

Latest Developments of fMRI

Peter A. Bandettini, Ph.D.

Section on Functional Imaging Methods

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

Laboratory of Brain and Cognition

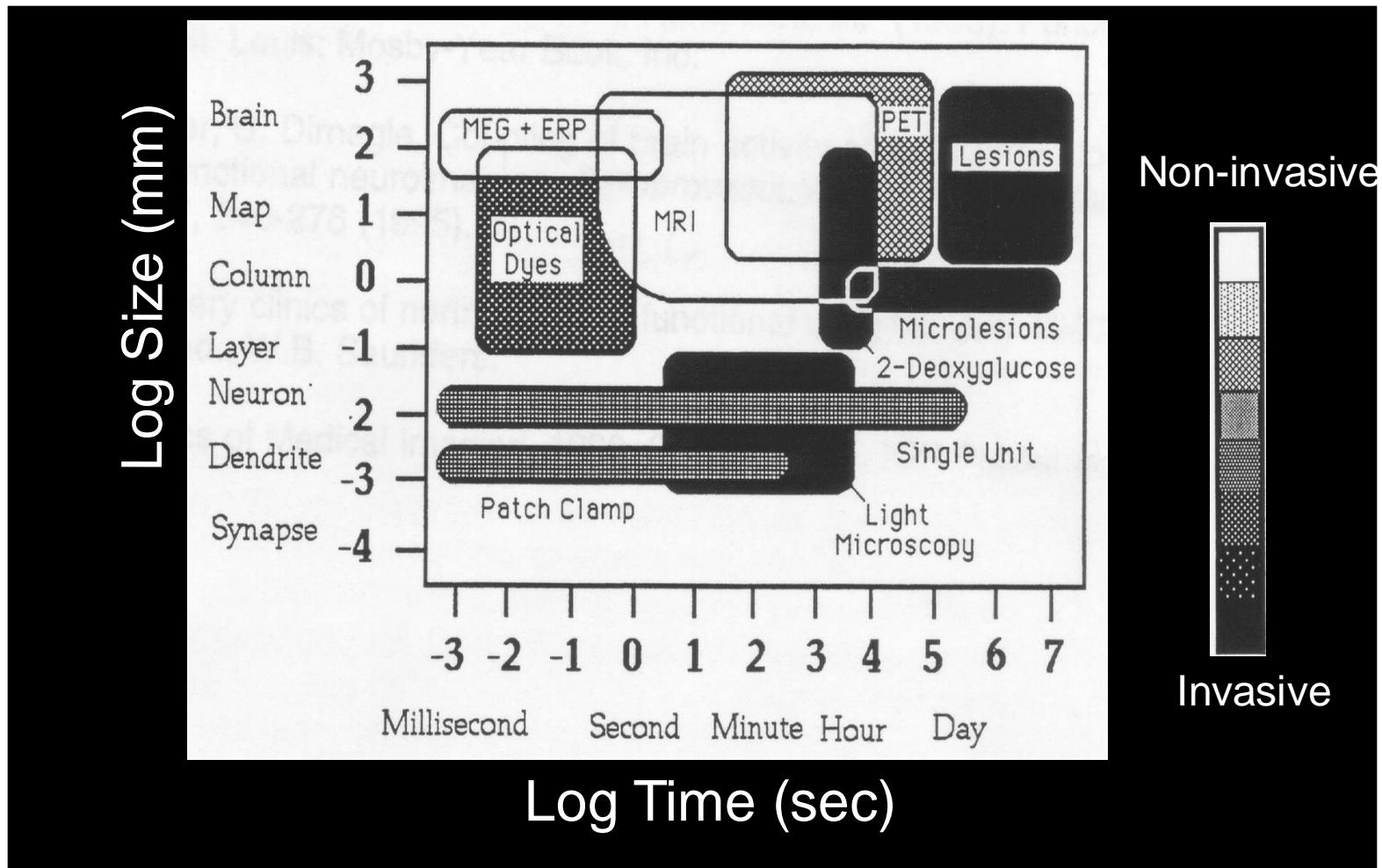
&

Functional MRI Facility

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



Functional Neuroimaging Techniques



A brief overview of the three main types of fMRI contrast

Volume

Flow or Perfusion

Oxygenation

Blood Volume

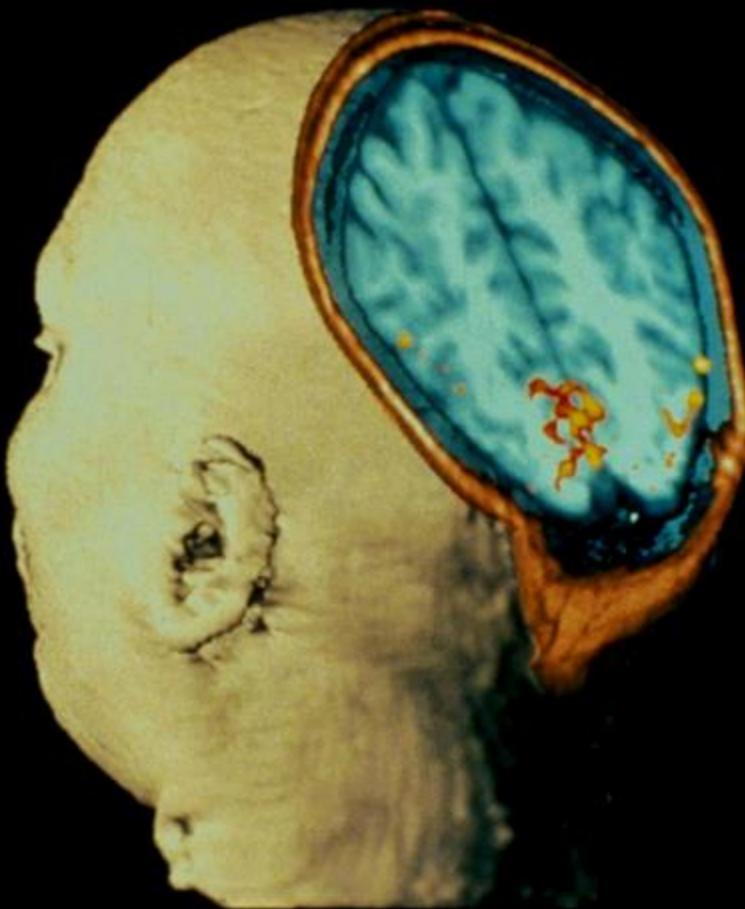
What started it all...

**Photic
Stimulation**

MRI Image showing
activation of the
Visual Cortex

From Belliveau, et al.
Science Nov 1991

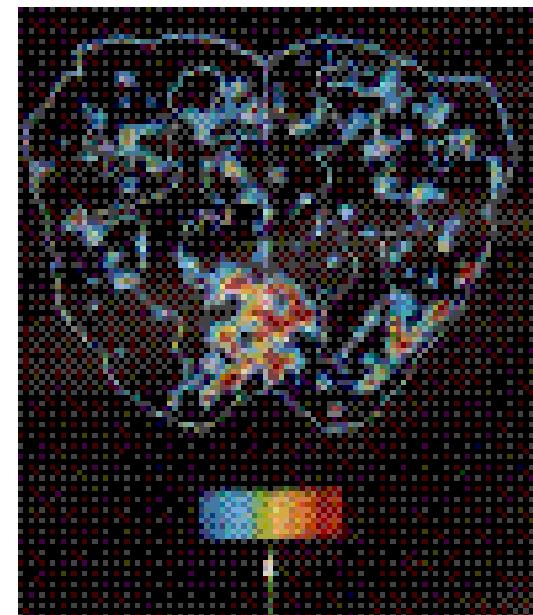
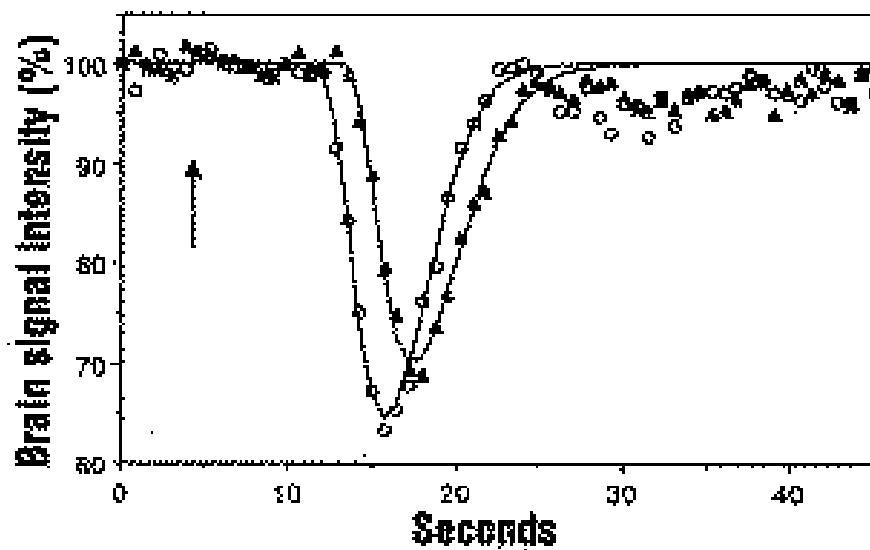
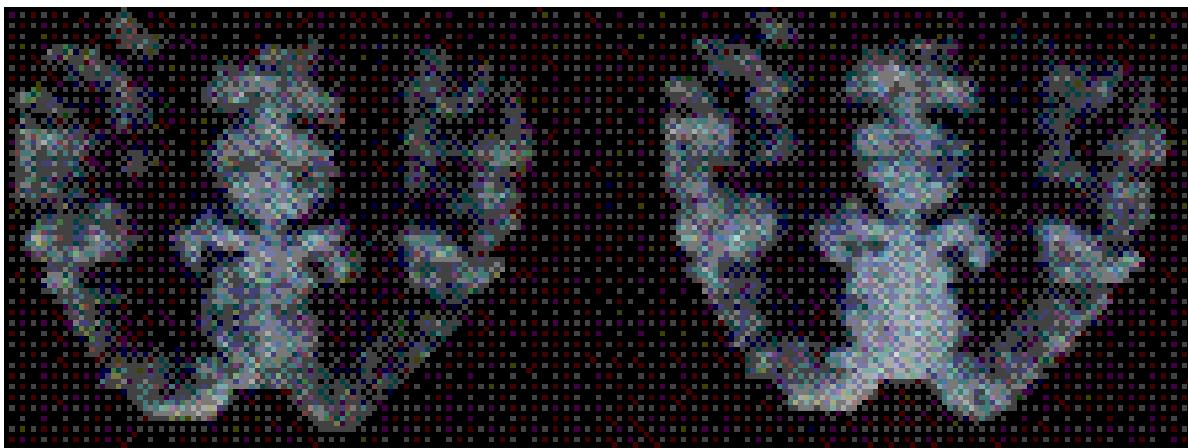
MSC - perfusion



Blood Volume

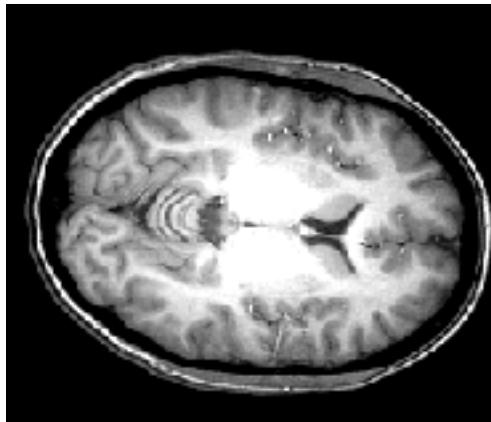
Resting

Active



MRI vs. fMRI

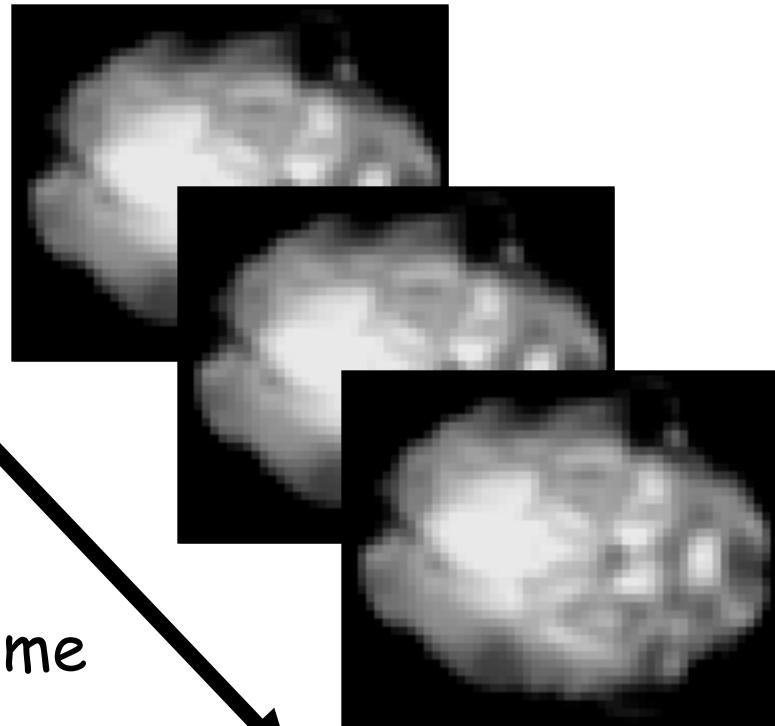
MRI



one image

high resolution
(1 mm or less)

fMRI



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

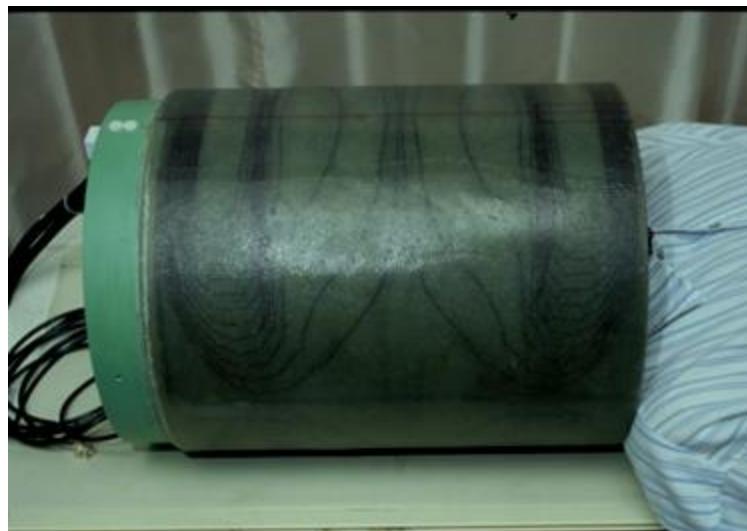
low resolution
(1.5 to 4 mm)



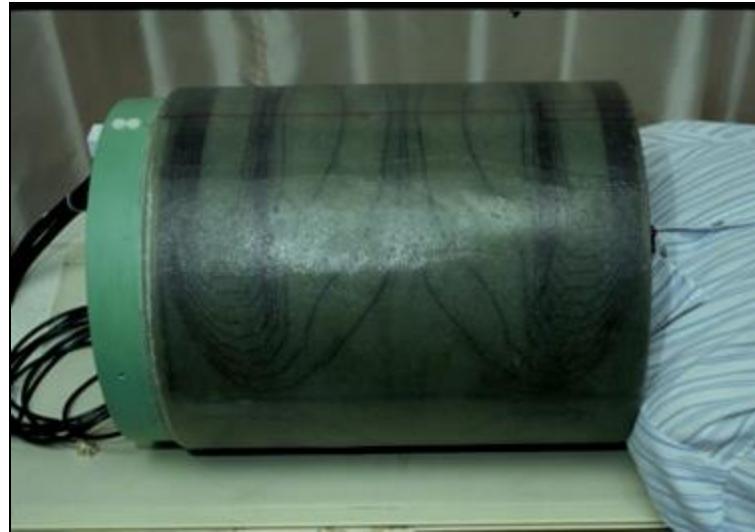
1991-1992



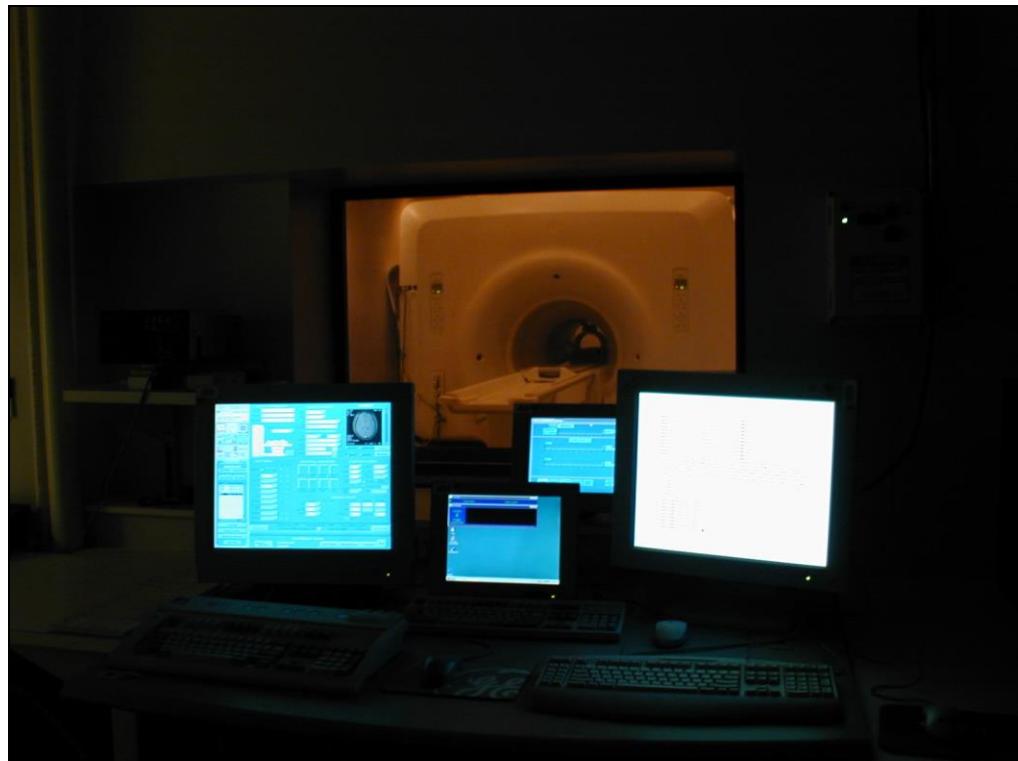
1992-1999



**Local Gradient Coil
(low inductance)**



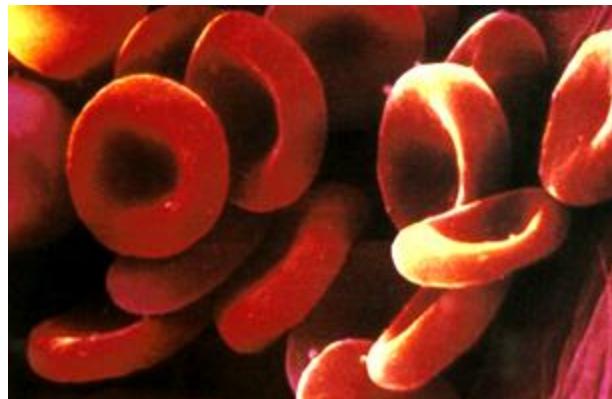
**Whole body gradients
(more powerful amplifiers)**





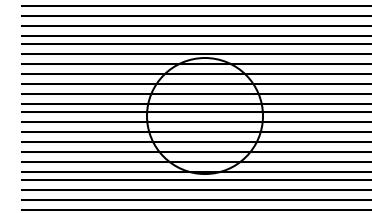
Blood Oxygenation

Oxygenated and deoxygenated red blood cells have different magnetic properties

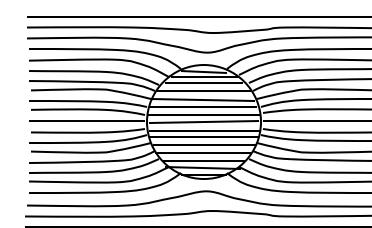


red blood cells

oxygenated



deoxygenated

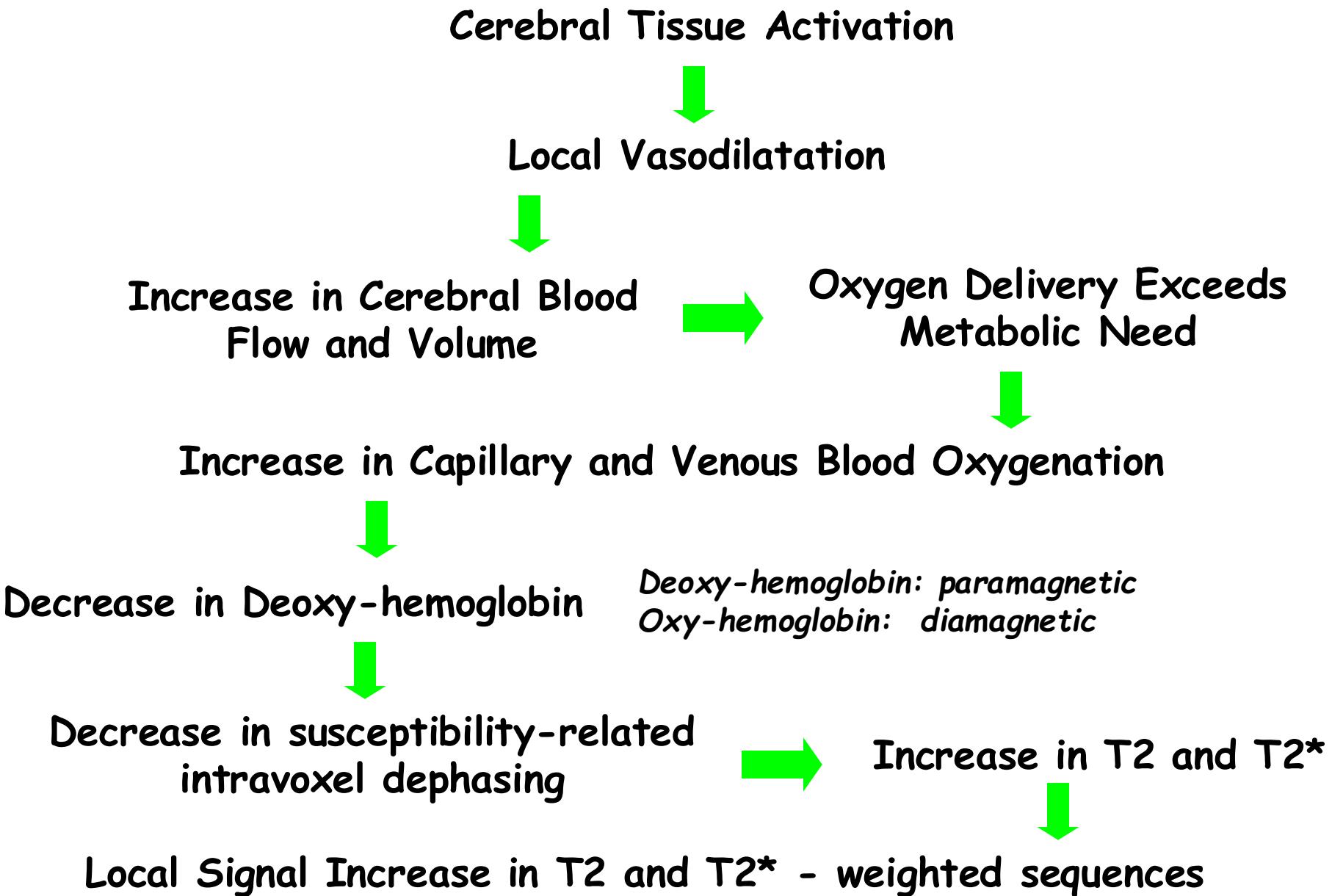


L. Pauling, C. D. Coryell, Proc. Natl. Acad. Sci. USA 22, 210-216, **1936**.

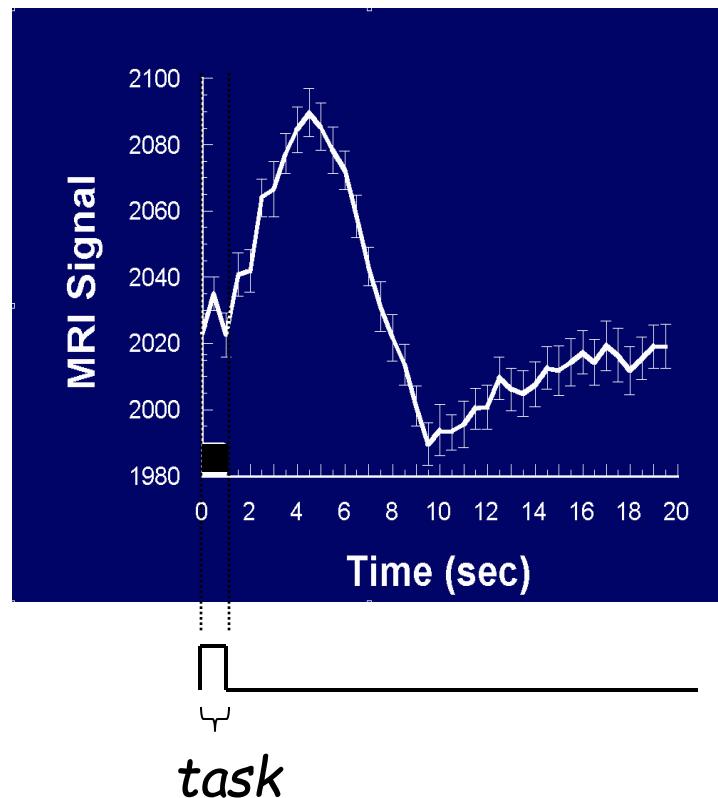
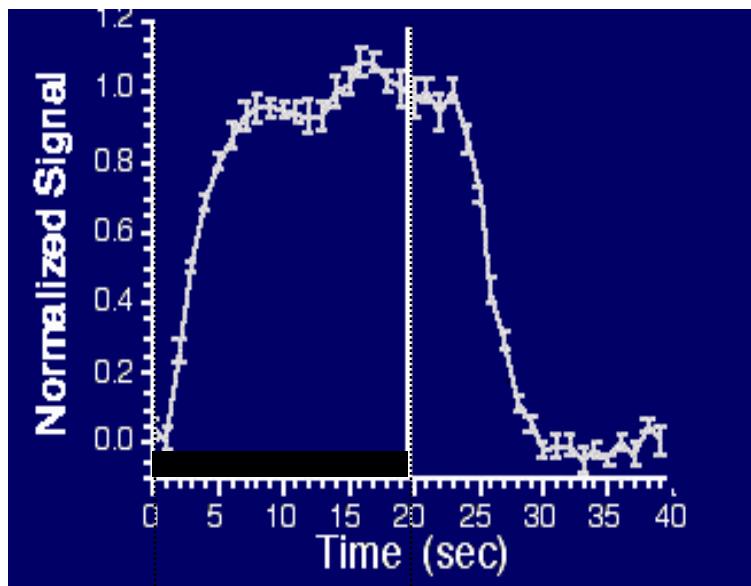
K.R. Thulborn, J. C. Waterton, et al., Biochim. Biophys. Acta. 714: 265-270, **1982**.

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

Blood Oxygenation



Blood Oxygenation

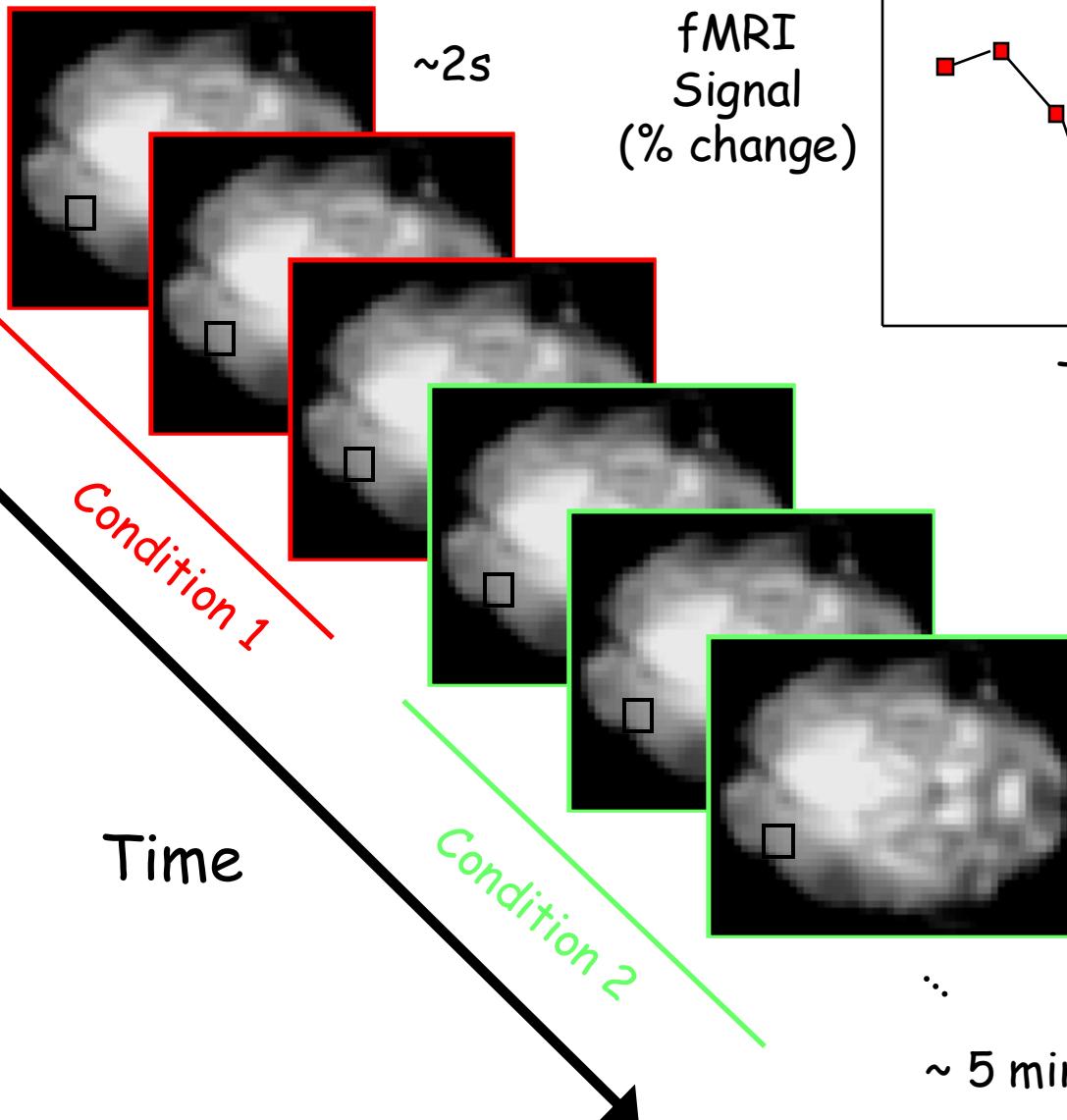




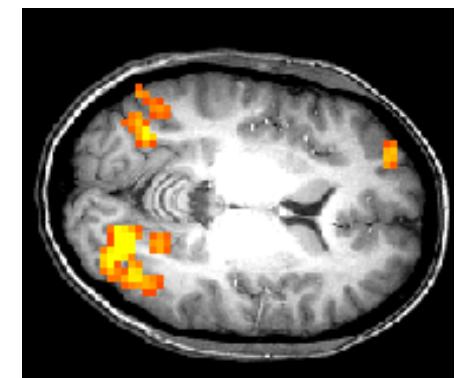
1991

Activation Statistics

Functional images



Statistical Map
superimposed on
anatomical MRI image

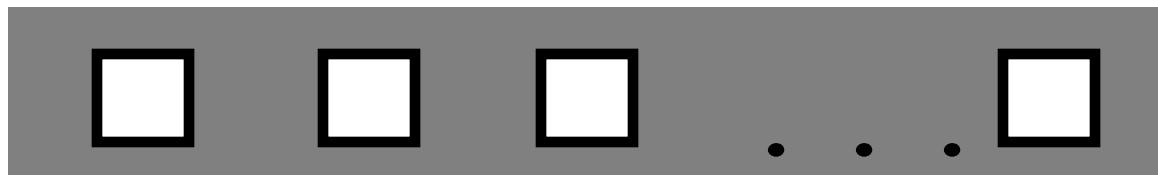
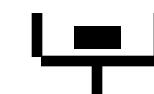
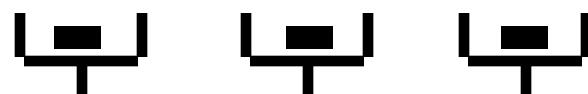
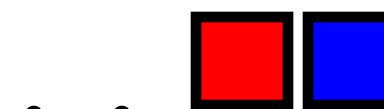
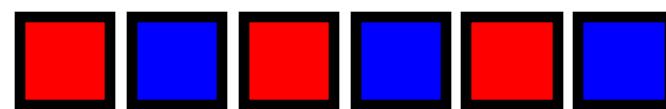
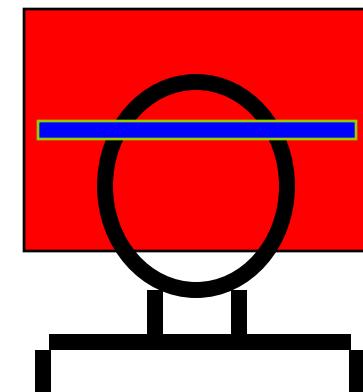
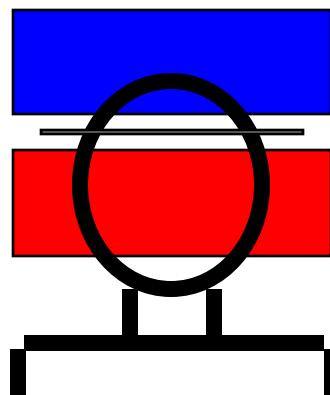


Courtesy, Robert Cox

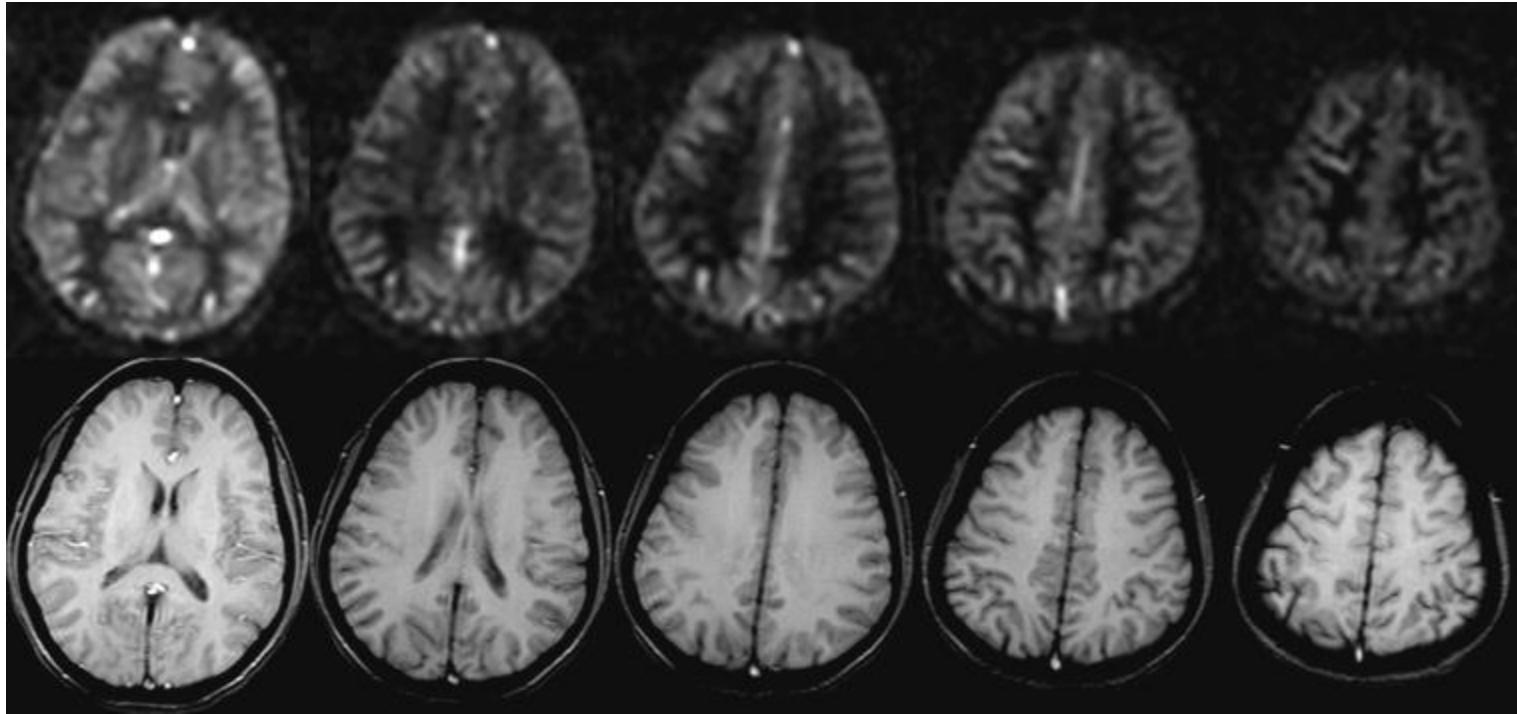
Perfusion

EPISTAR

FAIR



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 (EPICSTAR)." Radiology 192, 1-8.

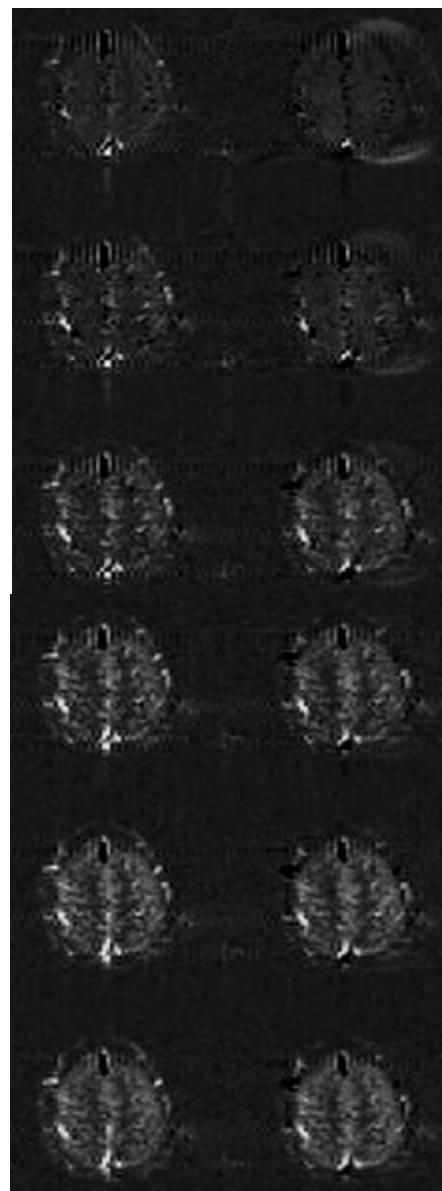
Kim, S.-G. (1995) "Quantification of relative cerebral blood flow change by flow-sensitive alternating inversion recovery (FAIR) technique: application to functional mapping." Magn. Reson. Med. 34, 293-301.

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



Overview of fMRI

Functional Contrast:

Blood volume

Blood flow/perfusion

Blood oxygenation

Spatial resolution:

Typical: 3 mm^3

Upper: 0.5 mm^3

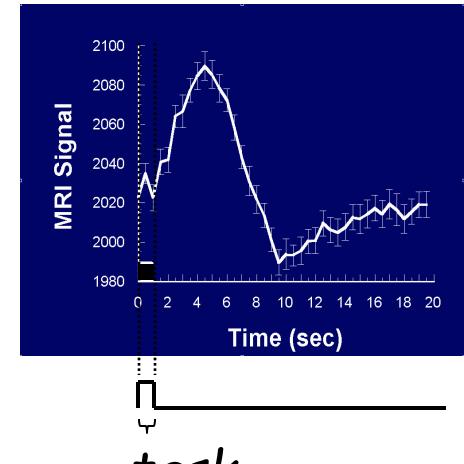
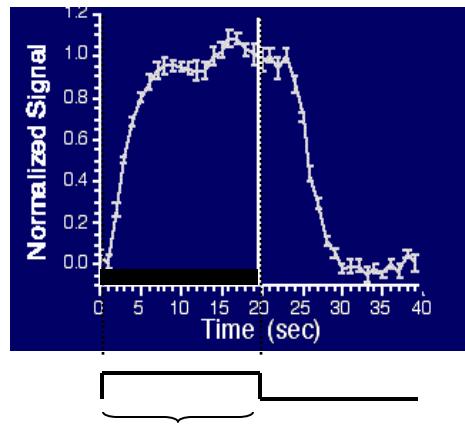
Temporal resolution:

Minimum duration: < 16 ms

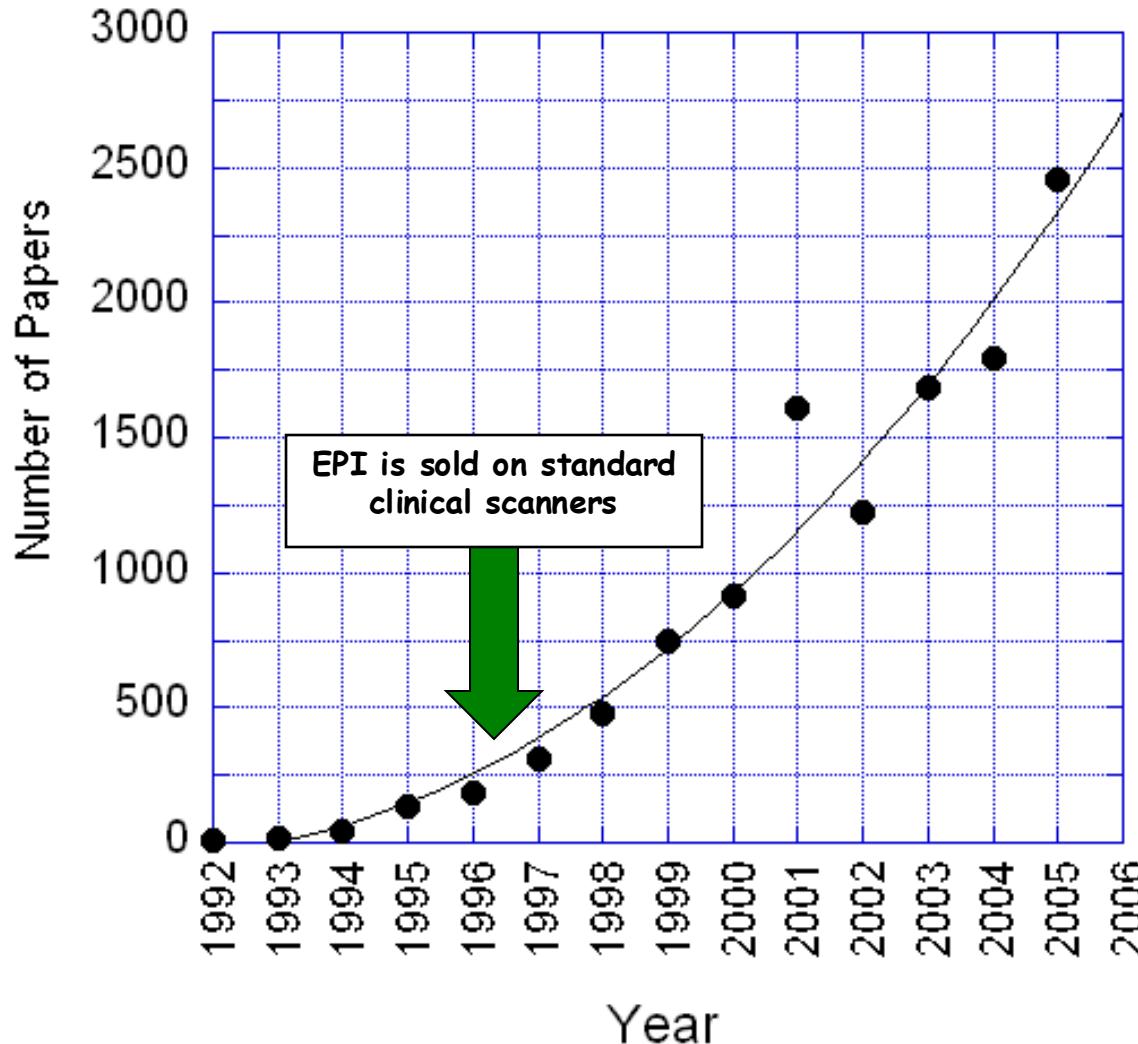
Minimum onset diff: 100 ms to 2 sec

Interpretability:

Neurovascular coupling, vascular sampling, blood, physiologic noise, motion and other artifacts, etc..

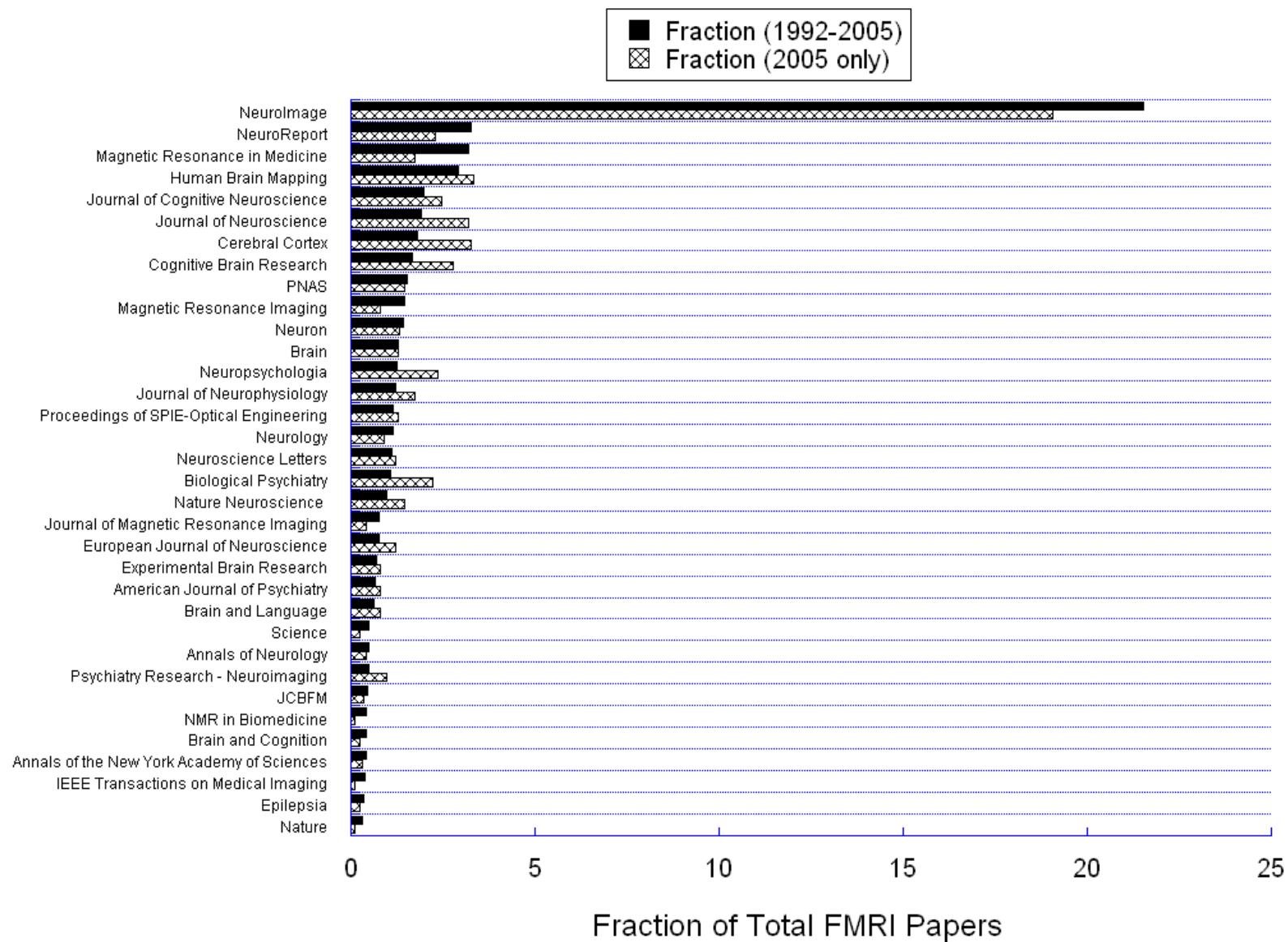


fMRI Papers Published per Year

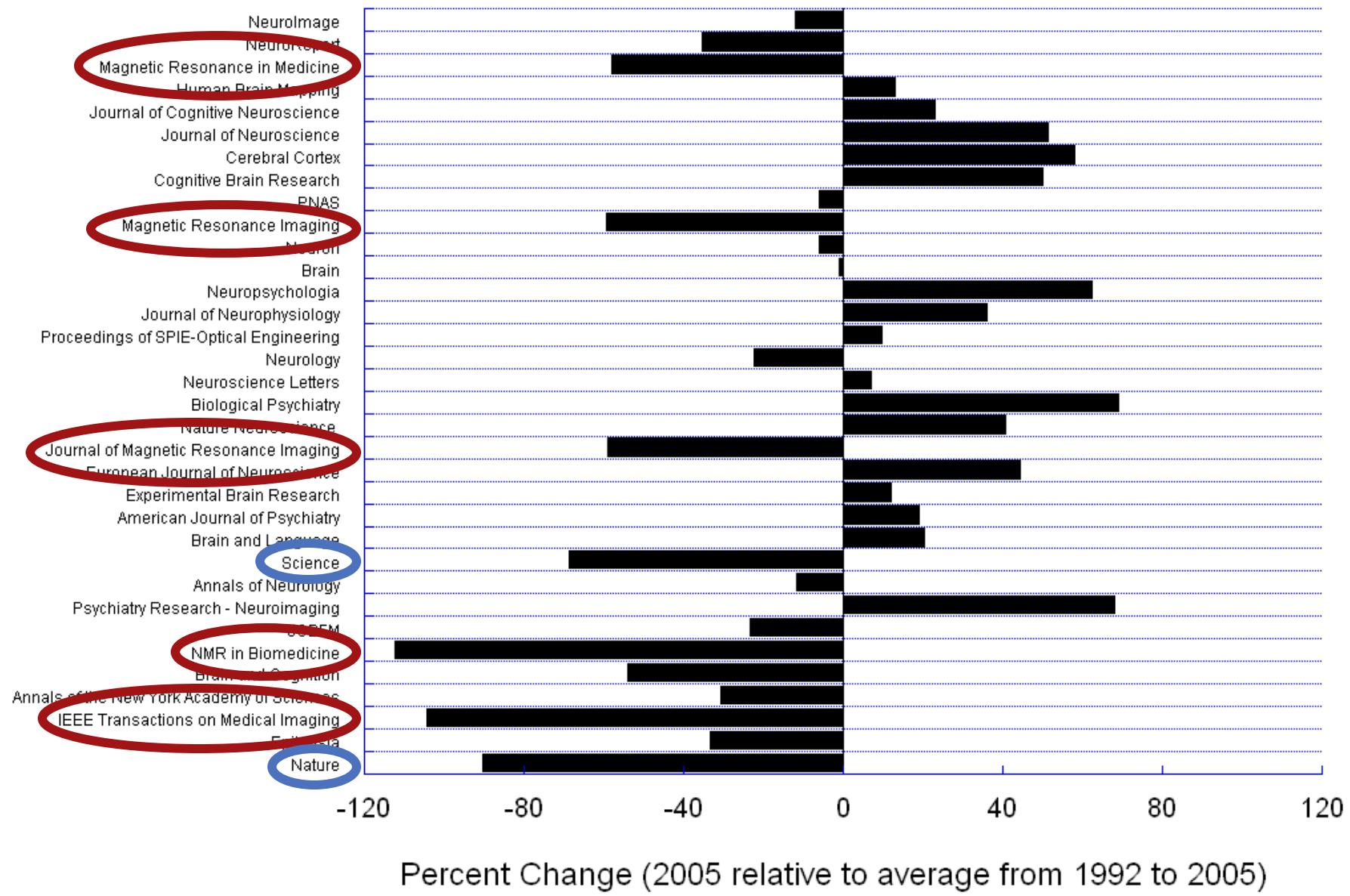


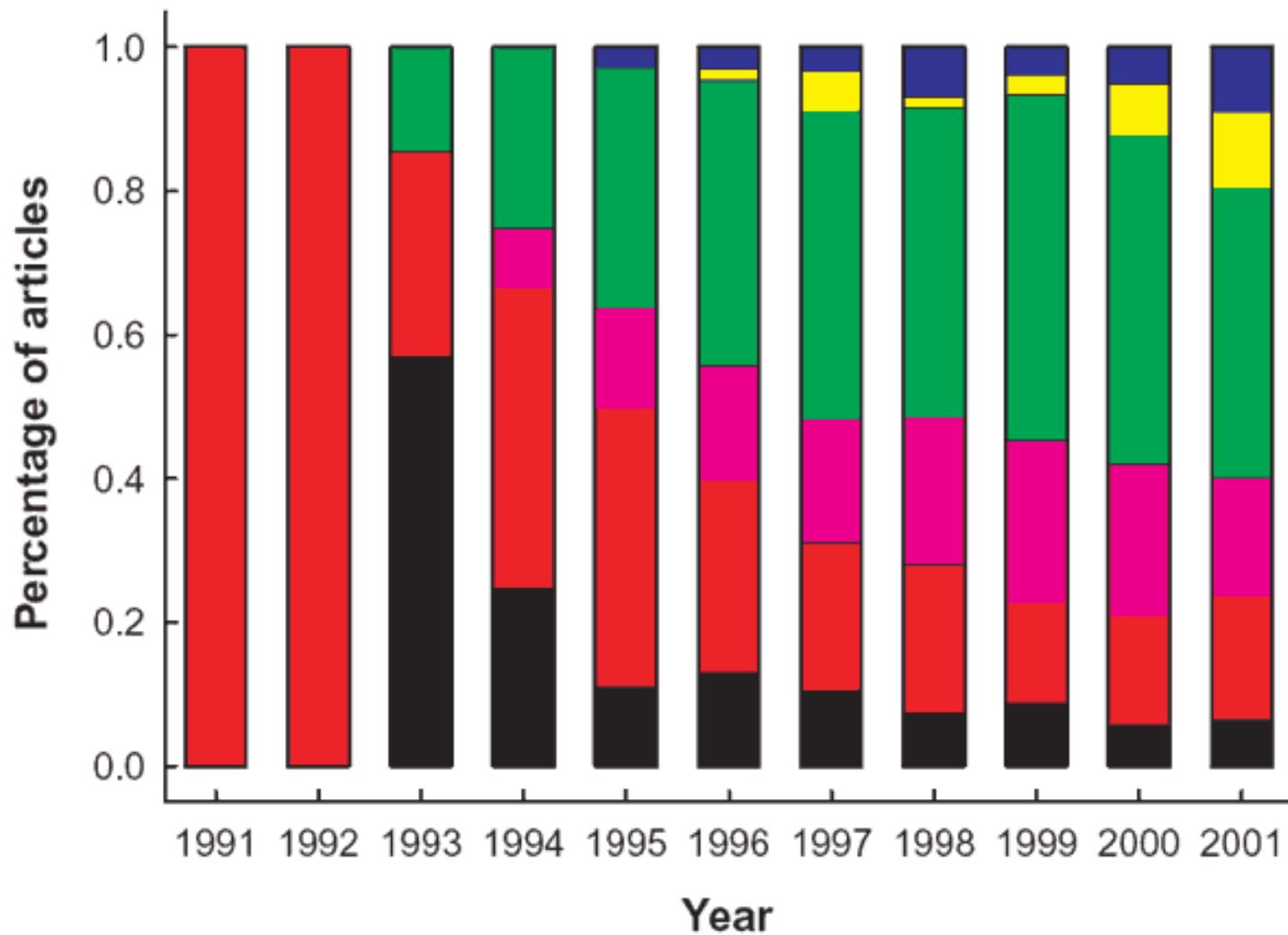
"fMRI" or "functional MRI"

Breakdown of fMRI papers by Journal



Percent Change in fMRI Publications of 2005 relative to Average (1992 - 2005) for Each Journal





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

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

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

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.

Technology

Coil arrays
High field strength
High resolution
Novel functional contrast

Methodology

Functional Connectivity Assessment
Multi-modal integration
Pattern classification
Real time feedback
Task design

Fluctuations
Dynamics
Cross - modal comparison

Basic Neuroscience
Behavior correlation/prediction
Pathology assessment

Interpretation

Applications

Technology

Coil arrays
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Methodology

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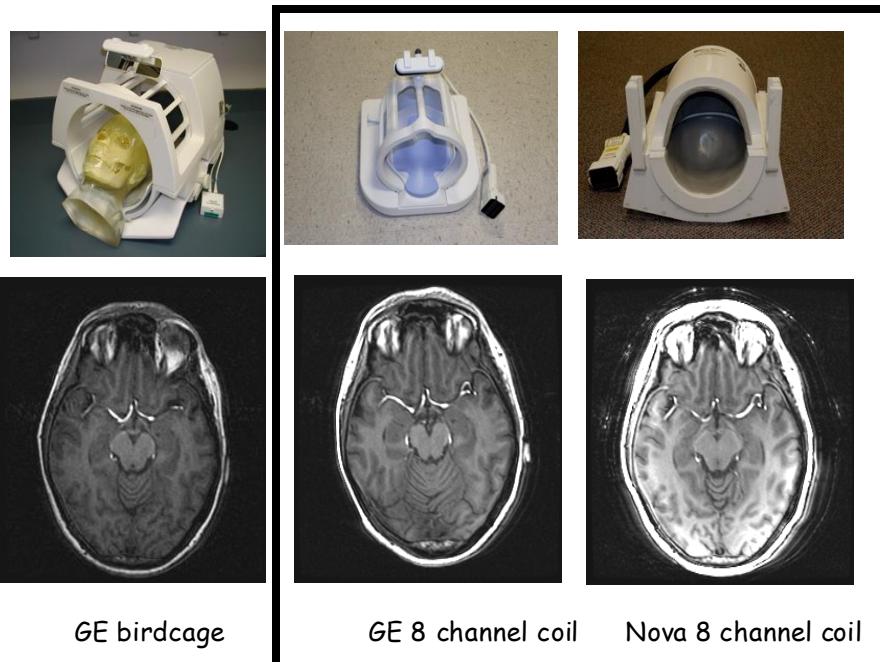
Basic Neuroscience
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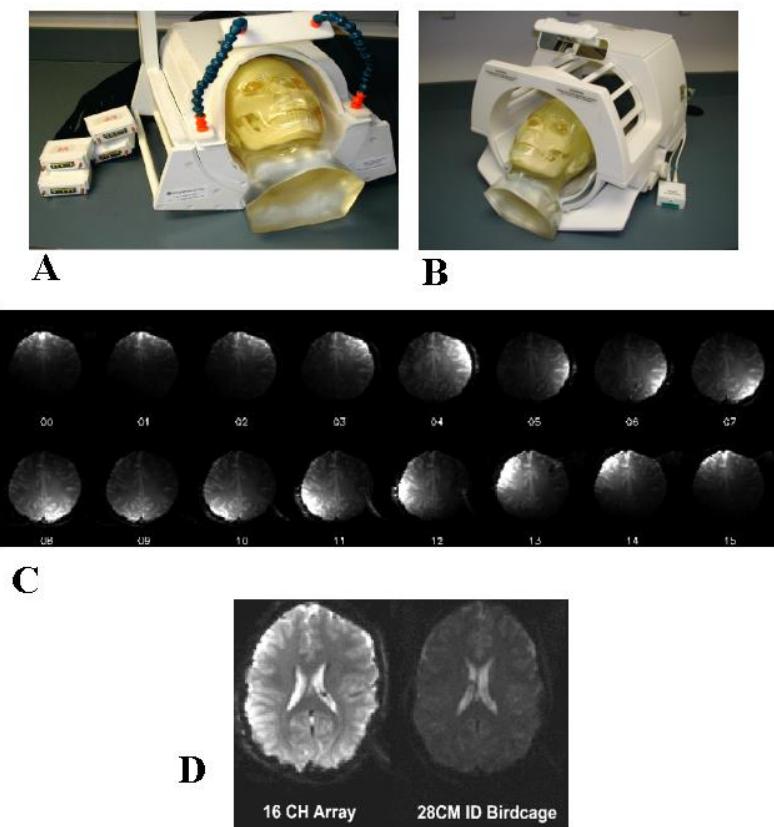
Applications

Technology

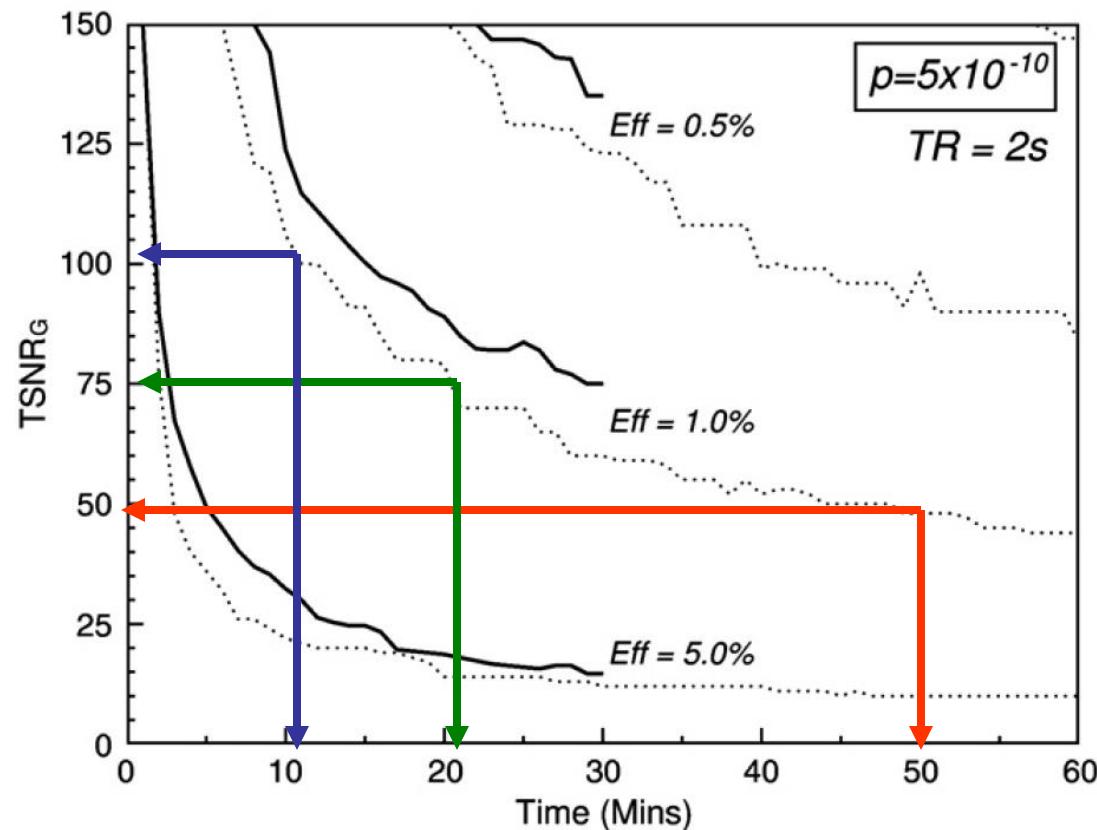
8 channel parallel receiver coil



16 channel parallel receiver coil



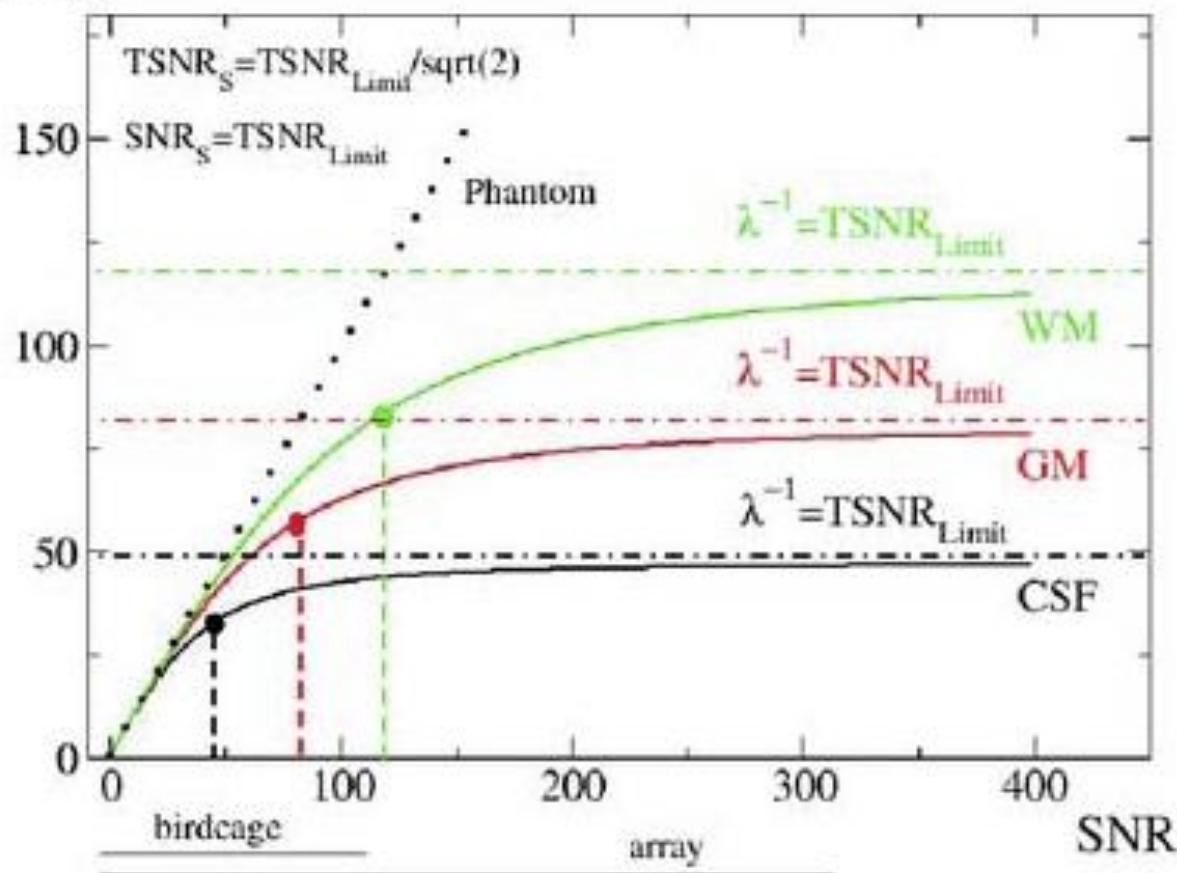
Technology



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)

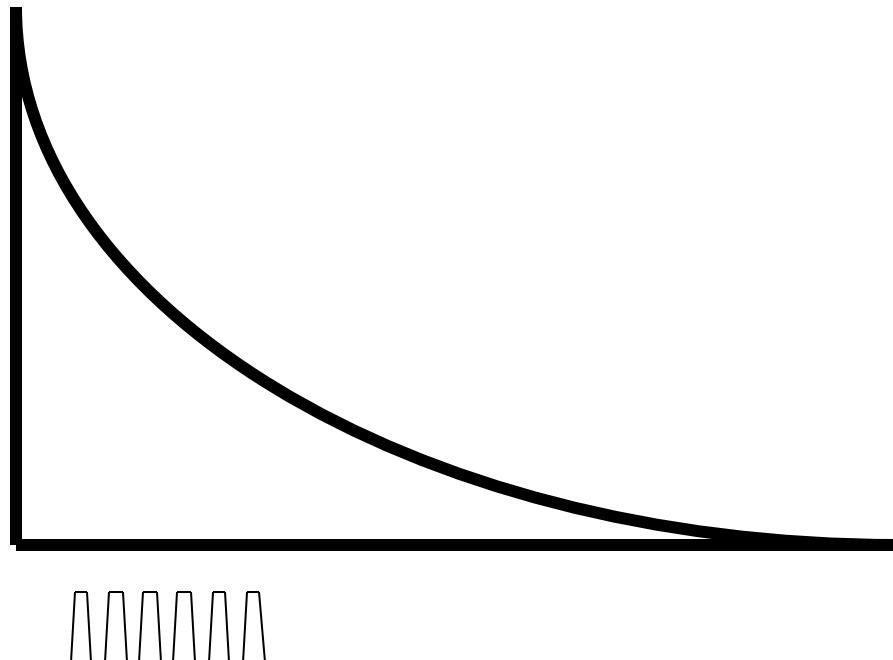
SNR vs TSNR

TSNR

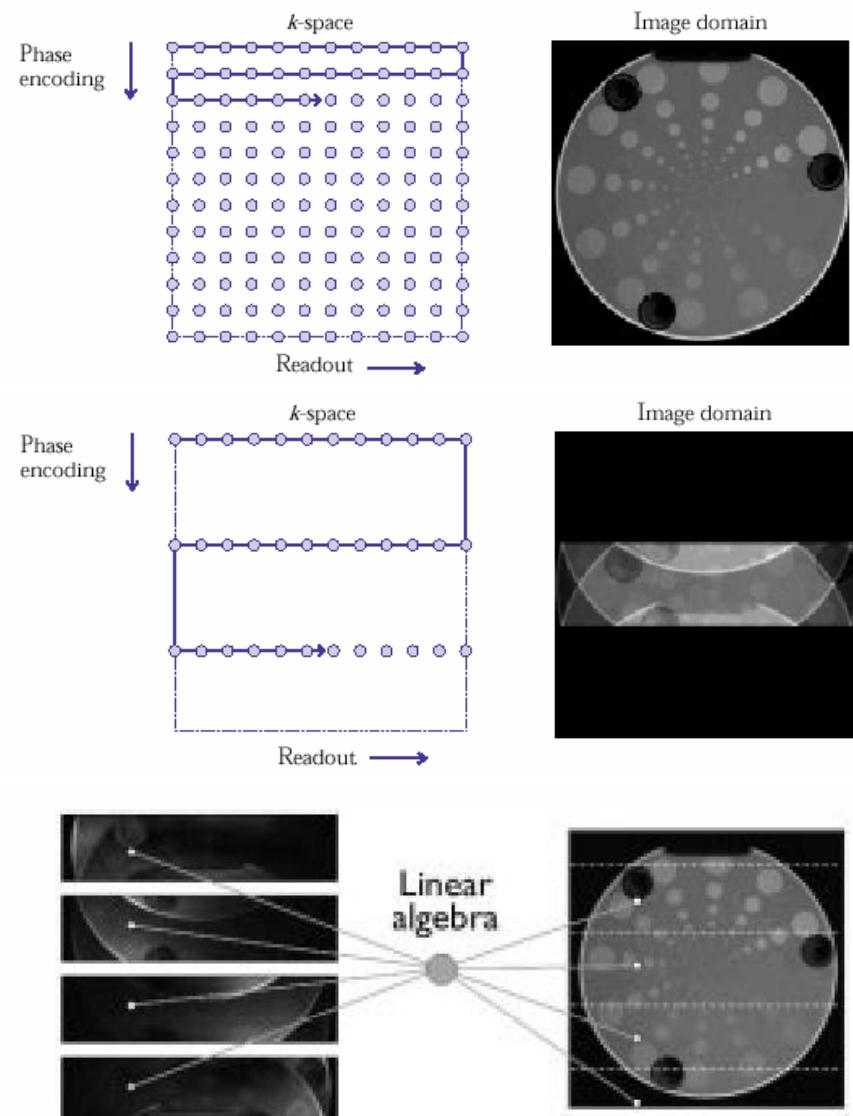


J. Bodurka, F. Ye, N Petridou, P. A. Bandettini, Mapping the MRI voxel volume in which thermal noise matches physiological noise - implications for fMRI. *NeuroImage*, 34, 542-549 (2007)

Technology

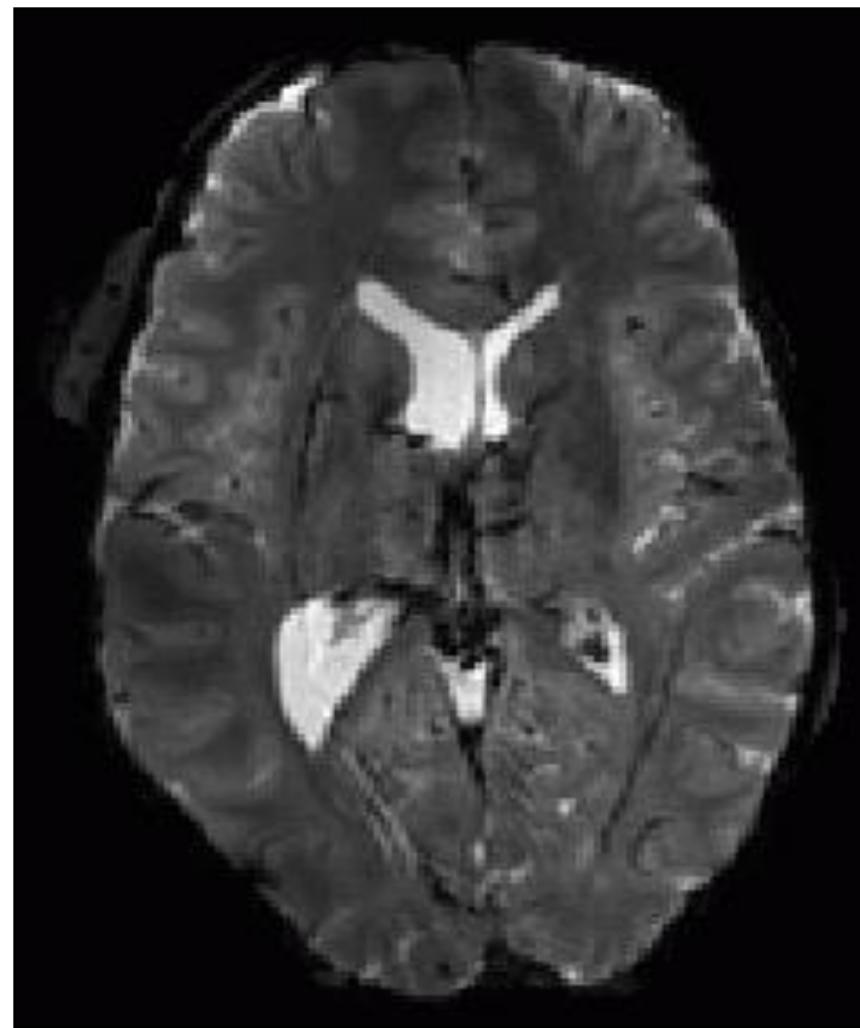


≈ 5 to 30 ms



Pruessmann, et al.

Technology



3T single-shot SENSE EPI using 16 channels: $1.25 \times 1.25 \times 2\text{mm}$

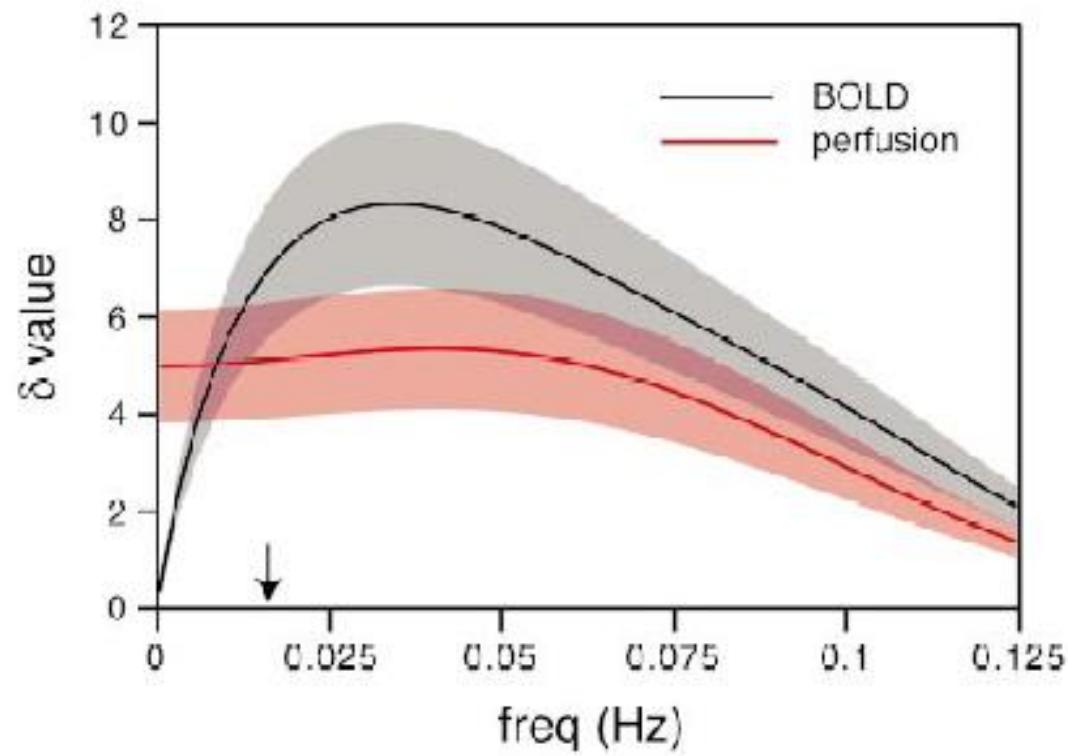
fMRI Contrast

- Volume (gadolinium)
- BOLD
- Perfusion (ASL)
- ΔCMRO_2
- $\Delta\text{Volume (VASO)}$
- Neuronal Currents
- Diffusion coefficient
- Temperature

Methodology

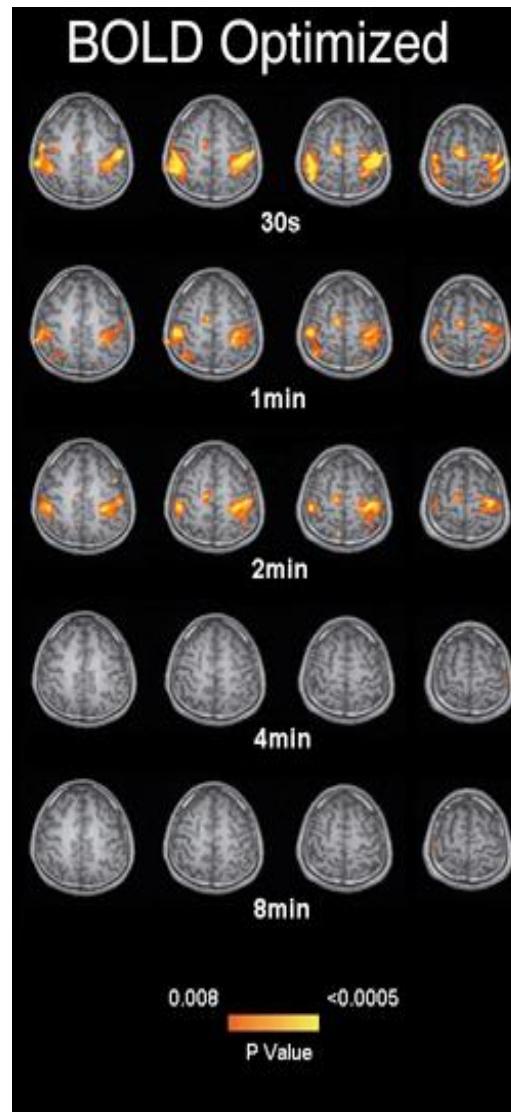
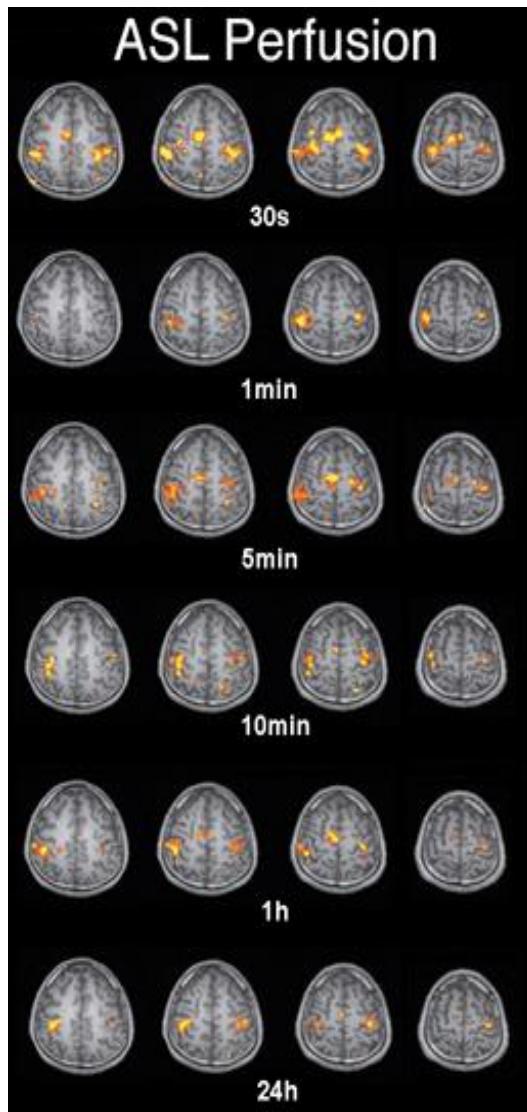
Perfusion (ASL)

Better than BOLD for long duration activation...

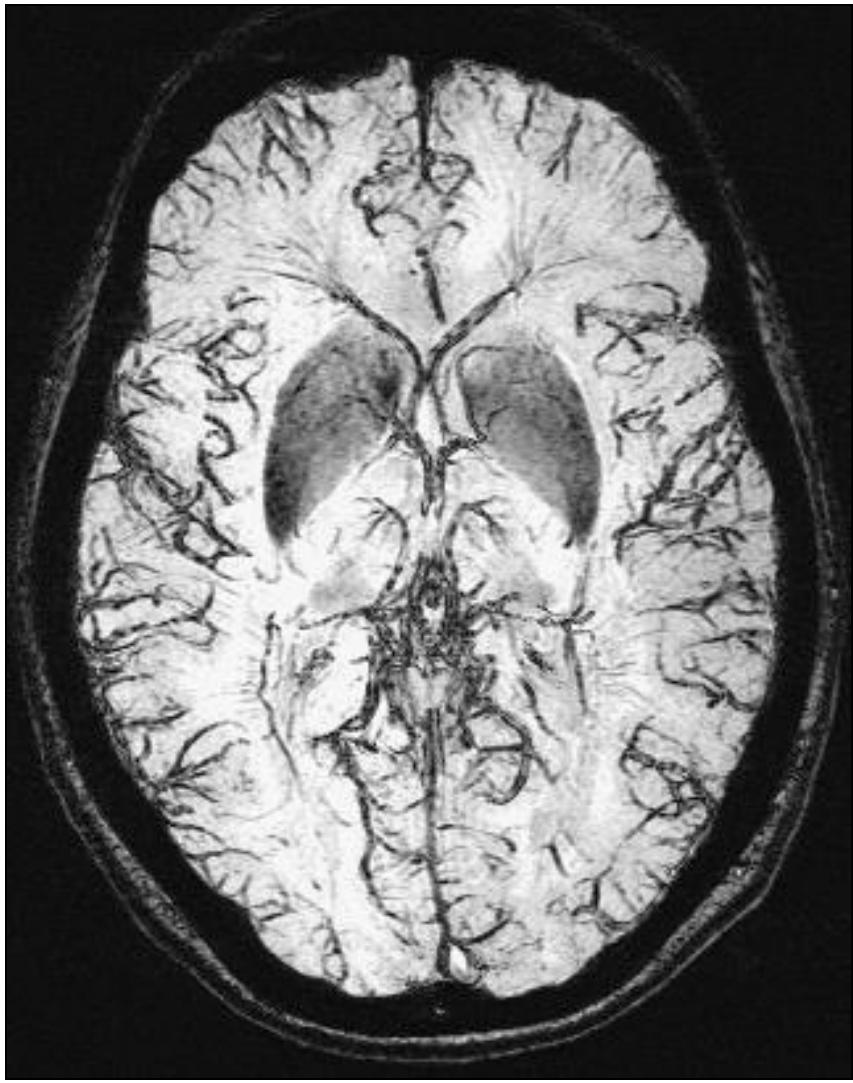


GK Aguirre et al, (2002) NeuroImage 15 (3): 488-500

Perfusion vs. BOLD: Low Task Frequency



BOLD effect to highlight veins: 3 Tesla



Bove-Bettis, et al (2004), SMRT

Technology

Coil arrays
High field strength
High resolution
Novel functional contrast

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Multi-modal integration
Pattern classification
Real time feedback
Task design

Fluctuations
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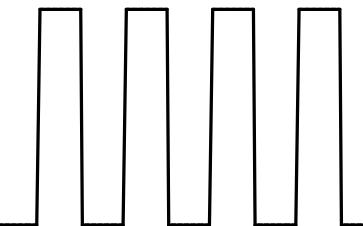
Interpretation

Applications

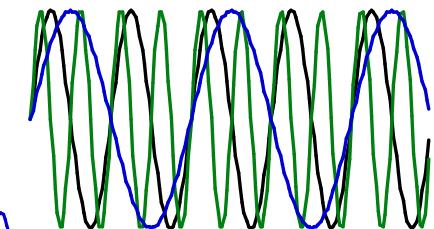
Methodology

Neuronal Activation Input Strategies

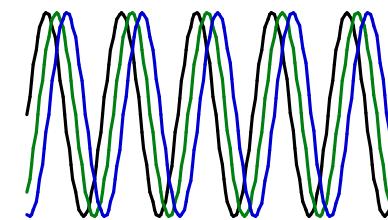
1. Block Design



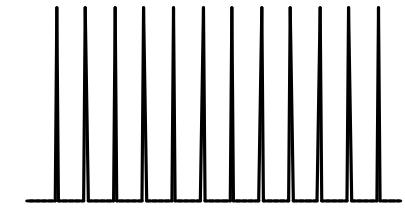
2. Frequency Encoding



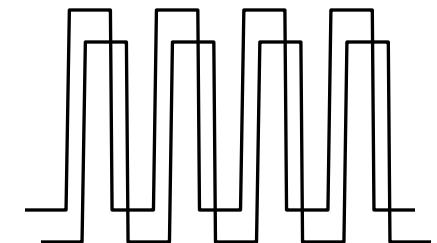
3. Phase Encoding



4. Event-Related



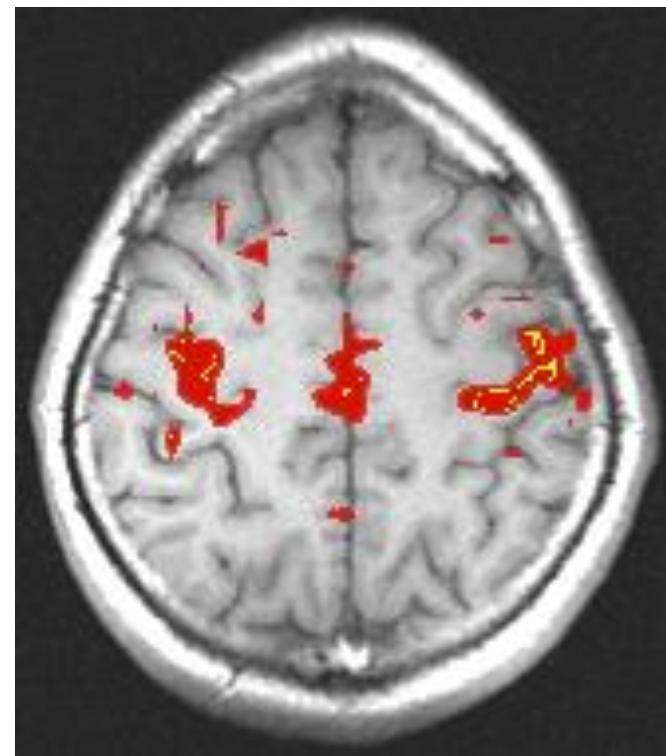
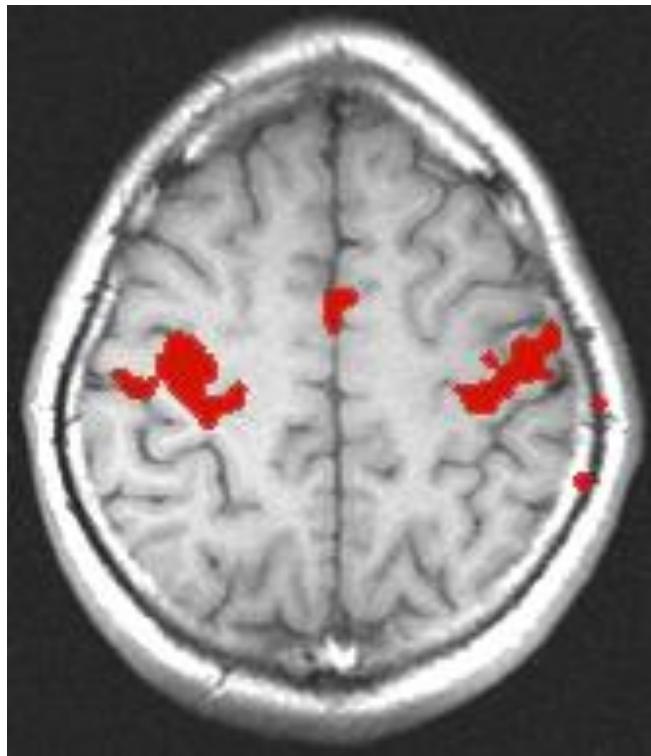
5. Orthogonal Block Design



6. Free Behavior Design.

Methodology

Resting State Correlations

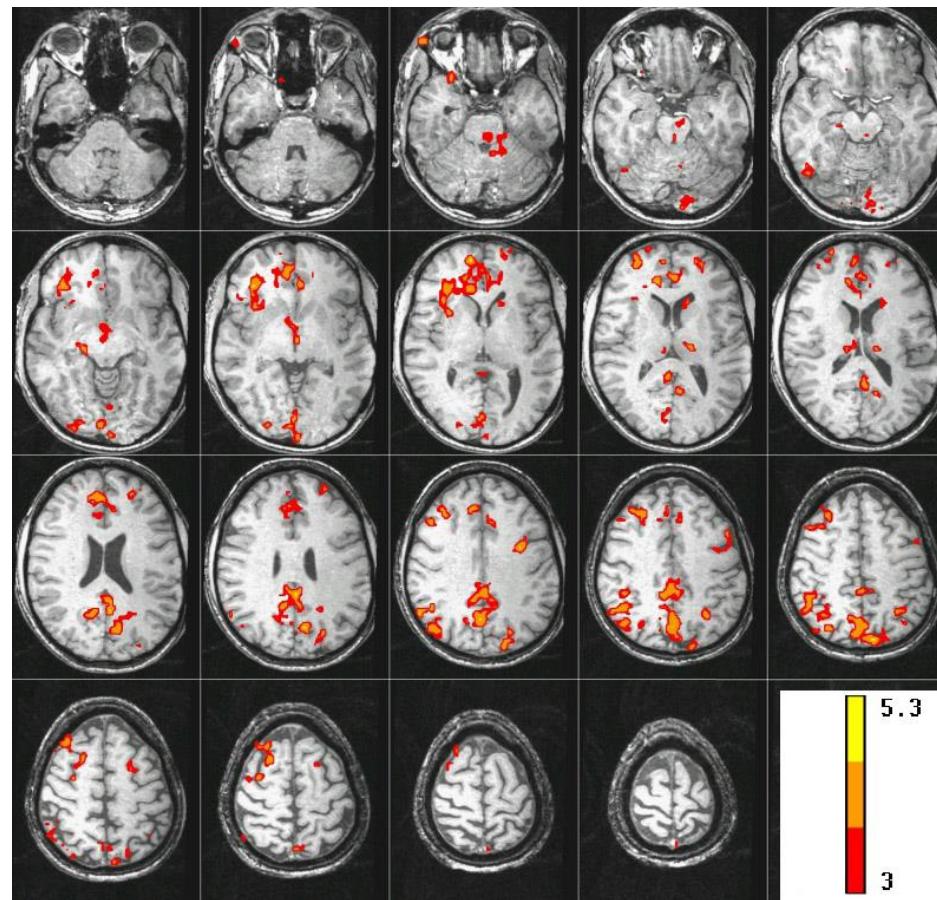


Activation:
correlation with reference function

Rest:
seed voxel in motor cortex

Methodology

BOLD correlated with SCR during "Rest"

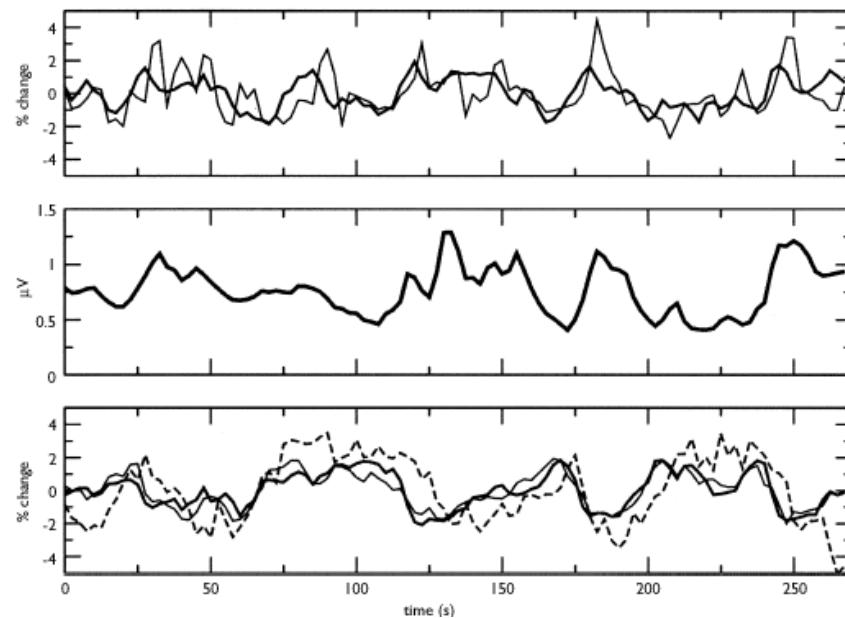


J. C. Patterson II, L. G. Ungerleider, and P. A
Bandettini, *NeuroImage* 17: 1787-1806, (2002).

Methodology

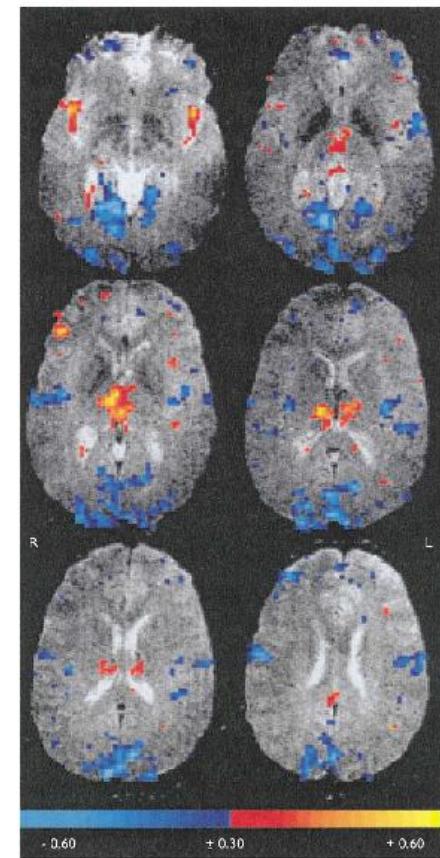
BOLD correlated with 10 Hz power during "Rest"

Positive
10 Hz power



Negative

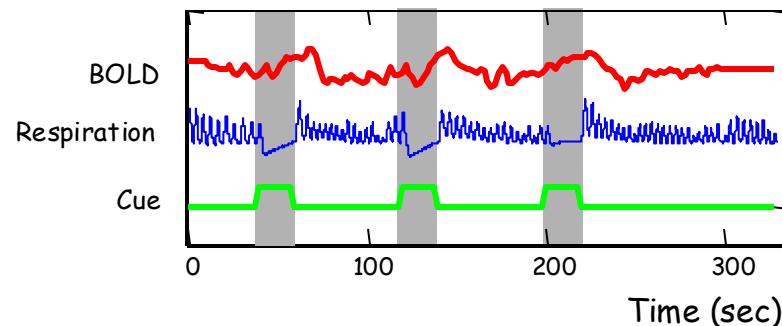
Goldman, et al (2002), Neuroreport



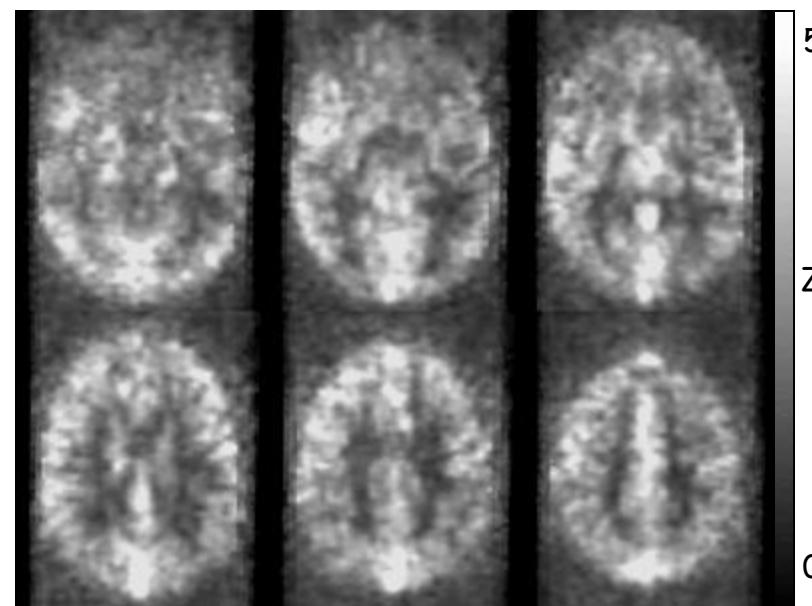
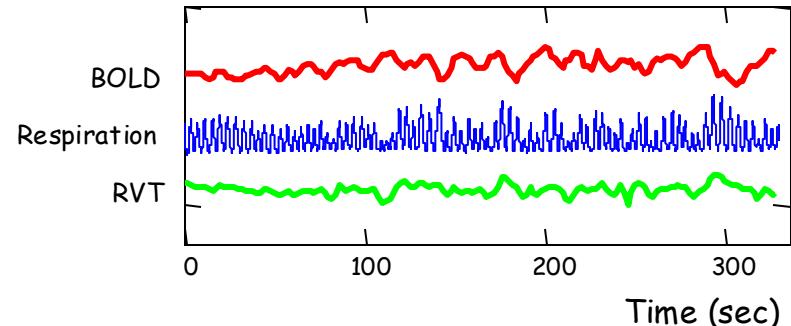
Methodology

Respiration induced signal changes

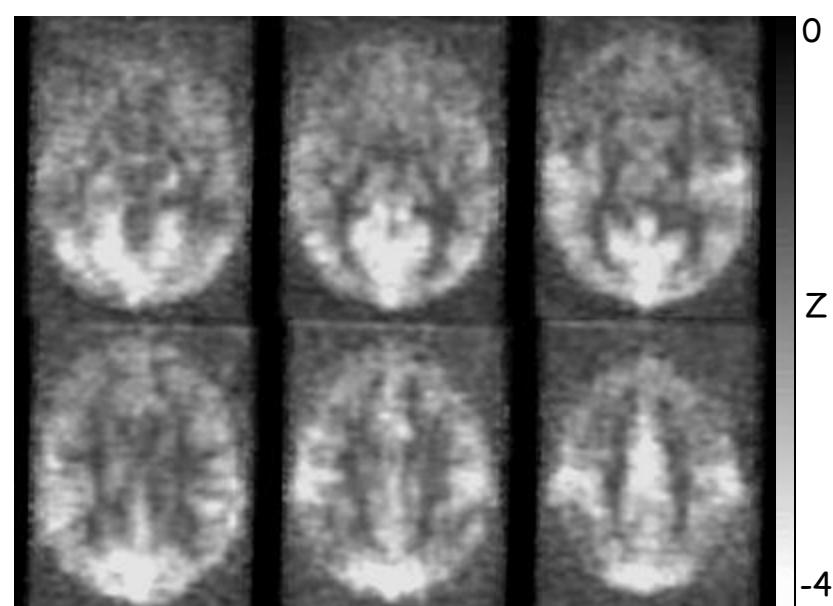
Breath-holding



Rest



(N=7)



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



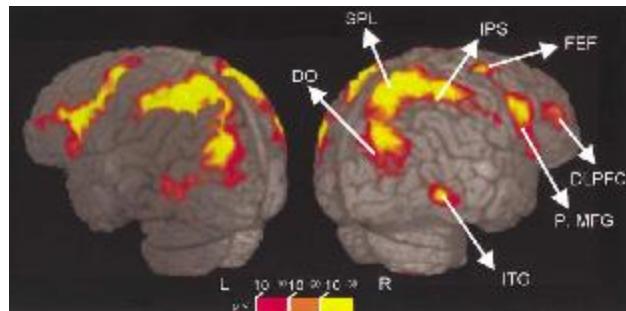
Mapping \leftrightarrow "Reading"

Methodology

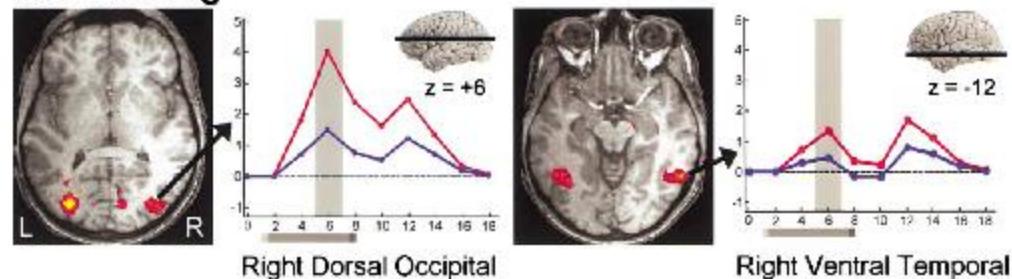
Neuron, Vol. 35, 975–987, August 29, 2002, Copyright ©2002 by Cell Press

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

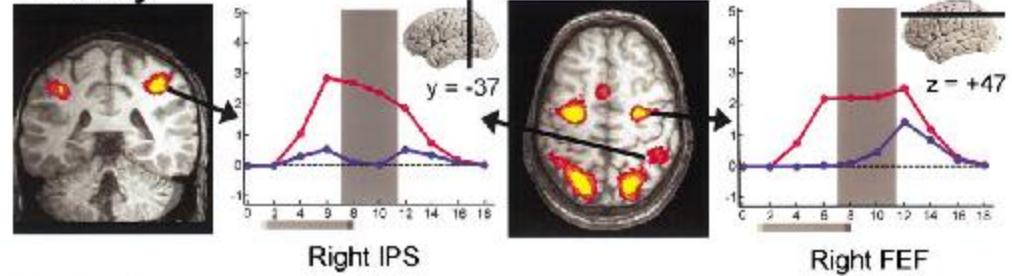
Luiz Pessoa,¹ Eva Gutierrez, Peter A. Bandettini,
and Leslie G. Ungerleider
Laboratory of Brain and Cognition
National Institute of Mental Health
National Institutes of Health
Bethesda, Maryland 20892



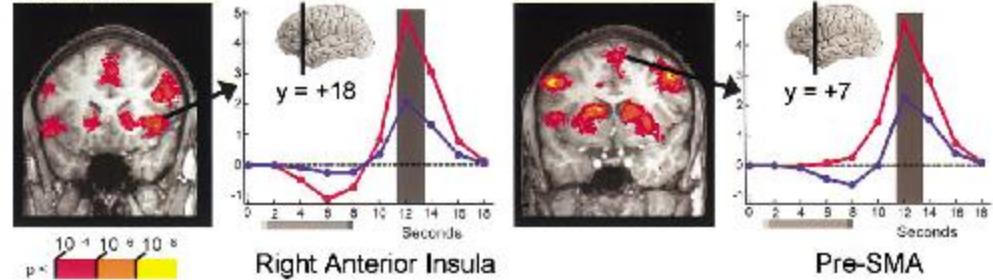
A. Encoding



B. Delay



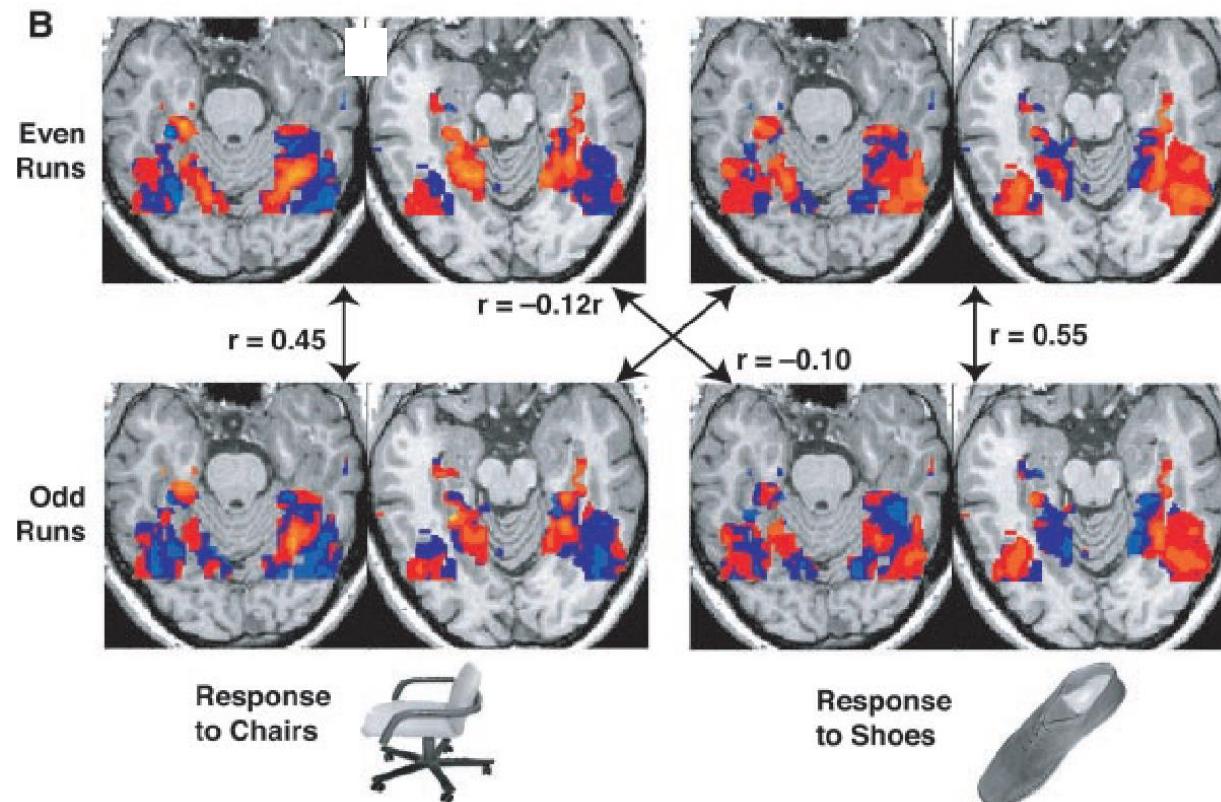
C. Test



Methodology

Ventral temporal category representations

Object categories are associated with distributed representations in ventral temporal cortex



Haxby et al. 2001

Methodology

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

David D. Cox^{a,b,*} and Robert L. Savoy^{a,b,c}

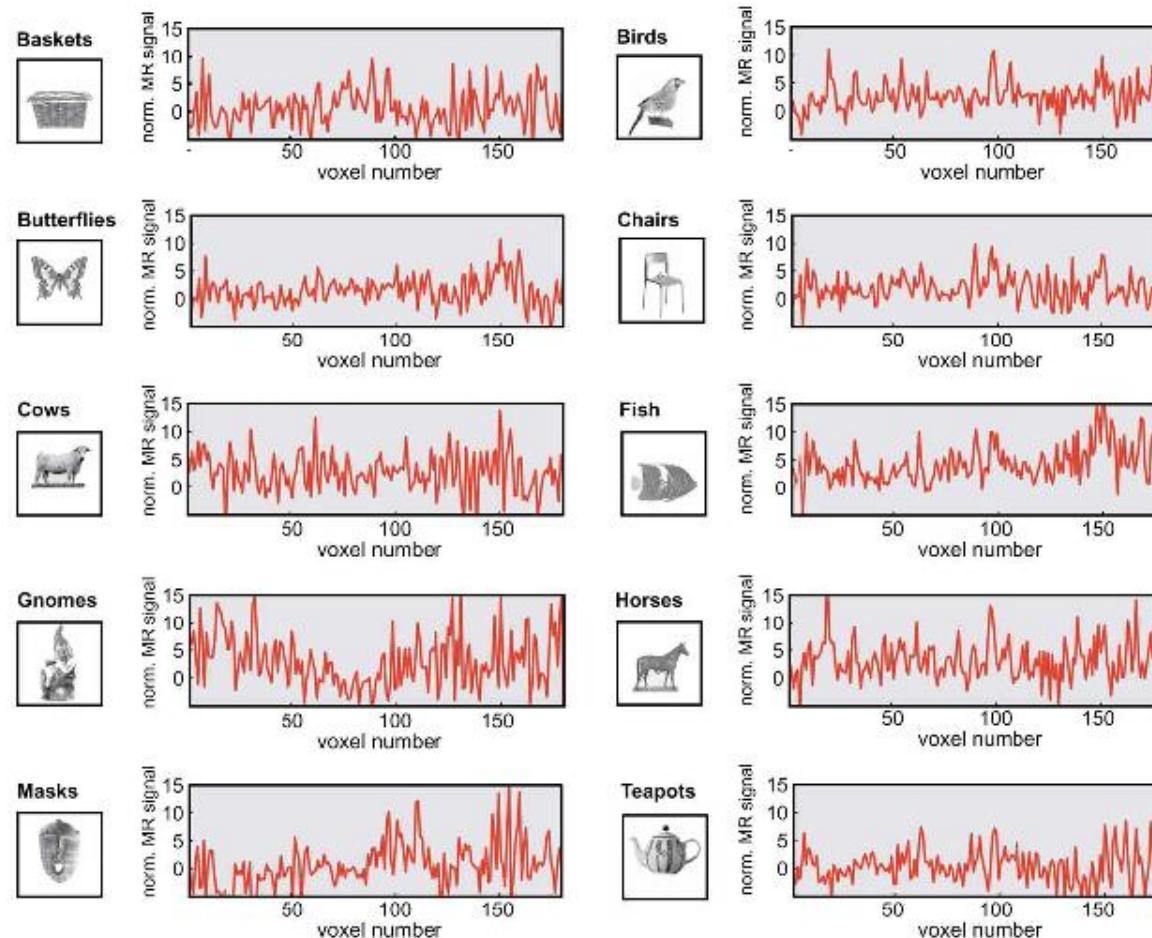
^a Rowland Institute for Science, Cambridge, MA 02142, USA

^b Athinoula A. Martinos Center for Structural and Functional Biomedical Imaging, Charlestown, MA 02129, USA

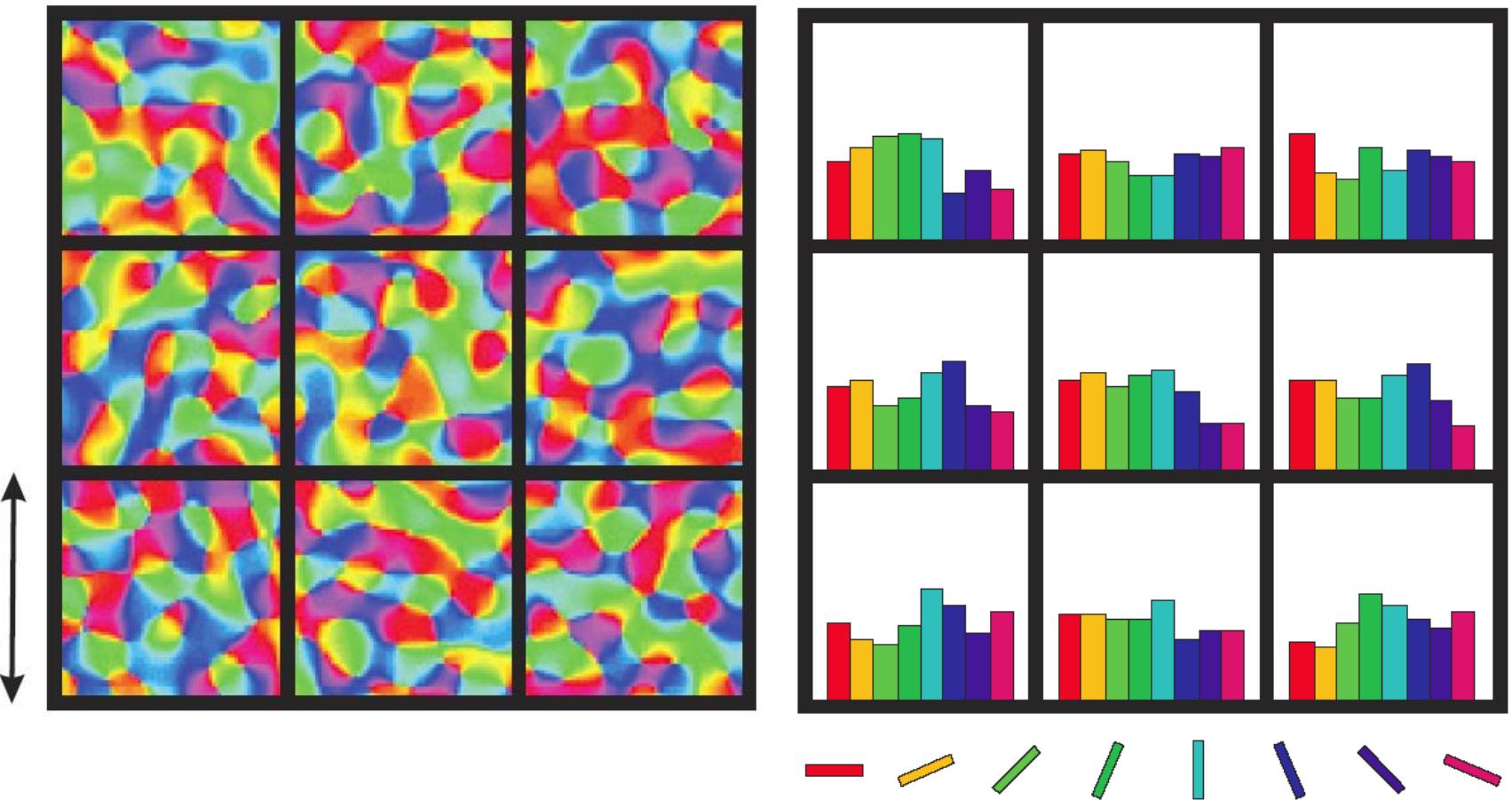
^c HyperVision, Inc., P.O. Box 158, Lexington, MA 02420, USA

Received 15 July 2002; accepted 10 December 2002

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



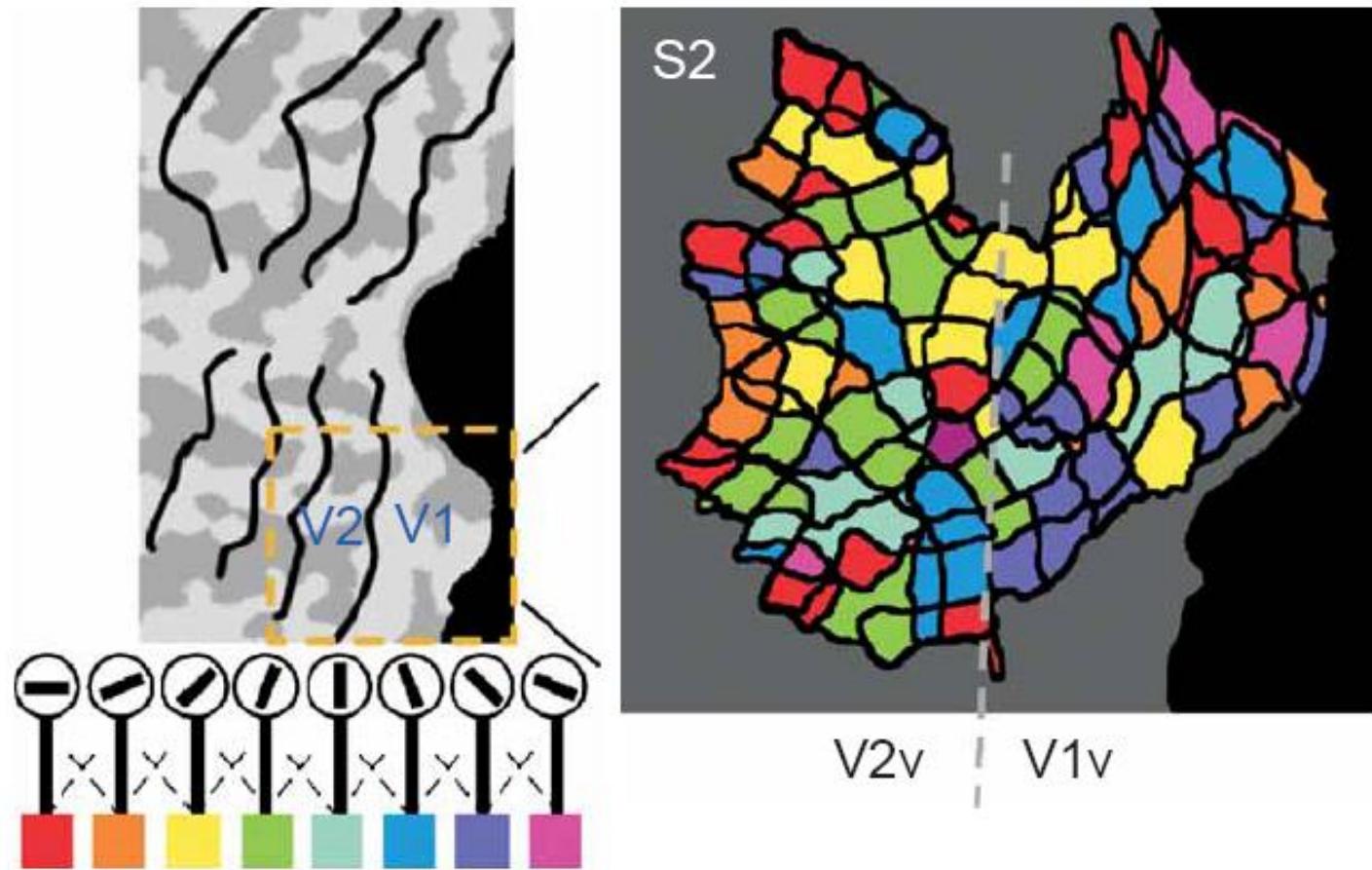
Methodology



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

Methodology

Lower spatial frequency clumping



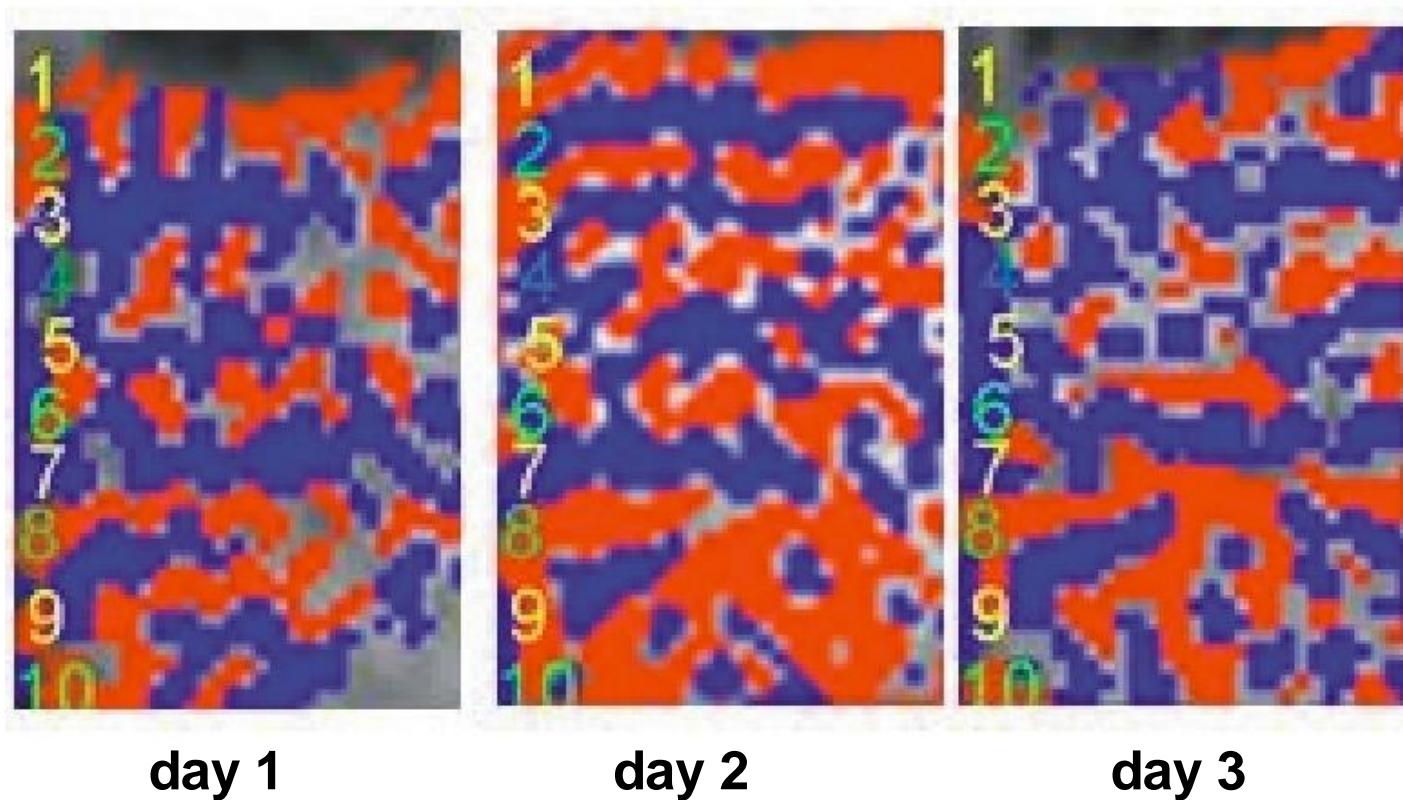
Kamitani & Tong (2005)

Methodology

What to do with high resolution data?

HSE-BOLD demonstration of ocular dominance columns

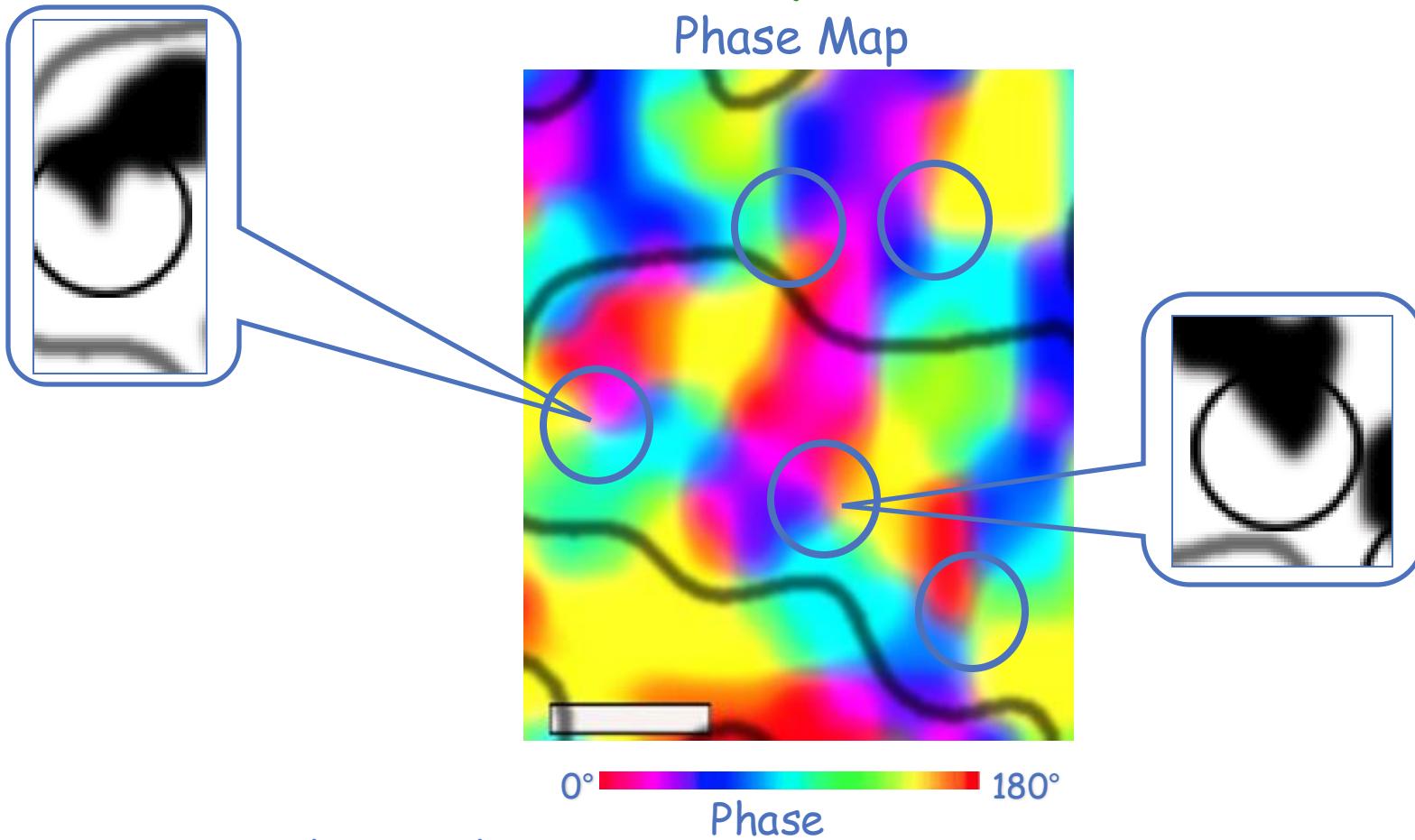
human, 7T, $0.5 \times 0.5 \times 3 \text{ mm}^3$



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

Methodology

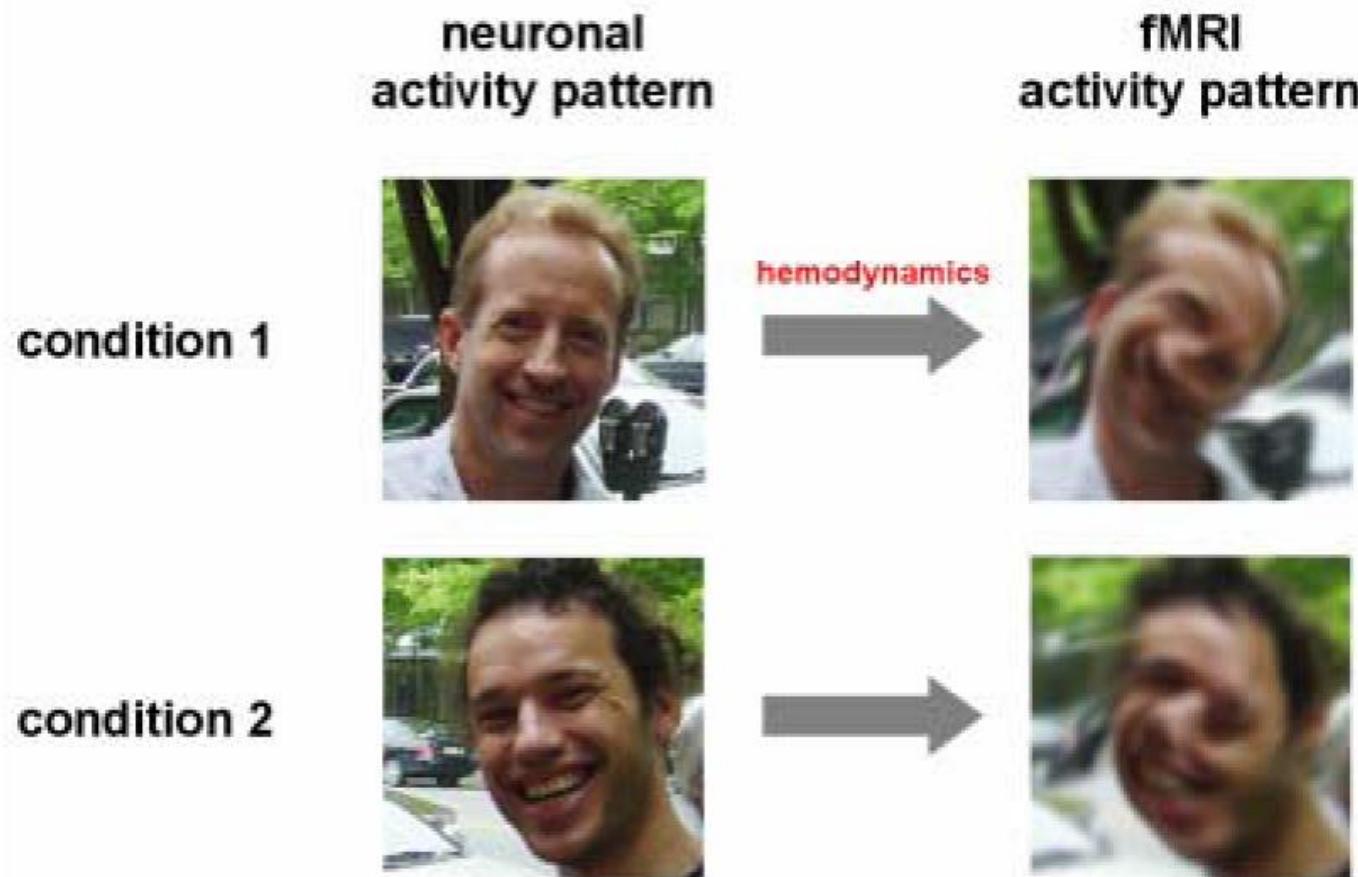
Orientation Columns in Human V1 as Revealed by fMRI at 7T



Yacoub, Ugurbil & Harel
University of Minnesota / CMRR
HBM 2006: Thursday, June 15, 2006 at 9:30

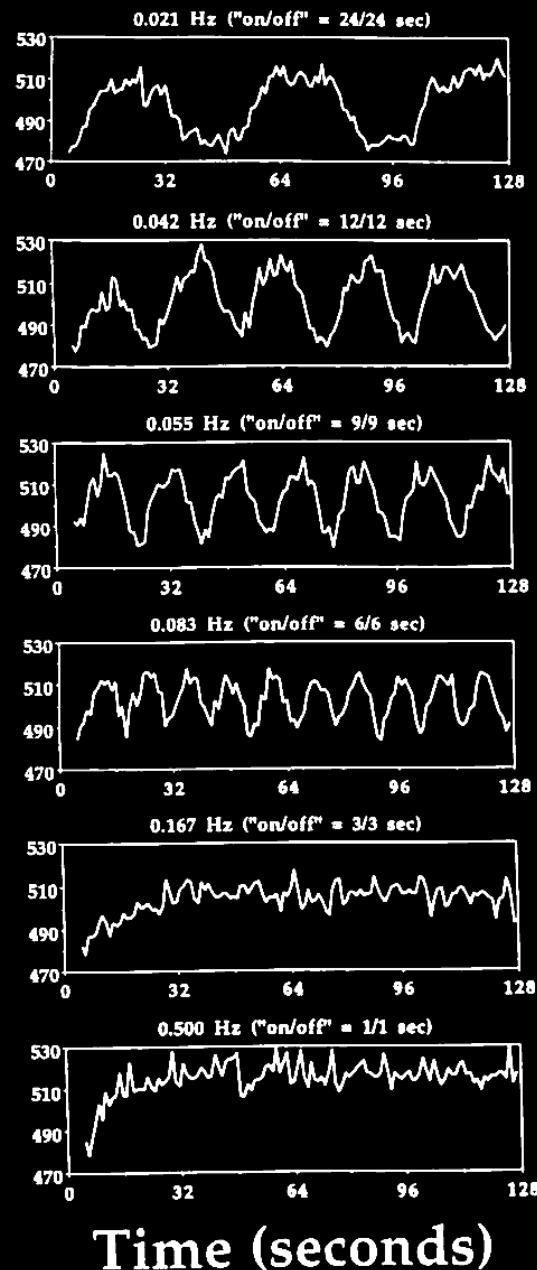
Scalebar = 0.5 mm

Methodology

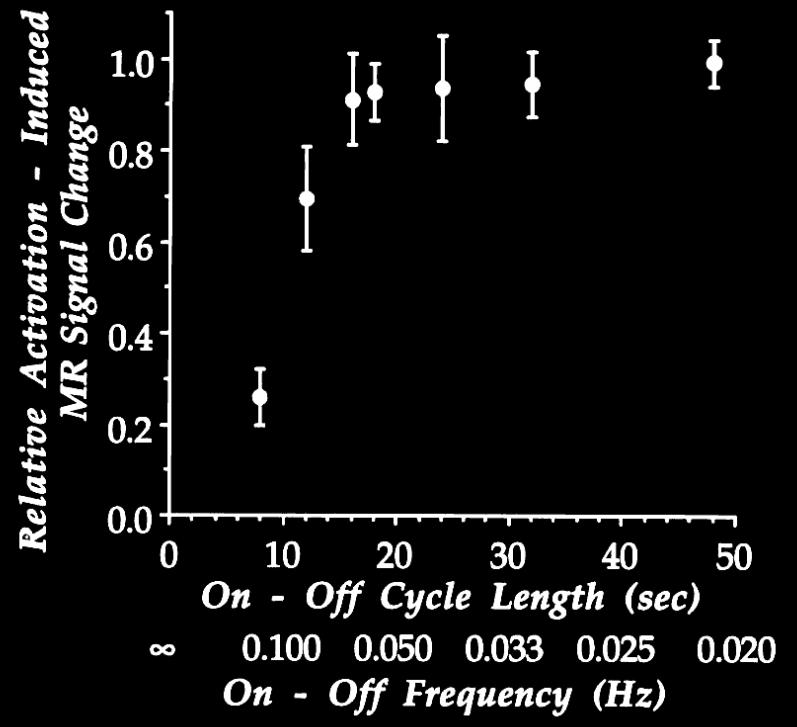


Methodology

MR Signal



Temporal Resolution

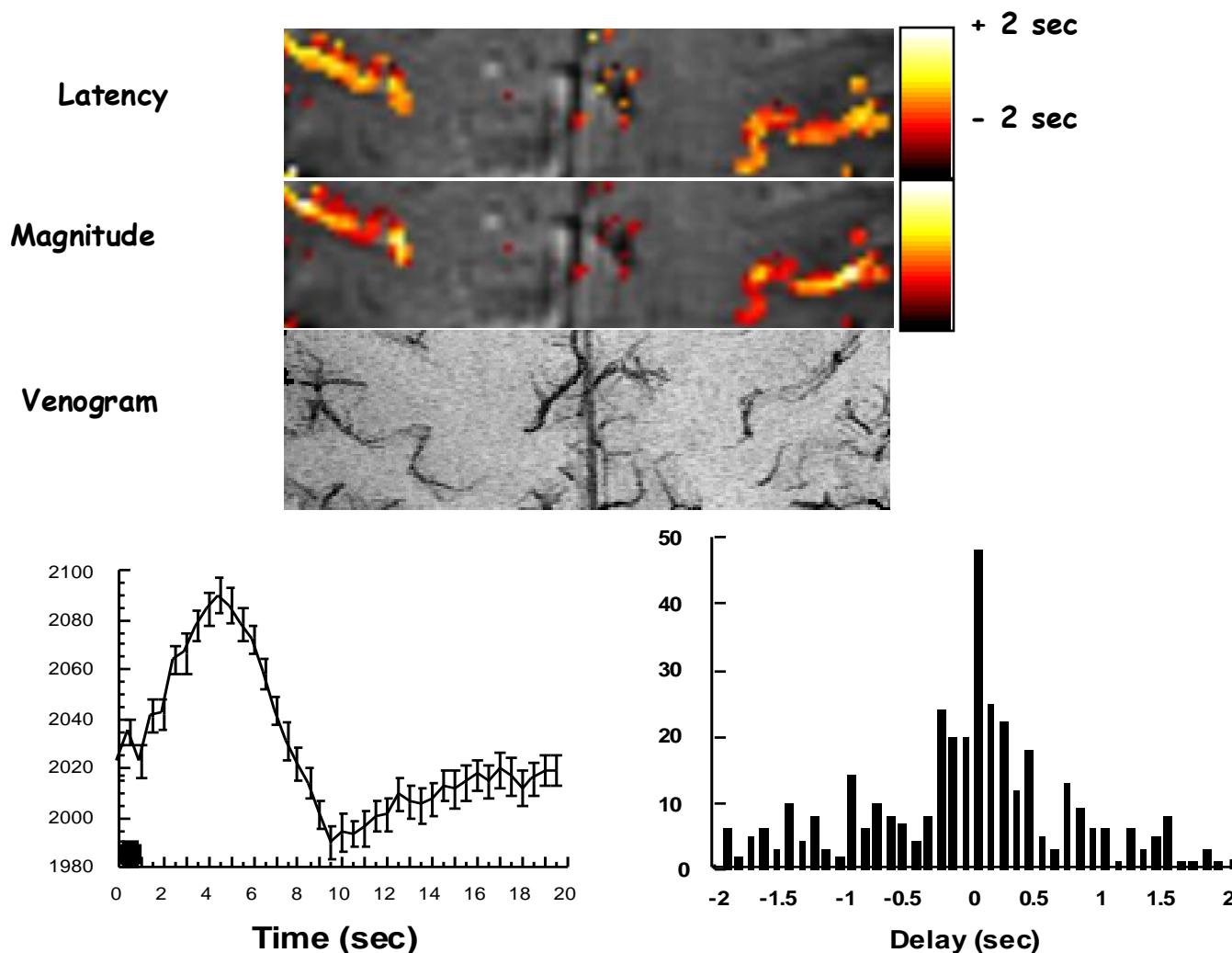


P. A. Bandettini, Functional MRI temporal resolution in "Functional MRI" (C. Moonen, and P. Bandettini., Eds.), p. 205-220, Springer - Verlag., 1999.

Methodology

Temporal Resolution

Latency Variation...



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

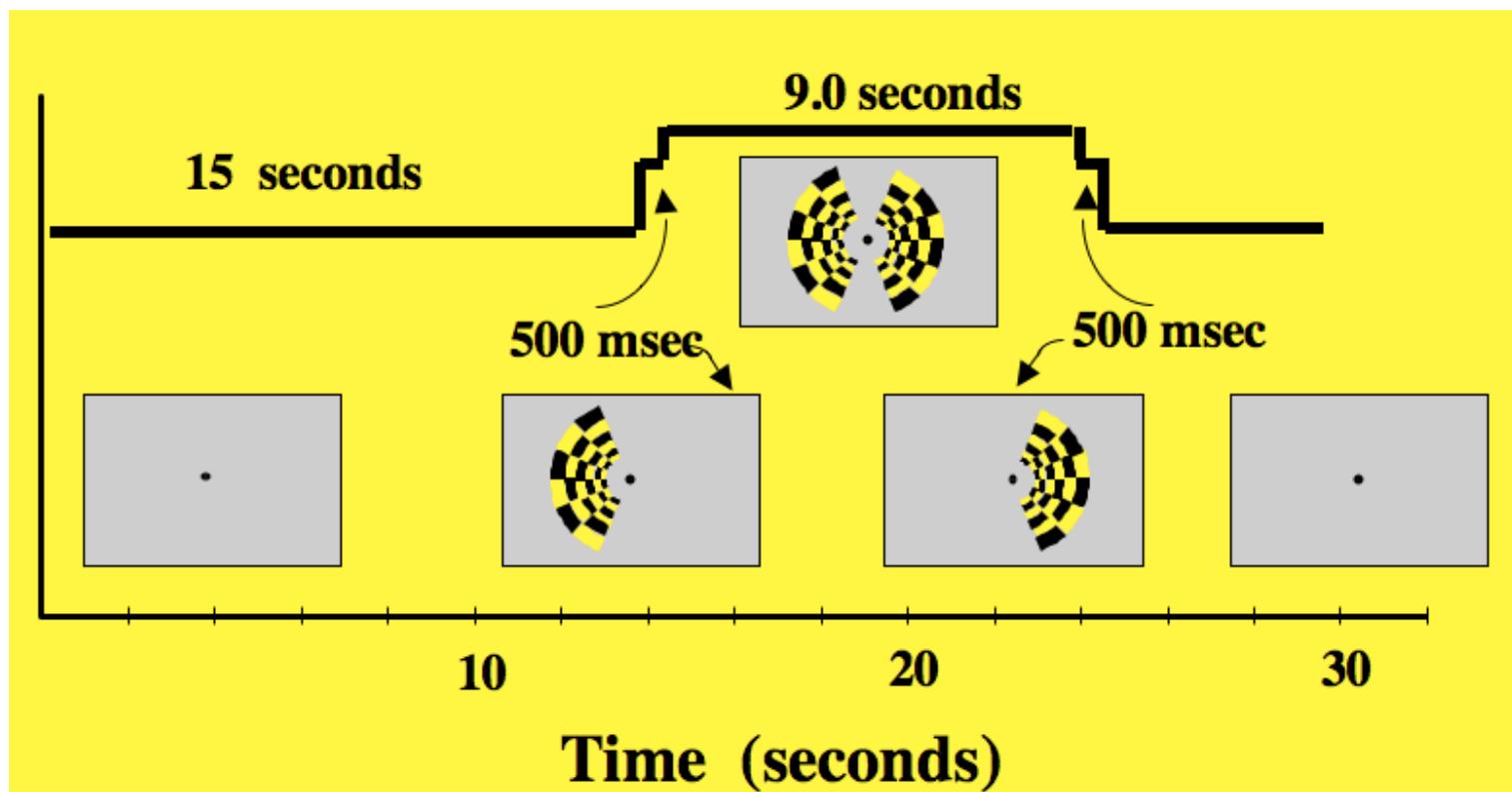
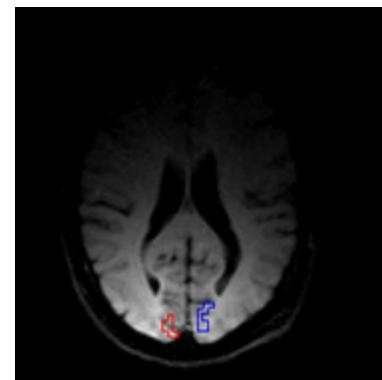
Methodology

Temporal Resolution

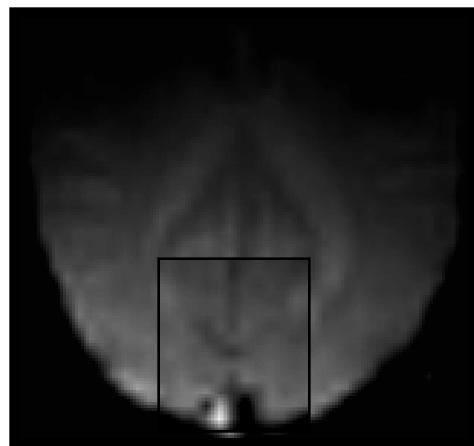
Hemi-Field Experiment

Right
Hemisphere

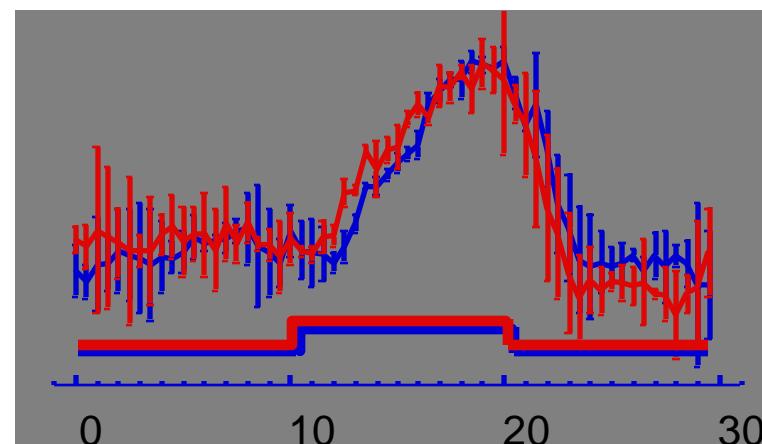
Left
Hemisphere



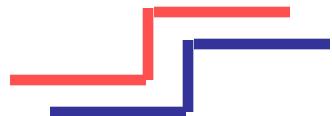
Methodology



Temporal Resolution



500 ms
||

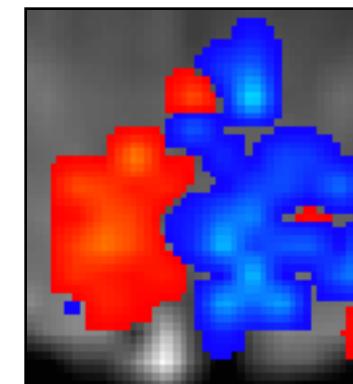
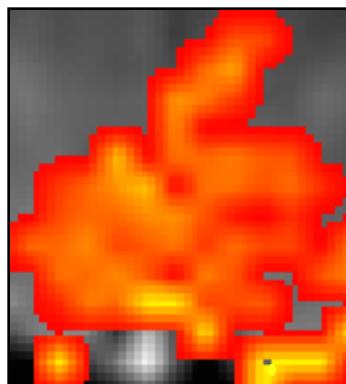
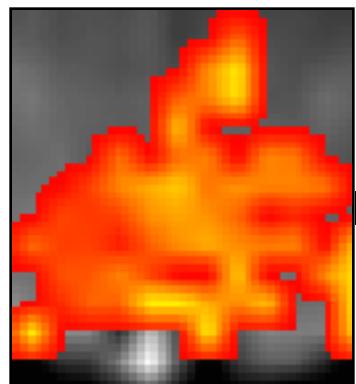


500 ms
||



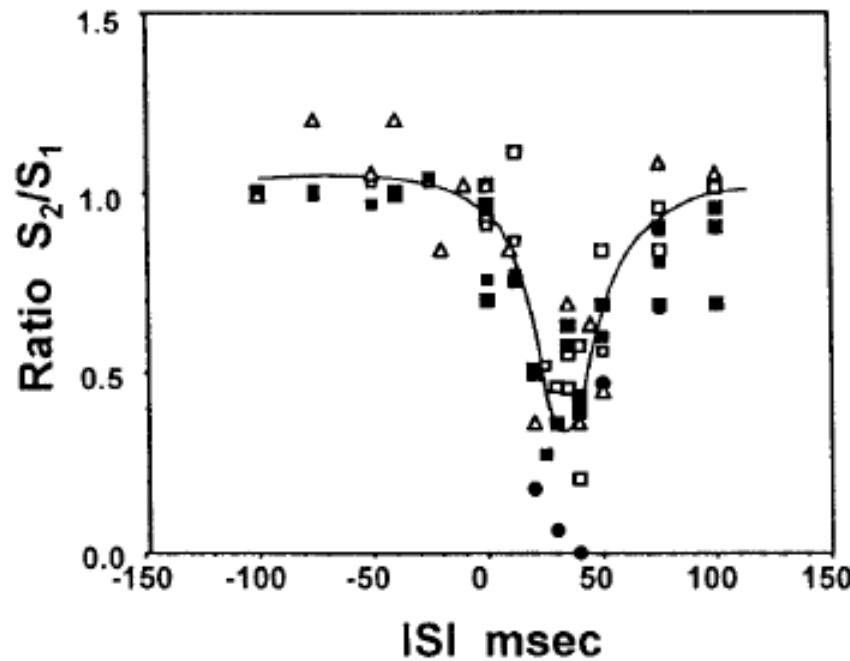
Right Hemifield

Left Hemifield



An approach to probe some neural systems interaction by functional MRI at neural time scale down to milliseconds

Seiji Ogawa^{†‡}, Tso-Ming Lee[†], Ray Stepnoski[†], Wei Chen[§], Xiao-Hong Zhu[§], and Kamil Ugurbil[§]



Technology

Coil arrays
High field strength
High resolution
Novel functional contrast

Methodology

Functional Connectivity Assessment
Multi-modal integration
Pattern classification
Real time feedback
Task design

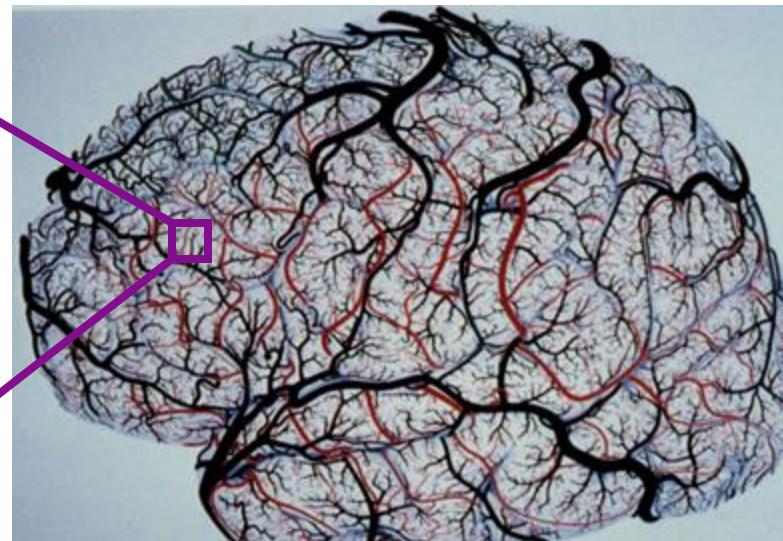
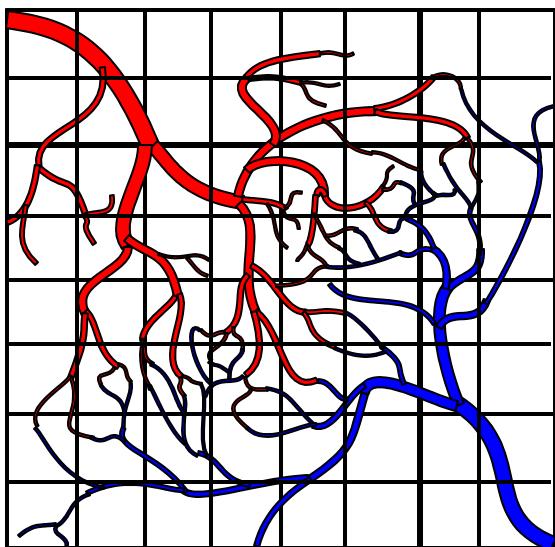
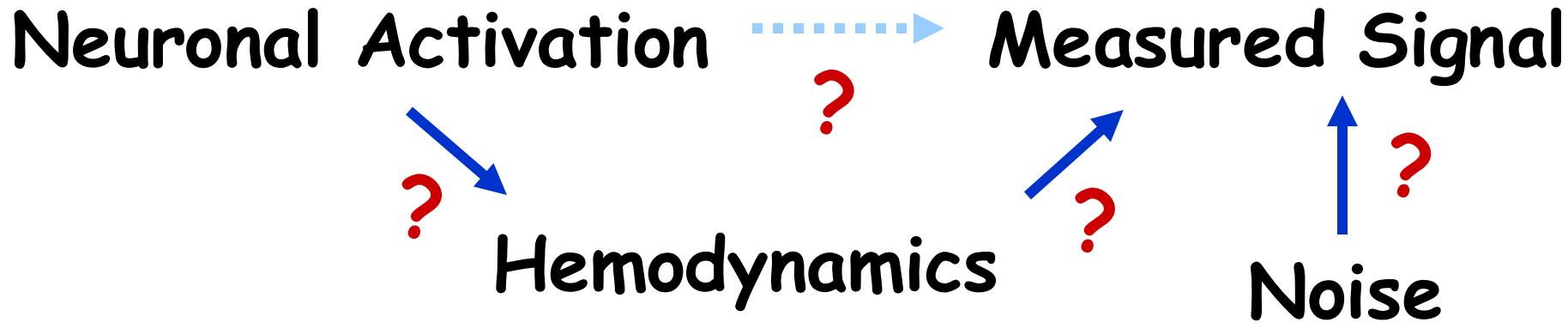
Fluctuations
Dynamics
Cross - modal comparison

Basic Neuroscience
Behavior correlation/prediction
Pathology assessment

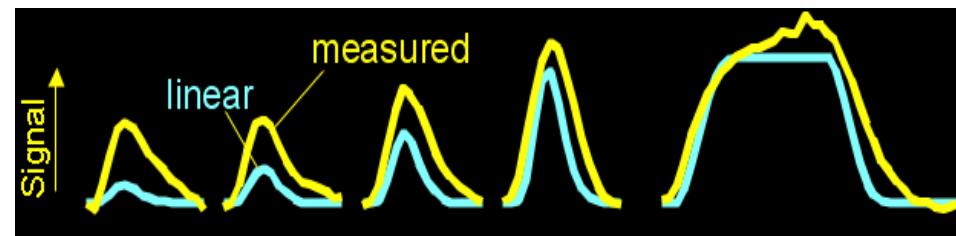
Interpretation

Applications

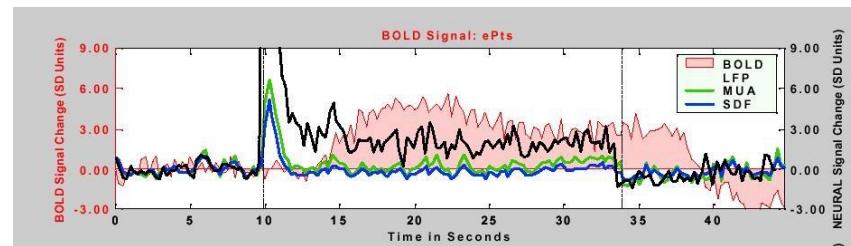
Interpretation



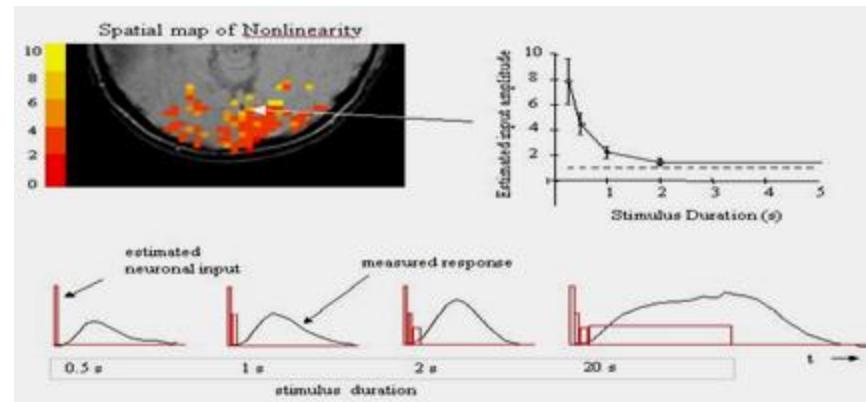
Interpretation



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



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

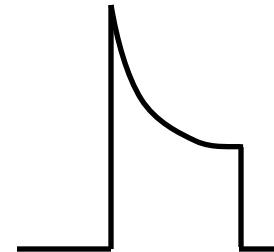
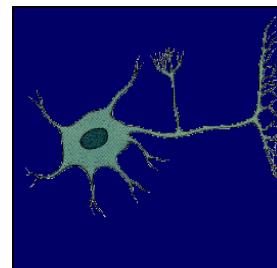


P. A. Bandettini et al, (2001) Nature Neuroscience, 4: 864-866.

Interpretation

Sources of this Nonlinearity

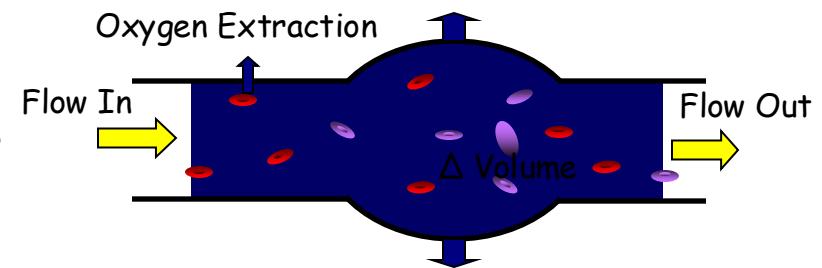
- Neuronal



- Hemodynamic

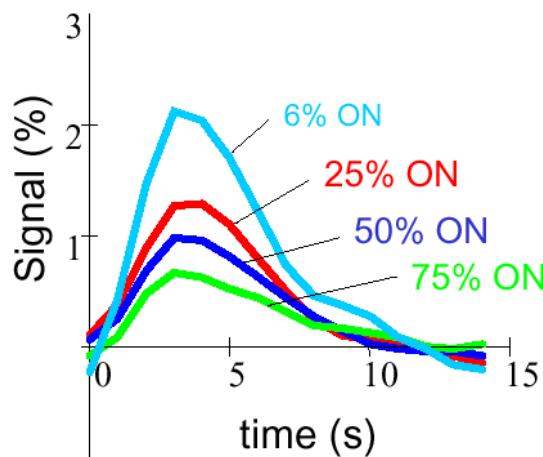
- Oxygen extraction

- Blood volume dynamics

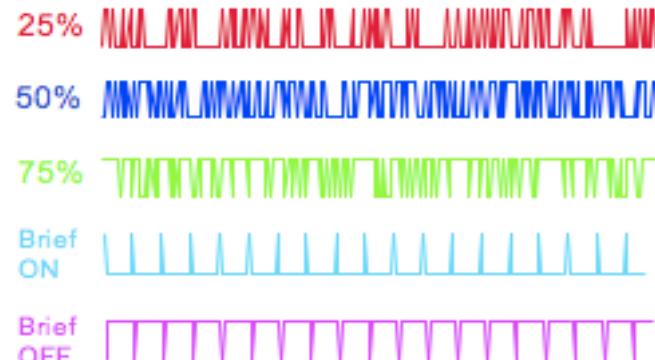
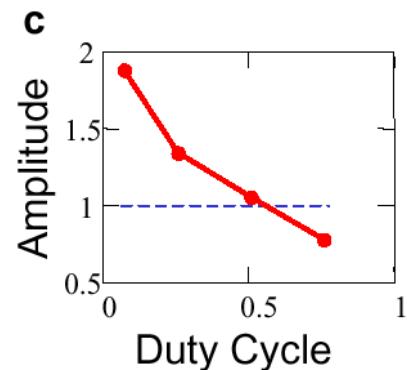
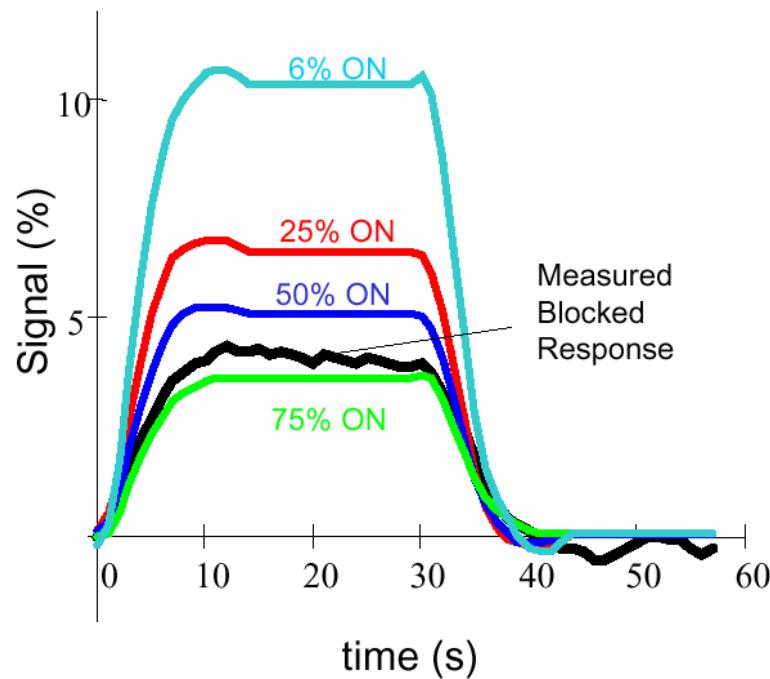


Interpretation Duty Cycle Effects

a Measured Event-related Responses



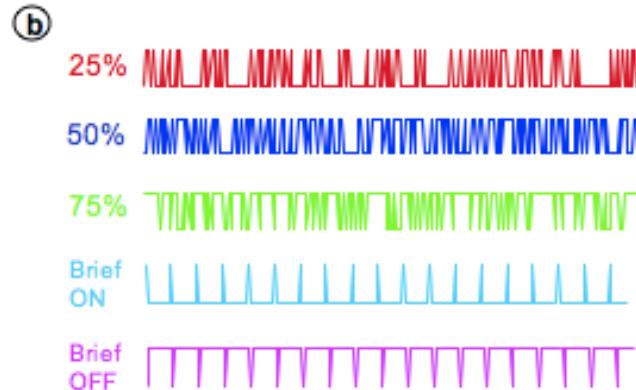
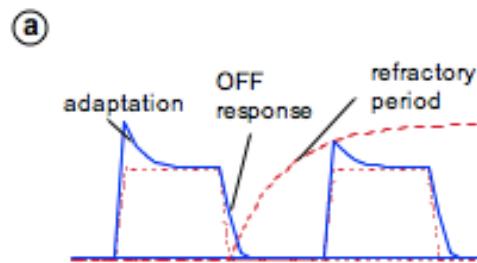
b Predicted Blocked Responses



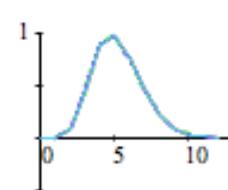
Interpretation

Linearity

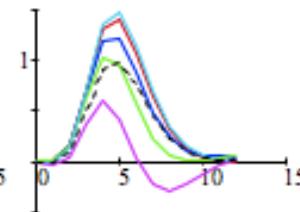
duty cycle effects



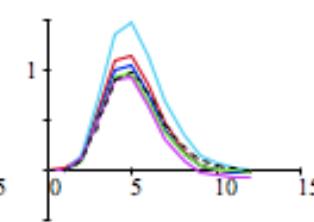
(c) Linear



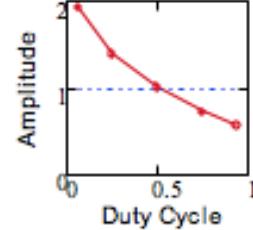
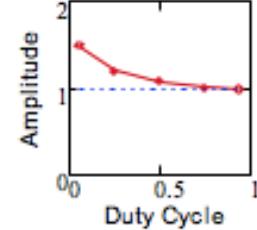
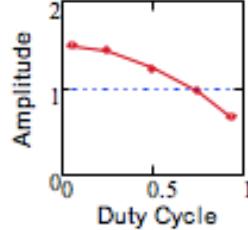
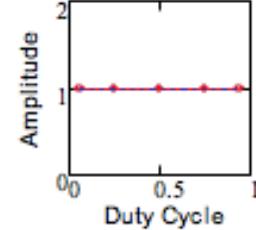
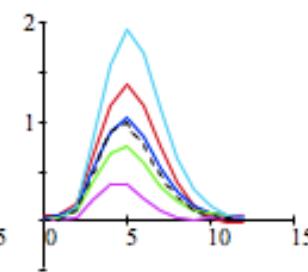
(d) Adaptation



(e) Adaptation + refractory



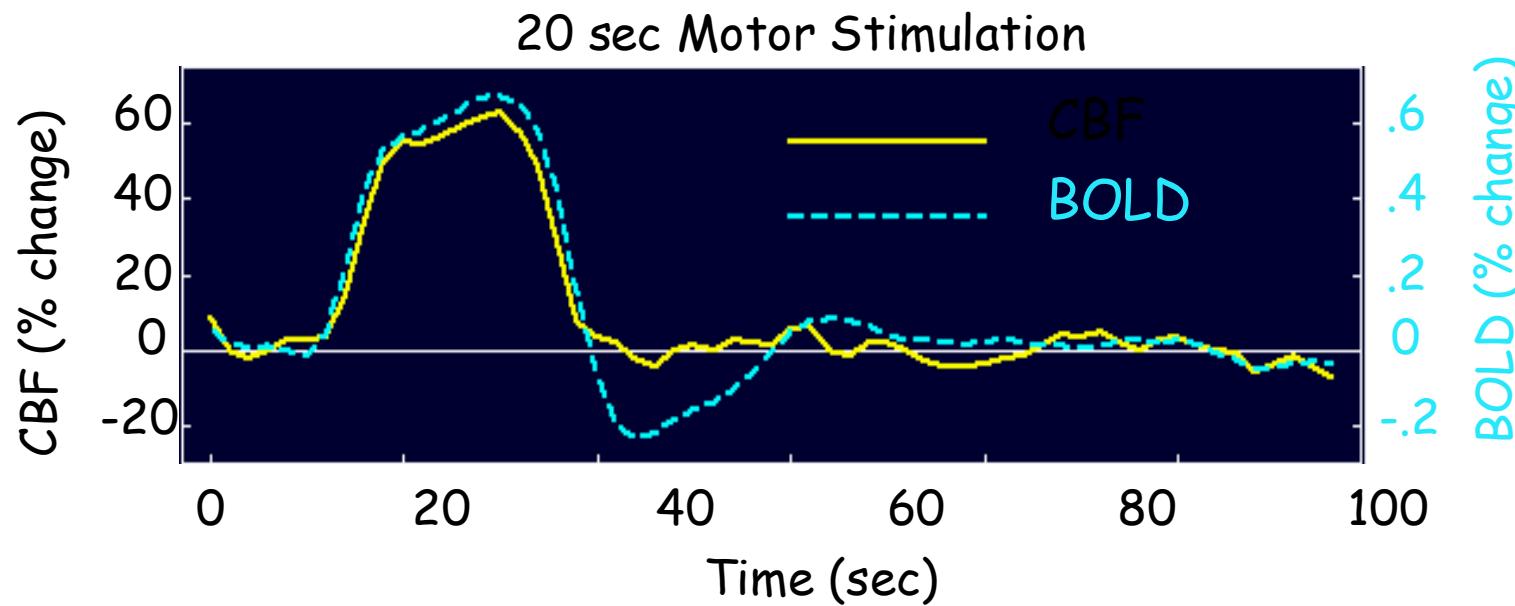
(f) Adaptation + refractory + OFF response



Interpretation

Post Undershoot

BOLD post-stimulus undershoot



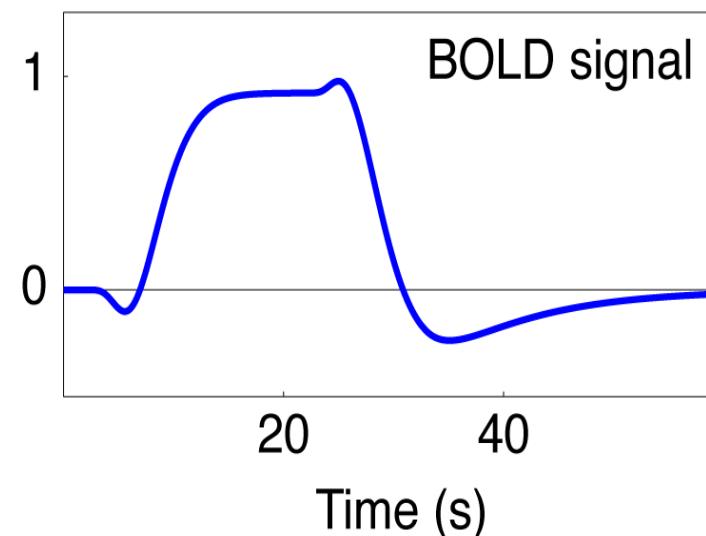
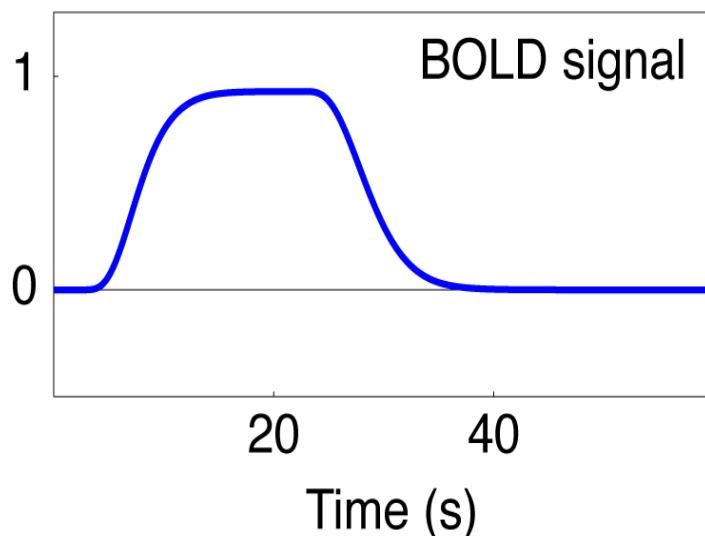
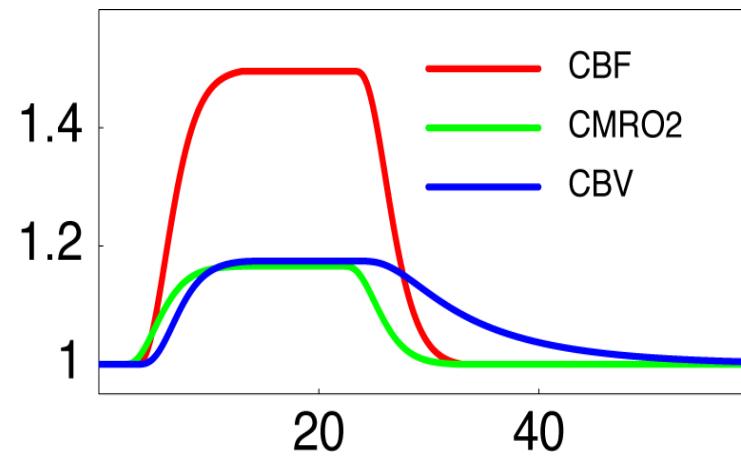
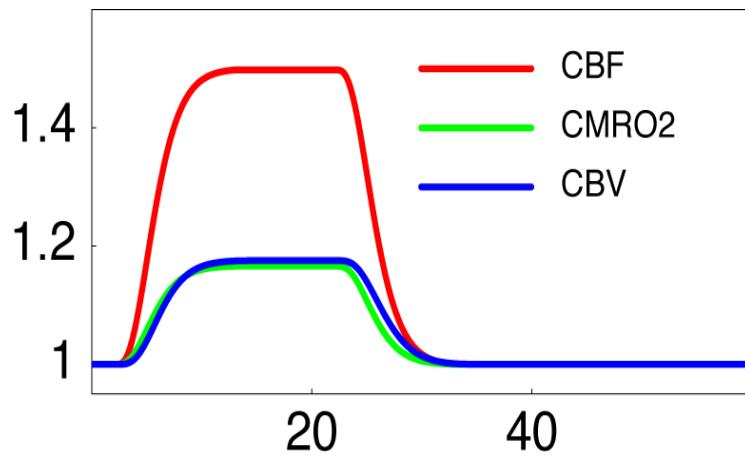
A BOLD undershoot without a CBF undershoot could be due to a slow return to baseline of either CBV or CMRO₂

Courtesy Rick Buxton

Interpretation

Post Undershoot

BOLD Signal Dynamics



Courtesy Rick Buxton

Technology

Coil arrays
High field strength
High resolution
Novel functional contrast

Methodology

Functional Connectivity Assessment
Multi-modal integration
Pattern classification
Real time feedback
Task design

Fluctuations
Dynamics
Cross - modal comparison

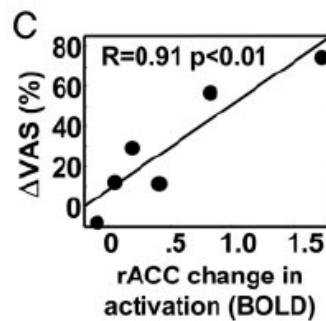
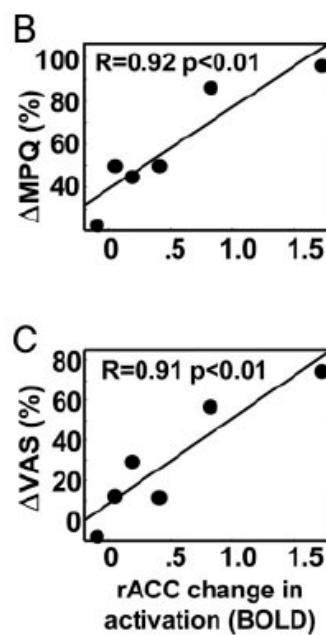
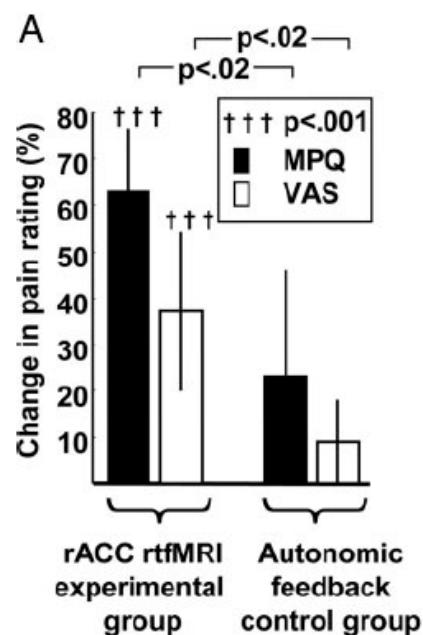
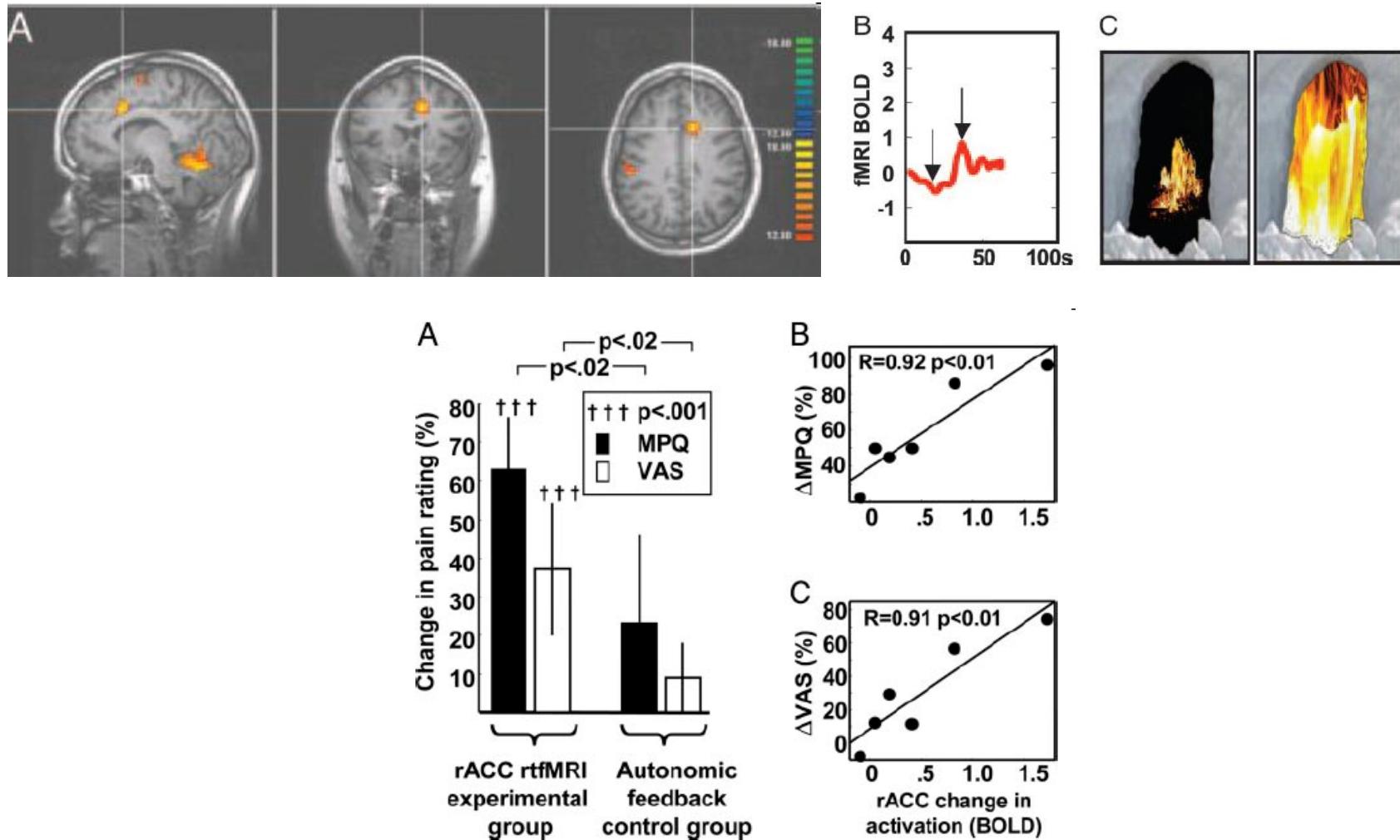
Basic Neuroscience
Behavior correlation/prediction
Pathology assessment

Interpretation

Applications

Applications

Real time fMRI feedback from Anterior Cingulate Cortex 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 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

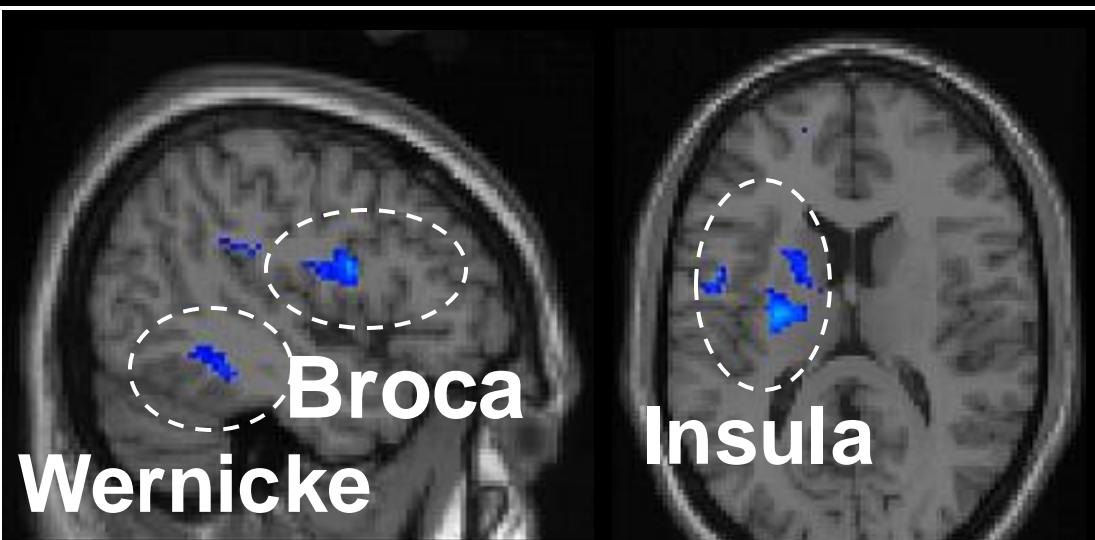
Middle row: Jon West, Kay Kuhns, Anthony Boemio, Peter Bandettini, Joey Dunsmoor, Doug Ruff, Kevin Murphy

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

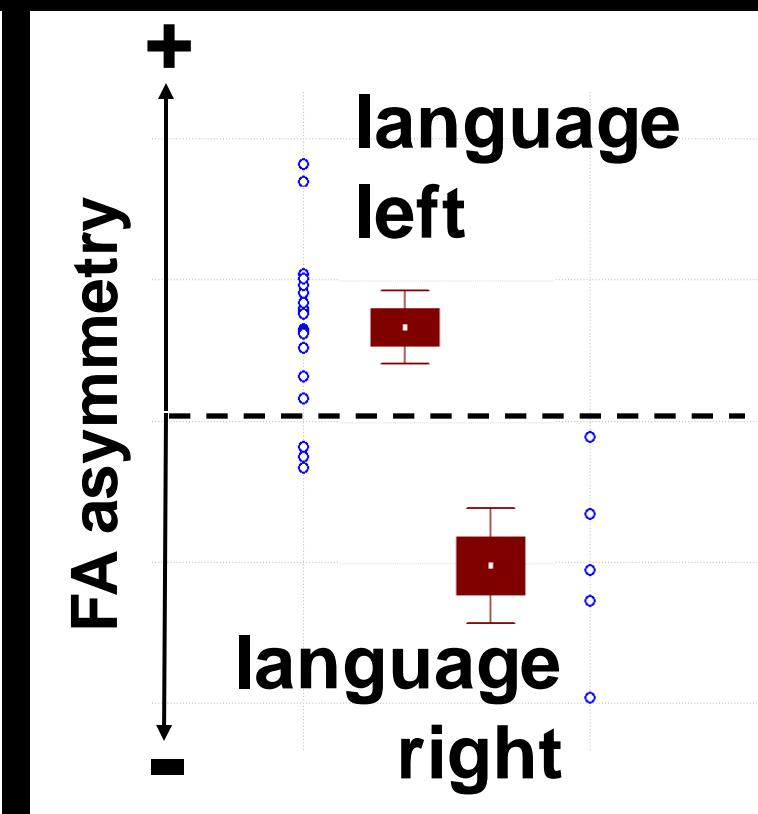
Anatomical correlates of right-hemispheric language processing: A DTI study

Monday – AM

S Mohammadi, A Jansen, W Schwindt, S Knecht, M Deppe
University of Münster, Germany



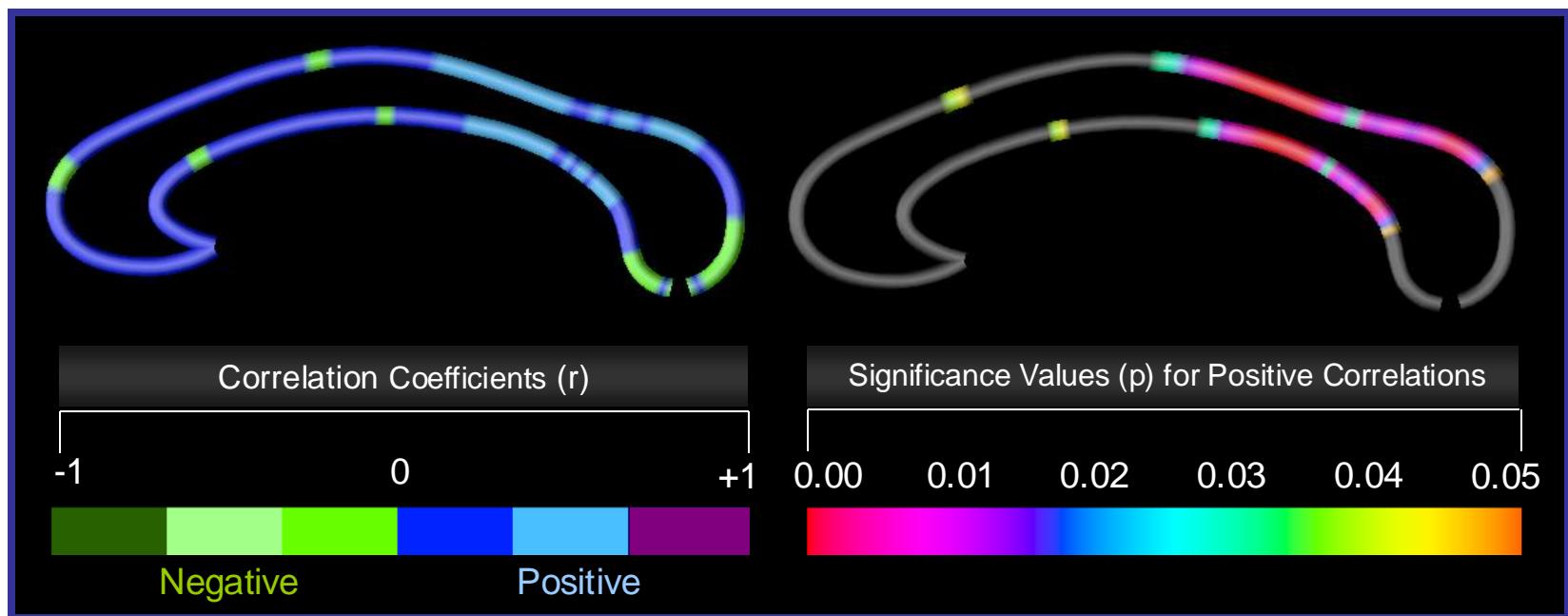
Blobs reflect FA differences between left and right hemispheric language dominant subjects



Results: language dominance was predicted by hemispheric FA asymmetry

Does callosal thickness correlate with intelligence?

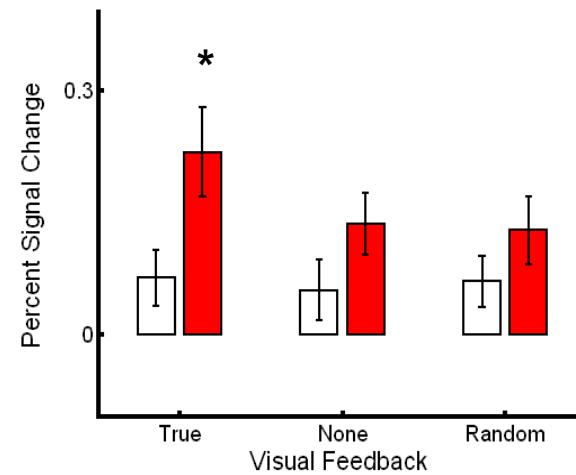
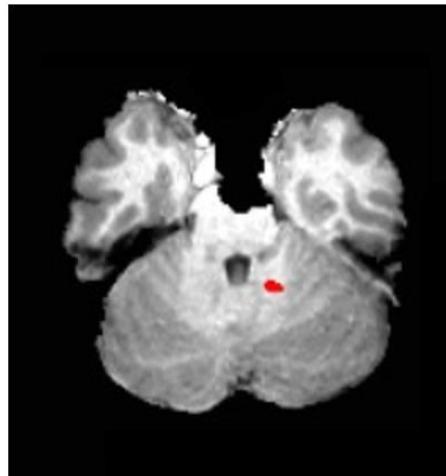
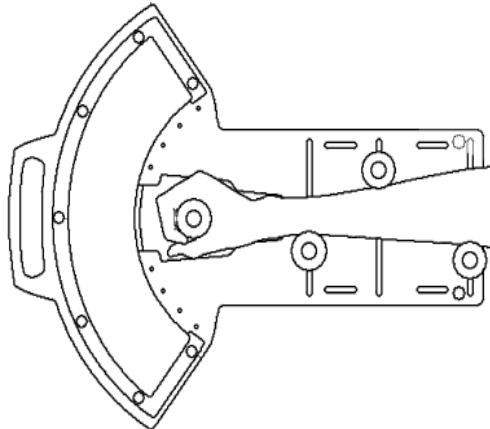
- (1) Intelligence and callosal thickness are correlated.
- (2) Only positive correlations are significant.
- (3) Positive correlations are most pronounced in the posterior half of the corpus callosum.



Cerebellar and posterior parietal involvement in the integration of visual and proprioceptive feedback during stabilization of the wrist

A.J. Suminski¹, S.M. Rao², and R.A. Scheidt¹

¹Marquette Univ., Milwaukee, WI; ²Medl College of Wisconsin, Milwaukee, WI



Activation in the **ipsilateral dentate nucleus** is **enhanced** when visual and proprioceptive feedback are **correlated** in time.

Reflective of its role in **integrating multiple sensory and feedforward estimates of limb state** thereby producing a unified limb state estimate that can be used to correct for movement errors.

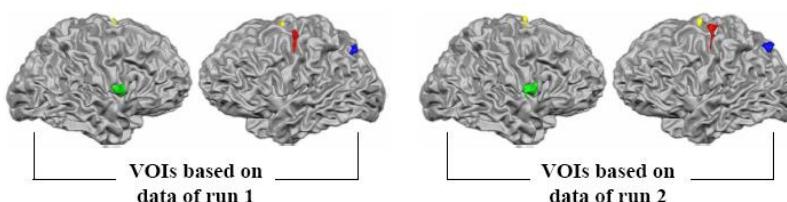
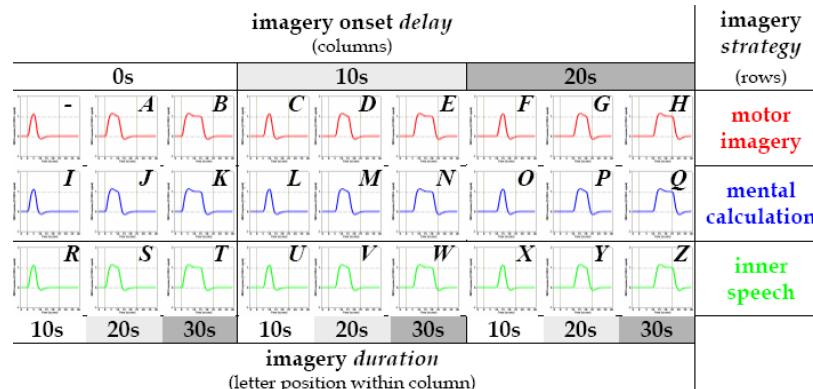


BOLD communication: When the brain speaks for itself



Bettina Sorger, Brigitte Dahmen, Joel Reithler, Rainer Goebel

Cognitive Neuroscience Department, Maastricht University, The Netherlands
Maastricht Brain Imaging Center (M-BIC), Maastricht University, The Netherlands



- parietal cortex
- dorsal premotor region
- yellow ● supplementary motor area
- green ● inferior frontal gyrus (Broca's area)

en-/decoder	guided letter encoding			letter	phrase encoding			letter
	paradigm	delay	duration		paradigm	delay	duration	
S1	.988	.981	.907	.889	.958	.986	.889	.840
S2	.938	.932	.870	.827	.784	.961	.804	.686
S3	.957	.981	.920	.883	.735	.931	.824	.608
R1	.966	.975	.932	.901	.869	.960	.869	.778
R2	.969	.957	.907	.877	.879	.990	.889	.788
R3	.944	.963	.858	.821	.800	.940	.798	.636
total	.961	.965	.899	.866	.852	.963	.852	.734

Fig. 3. Accuracy results of the guided letter and phrase encoding experiments (percentage of correct identification).

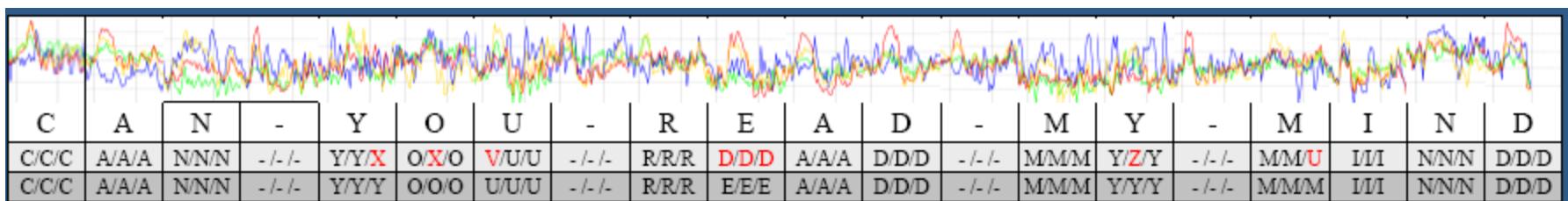


Fig. 4. Results of the phrase encoding experiment (subject S1).

The first row displays the single-trial time courses of the four VOIs generated by the subject while encoding the letters indicated in the second row. The third (light-grey) row displays letter decoding results obtained independently by three raters (R1/R2/R3) evaluating the time courses in randomized trial order (without word context information), whereas the fourth (dark-grey) row illustrates the results obtained by using the original trial order (providing word context information).

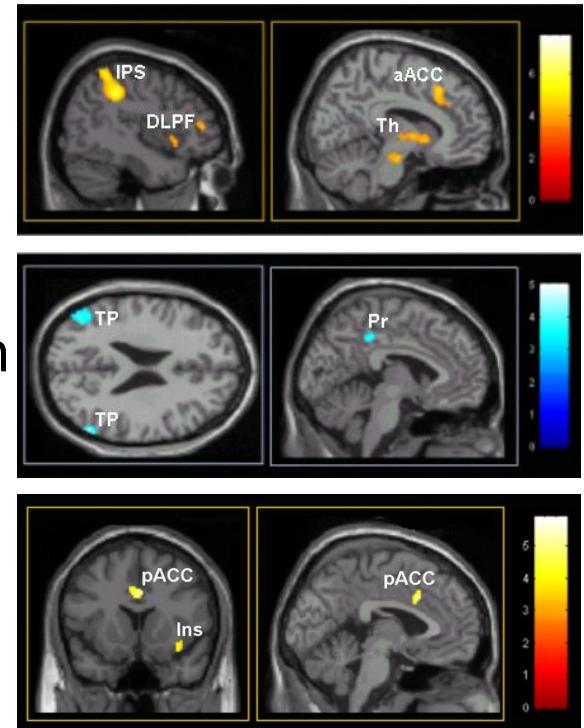
Baseline brain activity fluctuations predict somatosensory perception

M. Boly, E. Balteau, C. Schnakers, C. Degueldre, G. Moonen, A. Luxen, C. Phillips, P. Peigneux, P. Maquet, S. Laureys
Cyclotron Research Centre & Neurology Dept., University of Liège, Belgium

Poster #3 M-AM; Oral: "Cognition – Perception and Awareness" on Tuesday, June 12, 18:15.

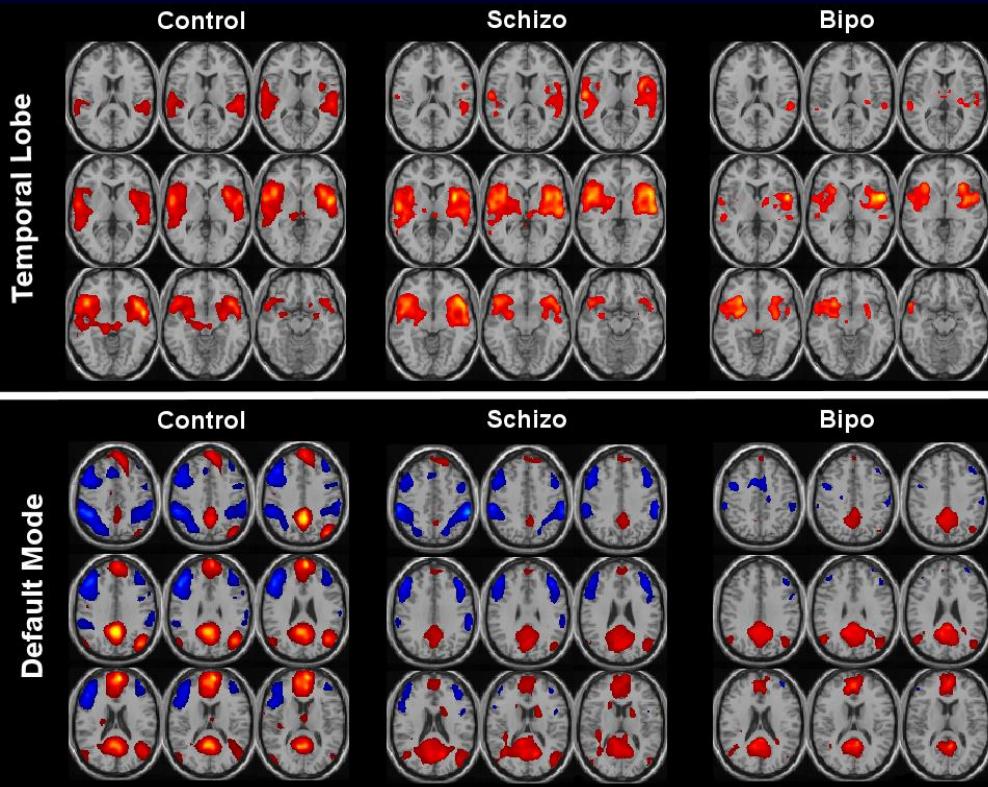
3 seconds before stimulation:

- Baseline *fronto-parietal* activity is high
⇒ stimulus will be *perceived*
- Baseline *default network* activity is high
⇒ stimulus will be *missed*
- Baseline *pain matrix* activity is high
⇒ stimulus will be *more painful*



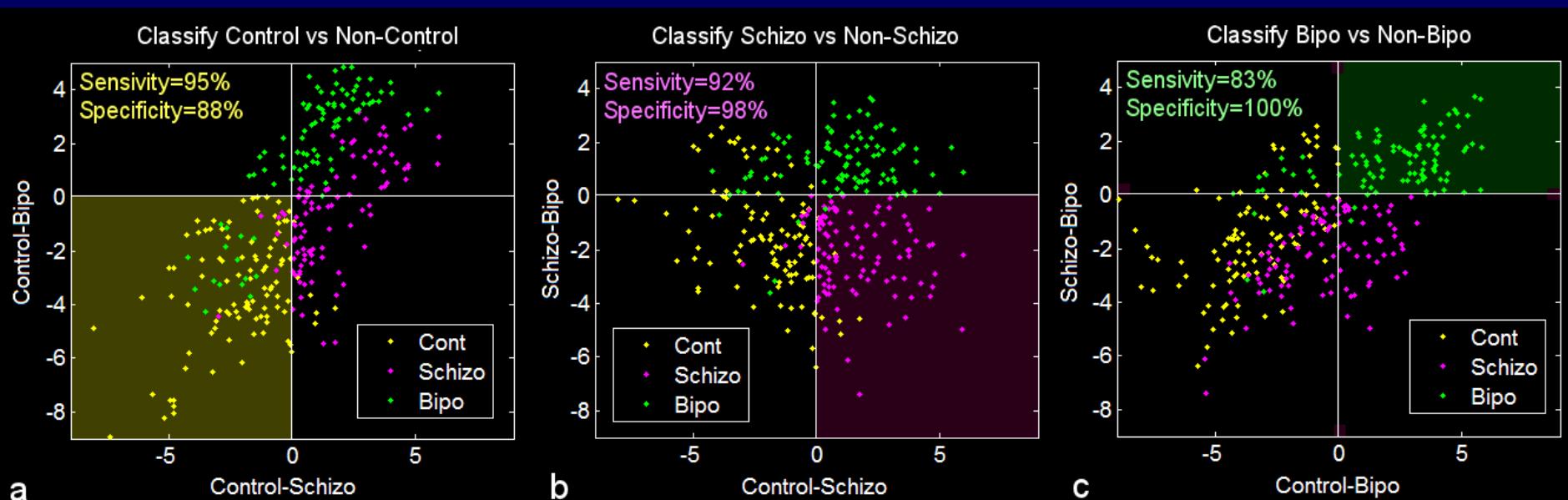
Spontaneous baseline activity fluctuations foretell sensory and pain intensity perception.

Vince Calhoun



Resting state ICA & classification for characterization of schizophrenia and bipolar patients

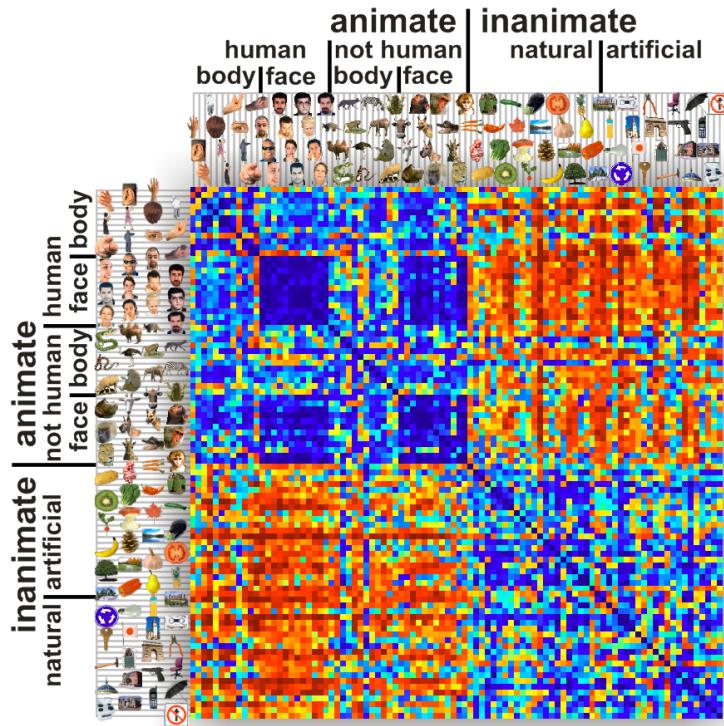
Results show a high average sensitivity (90%) and specificity (95%). Controls were correctly classified 95% of the time, schizophrenia patients 92%, and bipolar patients 81%.



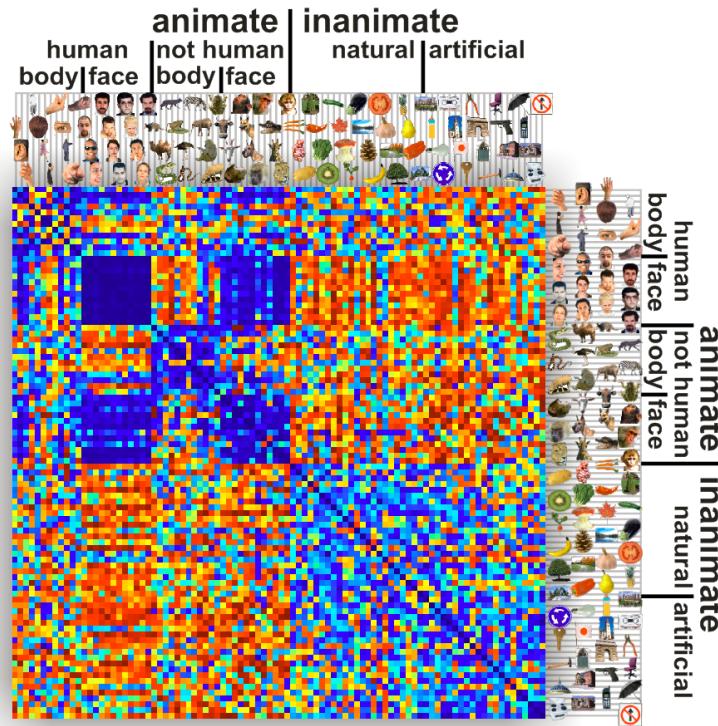
Matching categorical object representations in IT cortex of man & monkey

Kriegeskorte N, Mur M, Ruff D, Kiani R, Bodurka J, Bandettini P

dissimilarity matrices



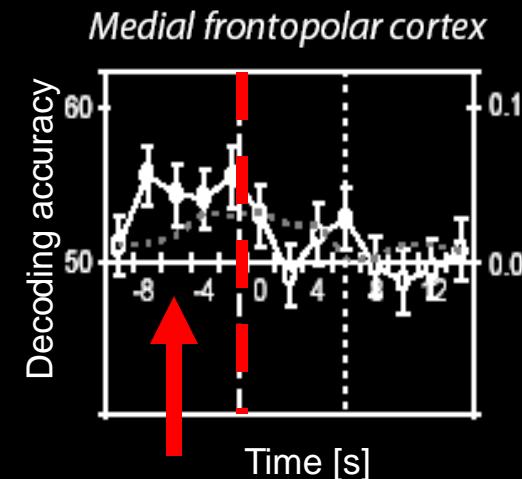
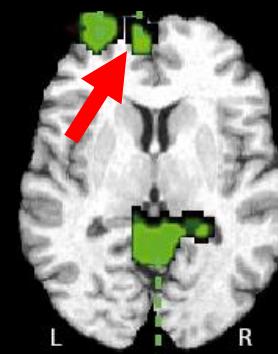
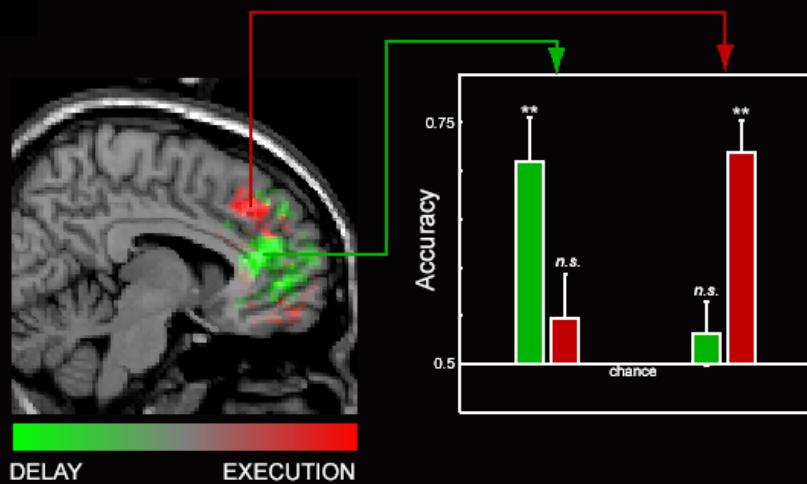
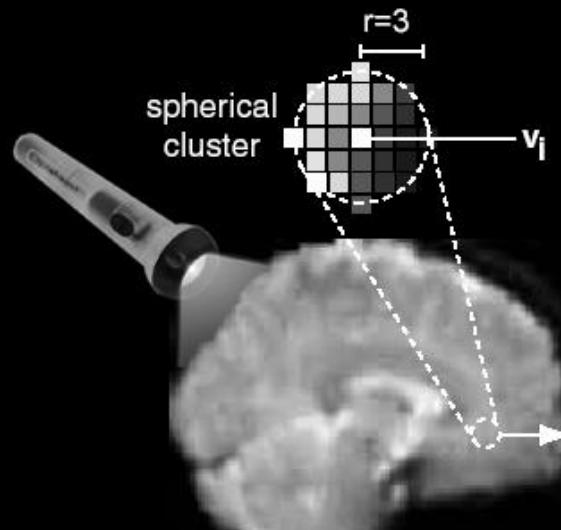
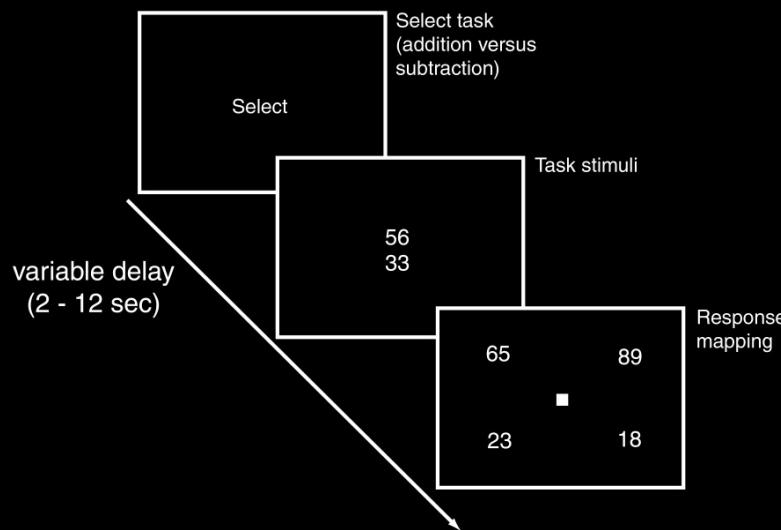
man



monkey

Reading hidden intentions in the human brain

Thu 9.45: Cognition – Representation and Processes

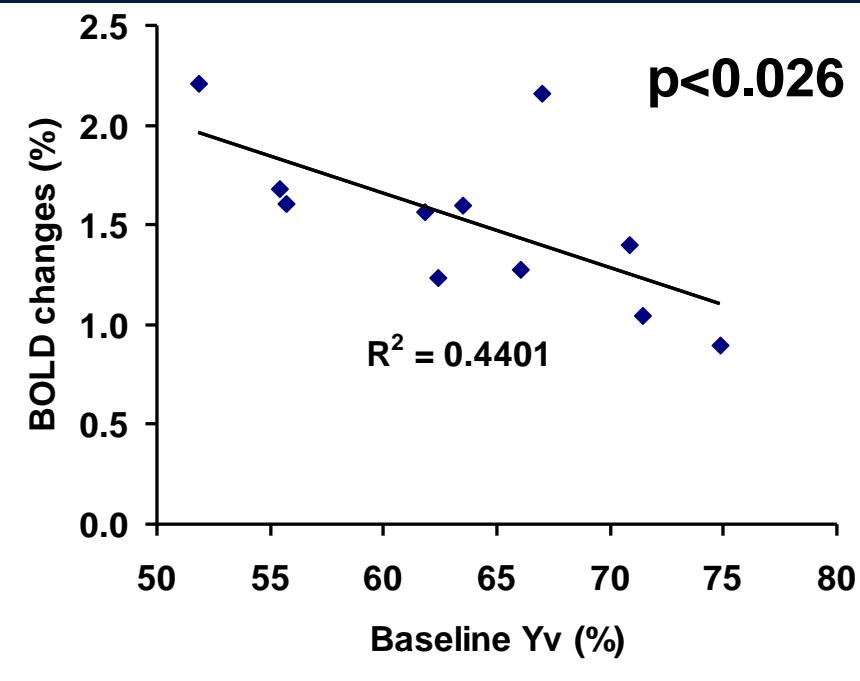


Haynes, Sakai, Rees, Gilbert, Frith & Passingham (Current Biology, 2007)

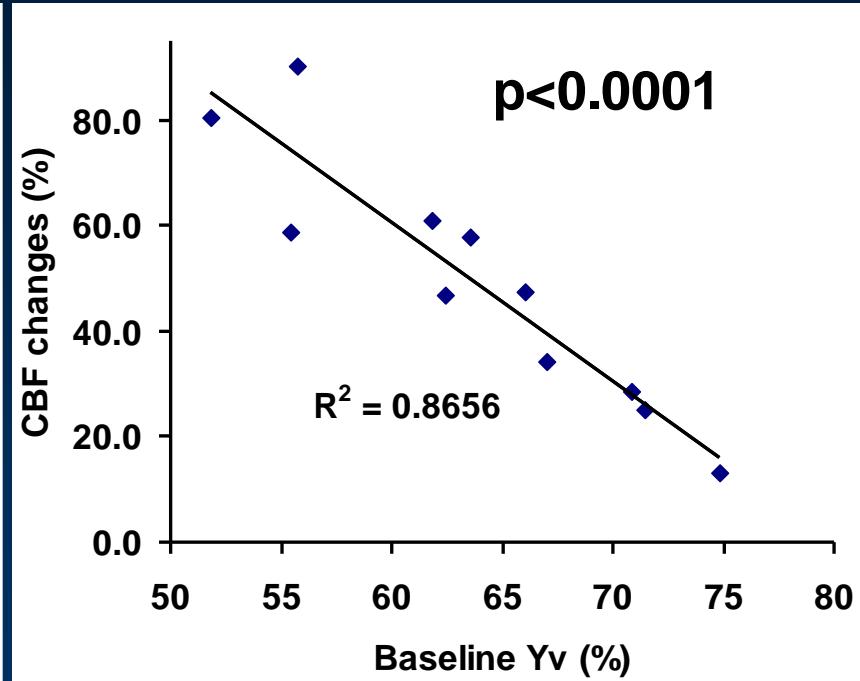
Soon, Brass, Heinze & Haynes (in preparation)

Baseline blood oxygenation modulates fMRI signals

BOLD fMRI



ASL fMRI



Individuals with higher baseline venous oxygenation tend to have smaller BOLD and CBF percentage signal changes