

The Biggest Unknowns in Functional MRI

Peter A. Bandettini, Ph.D

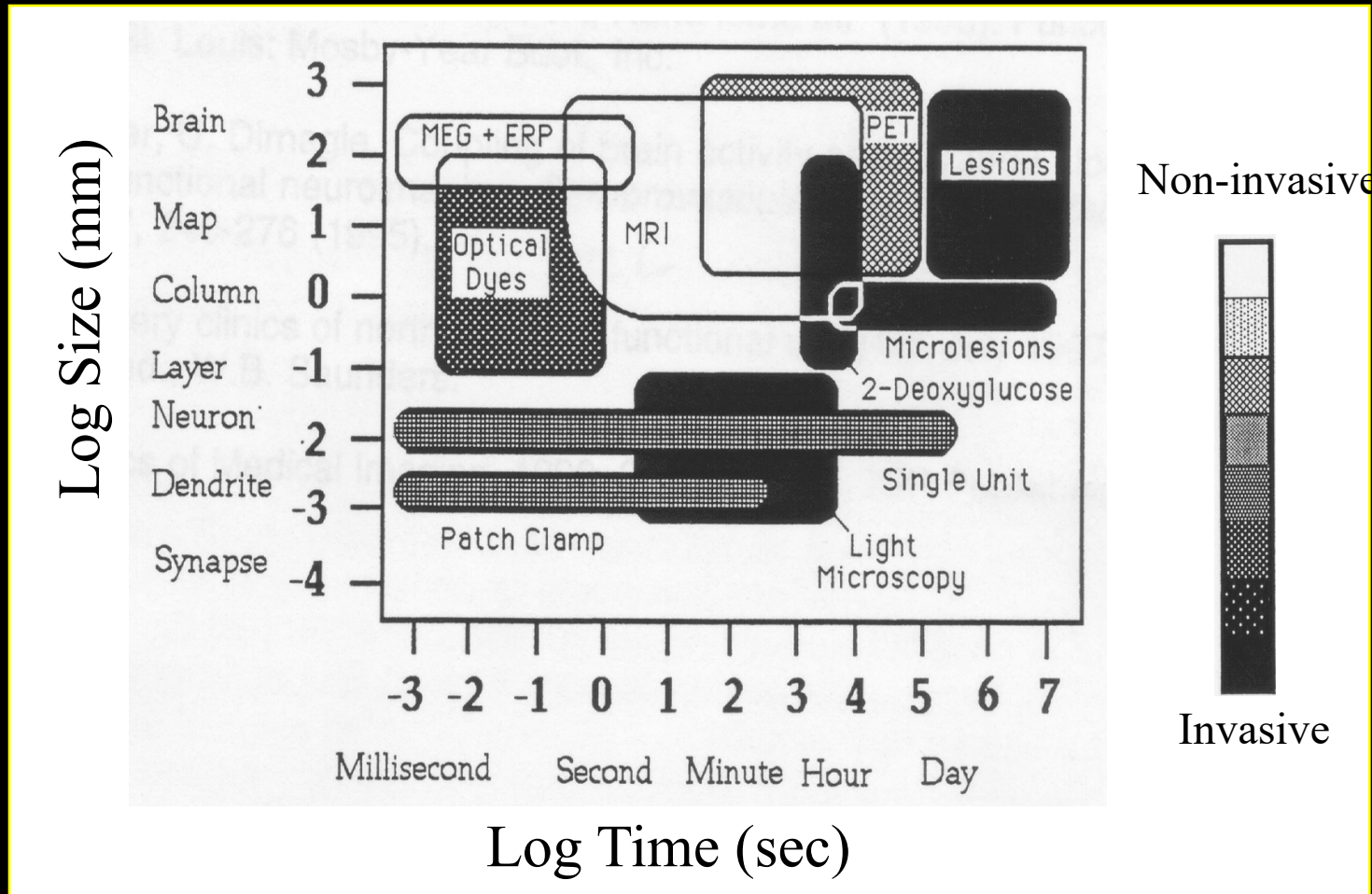
bandettini@nih.gov

Unit on Functional Imaging Methods
&
Functional MRI Facility

Laboratory of Brain and Cognition
National Institute of Mental Health



Functional Neuroimaging Techniques



Uses

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

Future Uses

Complementary use for clinical diagnosis

- utilization of clinical research results
- prediction of pathology

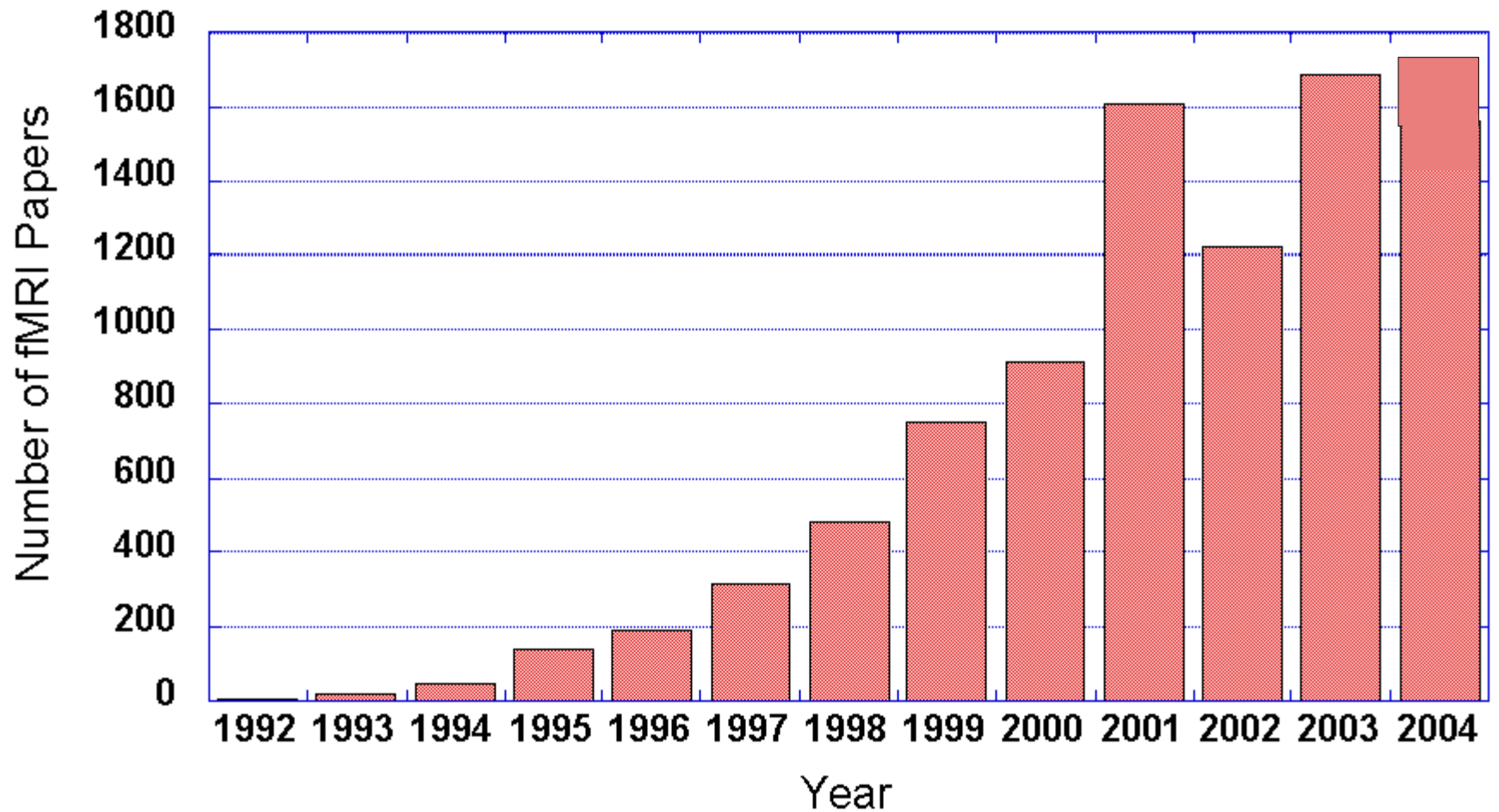
Clinical treatment and assessment

- drug, therapy, rehabilitation, biofeedback
- epileptic foci mapping
- drug effects

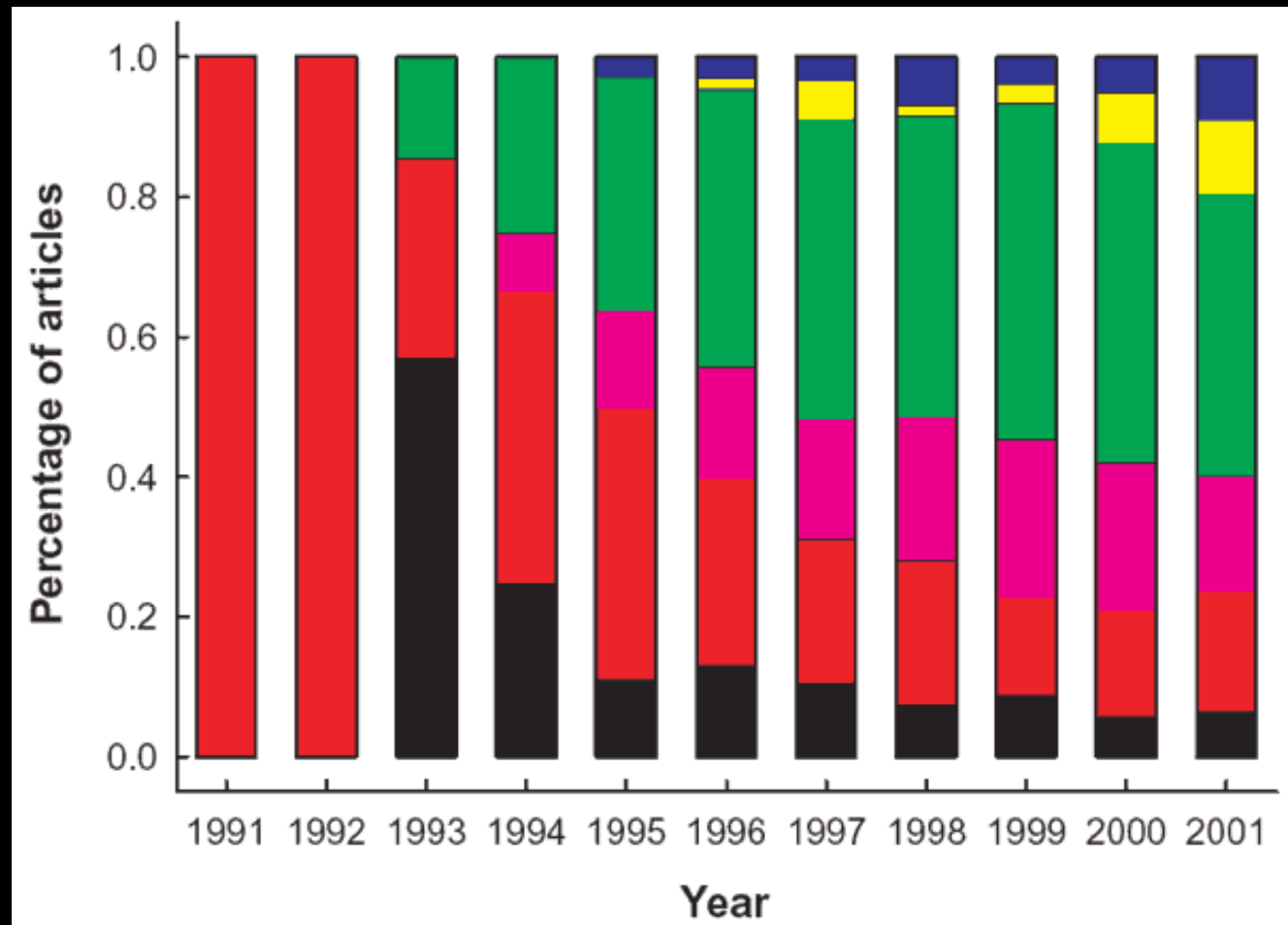
Non clinical uses

- complementary use with behavioral, anatomical, other modality results
- lie detection
- prediction of behavior tendencies
- brain/computer interface

Functional MRI Papers Published per Year



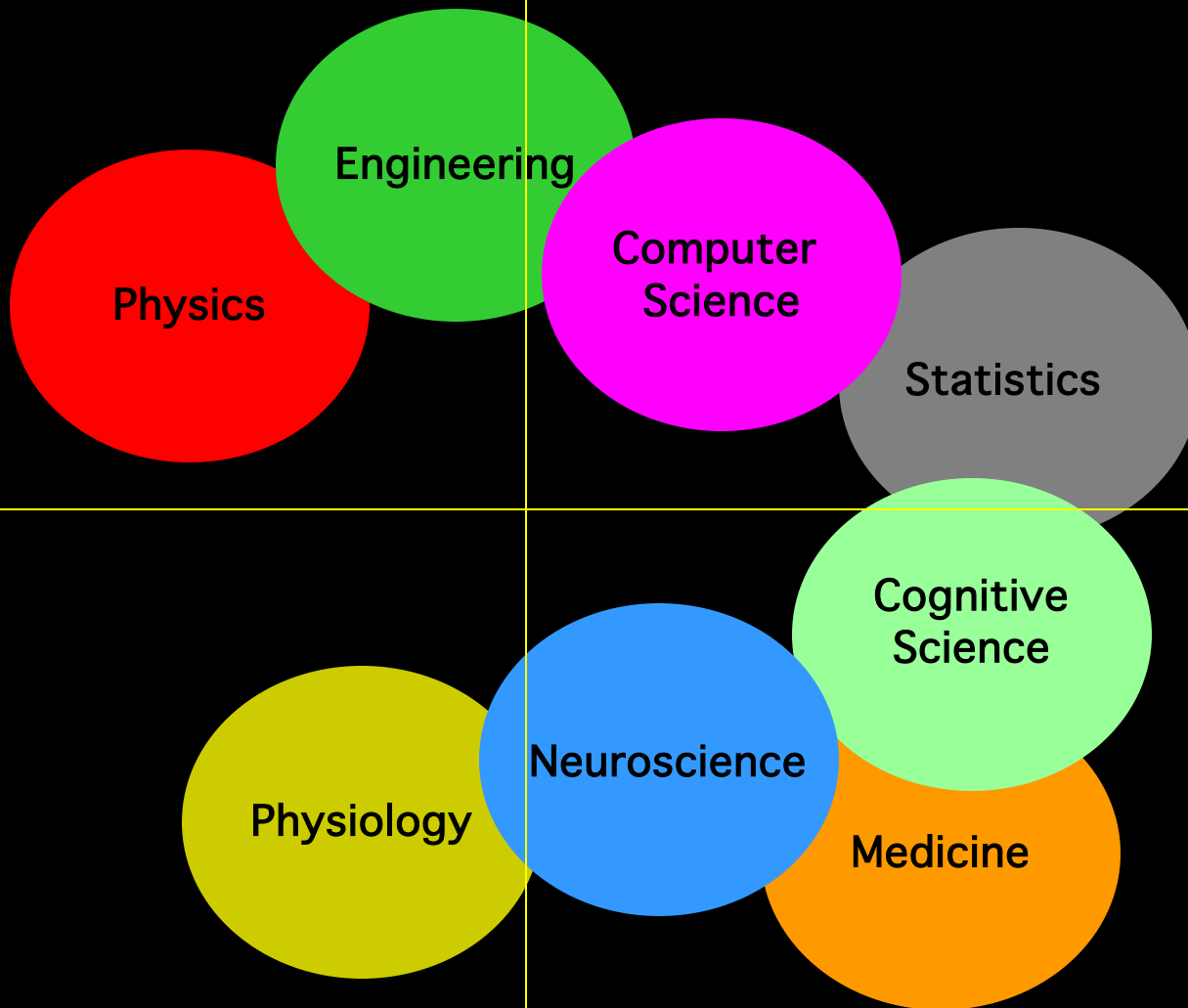
Type of fMRI research performed



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

Technology

Methodology



Interpretation

Applications

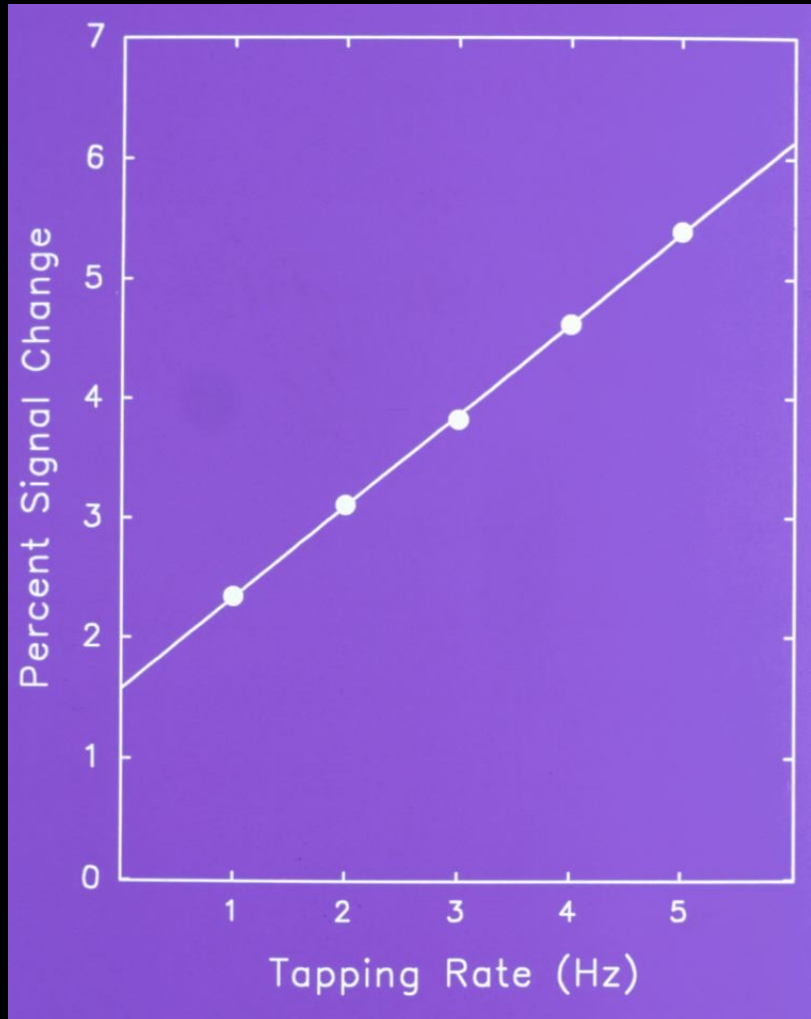
The Biggest Unknowns in Functional MRI

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

The Biggest Unknowns in Functional MRI

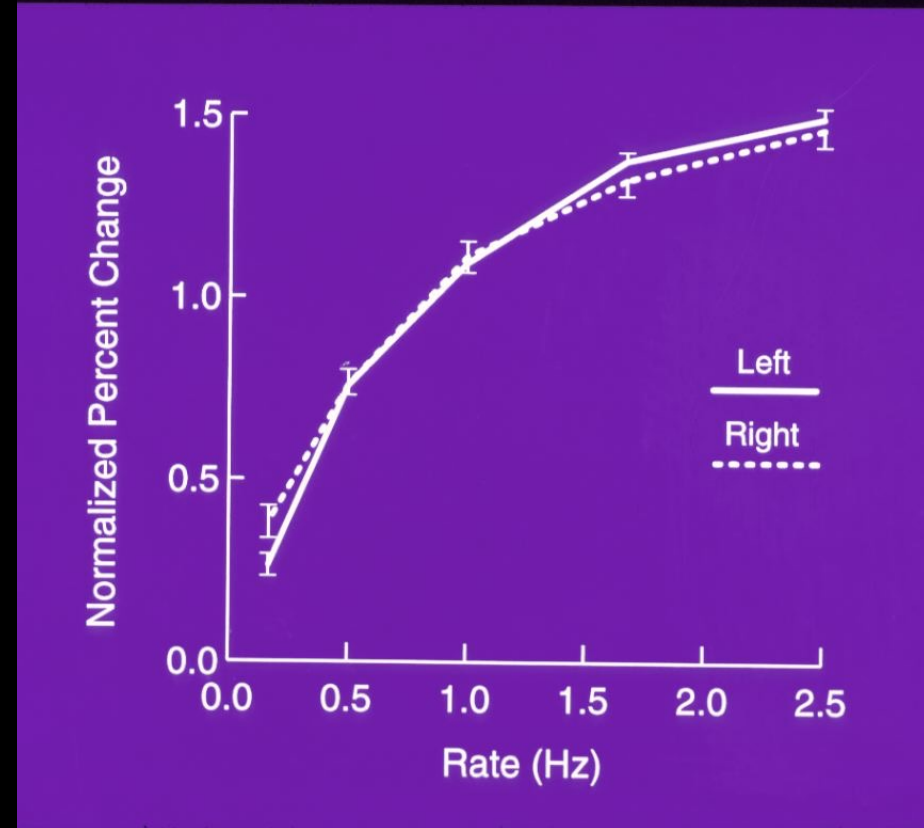
1. Relationship between neuronal activity and fMRI.
2. Sources of fMRI dynamic characteristics.
3. Sources of spatial and temporal variability.
4. What's really in the noise?
5. What's "resting" state?
6. Other sources of functional contrast?
7. Ultimate temporal resolution?
8. Ultimate spatial resolution?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

Motor Cortex



S. M. Rao et al, (1996) "Relationship between finger movement rate and functional magnetic resonance signal change in human primary motor cortex." *J. Cereb. Blood Flow and Met.* 16, 1250-1254.

Auditory Cortex

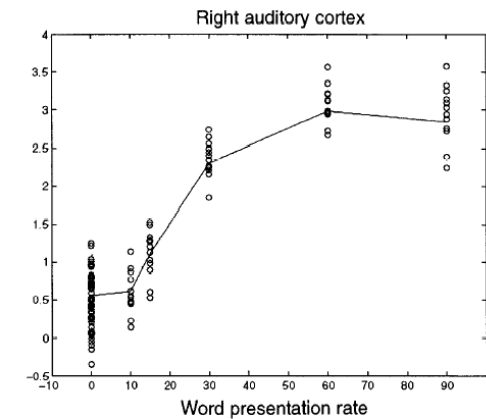
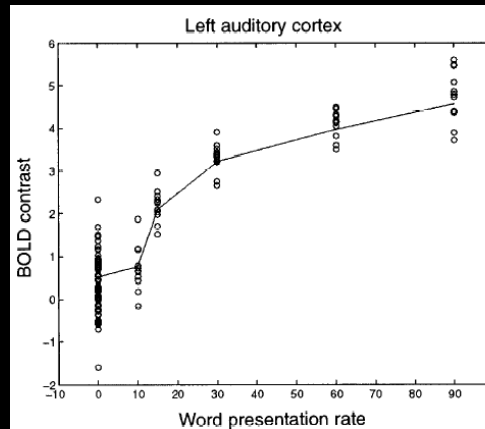
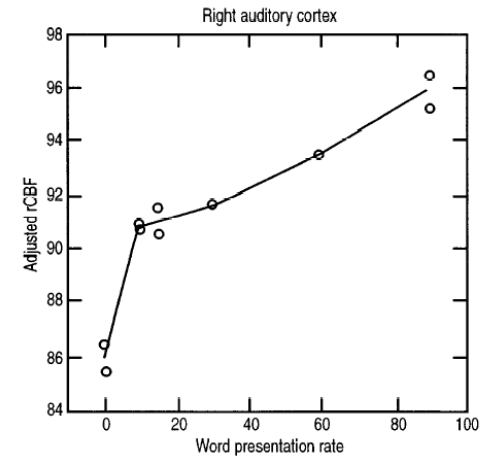


J. R. Binder, et al, (1994). "Effects of stimulus rate on signal response during functional magnetic resonance imaging of auditory cortex." *Cogn. Brain Res.* 2, 31-38

Characterizing the Relationship between BOLD Contrast and Regional Cerebral Blood Flow Measurements by Varying the Stimulus Presentation Rate

Geraint Rees, A. Howseman, O. Josephs, C. D. Frith, K. J. Friston, R. S. J. Frackowiak, and R. Turner
The Wellcome Department of Cognitive Neurology, Institute of Neurology, Queen Square, London WC1N 3BG, United Kingdom

Flow modulation is not necessarily the same as BOLD modulation



Simultaneous Recording of Evoked Potentials and T_2^* -Weighted MR Images During Somatosensory Stimulation of Rat

Gerrit Brinker, Christian Bock, Elmar Busch, Henning Krep, Konstantin-Alexander Hossmann, and Mathias Hoehn-Berlage

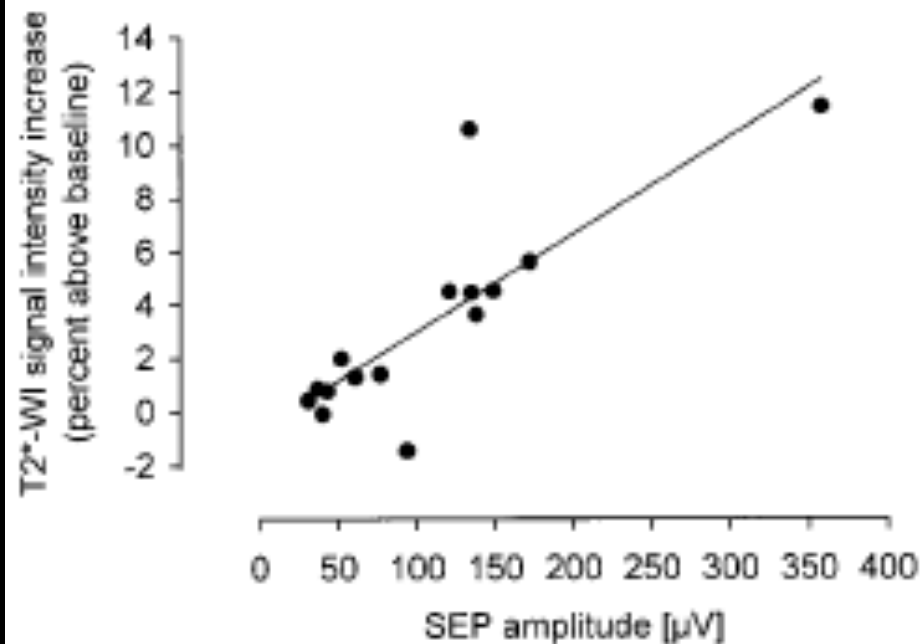
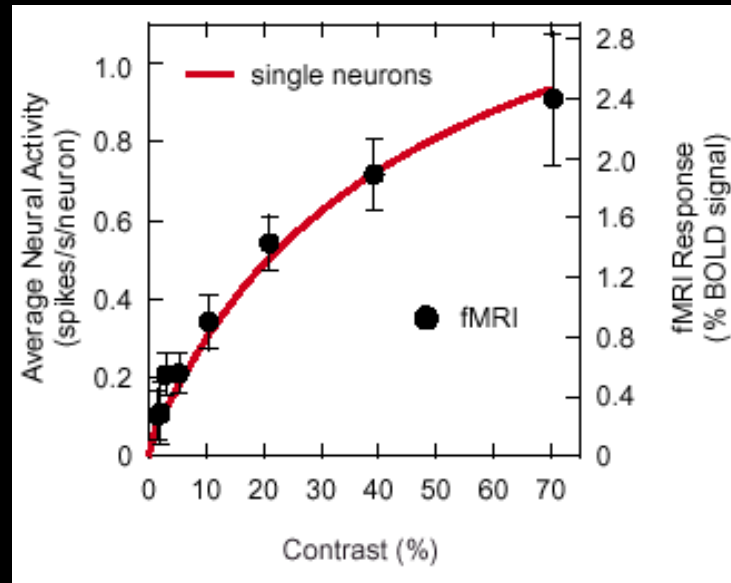


FIG. 3. Correlation of the increase of T_2^* -weighted imaging signal intensity with the peak-to-peak amplitude of the somatosensory evoked potential (SEP) during forepaw stimulation at increasing frequencies (data are from one individual animal; $r = 0.82$).

fMRI responses in human V1 are proportional to average firing rates in monkey V1?



Heeger, D. J., Huk, A. C., Geisler, W. S., and Albrecht, D. G. 2000. Spikes versus BOLD: What does neuroimaging tell us about neuronal activity? *Nat. Neurosci.* 3: 631–633.

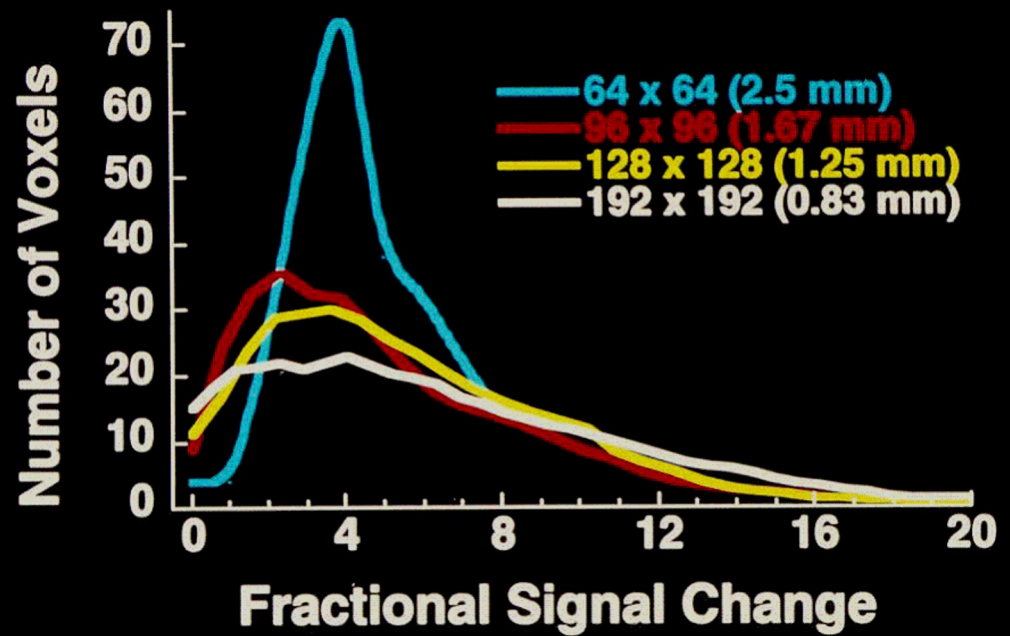
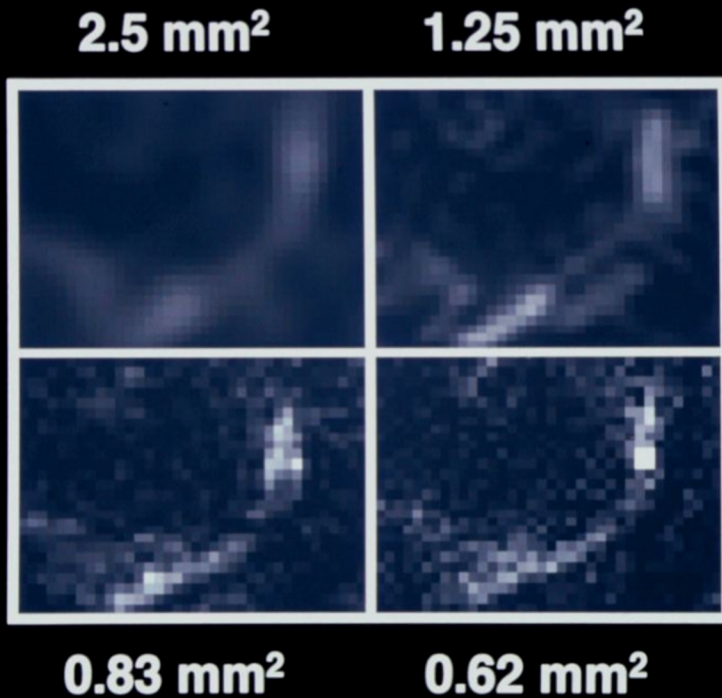
0.4 spikes/sec -> 1% BOLD

Rees, G., Friston, K., and Koch, C. 2000. A direct quantitative relationship between the functional properties of human and macaque V5. *Nat. Neurosci.* 3: 716–723.

9 spikes/sec -> 1% BOLD

Magnitude

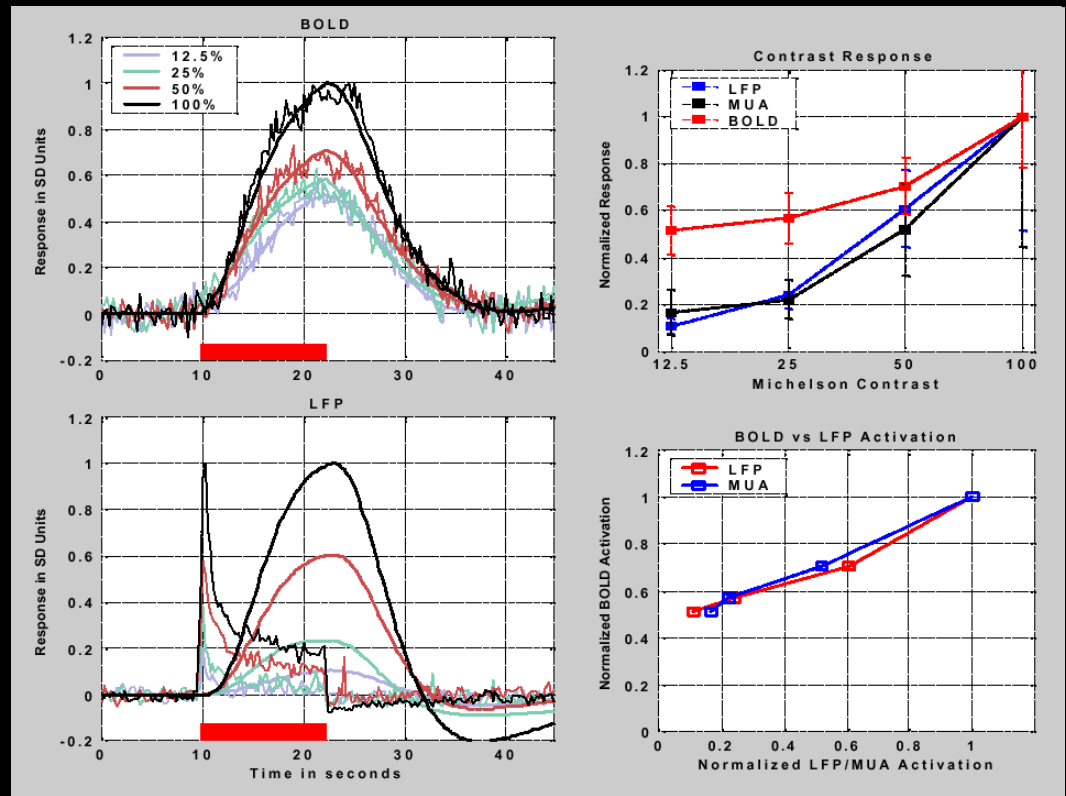
Fractional Signal Change



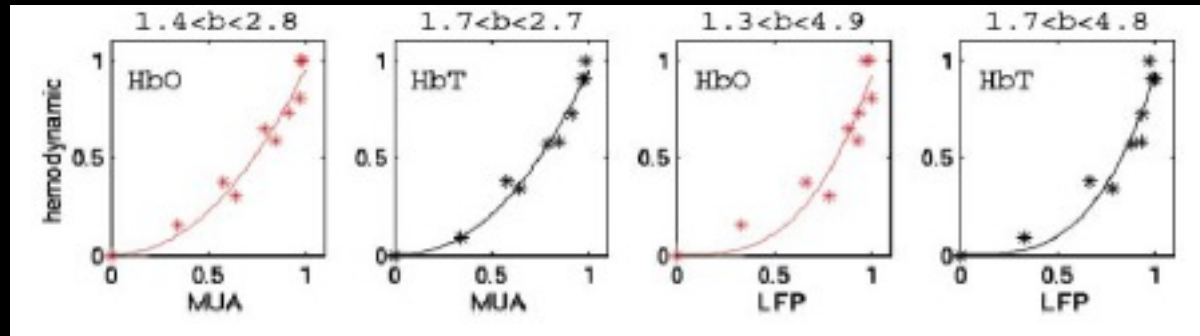
Jesmanowicz, P. A. Bandettini, J. S. Hyde, (1998) "Single shot half k-space high resolution EPI for fMRI at 3T." *Magn. Reson. Med.* 40, 754-762.

Relationship between neuronal activity and BOLD.

Magnitude



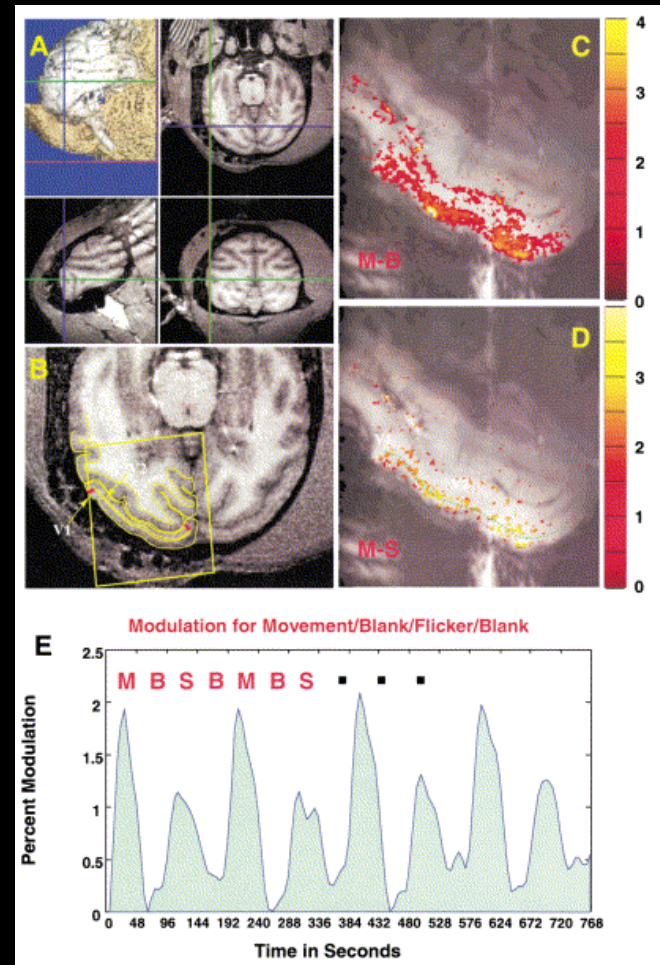
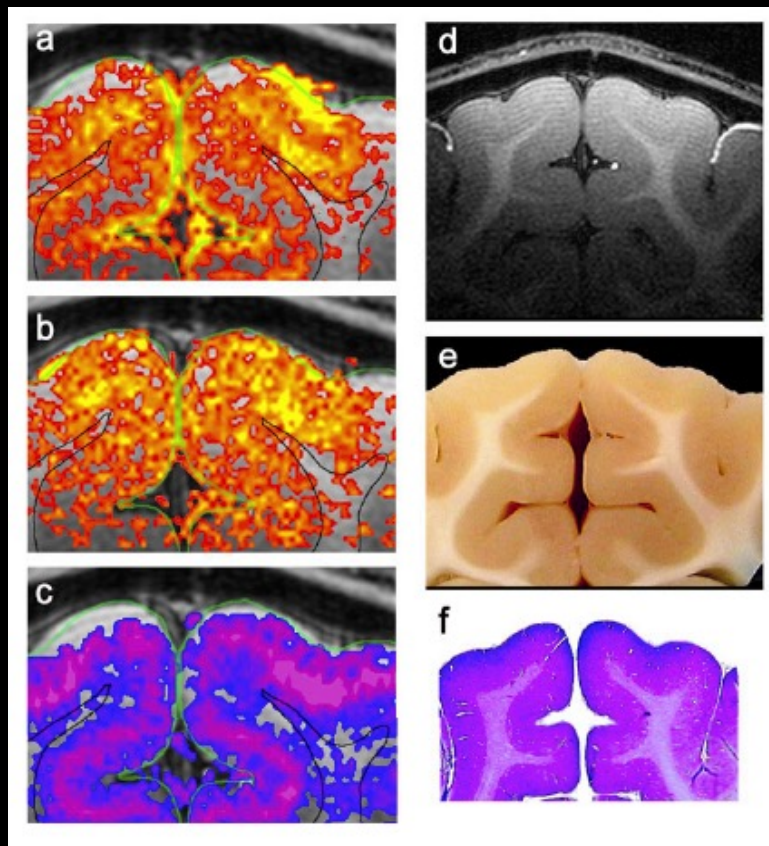
Logothetis et al. (2001) Nature, 412, 150-157



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

Relationship between neuronal activity and BOLD.

Location



Harel et al. (2004) ISMRM, 200

Logothetis et al. (2002) Neuron, 35, 227-242

T1 - weighted

Flow weighted



T2* weighted

BOLD weighted



T1 and T2* weighted

Flow and BOLD weighted

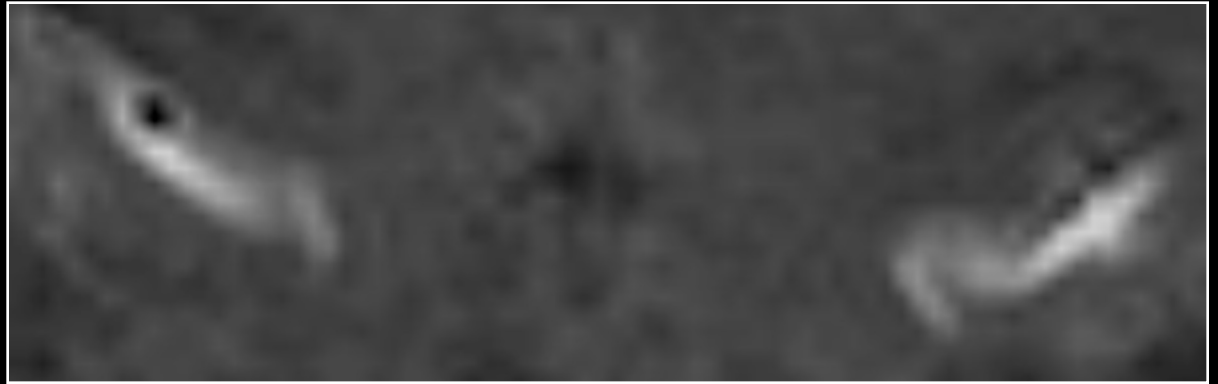


P. A. Bandettini, E. C. Wong, Echo - planar magnetic resonance imaging of human brain activation, in "Echo Planar Imaging: Theory, Technique, and Application" (F. Schmitt, M. Stehling, R. Turner, Eds.), p.493-530, Springer - Verlag, Berlin, 1997

Anatomy



BOLD



Perfusion



P. A. Bandettini, E. C. Wong, Magnetic resonance imaging of human brain function: principles, practicalities, and possibilities, *in* "Neurosurgery Clinics of North America: Functional Imaging" (M. Haglund, Ed.), p.345-371, W. B. Saunders Co., 1997.

The spatial extent of the BOLD response

Ziad S. Saad,^{a,b,*} Kristina M. Ropella,^b Edgar A. DeYoe,^c and Peter A. Bandettini^a

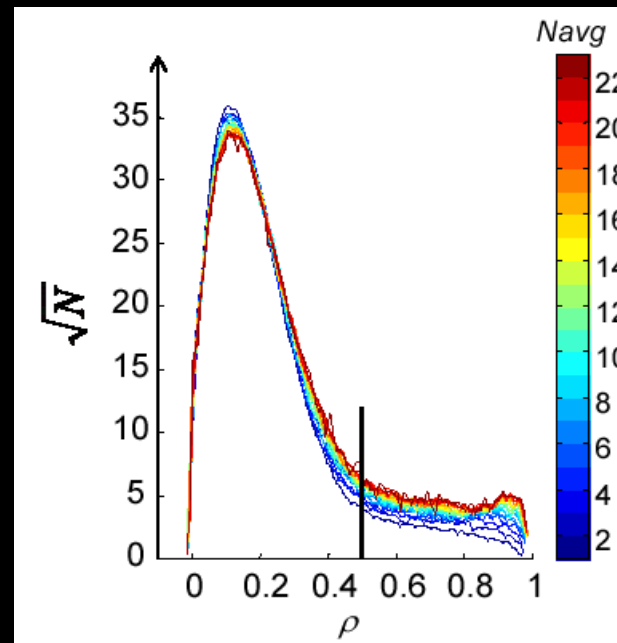
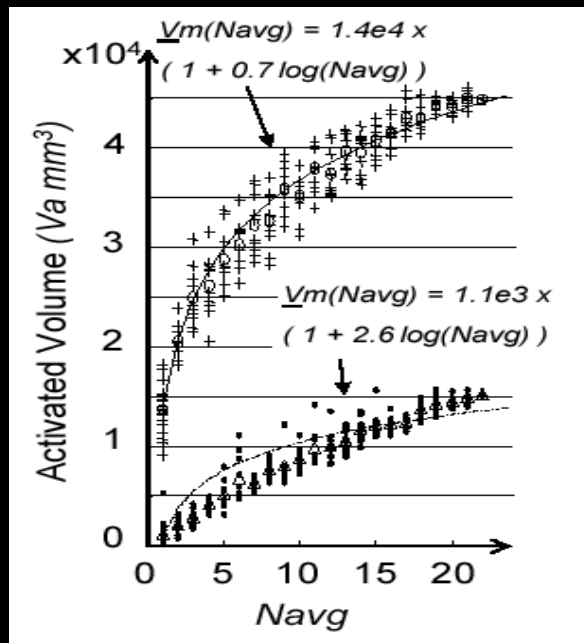
^aLaboratory of Brain and Cognition, National Institute of Mental Health, NIH, Bethesda, MD 20892-1148, USA

^bDepartment of Biomedical Engineering Marquette University, Milwaukee, WI 53233, USA

^cDepartment of Cell Biology, Neurobiology and Anatomy, Medical College of Wisconsin, Milwaukee, WI 53226, USA

Received 16 August 2002; revised 29 October 2002; accepted 21 November 2002

NeuroImage, 19: 132-144, (2003).



Task-Related Changes in Cortical Synchronization Are Spatially Coincident with the Hemodynamic Response

Krish D. Singh,*†‡ Gareth R. Barnes,* Arjan Hillebrand,* Emer M. E. Forde,* and Adrian L. Williams§

*The Wellcome Trust Laboratory for MEG Studies, Neurosciences Research Institute, Aston University, Birmingham, United Kingdom; †MARIARC, Liverpool University, Liverpool, United Kingdom; ‡Walton Centre for Neurology and Neurosurgery, Liverpool, United Kingdom; and §Department of Psychology, Royal Holloway, University of London, Egham, United Kingdom

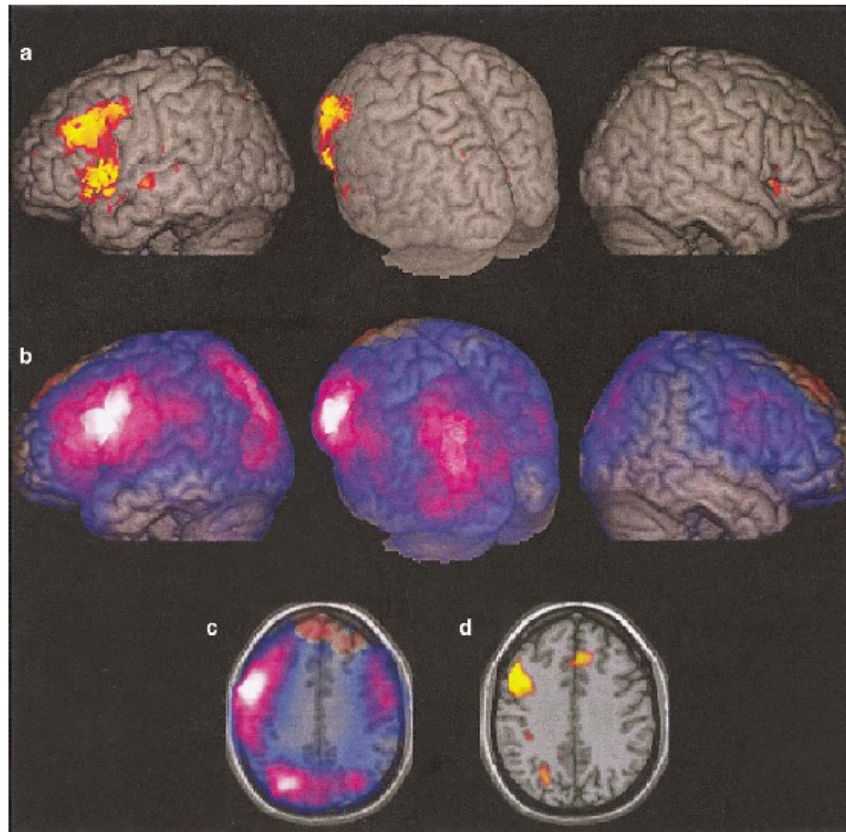


FIG. 2. The results of the group fMRI experiment and the group MEG experiment for the letter fluency task, superimposed on a template brain. The color scales are as described in the legend of Fig. 1. (a) Group fMRI data. Only those clusters significant at $P < 0.05$ (corrected) are shown. (b) The peak group SAM image. This shows the peak power increase or decrease at each voxel in the brain, irrespective of which frequency band the power change occurred in. This image can be thought of as an amalgam of Figs. 1b to 1f. (c) The peak group SAM data superimposed on a slice through the template brain at an MNI Z coordinate of +36. The image shows bilateral, but strongly left biased, activation within the dorsolateral prefrontal cortex (DLPFC) and posterior parietal cortex. (d) The group fMRI data superimposed on the $Z = +36$ slice. Note the left DLPFC and left posterior parietal activation which match the group SAM results. However, there is also a small cluster in a more anterior portion of the parietal lobe, and another in the medial frontal gyri, which are visible in the group fMRI data but not in the group MEG data.

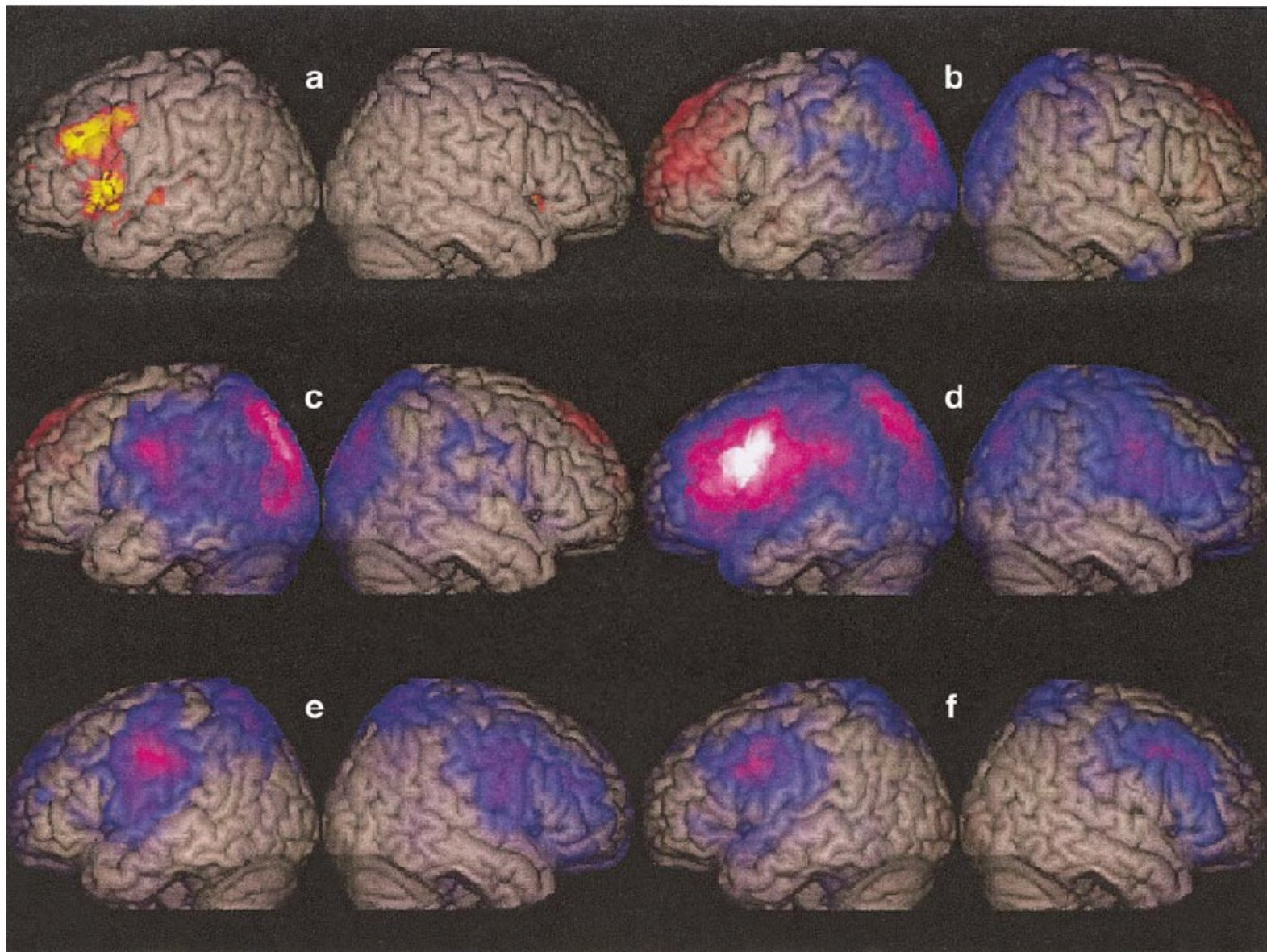
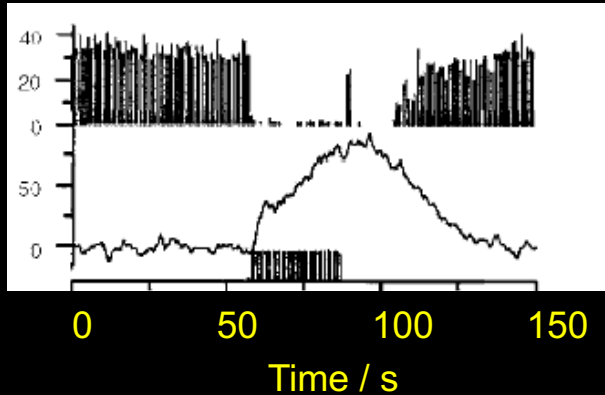


FIG. 1. The results of the group fMRI experiment and the group MEG experiment for the covert letter fluency task, superimposed on a template brain. (a) Group fMRI data. Only those clusters significant at $P < 0.05$ (corrected) are shown. The red–orange–yellow color scale depicts increasing BOLD amplitude. (b–f) The results of the group SAM analysis of the MEG data. Increases in signal power in the Active phase, compared to the Passive baseline are shown using a red–orange–yellow color scale. Decreases in signal power in the Active phase are shown using a blue–purple–white color scale. The power changes are in the following frequency bands (b) 1–10 Hz; (c) 5–15 Hz; (d) 15–25 Hz; (e) 25–35 Hz; and (f) 35–45 Hz.

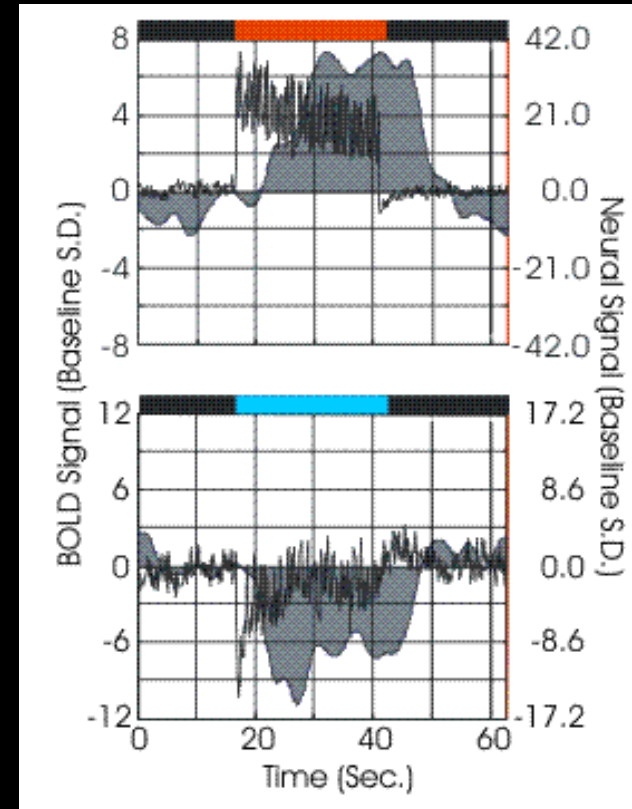
Relationship between neuronal activity and BOLD.

Inhibition



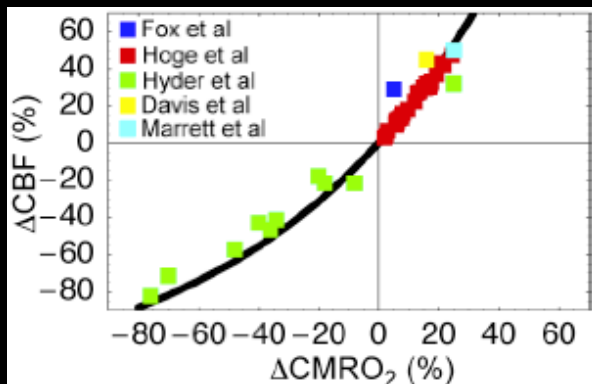
Mathiesen, et al (1998), J Physiol 512.2:555-566

Neg. BOLD



Schmuel et al. (2003) OHBM, 308

Why?



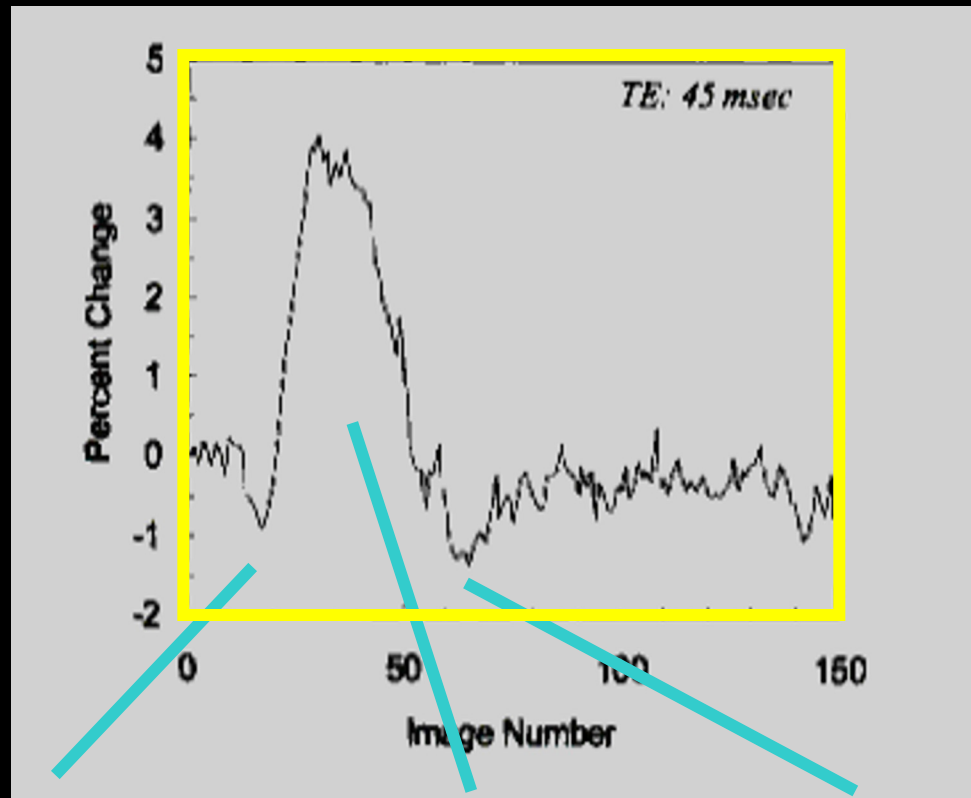
to preserve $[O_2]/[CO_2]$
at mitochondria?

Buxton (2004) ISMRM, 273

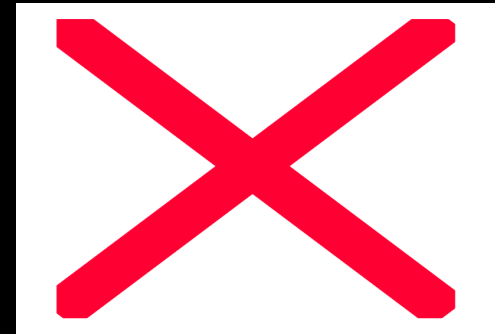
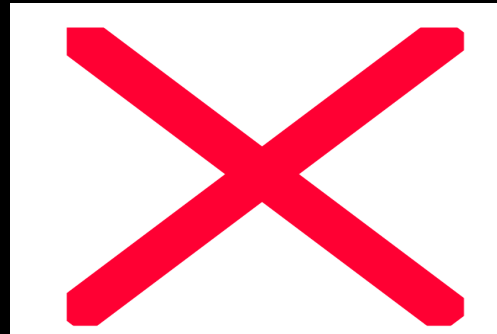
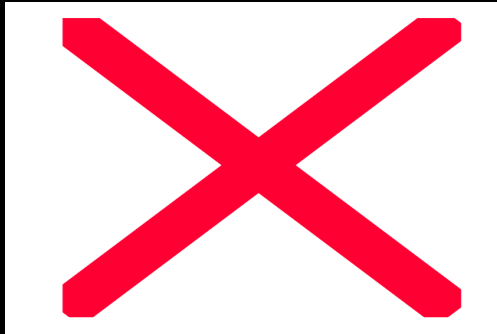
The Biggest Unknowns in Functional MRI

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

Sources of BOLD dynamic characteristics.

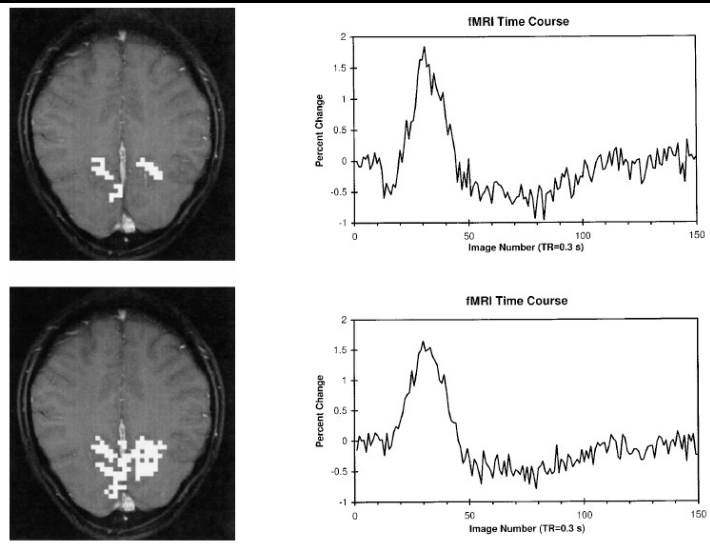


Yacoub E,
Le TH,
Ugurbil K,
Hu X
(1999)
Magn Res
Med
41(3):436-41

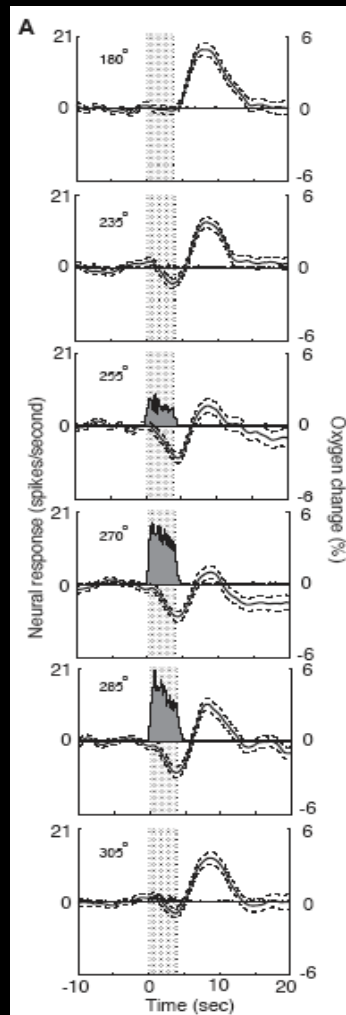


Sources of BOLD dynamic characteristics.

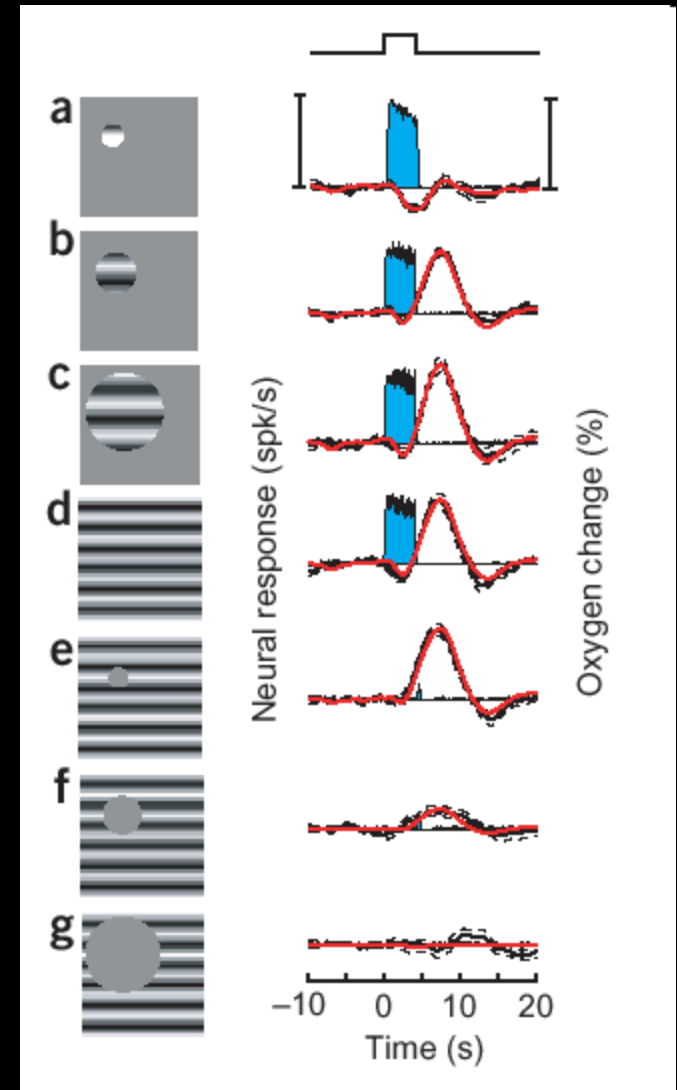
Pre-undershoot



Yacoub, et al (1999),
MRM 41, 1088-1092



Thompson, et al (2003),
Science 299, 1070-1072

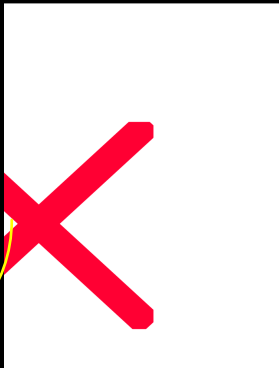


Thompson, et al (2004),
Nature Neuroscience
7, 919-920

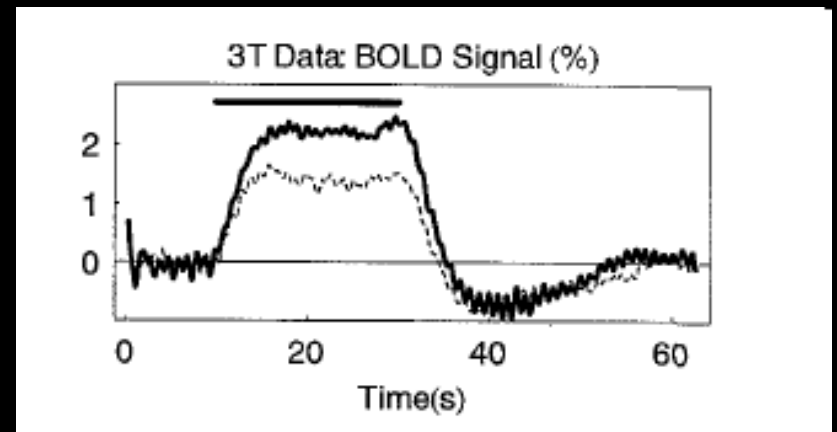
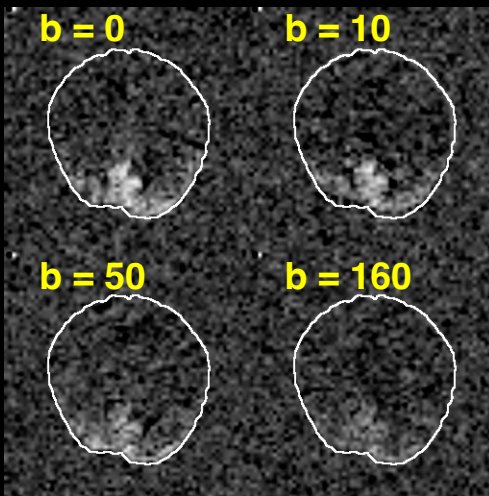
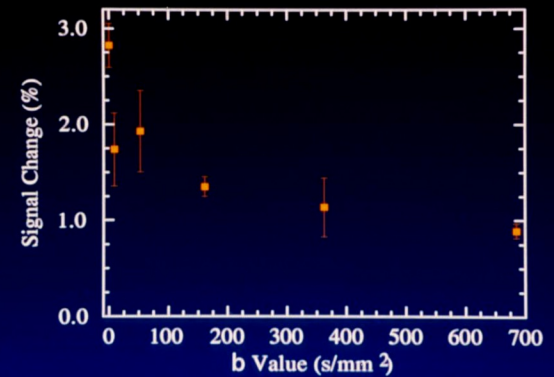
Post-undershoot

no diffusion weighting

diffusion weighting



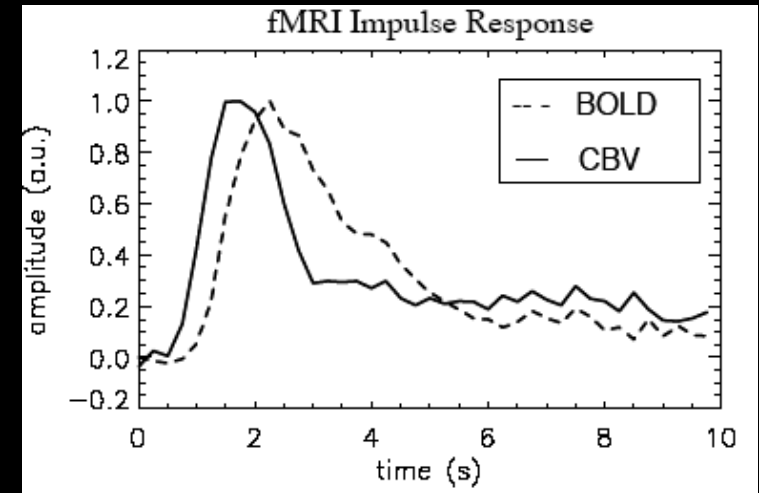
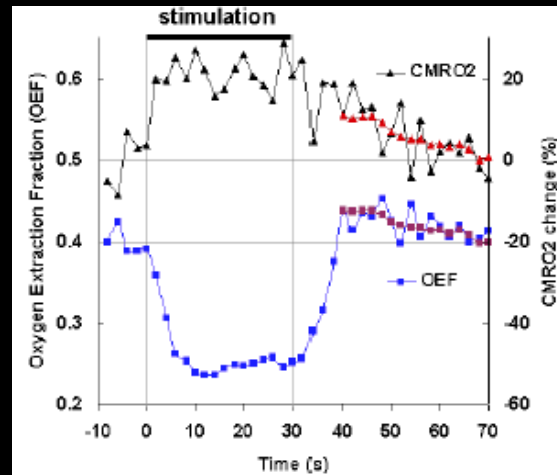
Summary of Diffusion-Weighted fMRI Data



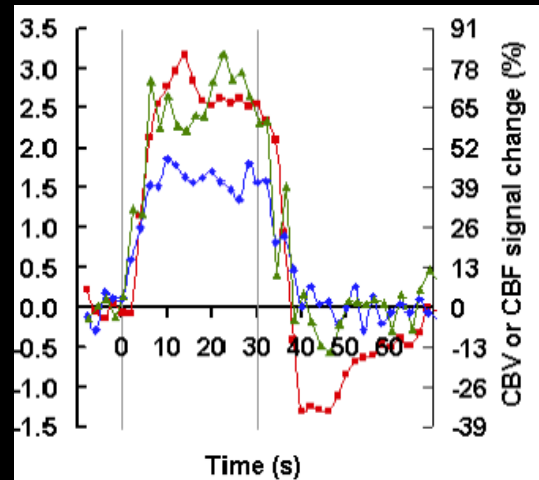
Buxton, et al (1998), ISMRM 7



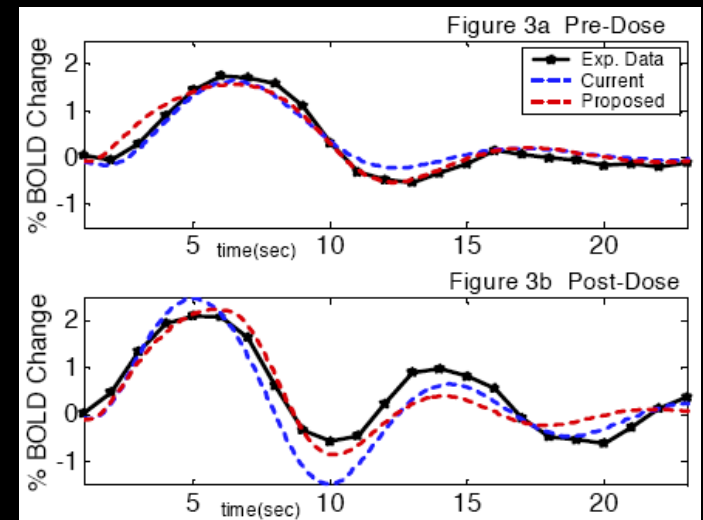
Sources of BOLD dynamic characteristics.



Silva, et al (2004), ISMRM 277



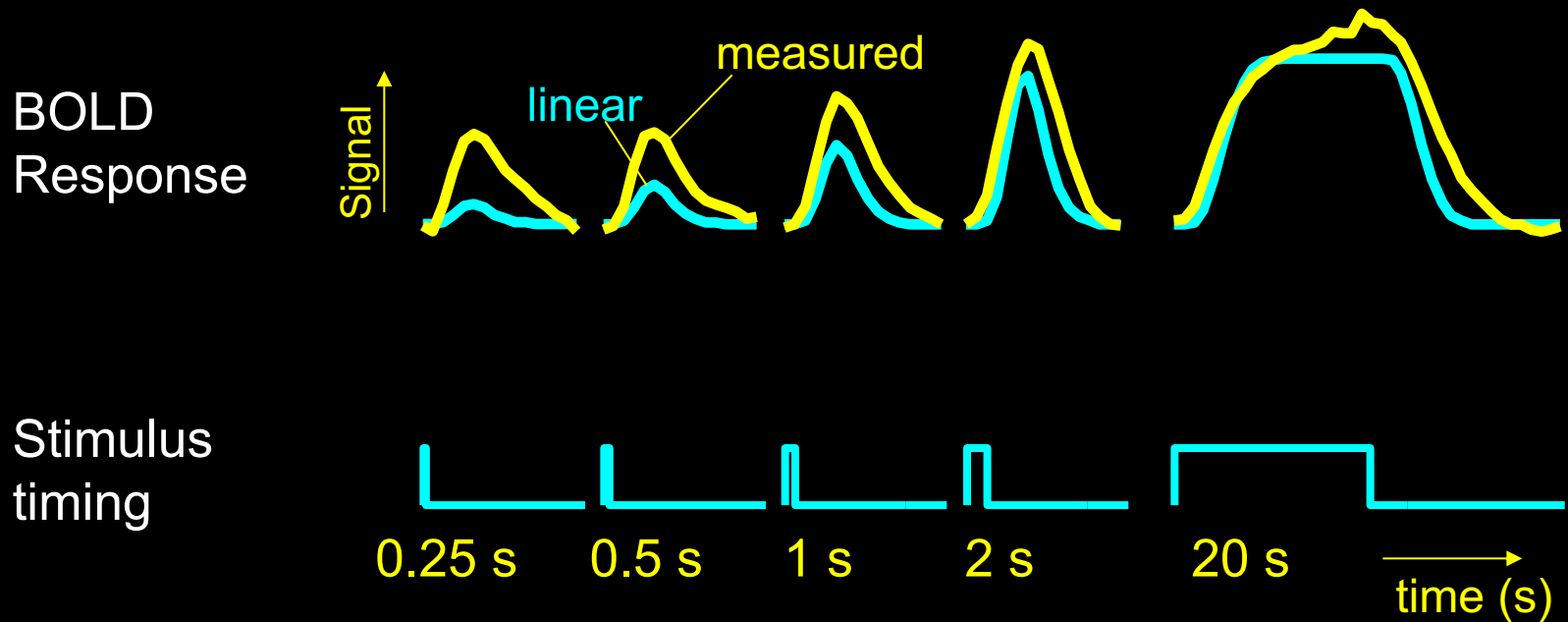
Lu, et al (2004), ISMRM 271



Behzadi, et al (2004), ISMRM 279

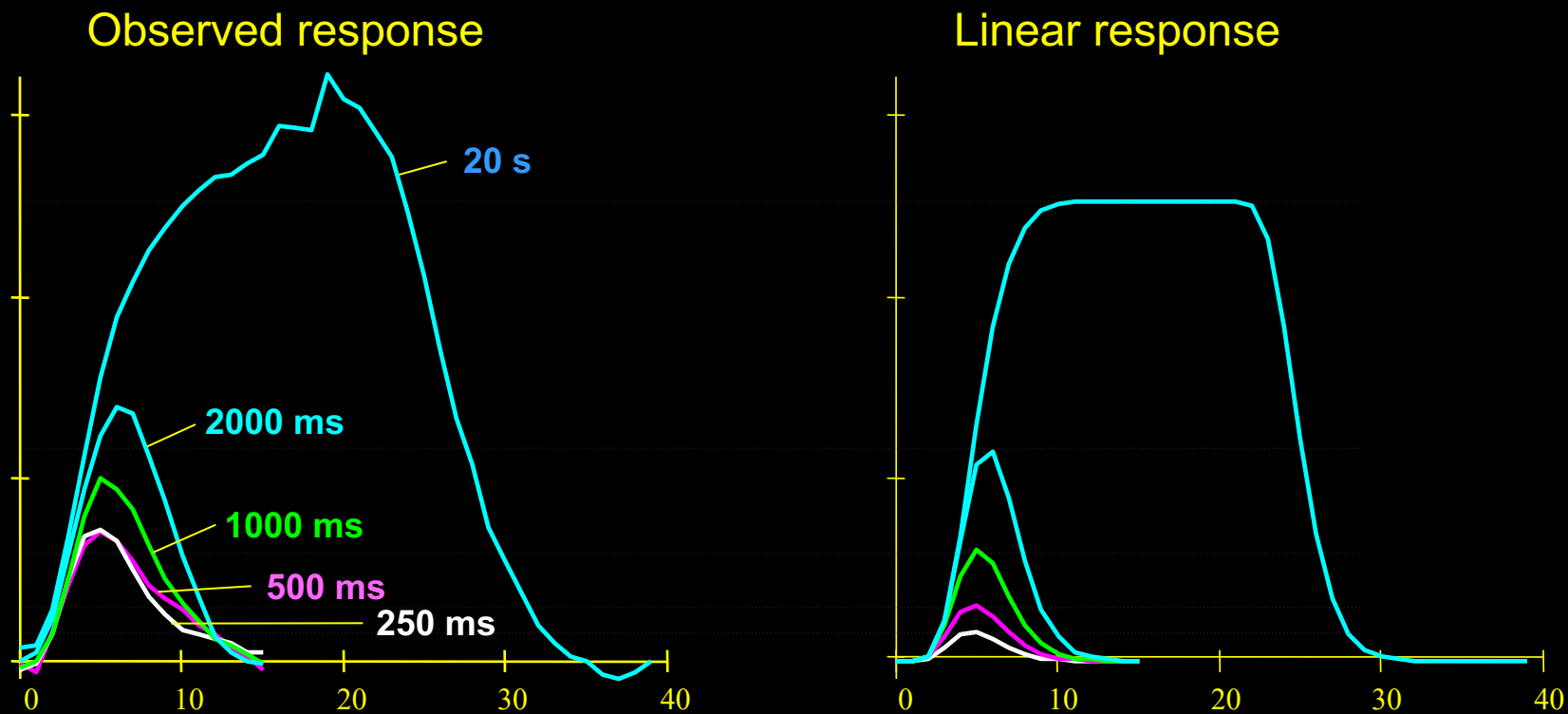
Dynamic Nonlinearity Assessment

Different stimulus “ON” periods



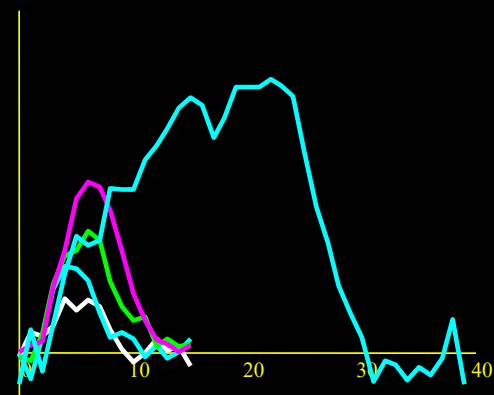
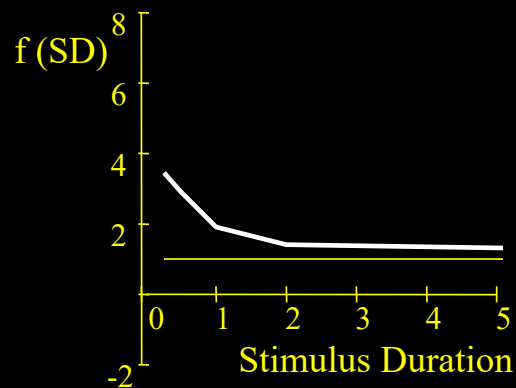
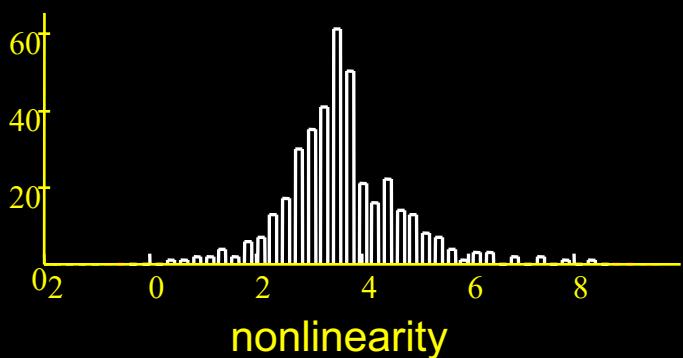
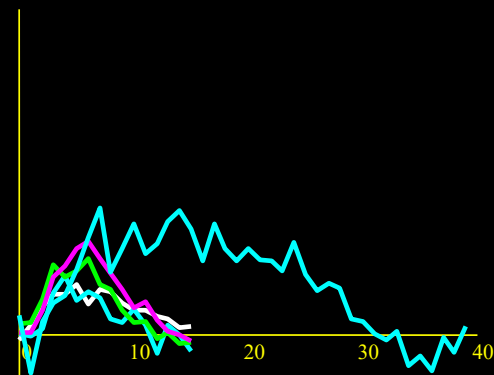
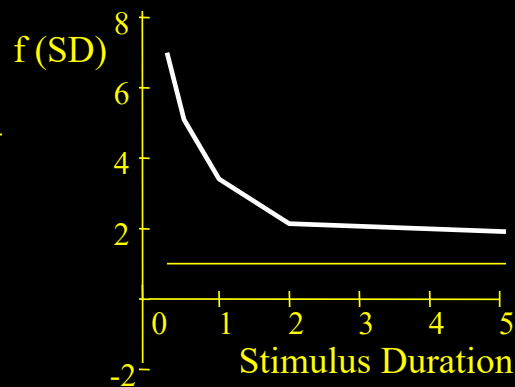
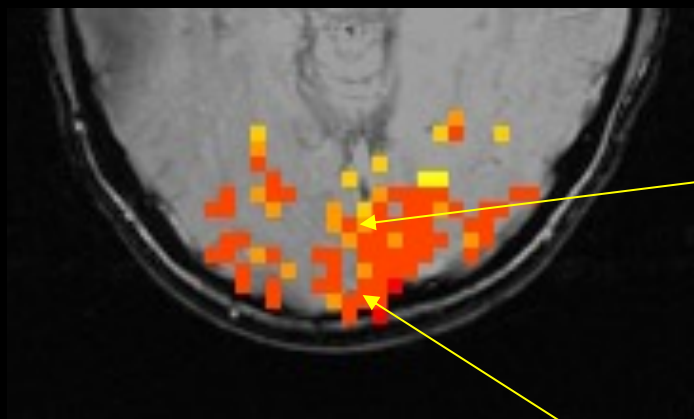
Brief stimuli produce larger responses than expected

BOLD response is nonlinear



Short duration stimuli produce larger responses than expected

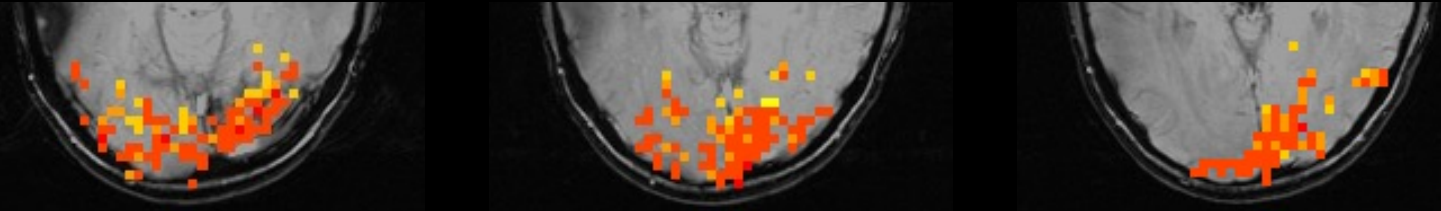
Spatial Heterogeneity of BOLD Nonlinearity



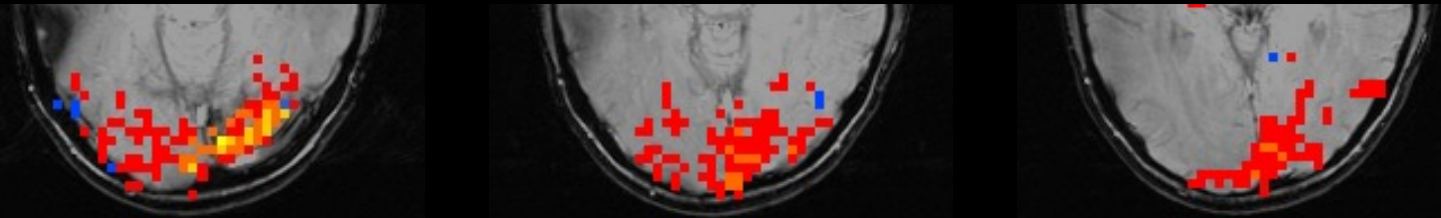
R. M. Birn, Z. Saad, P. A. Bandettini, (2001) "Spatial heterogeneity of the nonlinear dynamics in the fMRI BOLD response." *NeuroImage*, 14: 817-826.

Results – visual task

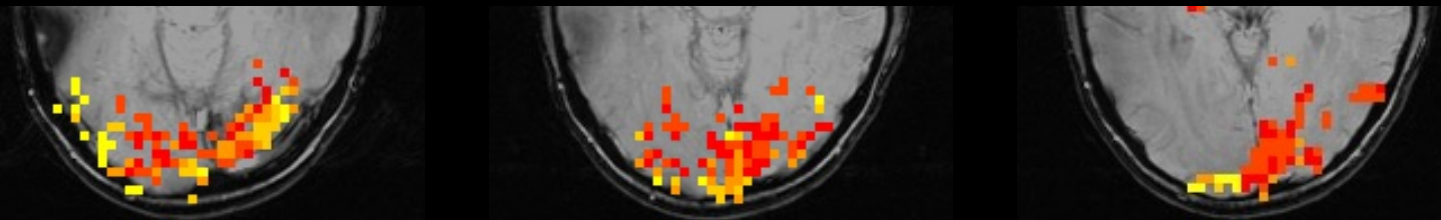
Nonlinearity



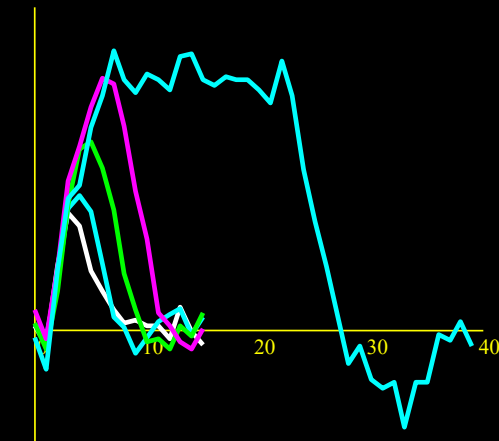
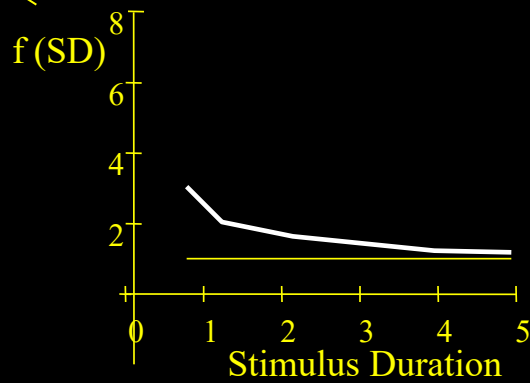
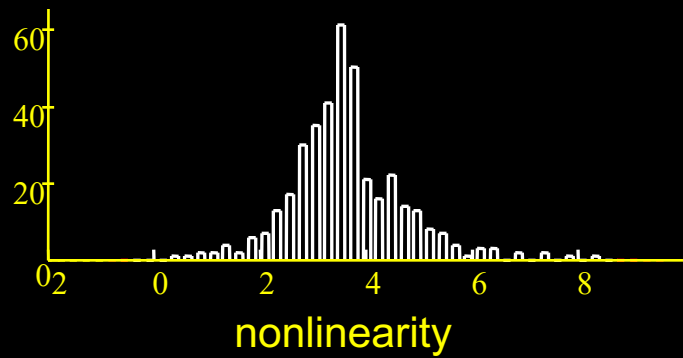
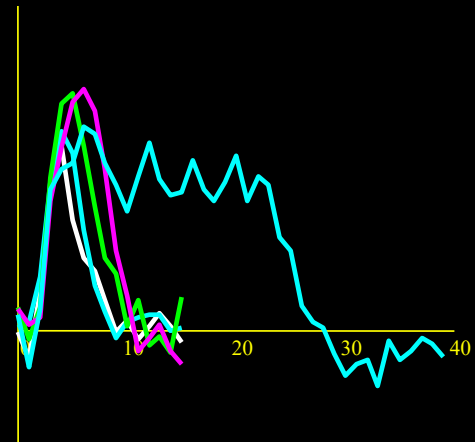
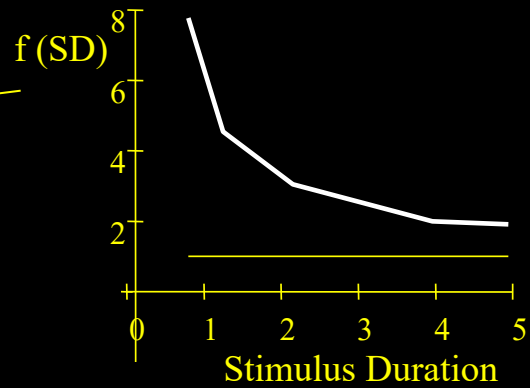
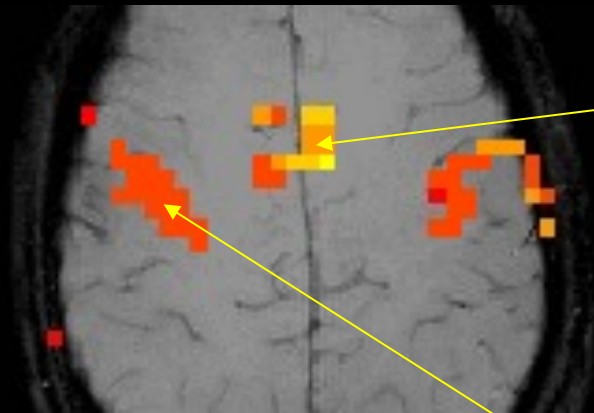
Magnitude



Latency

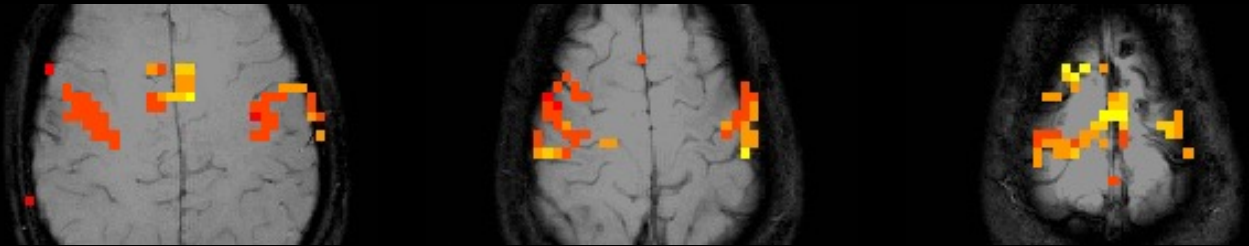


Results — motor task

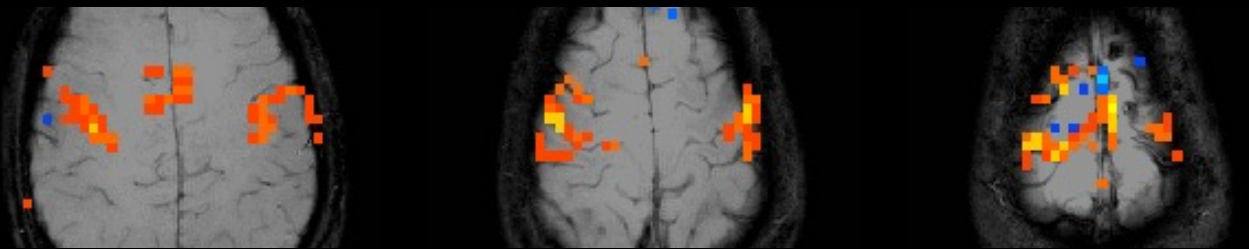


Results — motor task

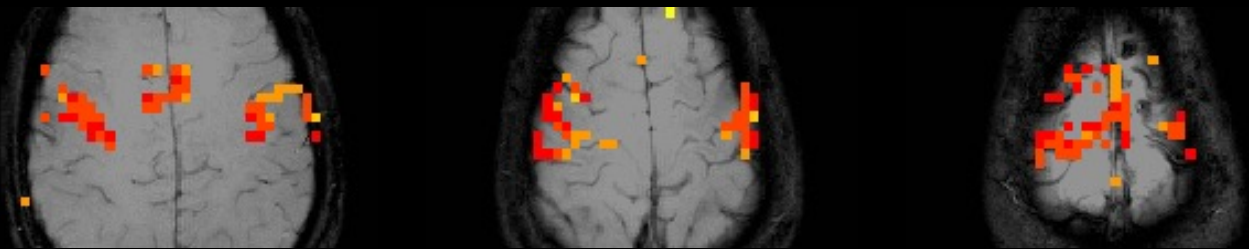
Nonlinearity



Magnitude

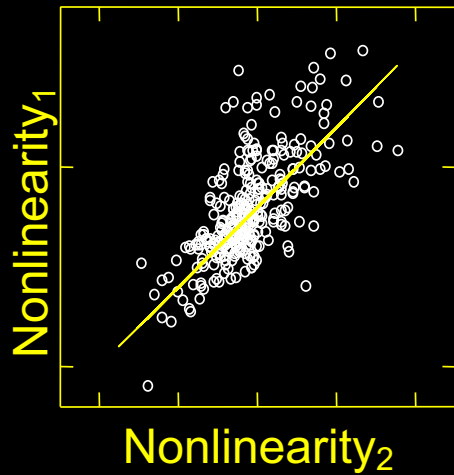


Latency

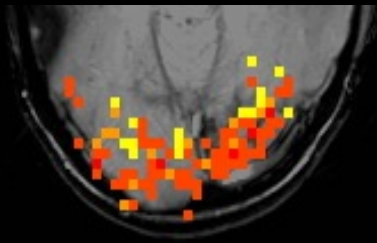
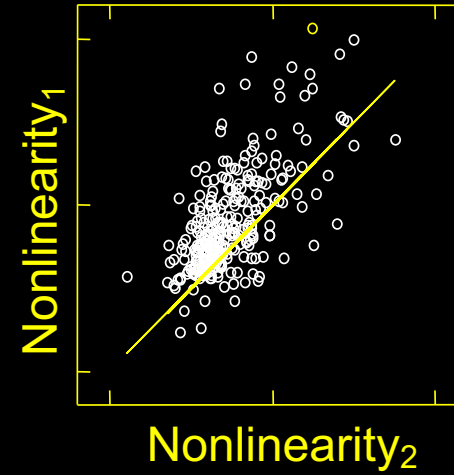


Reproducibility

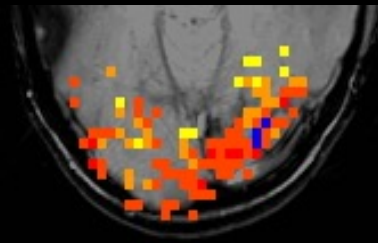
Visual task



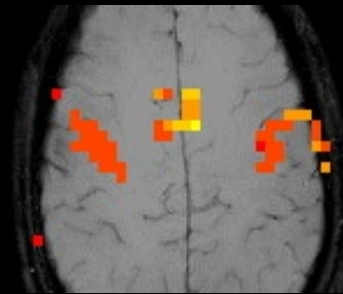
Motor task



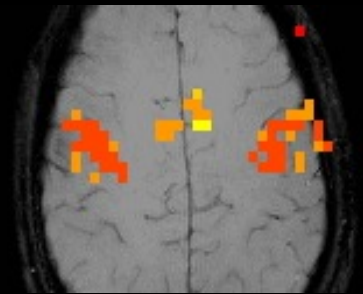
Experiment 1



Experiment 2

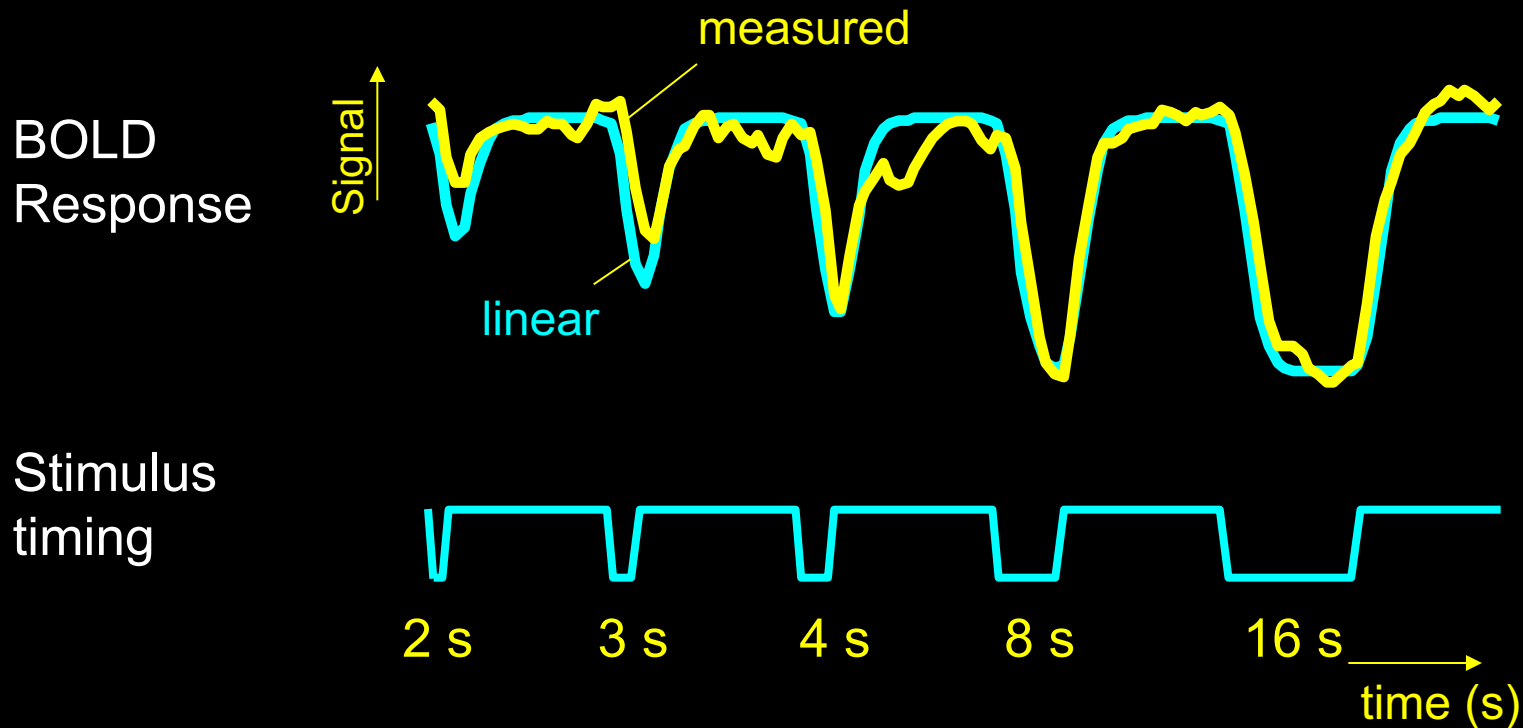


Experiment 1



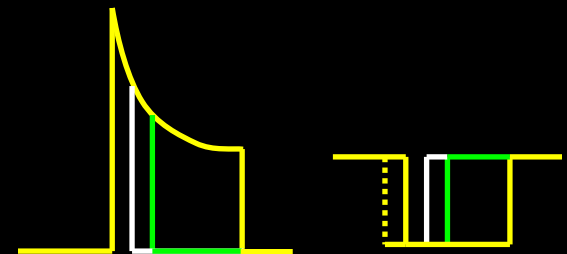
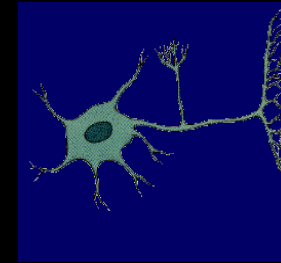
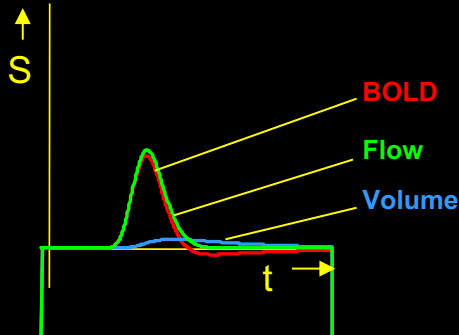
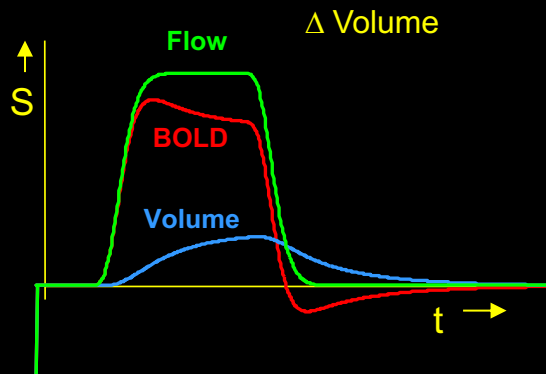
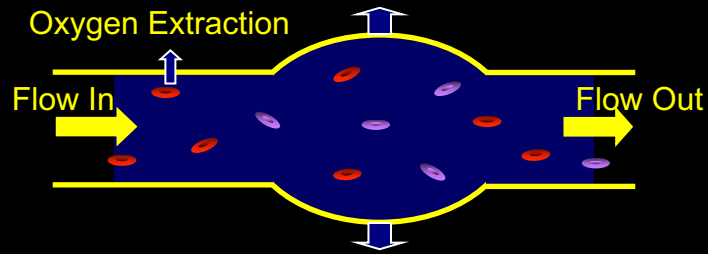
Experiment 2

Different stimulus “OFF” periods



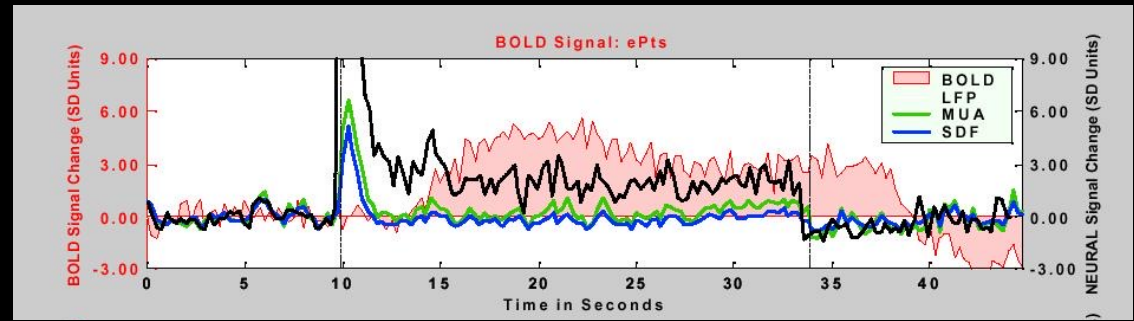
Brief stimulus OFF periods produce smaller decreases than expected

Sources of this Nonlinearity

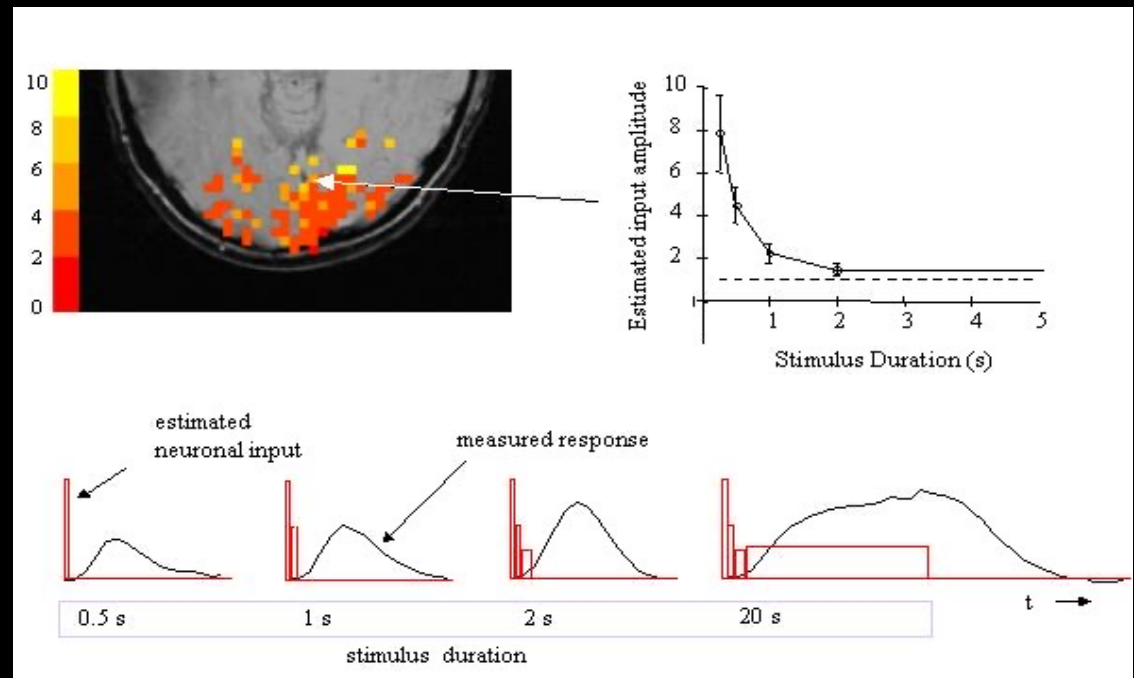


BOLD Correlation with Neuronal Activity

Logothetis et al. (2001)
“Neurophysiological investigation
of the basis of the fMRI signal”
Nature, 412, 150-157.



P. A. Bandettini and L. G. Ungerleider, (2001) “From neuron
to BOLD: new connections.”
Nature Neuroscience, 4: 864-866.



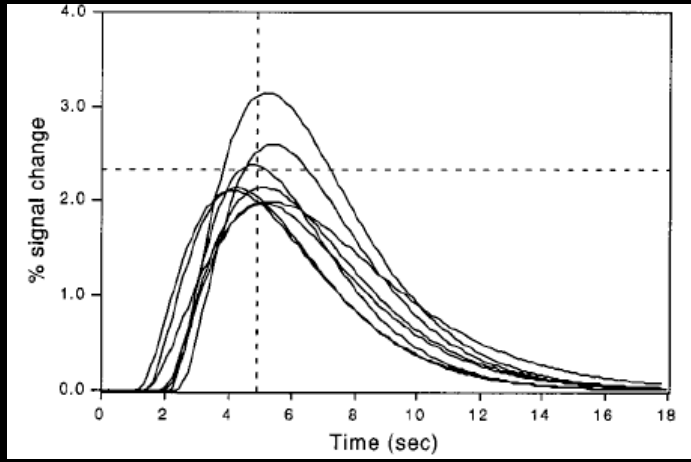
The Biggest Unknowns in Functional MRI

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

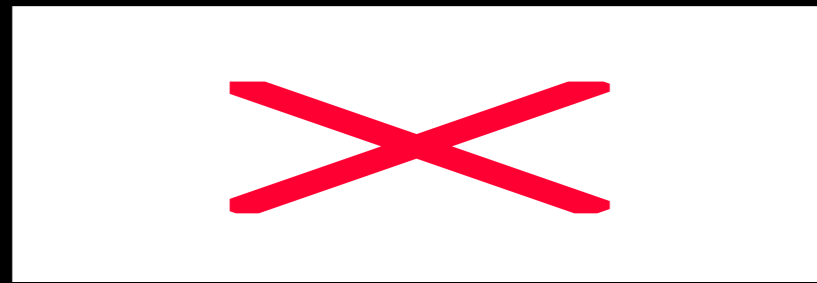
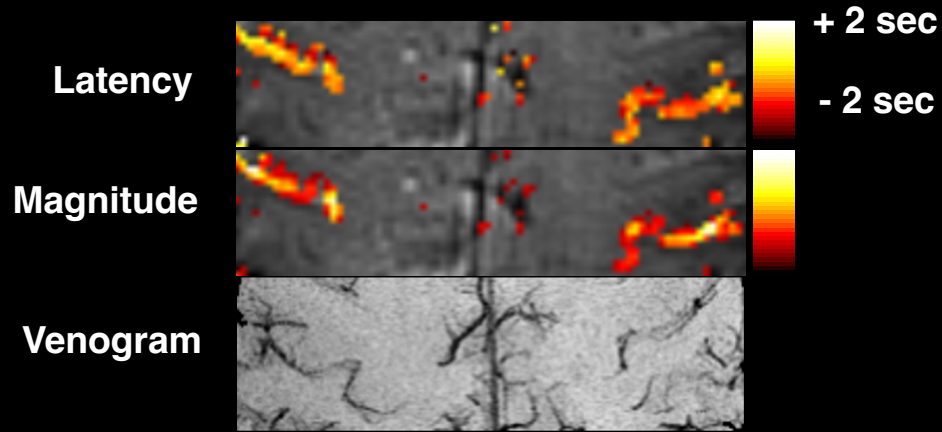
Sources of spatial and temporal variability.

Latency and Magnitude

From Subject to Voxel....



Miezin, et al (2000), NeuroImage 11, 735-759

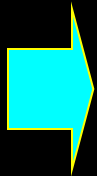
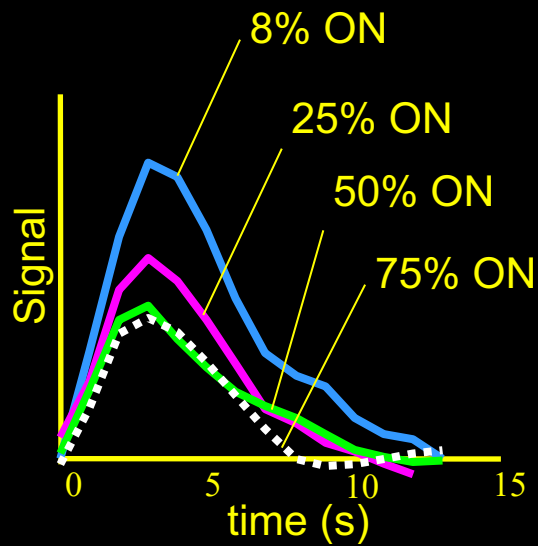


P. A. Bandettini, (1999) "Functional MRI" 205-220.

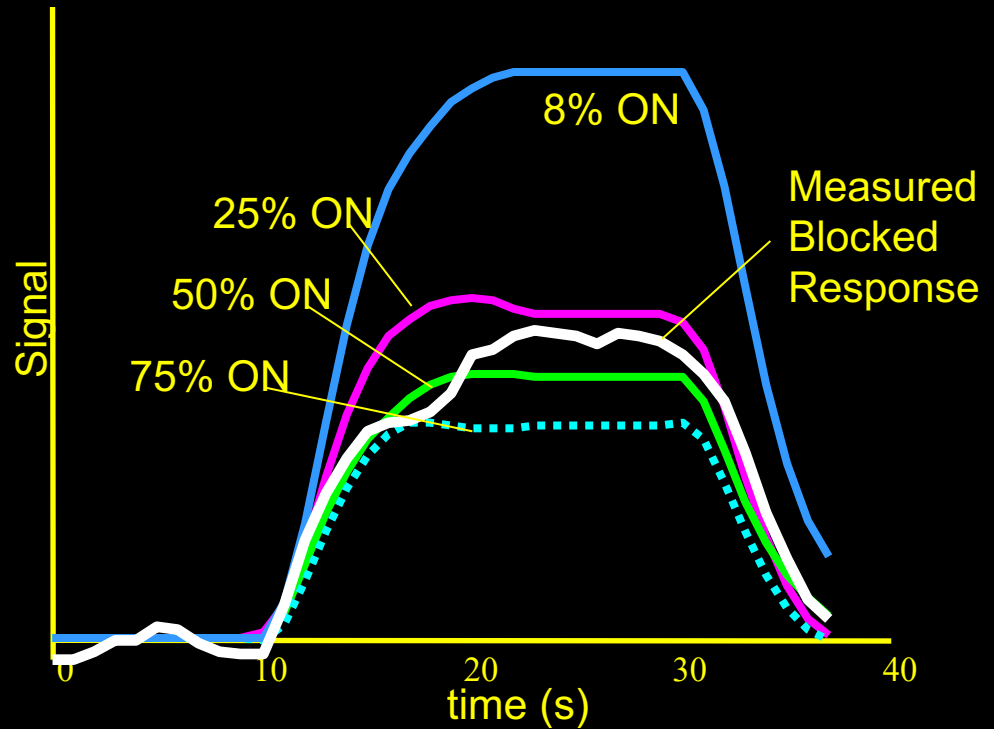
Rapid event-related design with varying ISI



*Estimated
Impulse Response*

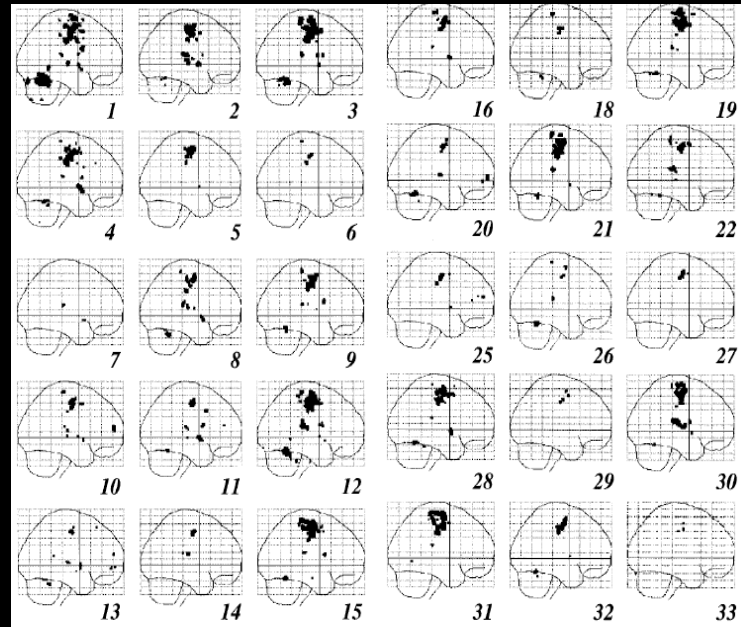


*Predicted Responses
to 20 s stimulation*

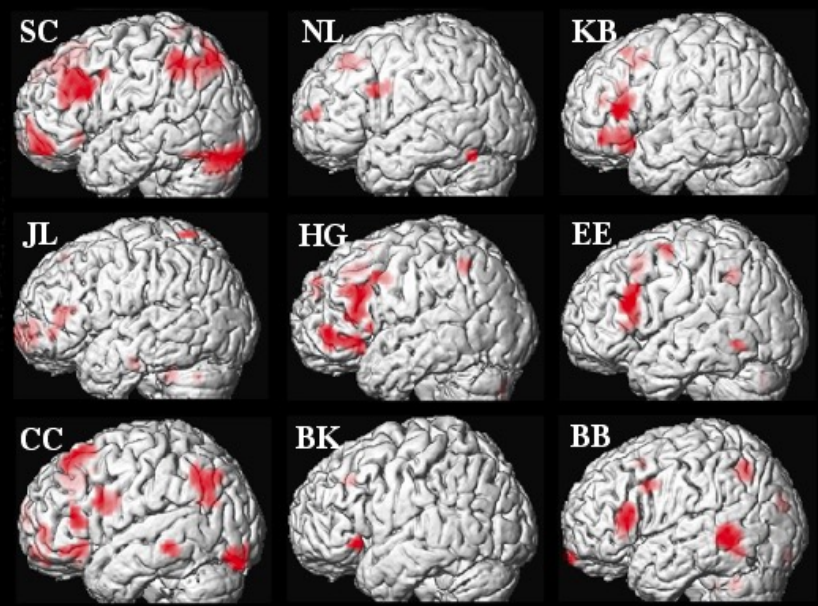
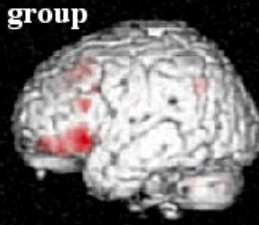


Sources of spatial and temporal variability.

Spatial Variation

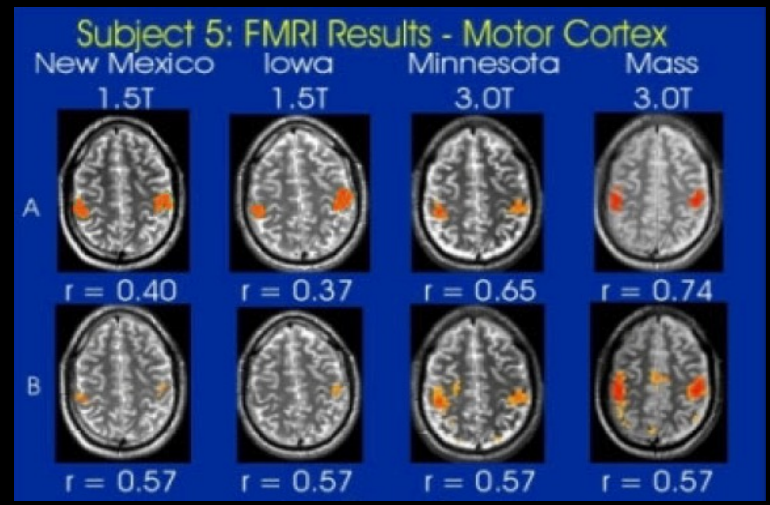


McGonigle, et al (2000),
NeuroImage 11, 708-734

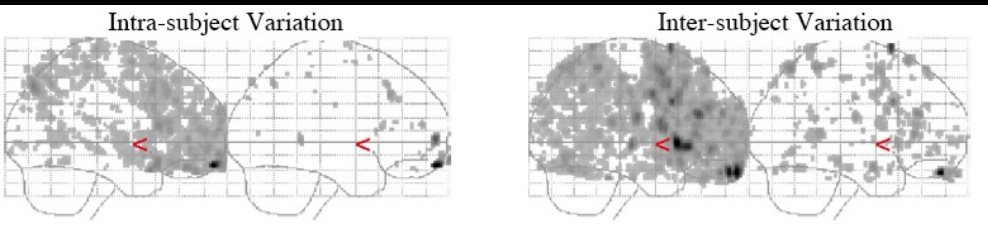


Courtesy, Mike Miller, UC Santa Barbara and
Jack Van Horn, fMRI Data Center, Dartmouth

F. BIRN
Biomedical
Informatics
Research
Network



L. Friedman, et al (2004), ISMRM 489

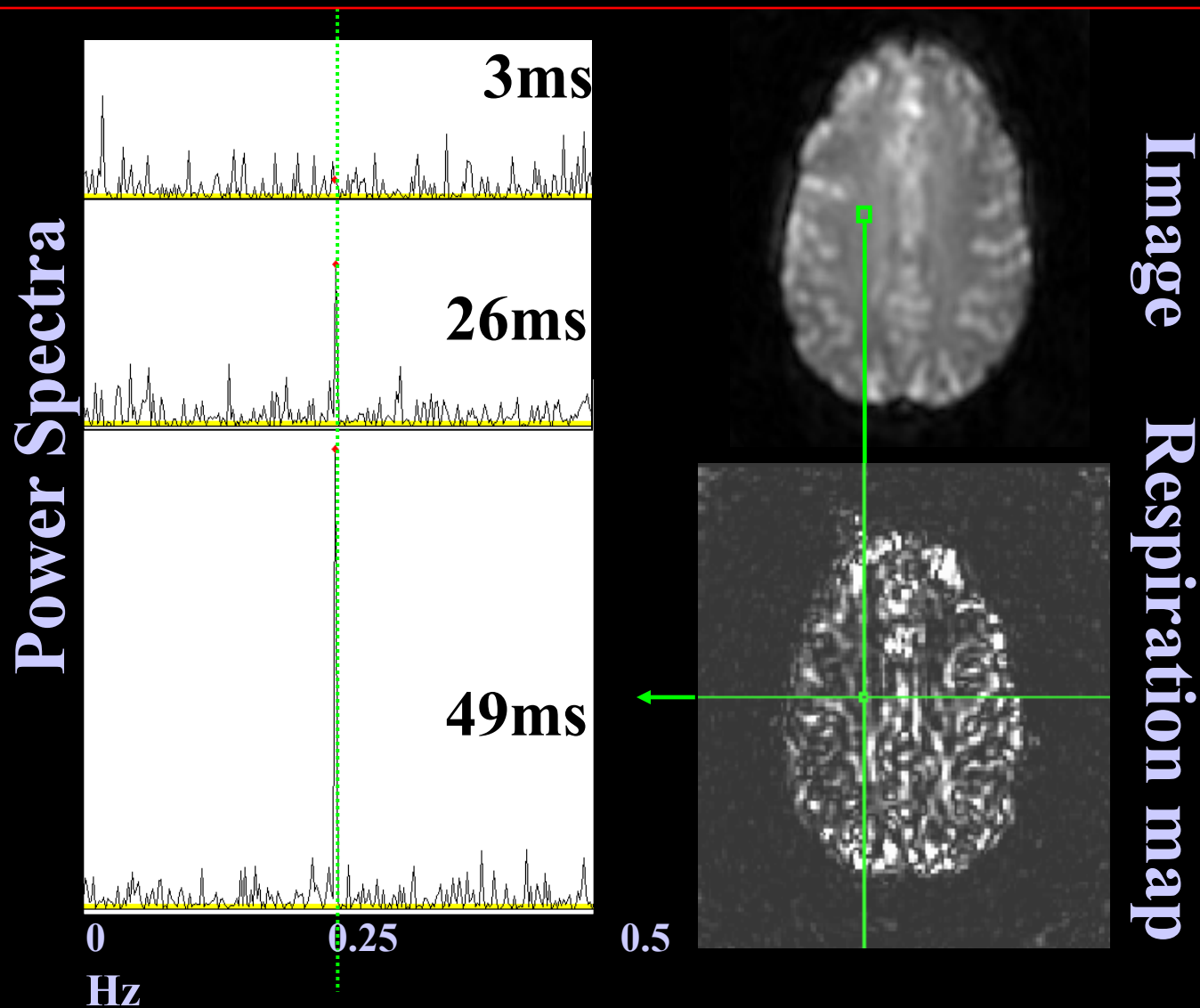


T.E. Lund, et al (2004), ISMRM 497a

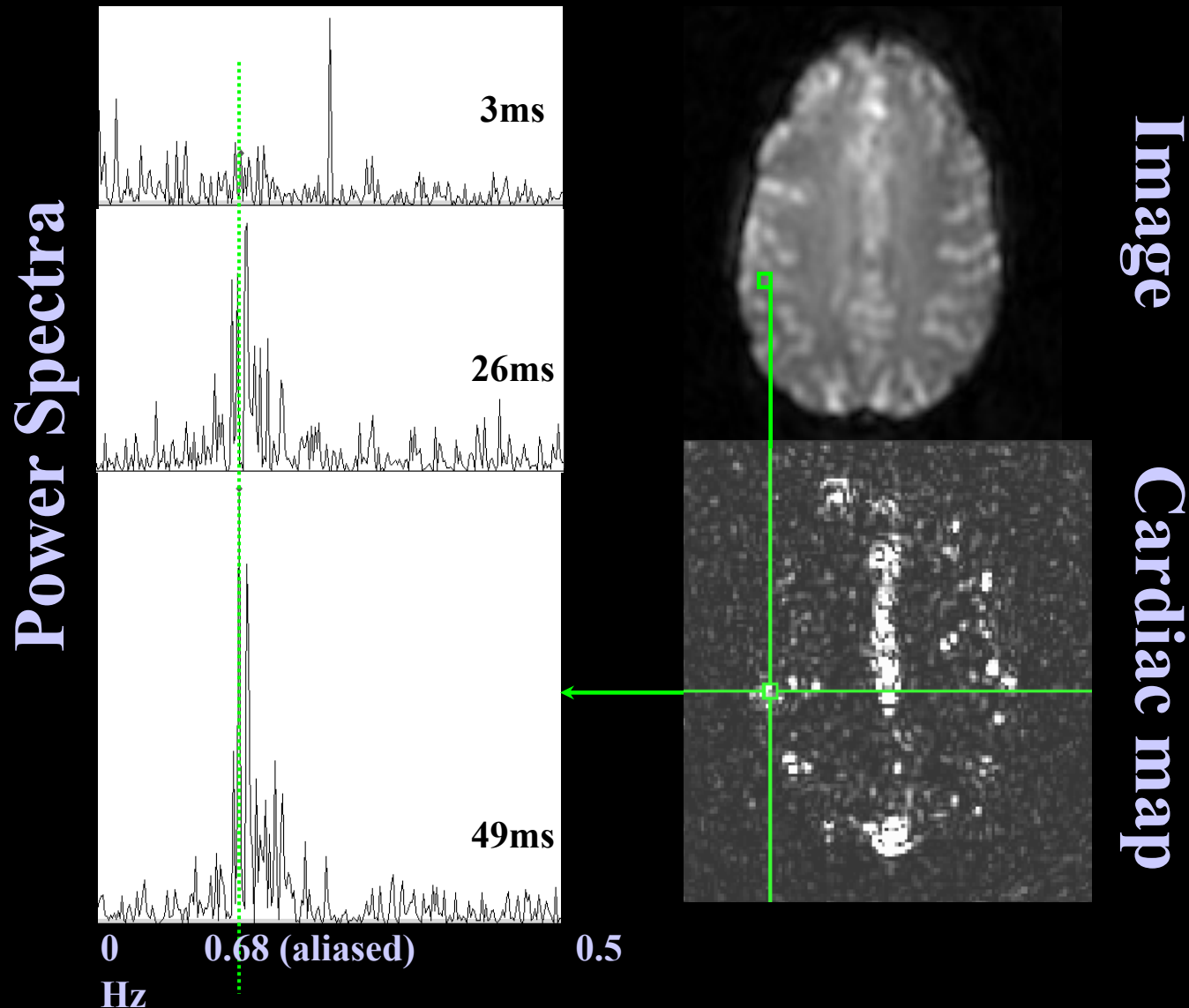
The Biggest Unknowns in Functional MRI

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

0.25 Hz Breathing at 3T

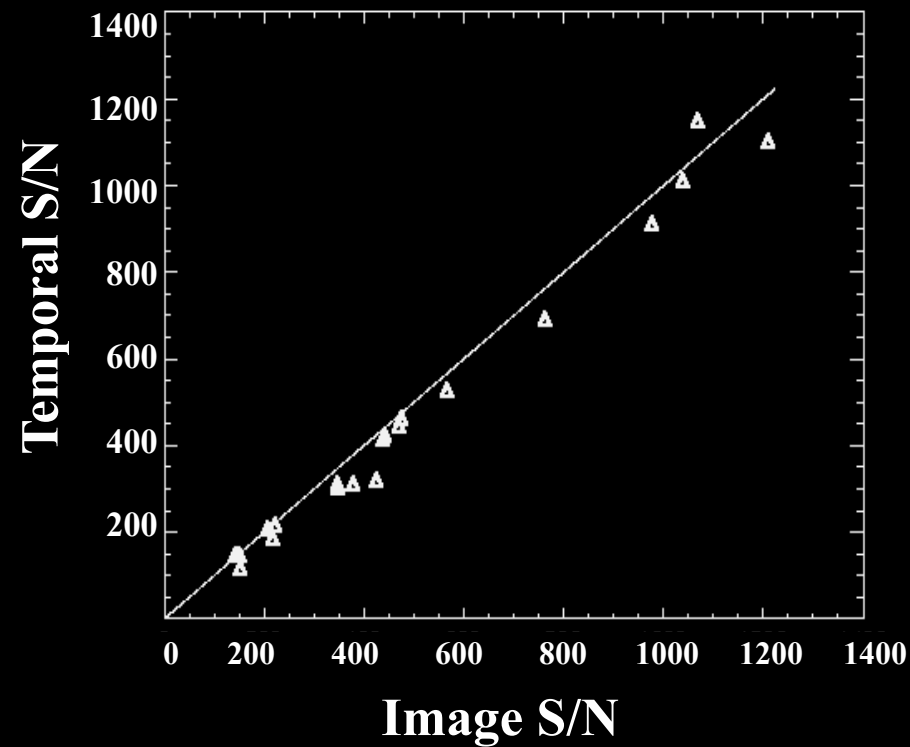


0.68 Hz Cardiac rate at 3T

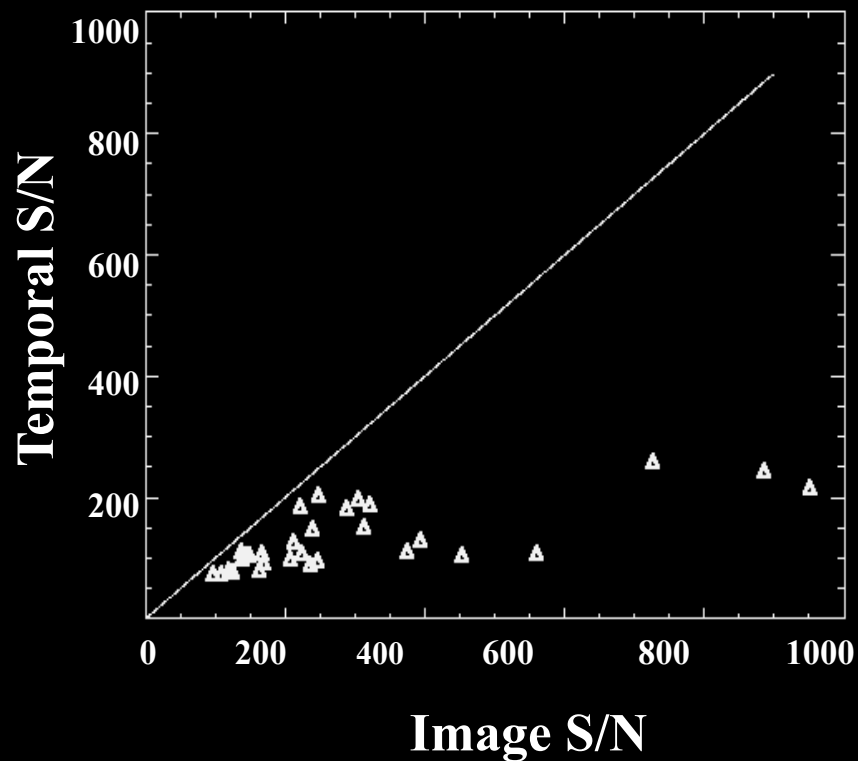


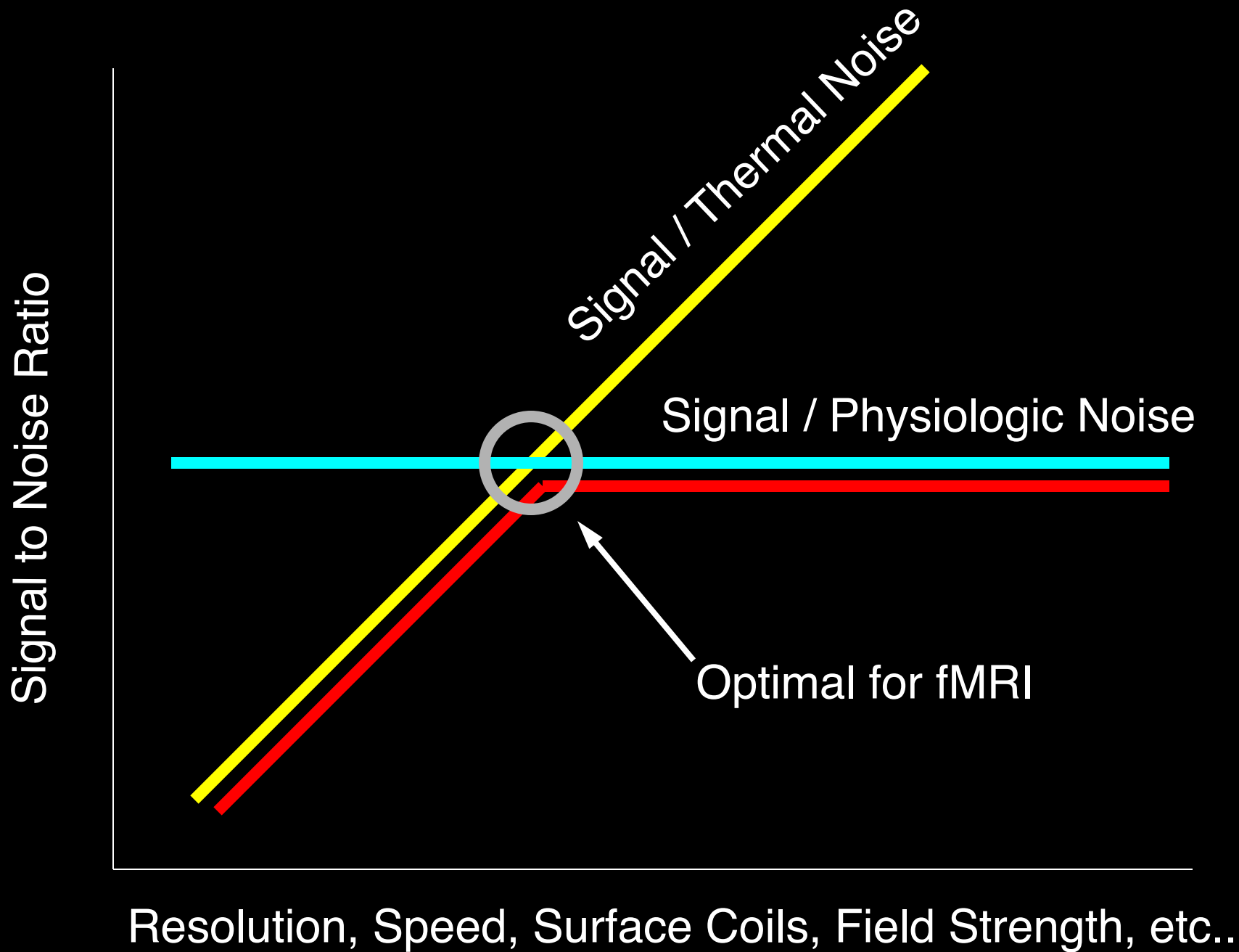
Temporal S/N vs. Image S/N

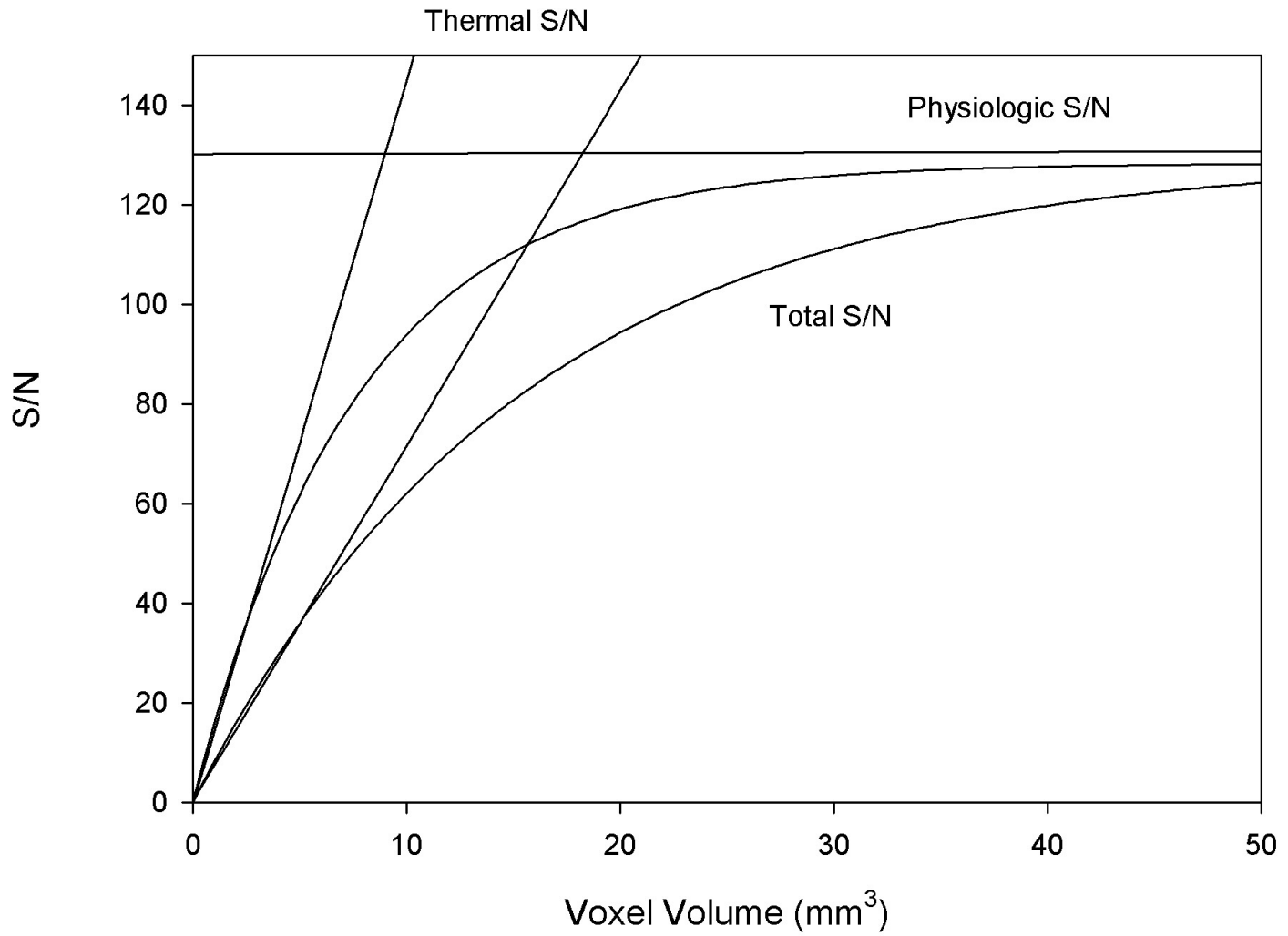
PHANTOMS

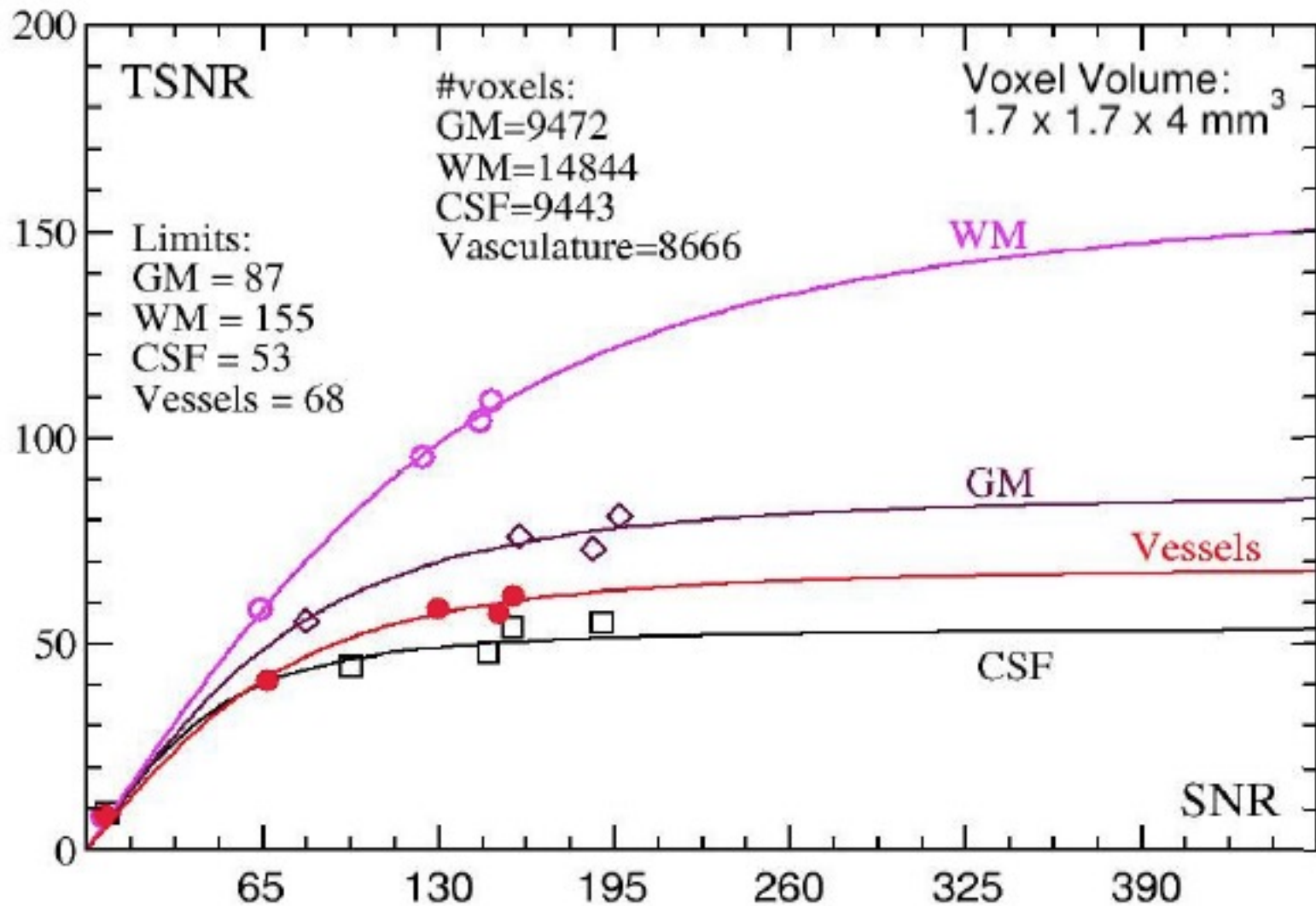


SUBJECTS



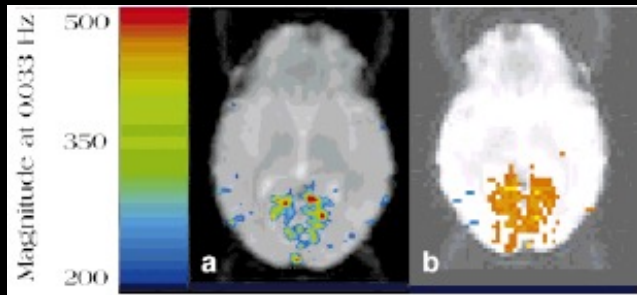
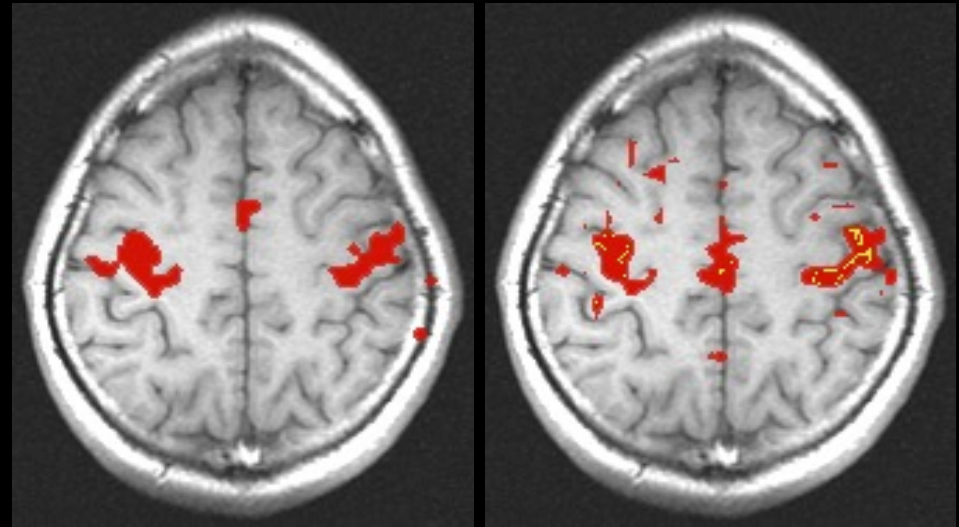
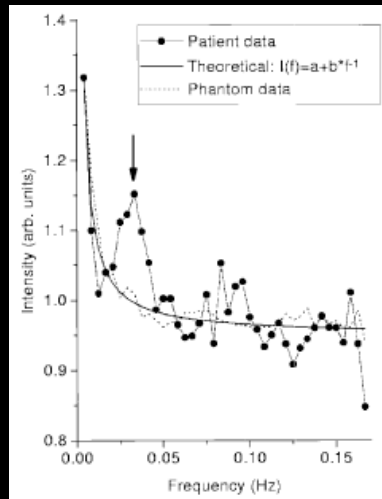






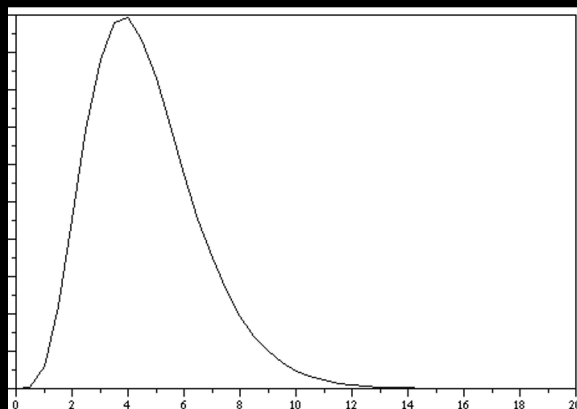
What's really in the noise?

Spontaneous Fluctuation Correlation

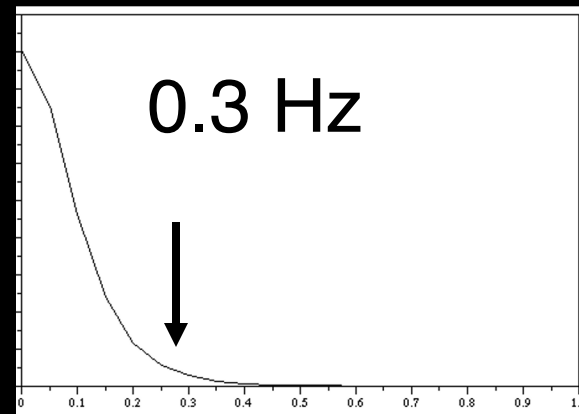


Kiviniemi, et al (2000), MRM 44, 373-378

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



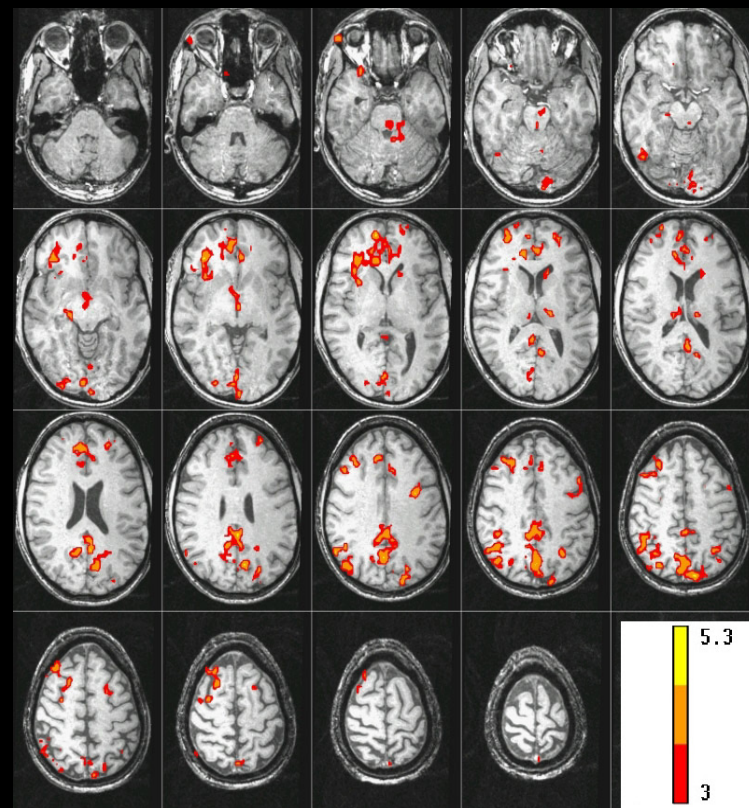
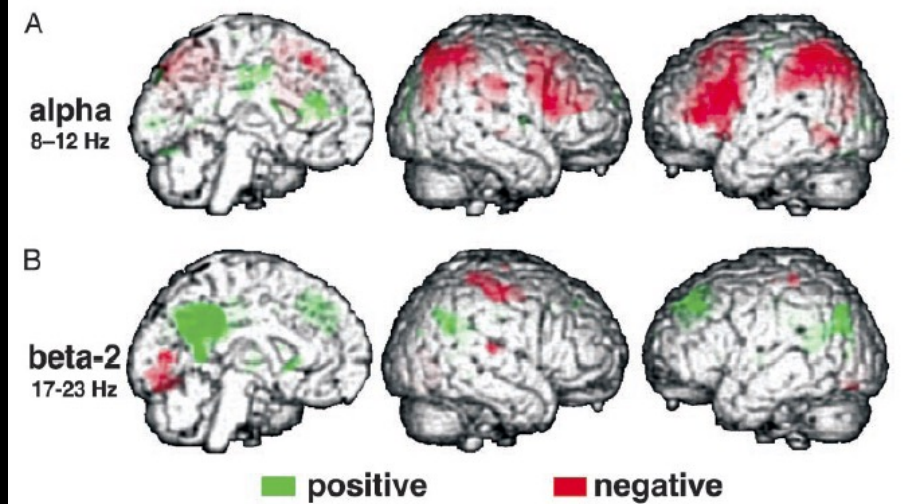
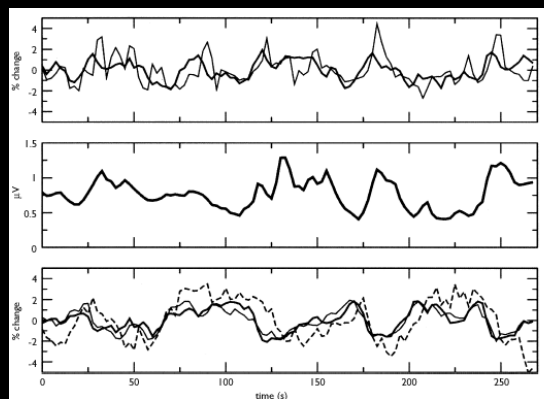
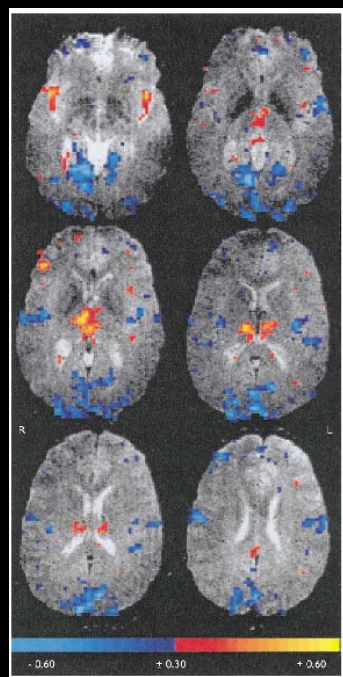
FFT
↔



What's really in the noise?

Laufs, et al
(2003), PNAS 100
(19), 11053-11058

Correlation with External Measures



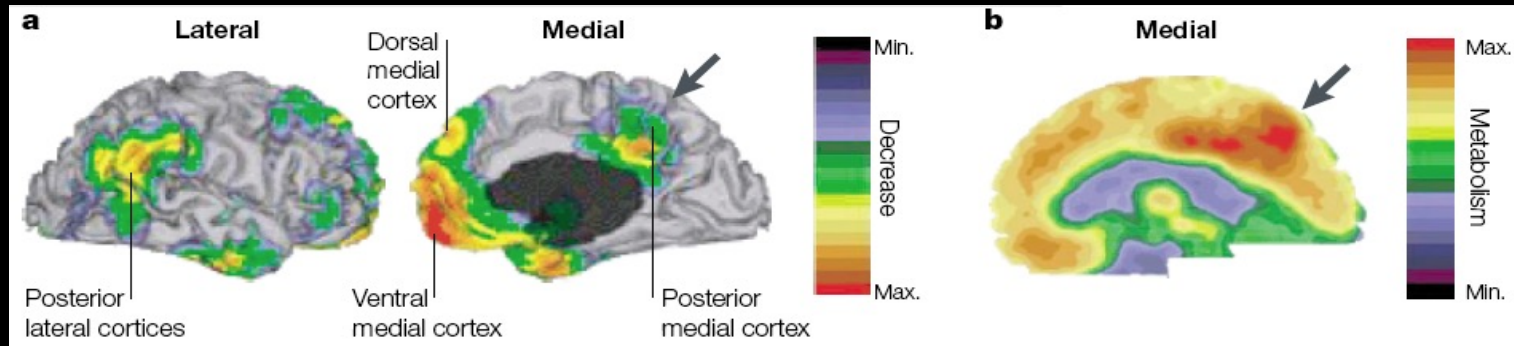
Goldman, et al (2002), Neuroreport

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

The Biggest Unknowns in Functional MRI

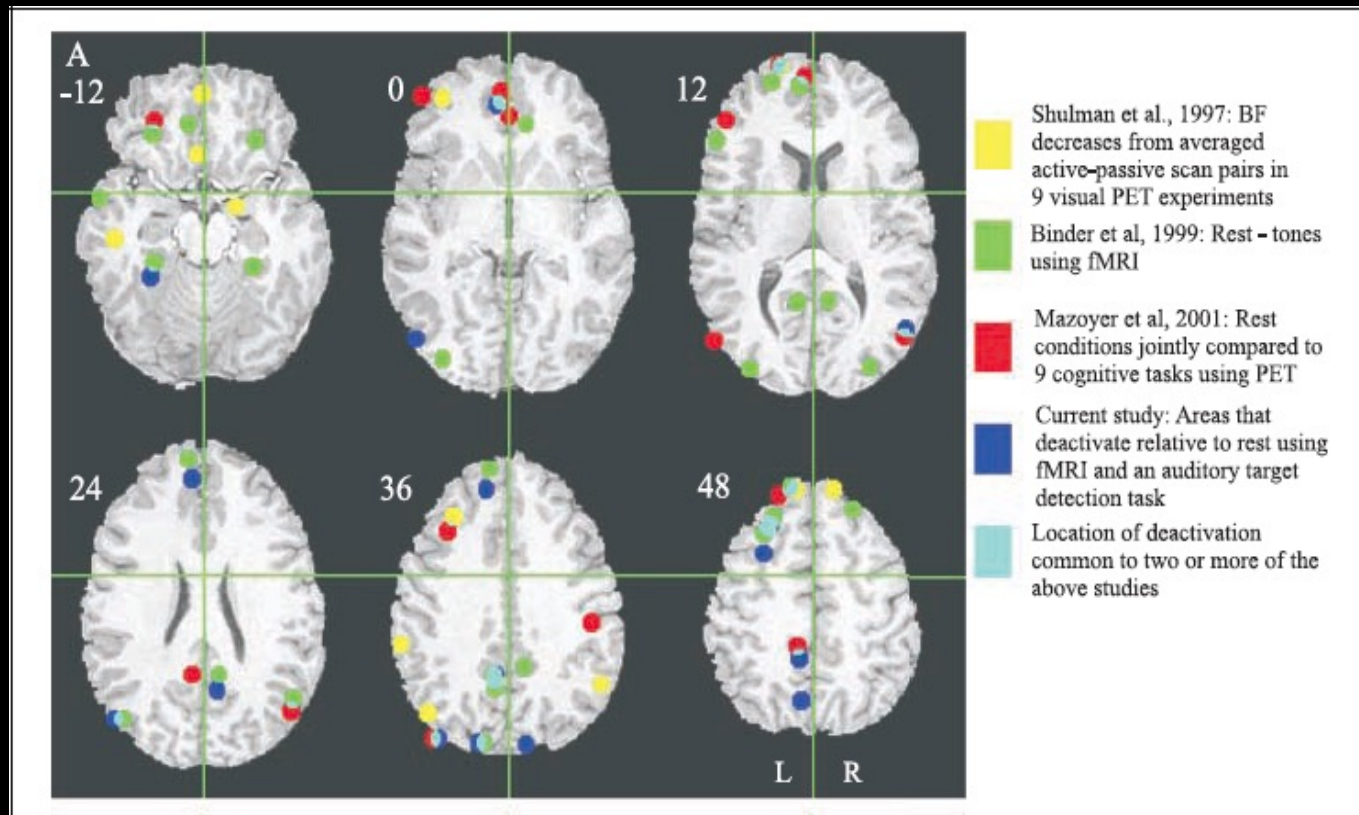
1. Relationship between neuronal activity and fMRI.
2. Sources of fMRI dynamic characteristics.
3. Sources of spatial and temporal variability.
4. What's really in the noise?
5. What's "resting" state?
6. Other sources of functional contrast?
7. Ultimate temporal resolution?
8. Ultimate spatial resolution?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

What is “resting” state?



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

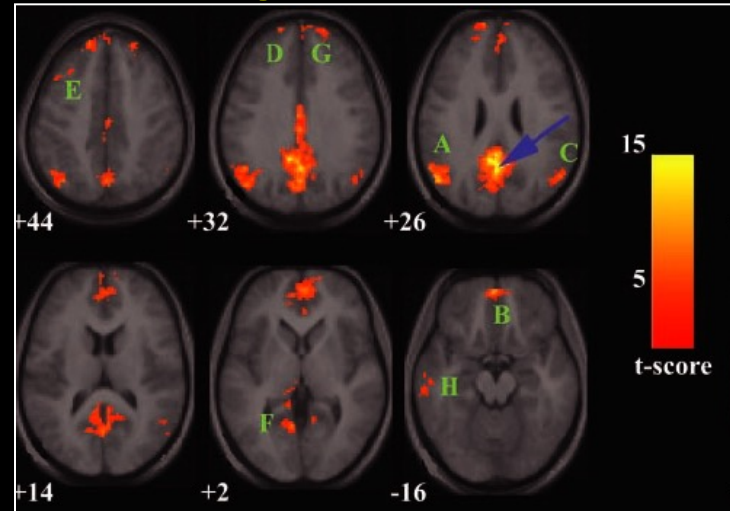
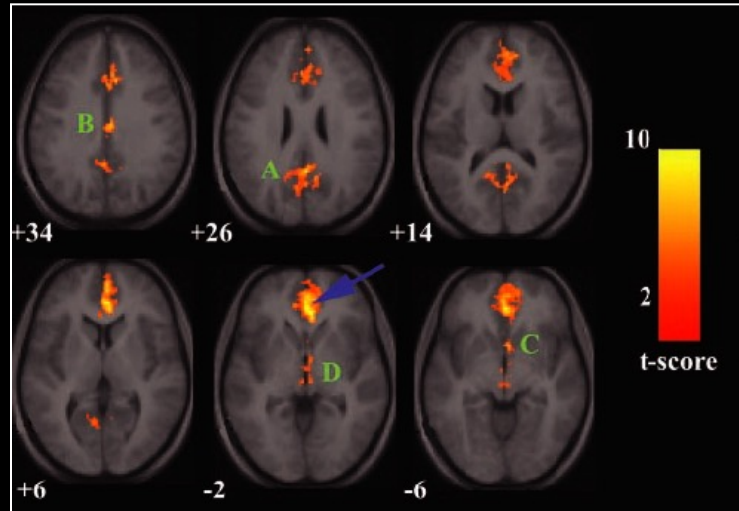
Decreases during activation



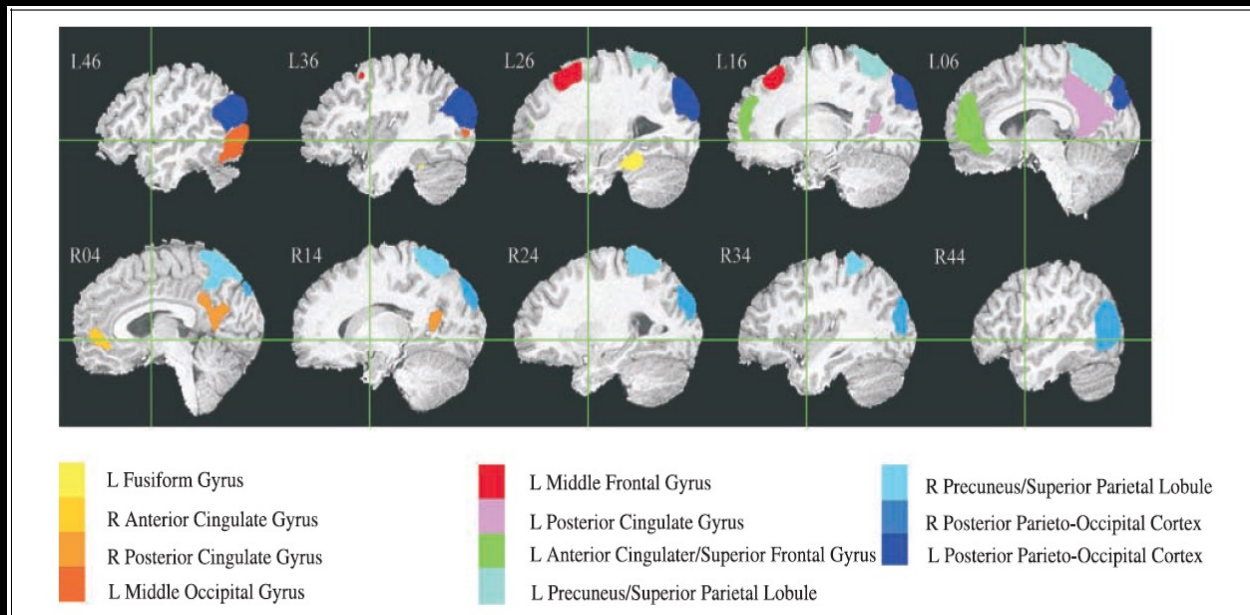
McKiernan, et al (2003), Journ. of Cog. Neurosci. 15 (3), 394-408

What is “resting” state?

Are decreases related to resting correlations?



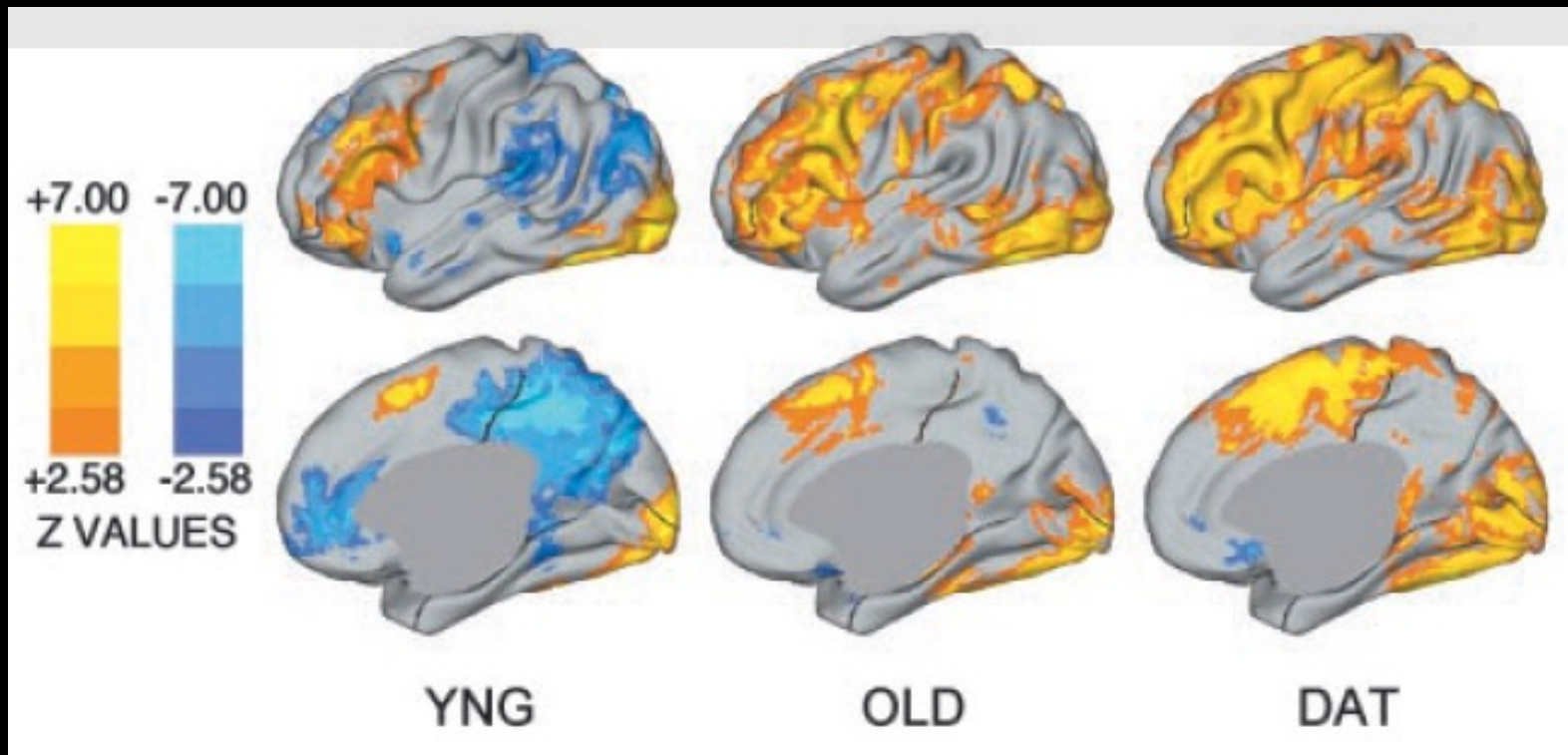
Greicius, et al (2003), PNAS 100 (1), 253-258



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

What is “resting” state?

Clinical applications?



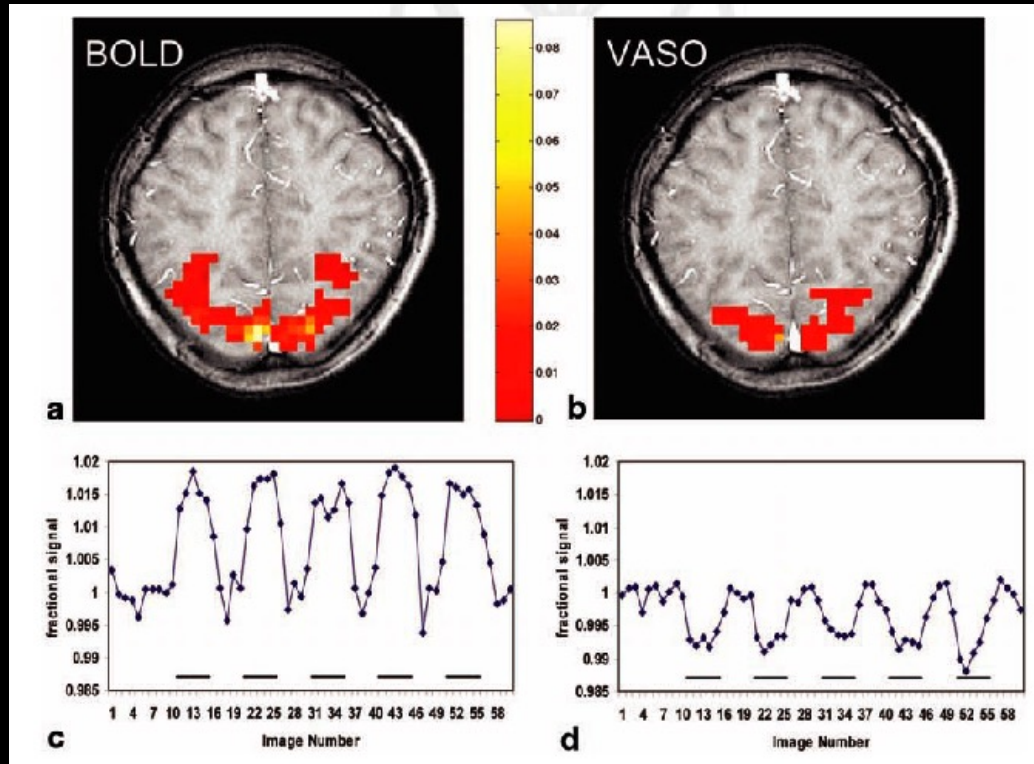
Lustig, et al (2003), PNAS 100 (19), 14504-14509

The Biggest Unknowns in Functional MRI

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

Other sources of functional contrast?

Blood Volume

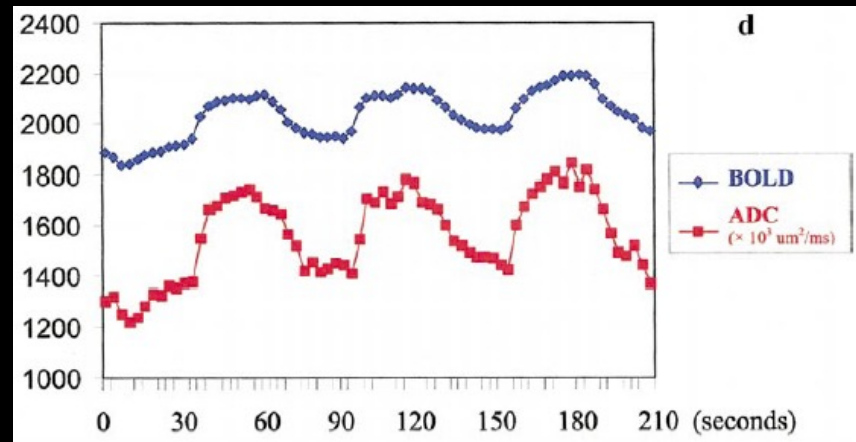


Lu, et al (2003) MRM 50 (2): 263-274

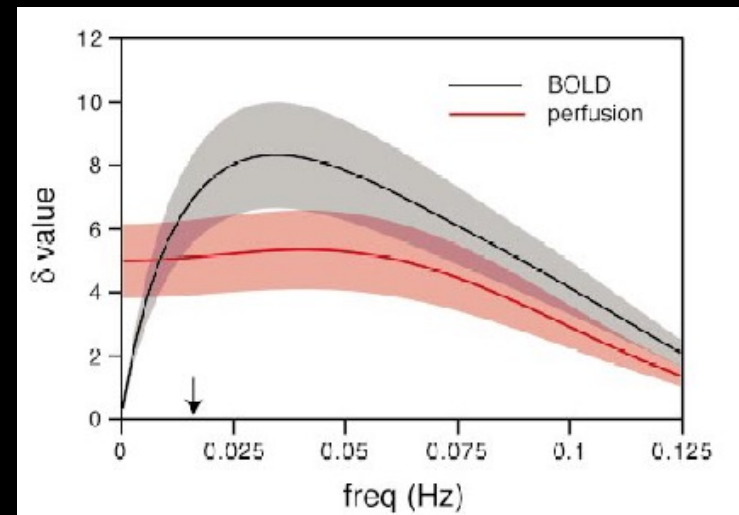
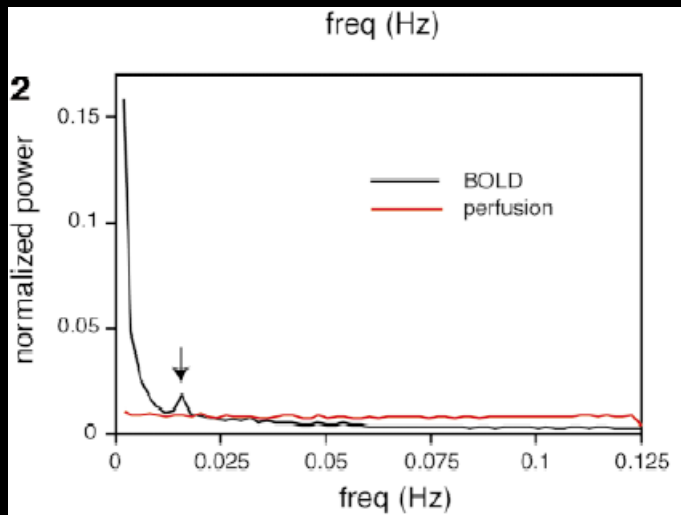
Other sources of functional contrast?

Non-ASL
Perfusion

Perfusion
Application



A. Song, et al (2002), NeuroImage 17, 742-750



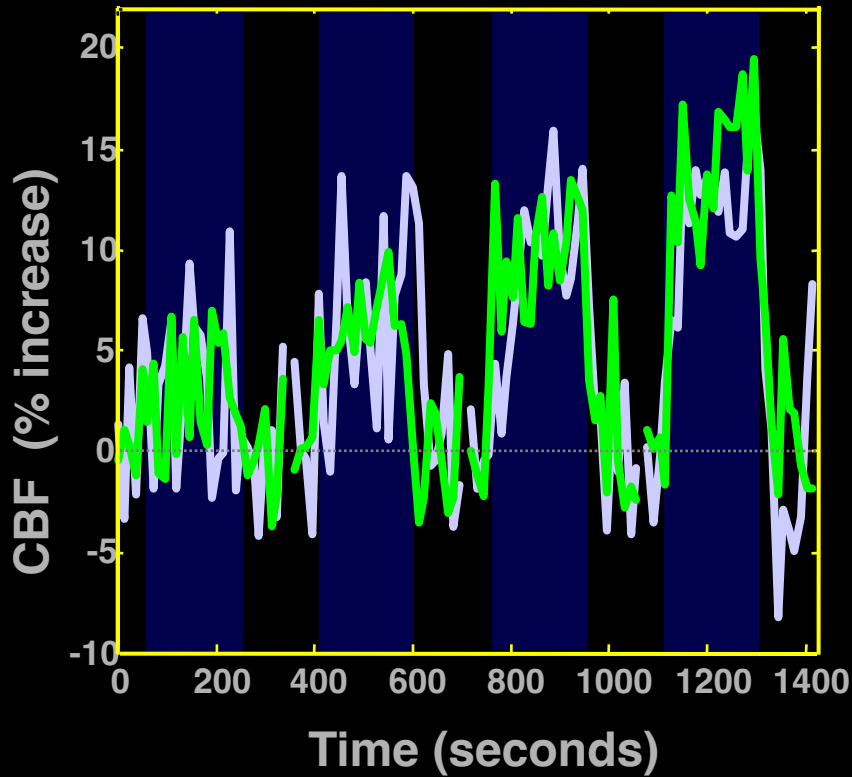
GK Aguirre et al, (2002) NeuroImage 15 (3): 488-500

Linear coupling between cerebral blood flow and oxygen consumption in activated human cortex

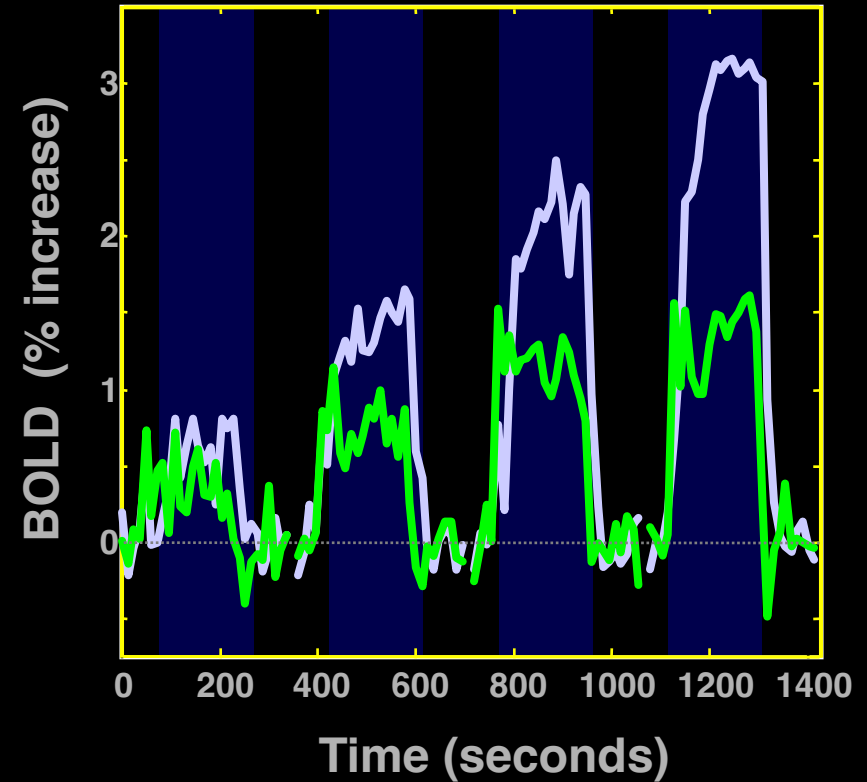
RICHARD D. HOGE^{*†}, JEFF ATKINSON^{*}, BRAD GILL^{*}, GÉRARD R. CRELIER^{*}, SEAN MARRETT[‡], AND G. BRUCE PIKE^{*}

^{*}Room WB325, McConnell Brain Imaging Centre, Montreal Neurological Institute, Quebec, Canada H3A 2B4; and [‡]Nuclear Magnetic Resonance Center, Massachusetts General Hospital, Building 149, 13th Street, Charlestown, MA 02129

CBF



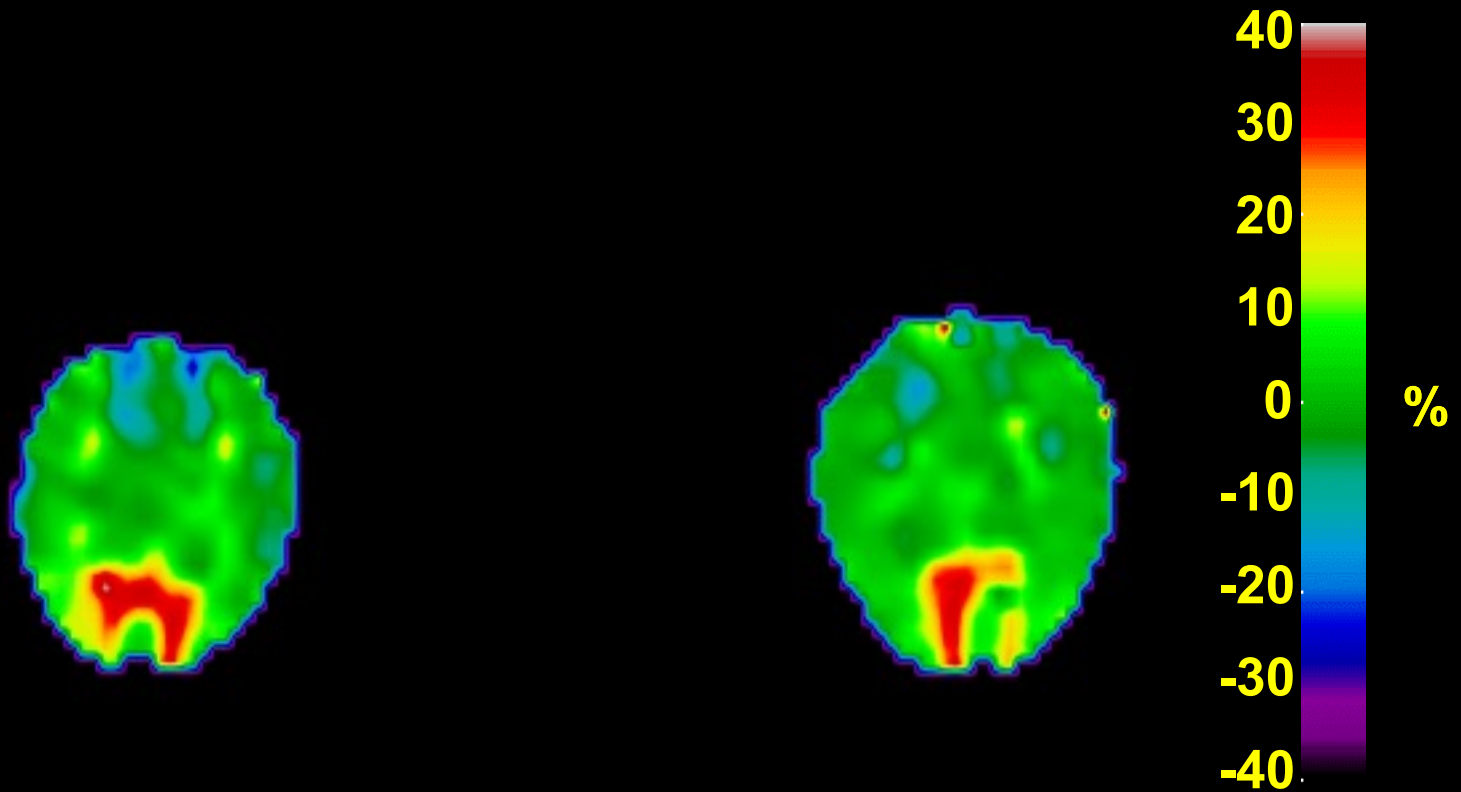
BOLD



Simultaneous Perfusion and BOLD imaging during
graded visual activation and hypercapnia

N=12

Computed CMRO₂ Changes

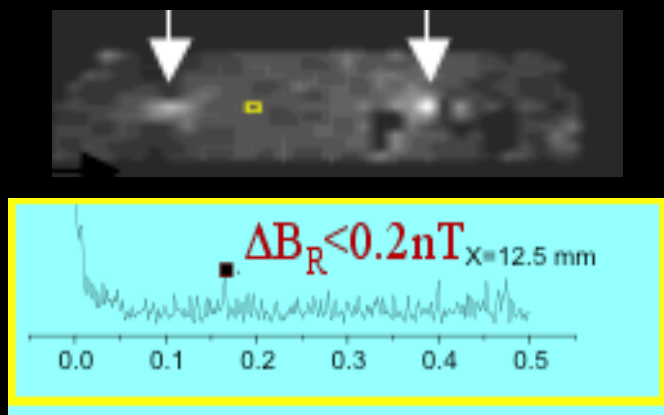


Subject 1

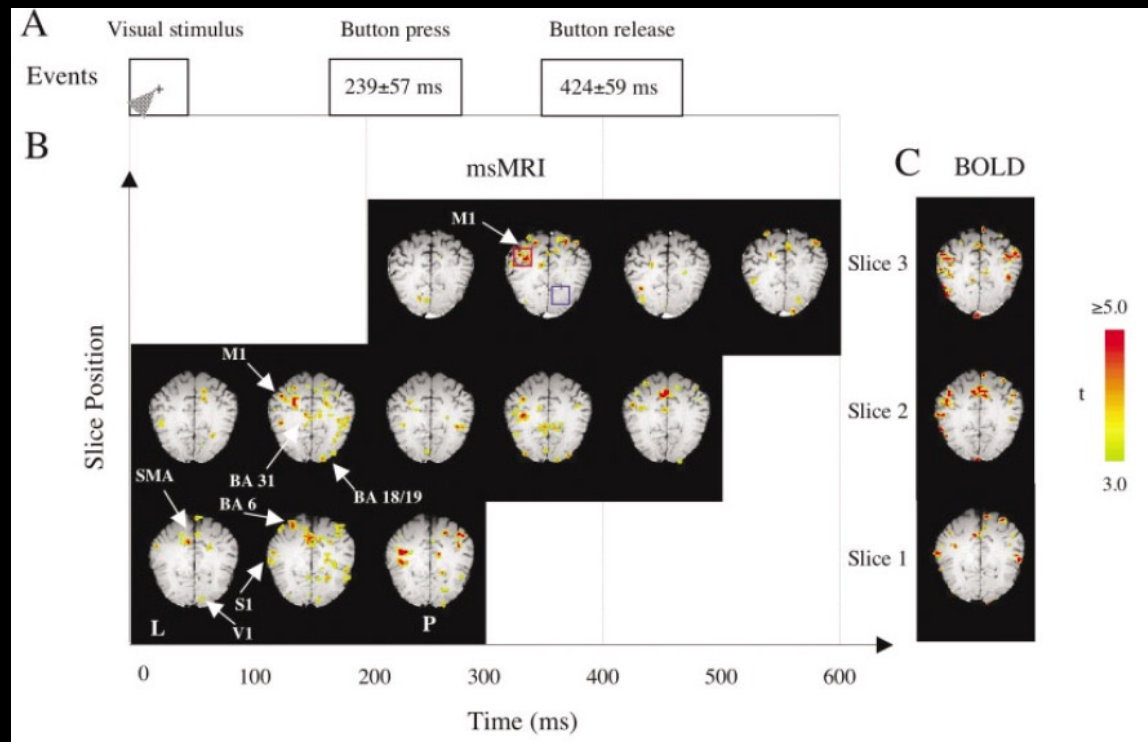
Subject 2

Other sources of functional contrast?

Direct Neuronal Current Imaging



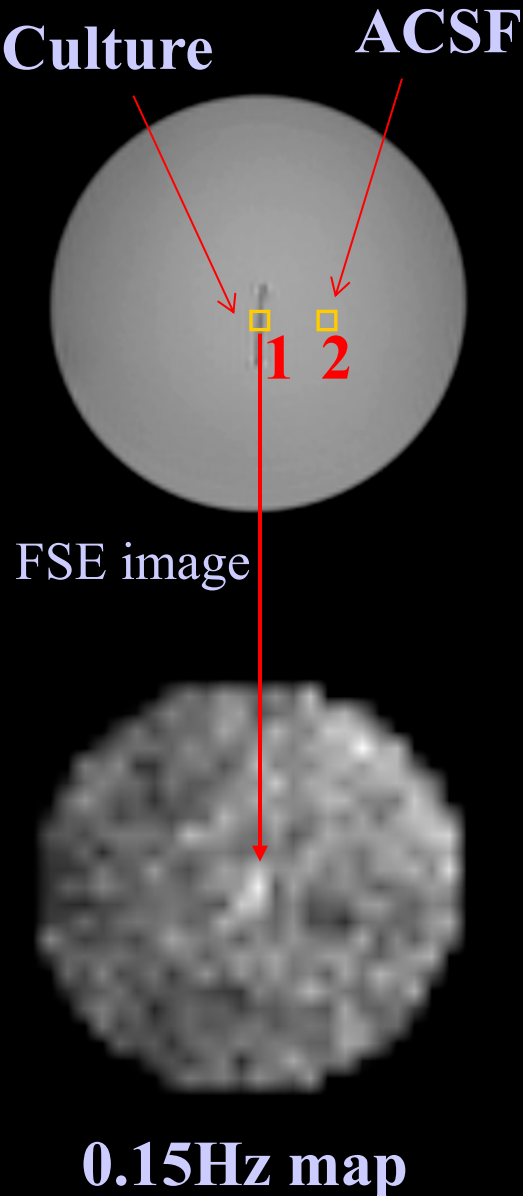
J. Bodurka, et al (2002).
MRM 47: 1052-1058.



J. Xiong, et al. (2003) HBM, 20: 41-49.

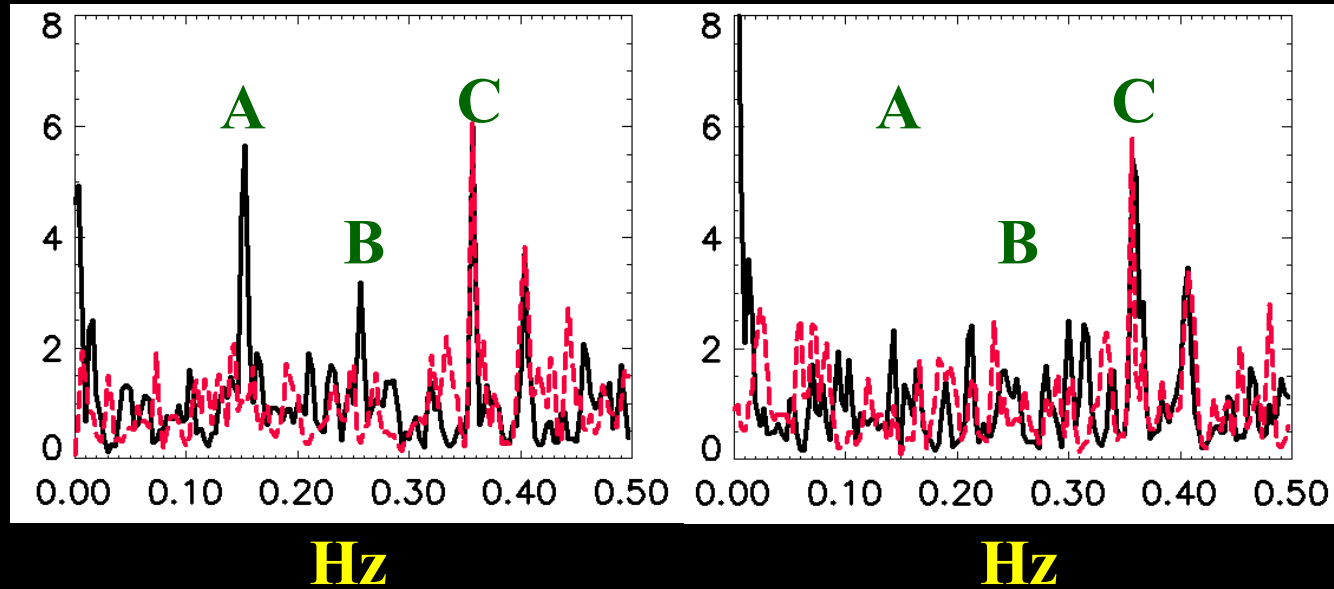
In Vitro Results

Other sources of functional contrast?



1: culture

2: ACSF



Active condition: black line

Inactive condition: red line

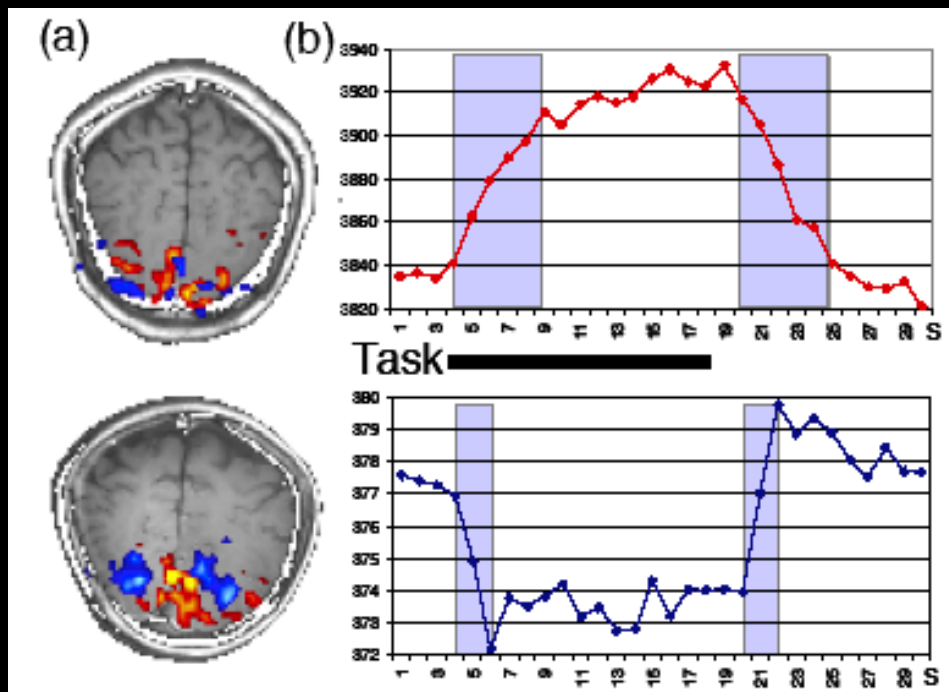
A: 0.15 Hz activity, on/off frequency

B: activity

C: scanner noise (cooling-pump)

Other sources of functional contrast?

Diffusion coefficient (high b-factor)



A. Song, et al (2004), ISMRM 1063

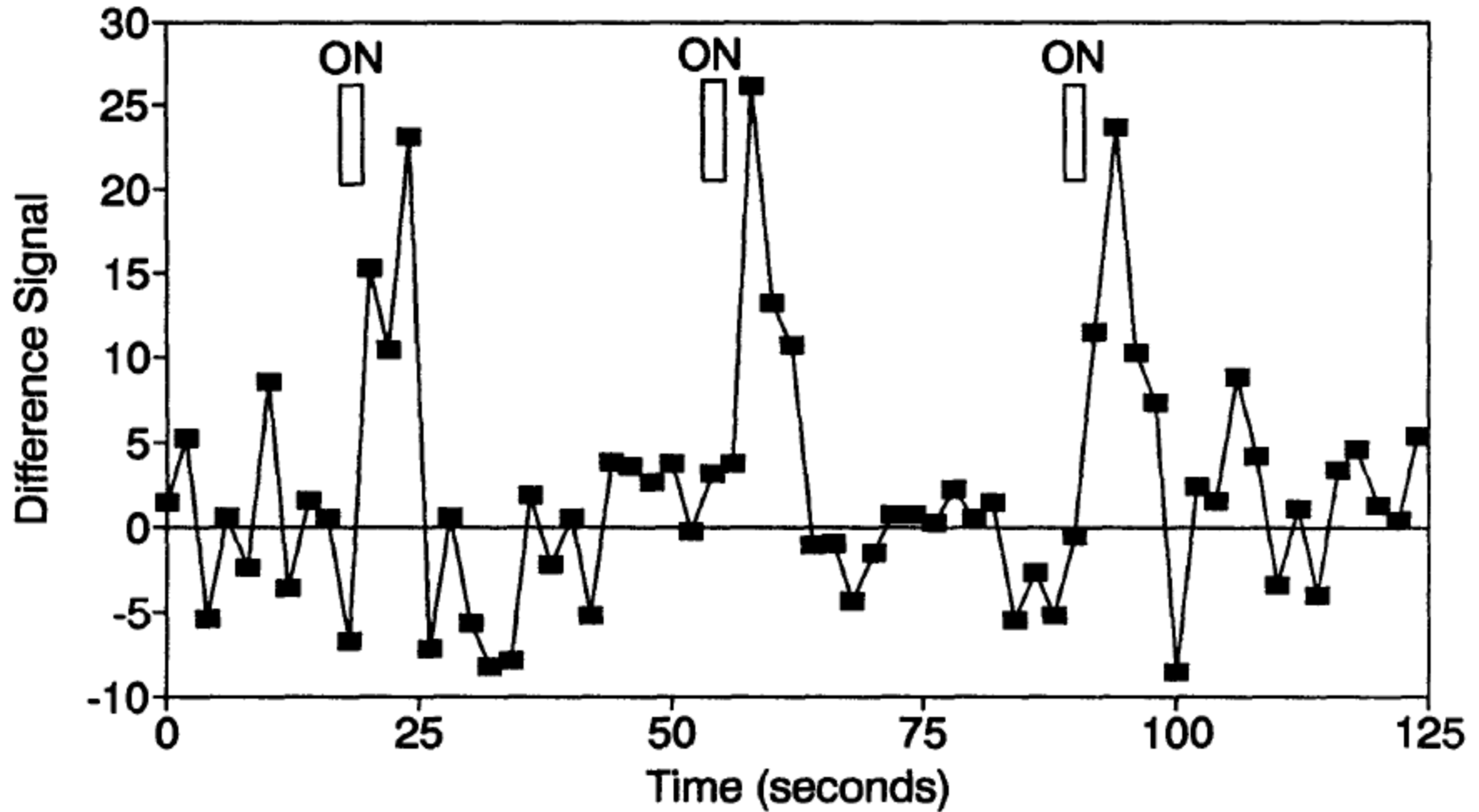
Temperature:

Yablonskiy, D. A., J. J. H. Ackerman, et al. (2000). "Coupling between changes in human brain temperature and oxidative metabolism during prolonged visual stimulation." Proceedings of the National Academy of Sciences of the United States of America 97(13): 7603-7608.

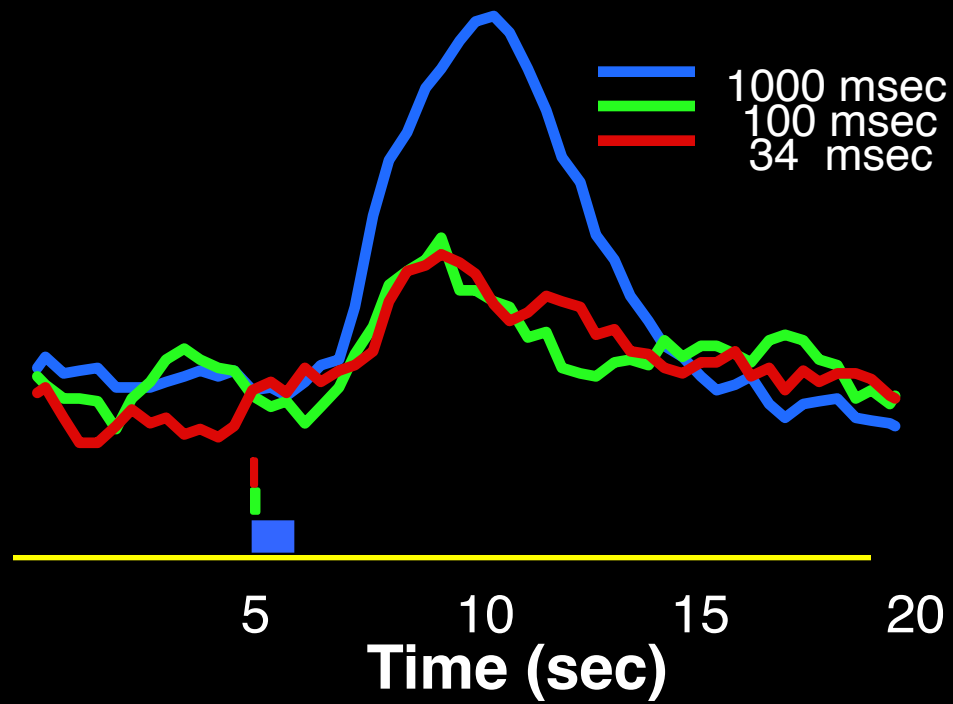
The Biggest Unknowns in Functional MRI

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

First Event-related fMRI Results

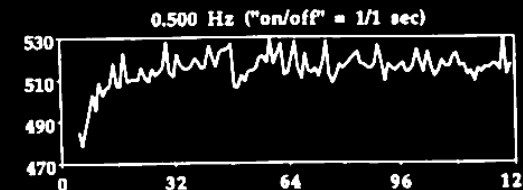
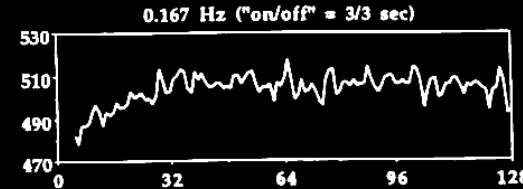
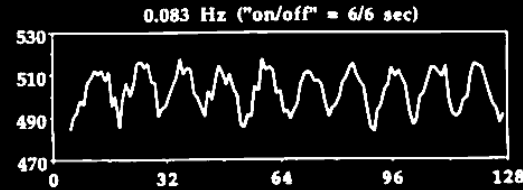
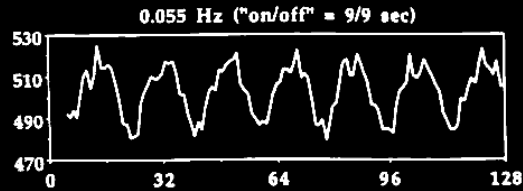
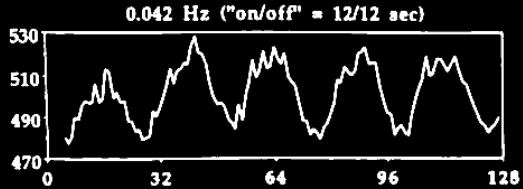
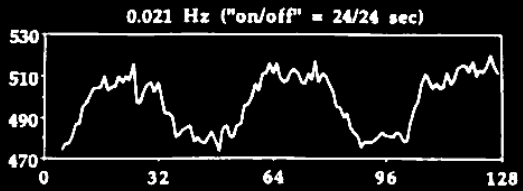


Blamire, A. M., et al. (1992). "Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging." *Proc. Natl. Acad. Sci. USA* 89: 11069-11073.

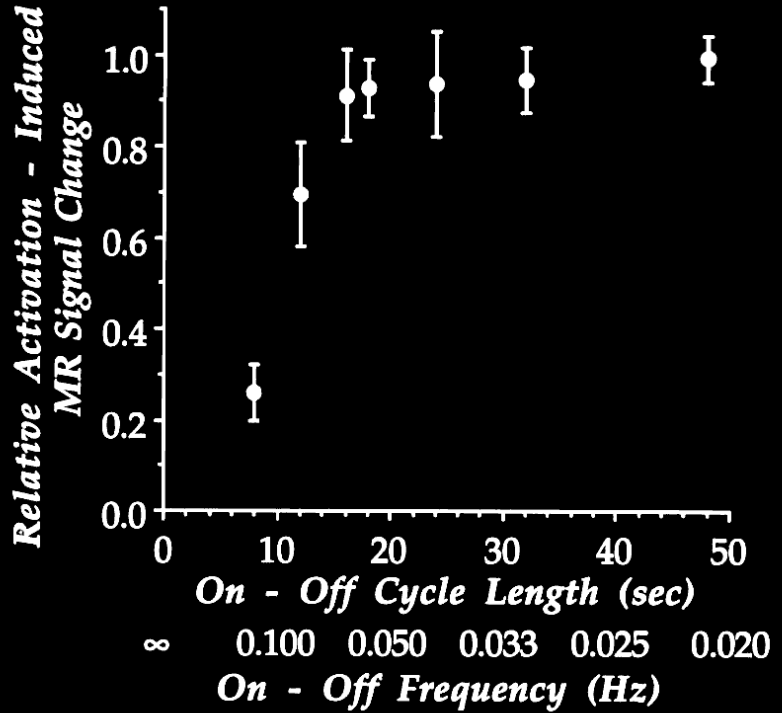


R. L. Savoy, et al., Pushing the temporal resolution of fMRI: studies of very brief visual stimuli, onset variability and asynchrony, and stimulus-correlated changes in noise [oral], 3rd Proc. Soc. Magn. Reson., Nice, p. 450. (1995).

MRI Signal



Time (seconds)

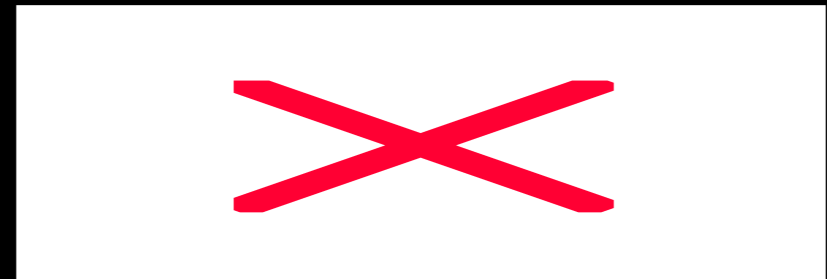
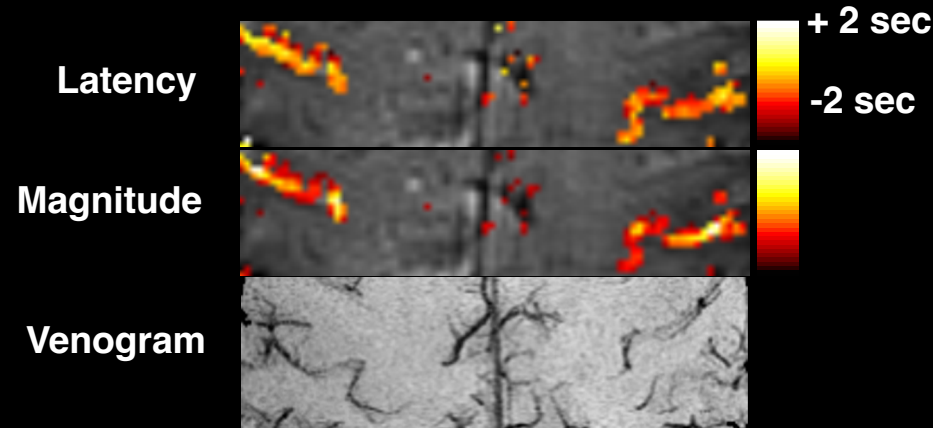


P. A. Bandettini, Functional MRI temporal resolution in "Functional MRI" (C. Moonen, and P. Bandettini., Eds.), p. 205-220, Springer - Verlag, 1999.

Ultimate temporal resolution?

Voxel-wise hemodynamic variation

Temporal resolution factors	Values for each factor
Fastest image acquisition rate	≈64 images/s
Minimum time for signal to significantly deviate from baseline	≈3 s
Fastest on-off rate in which amplitude-is not compromised	≈8 s on, 8 s off
Fastest on-off rate in which hemodynamic response keeps up	≈2 s on, 2 s off
Minimum activation duration	≈30 ms (no limit determined yet, but the response behaves similarly below 500 ms)
Standard deviation of baseline signal	≈1% (less if physiological fluctuations and system instabilities are filtered out)
Standard deviation of onset time estimation	≈450 ms
Standard deviation of return to baseline time estimation	≈1250 ms
Standard deviation of entire on-off response time estimation	≈650 ms
Range of latencies over space	± 2.5 s



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

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

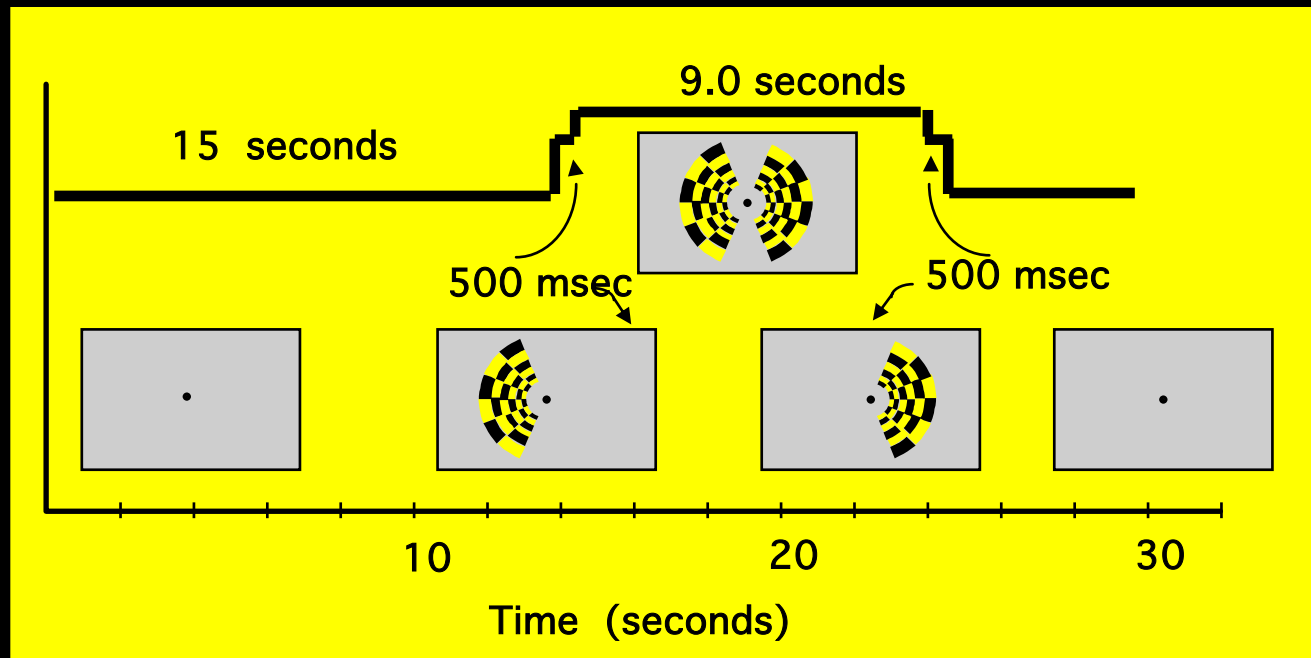
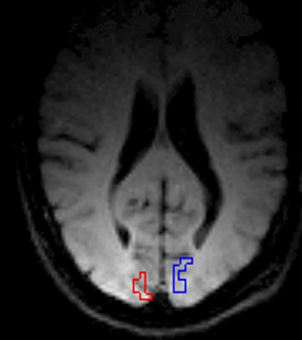
Relative dynamics obtained by precise activation timing modulation

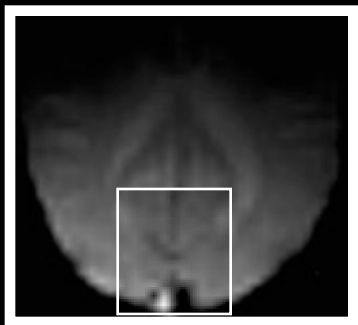
Preliminary results: (with Savoy et al. ~ 1995)

Hemi-Field Experiment

Left Hemisphere

Right Hemisphere





500 ms



500 ms



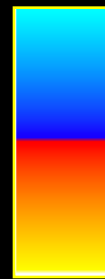
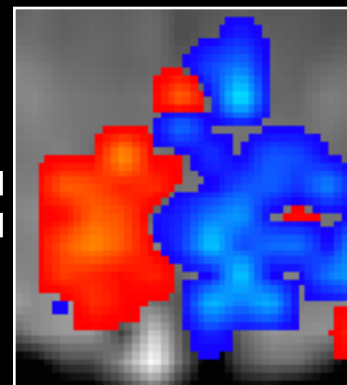
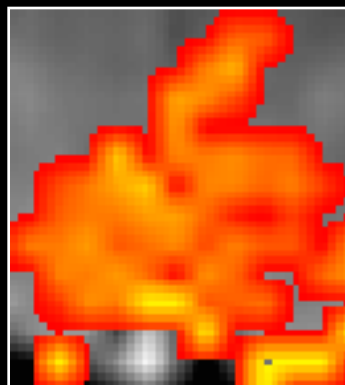
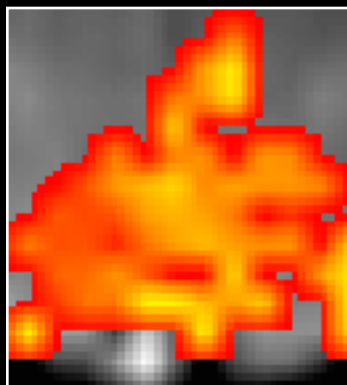
Right Hemifield

Left Hemifield

+ 2.5 s

0 s

- 2.5 s

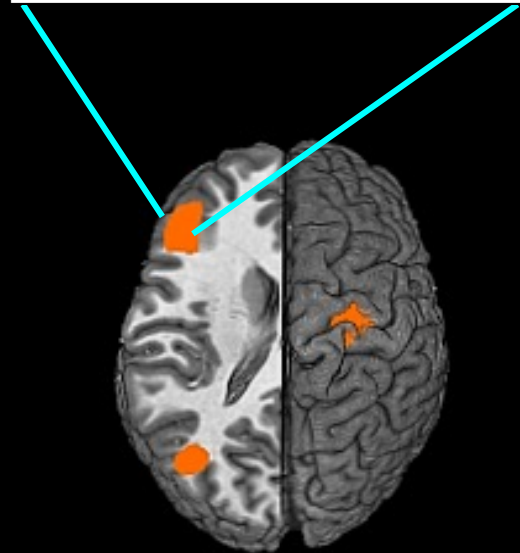
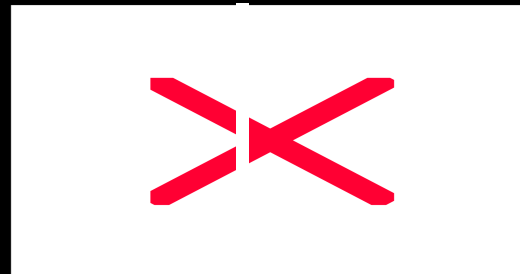


Ultimate temporal resolution?

Task Timing Modulation

Word vs. Non-word

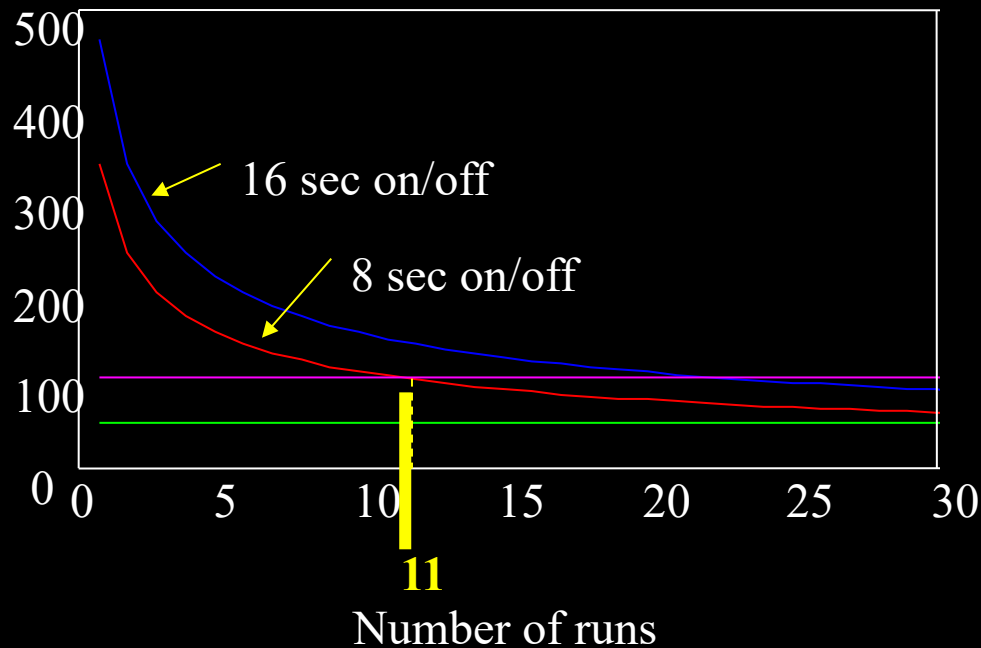
0°, 60°, 120° Rotation



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

Ultimate temporal resolution?

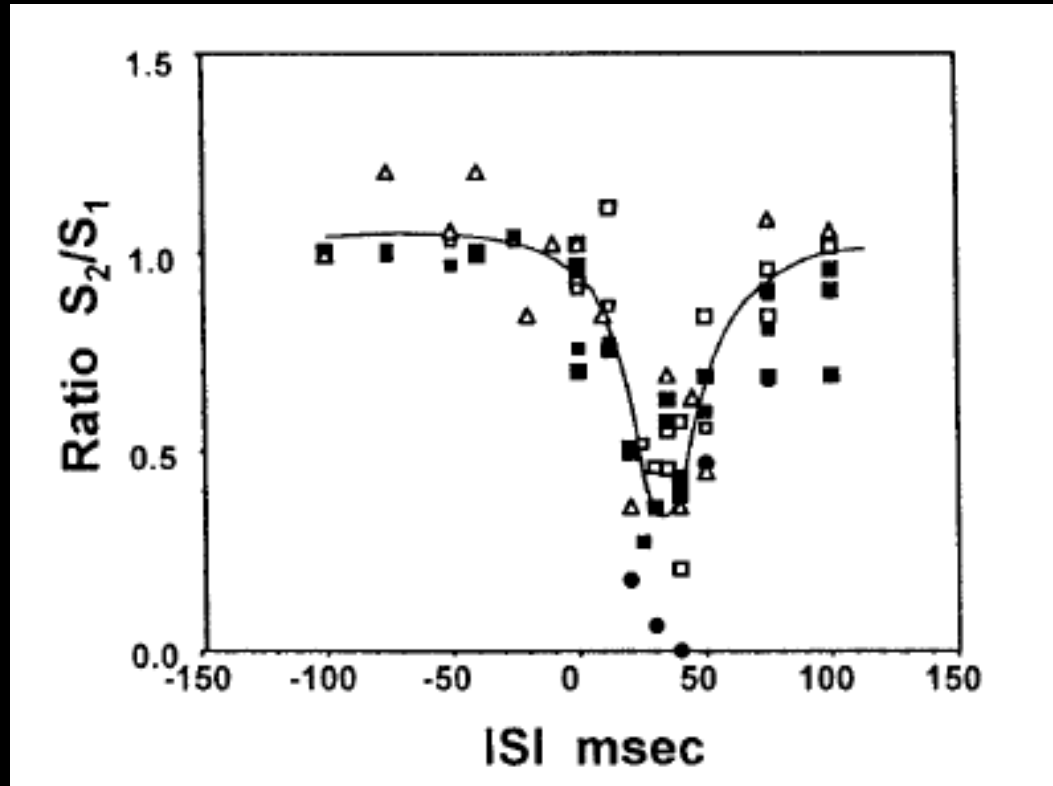
In an ideal world...no hemodynamic variation over space.



Smallest latency
Variation Detectable
(ms) ($p < 0.001$)

Ultimate temporal resolution?

Neuronal Communication Timing



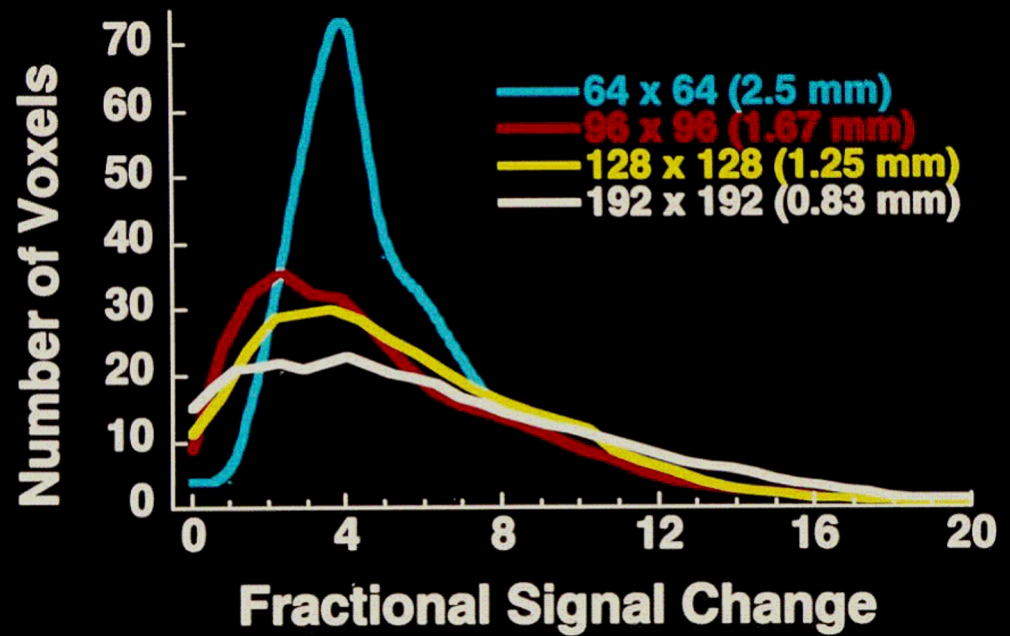
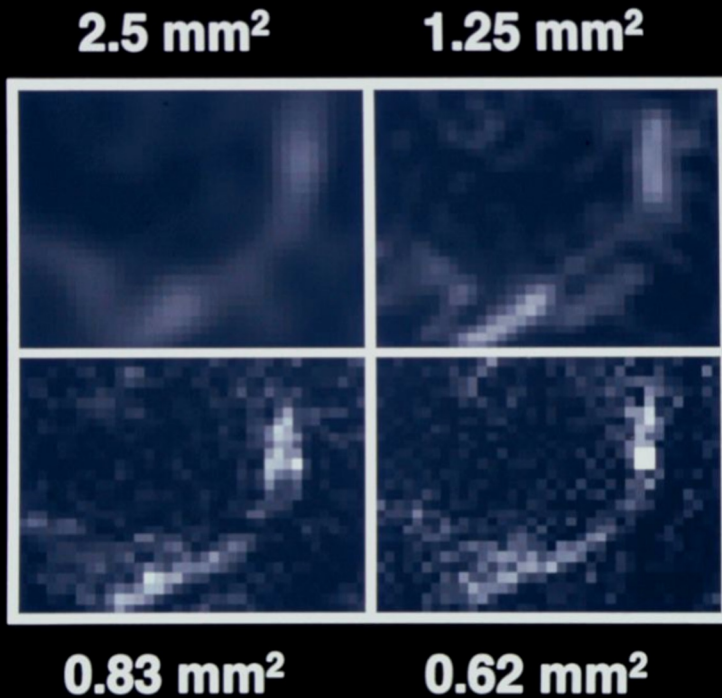
Ogawa, et al (2000), PNAS 97 (20)11026–11031

The Biggest Unknowns in Functional MRI

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

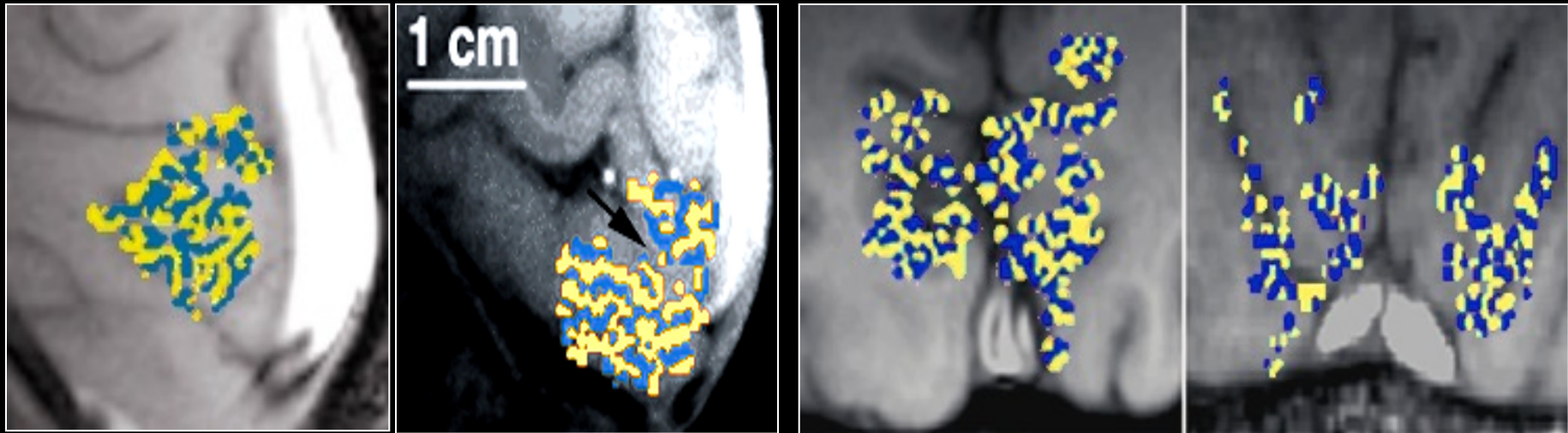
Magnitude

Fractional Signal Change

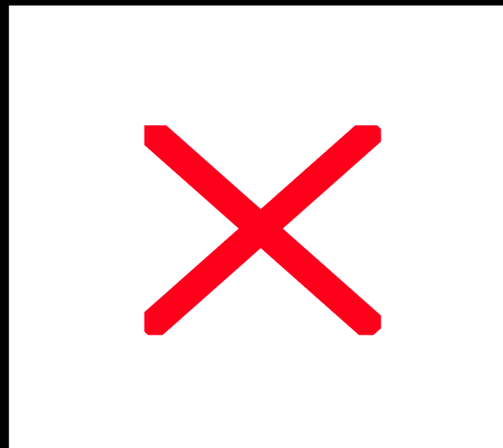


Jesmanowicz, P. A. Bandettini, J. S. Hyde, (1998) "Single shot half k-space high resolution EPI for fMRI at 3T." *Magn. Reson. Med.* 40, 754-762.

Ocular Dominance Column Mapping using fMRI



Menon, R. S., S. Ogawa, et al. (1997). "Ocular dominance in human V1 demonstrated by functional magnetic resonance imaging." *J Neurophysiol* 77(5): 2780-7.



Optical Imaging

R. D. Frostig et. al, PNAS 87: 6082-6086, (1990).

Human Ocular Dominance Columns as Revealed by High-Field Functional Magnetic Resonance Imaging

Kang Cheng,¹ R. Allen Waggoner, and Keiji Tanaka

Laboratory for Cognitive Brain Mapping

RIKEN Brain Science Institute and

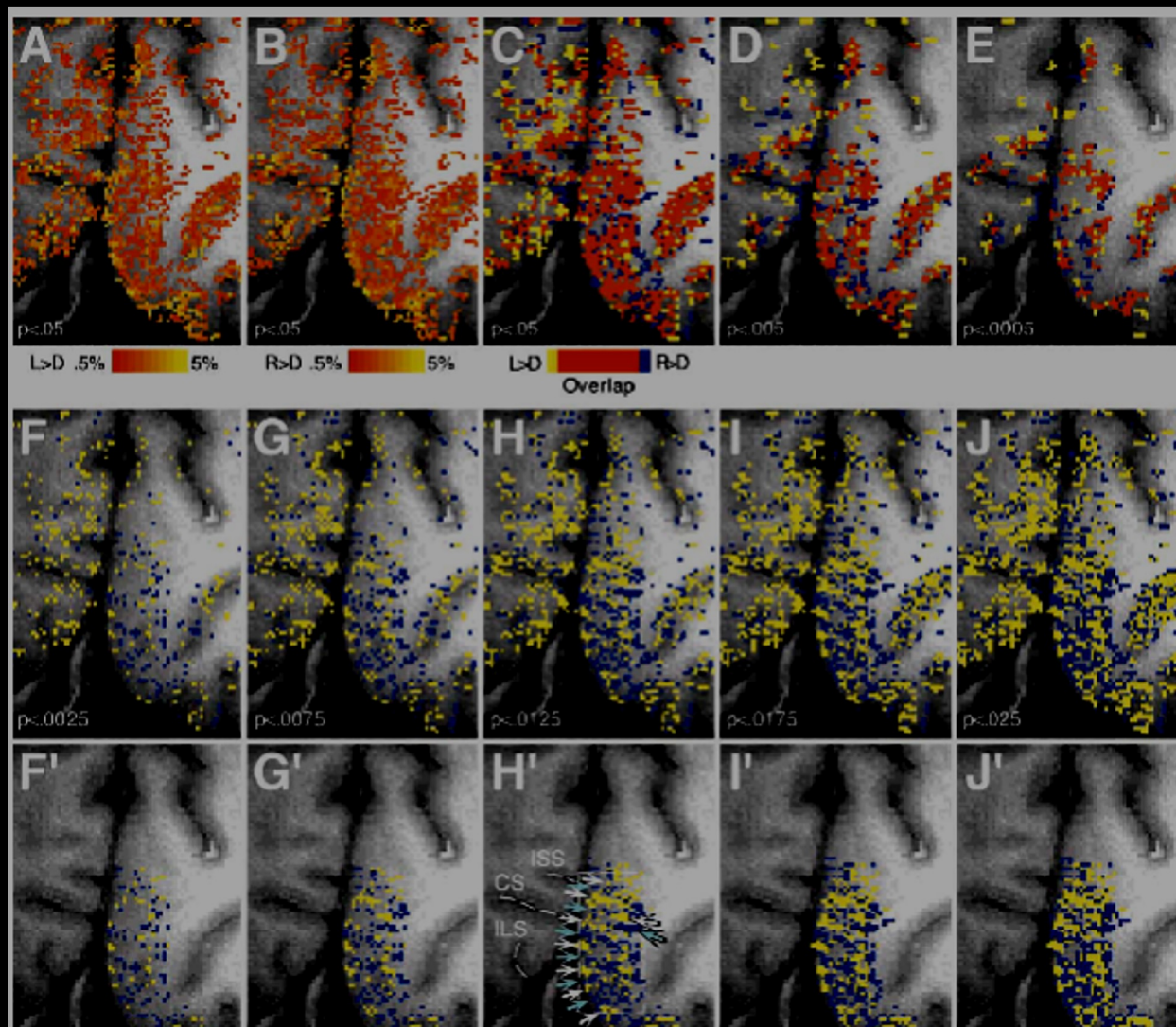
CREST

Japan Science and Technology Corporation

2-1 Hirosawa

Wako, Saitama 351-0198

Japan



Parallel acquisition (16 radio frequency channels)

Custom-built
Radio-frequency
(RF) coil



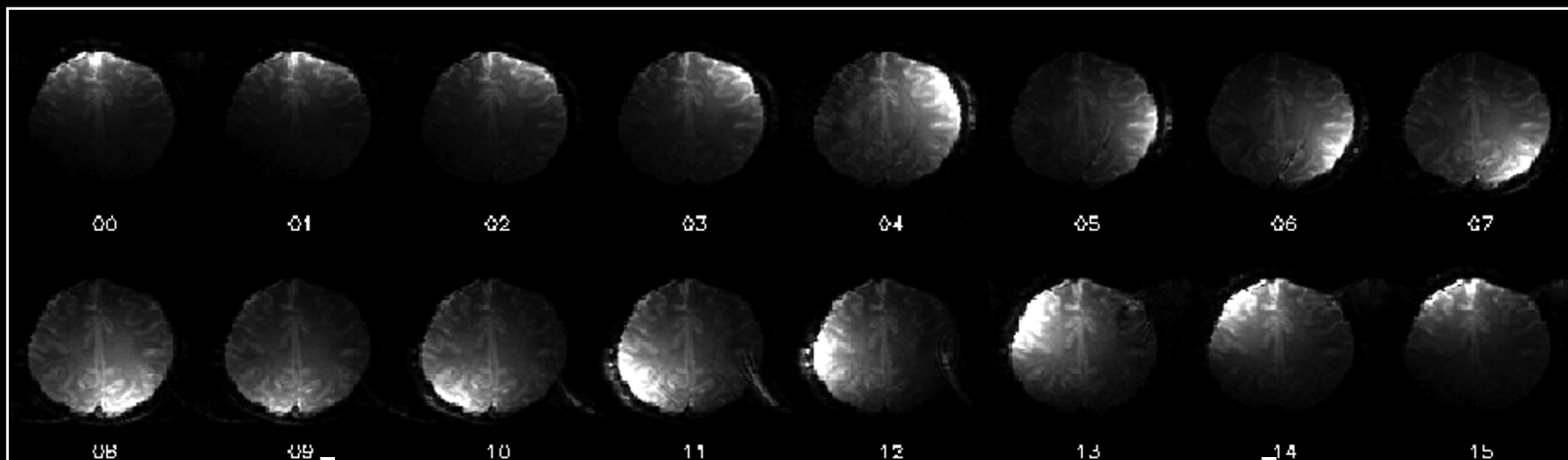
Nova Medical, Inc.

Parallel acquisition (16 radio frequency channels)

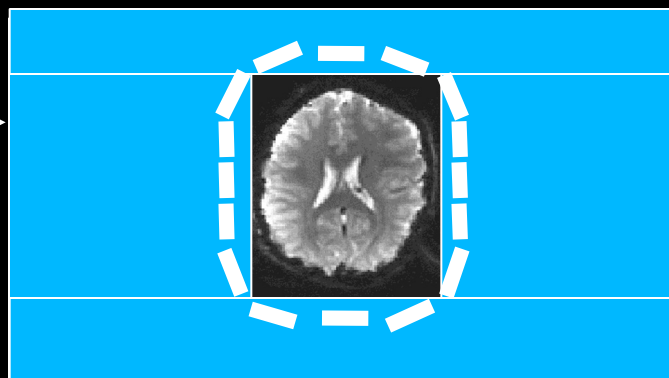
Receiver
Hardware



Individual coil images

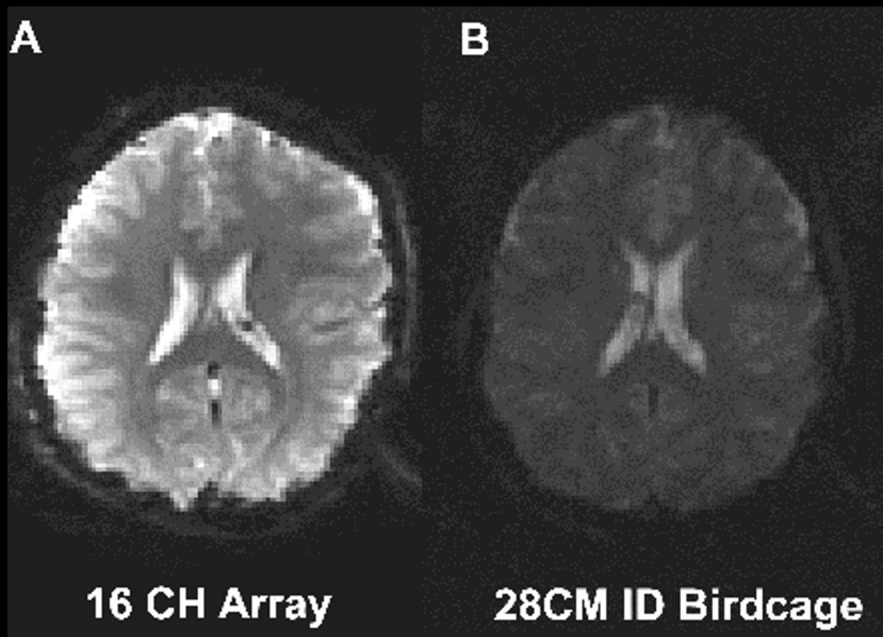


Single combined image



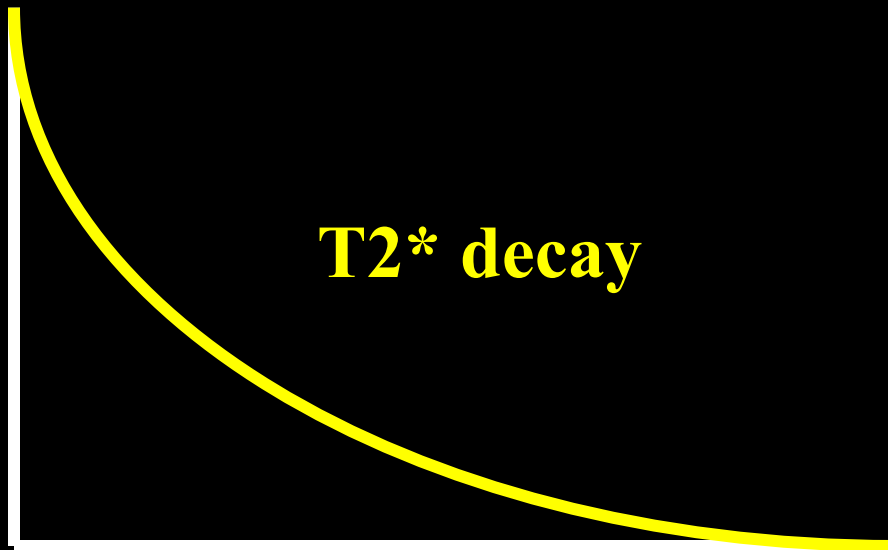
Parallel acquisition (16 radio frequency channels)

Large improvement in signal-to-noise ratio (SNR)



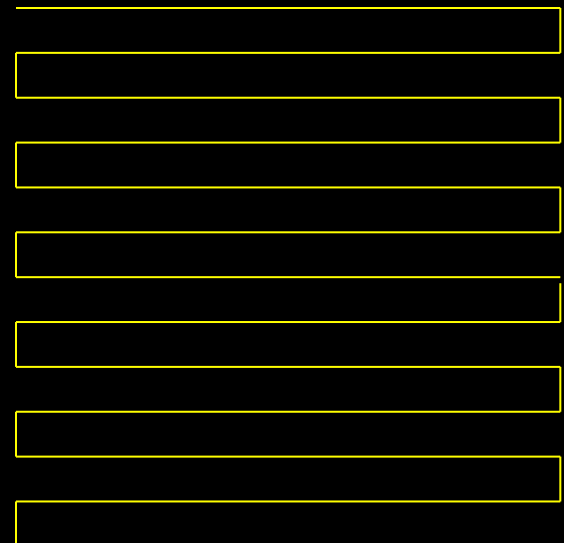
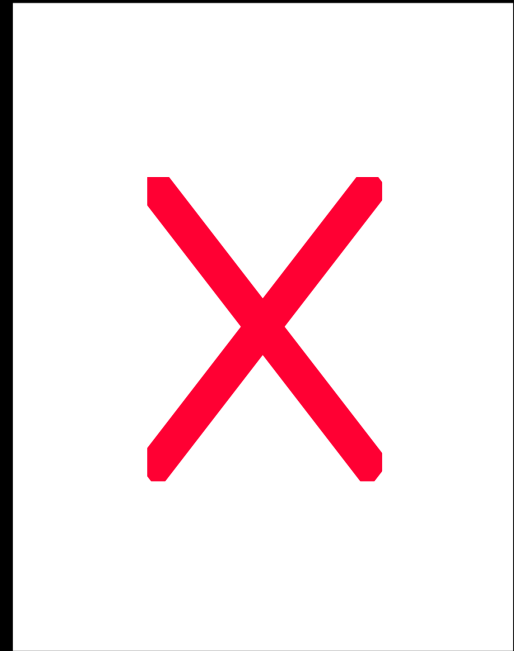
- Increased resolution
- Increased imaging speed
- Increased sensitivity

Single Shot EPI



EPI Readout Window

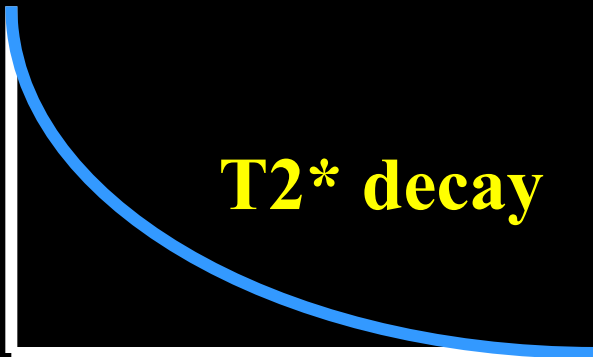
≈ 20 to 40 ms



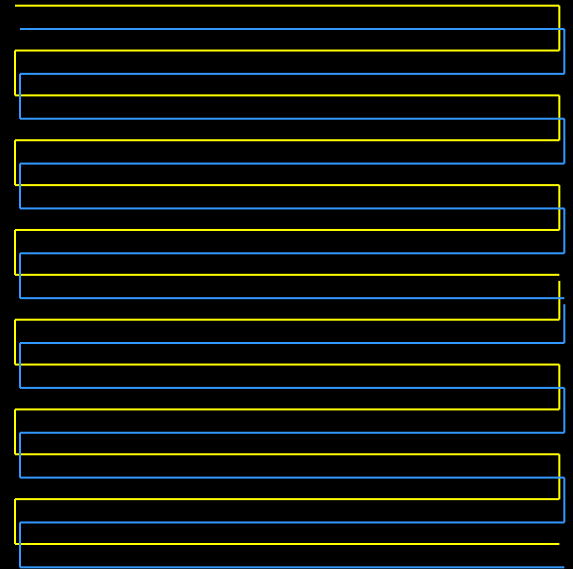
Multishot Imaging



EPI Window 1



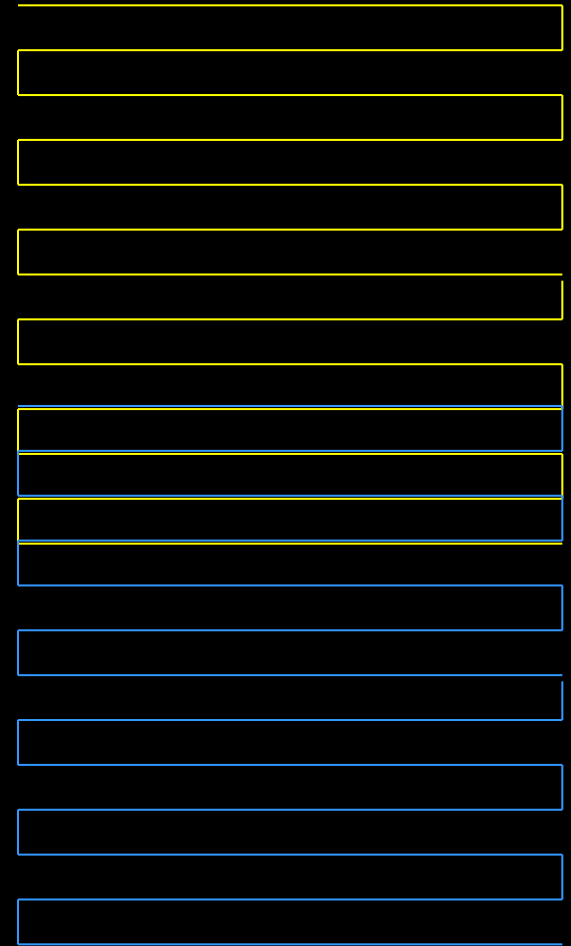
EPI Window 2



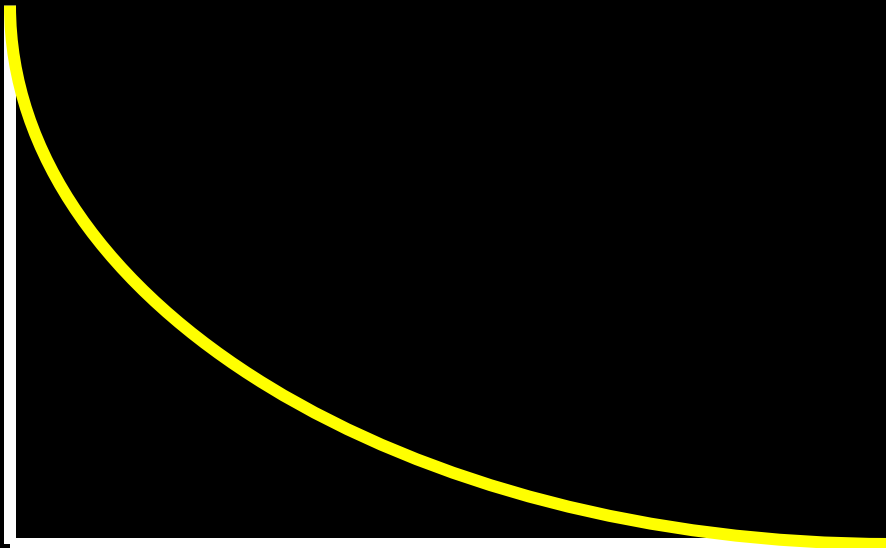
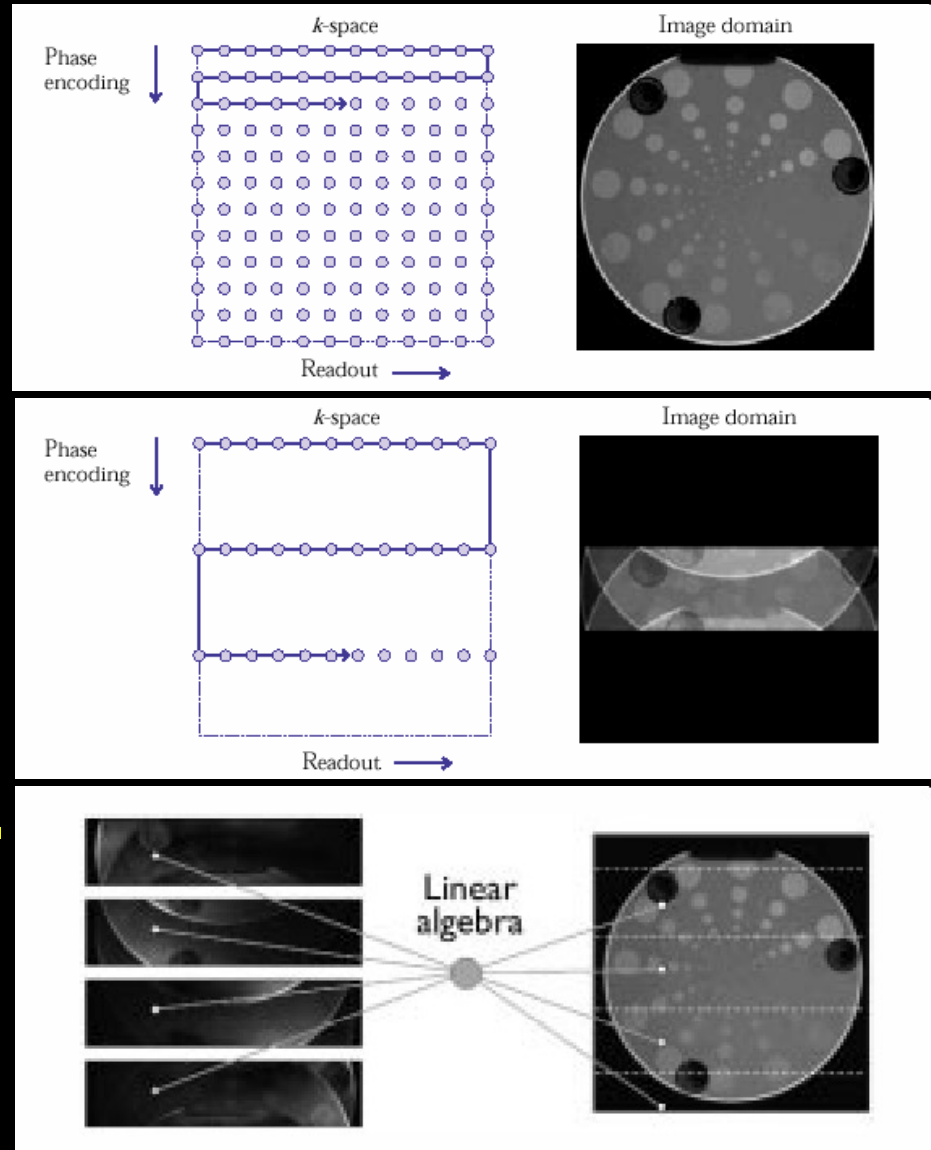
Partial k-space imaging



EPI Window



SENSE Imaging



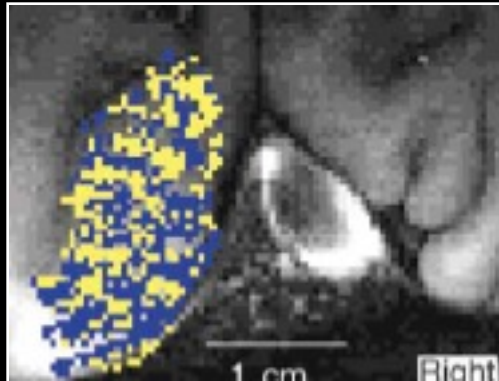
≈ 5 to 30 ms

Pruessmann, et al.

Ultimate spatial resolution?

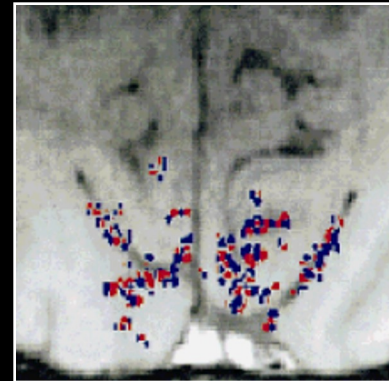
Resolving columns with single shot EPI is a goal..

0.47 x 0.47 in plane resolution



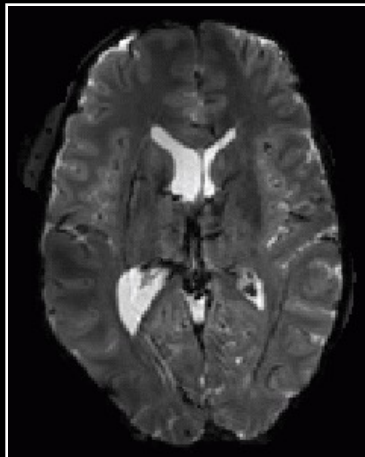
Cheng, et al. (2001) Neuron,32:359-374

0.54 x 0.54 in plane resolution



Menon et al, (1999) MRM 41 (2): 230-235

Multi-shot with
navigator pulse



...using SENSE, 32 channels, 7T,
and perhaps partial k-space we might get to 0.5 mm³

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

The Biggest Unknowns in Functional MRI

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

Ultimate clinical utility?

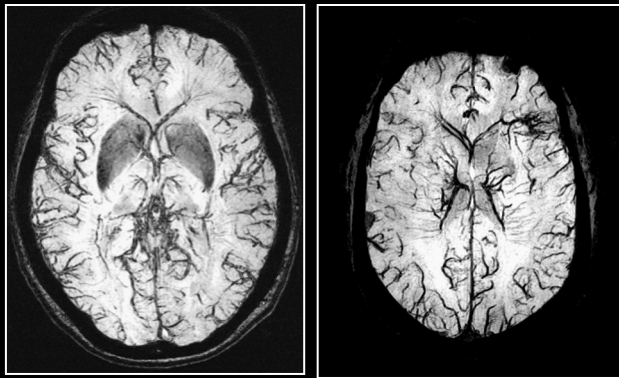
Needs:

Real time feedback

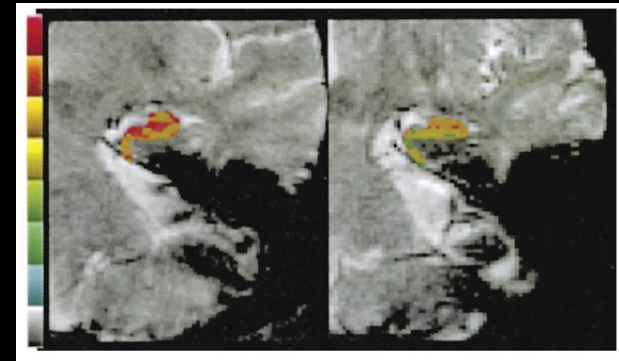
Characterization of confounding effects

Robust yet incisive set of probe tasks

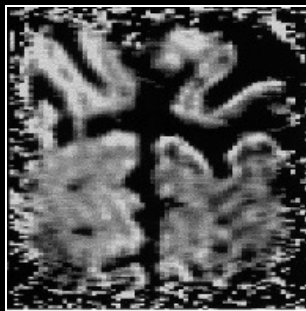
Baseline information?



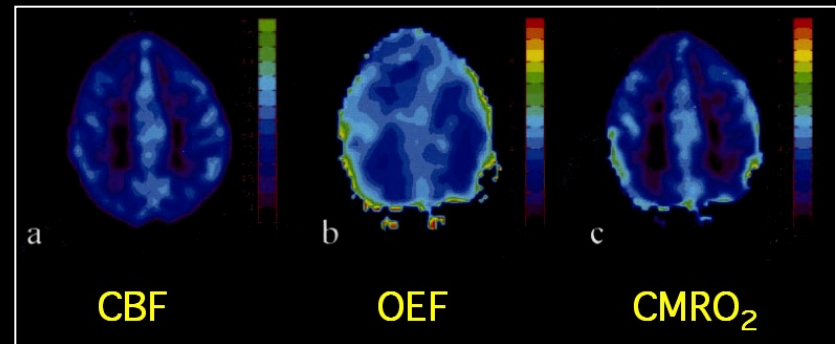
Bove-Bettis, et al (2004), SMRT



Small, et al (2001), Neuron 28:853-664



Bartha, et al (2002), MRM 47:742-750



An, et al (2001), NMR in Biomedicine 14:441-447

The Biggest Unknowns in Functional MRI

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

Best processing and display methods?

Processing

fMRI data, and noise is time and space varying in predictable and unpredictable ways over several temporal and spatial scales...

Signal and noise models...

Model free, open ended, methods?

Classification methods?

Multivariate methods?

Connectivity (across time and space scales?)

Best processing and display methods?

Display

To convey:

- collapsed multidimensional data
- sense of data quality

Surface

Glass brain

ROI

Time courses

Example slices

Connectivity maps?

“Quality” index?

The Biggest Unknowns in Functional MRI

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

Optimal Field Strength?

Utility vs. Difficulty

Difficulty:

Shimming (generally lower T2 and T2*)
RF penetration effects
Stability

Utility:

Higher SNR
Better susceptibility contrast
Better ASL perfusion contrast (longer T1)



Functional Imaging Methods Unit &



Functional MRI Facility

Computer Specialist:

Adam Thomas

Staff Scientists:

Sean Marrett

Jerzy Bodurka

Frank Ye

Wen-Ming Luh

Rasmus Birn

Post Docs:

Hauke Heekeren

David Knight

Anthony Boemio

Niko Kriegeskorte

Scanning Technologists:

Karen Bove-Bettis

Paula Rowser

Alda Ottley

Ellen Condon

Program Assistant:

Kay Kuhns

Graduate Student:

Natalia Petridou

Unit on Functional Imaging & FMRI Core Facility



<http://sodium.nimh.nih.gov/upload>
T165.ppt