

Seventeen Years of Functional MRI

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&

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



1. A bit of history of MRI and functional MRI
2. Basics of functional MRI contrast and methods
3. Basic and cutting edge applications

Magnetic Resonance Imaging

1984

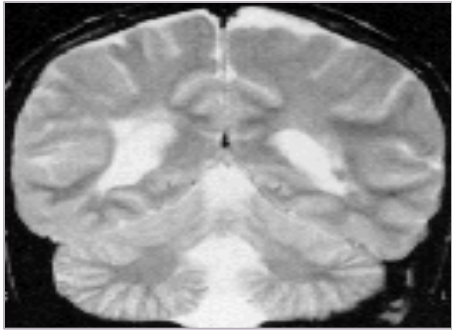


Water: 42 MHz/Tesla

1.5 Tesla = 63 MHz

3 Tesla = 126 MHz

7 Tesla = 294 MHz



MRI Images with Different Contrast Weighting

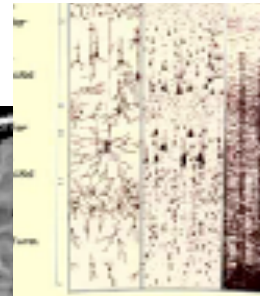
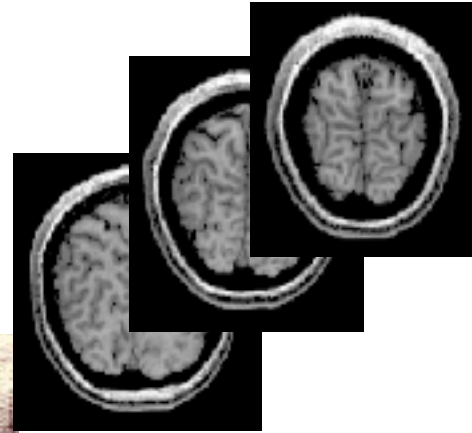
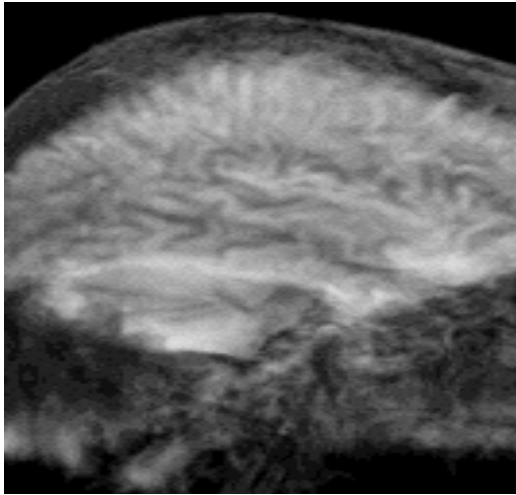
T1 Weighted

T2 Weighted

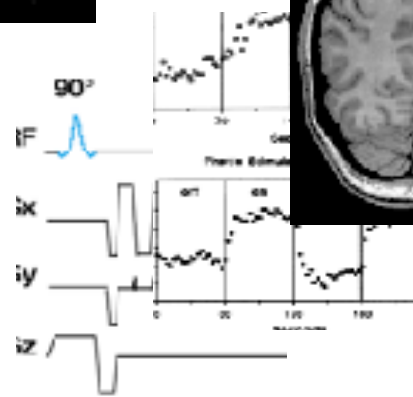
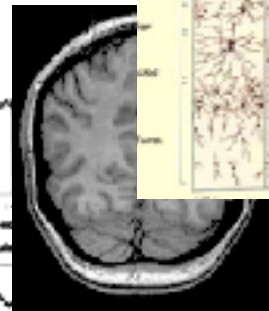
Venography



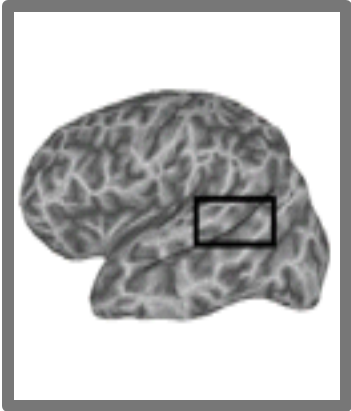
Fiber Track Imaging



Anatomy



Angiography



Perfusion

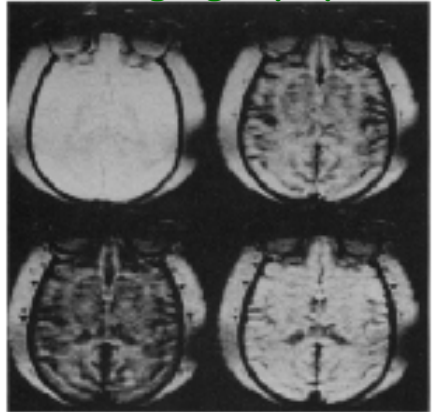
- 1. A bit of history of MRI and functional MRI**
- 2. Basics of functional MRI contrast and methods**
- 3. Basic and cutting edge applications**

metabolic imaging (NAA)

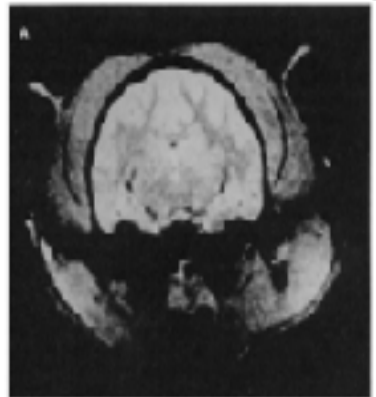


(1990) Science, 250, 53-61.

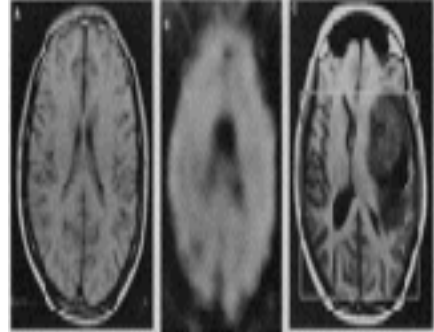
angiography



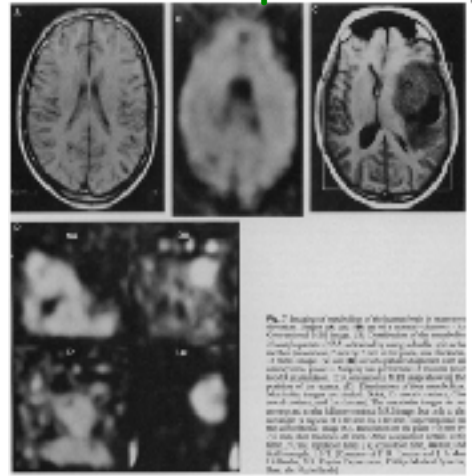
Diffusion



magnetization transfer



Gadolinium perfusion



NAA	choline
creatine	lactate

Magnetic Properties of Blood

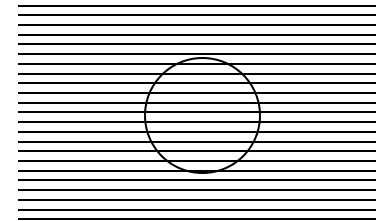
L. Pauling, C. D. Coryell, *Proc. Natl. Acad. Sci. USA* 22, 210-216, **1936**.

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

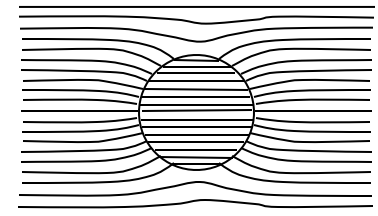
S. Ogawa, T. M. Lee, A. R. Kay, D. W. Tank, *Proc. Natl. Acad. Sci. USA* 87, 9868-9872, **1990**.

Turner, R., Lebihan, D., Moonen, C. T. W., Despres, D. & Frank, J. *Magnetic Resonance in Medicine*, 22, 159-166, **1991**.

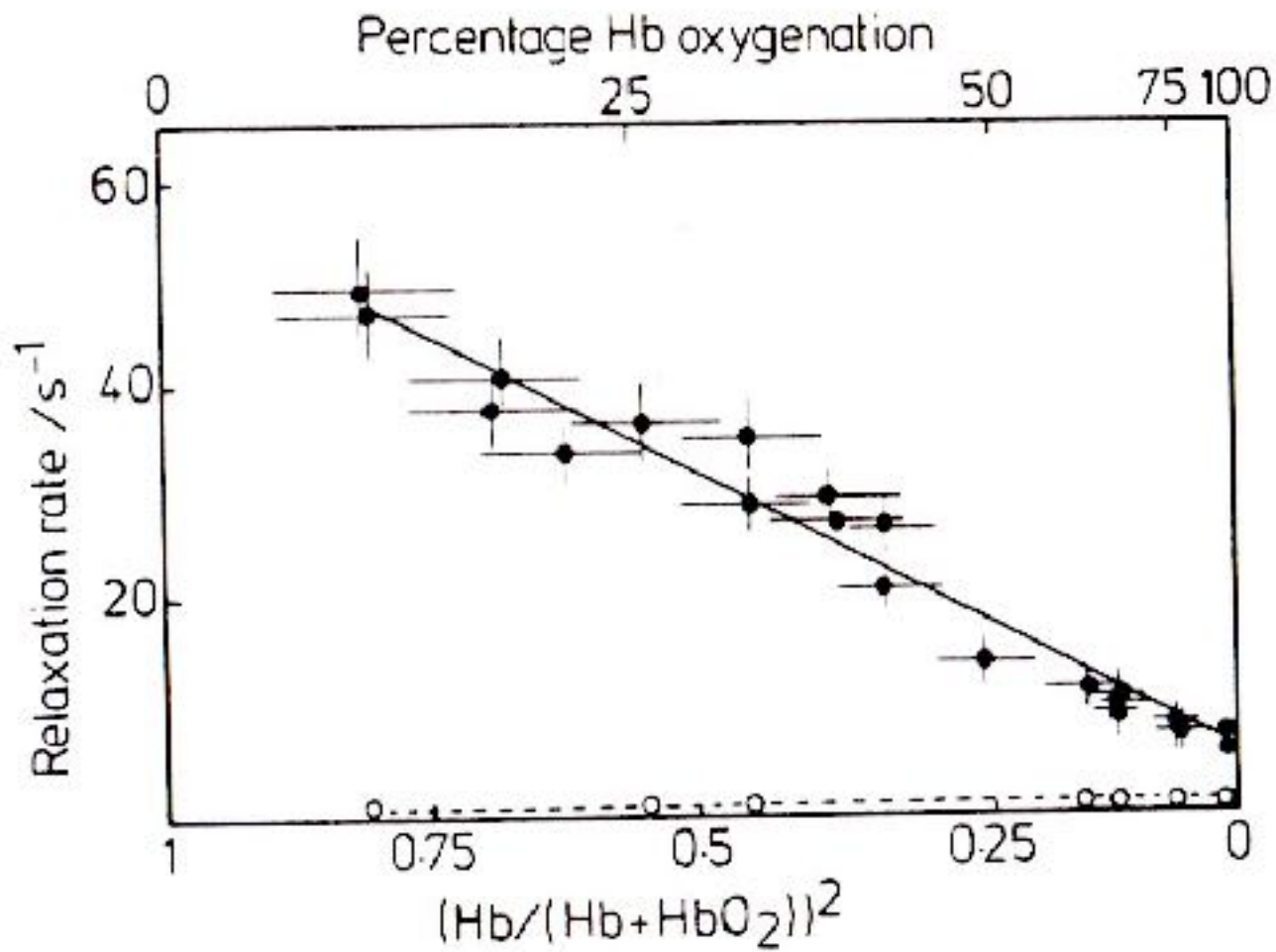
oxygenated



deoxygenated



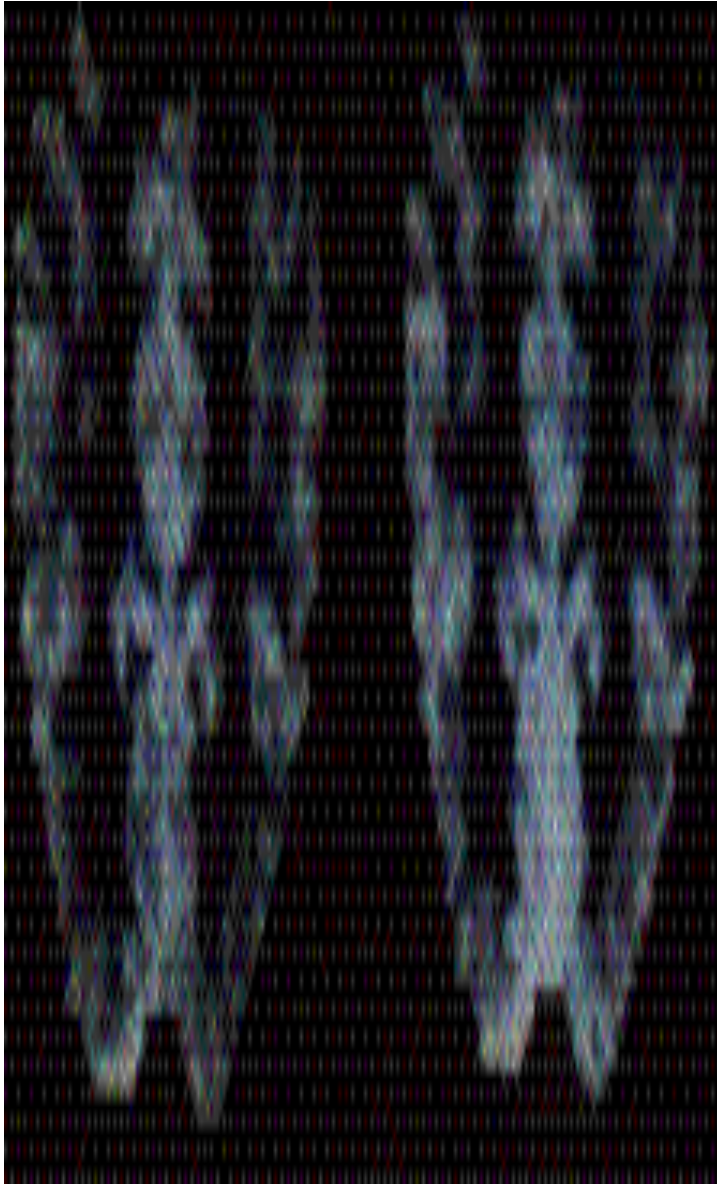
red blood cells



K. R. Thulborn, J. C. Waterton, P. M. Matthews, G. K. Radda, *Biochimica et Biophysica Acta*, 714, 265-270, 1982

in vivo

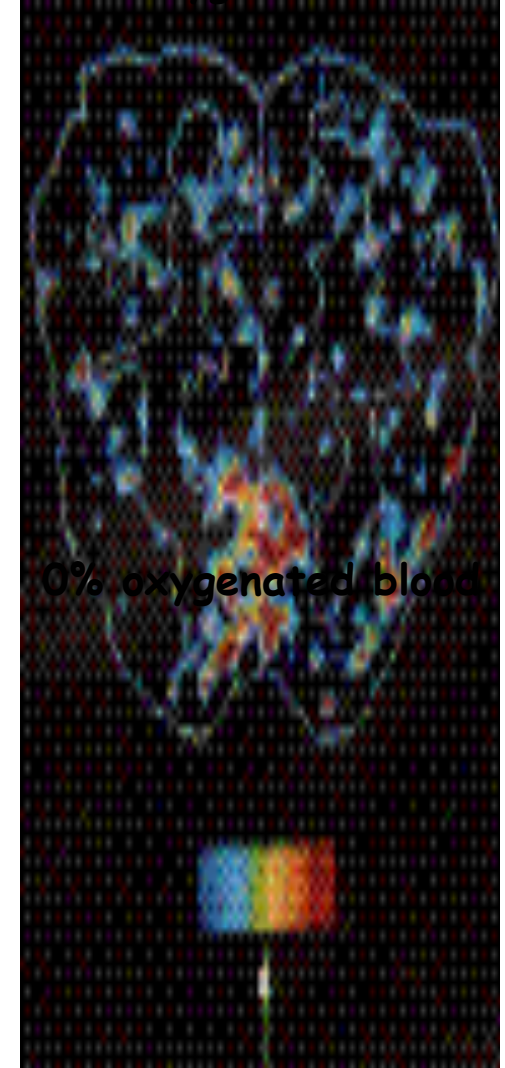
100% O₂



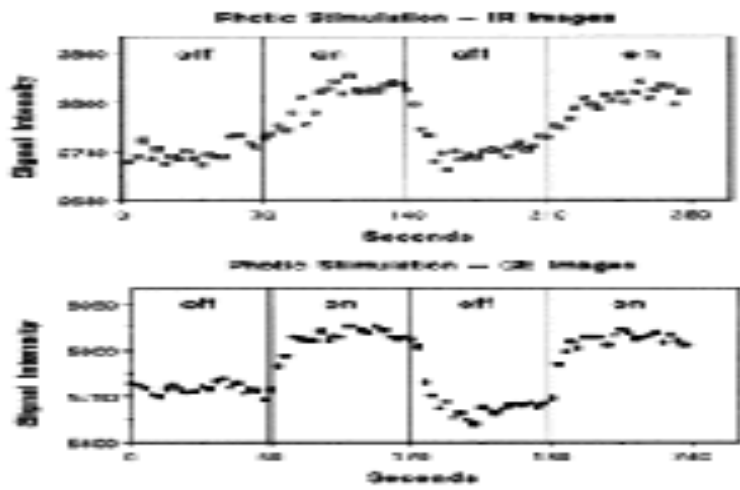
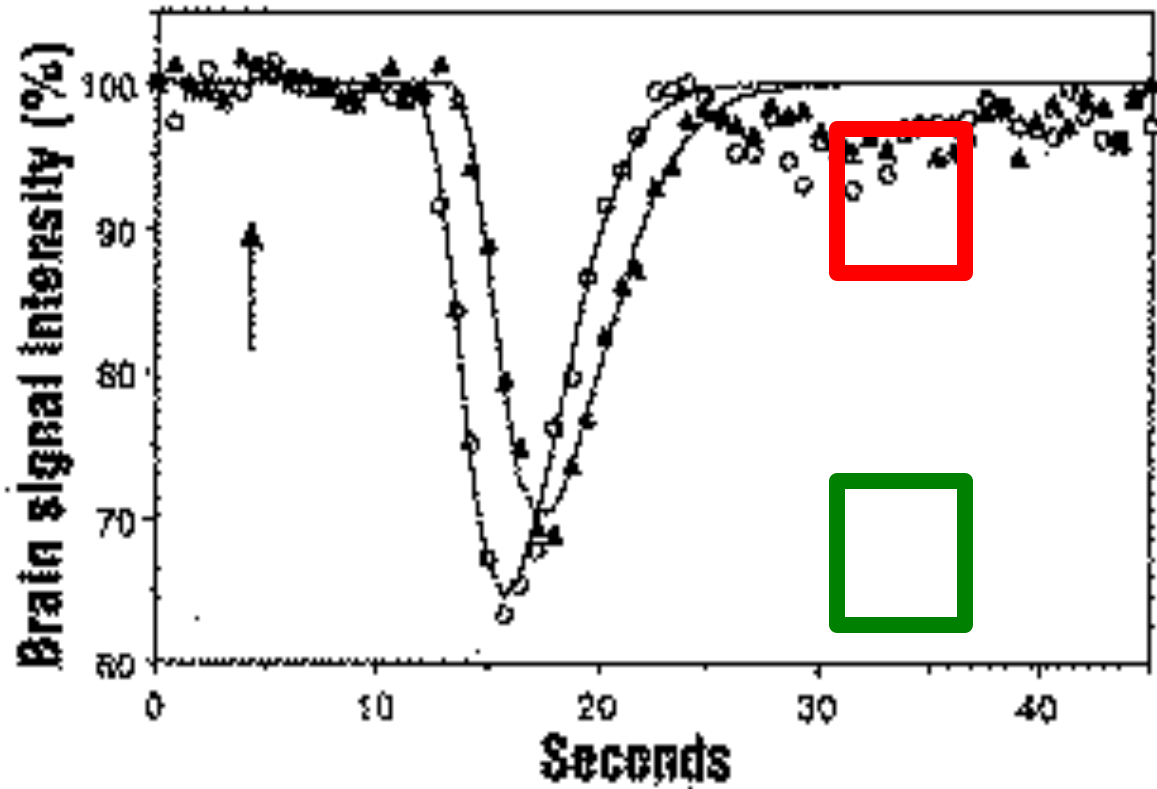
20% O₂

in vitro

100% oxygenated blood



0% oxygenated blood

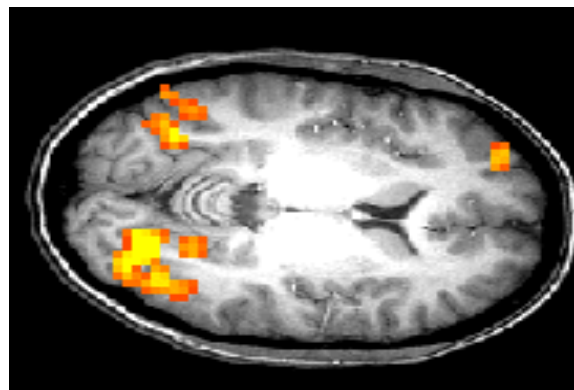
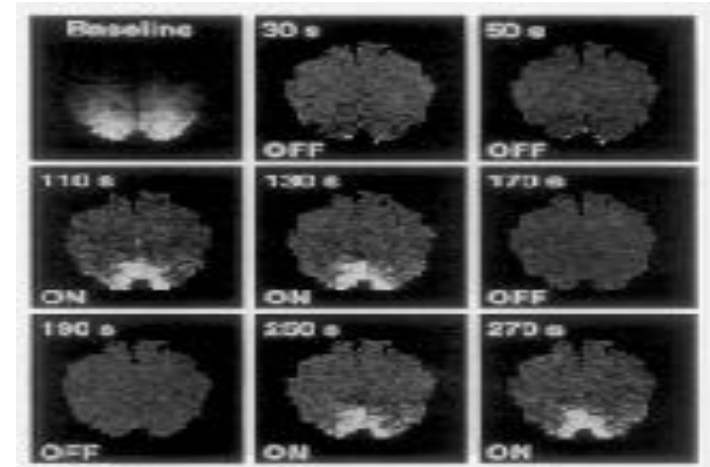


R. Turner, D. LeBihan, C.T.W. Moonen, D. Despres, J. Frank, Magn. Reson. Med, 22, 159-166 (1991)

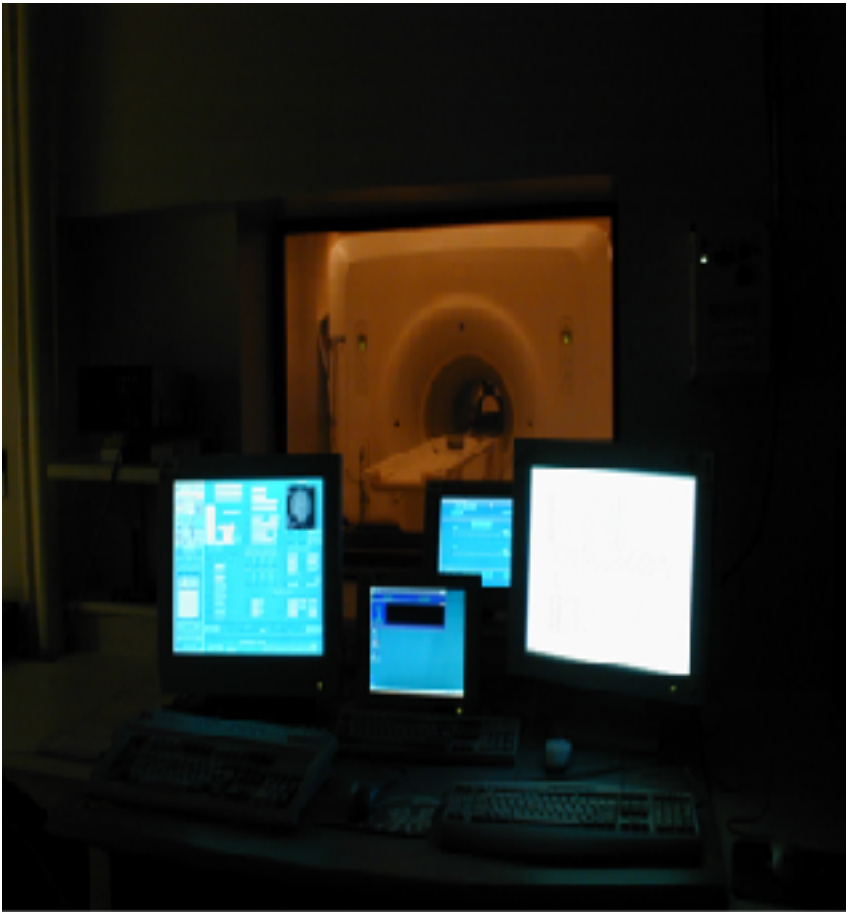
1. A bit of history of MRI and functional MRI
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Blood Volume Imaging

Susceptibility Contrast agent bolus injection and time series collection of T2* or T2 - weighted images

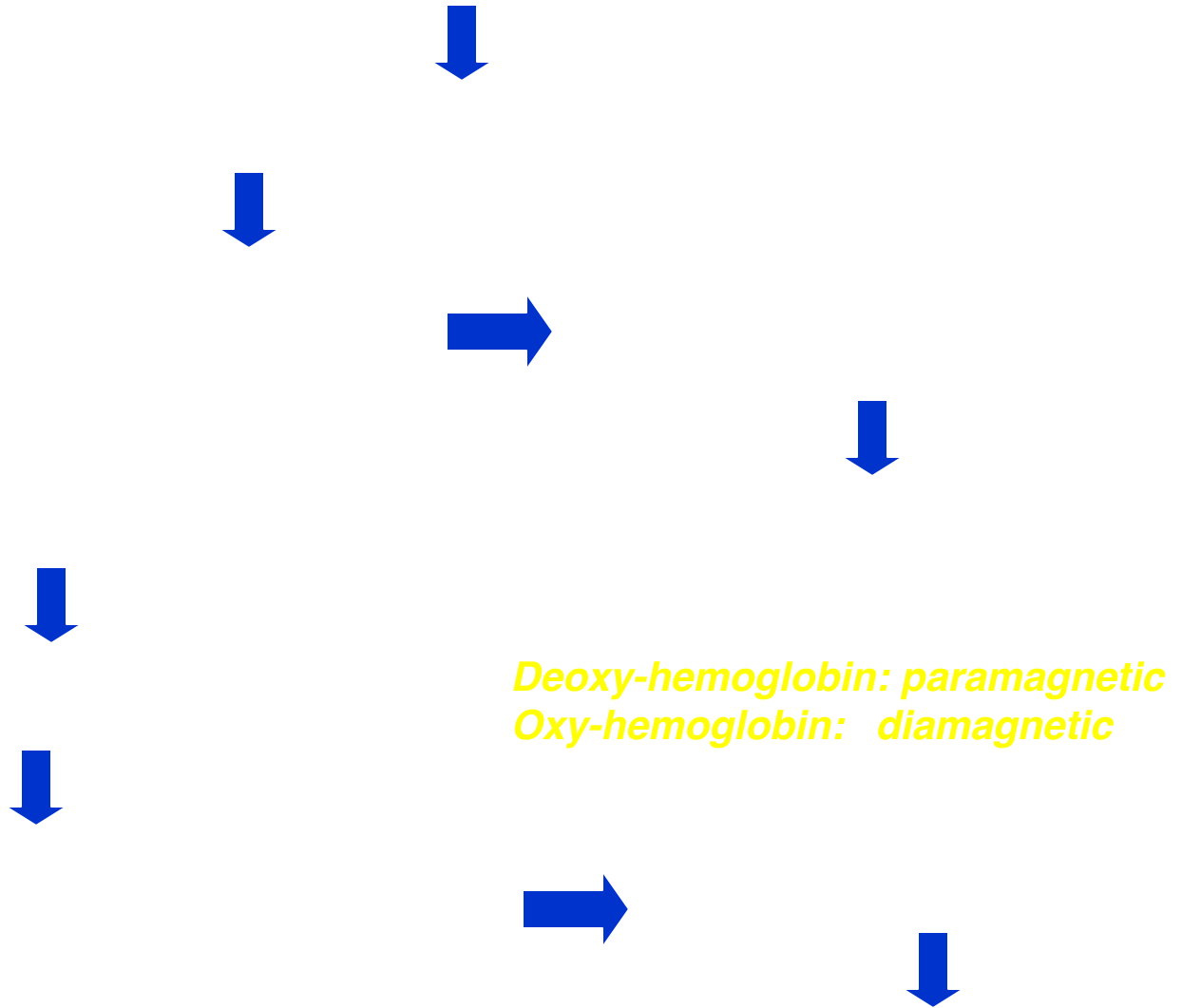


Kwong et al.

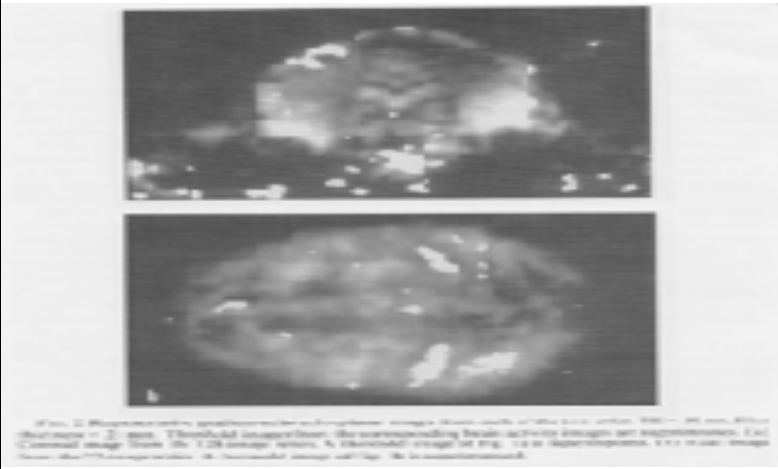
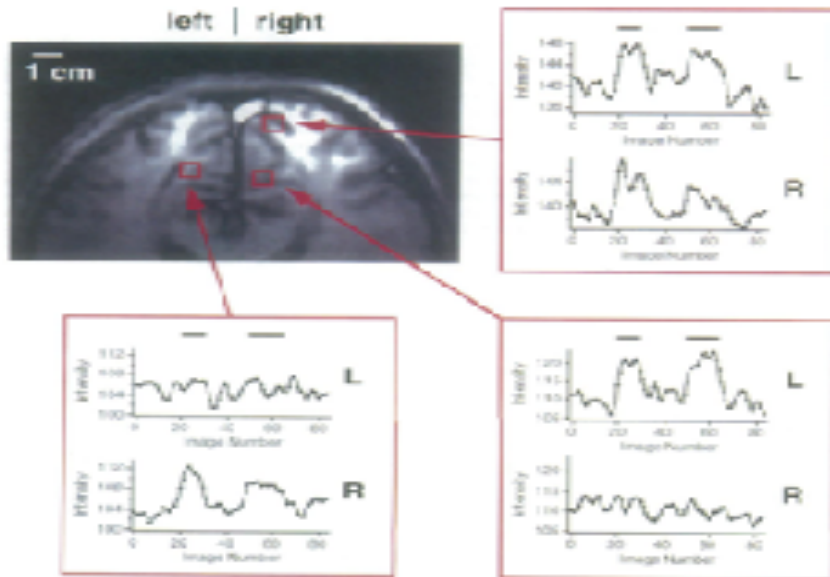


K. K. Kwong, et al, (1992) "Dynamic magnetic resonance imaging of human brain activity during primary sensory stimulation." Proc. Natl. Acad. Sci. USA. 89, 5675-5679.

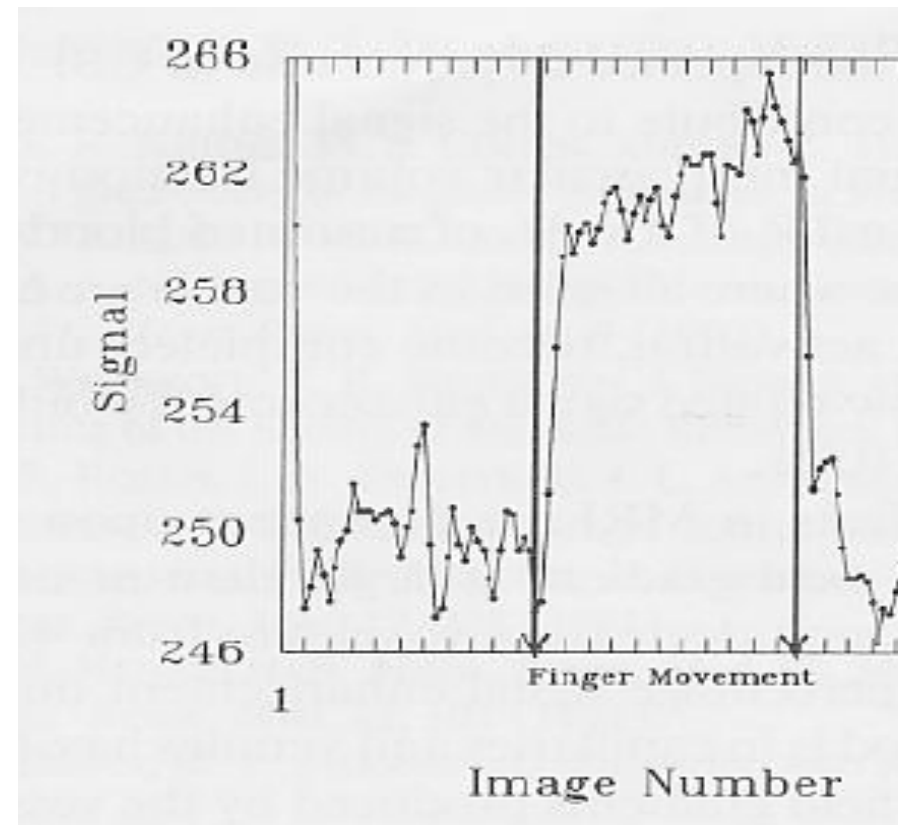
BOLD Contrast in the Detection of Neuronal Activity



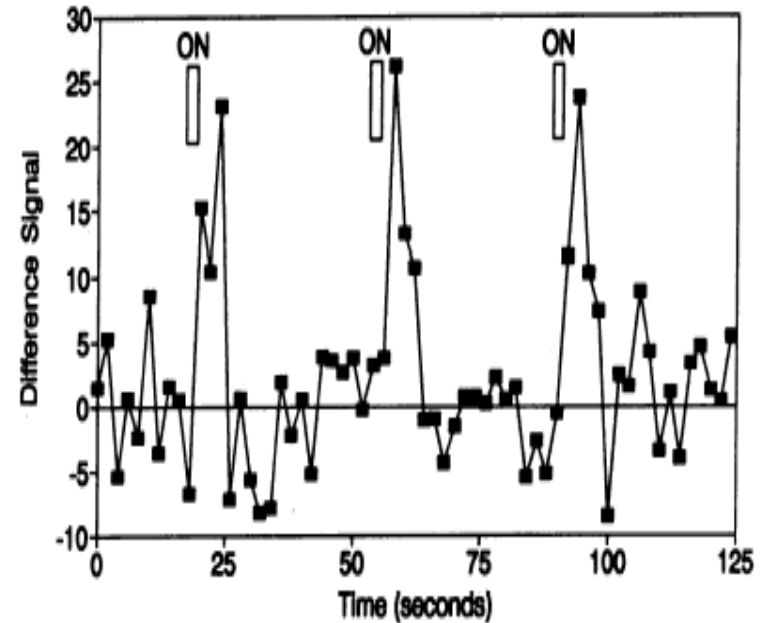
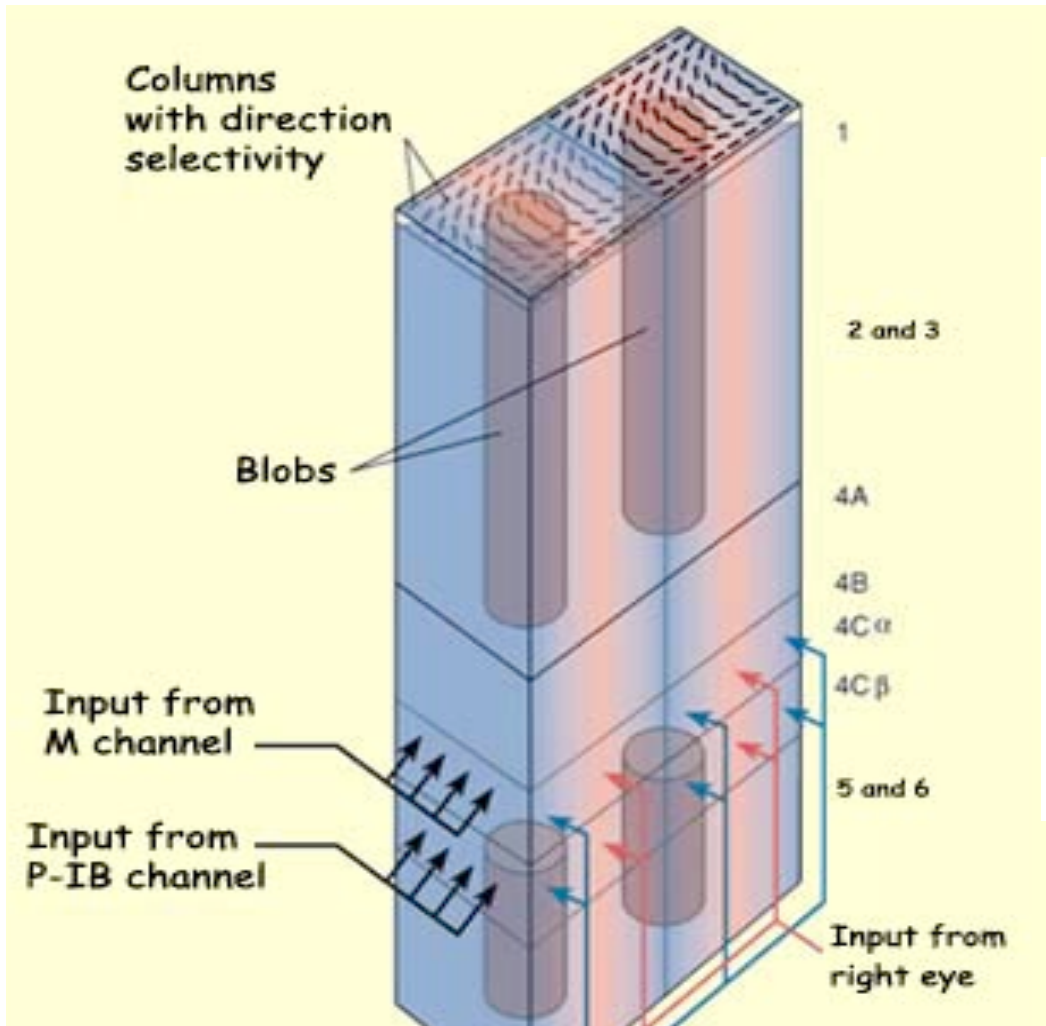
Ogawa et al.



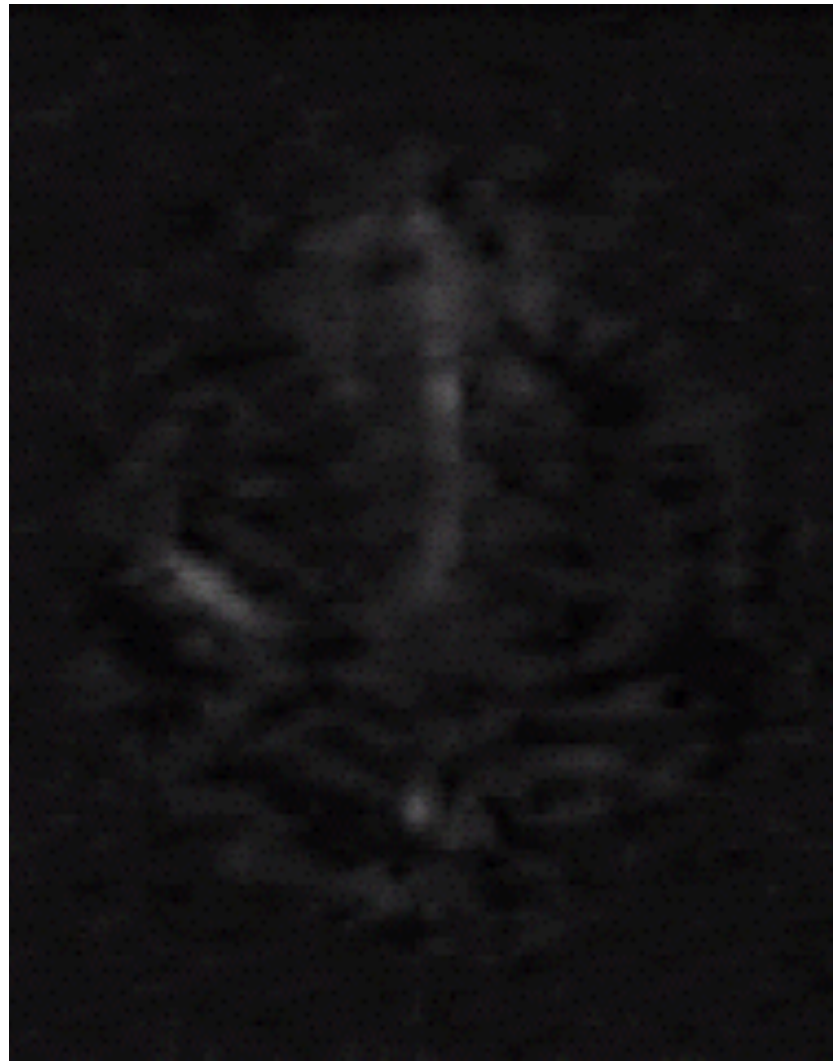
S. Ogawa, et al., (1992) "Intrinsic signal changes accompanying sensory stimulation: functional brain mapping with magnetic resonance imaging. Proc. Natl. Acad. Sci. USA." 89, 5951-5955.



Bandettini et al.



P. A. Bandettini, et al., (1992) "Time course EPI of human brain function during task activation." *Magn. Reson. Med* 25, 390-397.



1991

Blamire et al.



·Blamire, A. M., et al. (1992). "Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging." Proc. Natl. Acad. Sci. USA 89: 11069-11073.

1992...Perfusion using Arterial Spin Labeling

- 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.

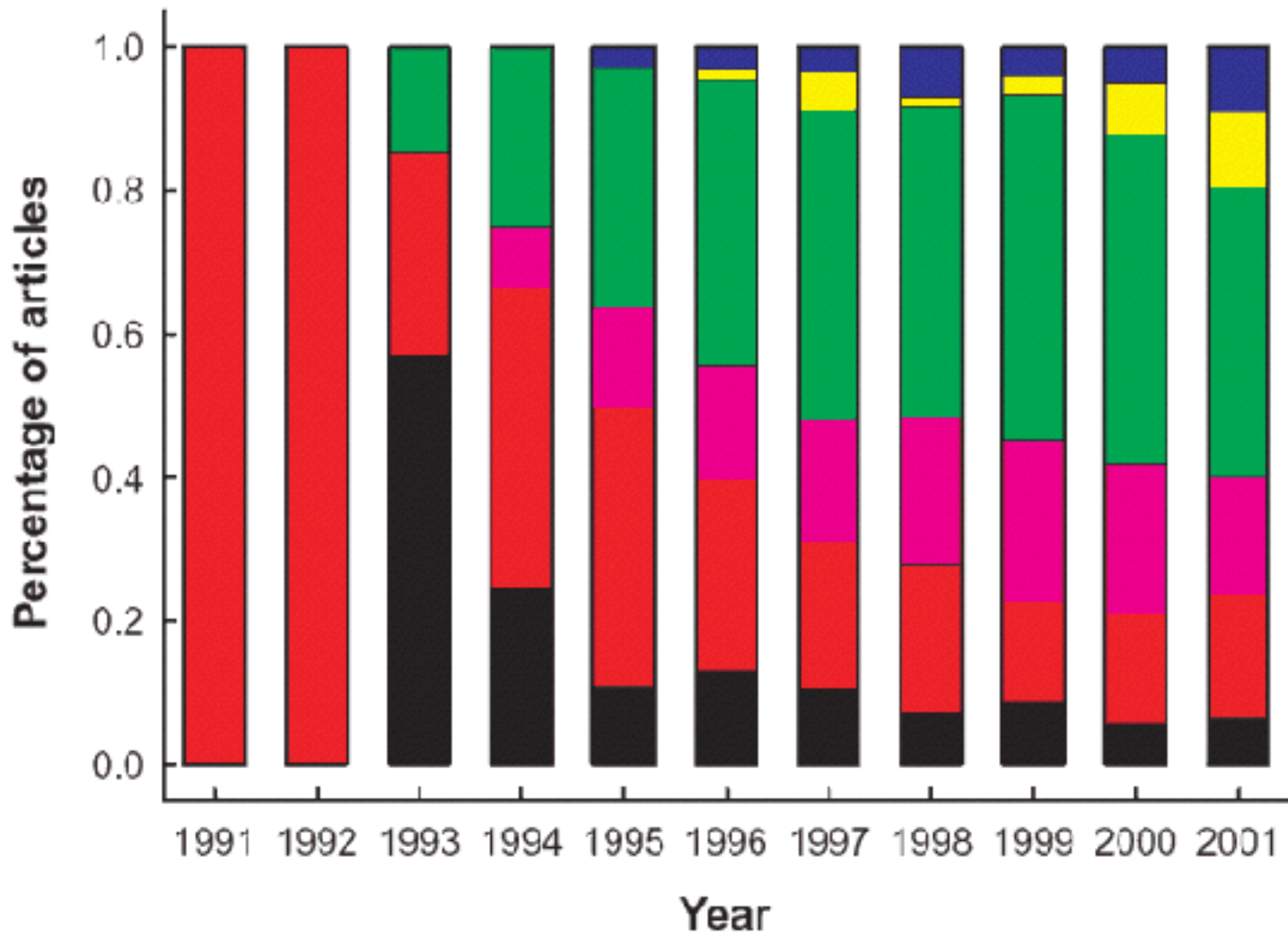
Contrast in Functional MRI

- Blood Volume
- Blood Oxygenation Changes
- Blood Perfusion

fMRI Contrast

- Volume (gadolinium)
- BOLD
- Perfusion (ASL)
- $\Delta CMRO_2$
- Δ Volume (VASO)
- Neuronal Currents
- Diffusion coefficient
- Temperature

Scopus: **Articles** or **Reviews** 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

Log Size (mm)

Brain

Map

Column

Layer

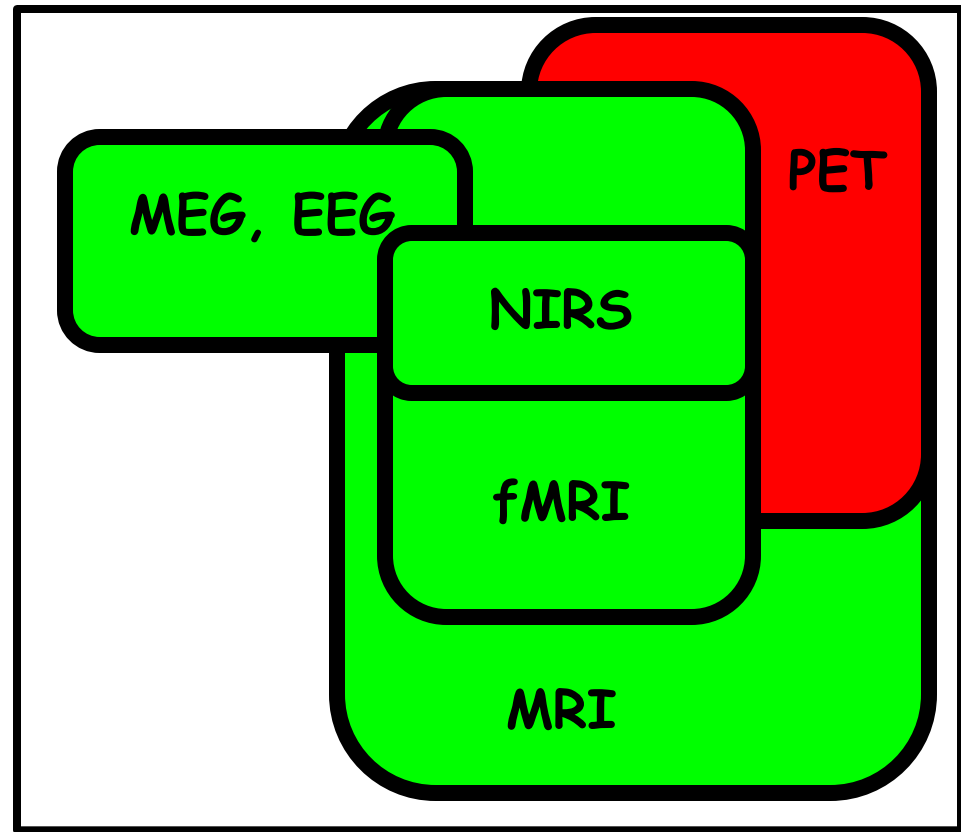
3

2

1

0

-1



-3 -2 -1 0 1 2 3 4 5 6 7

Millisecond Second Minute Hour Day

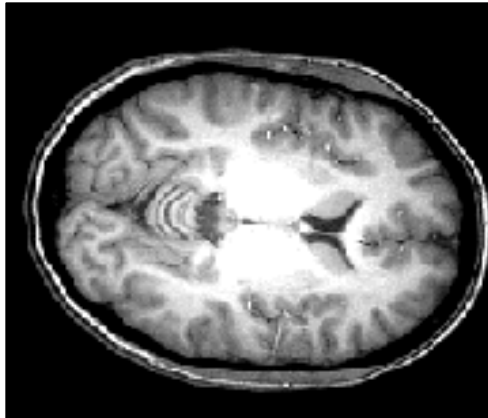
Log Time (sec)

Topics Studied with fMRI at the NIH

- Epilepsy
- Visual processing
- Mood disorders
- Learning
- Habituation
- Plasticity
- Motor Function
- Auditory processing
- Attention
- Language
- Speech
- Stroke
- Social Interaction
- Development
- Aging

MRI vs. fMRI

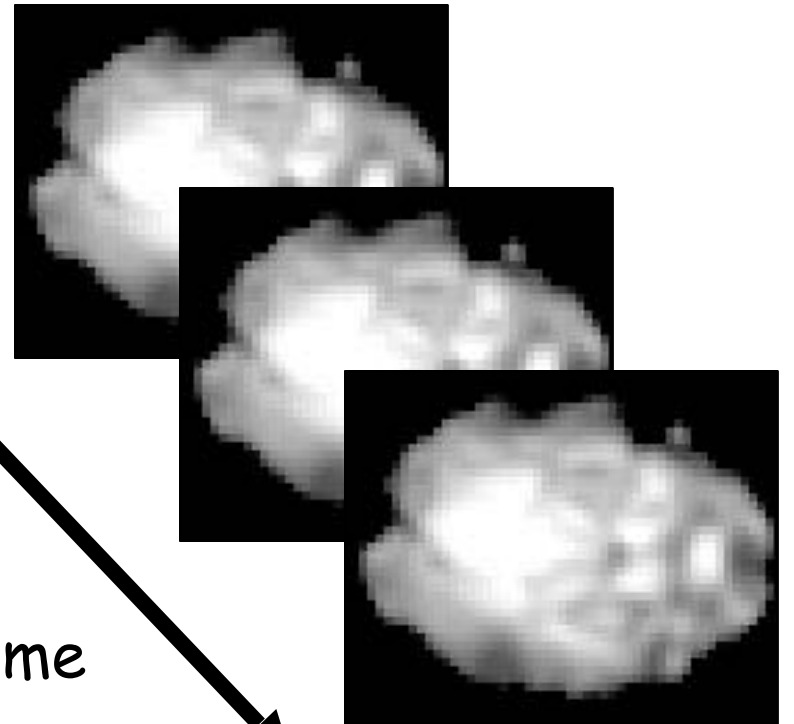
MRI



one image

high resolution
(1 mm or less)

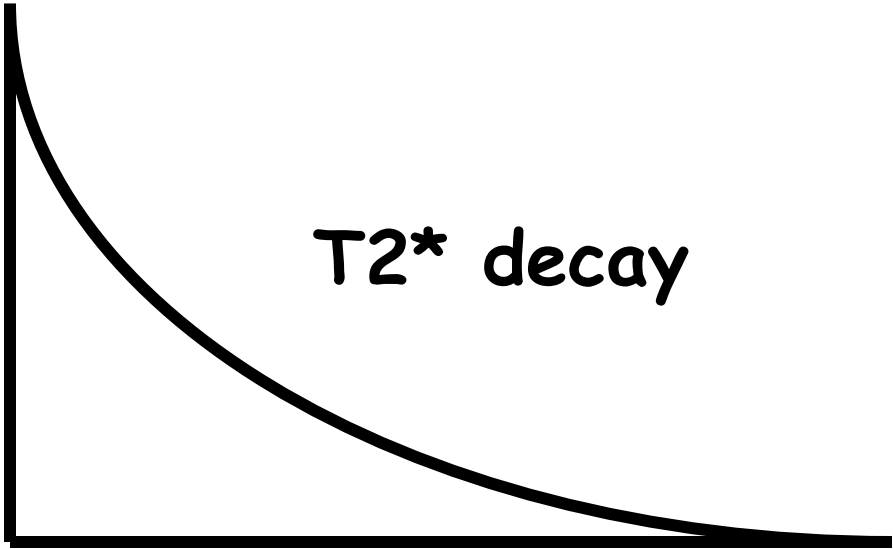
fMRI



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

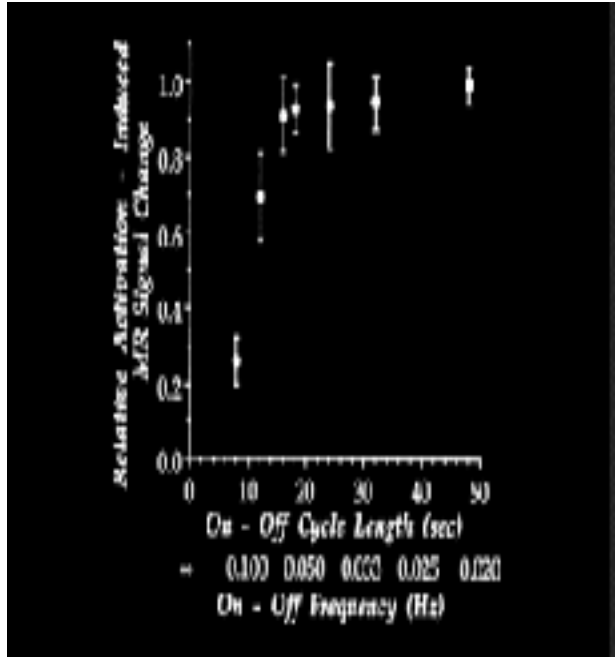
low resolution
(1.5 to 4 mm)

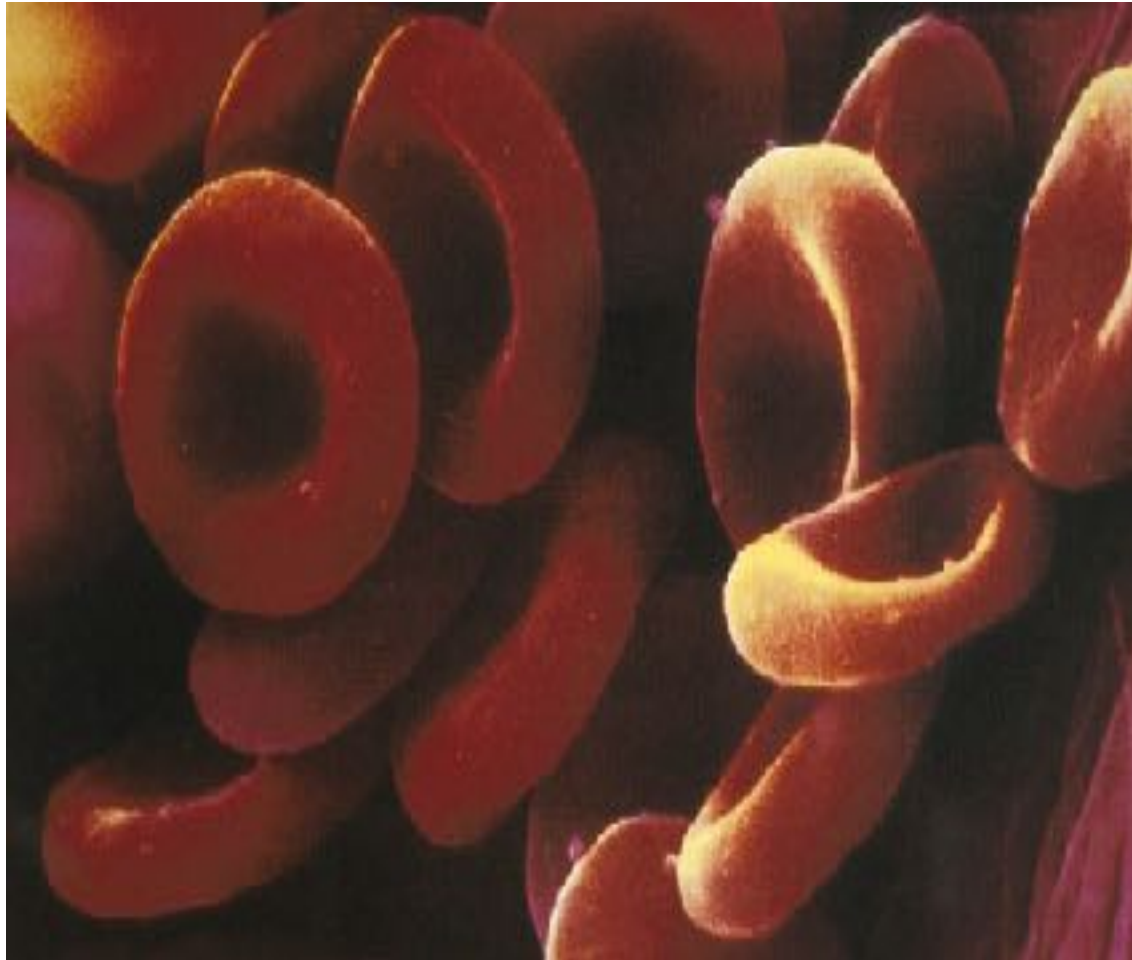
Single Shot Echo Planar Imaging (EPI)



EPI Readout Window

≈ 20 to 40 ms

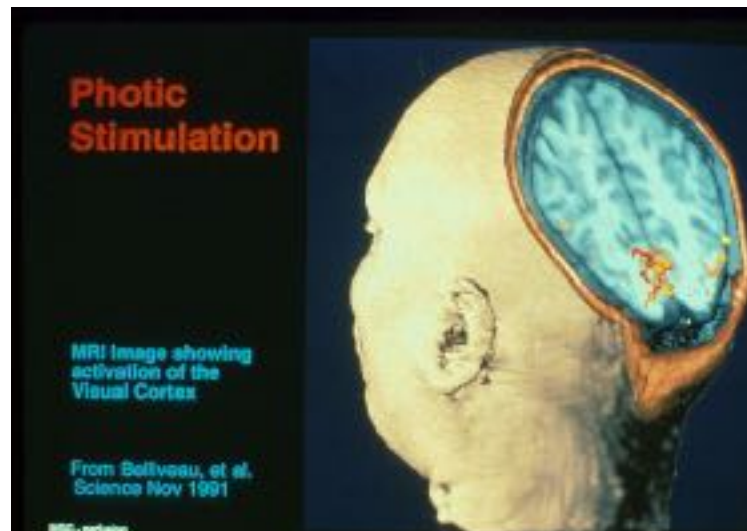




August, 1991

1991-1992

1992-1999



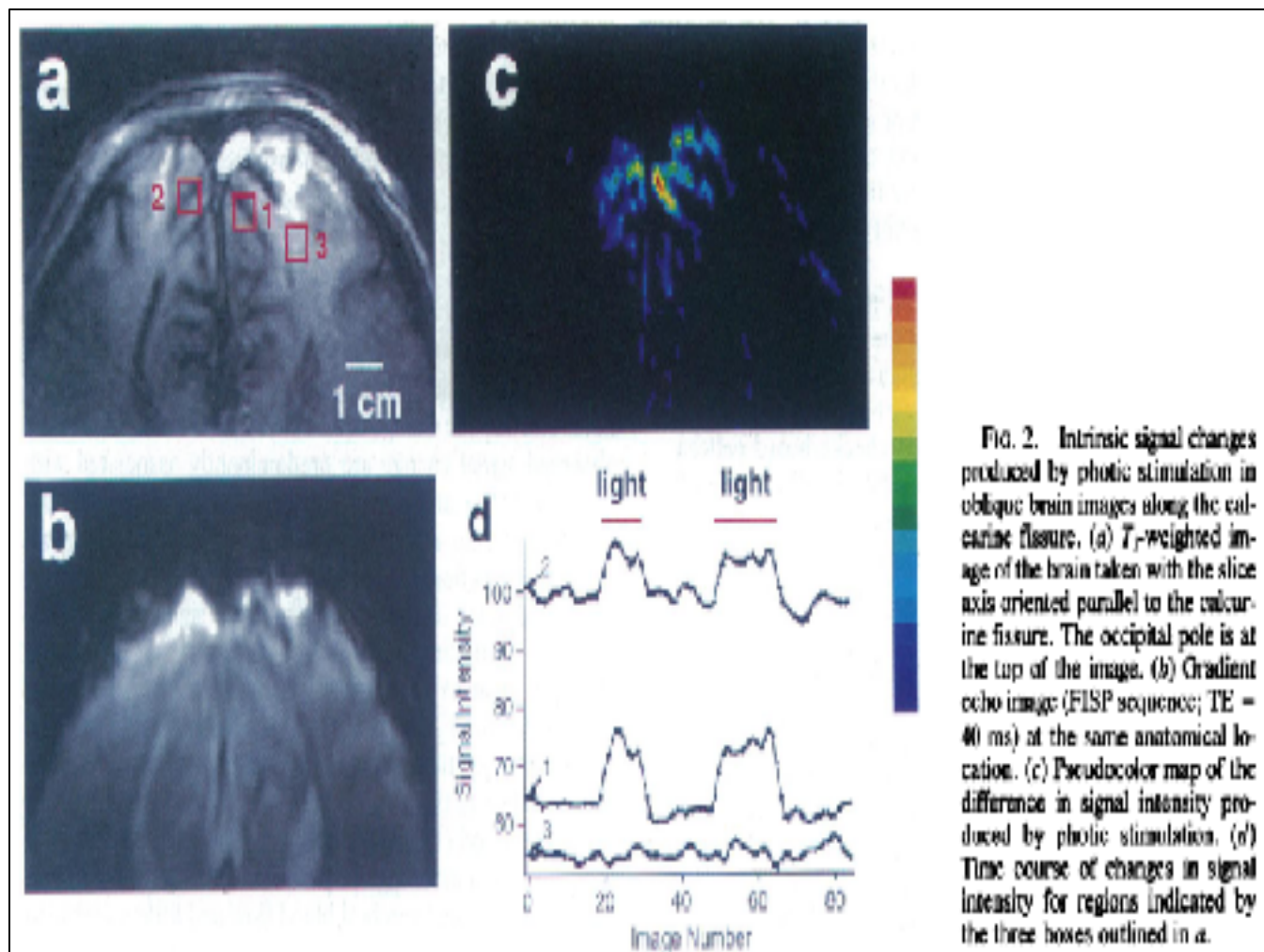
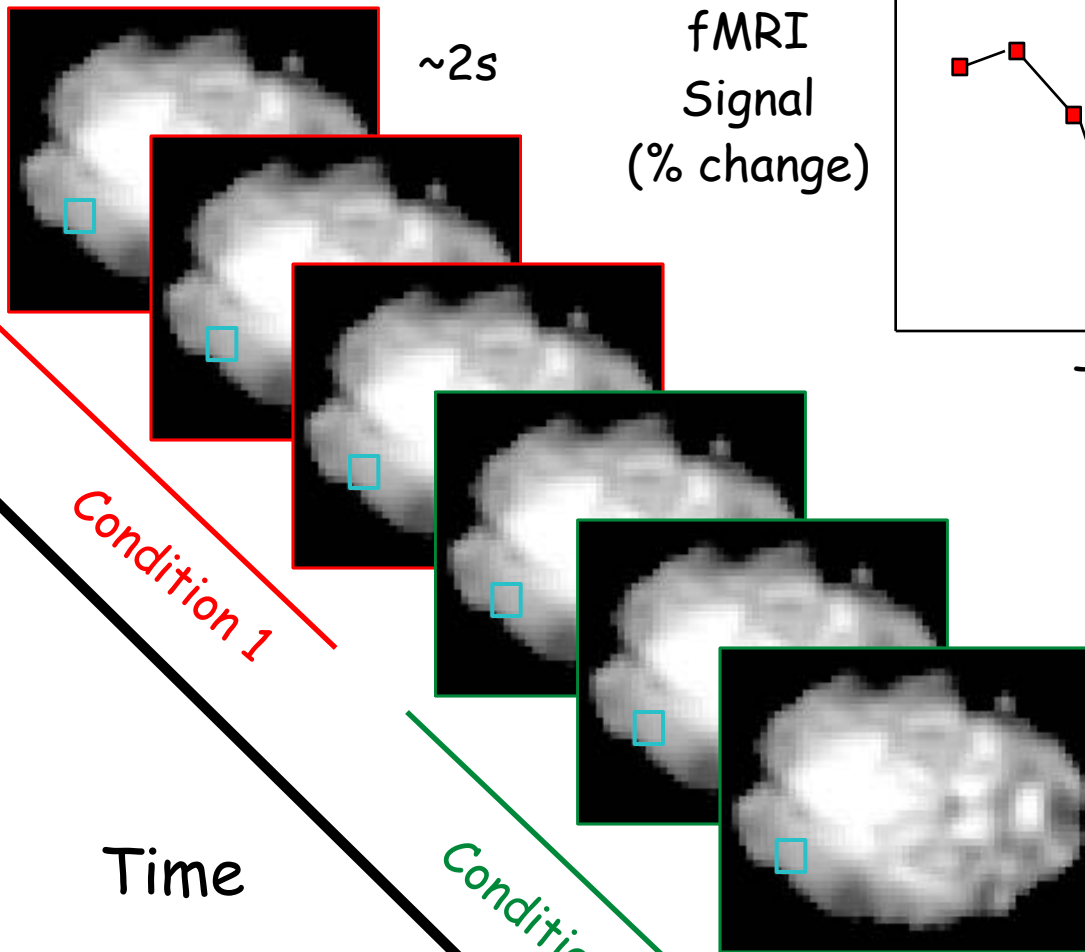


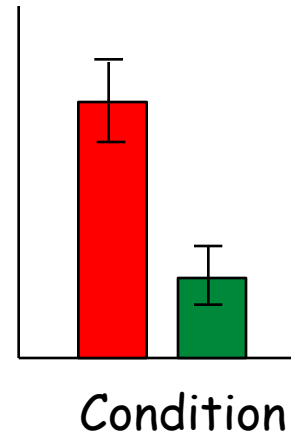
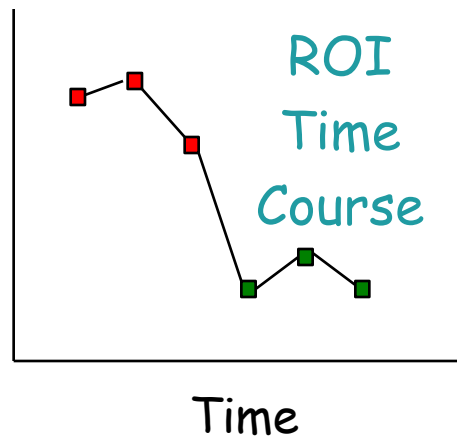
FIG. 2. Intrinsic signal changes produced by photic stimulation in oblique brain images along the calcarine fissure. (a) T_1 -weighted image of the brain taken with the slice axis oriented parallel to the calcarine fissure. The occipital pole is at the top of the image. (b) Gradient echo image (FLSP sequence; TE = 40 ms) at the same anatomical location. (c) Pseudocolor map of the difference in signal intensity produced by photic stimulation. (d) Time course of changes in signal intensity for regions indicated by the three boxes outlined in a.

Activation Statistics

Functional images



fMRI
Signal
(% change)



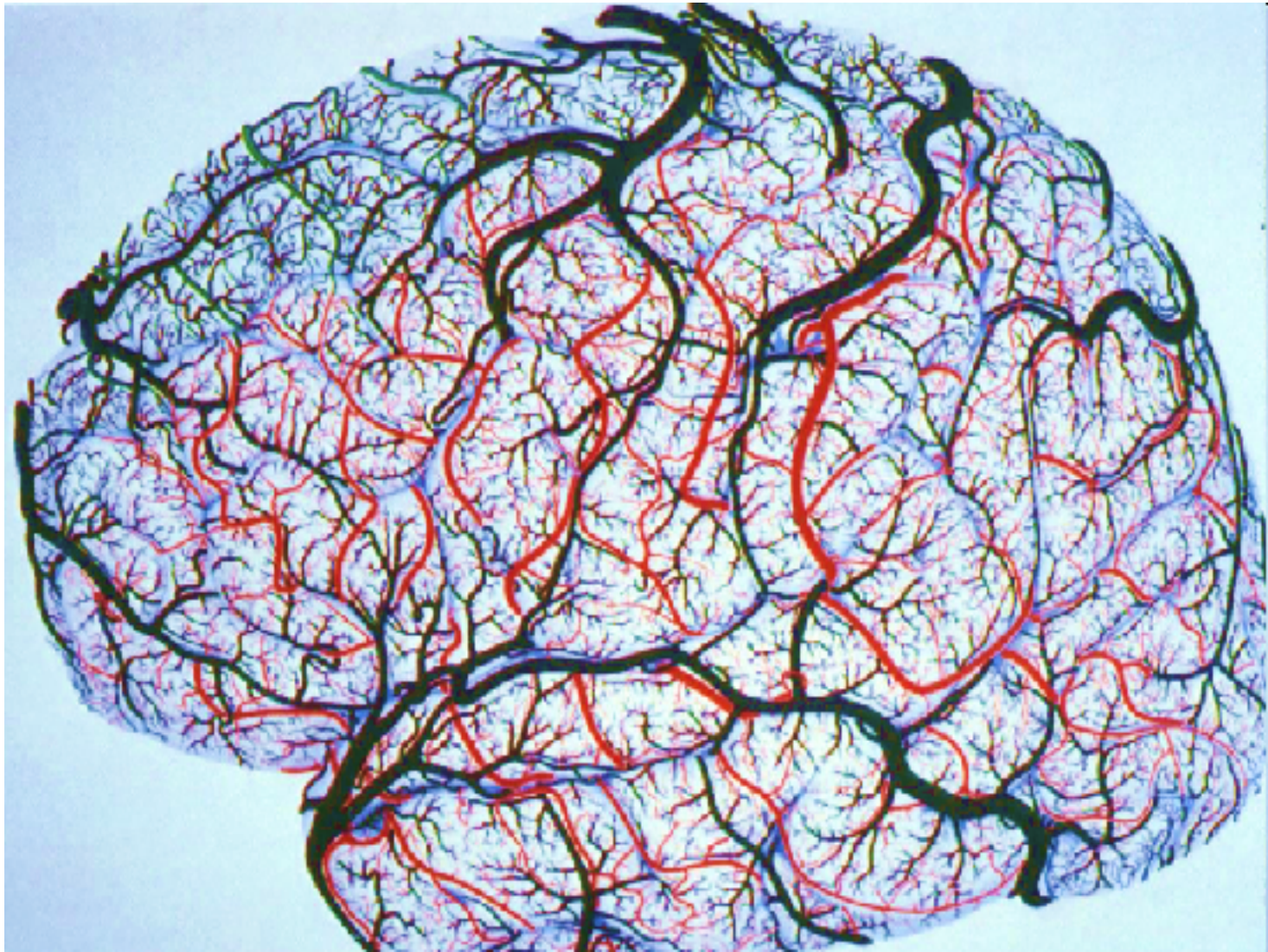
Statistical Map
superimposed on
anatomical MRI image



fMRI Setup



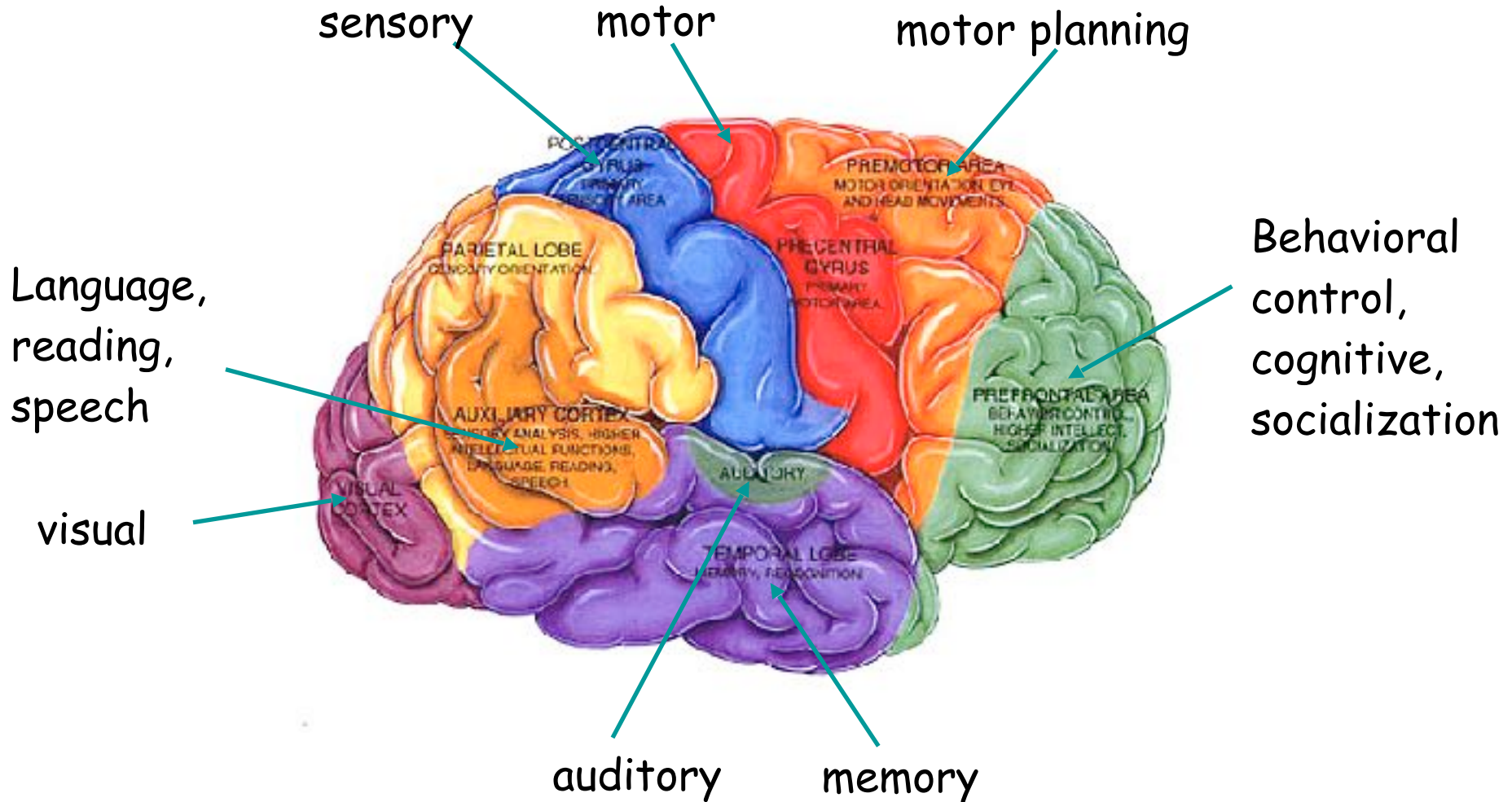
Courtesy, Robert Cox,
Scientific and Statistical
Computing Core Facility,
NIMH

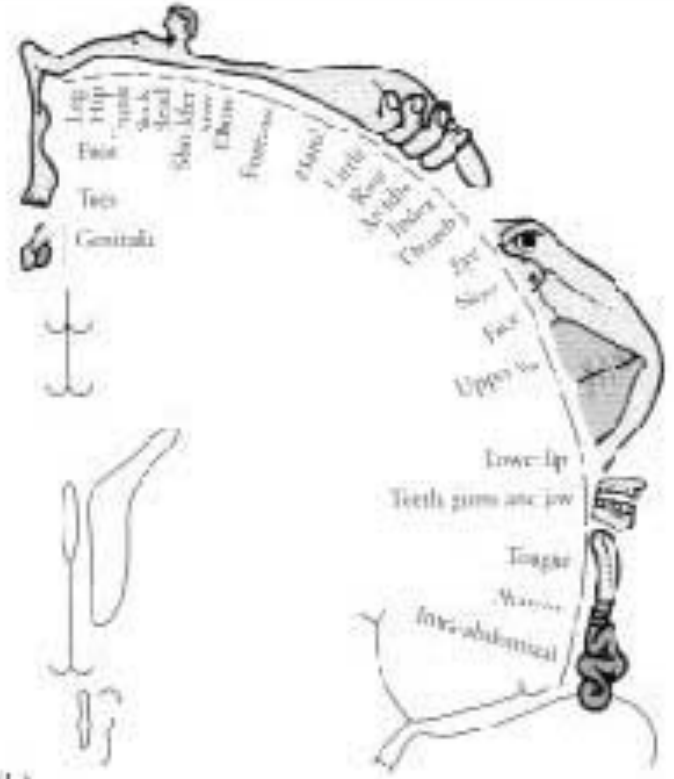
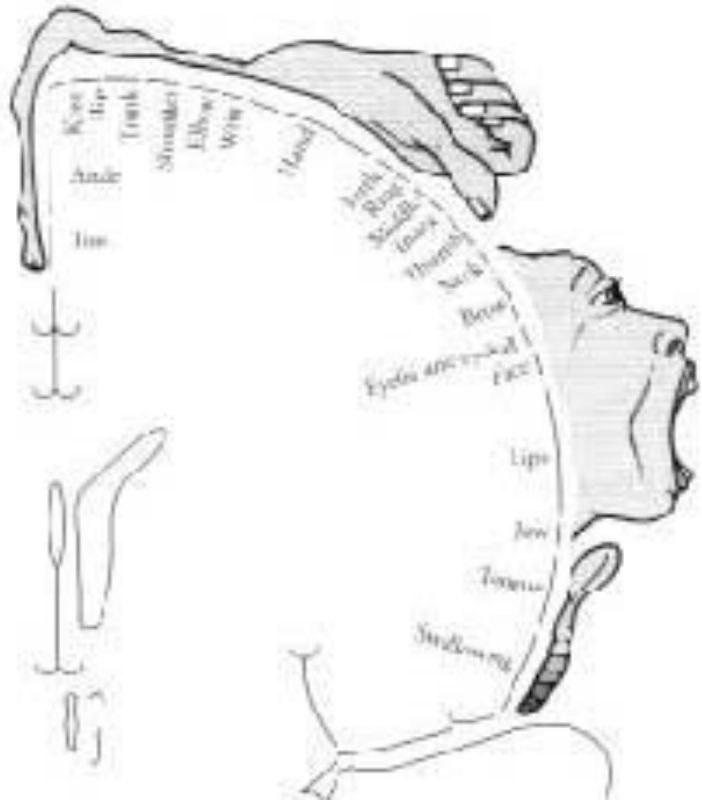


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Motor Cortex

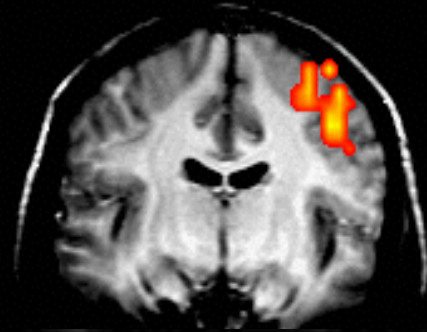
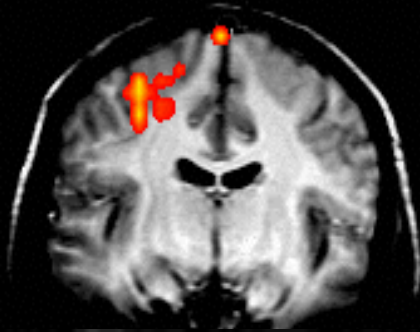
Brain Function



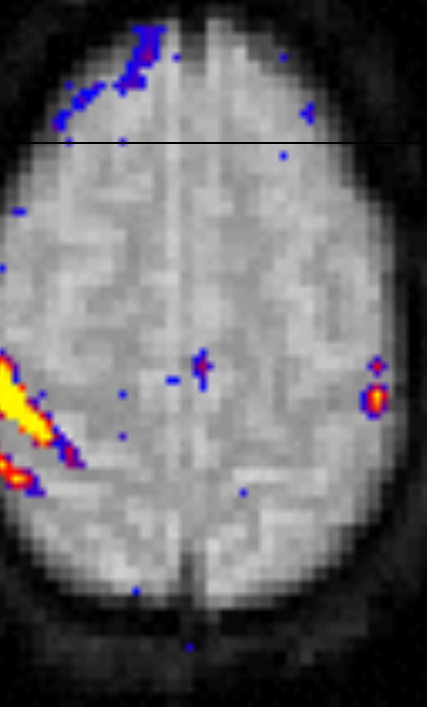
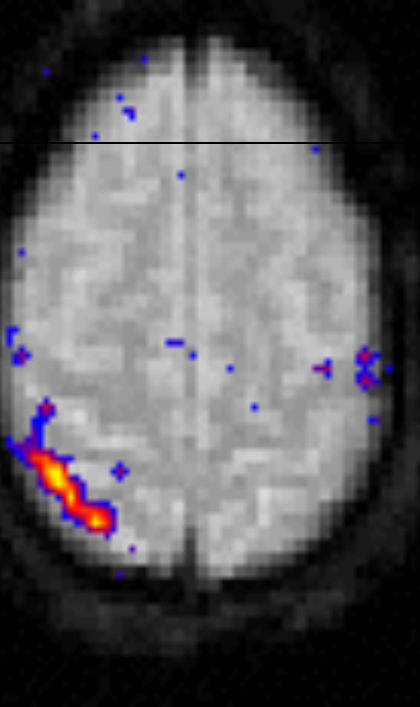


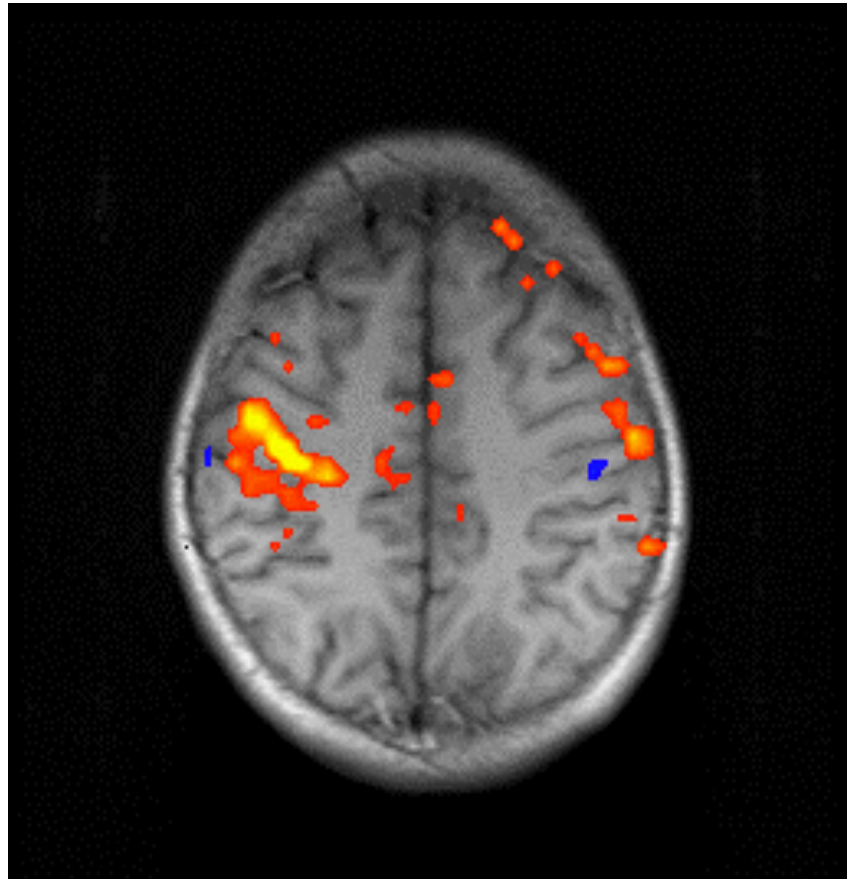
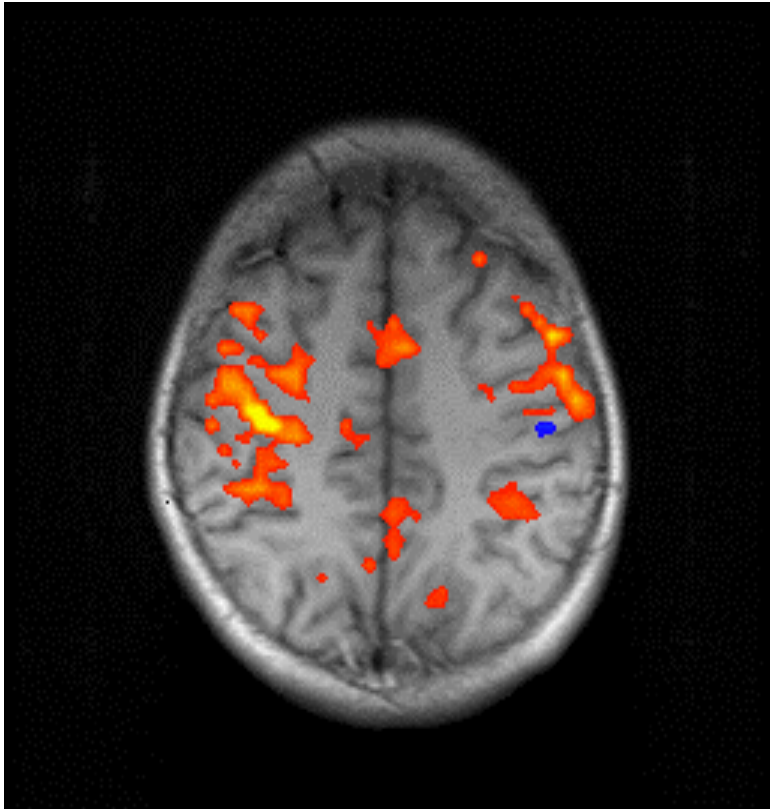
(b)

Toe movement

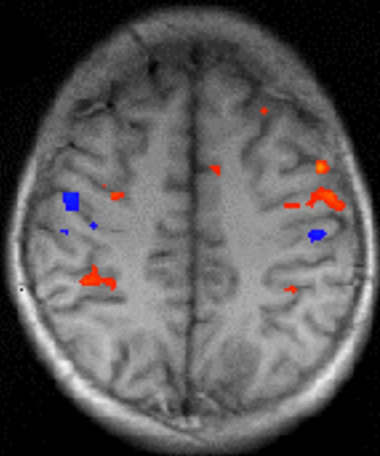


Finger movement

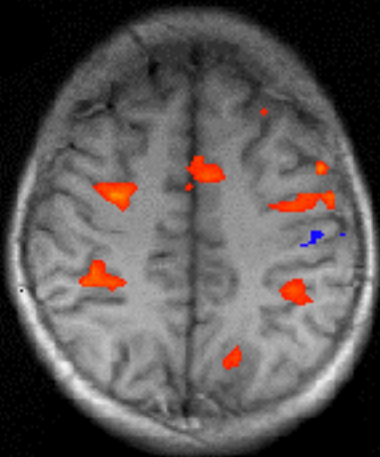




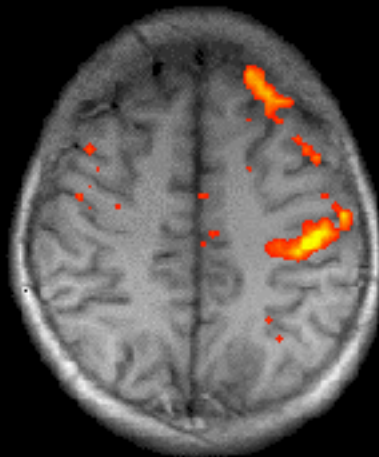
Simple Right



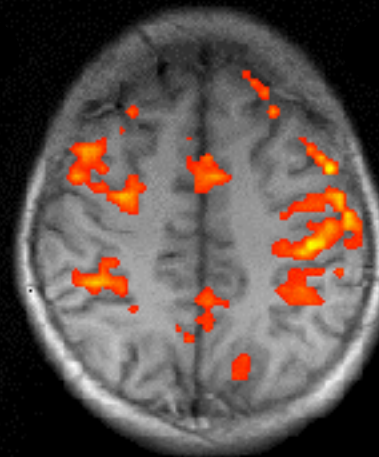
Simple Left



Complex Left



Imagined
Complex Left



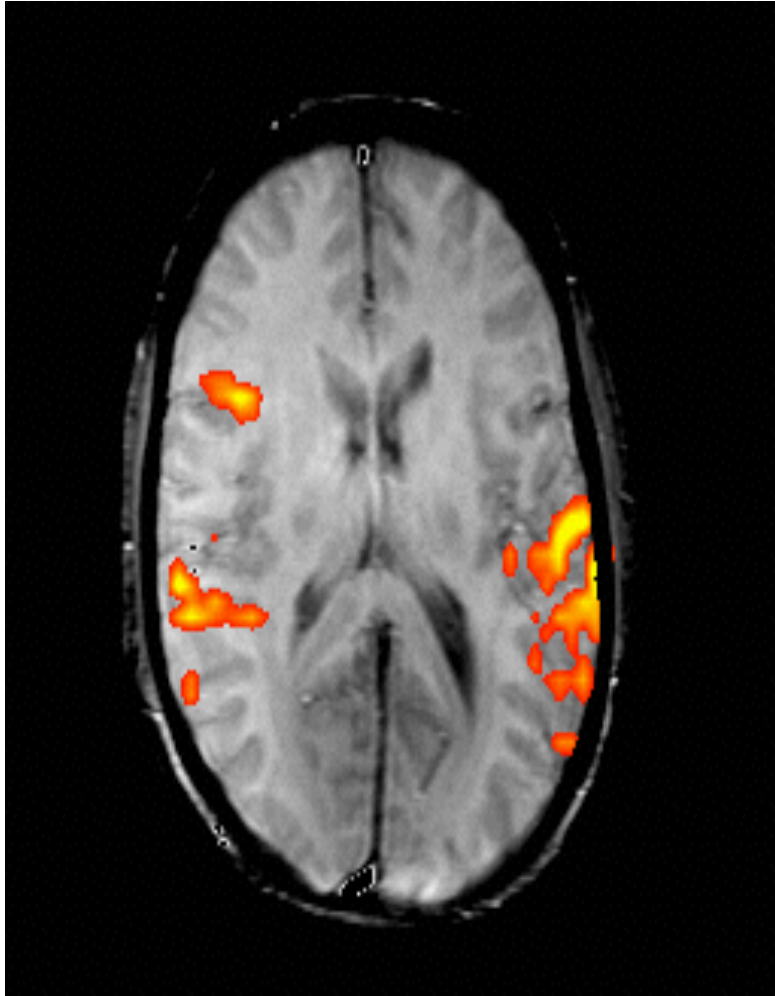
Presurgical Mapping

Left Foot

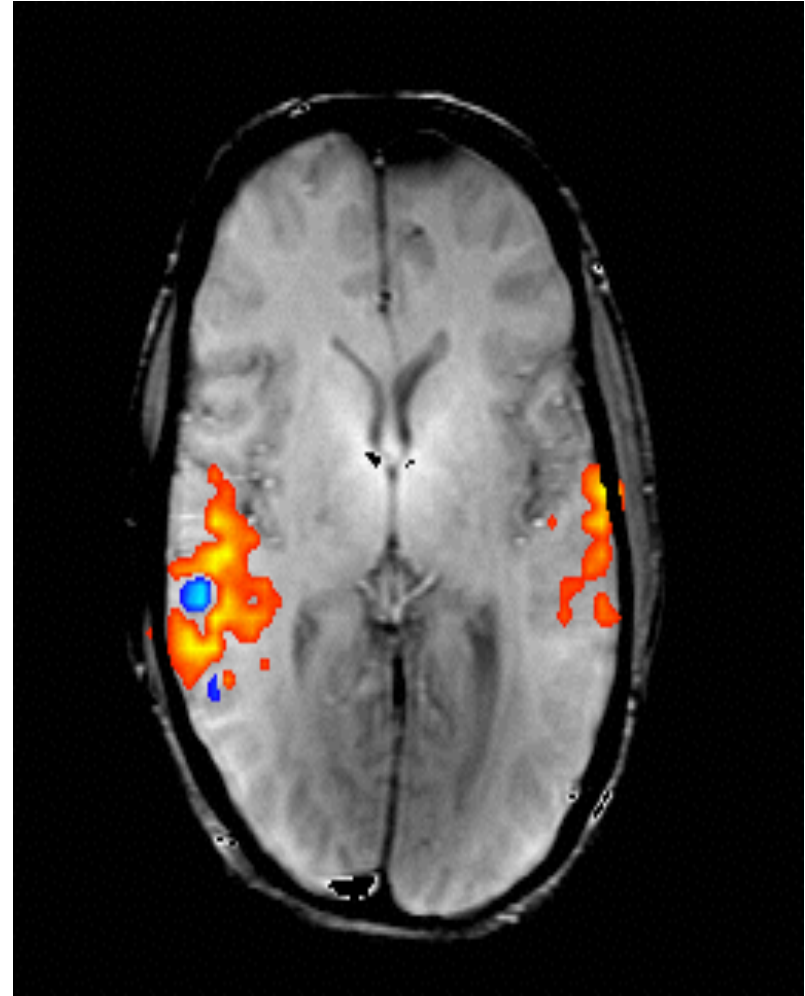
Tumor

Right Foot

Right Hand



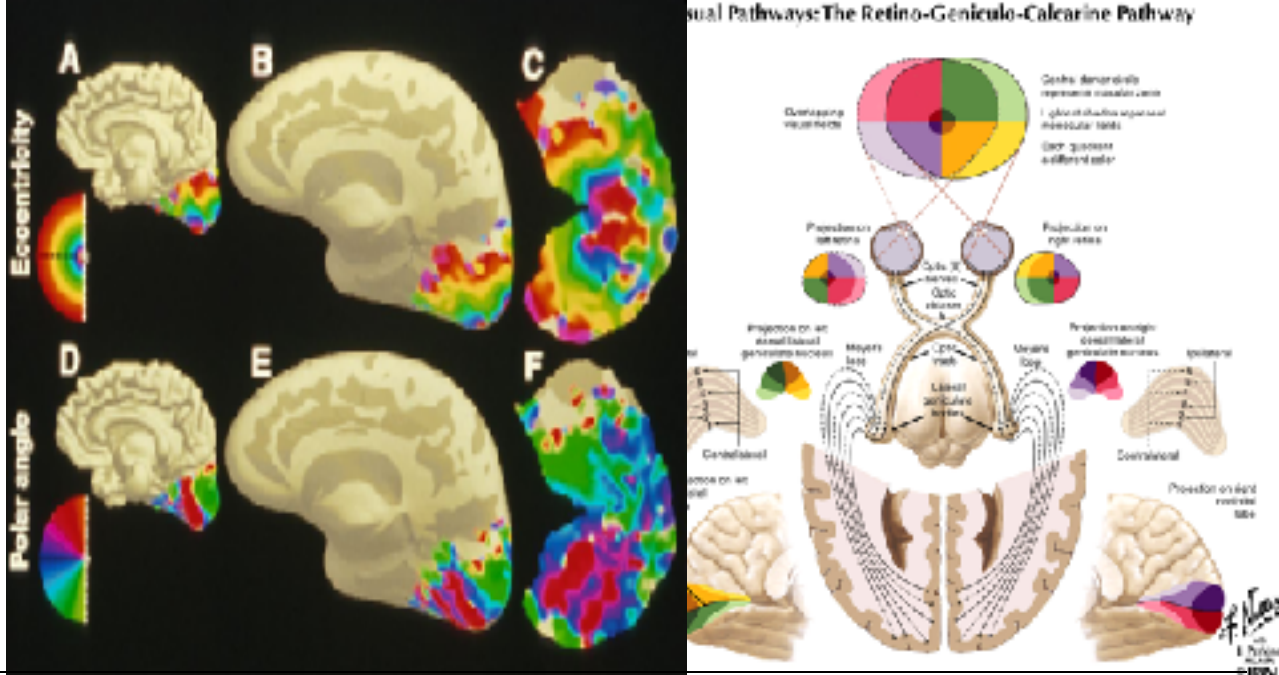
fMRI



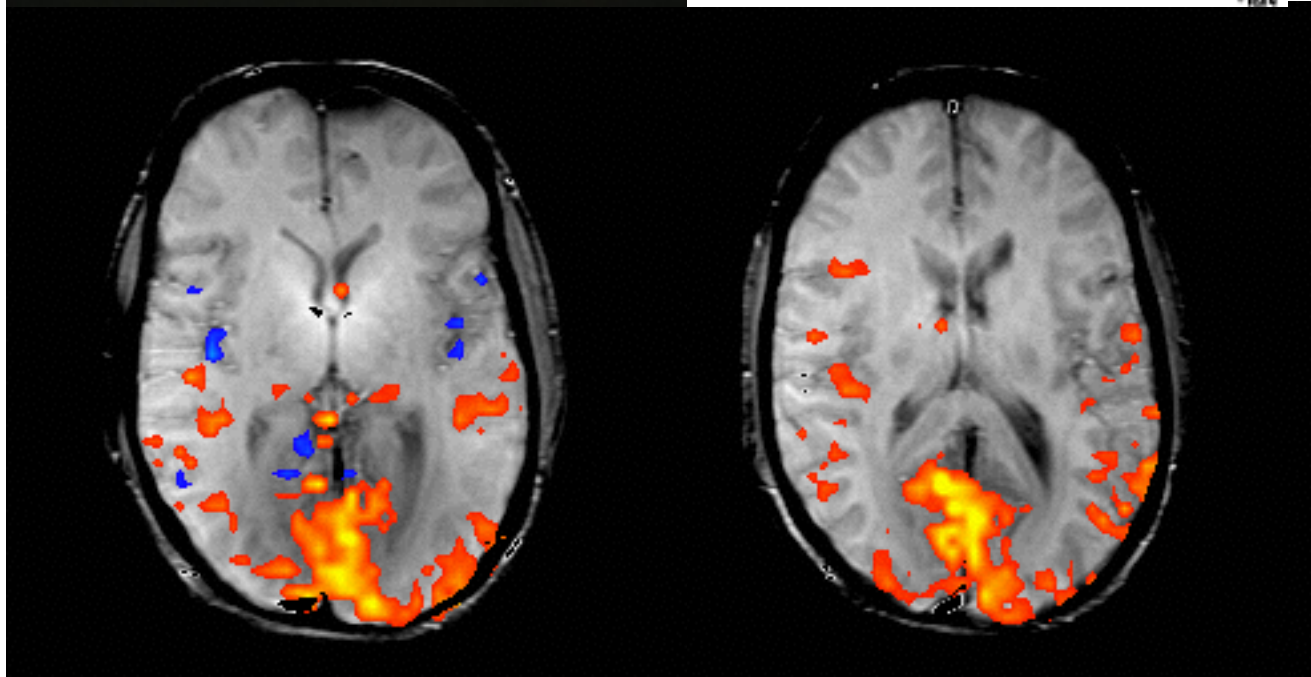
O-15 PET

Visual and Auditory Processing

Reading






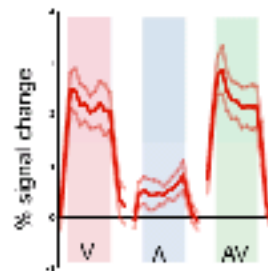
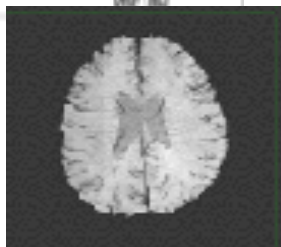
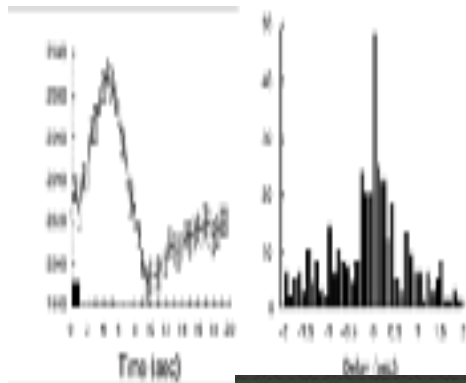
Listening

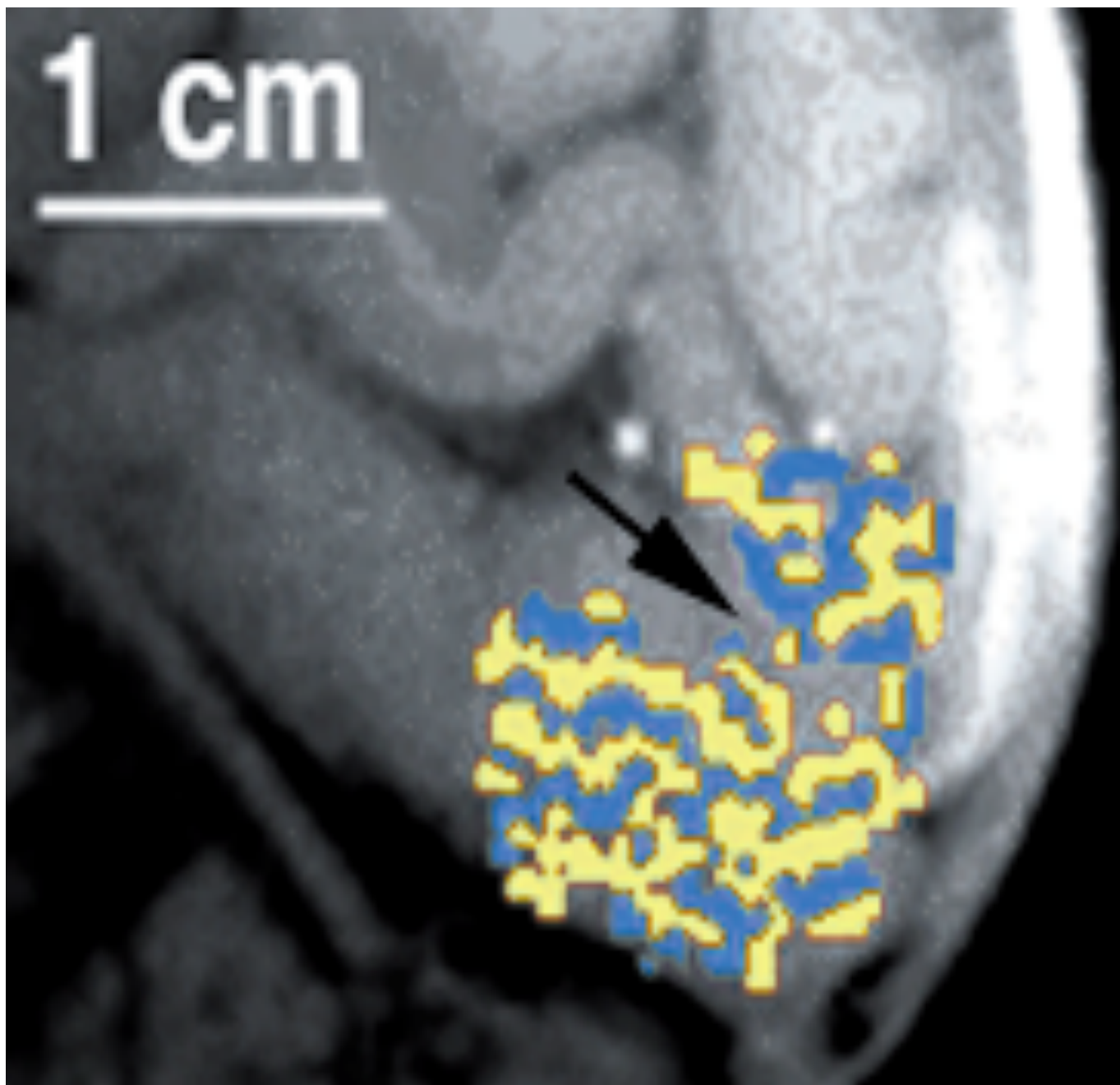


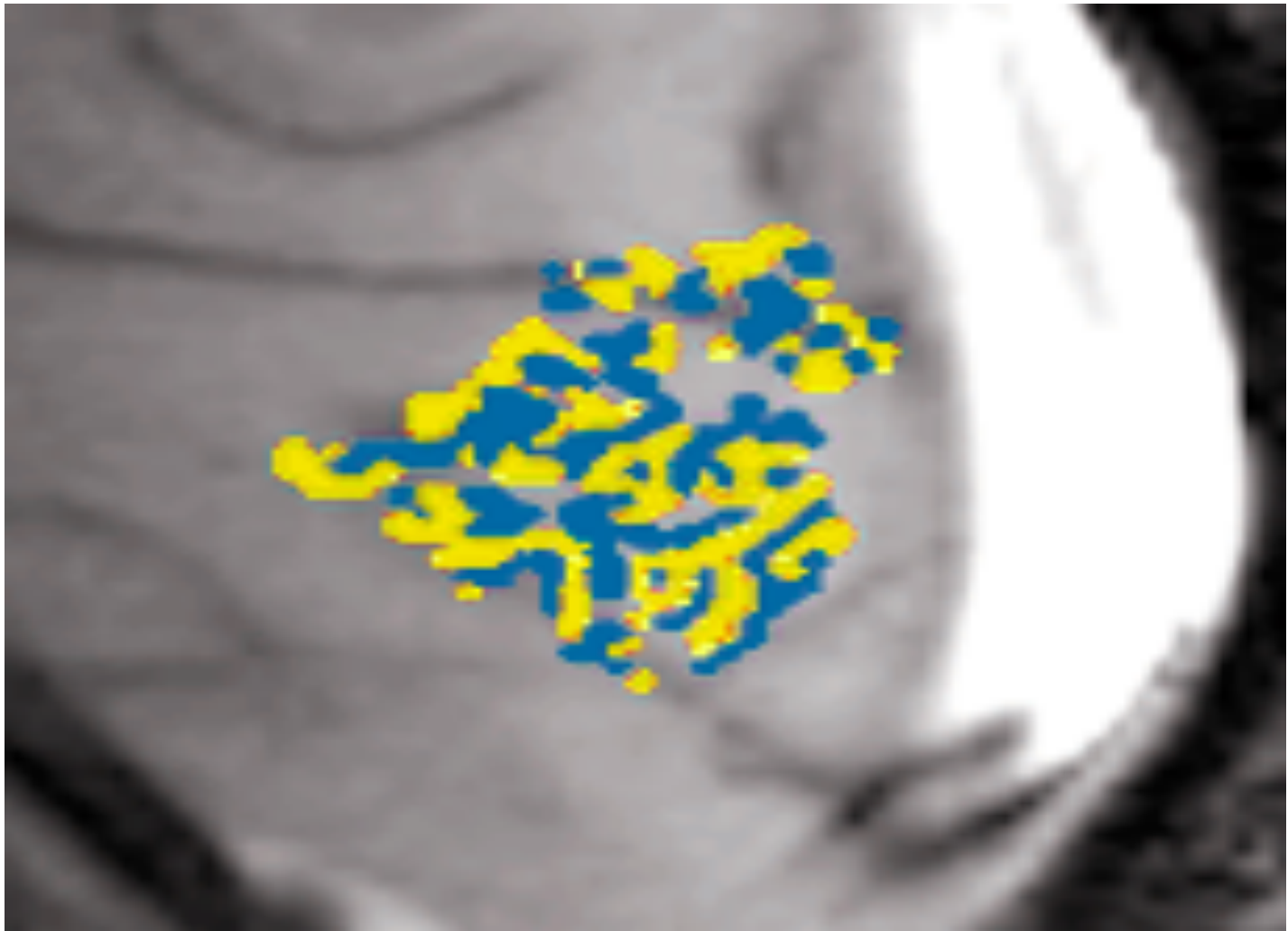
Multi-sensory integration

M.S. Beauchamp et al.,

-  Visual
-  Auditory
-  Multisensory



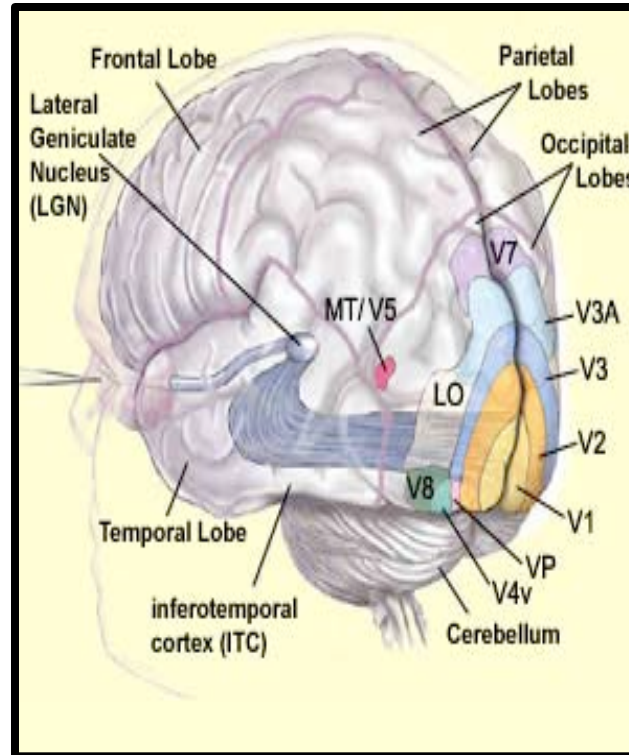
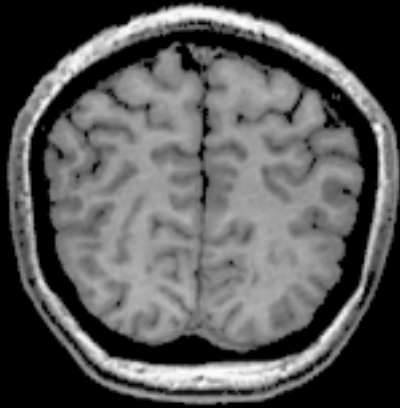




Visual Processing

(spatial resolution)

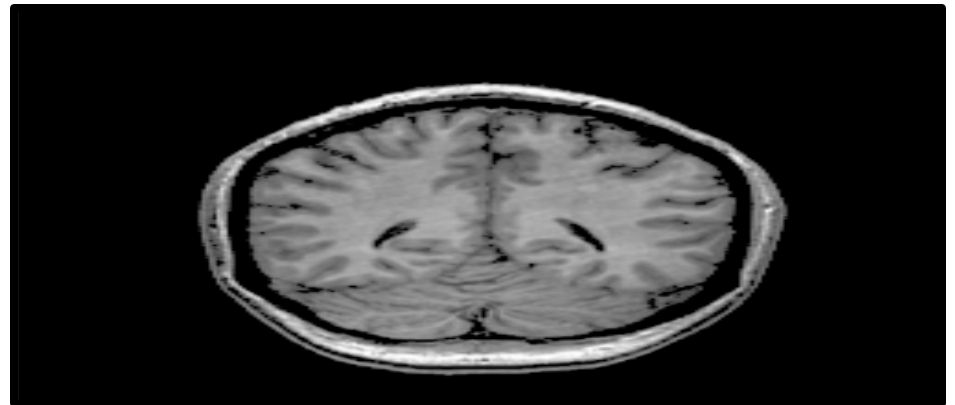
Visual Cortex Organization



Functional Magnetic Resonance Imaging in Medicine and Physiology

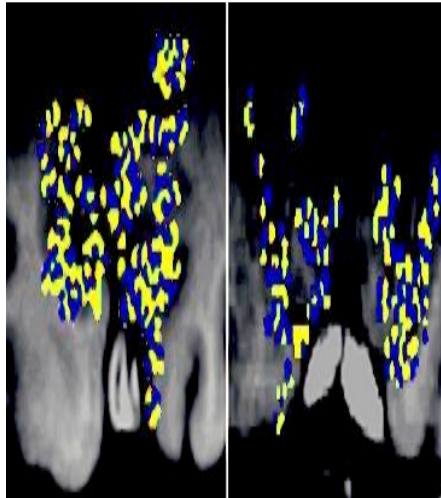
CHRIST T. W. MOONEN, PETER C. M. VAN ZIJL, JOSEPH A. FRANK,
DENIS LE BIHAN, EDWIN D. BECKER

<http://www.thebrain.mcgill.ca>



ODC Maps using fMRI

calcarine



1 cm

- Identical in size, orientation, and appearance to those obtained by optical imaging¹ and histology^{3,4}.

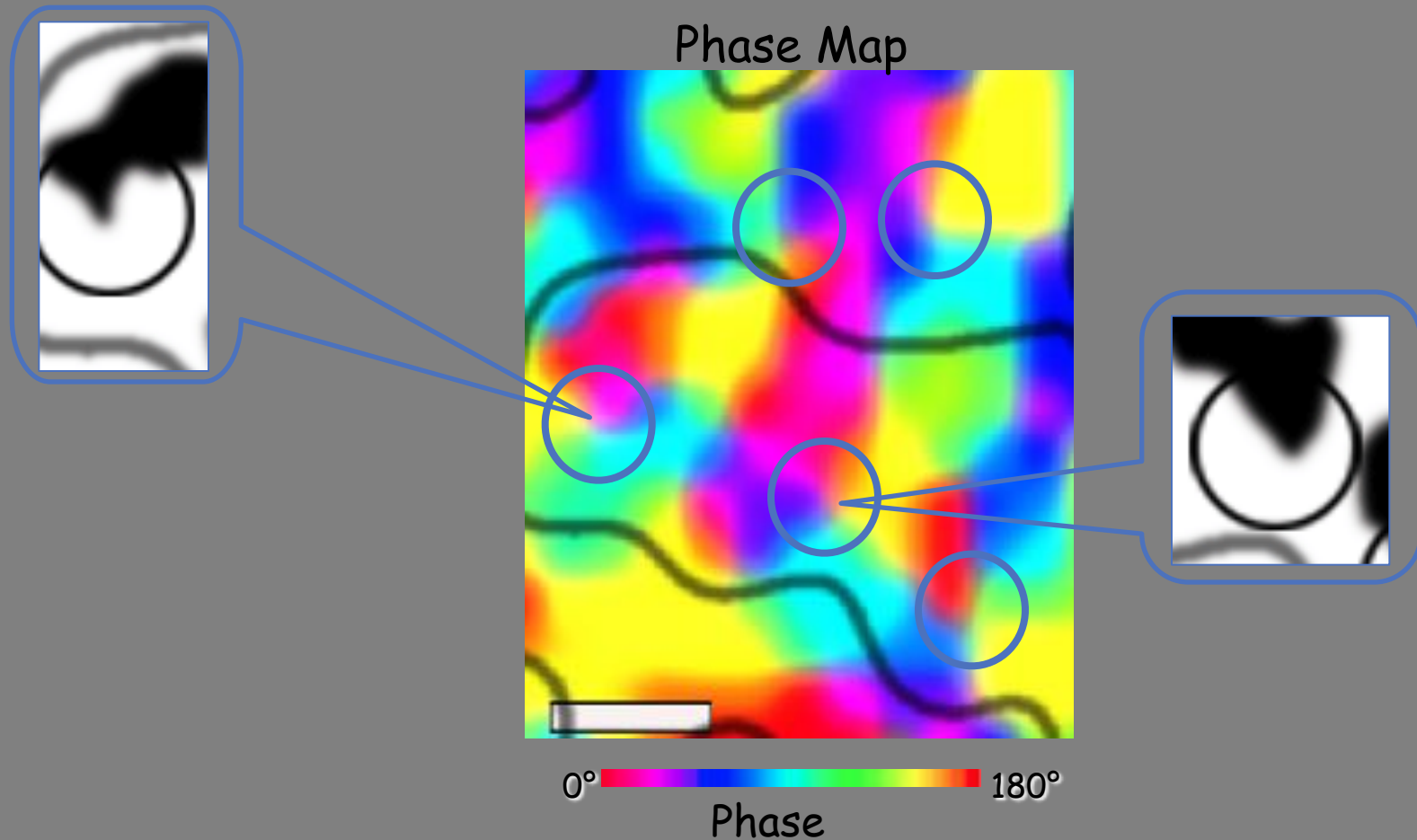
Menon et al.

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

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

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

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



Yacoub, Ugurbil & Harel

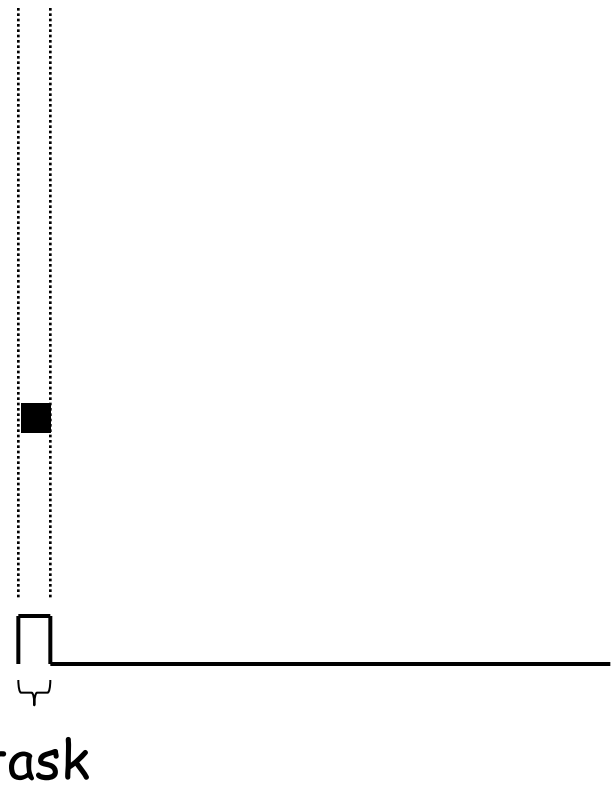
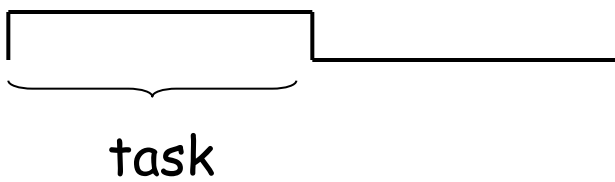
University of Minnesota / CMRR

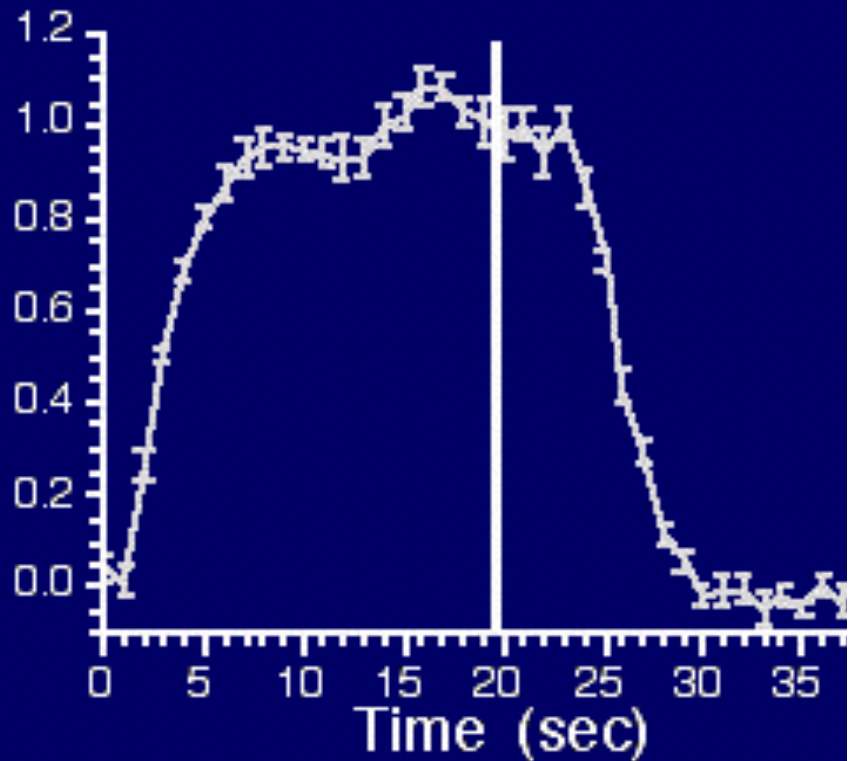
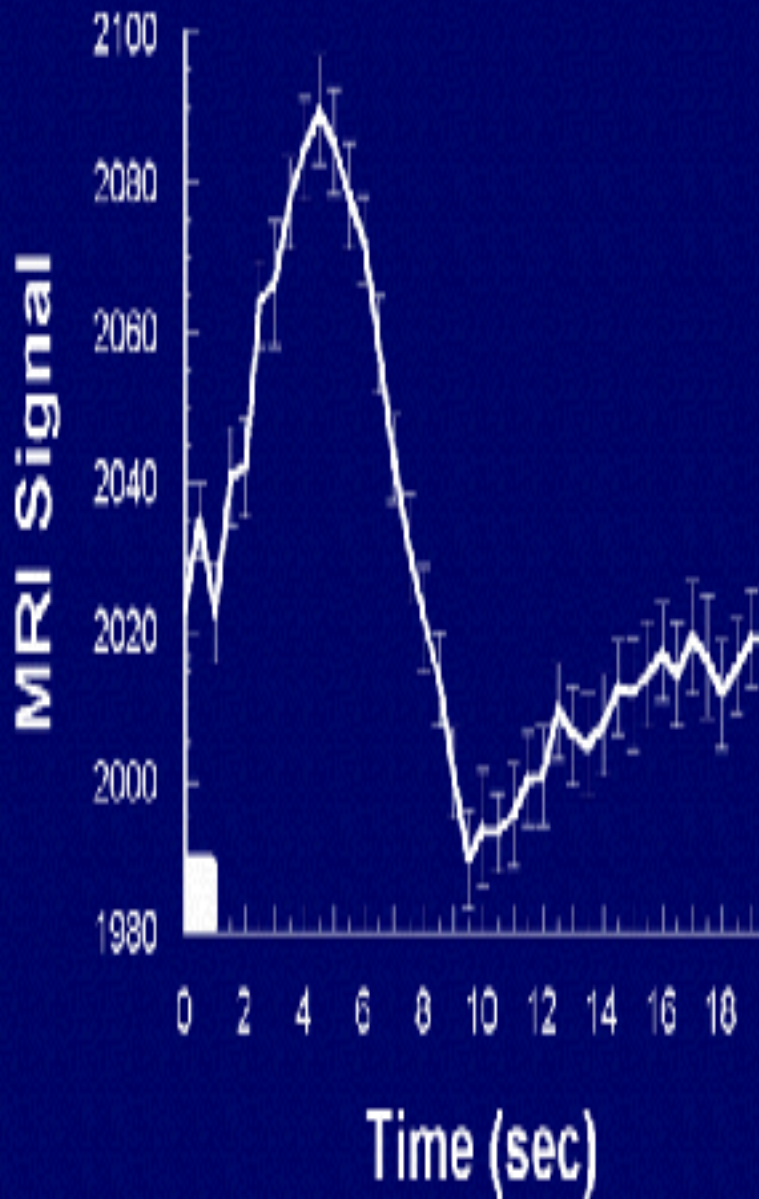
HBM 2006: Thursday, June 15, 2006 at 9:30

Temporal Resolution

The BOLD Signal

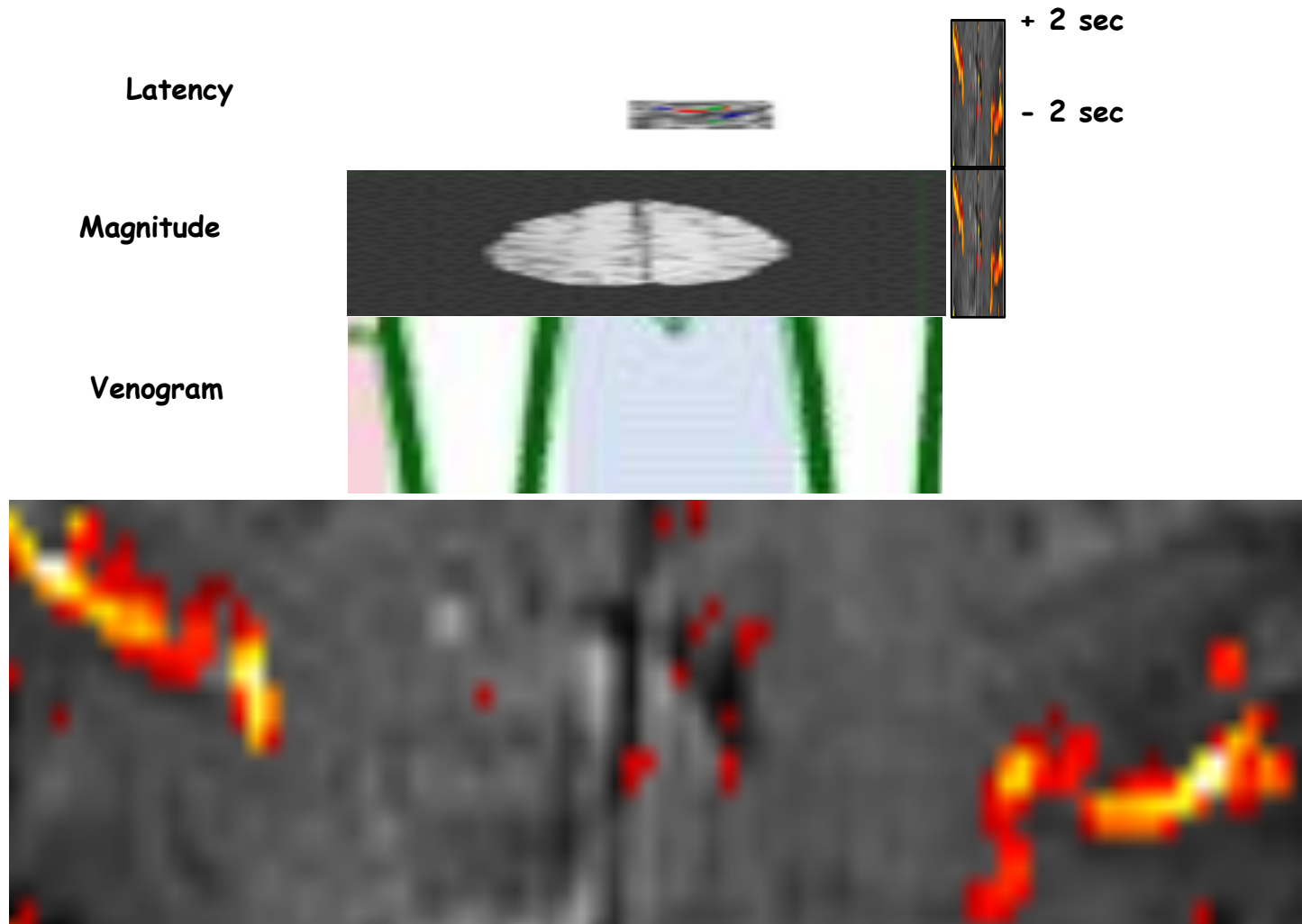
Blood Oxygenation Level Dependent (BOLD) signal changes





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

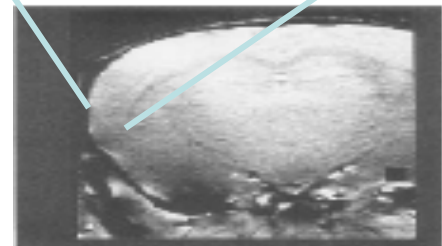
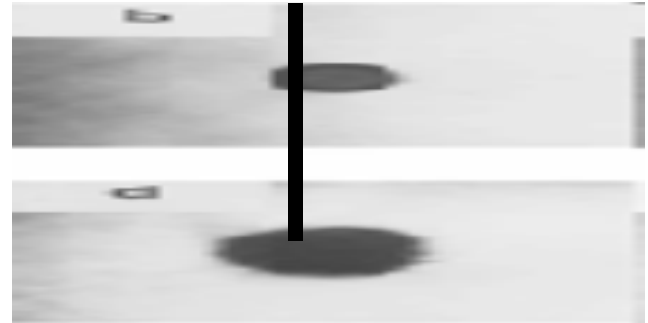
Latency Variation...



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

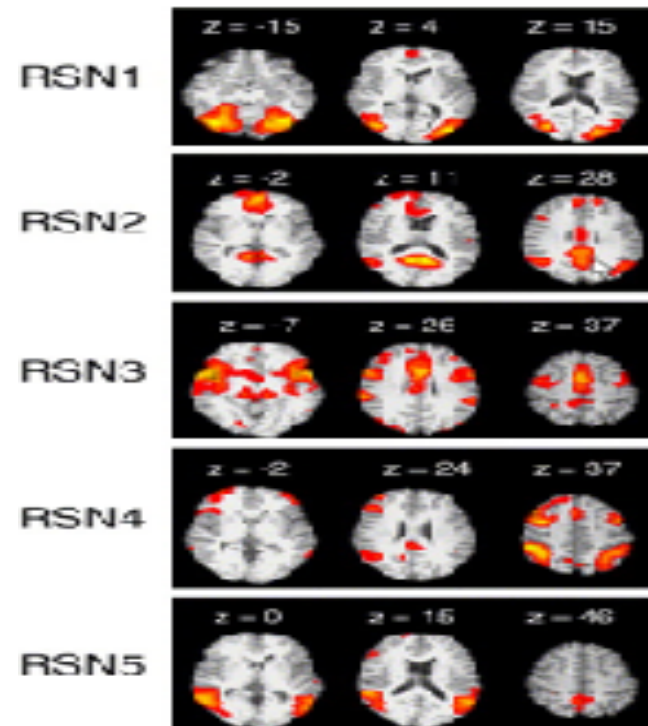
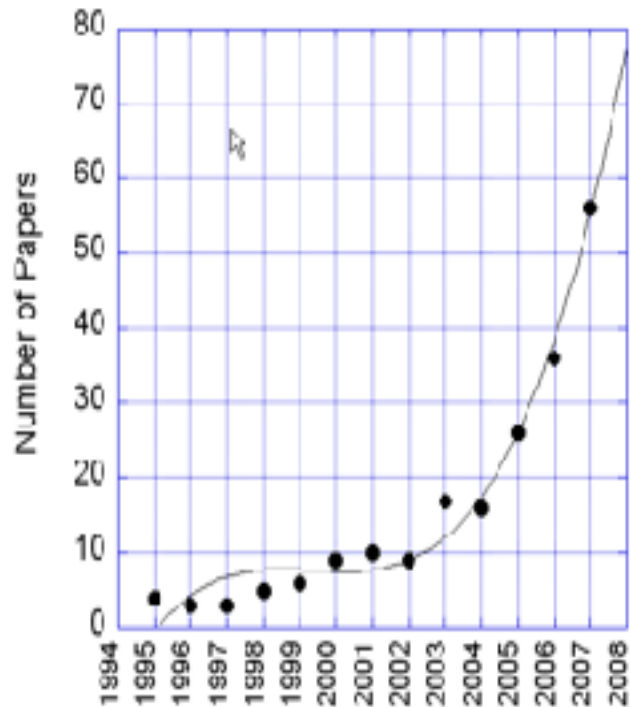
Word vs. Non-word

0°, 60°, 120° Rotation



"Resting State"
Fluctuations

Resting State Correlations



Activation:

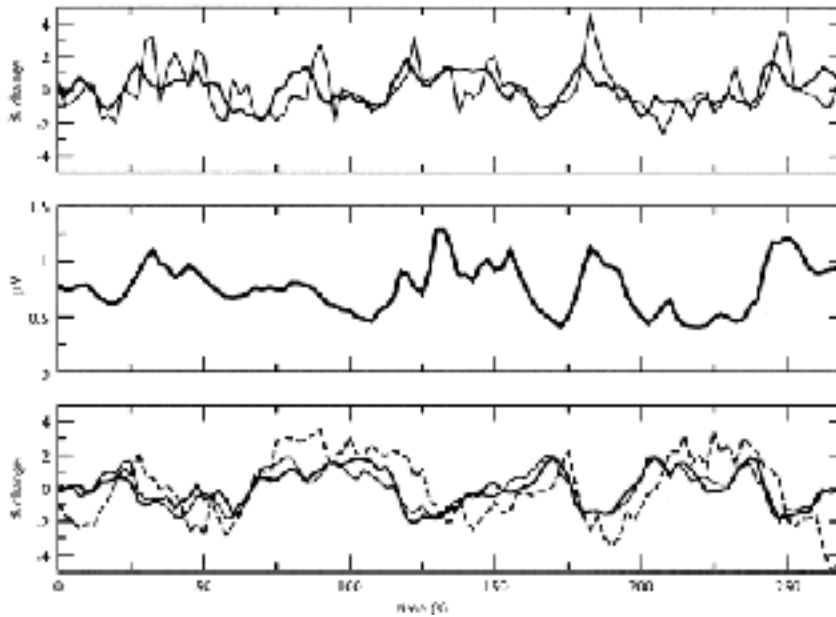
correlation with reference function

Rest:

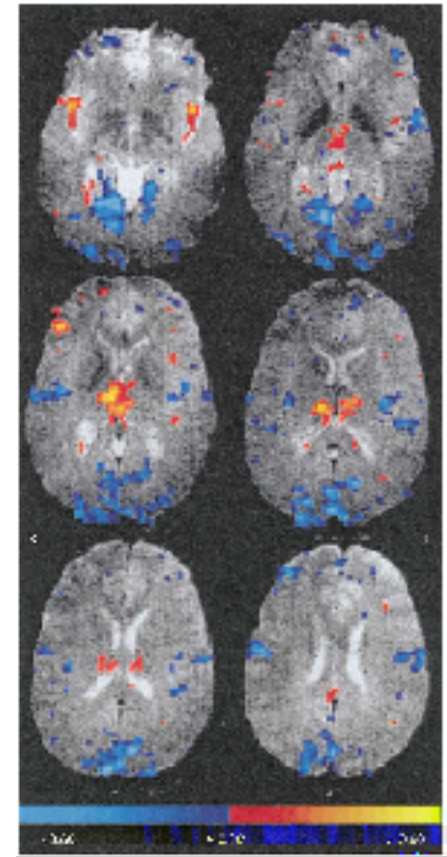
seed voxel in motor cortex

BOLD correlated with 10 Hz power during "Rest"

Positive
10 Hz power
Negative



Goldman, et al (2002), Neuroreport

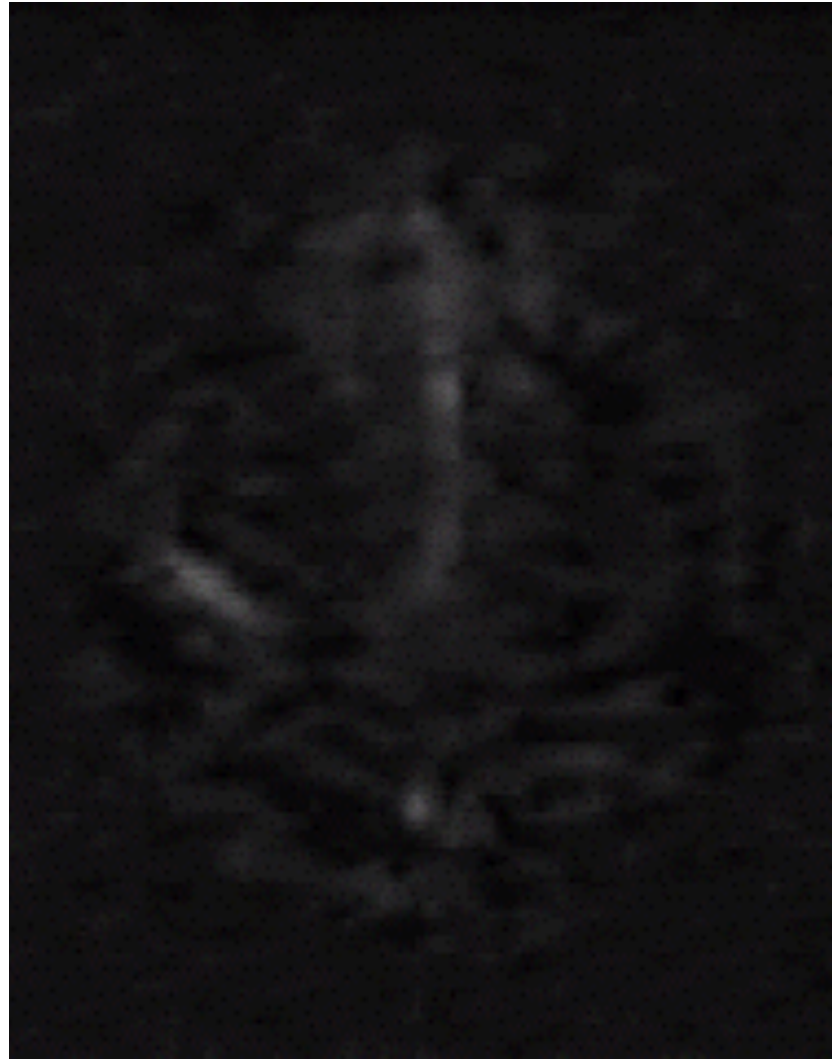


Resting state networks identified with ICA

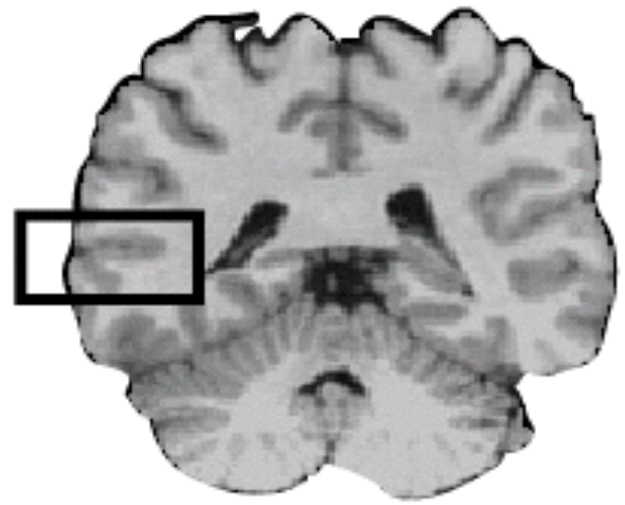
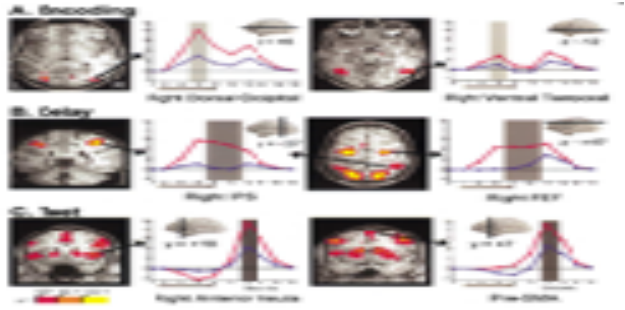
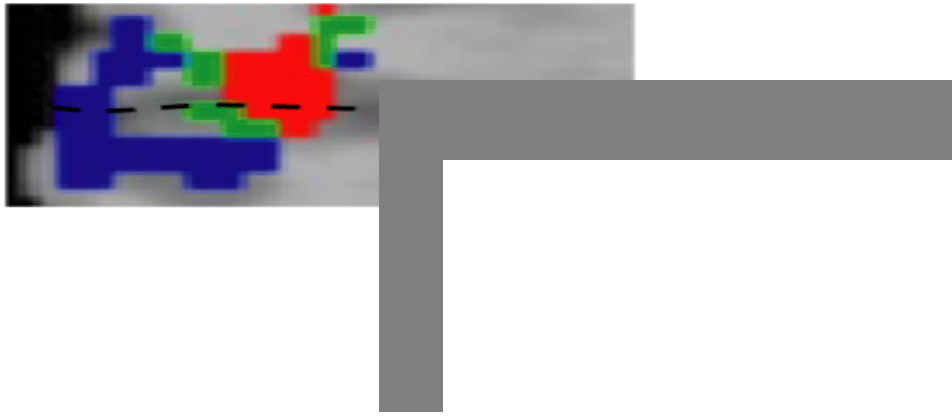
M. DeLuca, C.F. Beckmann, N. De Stefano,
P.M. Matthews, S.M. Smith, **fMRI resting state
networks define distinct modes of long-distance
interactions in the human brain.** *NeuroImage*, 29,
1359-1367

"Brain Reading"

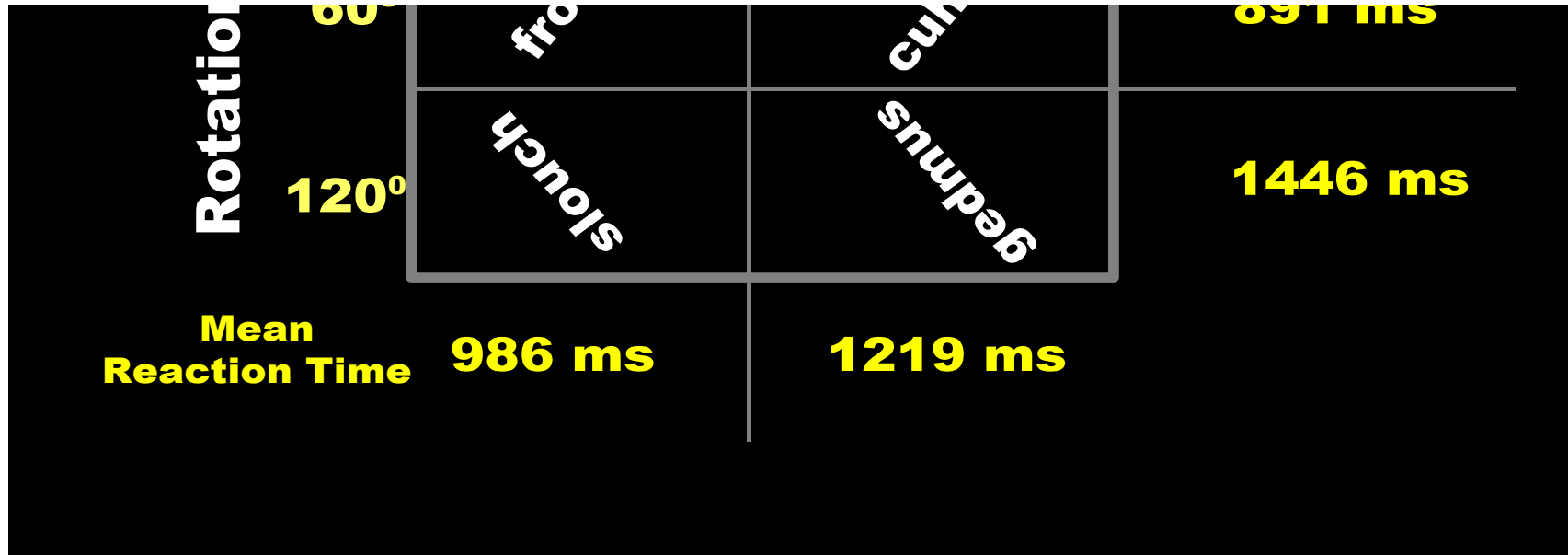
Methodology



Mapping ↔ "Reading"



Visual object categories distinguished by widely distributed inferotemporal activity pattern



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^{1,2,3*} and Robert L. Savoy^{1,2,3*}

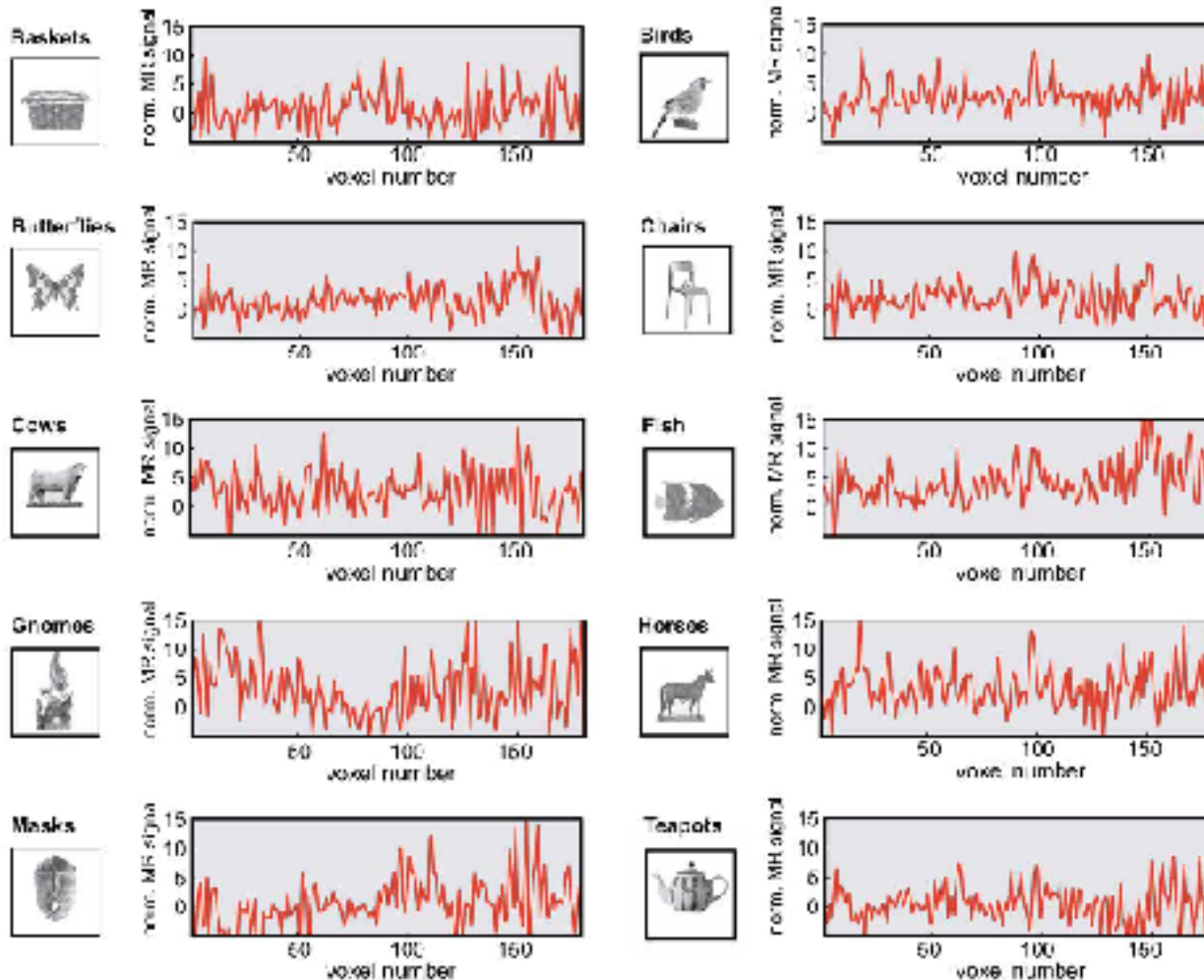
¹Harvard Institute for Learning, Cambridge, MA 02142, USA

²Department of Neuroimaging and Anatomical Functional Imaging, Charlestown, MA 02128, USA

³Alphas Pictures, Inc., P.O. Box 158, Arlington, MA 02478, USA

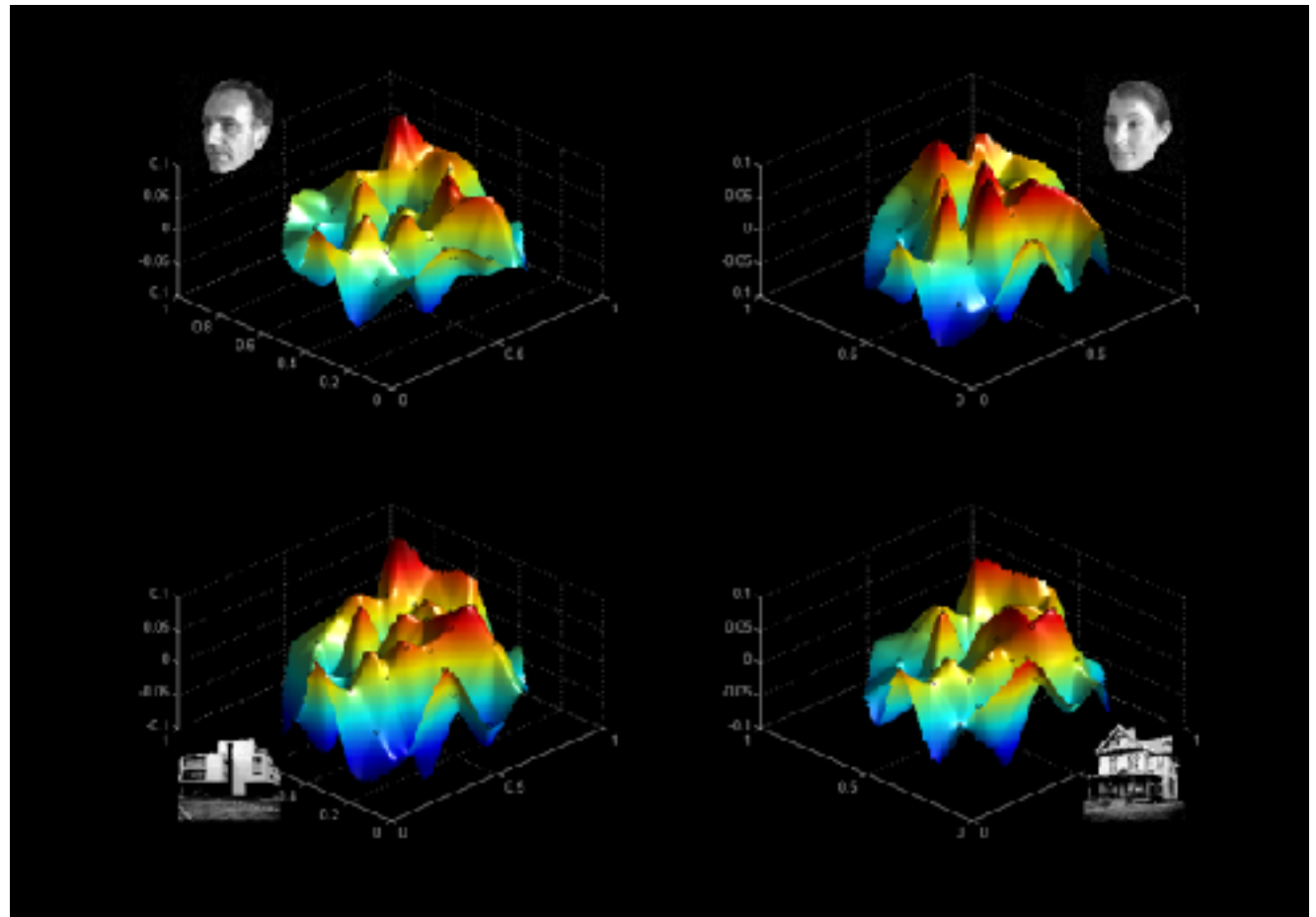
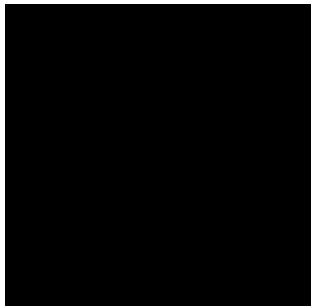
Received 15 July 2002; accepted 10 December 2002

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

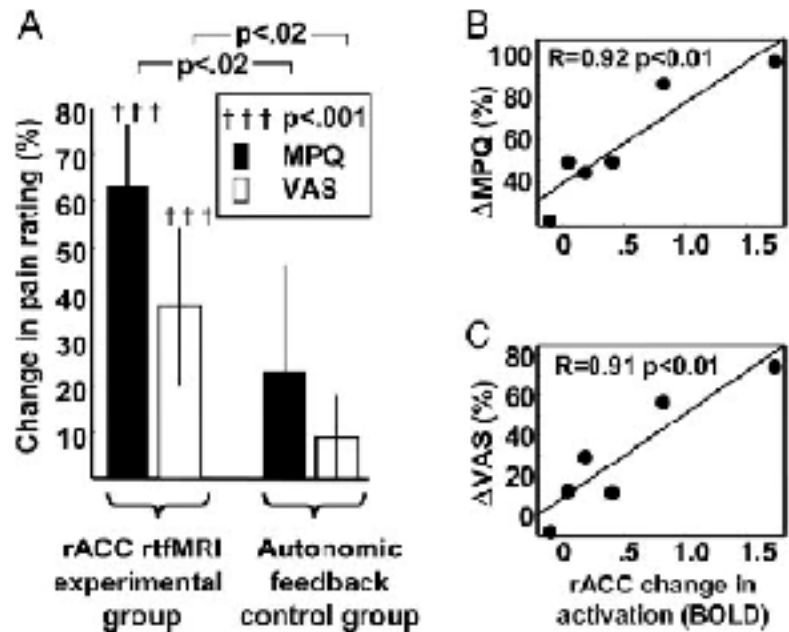
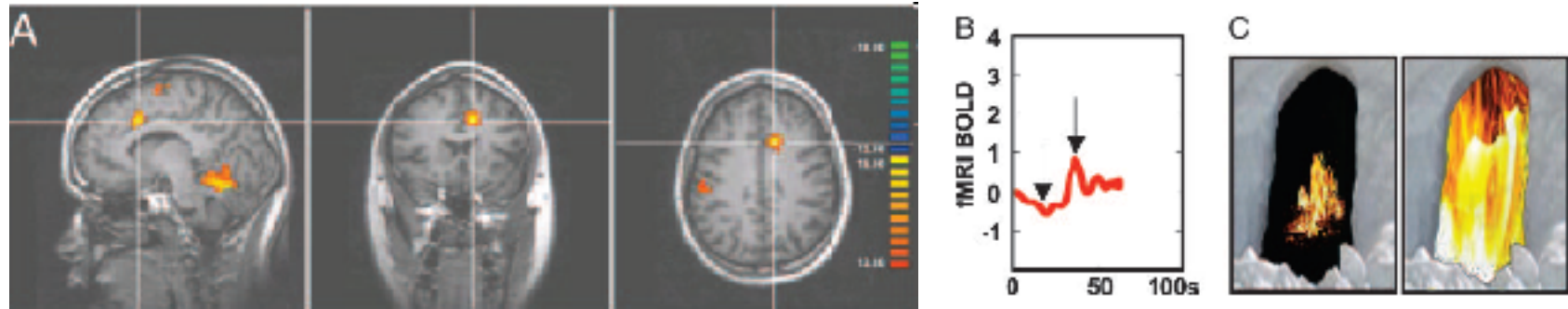


"searchlight" ROI

Multivariate analysis



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)

Current Uses of fMRI

Understanding normal brain organization and changes

- networks involved with specific tasks (low to high level processing)
- changes over time (seconds to years)
- correlates of behavior (response accuracy, performance changes...)

Clinical research

- correlates of specifically activated networks to clinical populations
- presurgical mapping
- epileptic foci mapping
- drug effects

Potential uses of fMRI

Complementary use for clinical diagnosis

- utilization of clinical research results

Clinical treatment and assessment

- drug, therapy, rehabilitation, biofeedback

Non clinical uses

- complementary use with behavioral results
- lie detection
- prediction of behavior tendencies (many contexts)
- brain/computer interface

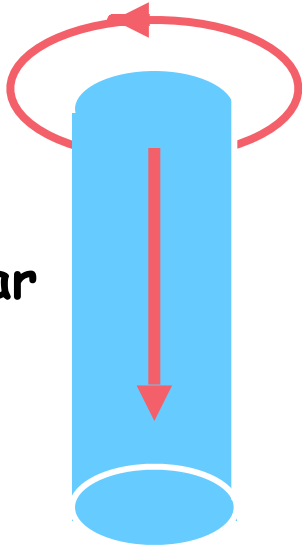
Section on Functional Imaging Methods & Functional MRI Facility Jan 19, 2007

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**

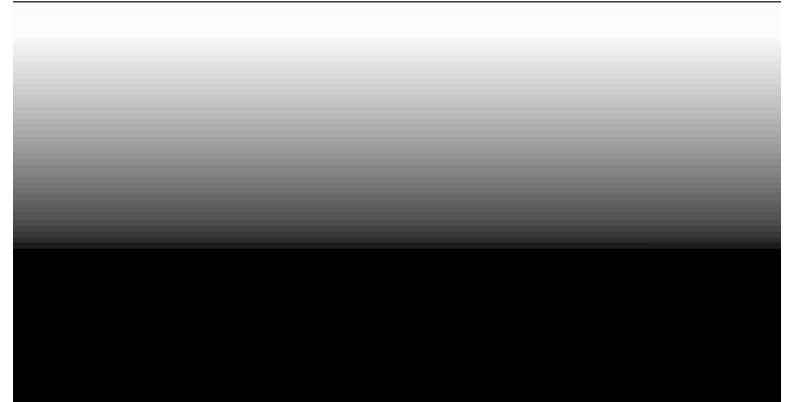
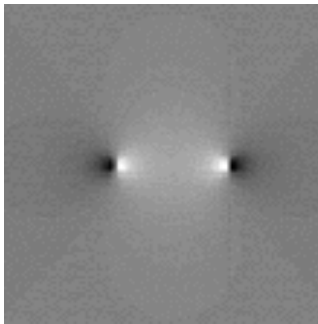
Front row: **Dorian Van Tassel**, **Jerzy Bodurka**, **Adam Thomas**, **Marieke Mur**, **David Knight**

Magnetic Field



**Intracellular
Current**

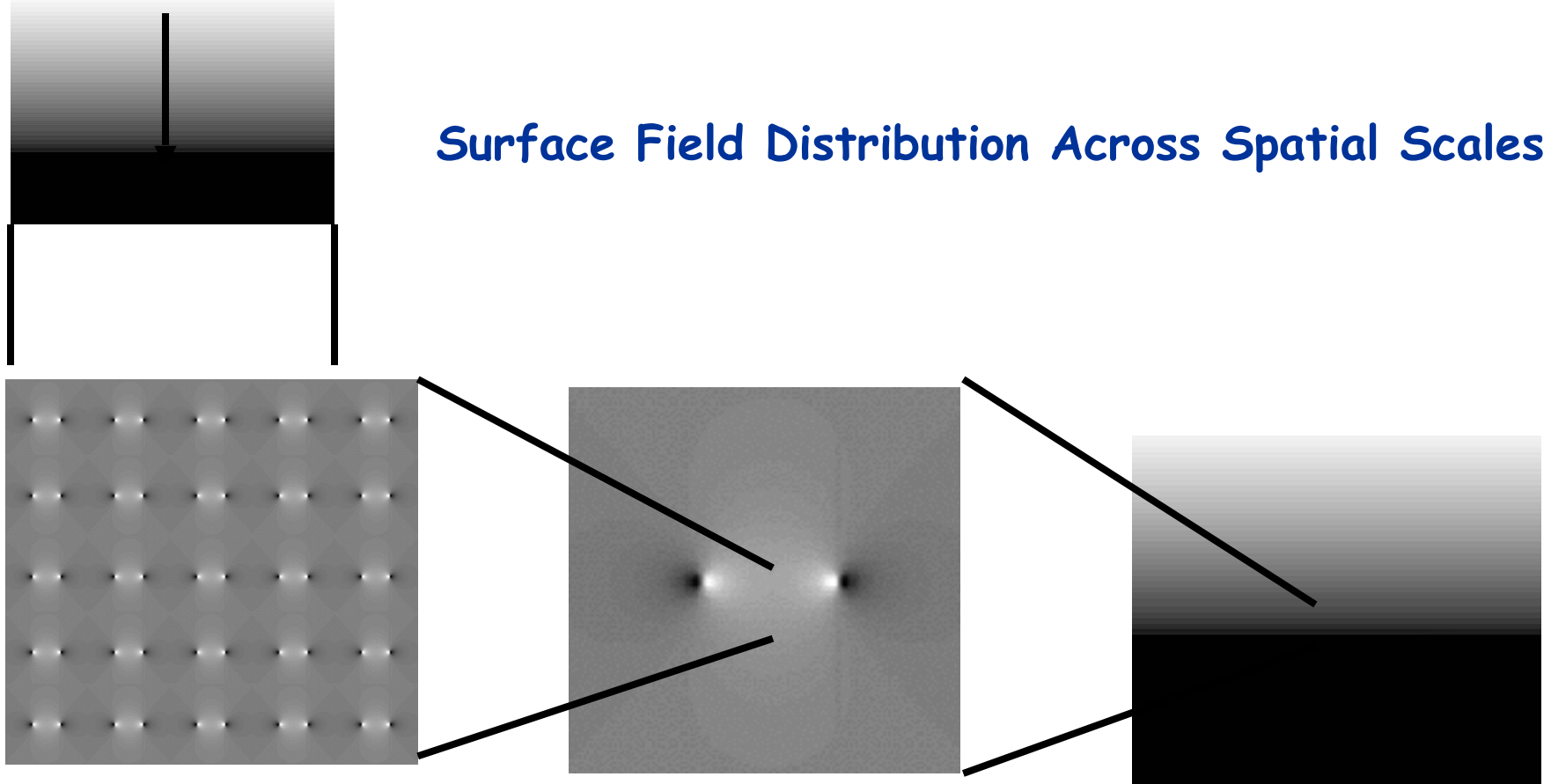
Surface Fields



100 fT at on surface of skull

J.P. Wikswo Jr et al. *J Clin
Neuronphy* 8(2): 170-188, 1991

Surface Field Distribution Across Spatial Scales

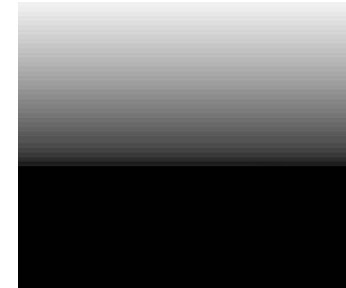


Adapted from: J.P. Wikswo Jr et al.

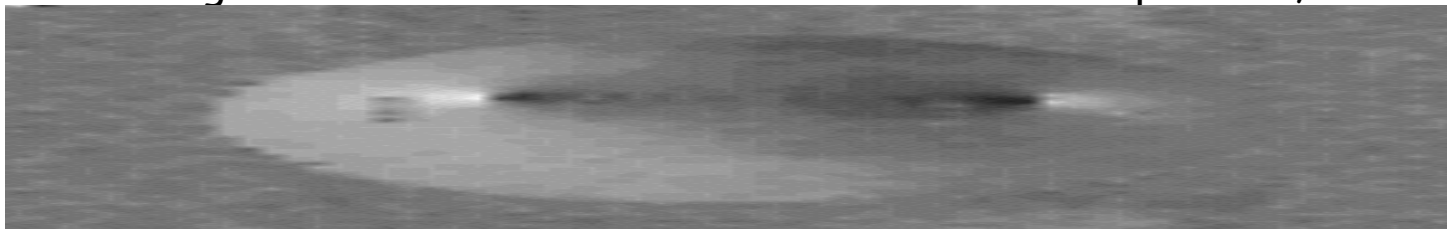
J Clin Neurophys 8(2): 170-188, 1991

Magnetic field associated with a bundle of dendrites

Because $B_{MEG} = 100 \text{ fT}$ is measured by MEG on the scalp

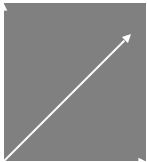
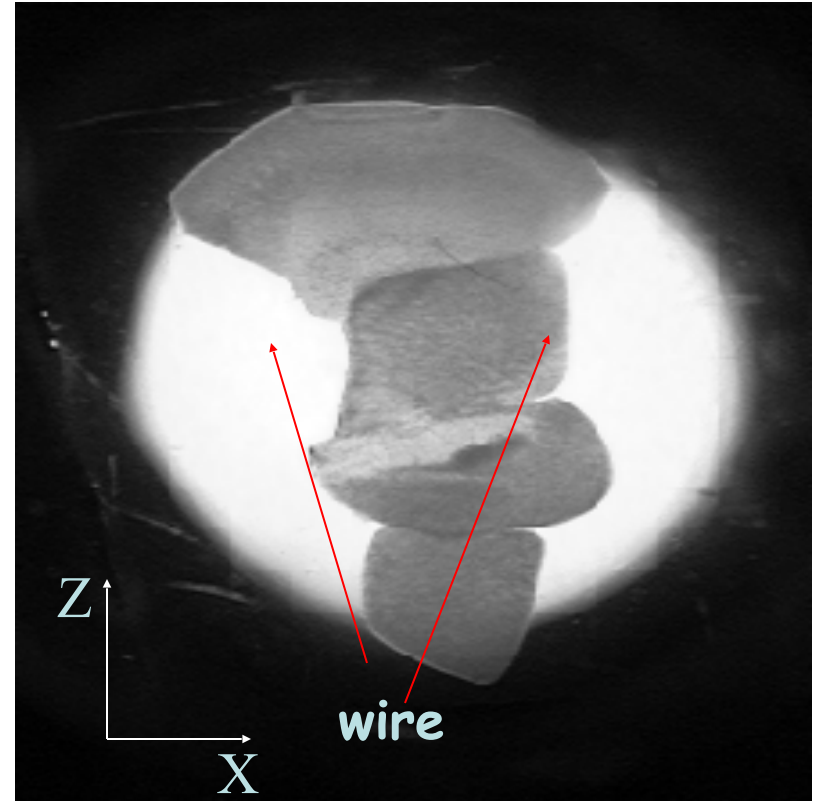
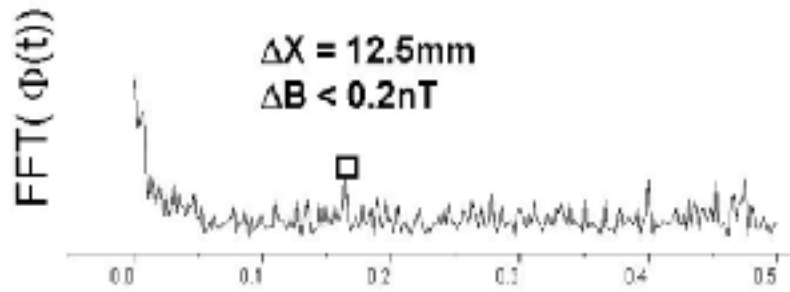
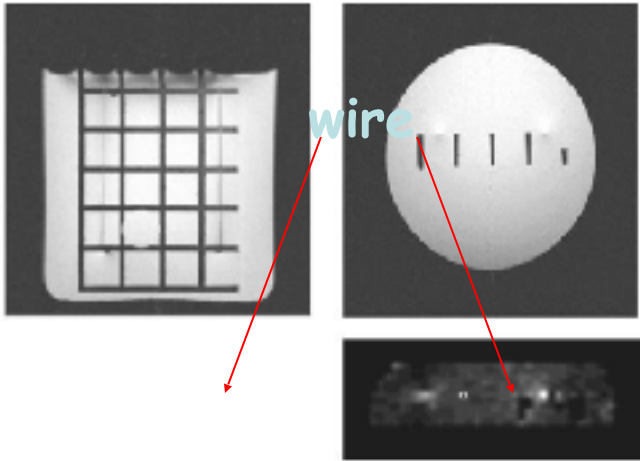


at least 50,000 neurons (0.002 fT (per dendrite) $\times 50,000 = 100 \text{ fT}$), must coherently act to generate such field. These bundles of neurons produce, within a typical voxel, 1



$B_{MRI} \approx 0.2 \text{ nT}$

Current Phantom Experiment



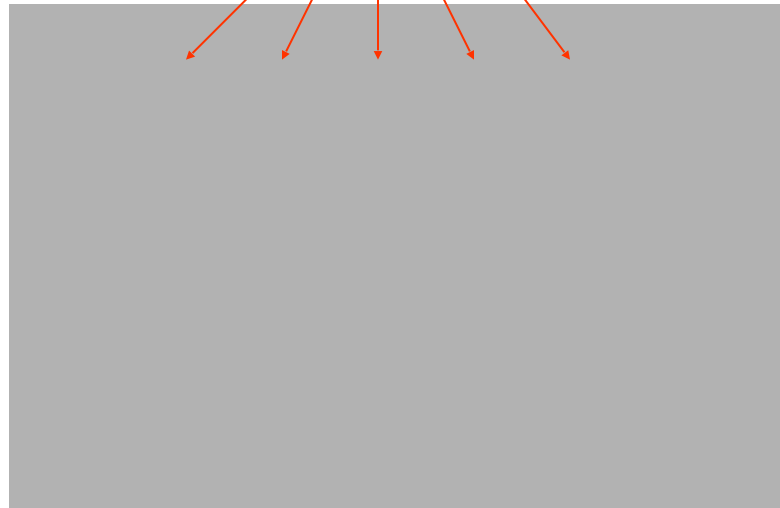
calculated $B_c \parallel B_0$

Measurement



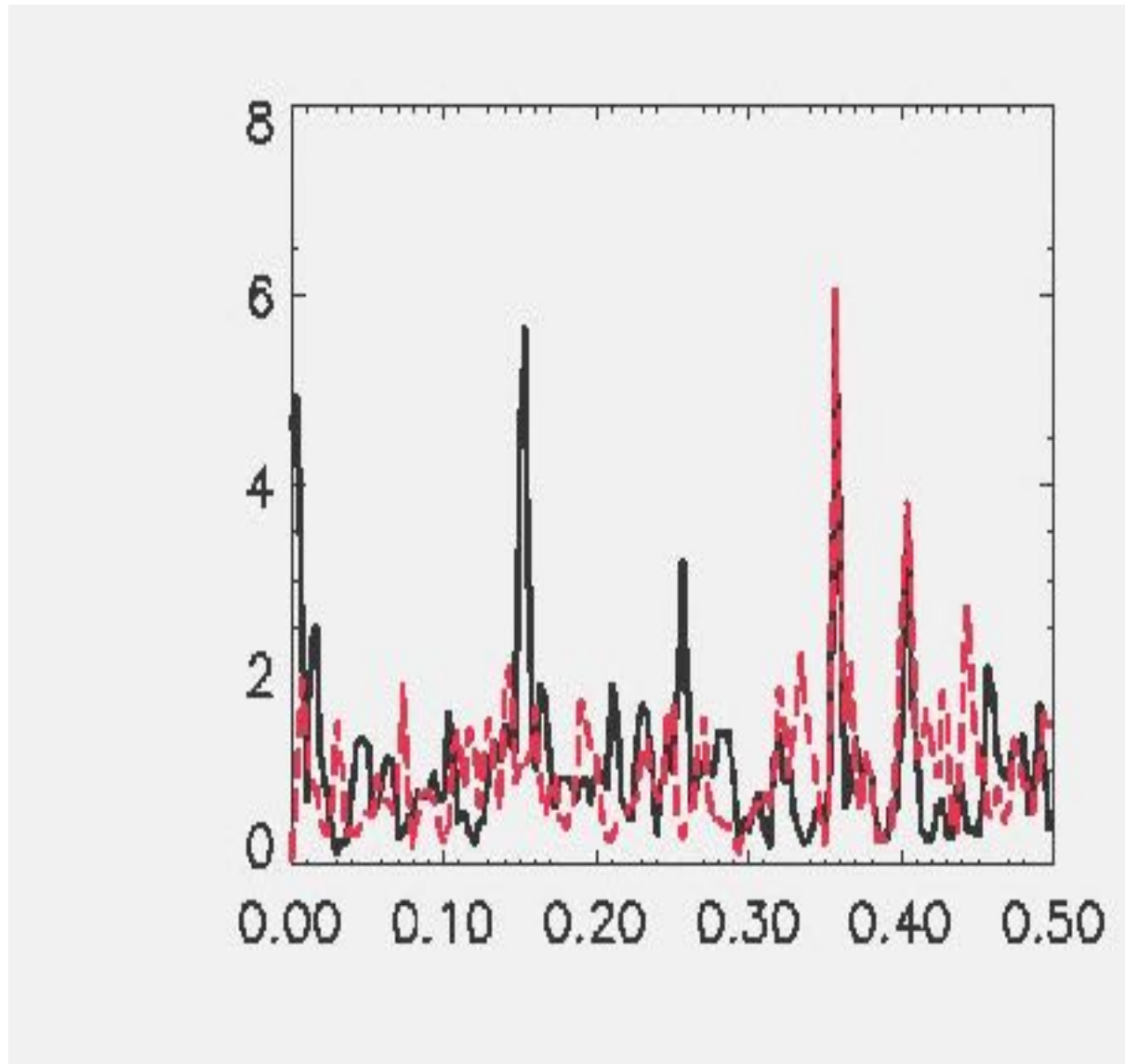
$$\Delta\phi \cong 20^\circ$$

70 μA current



Single shot GE EPI

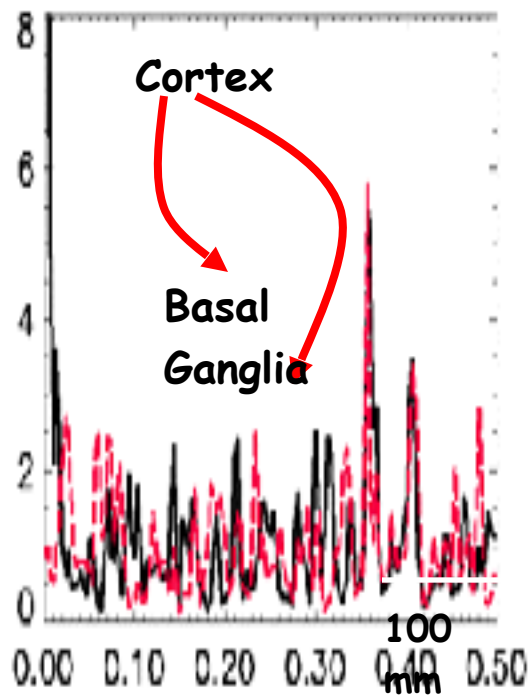
Correlation image



J. Bodurka, P. A. Bandettini. *Magn. Reson. Med.* 47: 1052-1058, (2002).

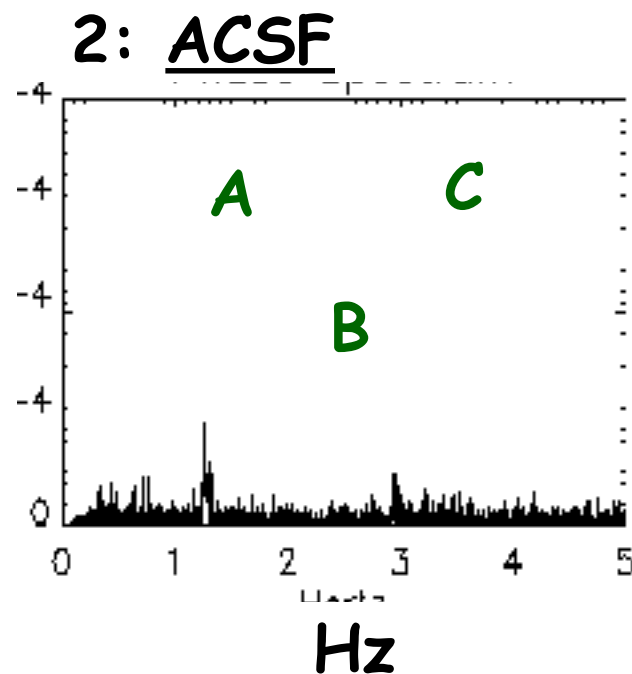
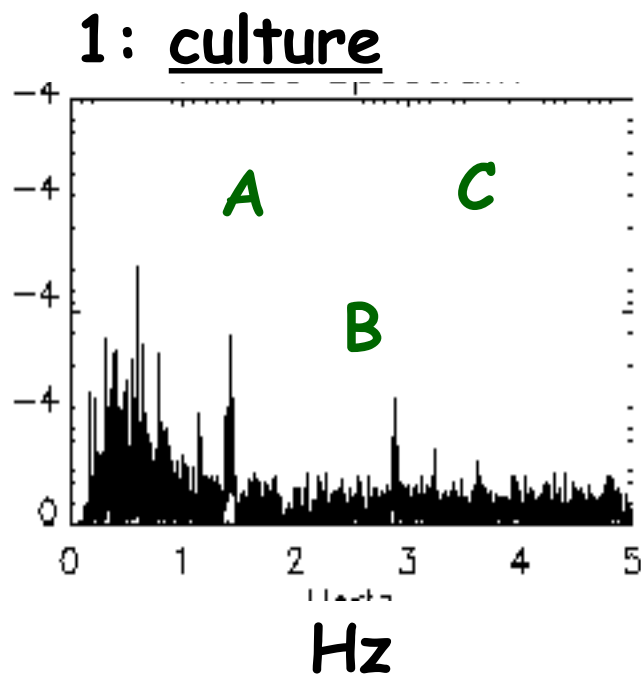
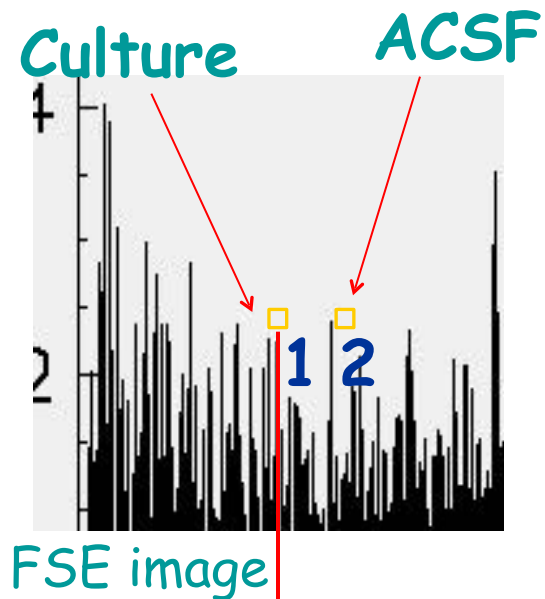
in vitro model

Organotypic (no blood supply or hemoglobin traces) sections of newborn-rat somato-sensory Cortex & Basal Ganglia



- Size: in-plane: $\sim 1-2\text{mm}^2$, thickness: $60-100\mu\text{m}$
- Neuronal Population: 10,000-100,000
- Spontaneous synchronized activity $< 2\text{Hz}$
- Epileptiform activity
- Spontaneous beta freq. activity (20-30Hz)
- Network Activity Range: $\sim 0.5-15\mu\text{V}$

3 Tesla data



Active condition: black line

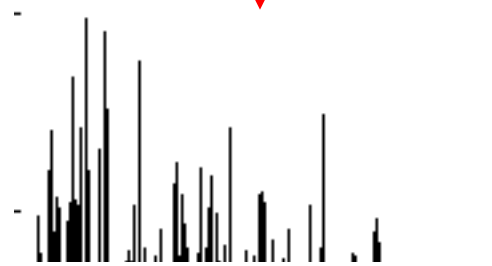
Inactive condition: red line

A: 0.15 Hz activity, on/off frequency

B: activity

C: scanner noise (cooling-pump)

0.15Hz map



7 Tesla data

TTX

Power decrease between PRE
& TTX EEG : ~ 81%

Decrease between PRE &
TTX MR phase: ~ 70%

Decrease between PRE &
TTX MR magnitude: ~ 8%

N. Petridou, D. Plenz, A. C. Silva, J. Bodurka, M. Loew, P. A.
Bandettini, Proc. Nat'l. Acad. Sci. USA. 103, 16015-16020 (2006).