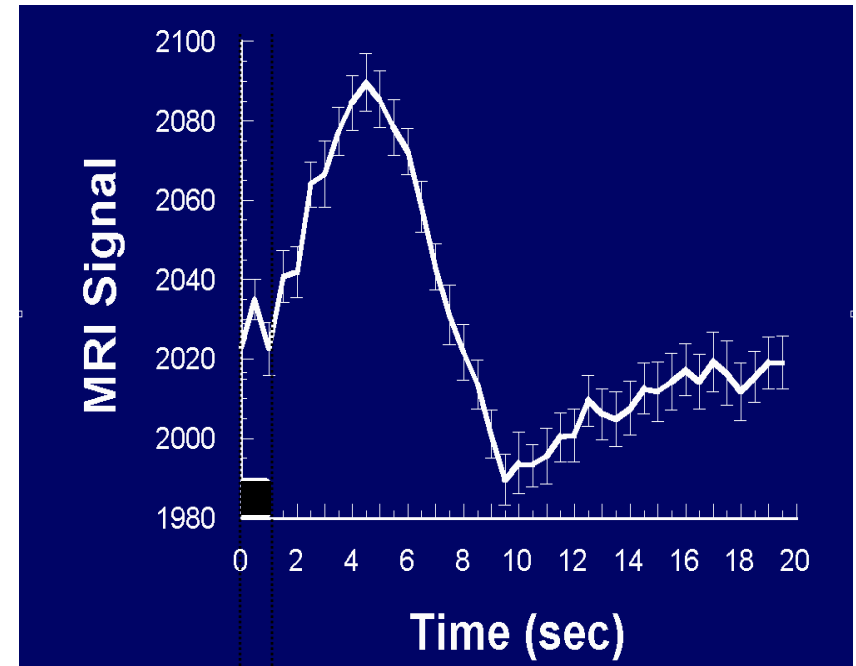
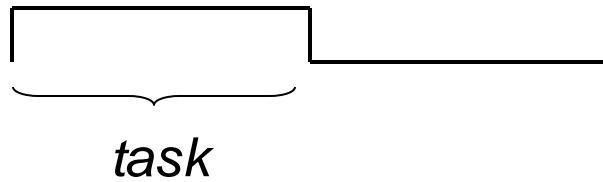
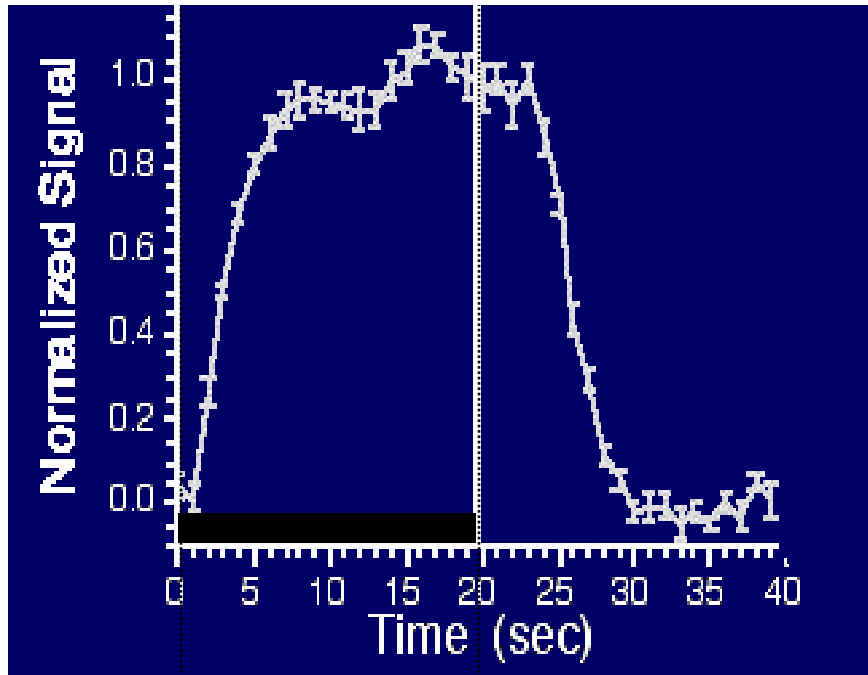
Basic Neuroscience  
Behavior correlation/prediction  
Pathology correlation

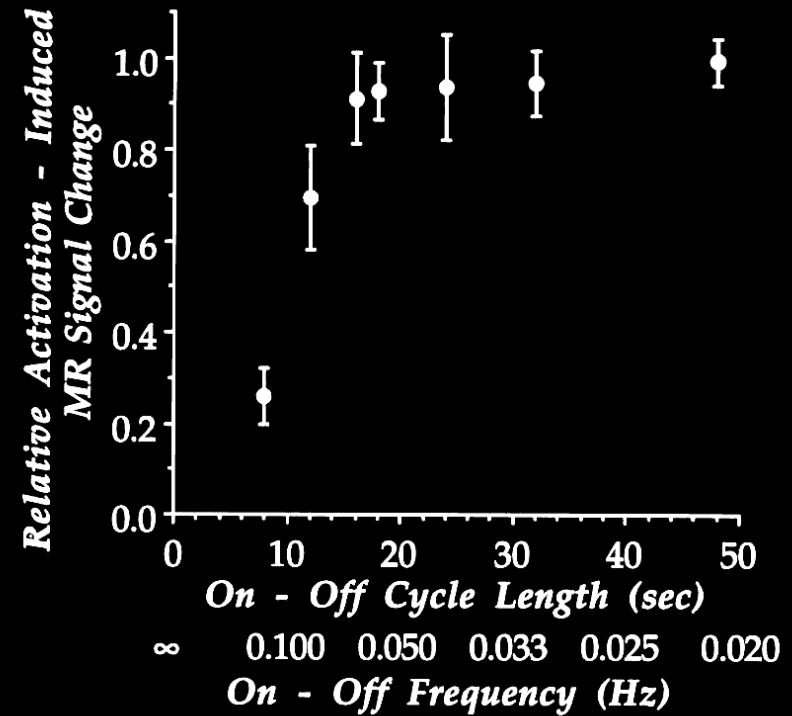
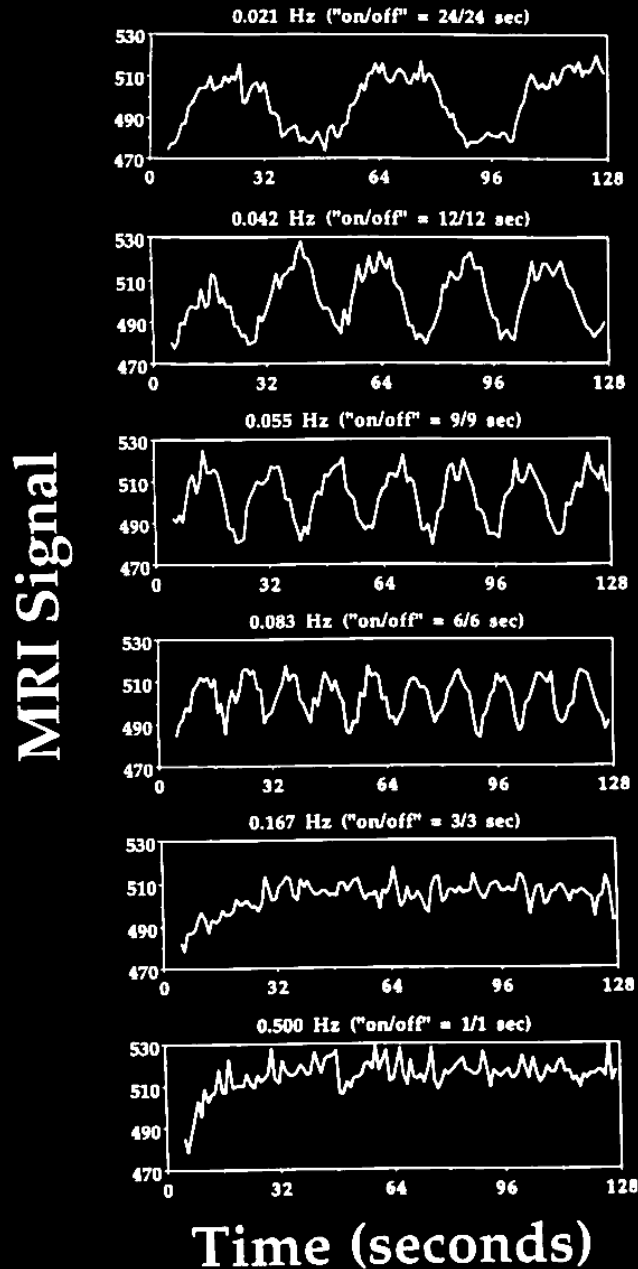
# Interpretation

# Applications

# Methodology

# Temporal Resolution



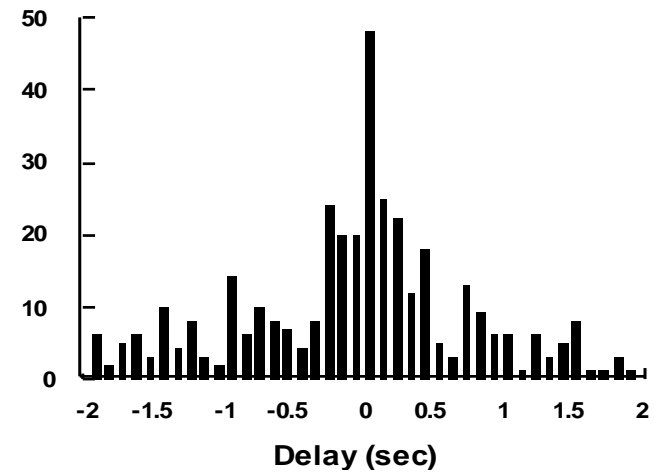
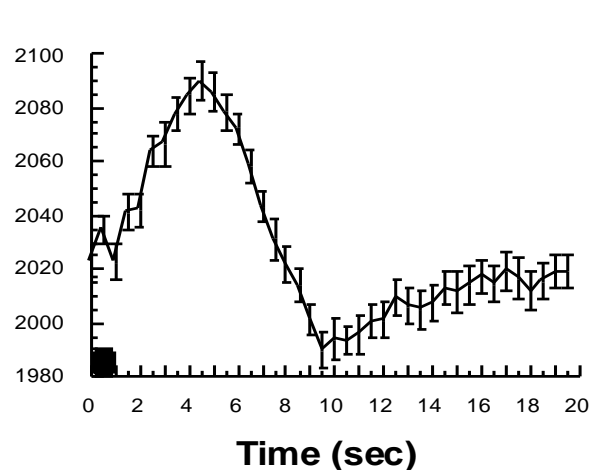
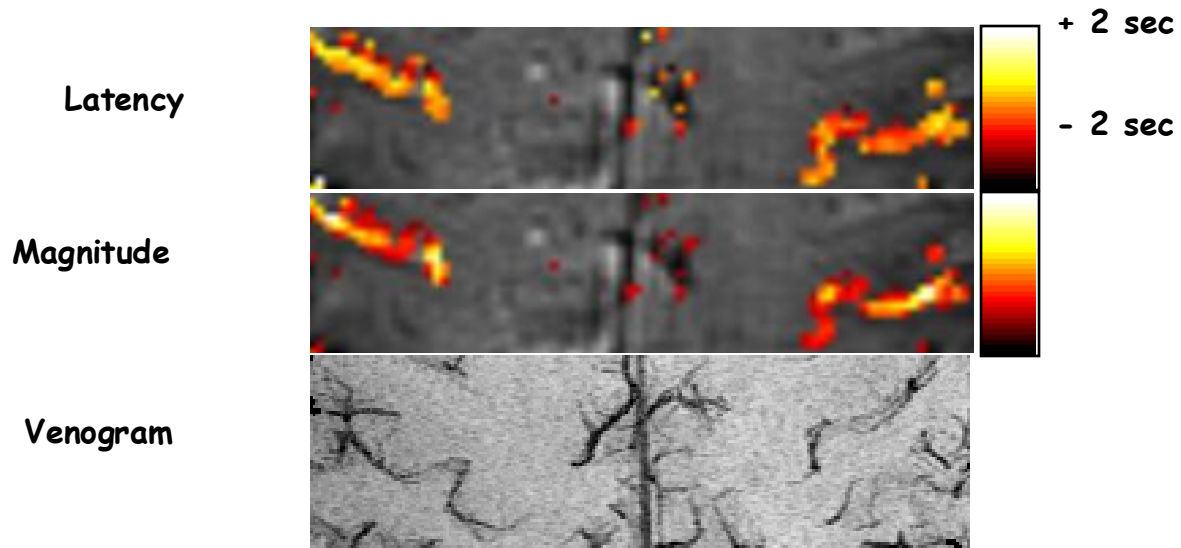


P. A. Bandettini, Functional MRI temporal resolution in "Functional MRI" (C. Moonen, and P. Bandettini., Eds.), p. 205-220, Springer - Verlag, . 1999.

# Methodology

# Temporal Resolution

## Latency Variation...



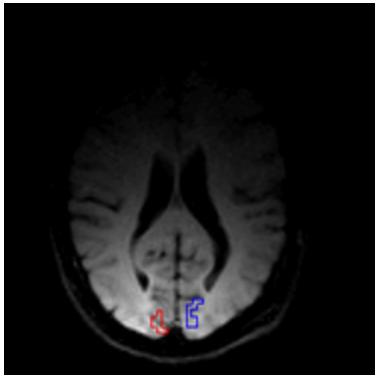
P. A. Bandettini, (1999) "Functional MRI" 205-220.

# Methodology

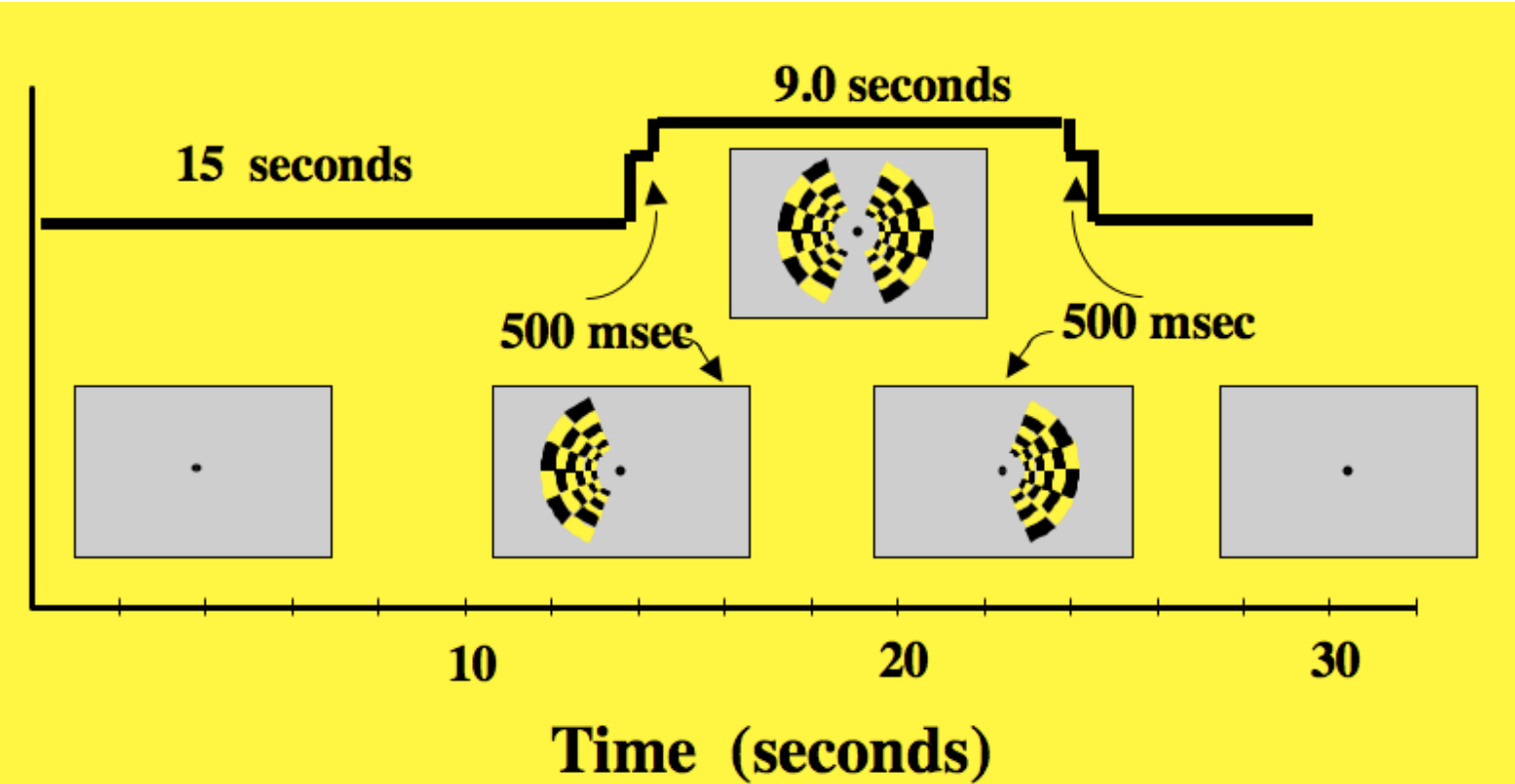
# Temporal Resolution

## Hemi-Field Experiment

Right Hemisphere

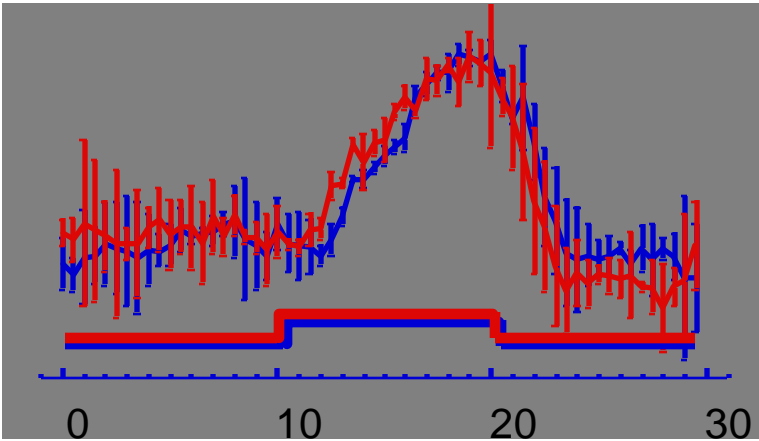
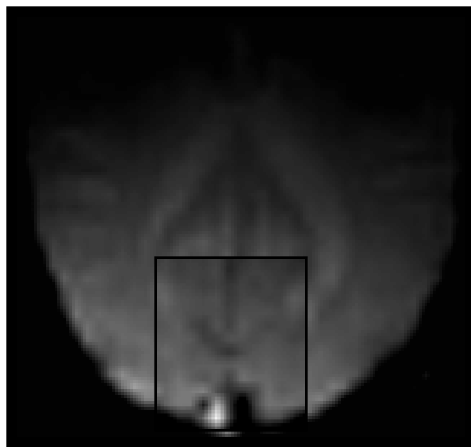


Left Hemisphere



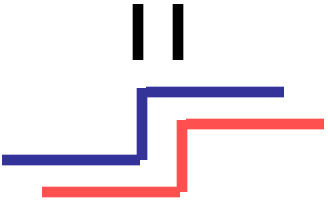
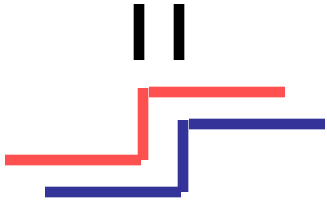
# Methodology

# Temporal Resolution



500 ms

500 ms



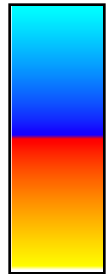
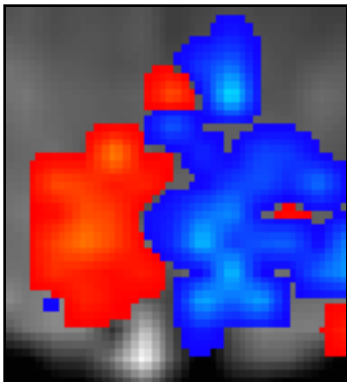
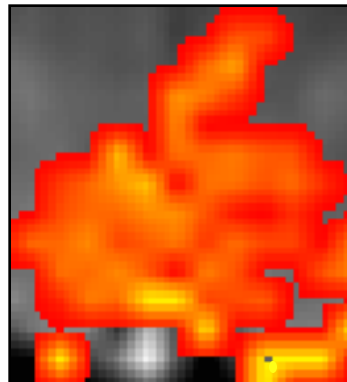
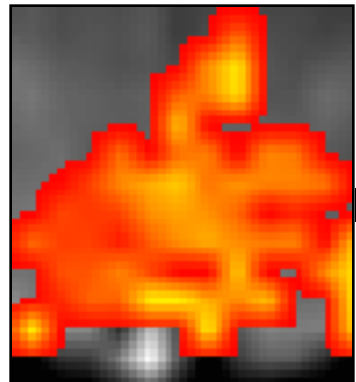
Right Hemifield

Left Hemifield

+ 2.5 s

0 s

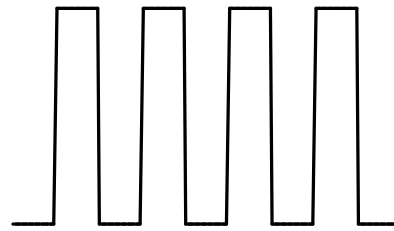
- 2.5 s



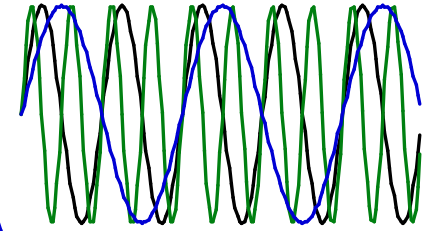


# Neuronal Activation Input Strategies

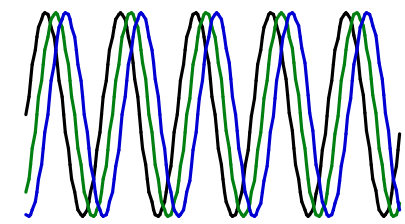
1. Block Design



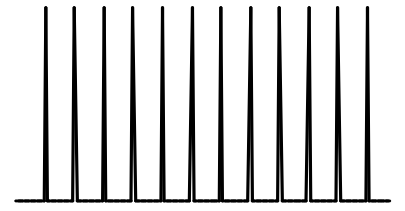
2. Frequency Encoding



3. Phase Encoding

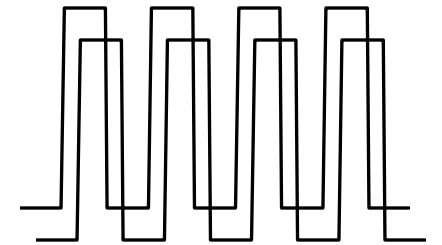


4. Event-Related

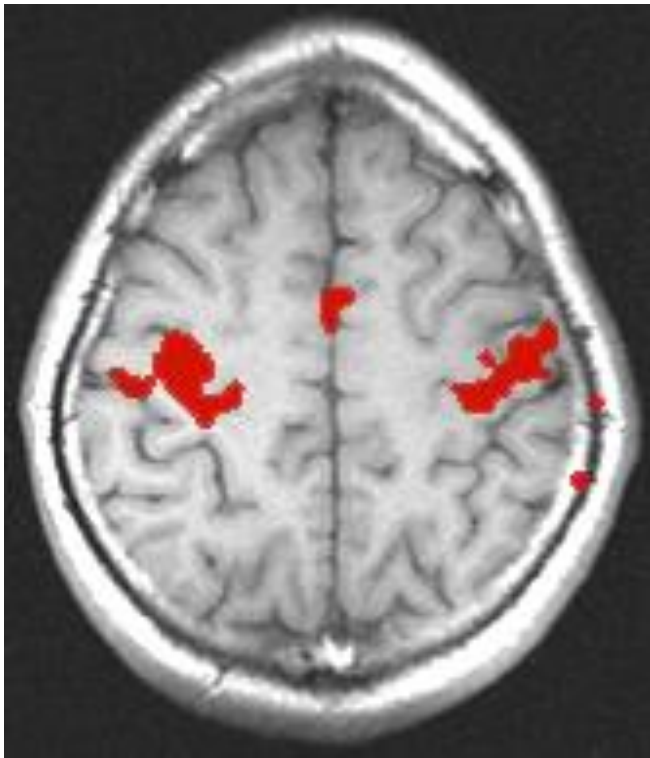


5. Orthogonal Block Design

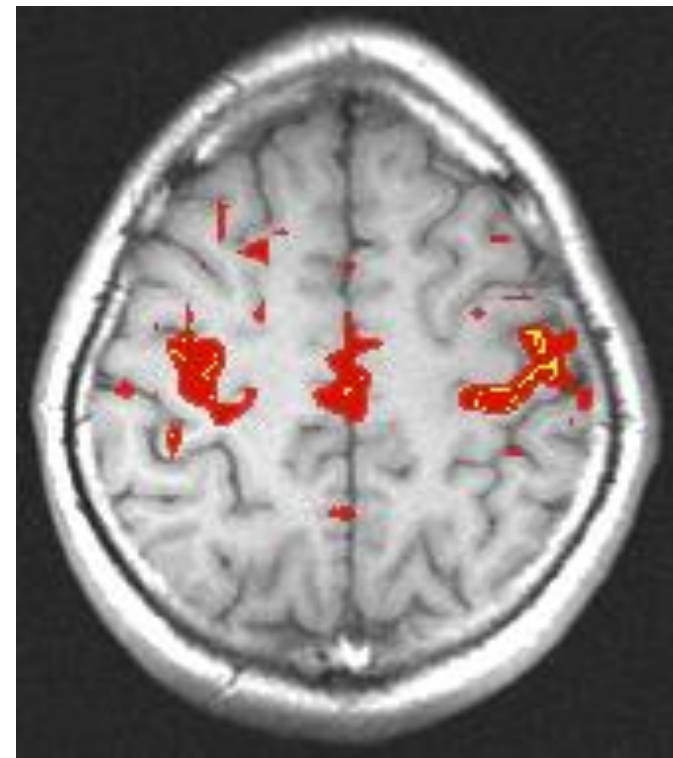
6. Free Behavior Design.



## Resting State Correlations



Activation:  
correlation with reference function

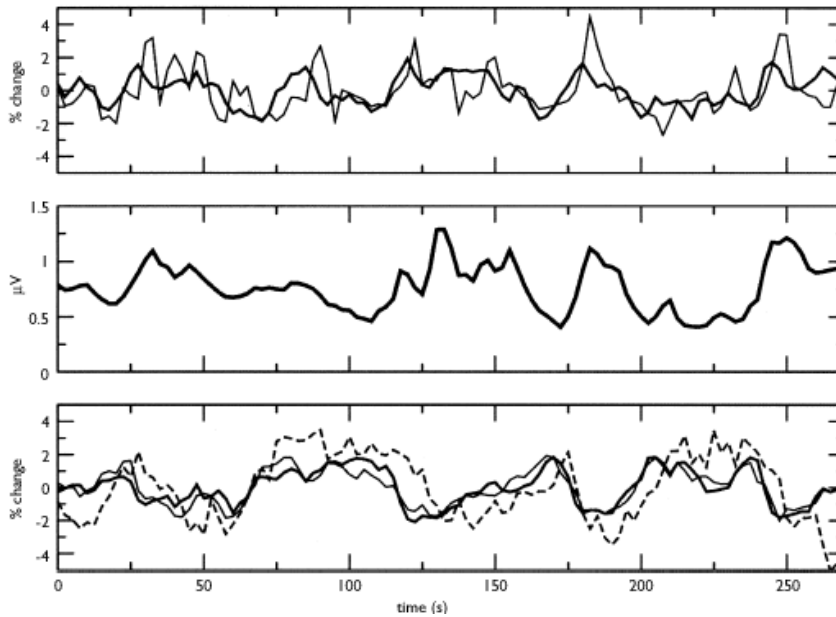


Rest:  
seed voxel in motor cortex

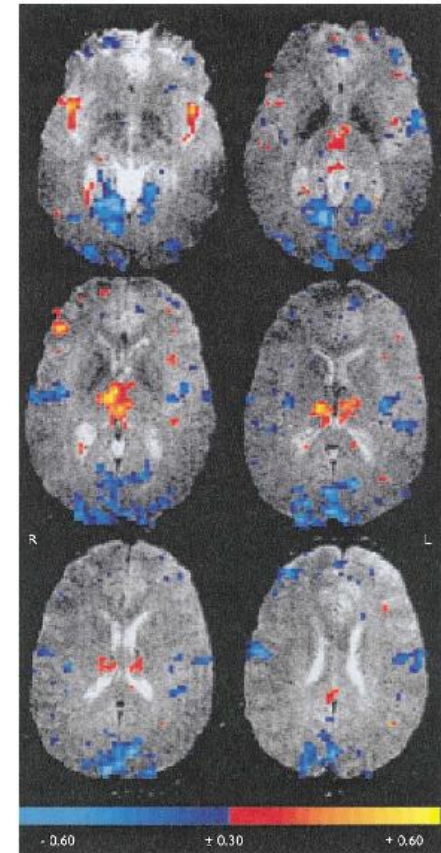
# Methodology

BOLD correlated with 10 Hz power during "Rest"

Positive  
10 Hz power  
Negative

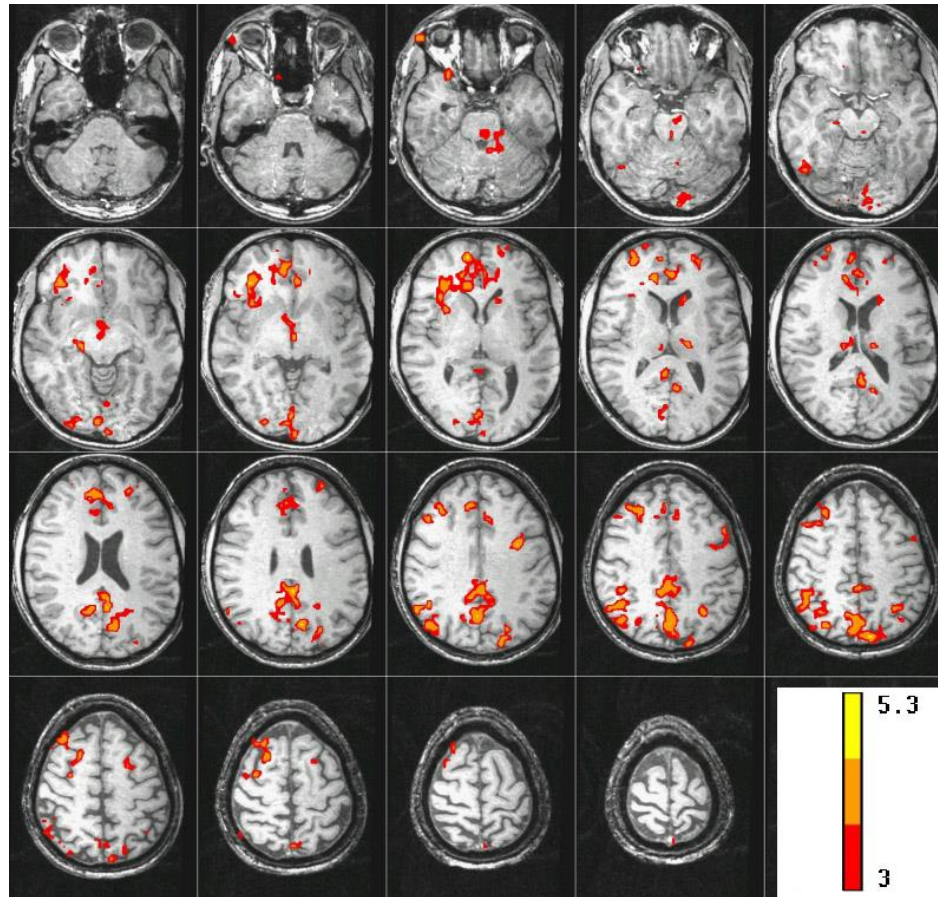


Goldman, et al (2002), Neuroreport



# Methodology

BOLD correlated with SCR during "Rest"

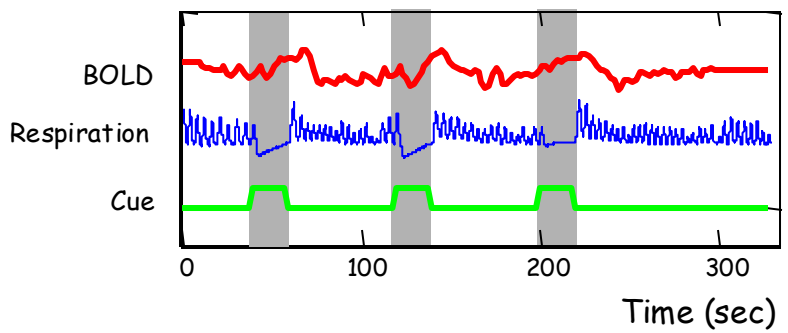


J. C. Patterson II, L. G. Ungerleider, and P. A. Bandettini, *NeuroImage* 17: 1787-1806, (2002).

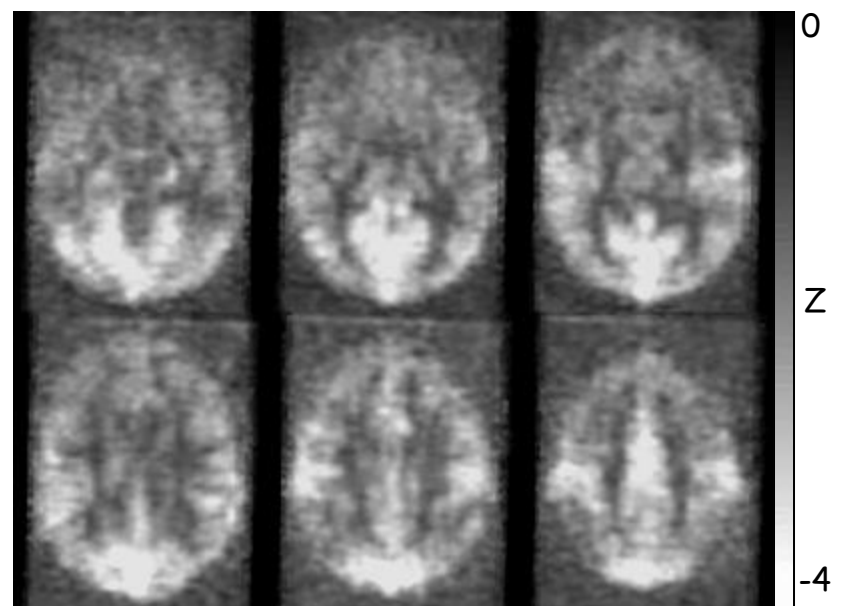
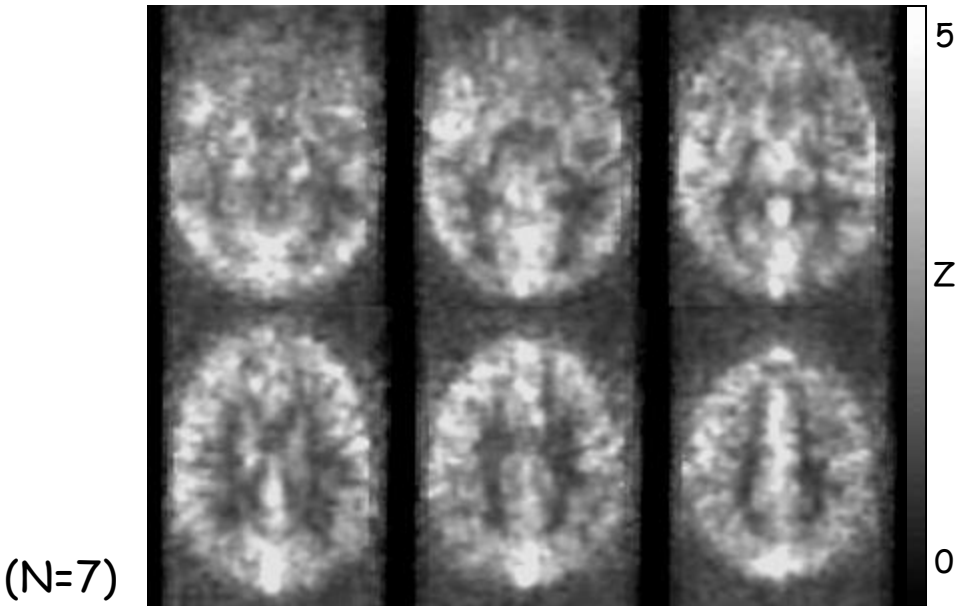
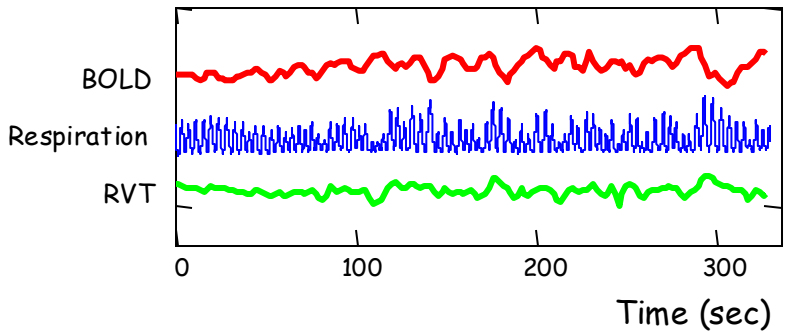
# Methodology

## Respiration induced signal changes

### Breath-holding



### Rest



R. M. Birn, J. B. Diamond, M. A. Smith, P. A. Bandettini, Separating respiratory variation-related fluctuations from neuronal activity-related fluctuations in fMRI, NeuroImage 31, 1536-1548 (2006)

# Methodology



Mapping ↔ "Reading"

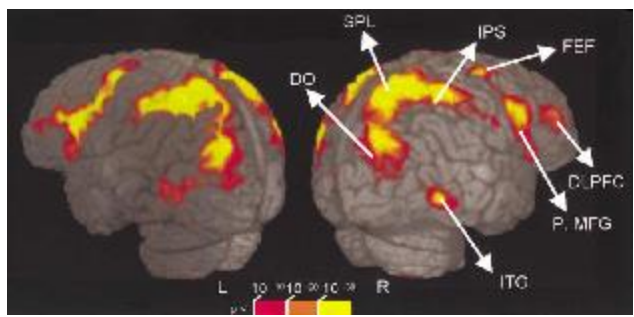


# Methodology

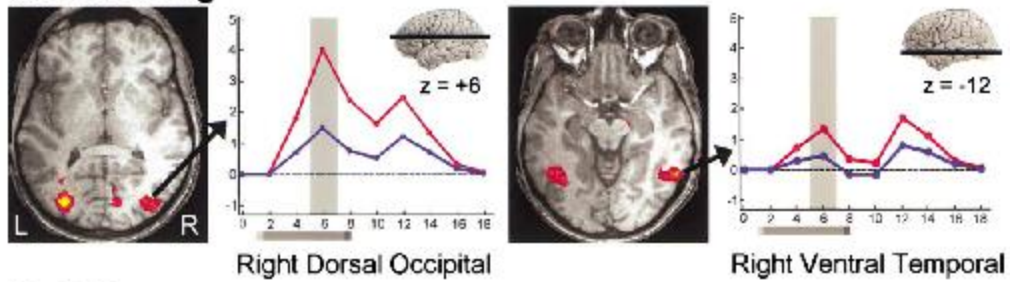
Neuron, Vol. 35, 975-987, August 29, 2002, Copyright ©2002 by Cell Press

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

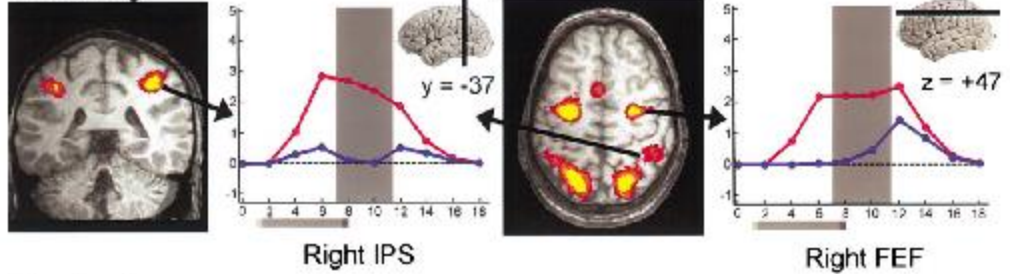
Luiz Pessoa,<sup>1</sup> Eva Gutierrez, Peter A. Bandettini, and Leslie G. Ungerleider  
 Laboratory of Brain and Cognition  
 National Institute of Mental Health  
 National Institutes of Health  
 Bethesda, Maryland 20892



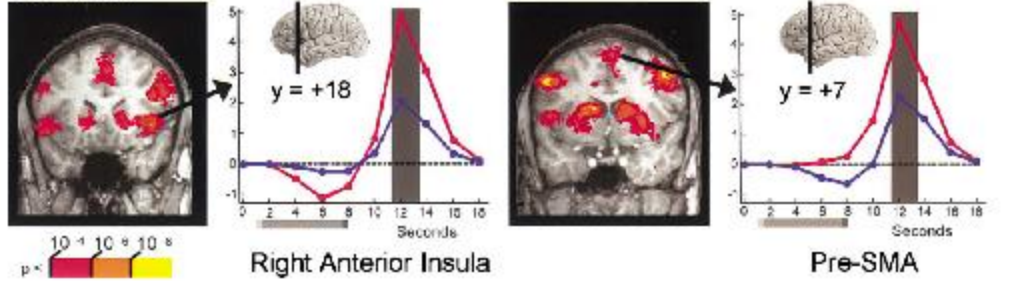
### A. Encoding



### B. Delay



### C. Test

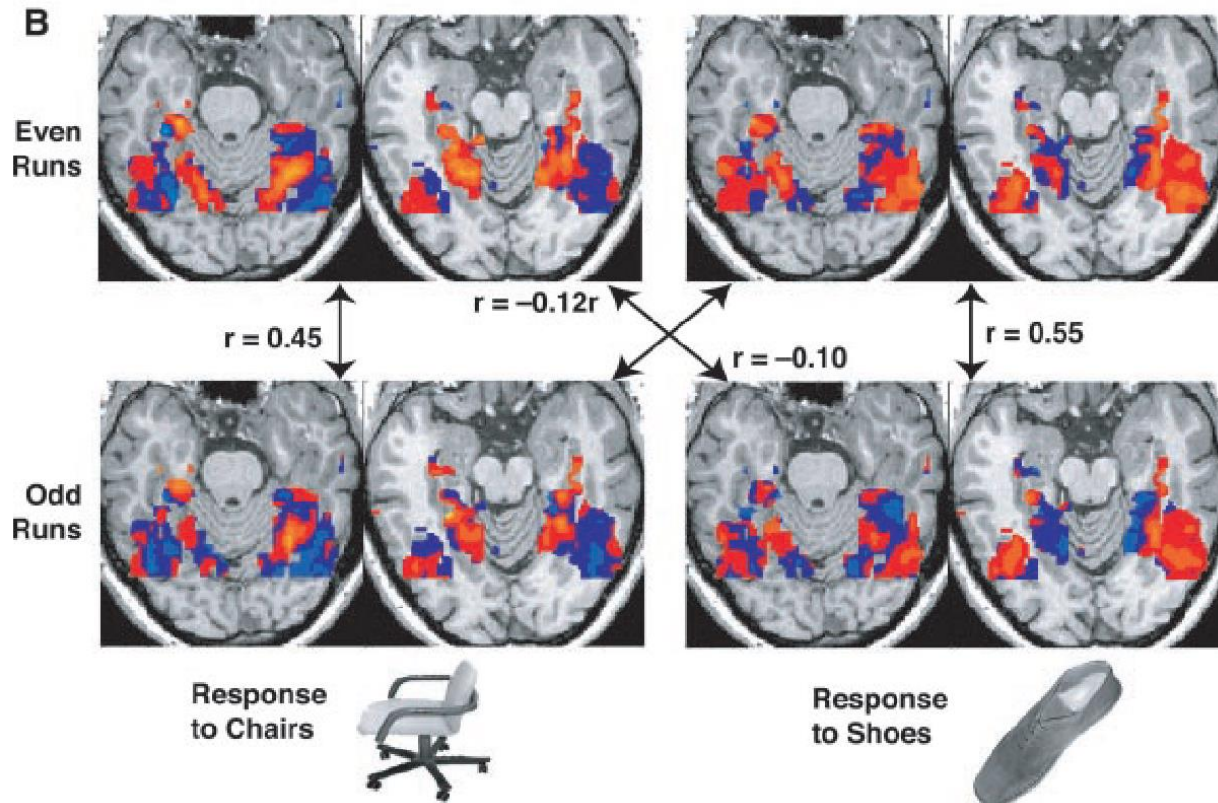


$10^{-4}$   $10^{-6}$   $10^{-8}$

# Methodology

## Ventral temporal category representations

Object categories are associated with distributed representations in ventral temporal cortex



Haxby et al. 2001



## 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>

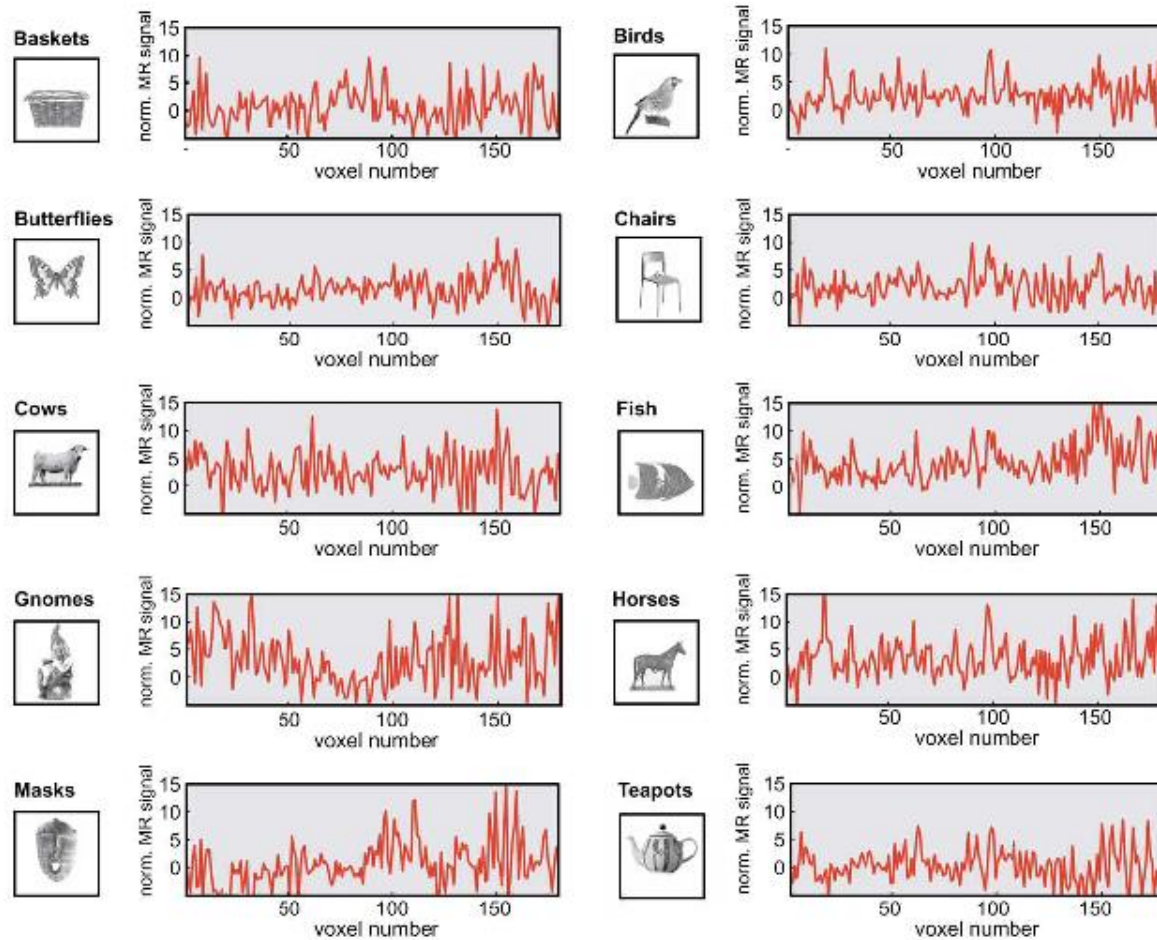
<sup>a</sup> Rowland Institute for Science, Cambridge, MA 02142, USA

<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

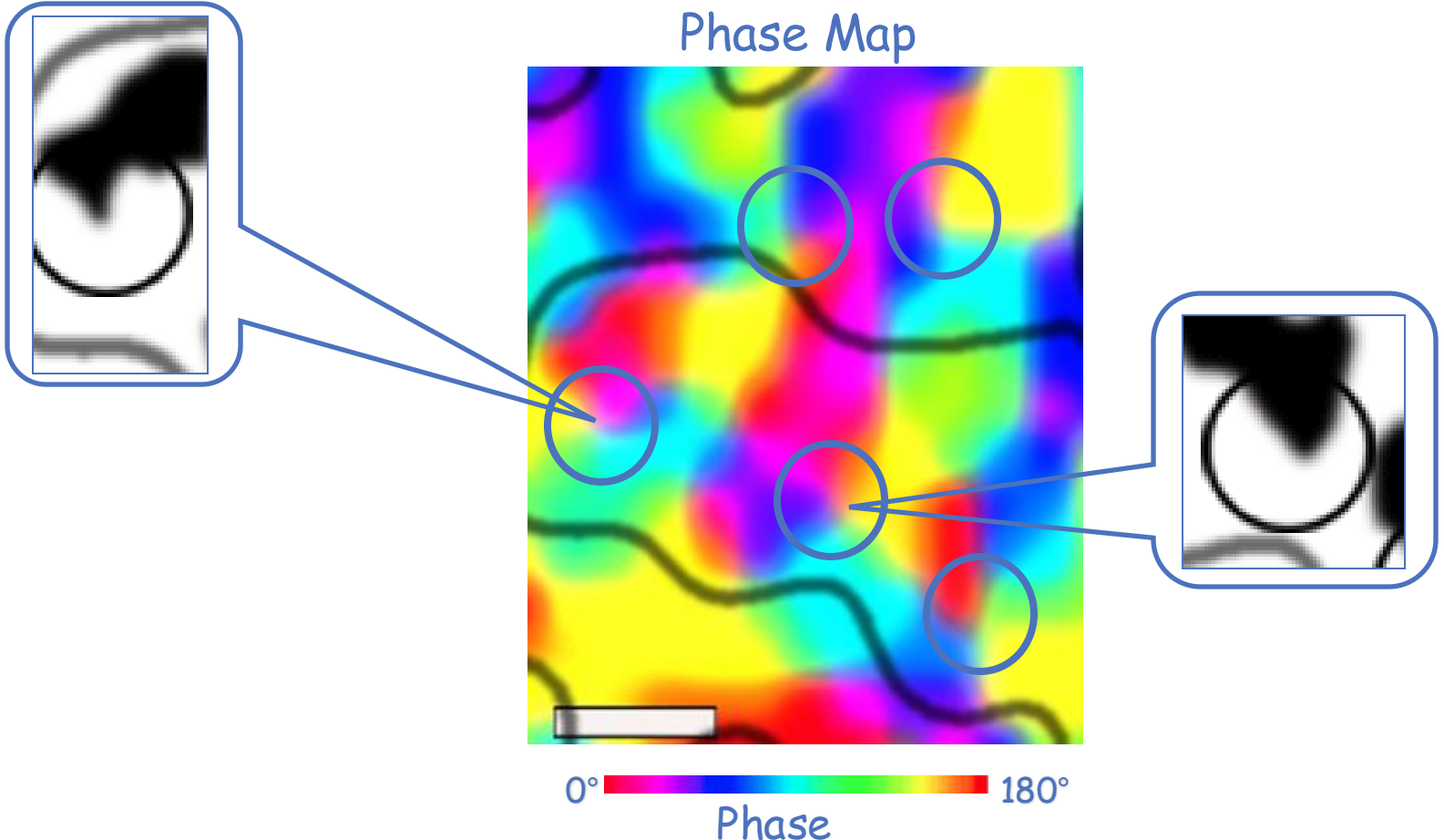
## NEUROIMAGE 19 (2): 261-270 Part 1 JUN 2003



# Methodology

## Orientation Columns in Human V1 as Revealed by fMRI at 7T

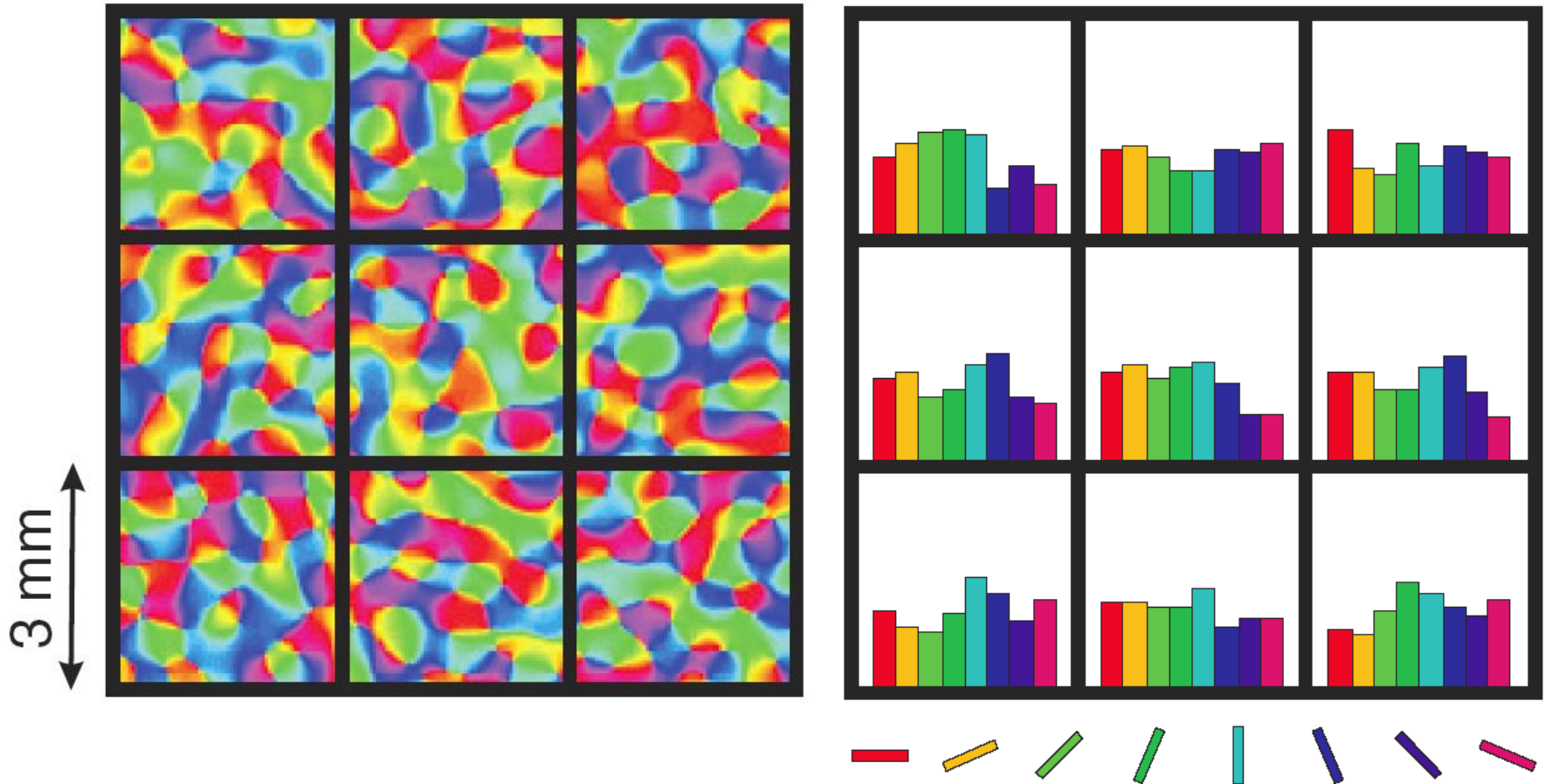
Phase Map



Yacoub, Ugurbil & Harel  
University of Minnesota / CMRR  
HBM 2006

Scalebar = 0.5 mm

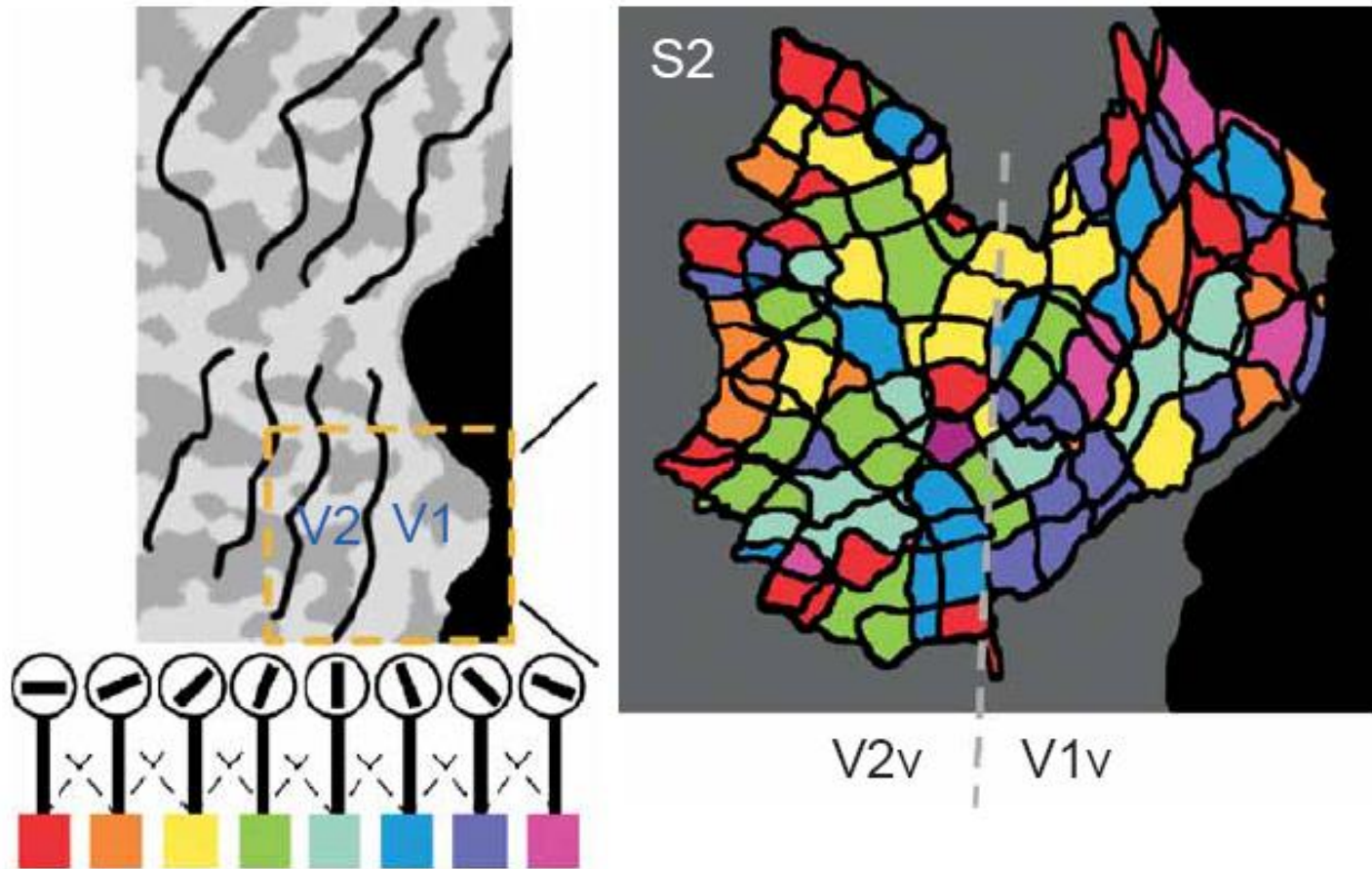
# Methodology



Boynton (2005), News & Views on Kamitani & Tong (2005) and Haynes & Rees (2005)

# Methodology

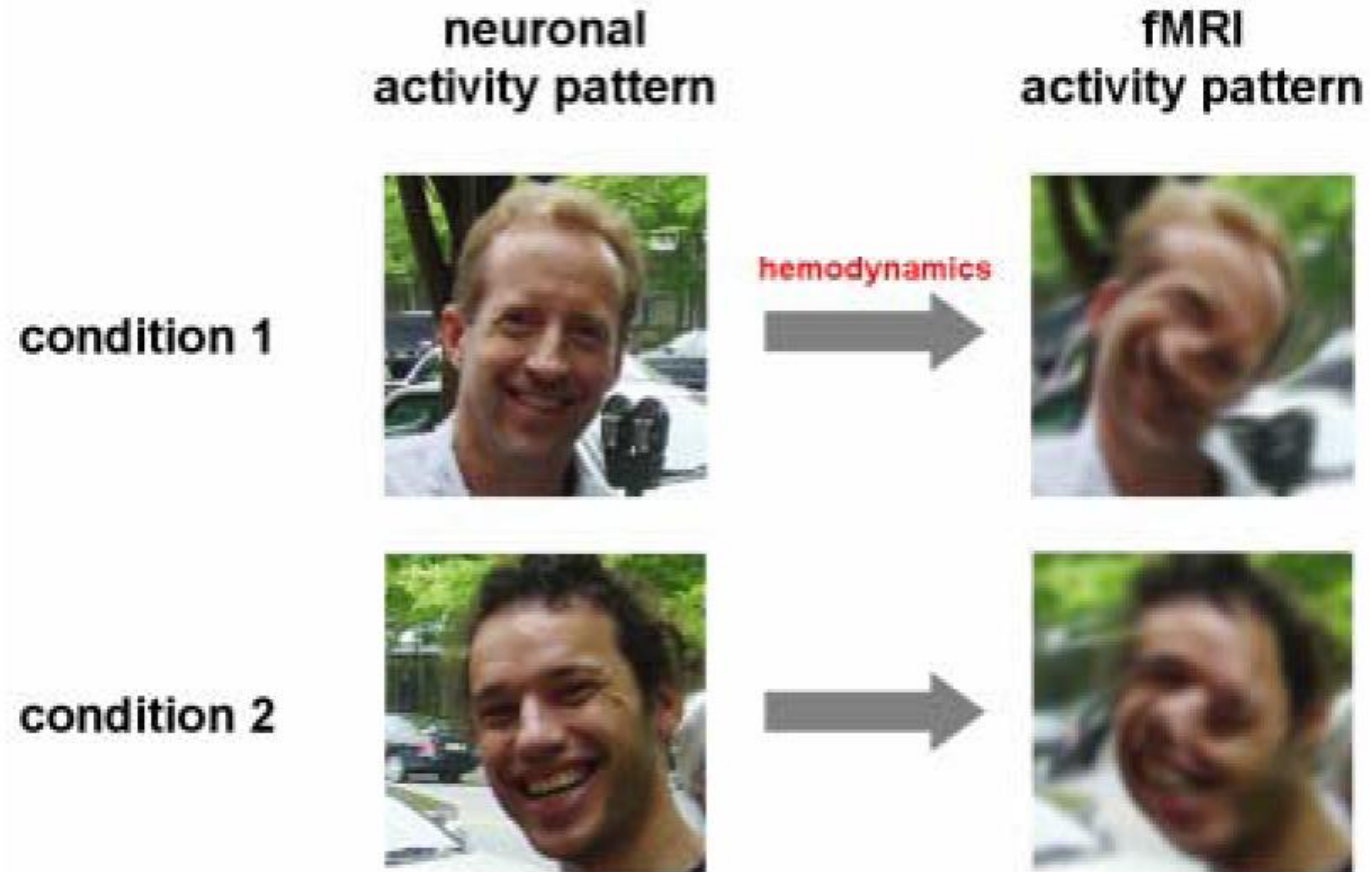
## Lower spatial frequency clumping



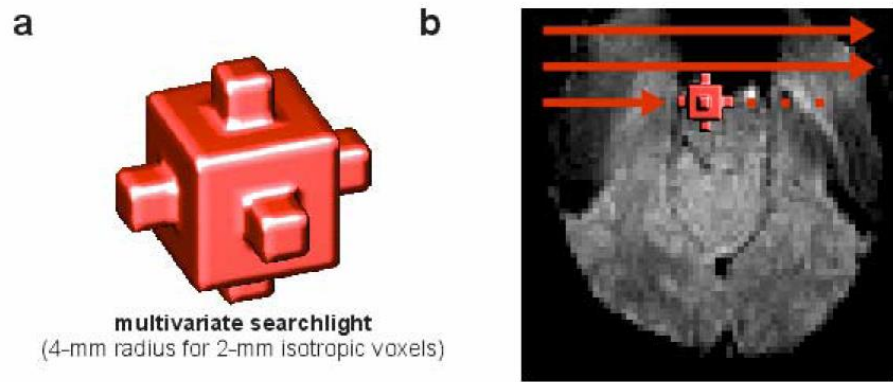
Kamitani & Tong (2005)



# Methodology



# Methodology



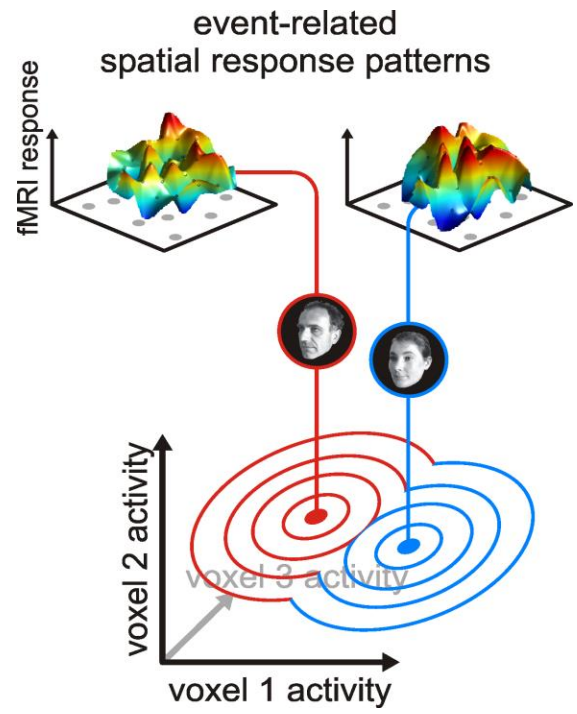
**multivariate searchlight**  
(4-mm radius for 2-mm isotropic voxels)



Information-based searchlight map with t-map texture (FDR  $q < 0.05$ )



Unsmoothed-data t map (same number of voxels marked)



N. Kriegeskorte, R. Goebel, P. Bandettini, Information-based functional brain mapping. *Proc. Nat'l. Acad. Sci. USA*, 103, 3863-3868 (2006).

# Technology

Coil arrays  
High field strength  
High resolution  
Novel functional contrast

# Methodology

Connectivity assessment  
Multi-modal integration  
Pattern classification  
Task design

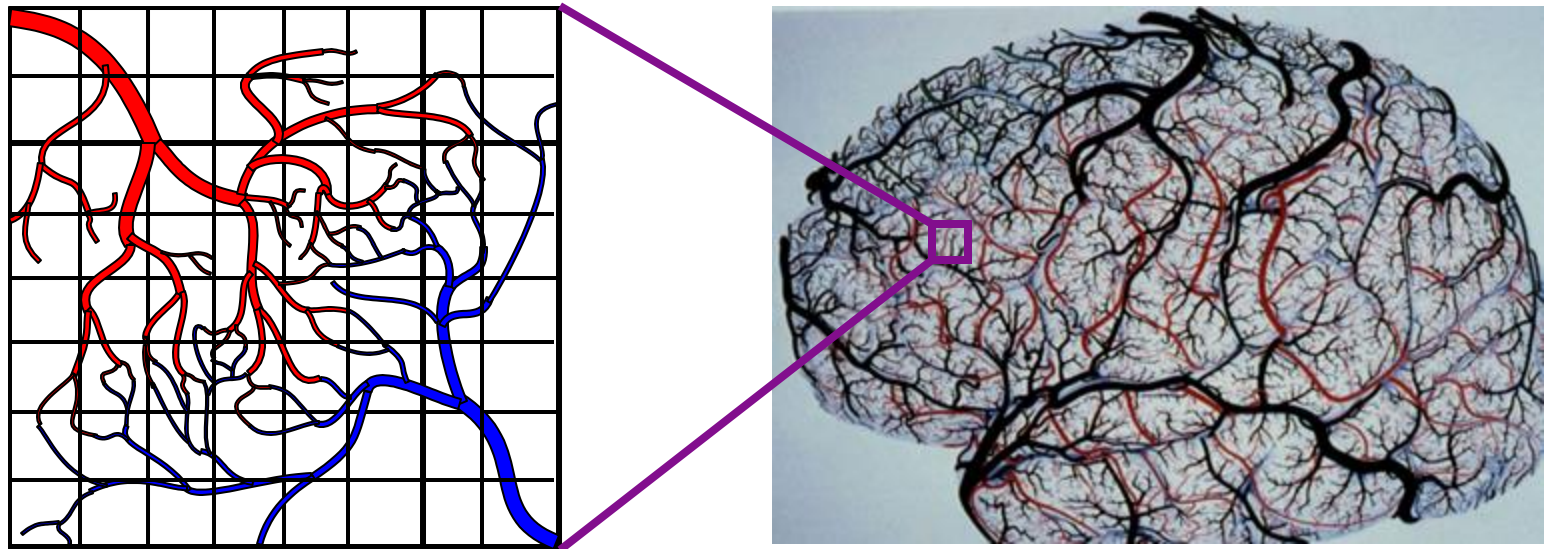
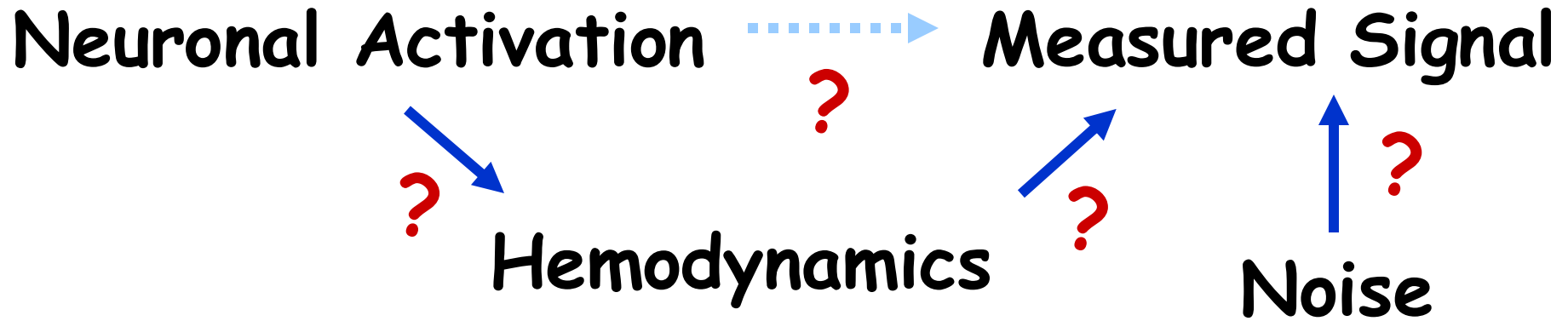
Fluctuations  
Dynamics  
Cross - modal comparison

Basic Neuroscience  
Behavior correlation/prediction  
Pathology correlation

# Interpretation

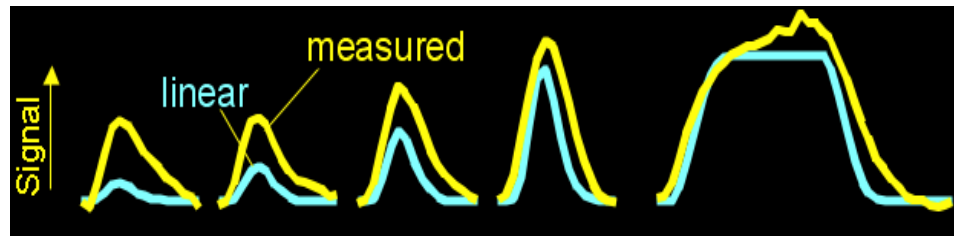
# Applications

# Interpretation

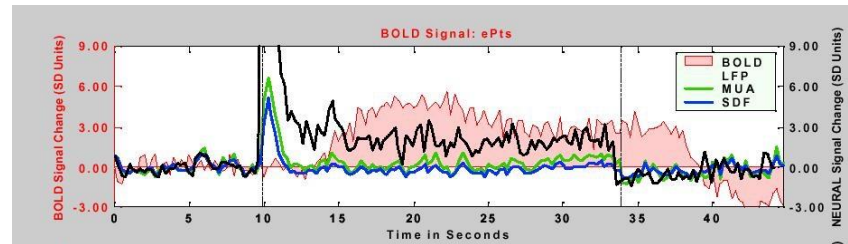




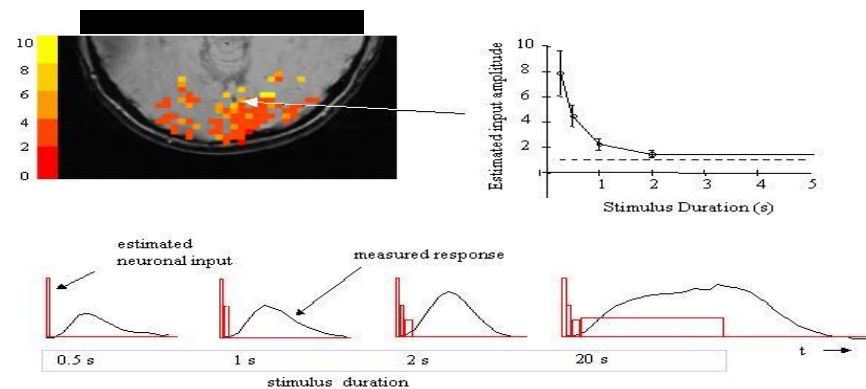
# Interpretation



R. M. Birn, (2001) *NeuroImage*, 14: 817-826.



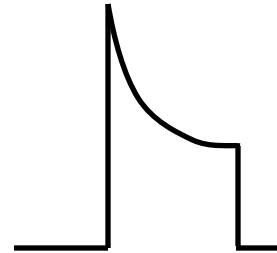
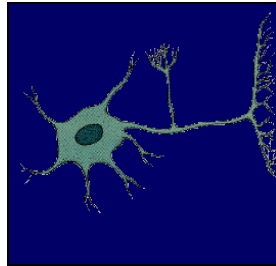
Logothetis et al. (2001) *Nature*, 412, 150-157.



P. A. Bandettini et al, (2001) *Nature Neuroscience*, 4: 864-866.

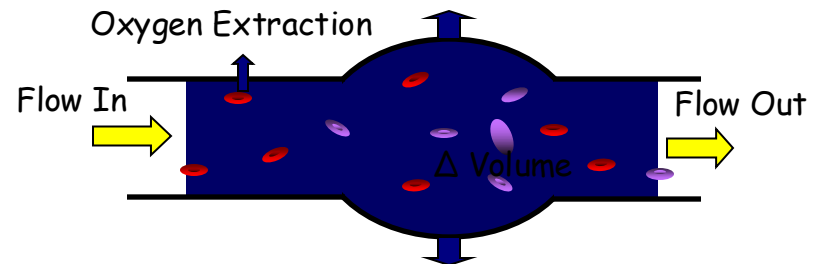
## Sources of this Nonlinearity

- Neuronal



- Hemodynamic

- Oxygen extraction
- Blood volume dynamics



# Technology

Coil arrays  
High field strength  
High resolution  
Novel functional contrast

# Methodology

Connectivity assessment  
Multi-modal integration  
Pattern classification  
Task design

Fluctuations  
Dynamics  
Cross - modal comparison

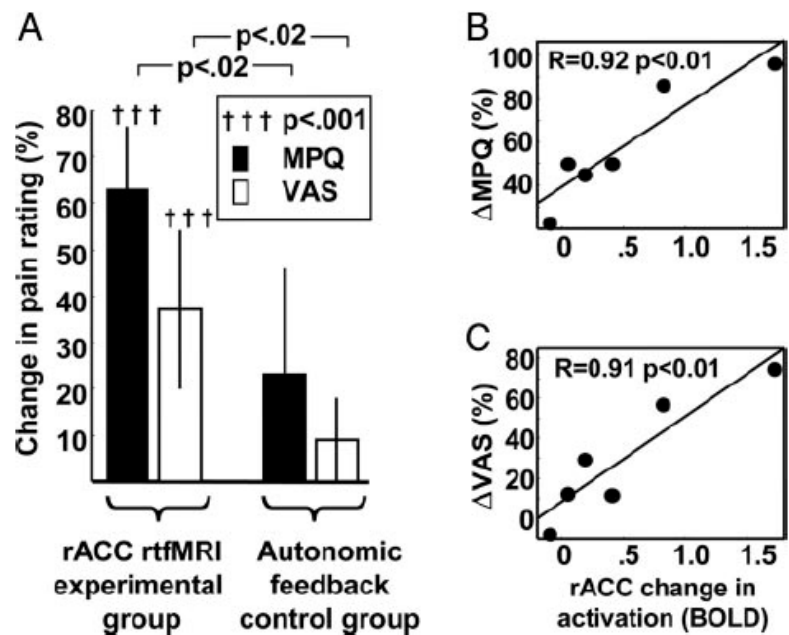
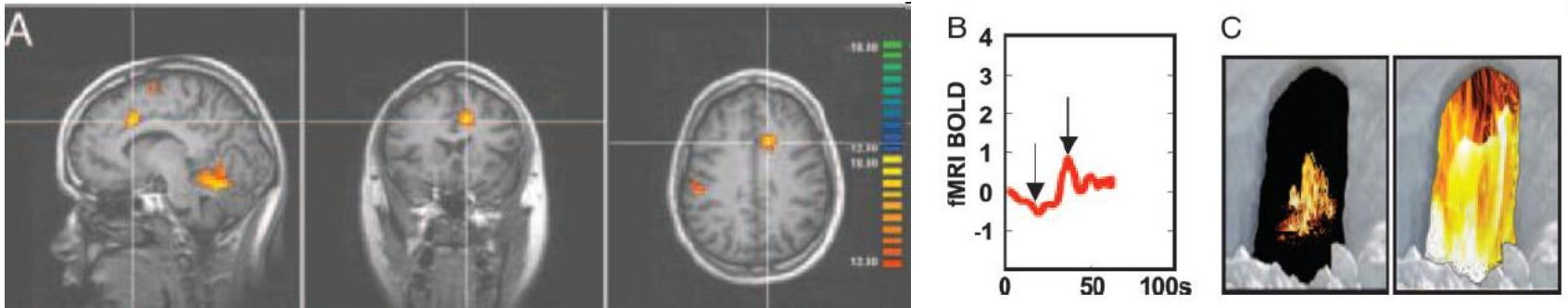
Basic Neuroscience  
Behavior correlation/prediction  
Pathology correlation

# Interpretation

# Applications

# Applications

## Real time fMRI feedback to reduce chronic pain



Control over brain activation and pain learned by using real-time functional MRI, R. C. deCharms, et al. PNAS, 102; 18626-18631 (2005)

# What fMRI Can Do (routine fMRI)

## Help in understanding healthy brain organization

- map networks involved with specific behavior, stimulus, or performance
- characterize changes over time (seconds to years)
- determine correlates of behavior (response accuracy, etc...)

## Current Clinical Applications

- presurgical mapping (CPT code in place as of Jan, 2007)

## Current Clinical Research

- assessment of recovery and plasticity
- clinical population characterization with probe task or resting state

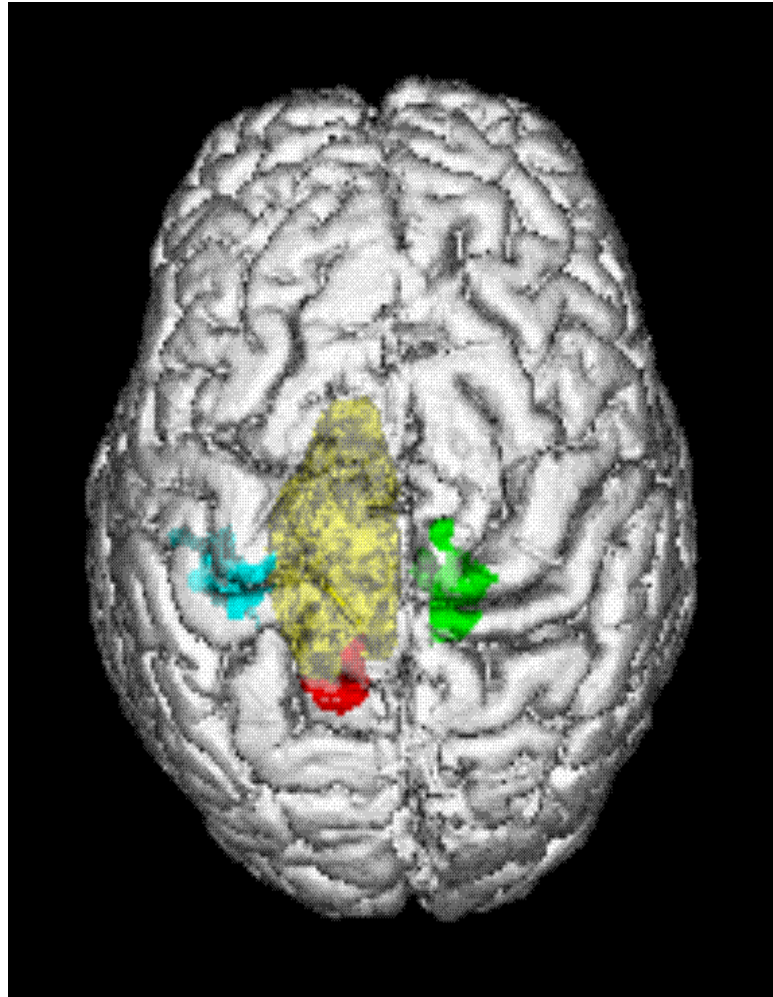
# Presurgical Mapping

Left Foot

Tumor

Right Foot

Right Hand



# What fMRI Can't Do (what are the problems with fMRI?)

- Too low SNR for routine clinical use (takes too long)
- Requires patient cooperation (too sensitive to motion)
- Too low spatial resolution (each voxel has several million neurons)
- Too low temporal resolution (hemodynamics are variable and sluggish)
- Too indirectly related to neuronal activity
- Too many physiologic variables influence signal
- Requires a task (BOLD cannot look at baseline maps)
- Too confined space and high acoustic noise.

# What fMRI Might Do

## Complementary use for clinical diagnoses

- utilization of clinical research results for diagnoses
- prediction of pathology

## Clinical treatment and assessment of therapy

- better understanding mechanism of pathology for focused therapy
- drug effect assessment
- assessment of therapy progress, biofeedback
- epileptic foci mapping
- neurovascular physiology assessment

## Non clinical uses

- lie detection
- prediction of behavior tendencies
- brain/computer interface



# Section on Functional Imaging Methods & Functional MRI Facility Jan 19, 2007



Back row: **Wenming Luh**, **Niko Kriegeskorte**, **Rasmus Birn**, **Tyler Jones**, **Sean Marrett**  
Middle row: **Jon West**, **Kay Kuhns**, **Anthony Boemio**, **Peter Bandettini**, **Joey Dunsmoor**, **Doug Ruff**, **Kevin Murphy**  
Front row: **Dorian Van Tassel**, **Jerzy Bodurka**, **Adam Thomas**, **Marieke Mur**, **David Knight**