

fMRI: Past, Present, and Future

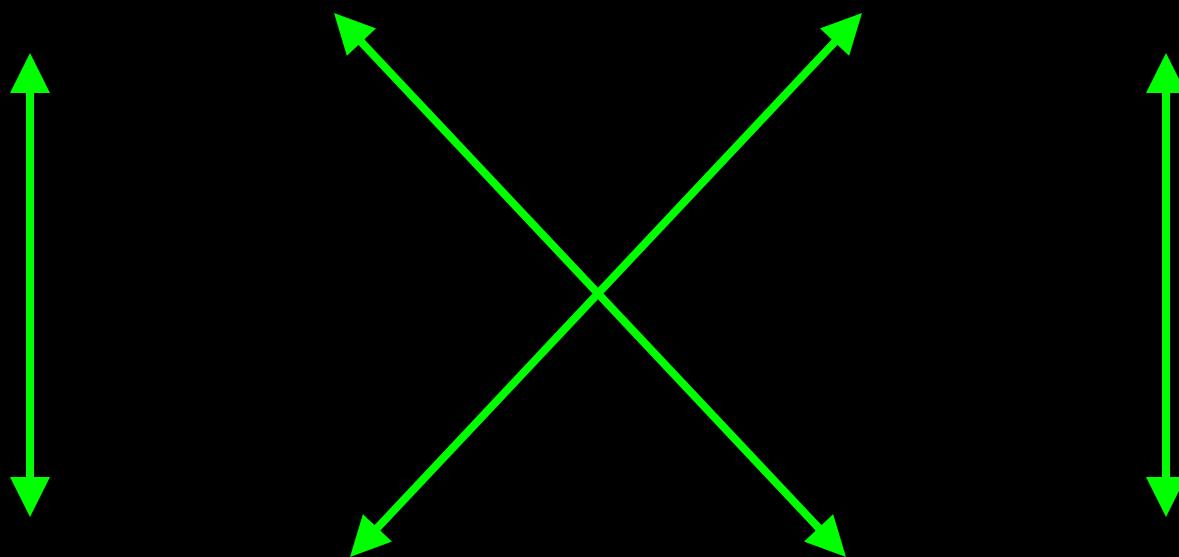
Peter A. Bandettini, Ph.D

Unit on Functional Imaging Methods
&
3T Neuroimaging Core Facility

Laboratory of Brain and Cognition
National Institute of Mental Health



Technology ↔ Methodology



Interpretation ↔ Applications

Technology

Methodology

Engineers

Statisticians

Physicists

Mathematicians

Neuroscientists

Physiologists

Clinicians

Interpretation

Applications

Technology

MRI	EPI	1.5T,3T, 4T	EPI on Clin. Syst.	Diff. tensor	Mg ⁺	7T
		Local Human Head Gradient Coils		Real time fMRI	Venography	
	ASL	Spiral EPI	Nav. pulses	Quant. ASL	Z-shim	SENSE
	BOLD		Multi-shot fMRI	Dynamic IV volume	Simultaneous ASL and BOLD	Baseline Susceptibility
						Current Imaging?

Methodology

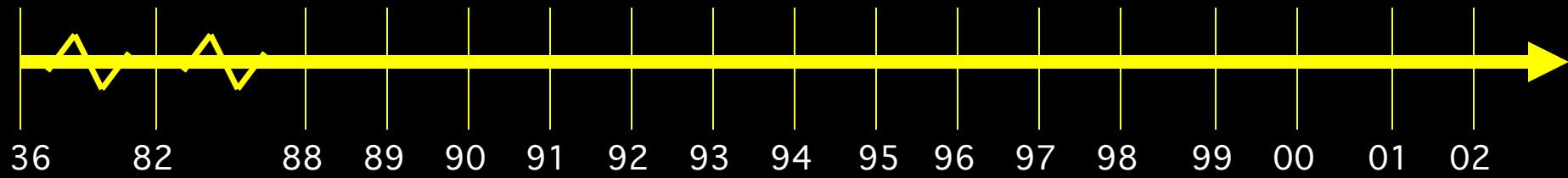
IVIM	Baseline Volume	Correlation Analysis	Motion Correction	CO ₂ Calibration
		Parametric Design		Multi-Modal Mapping
		Surface Mapping		Free-behavior Designs
		Phase Mapping		
		Linear Regression	Mental Chronometry	
		Event-related	Deconvolution	

Interpretation

Blood T2	BOLD models	PET correlation		
	B ₀ dep.	IV vs EV	ASL vs. BOLD	
		Pre-undershoot	PSF of BOLD	
	TE dep	Resolution Dep.	Extended Stim.	
		Post-undershoot	Linearity	Metab. Correlation
Hemoglobin	SE vs. GE	CO ₂ effect	Fluctuations	Optical Im. Correlation
		NIRS Correlation	Balloon Model	Electrophys. correlation
	Veins	Inflow		

Applications

Volume - Stroke	Complex motor			
	Language	Imagery	Memory	Emotion
	Motor learning	Children	Tumor vasc.	Drug effects
	BOLD -V1, M1, A1	Presurgical	Attention	Ocular Dominance
		V1, V2..mapping	Priming/Learning	Clinical Populations
	△ Volume-V1		Plasticity	Face recognition
				Performance prediction



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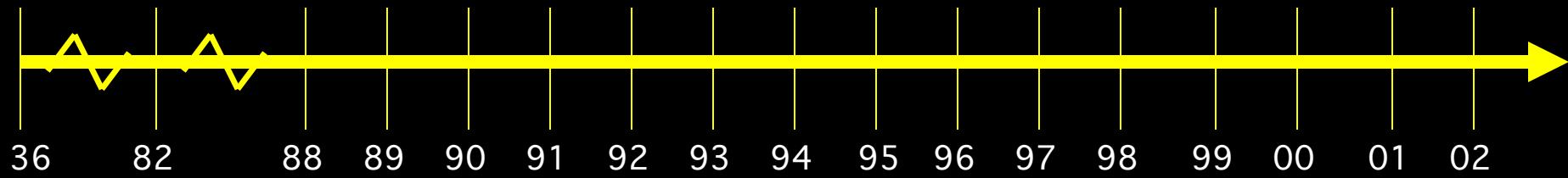
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L. Pauling, C. D. Coryell, (1936) "The magnetic properties and structure of hemoglobin, oxyhemoglobin, and carbonmonoxyhemoglobin." Proc.Natl. Acad. Sci. USA 22, 210-216.

Thulborn, K. R., J. C. Waterton, et al. (1982). "Oxygenation dependence of the transverse relaxation time of water protons in whole blood at high field." Biochim. Biophys. Acta. 714: 265-270.

S. Ogawa, T. M. Lee, A. R. Kay, D. W. Tank, (1990) "Brain magnetic resonance imaging with contrast dependent on blood oxygenation." Proc. Natl. Acad. Sci. USA 87, 9868-9872.

R. Turner, D. LeBihan, C. T. W. Moonen, D. Despres, J. Frank, (1991). Echo-planar time course MRI of cat brain oxygenation changes. Magn. Reson. Med. 27, 159-166.

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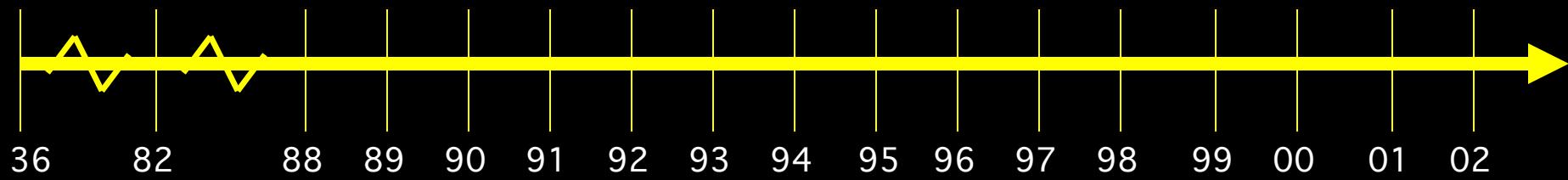
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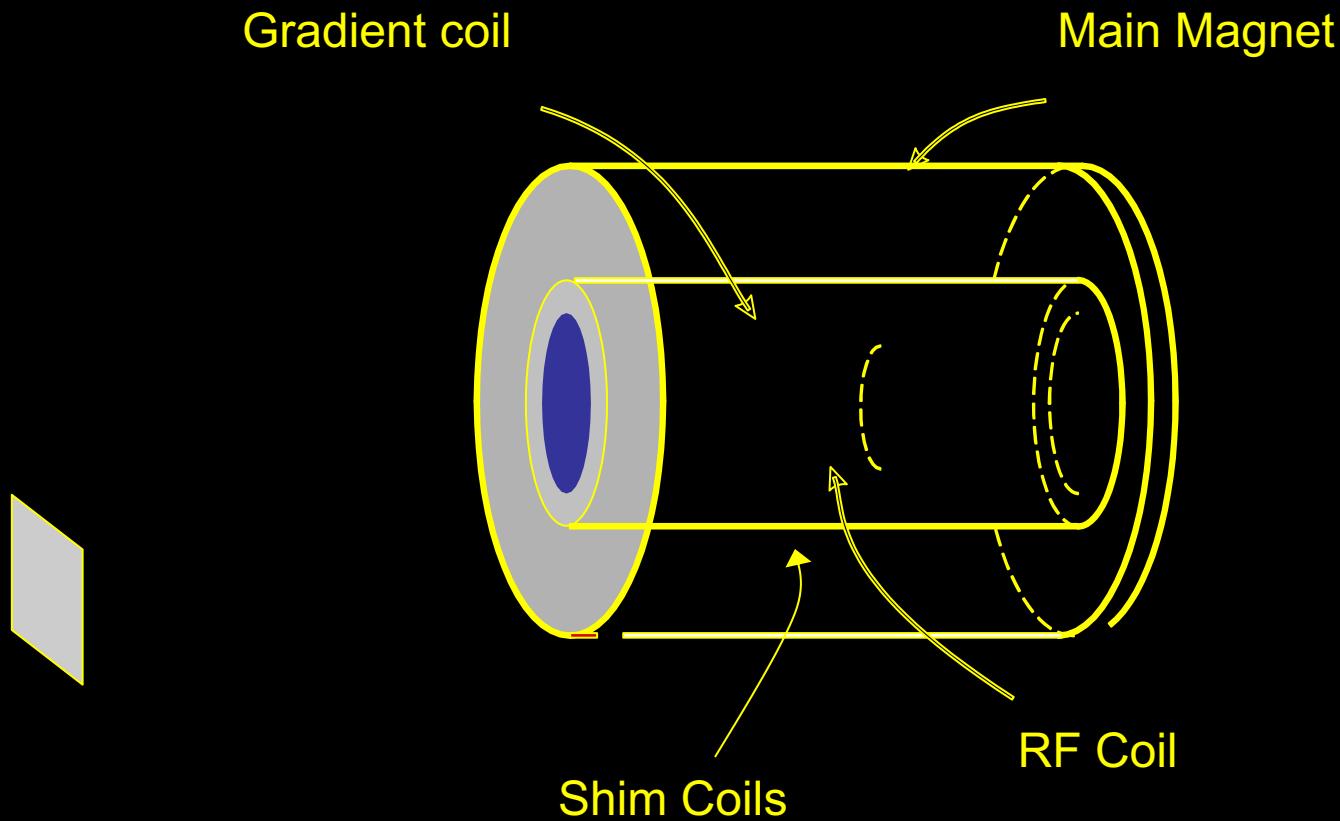


$$f_o = \gamma B_o$$

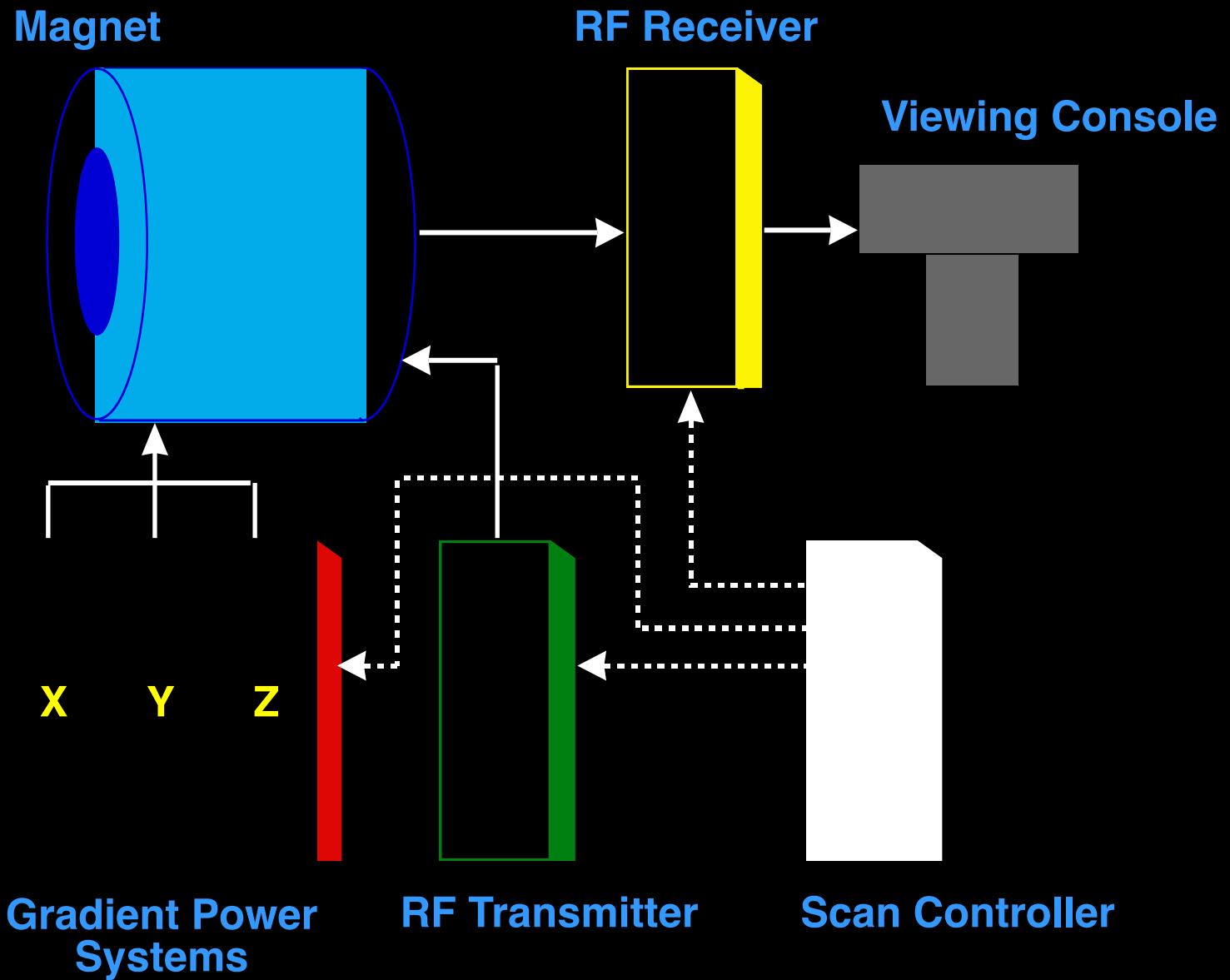
f_o = Larmor frequency

γ = gyromagnetic ratio (42.6 MHz/Tesla)

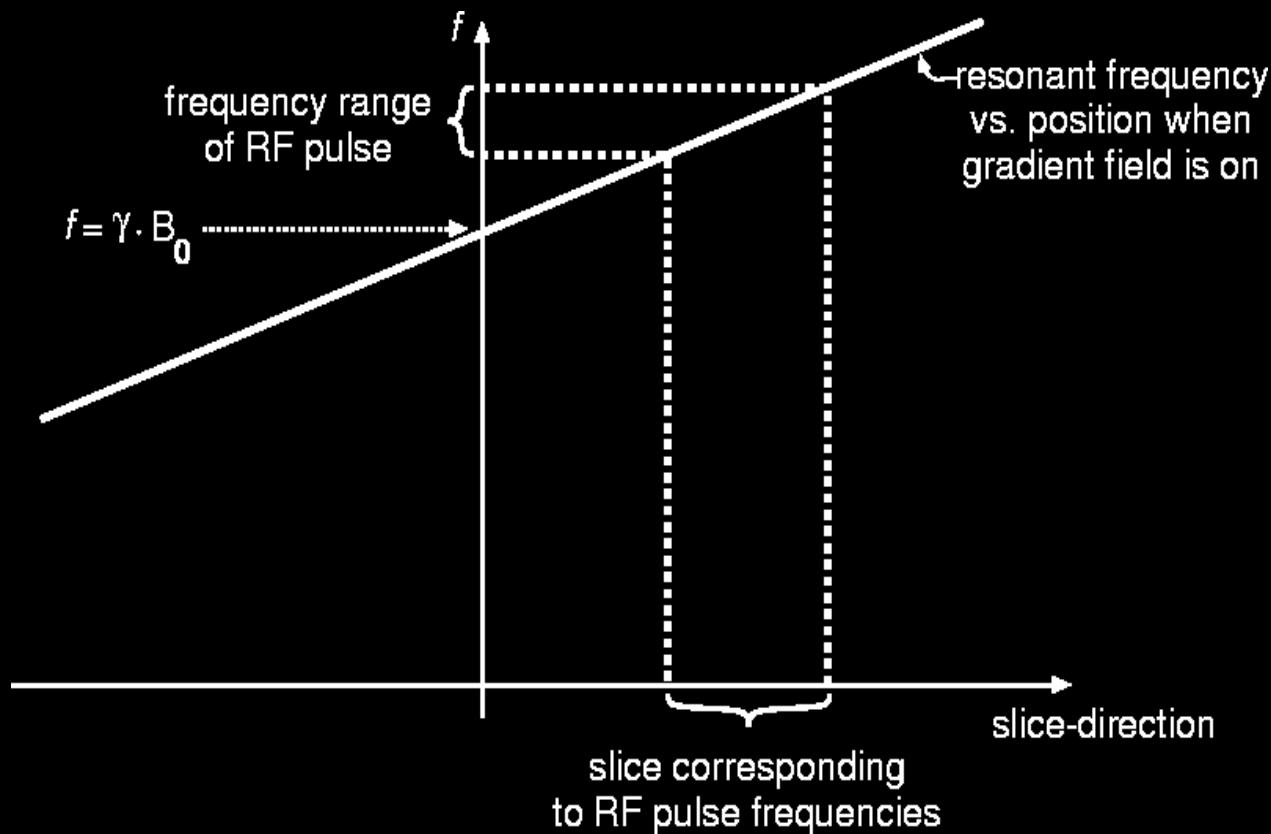
B_o = magnetic field strength (Tesla)



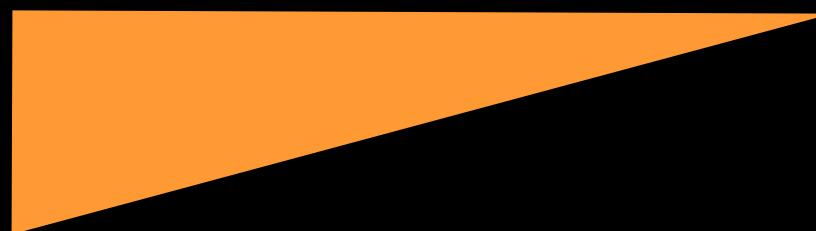
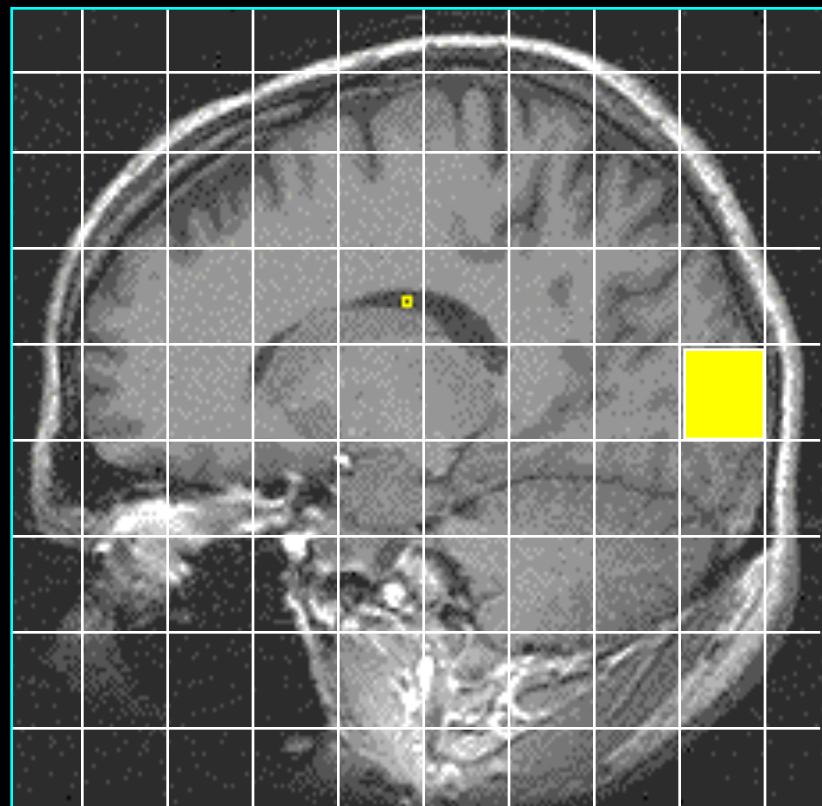
Imaging System Components

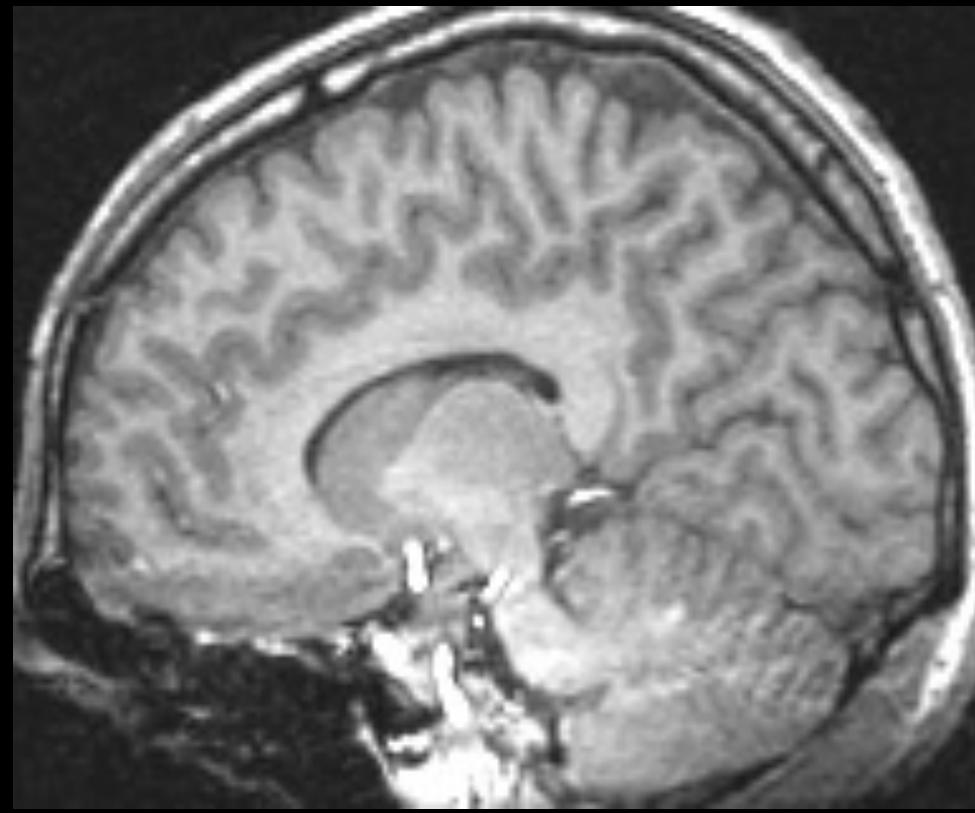
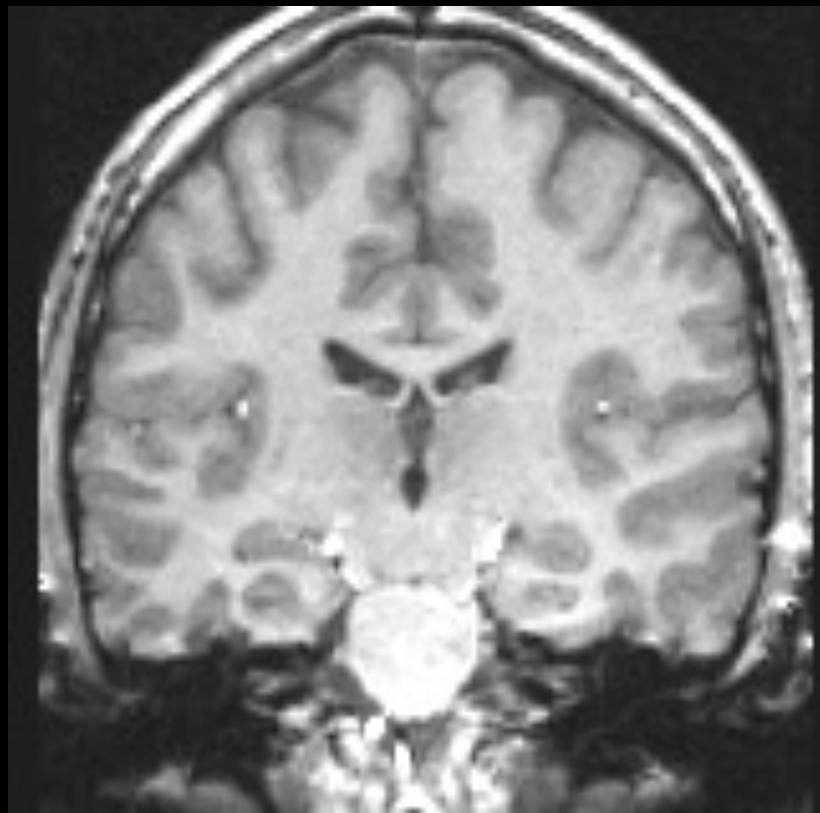


Slice Selection



In – Plane Spatial Localization



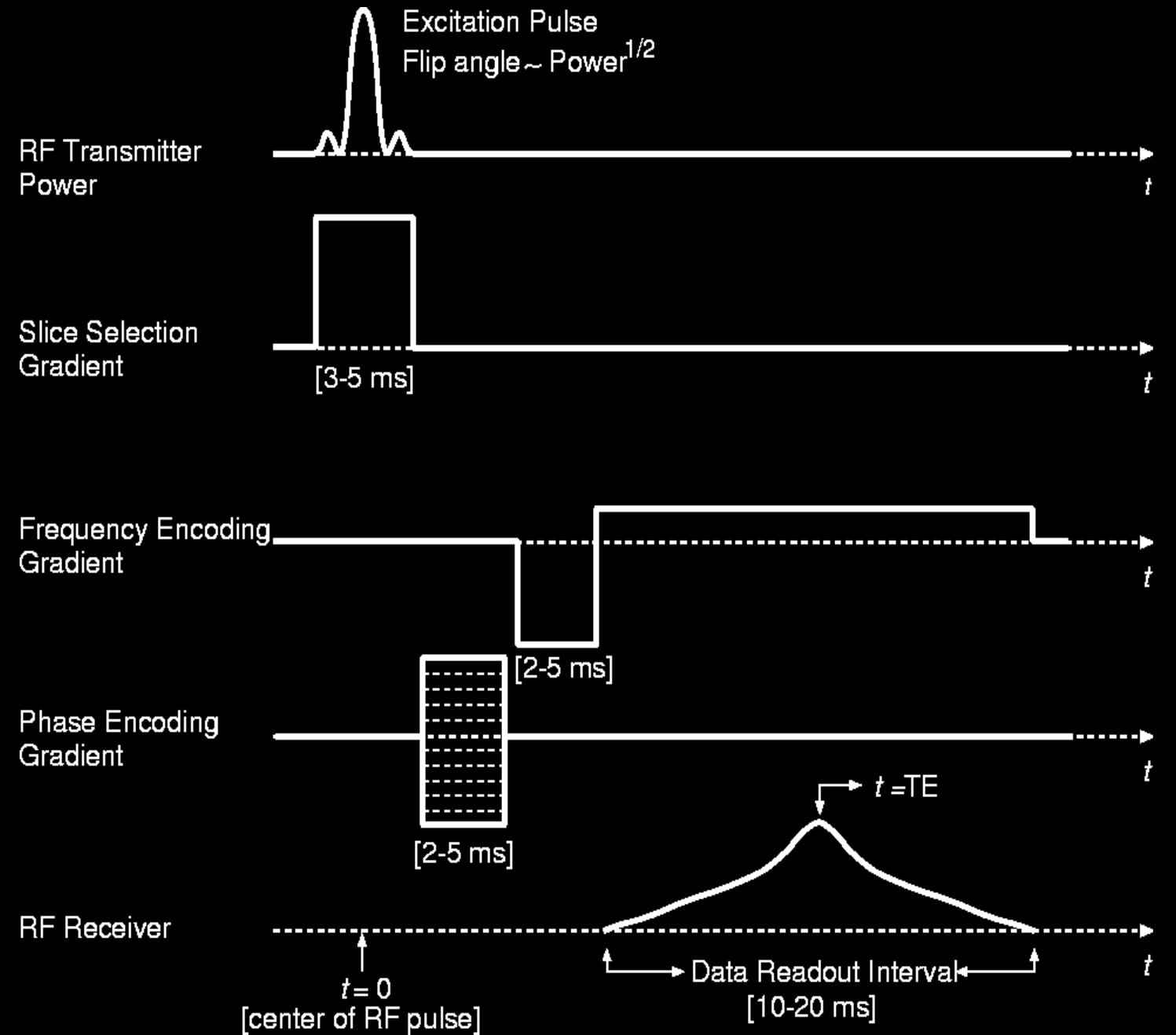


Source of Anatomical Contrast: MRI Parameters



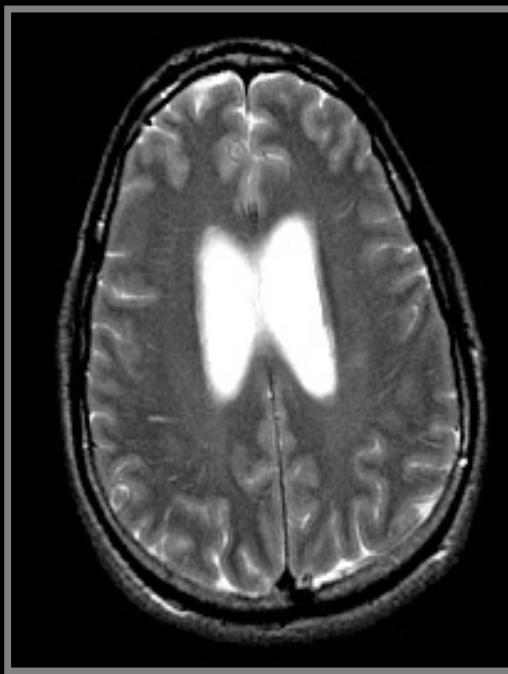
- Spin (Proton) Density
- T1 Relaxation Time
- T2, T2* Relaxation Times

MRI Pulse Sequence for Gradient Echo Imaging

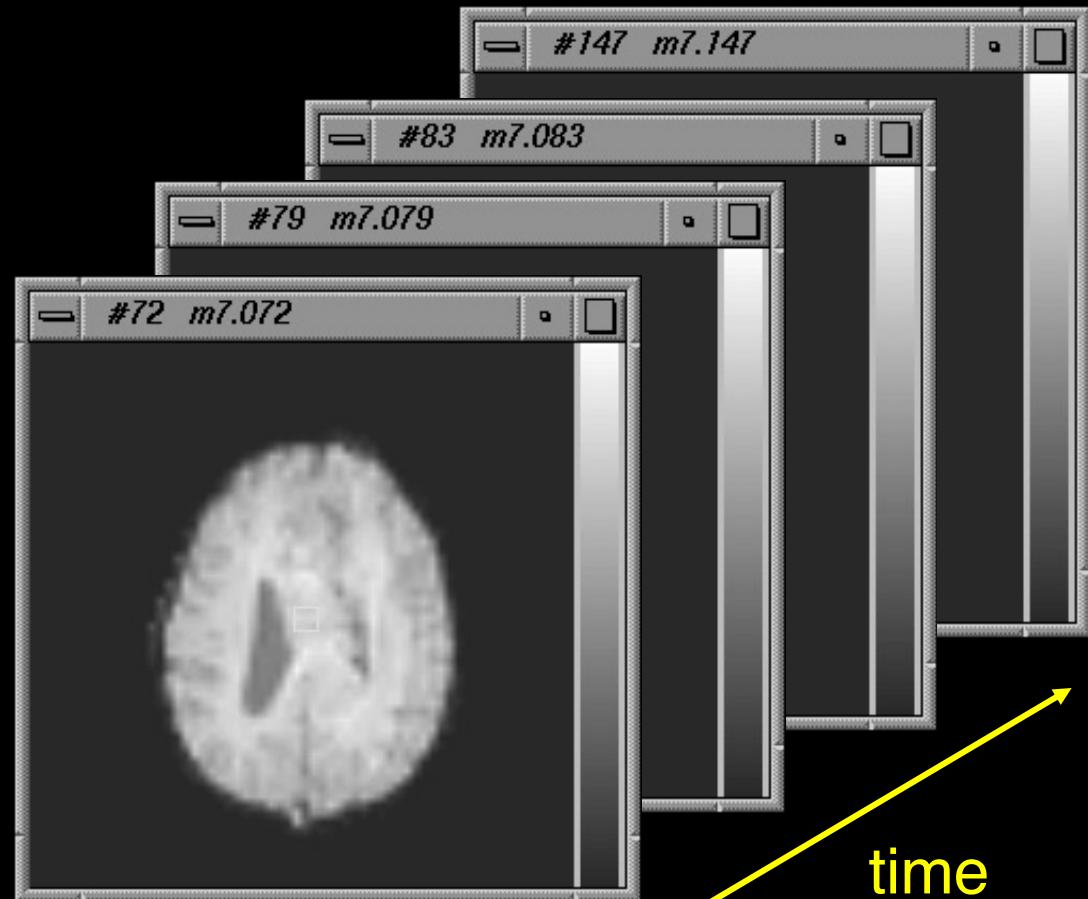


Illustrates sequence of events during scanning

As shown, this method (FLASH) takes 35 ms per RF shot, so would take 2.25 s for a 64×64 image

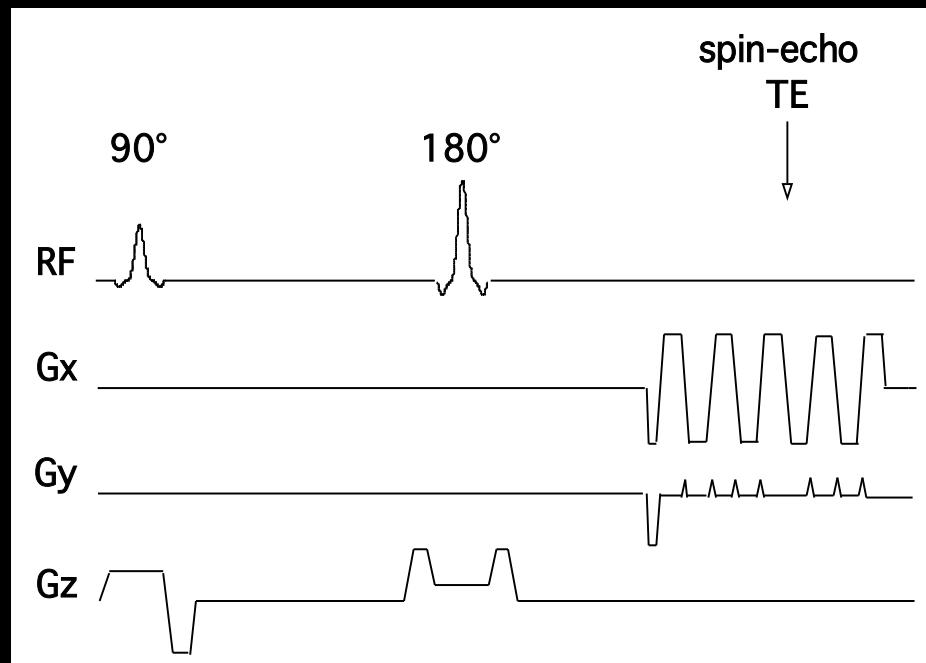
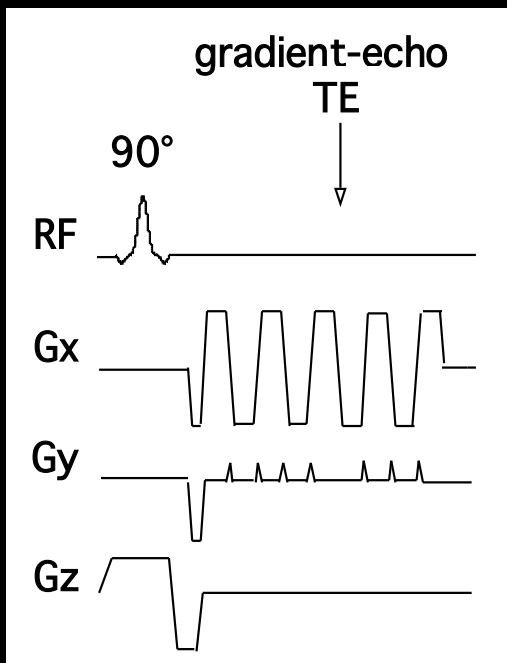


Anatomic



Functional

Echo-Planar Imaging

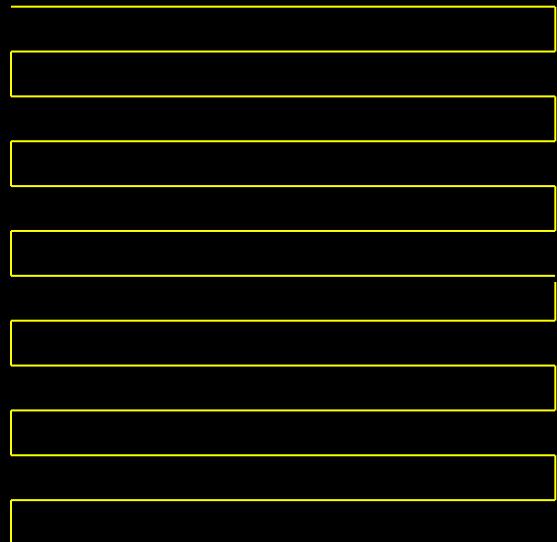


Single Shot Imaging

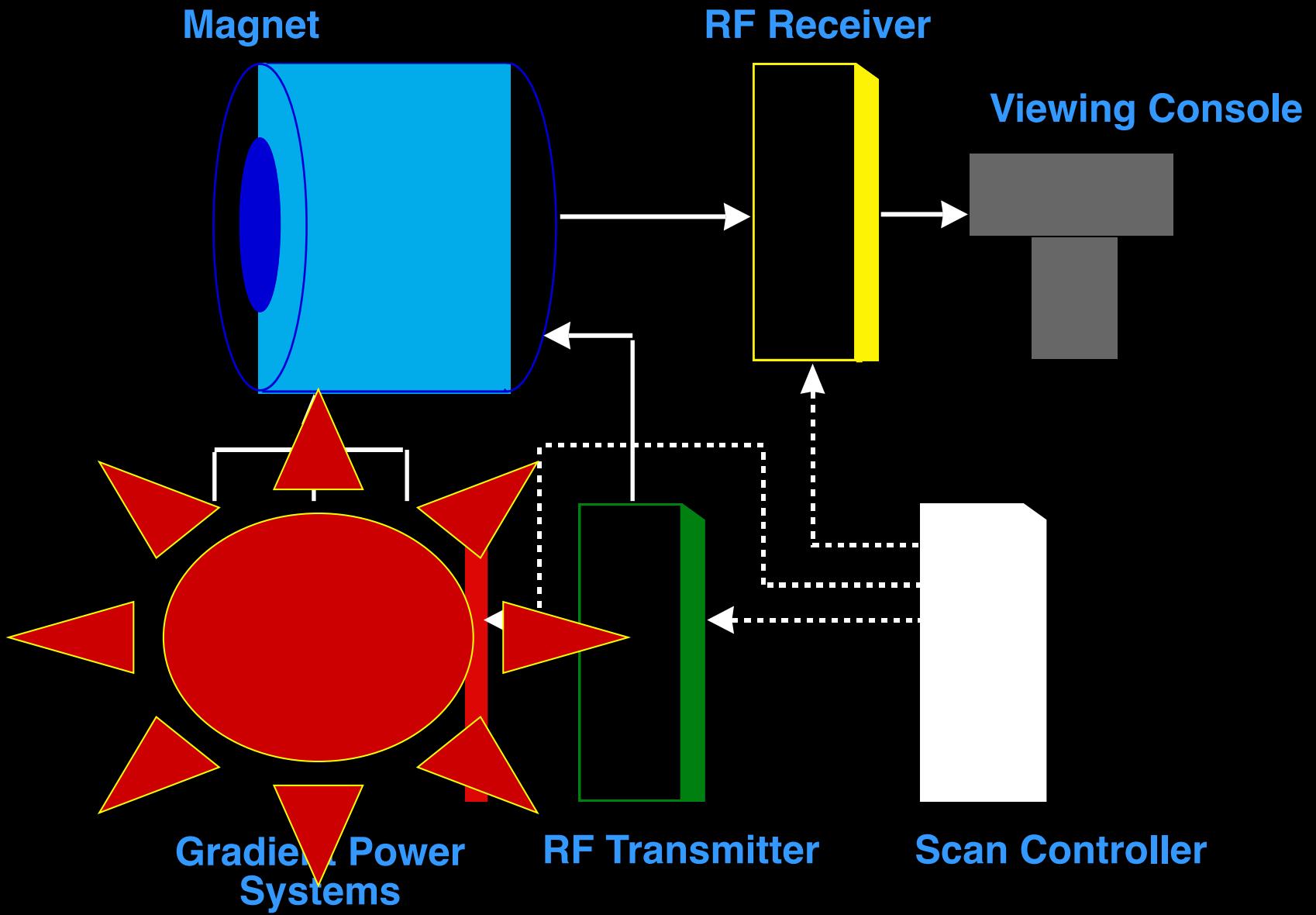


EPI Readout Window

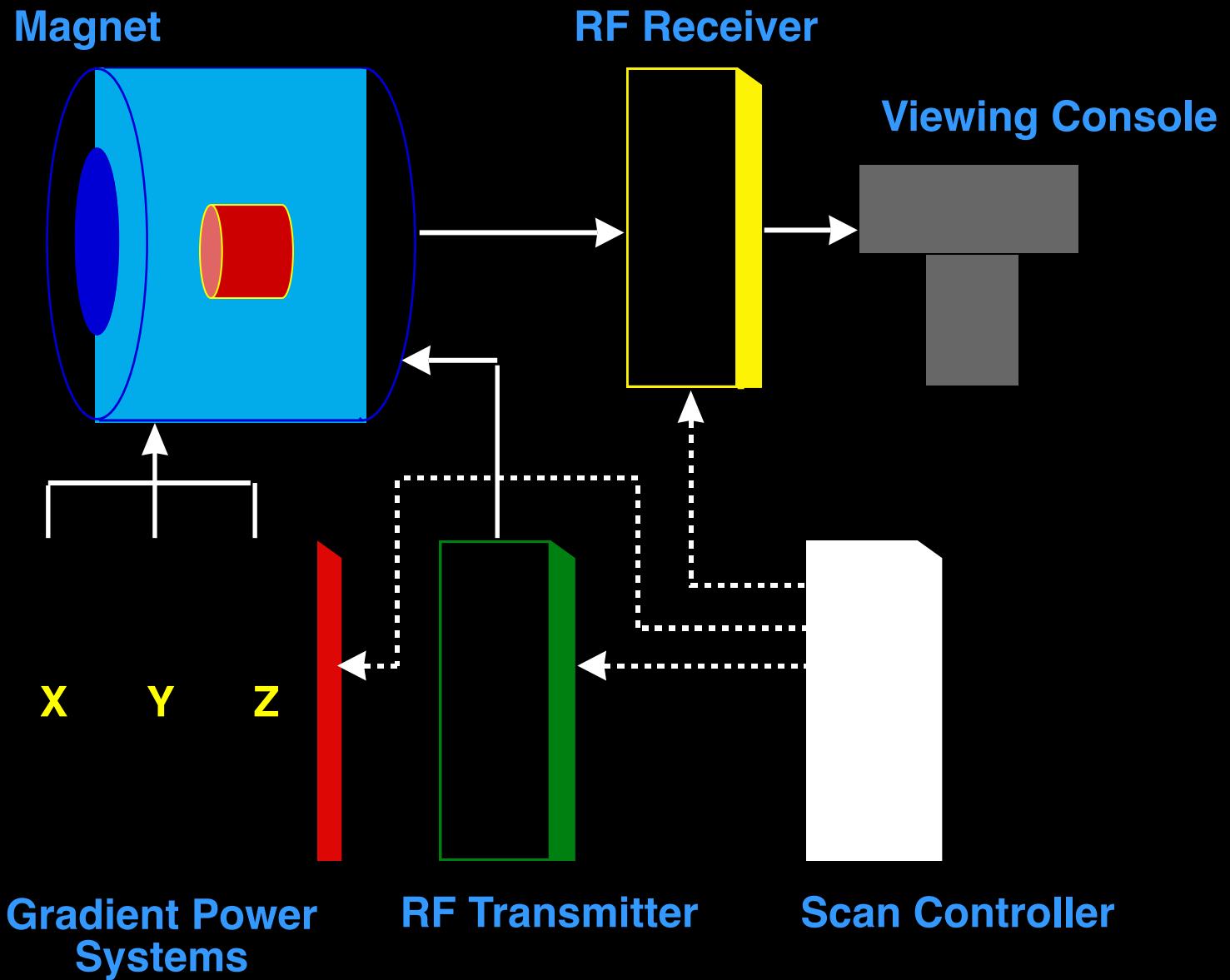
≈ 20 to 40 ms



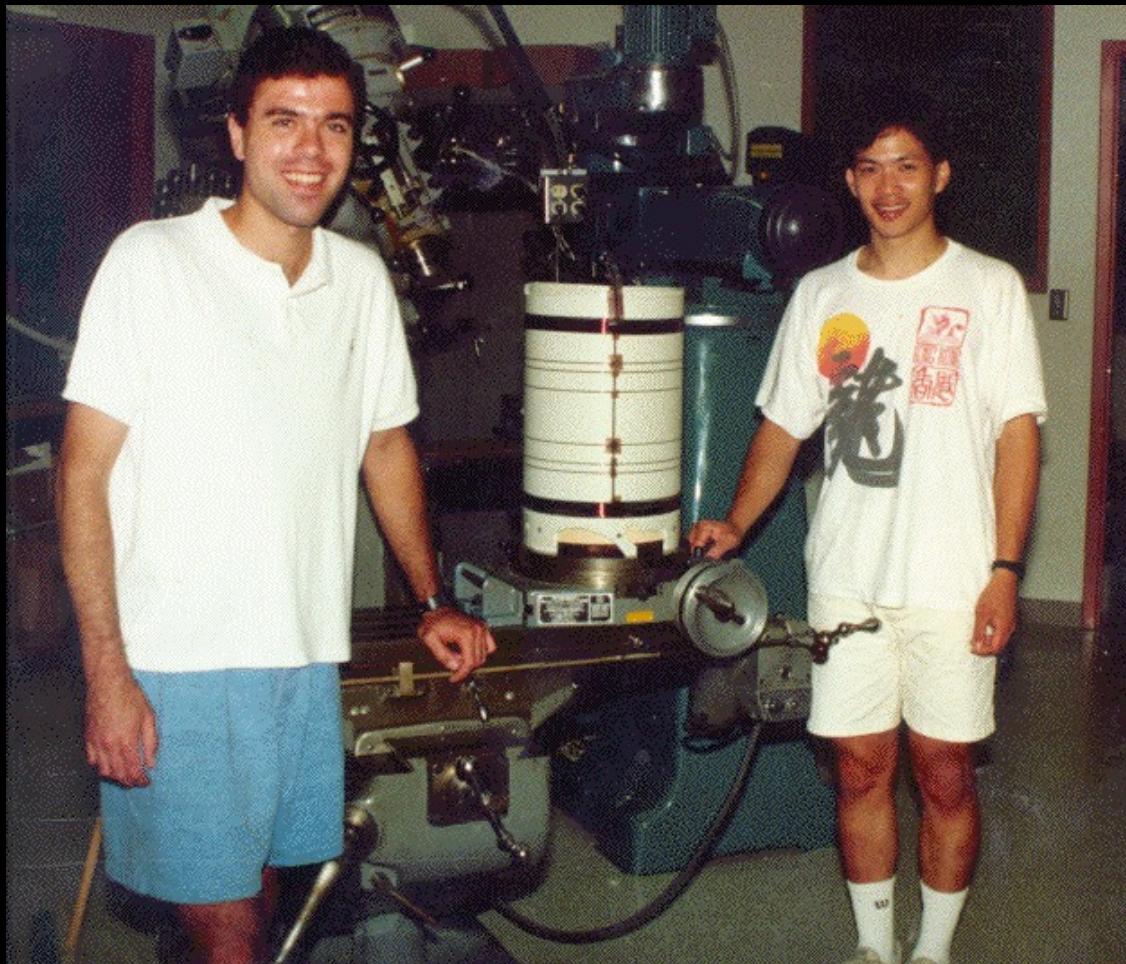
Imaging System Components



Imaging System Components



Local gradients solved the problem



August, 1991

1991-1992



1992-1999



Functional MRI Methods

Blood Volume Imaging

BOLD Contrast

Arterial Spin Labeling

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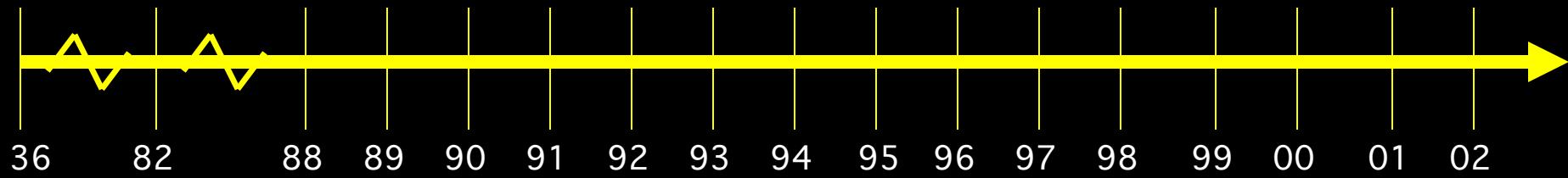
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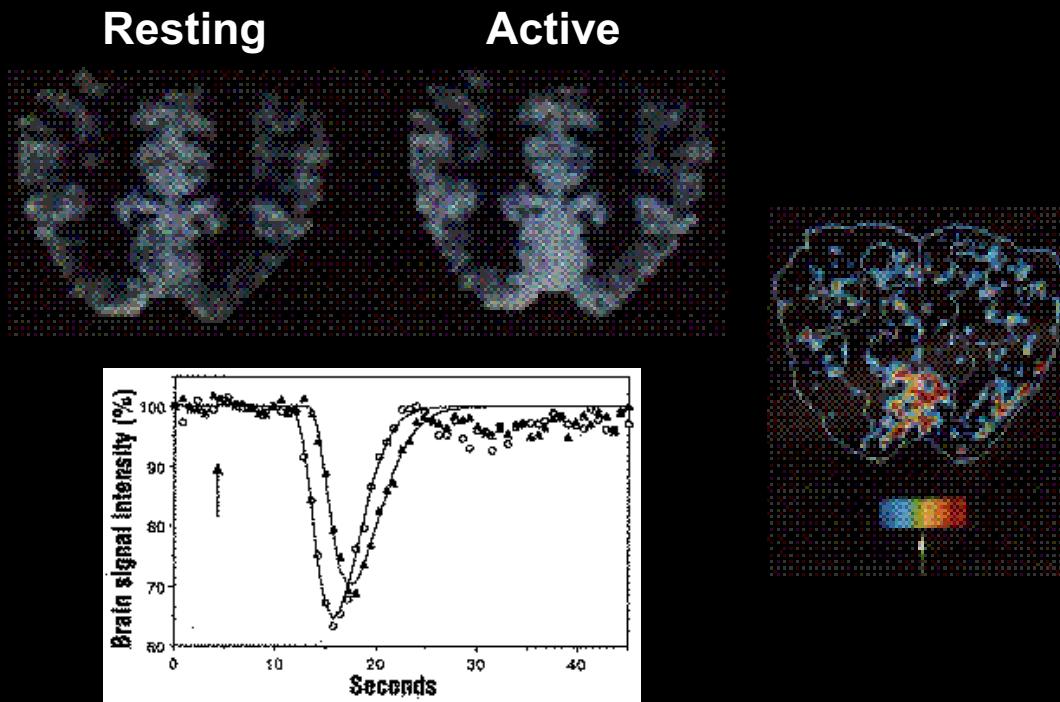
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Blood Volume Imaging

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



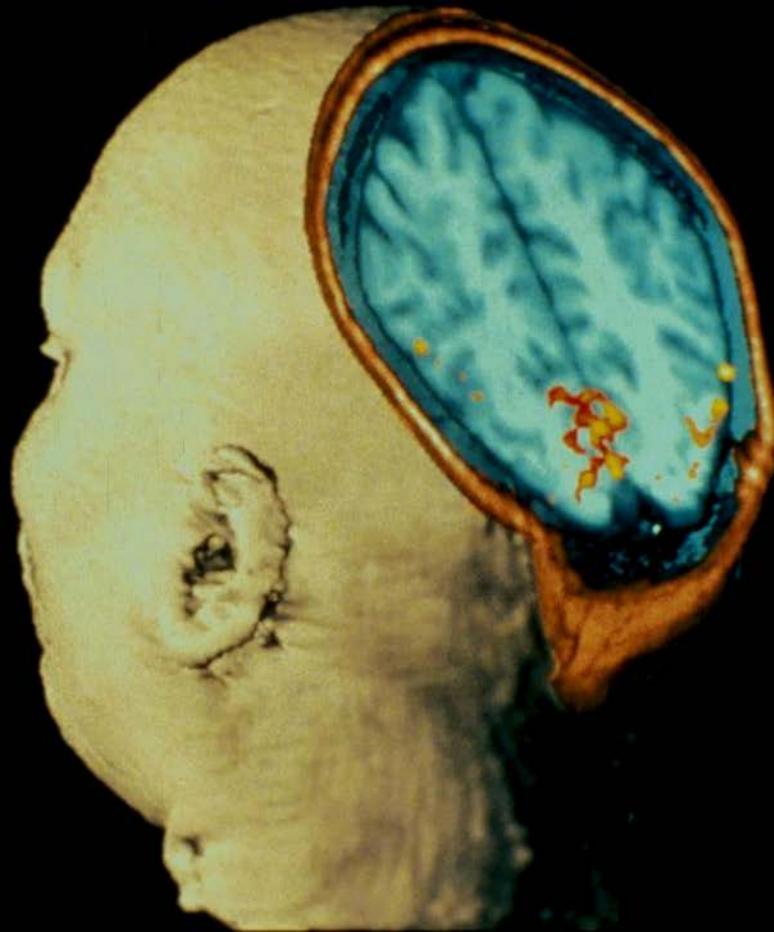
Blood Volume

**Photic
Stimulation**

**MRI Image showing
activation of the
Visual Cortex**

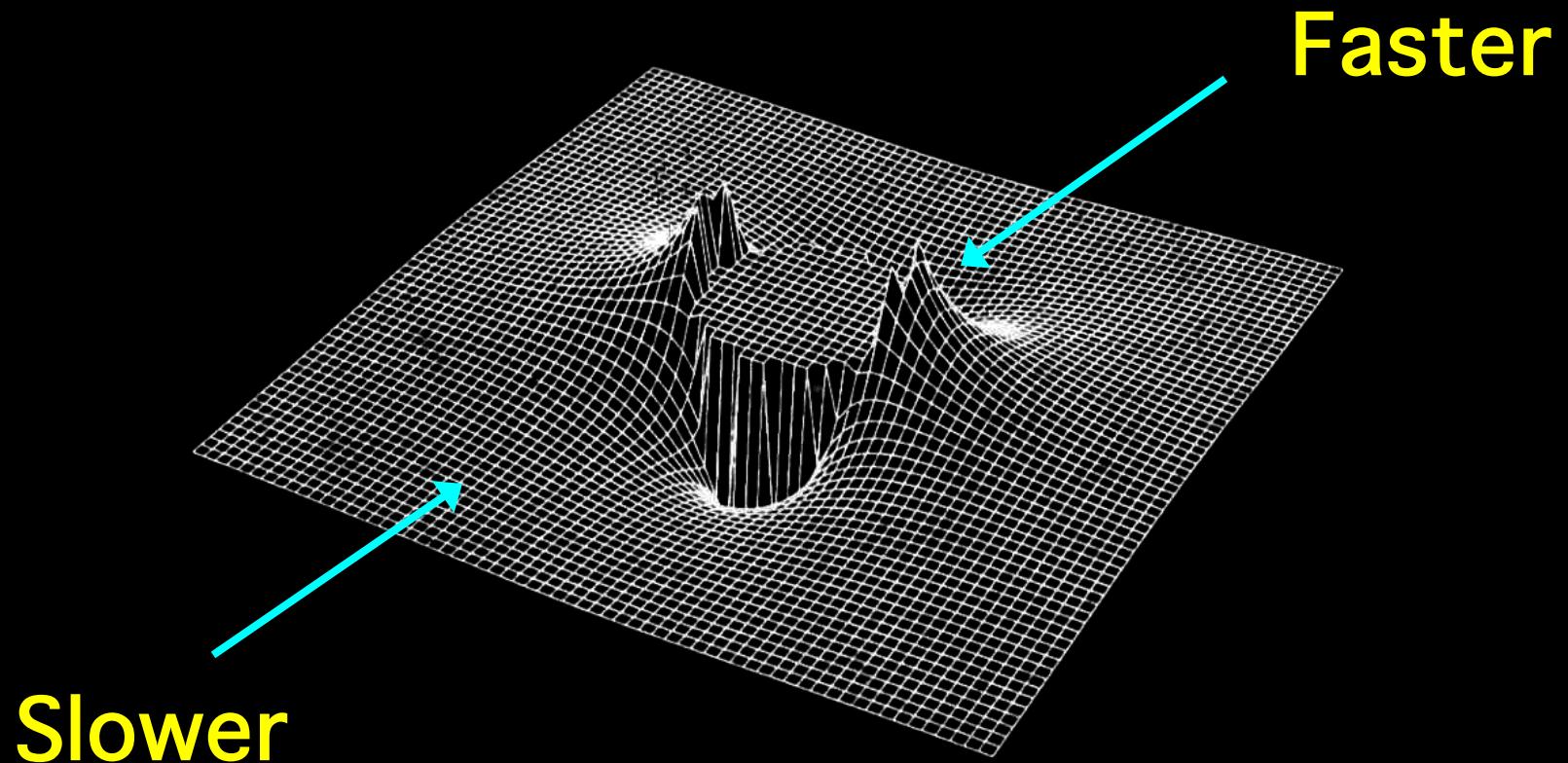
**From Belliveau, et al.
Science Nov 1991**

MSC - perfusion



Susceptibility Contrast

Susceptibility-Induced Field Distortion in the
Vicinity of a Microvessel \perp to B_0 .



Alternating Left and Right Finger Tapping



~ 1992

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

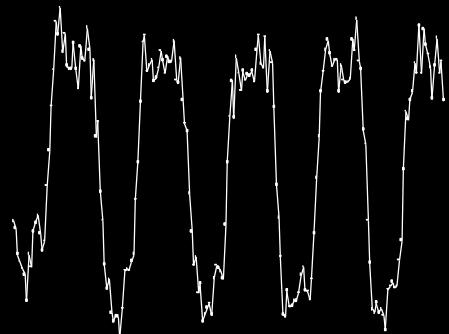
S. Ogawa, et al., (1992) “Intrinsic signal changes accompanying sensory stimulation: functional brain mapping with magnetic resonance imaging. Proc. Natl. Acad. Sci. USA.” 89, 5951-5955.

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

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.

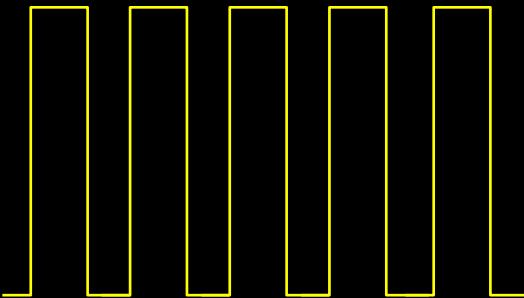
Creating a Functional Image

ON ON ON ON ON



Signal Time Course

X



Reference Function

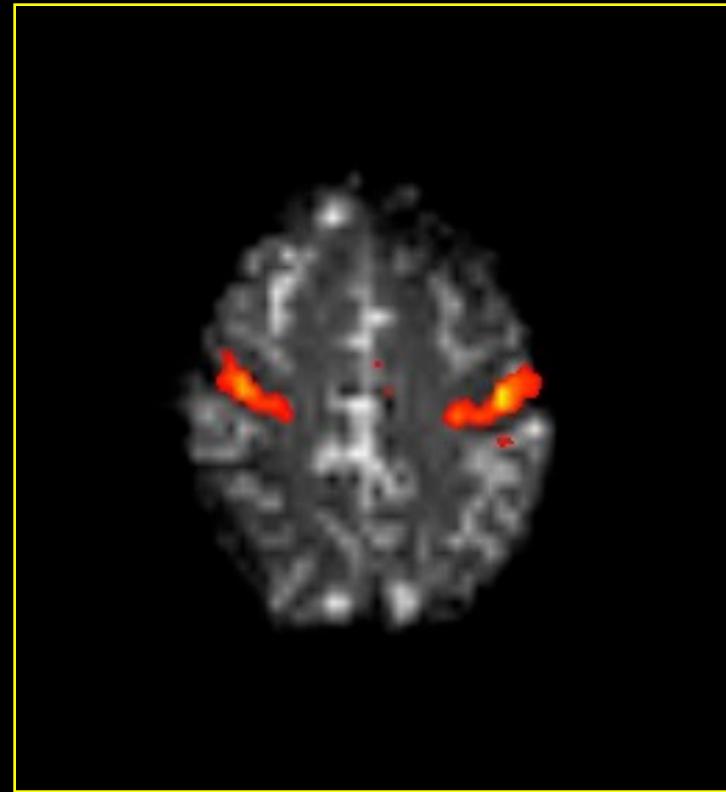
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P. A. Bandettini, A. Jesmanowicz, E. C. Wong, J. S. Hyde, Processing strategies for time-course data sets in functional MRI of the human brain. *Magn. Reson. Med.* **30**, 161-173 (1993).



Cross Correlation Image



Cross Correlation Image
Anatomical Image

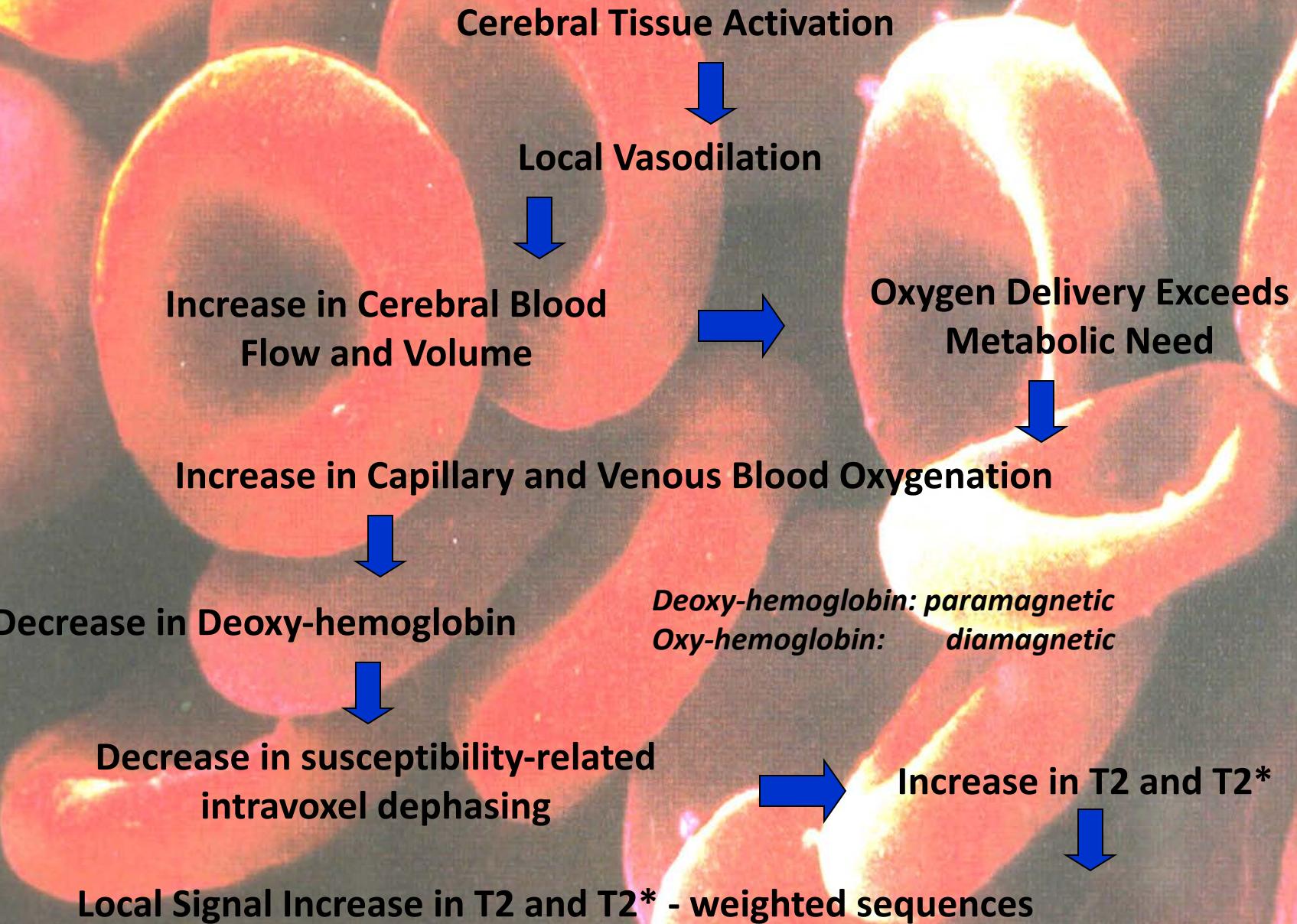
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**Correlation analysis, Fourier analysis, t-test, f-test...
SPM, AFNI, brain voyager, FIASCO, FSL, free surfer...**



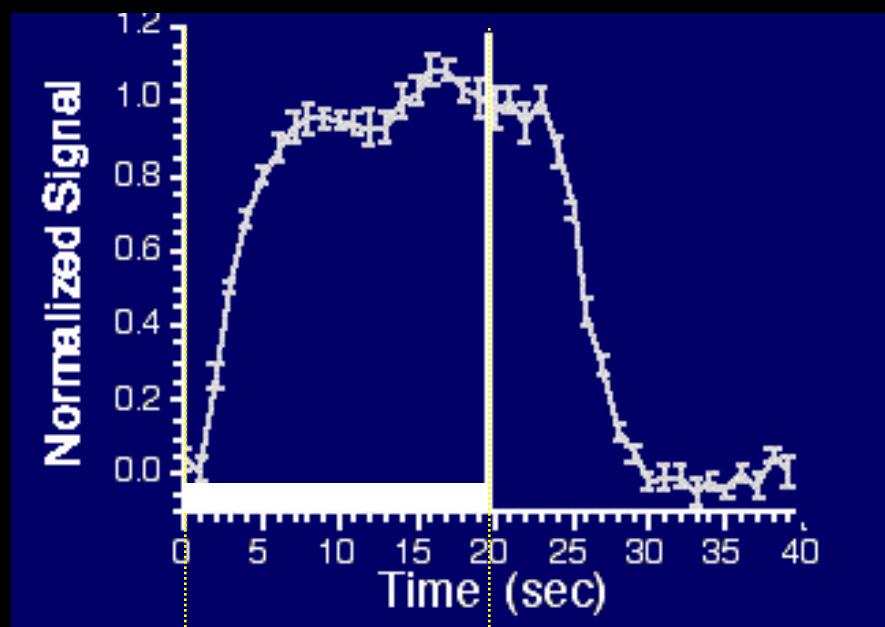
*Quality of results and importance of the findings depends on
type of question asked, experimental method, and analysis method...*

BOLD Contrast in the Detection of Neuronal Activity

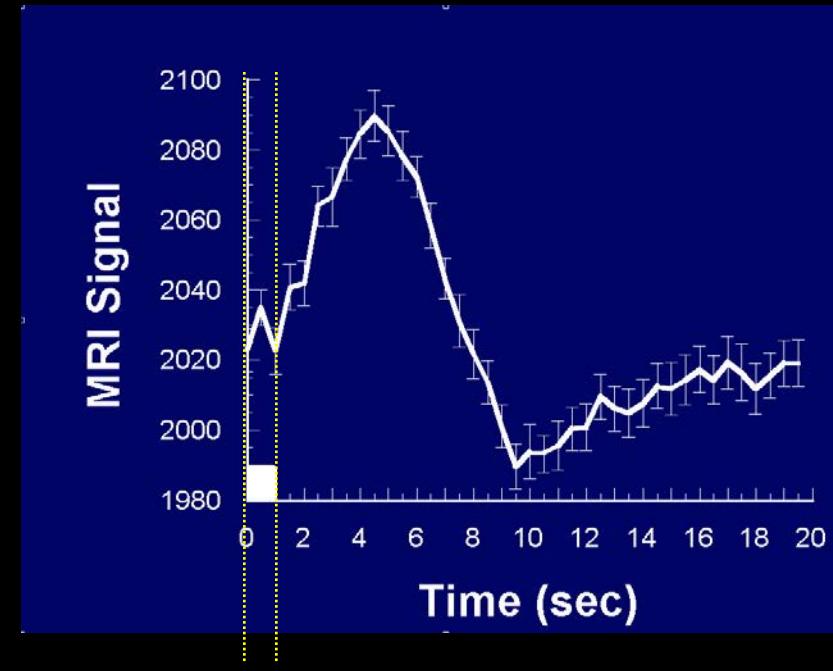


The BOLD Signal

Blood Oxxygenation Level Dependent (BOLD) signal changes

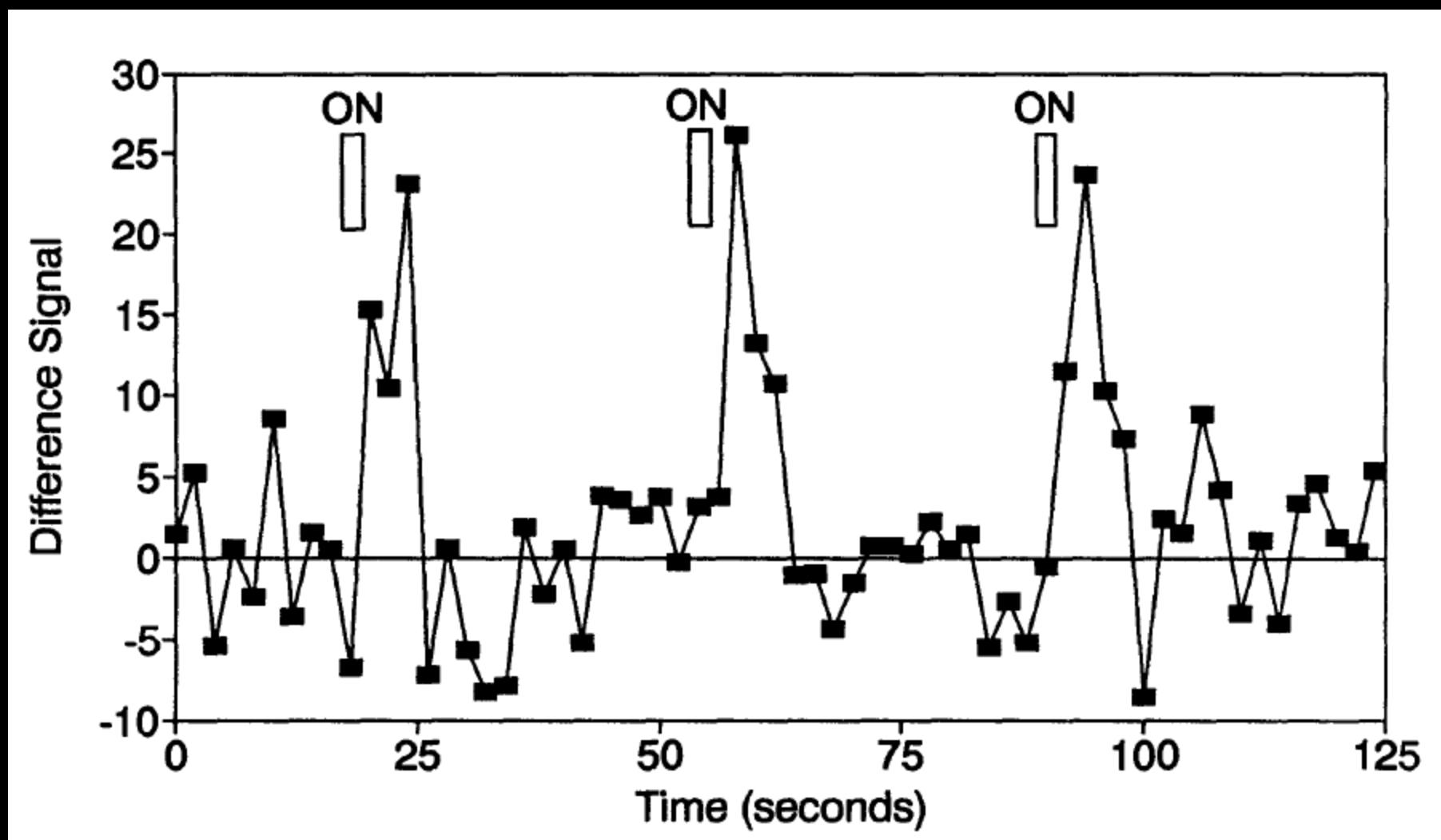


task



task

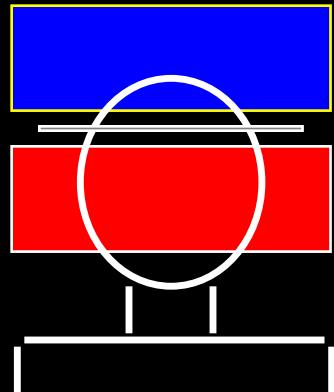
First Event-related fMRI Results



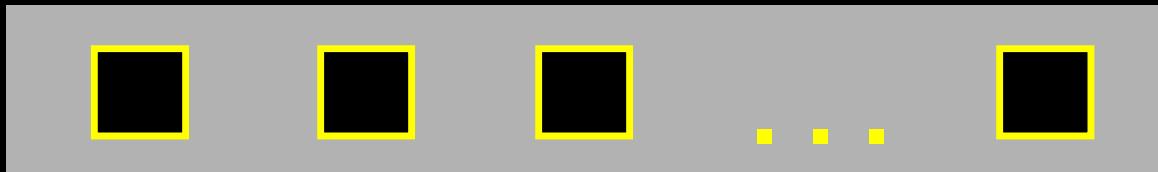
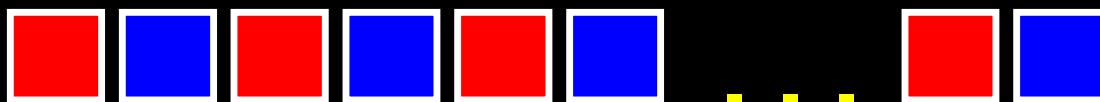
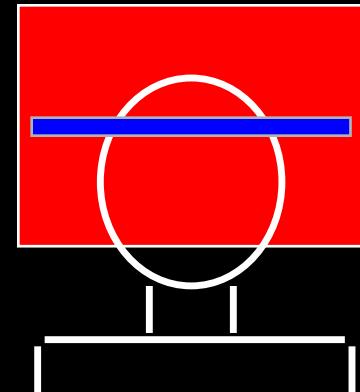
Blamire, A. M., et al. (1992). "Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging." Proc. Natl. Acad. Sci. USA 89: 11069-11073.

Blood Perfusion

EPISTAR



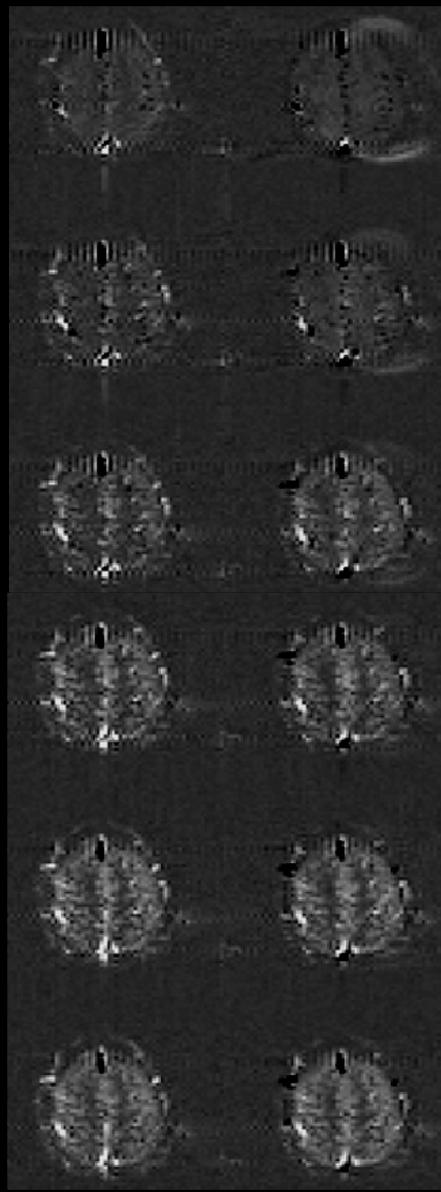
FAIR



Perfusion
Time Series

TI (ms) FAIR EPISTAR

200



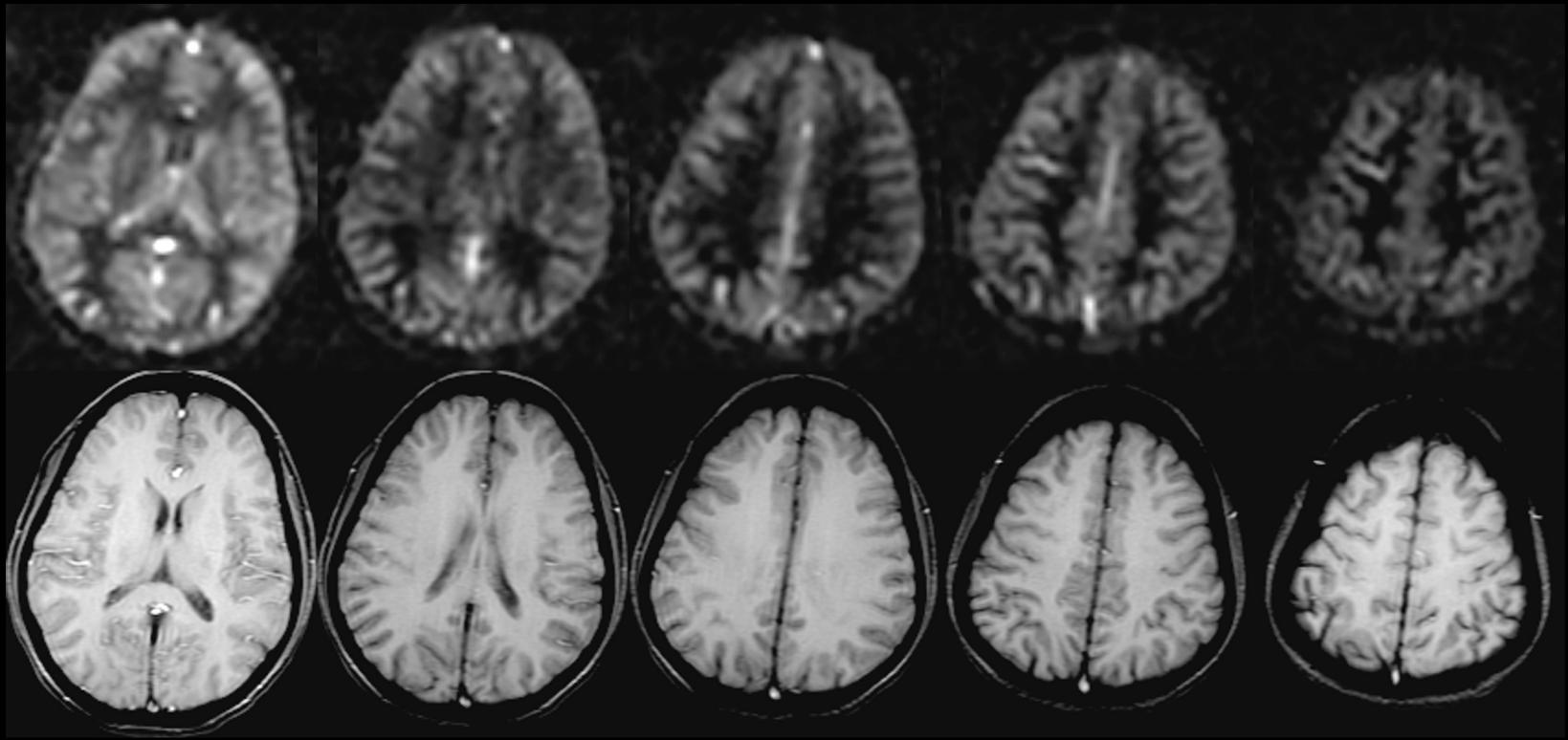
400

600

800

1000

1200



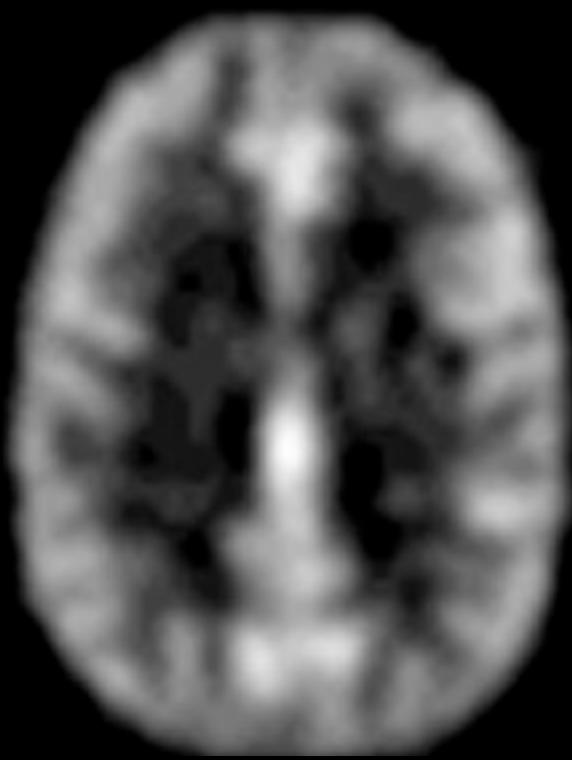
Williams, D. S., Detre, J. A., Leigh, J. S. & Koretsky, A. S. (1992) "Magnetic resonance imaging of perfusion using spin-inversion of arterial water." Proc. Natl. Acad. Sci. USA 89, 212-216.

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

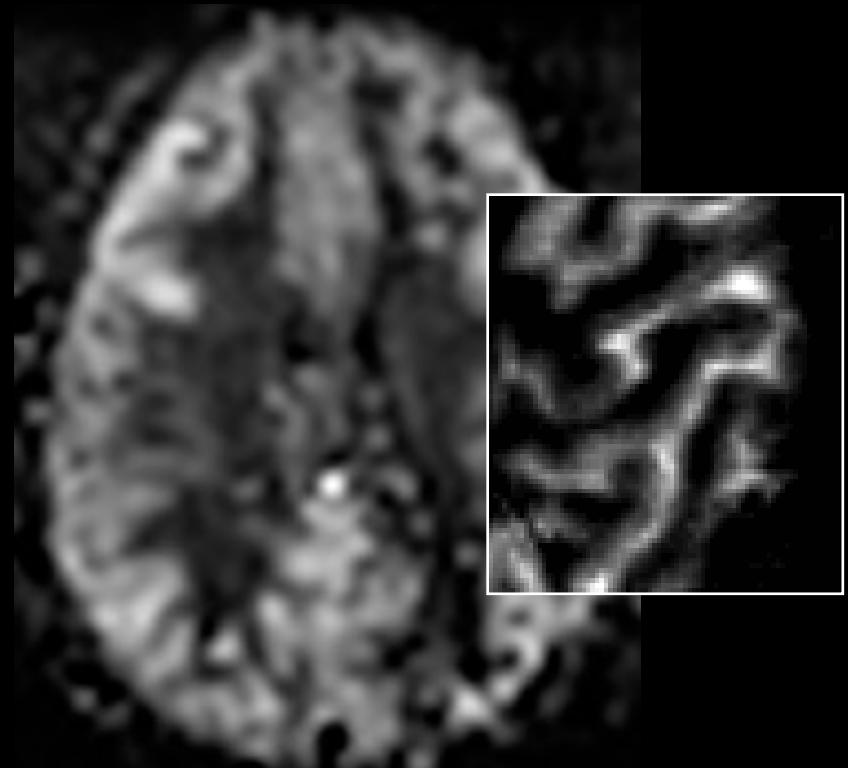
Kim, S.-G. (1995) "Quantification of relative cerebral blood flow change by flow-sensitive alternating inversion recovery (FAIR) technique: application to functional mapping." Magn. Reson. Med. 34, 293-301.

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

Comparison with Positron Emission Tomography

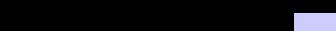


PET: H_2^{15}O



MRI: ASL

Volume



- unique information
- baseline information
- multislice trivial

- invasive
- low C / N for func.

BOLD

- highest C / N
- easy to implement
- multislice trivial
- non invasive
- highest temp. res.

- complicated signal
- no baseline info.

Perfusion

- unique information
- control over ves. size
- baseline information
- non invasive

- multislice non trivial
- lower temp. res.
- low C / N

Refinements

BOLD Contrast Interpretation

Dynamics, Paradigm Design and Processing

Applications

Refinements

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Dynamics, Paradigm Design and Processing

Applications

The Neuroscientists' Challenge:

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

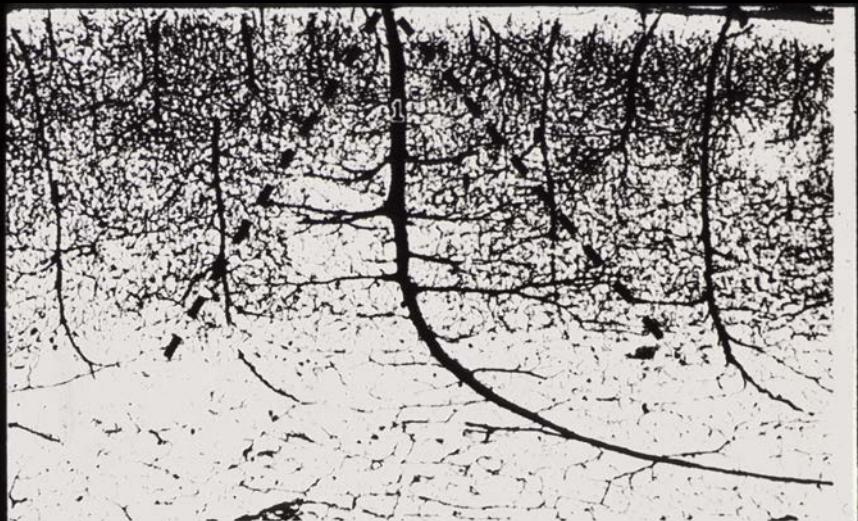
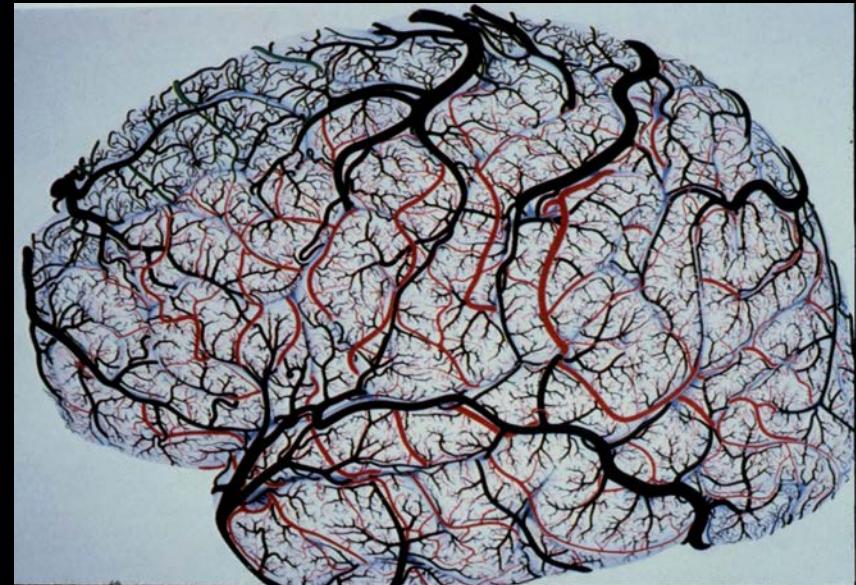
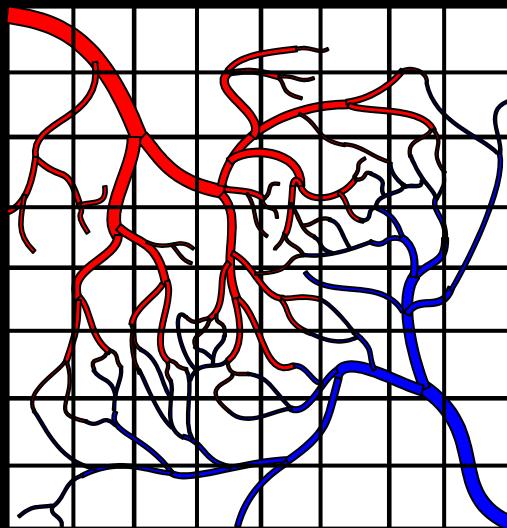


FIG. 43. Middle temporal gyrus. Female, 60 years. (1) Principal intracortical vein. The branches length regularly decreases from deep towards superficial cortical regions, thus the vascular territory of the principal vein has a conical appearance (dotted line) ($\times 28$)



Neuronal
Activation



Measured
Signal

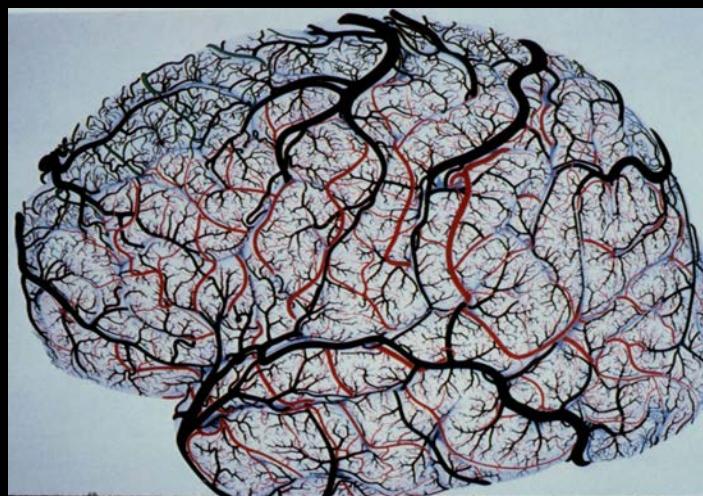
Hemodynamics

?

?

?

Noise



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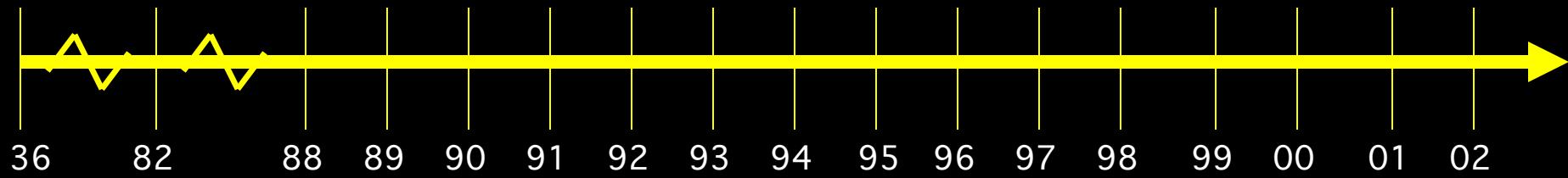
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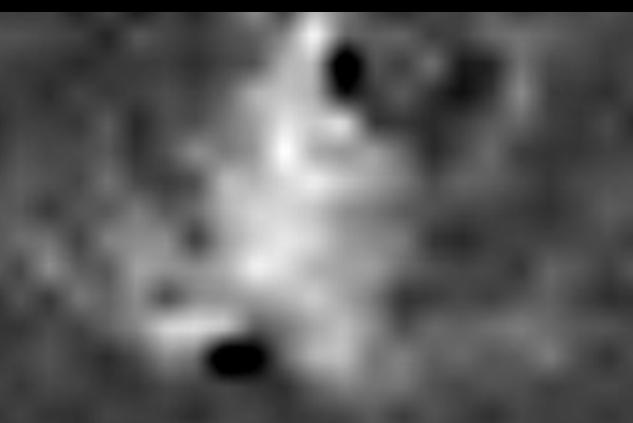
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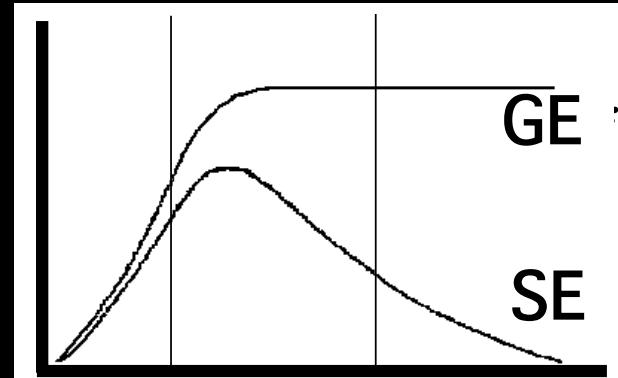
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	△ Volume-V1		Plasticity	Face recognition
				Performance prediction





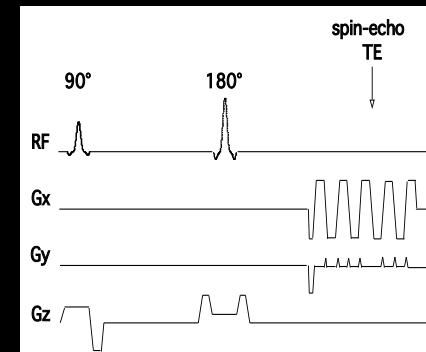
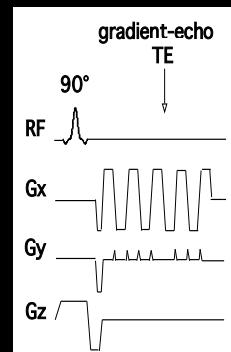
Contrast



2.5 to 3 μm 3 to 15 μm 15 to ∞ μm

compartment size

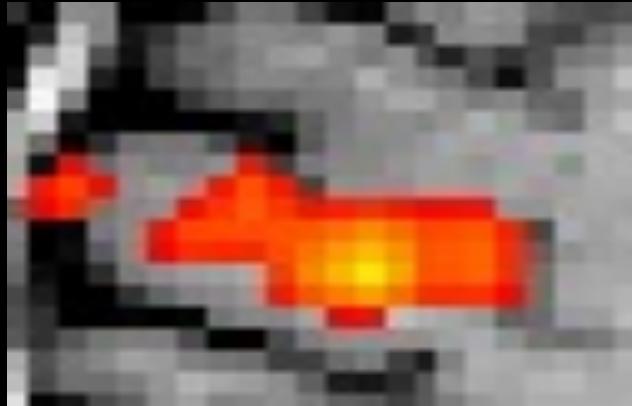
Gradient - Echo



Spin - Echo

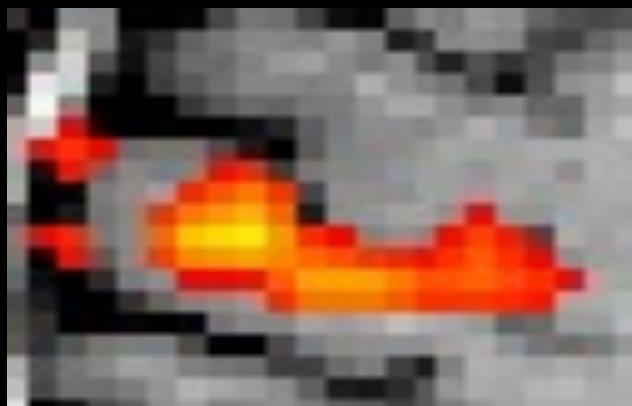
T1 - weighted

Flow weighted



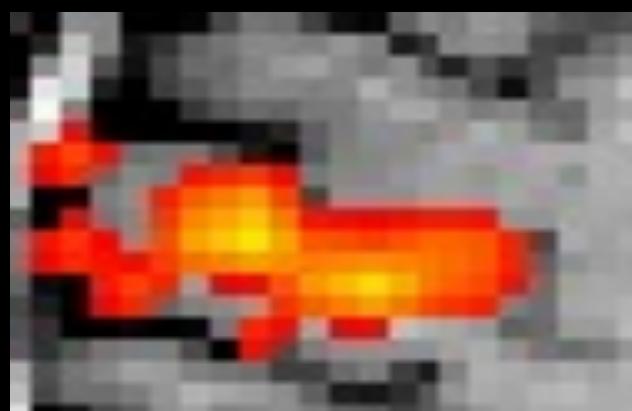
T2* weighted

BOLD weighted



T1 and T2* weighted

Flow and BOLD weighted



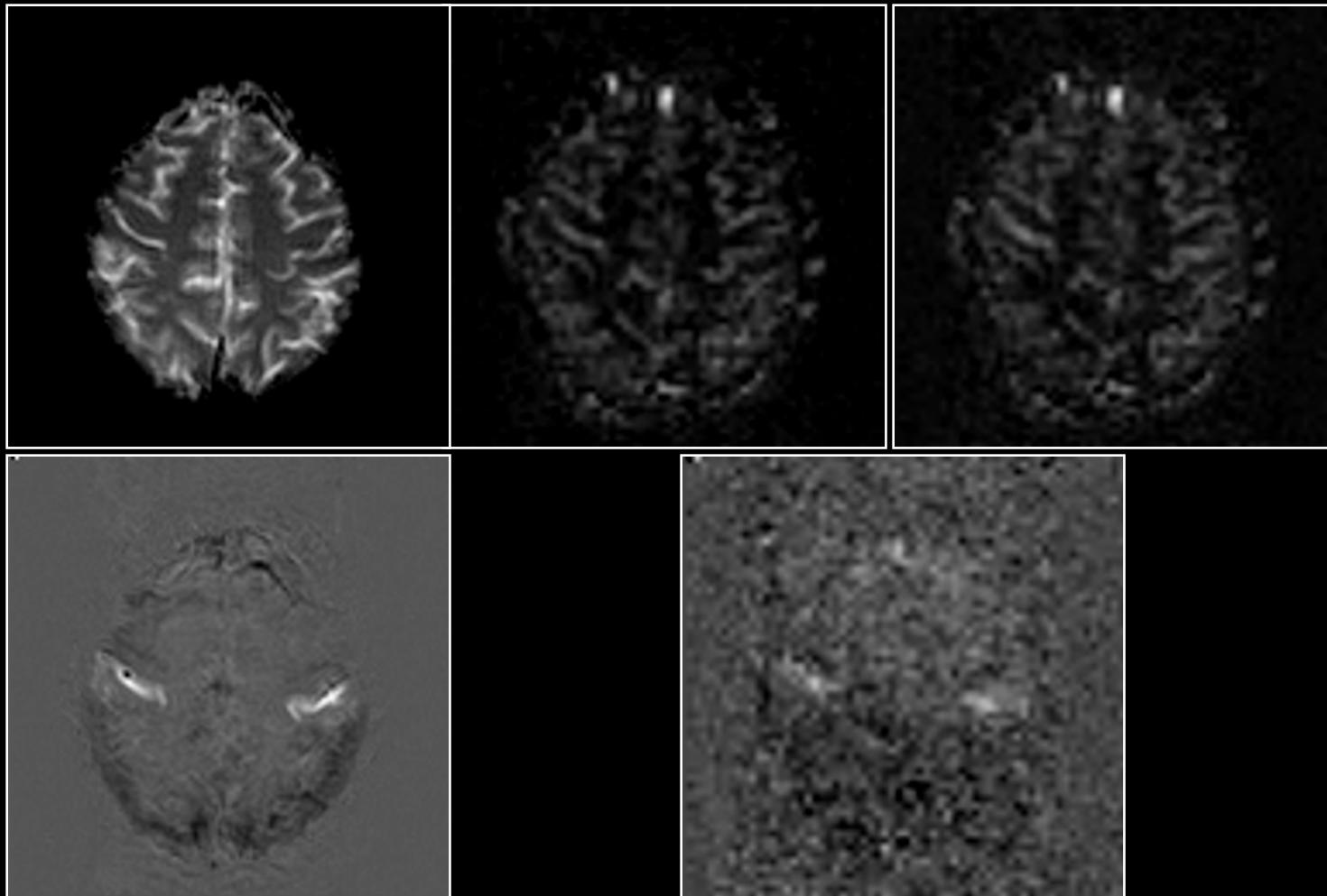
P. A. Bandettini, E. C. Wong, Echo - planar magnetic resonance imaging of human brain activation, *in* "Echo Planar Imaging: Theory, Technique, and Application" (F. Schmitt, M. Stehling, R. Turner, Eds.), p.493-530, Springer - Verlag, Berlin, 1997

BOLD

Rest

Perfusion

Activation



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.

Anatomy



BOLD



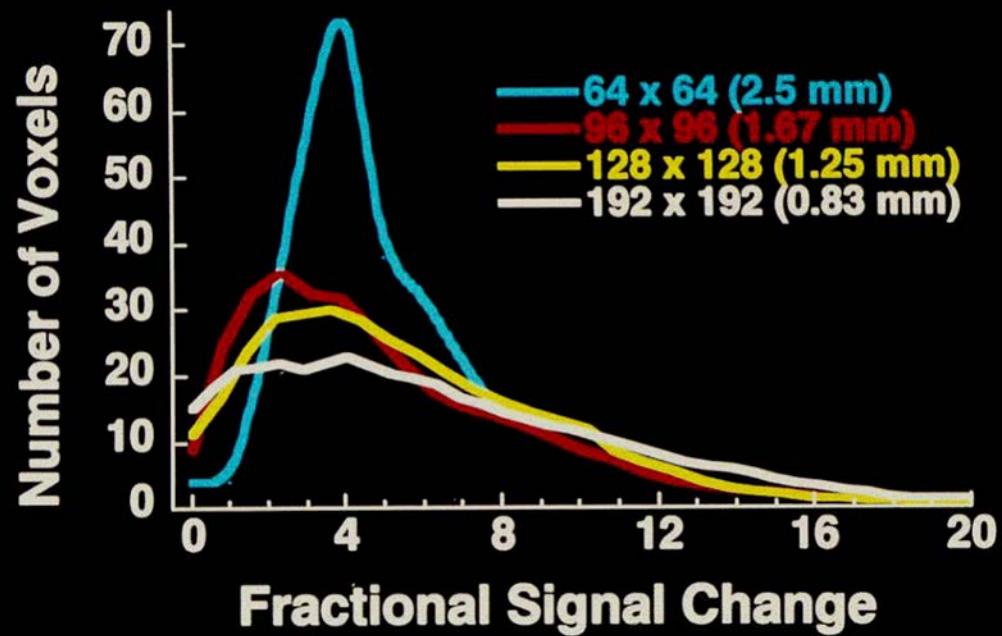
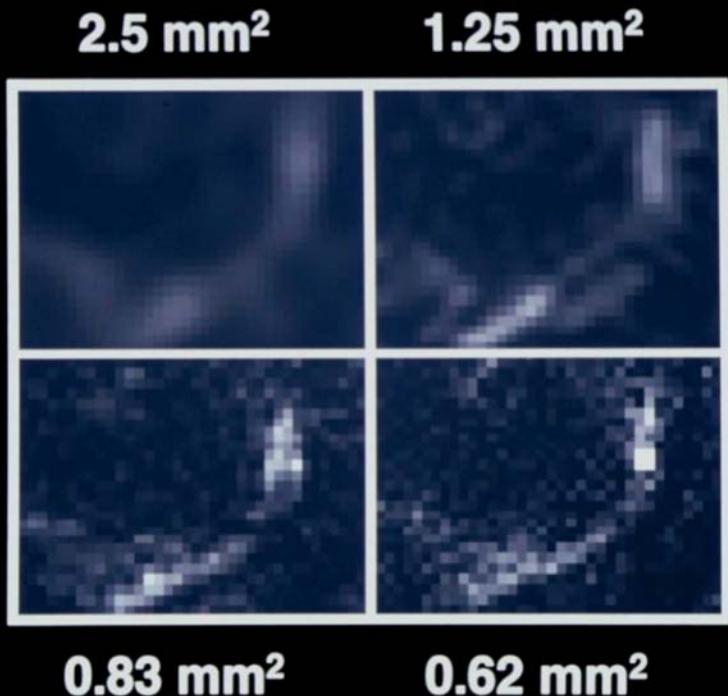
Perfusion



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.

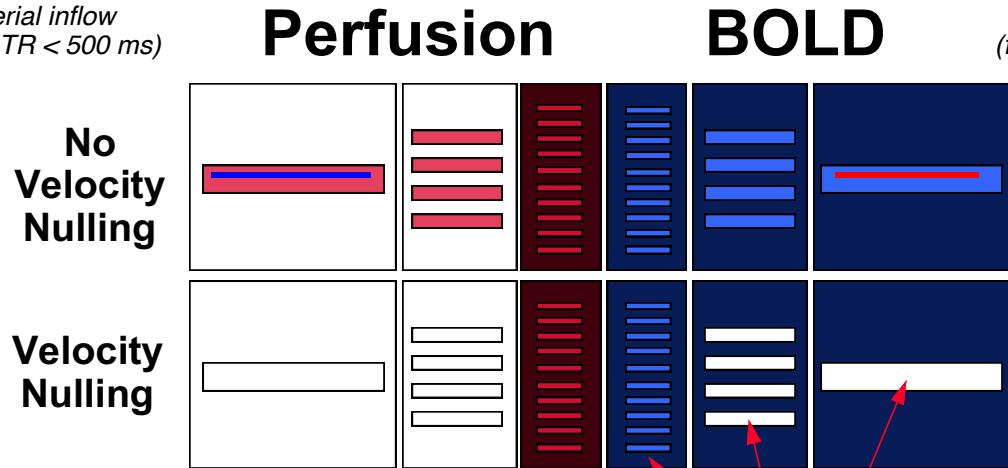
Partial k-space imaging

Fractional Signal Change



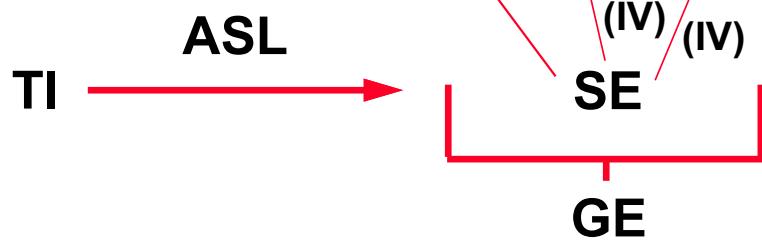
Jesmanowicz, P. A. Bandettini, J. S. Hyde, (1998) "Single shot half k-space high resolution EPI for fMRI at 3T." *Magn. Reson. Med.* 40, 754-762.

Arterial inflow
(*BOLD TR < 500 ms*)

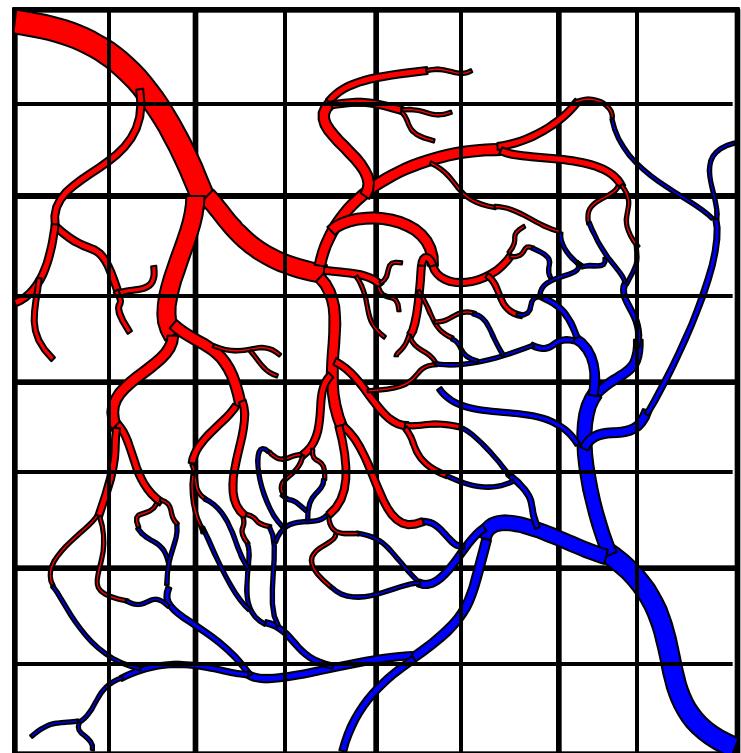


Venous inflow
(for ASL, w/ no VN)

Pulse Sequence
Sensitivity



Spatial
Heterogeneity



Refinements

BOLD Contrast Interpretation

Dynamics, Paradigm Design and Processing

Applications

Technology

MRI	EPI	1.5T,3T, 4T	EPI on Clin. Syst. Nav. pulses	Diff. tensor	Mg ⁺	7T Venography SENSE
		Local Human Head Gradient Coils		Real time fMRI		
		ASL		Quant. ASL	Z-shim	
		Spiral EPI		Dynamic IV volume		
		BOLD		Multi-shot fMRI	Simultaneous ASL and BOLD	Baseline Susceptibility Current Imaging?

Methodology

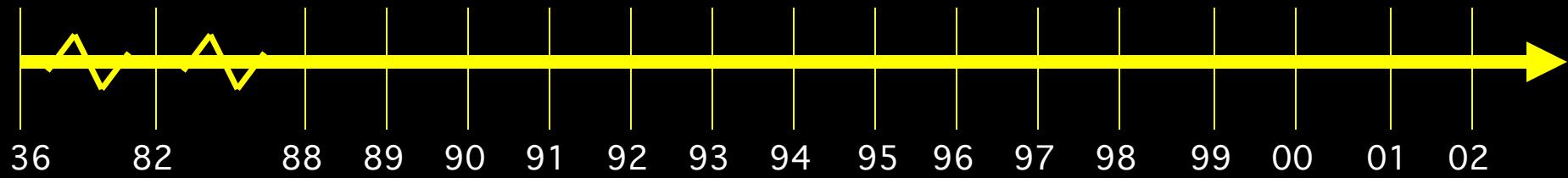
Baseline Volume	IVIM	Correlation Analysis	Motion Correction Multi-Modal Mapping Free-behavior Designs	CO ₂ Calibration
		Parametric Design		Multi-Modal Mapping
		Surface Mapping		
		Phase Mapping		Free-behavior Designs
		Linear Regression		Mental Chronometry
		Event-related		Deconvolution

Interpretation

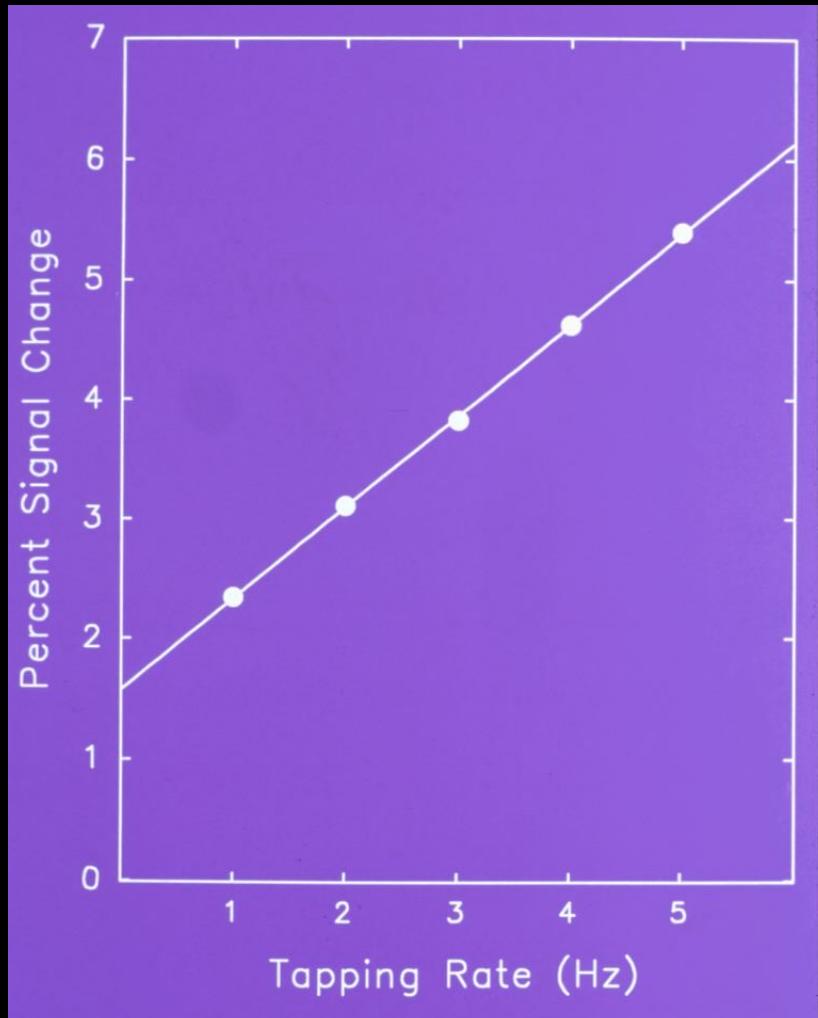
Blood T2	Hemoglobin	BOLD models	PET correlation	
		B ₀ dep.	IV vs EV	ASL vs. BOLD
		TE dep	Pre-undershoot	PSF of BOLD
		SE vs. GE	Resolution Dep.	Extended Stim.
		NIRS Correlation	Post-undershoot	Linearity
		Veins	CO ₂ effect	Metab. Correlation
		Inflow	Fluctuations	Optical Im. Correlation
			Balloon Model	Electrophys. correlation

Applications

Volume - Stroke	△ Volume-V1	Complex motor	Memory	Emotion
		Language		
		Imagery		
		Motor learning		
		Children		
		Tumor vasc.		Drug effects
		Ocular Dominance		
		Clinical Populations		
		Performance prediction		
		Face recognition		
		Plasticity		

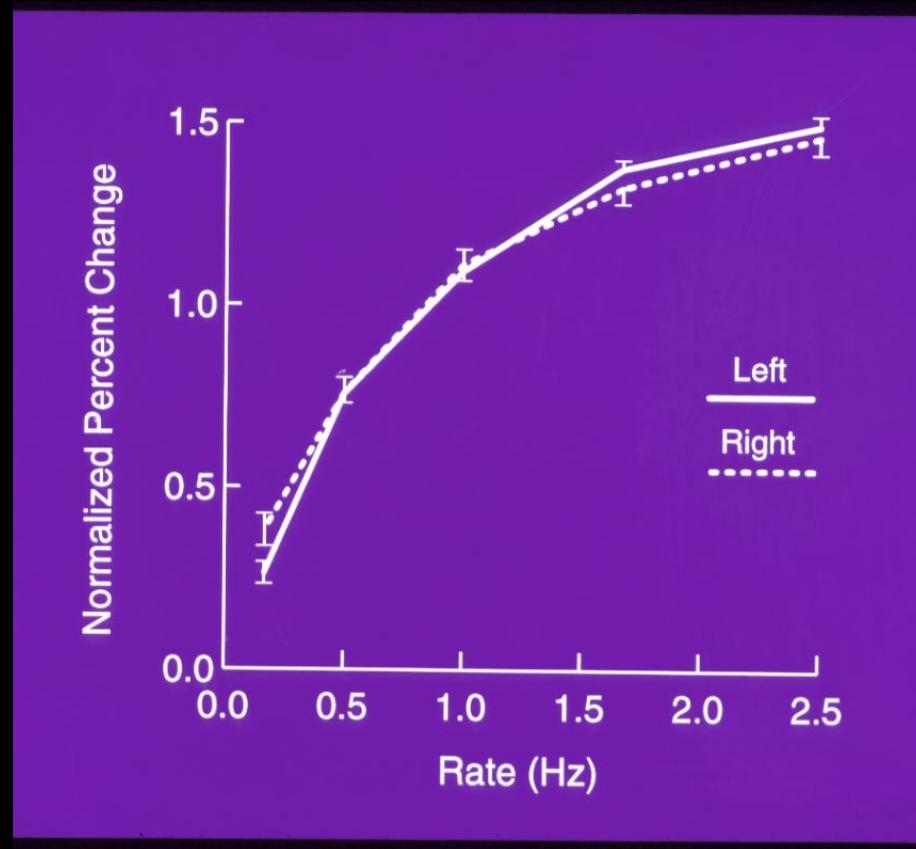


Motor Cortex



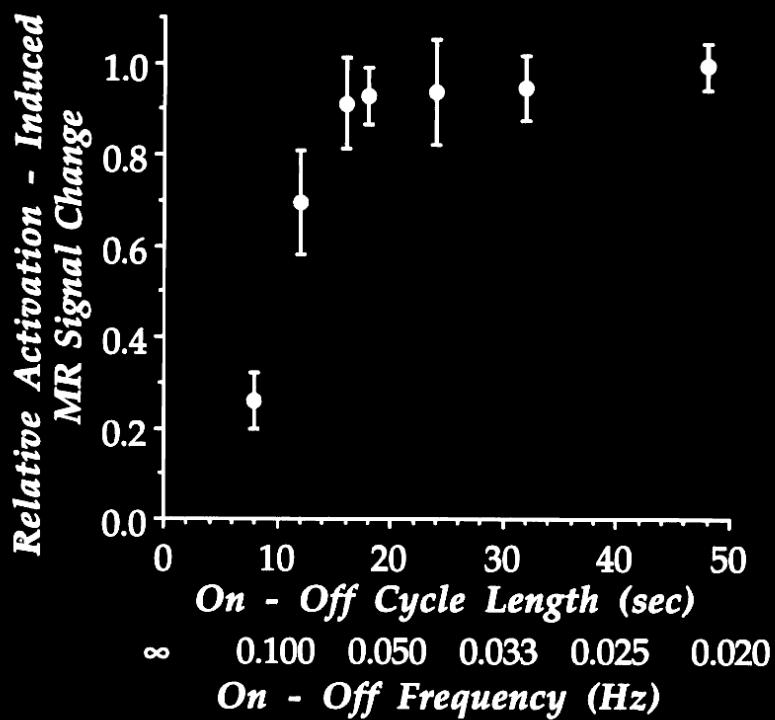
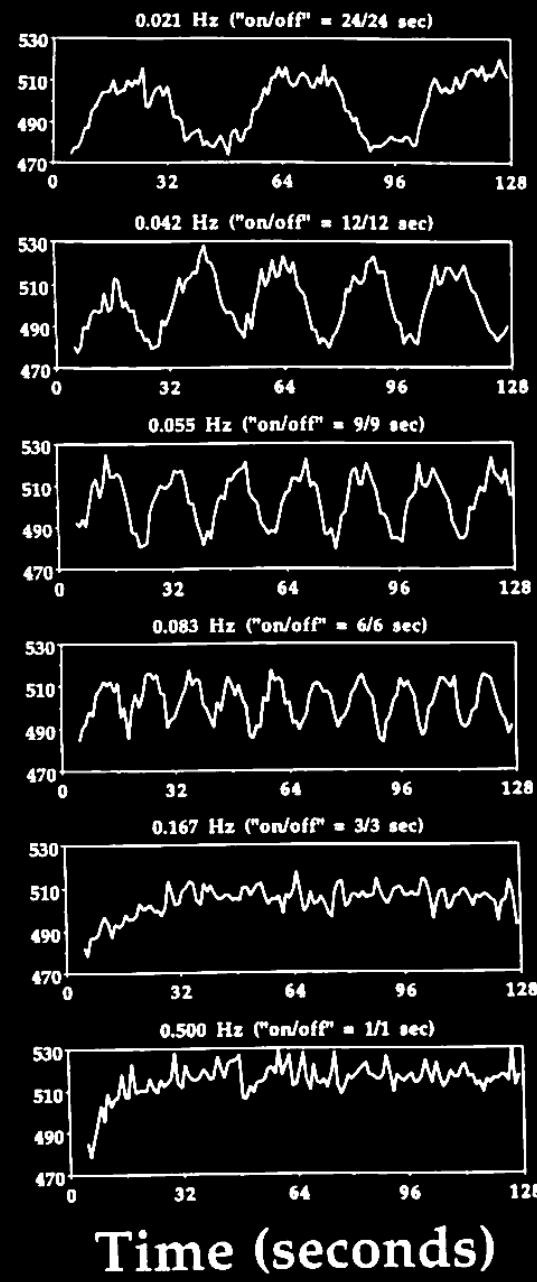
S. M. Rao et al, (1996) “Relationship between finger movement rate and functional magnetic resonance signal change in human primary motor cortex.” *J. Cereb. Blood Flow and Met.* 16, 1250-1254.

Auditory Cortex



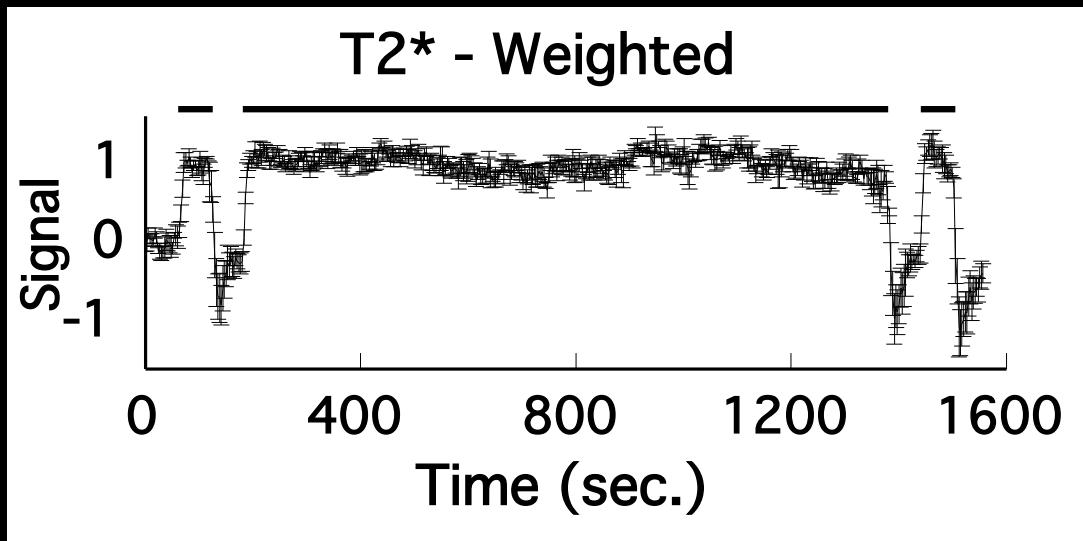
J. R. Binder, et al, (1994). “Effects of stimulus rate on signal response during functional magnetic resonance imaging of auditory cortex.” *Cogn. Brain Res.* 2, 31-38

MRI Signal

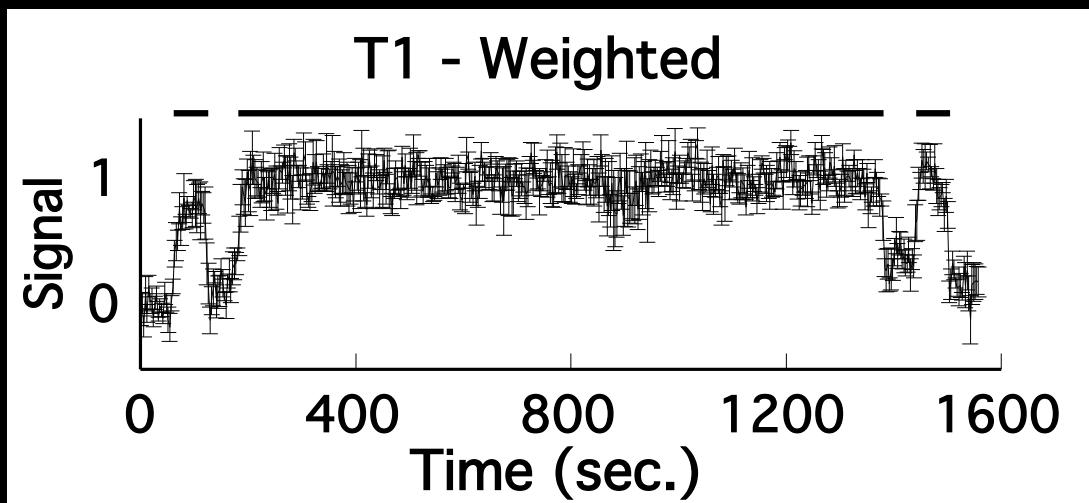


P. A. Bandettini, Functional MRI
temporal resolution in "Functional
MRI" (C. Moonen, and P. Bandettini.,
Eds.), p. 205-220, Springer - Verlag.,
1999.

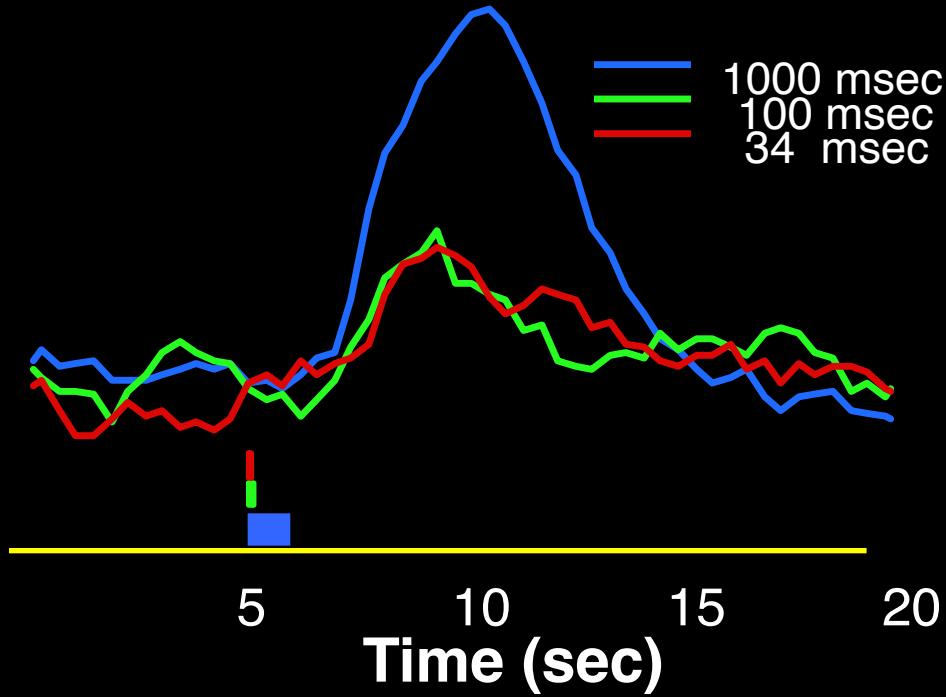
BOLD



Flow

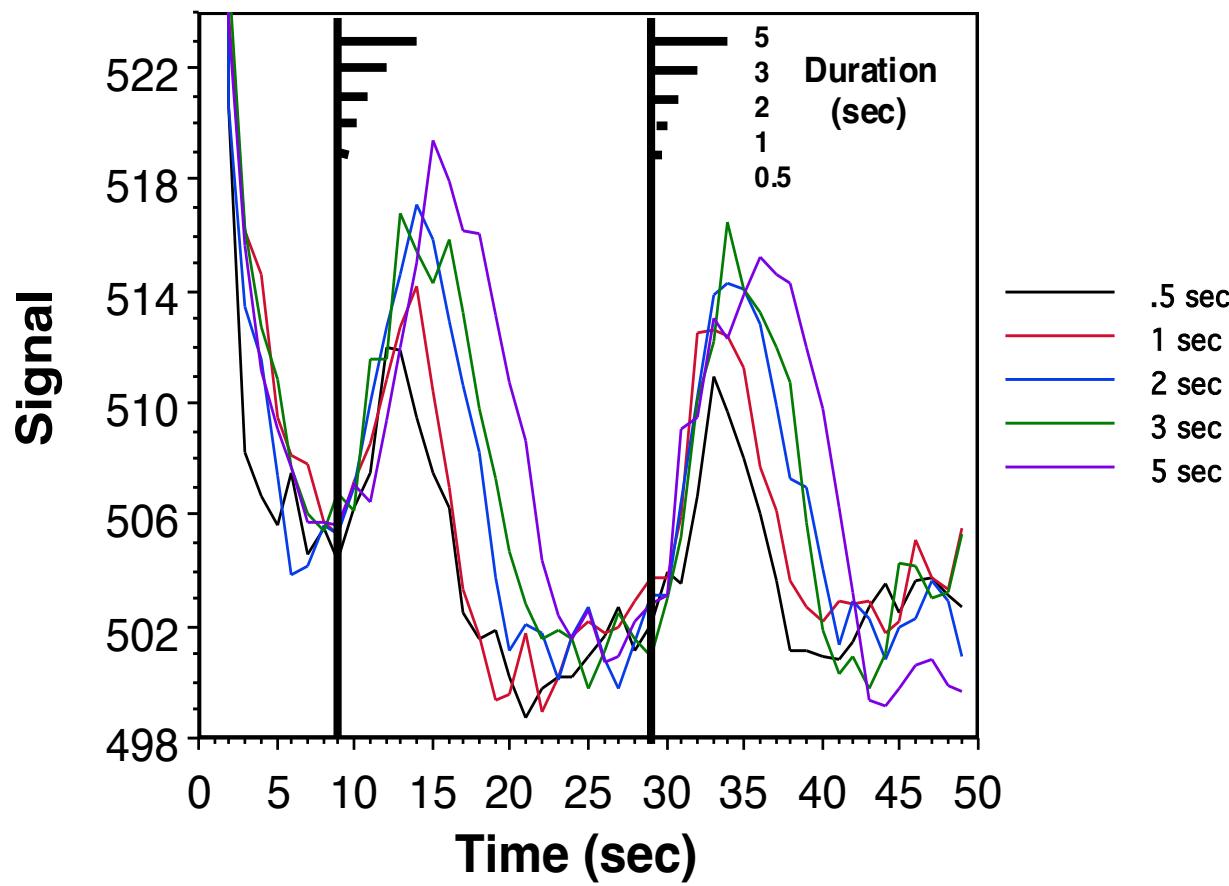


P. A. Bandettini, K. K. Kwong, T. L. Davis, R. B. H. Tootell, E. C. Wong, P. T. Fox, J. W. Belliveau, R. M. Weisskoff, B. R. Rosen, (1997). “Characterization of cerebral blood oxygenation and flow changes during prolonged brain activation.” *Human Brain Mapping* 5, 93-109.

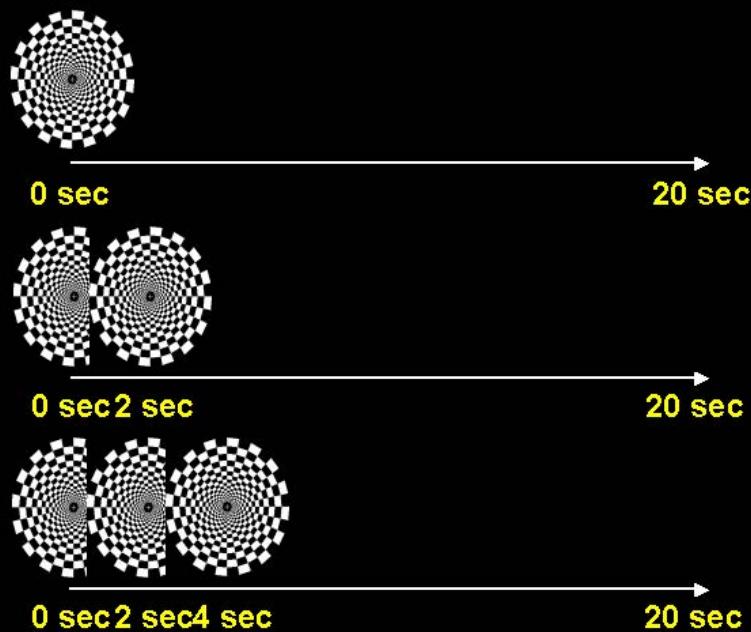


R. L. Savoy, et al., Pushing the temporal resolution of fMRI: studies of very brief visual stimuli, onset variability and asynchrony, and stimulus-correlated changes in noise [oral], 3'rd Proc. Soc. Magn. Reson., Nice, p. 450. (1995).

Motor Cortex



Bandettini, et al., The functional dynamics of blood oxygenation level contrast in the motor cortex, 12'th Proc. Soc. Magn. Reson. Med., New York, p. 1382. (1993).

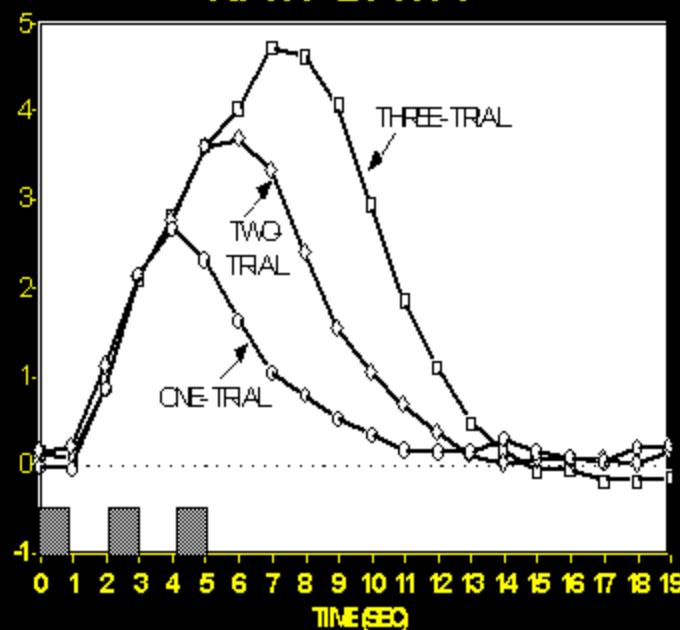


♦ Human Brain Mapping 5:329–340(1997)

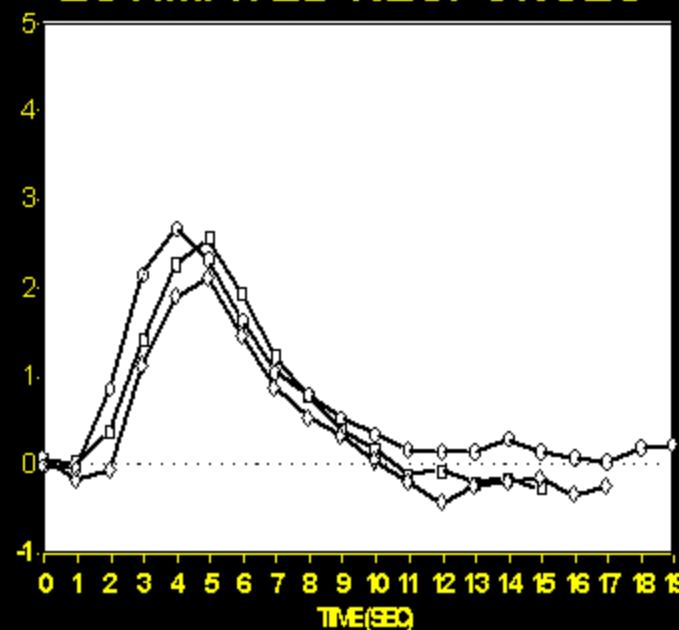
Selective Averaging of Rapidly Presented Individual Trials Using fMRI

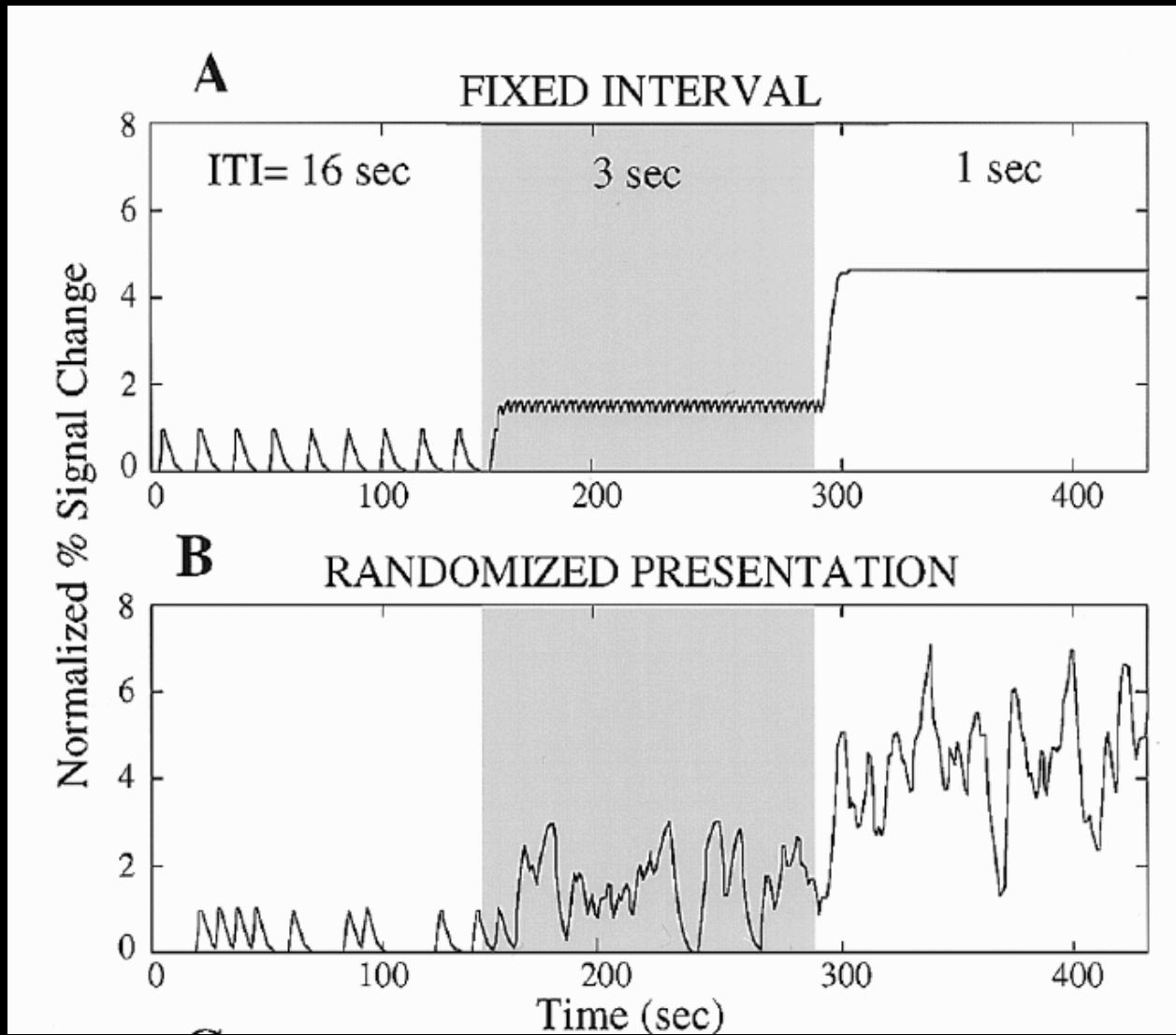
Anders M. Dale* and Randy L. Buckner

RAW DATA

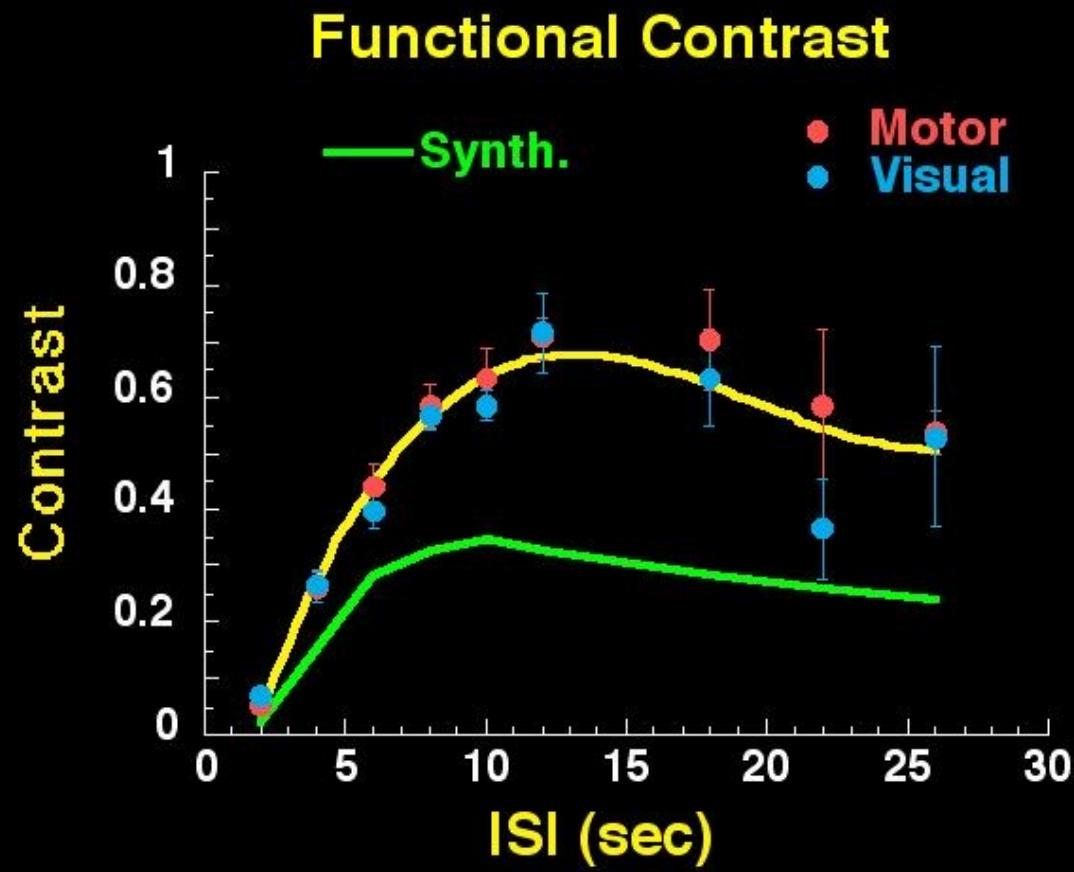


ESTIMATED RESPONSES





M.A. Burock et al. *NeuroReport*, 9, 3735-9 (1998)



(Block design = 1)

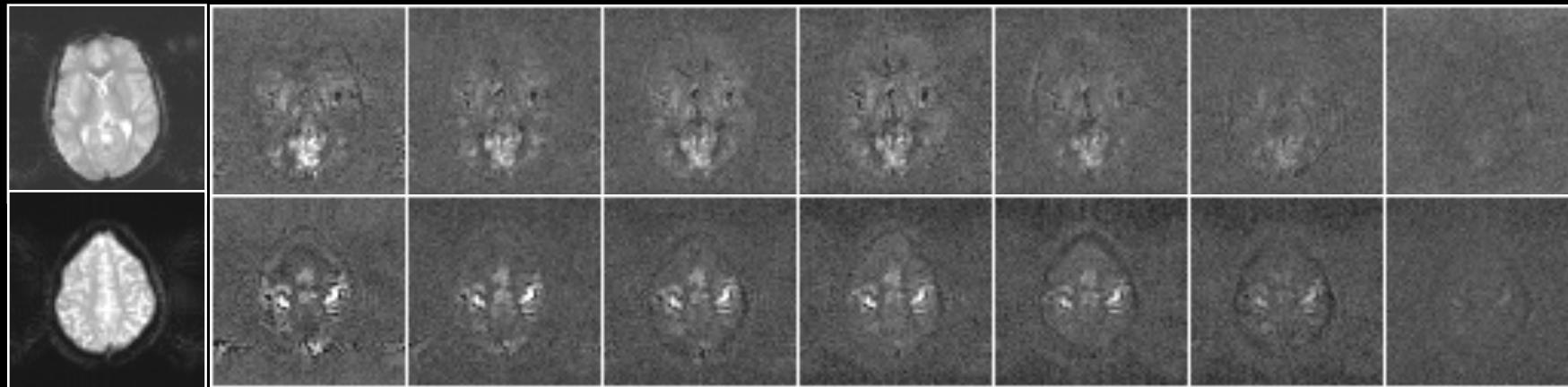
P. A. Bandettini, R. W. Cox. Functional contrast in constant interstimulus interval event - related fMRI: theory and experiment. *Magn. Reson. Med.* 43: 540-548 (2000).

Contrast to Noise Images

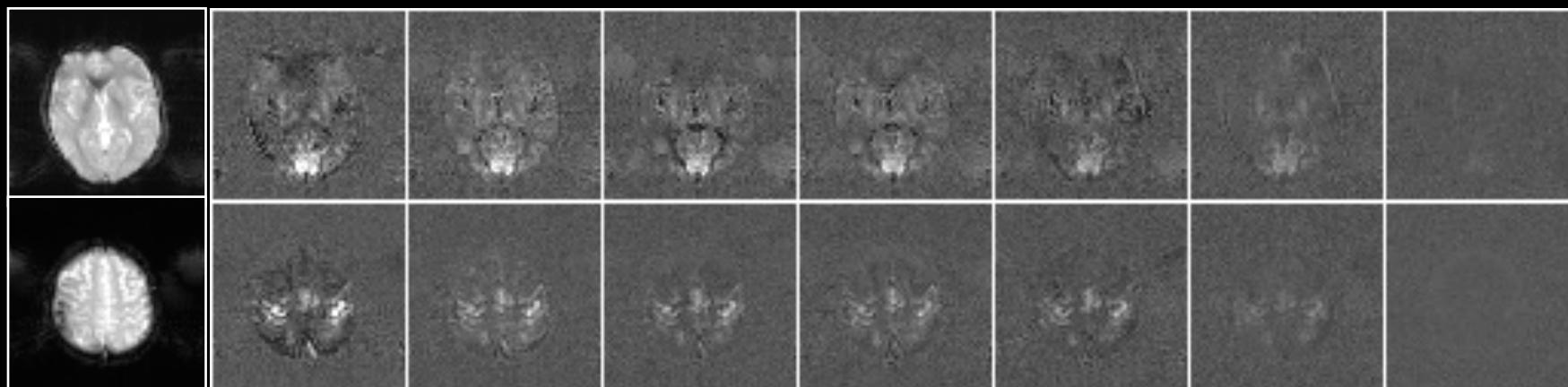
(ISI, SD)

20, 20 12, 2 10, 2 8, 2 6, 2 4, 2 2, 2

S1



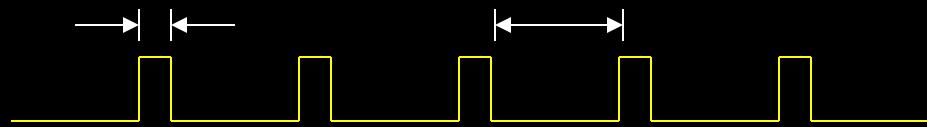
S2



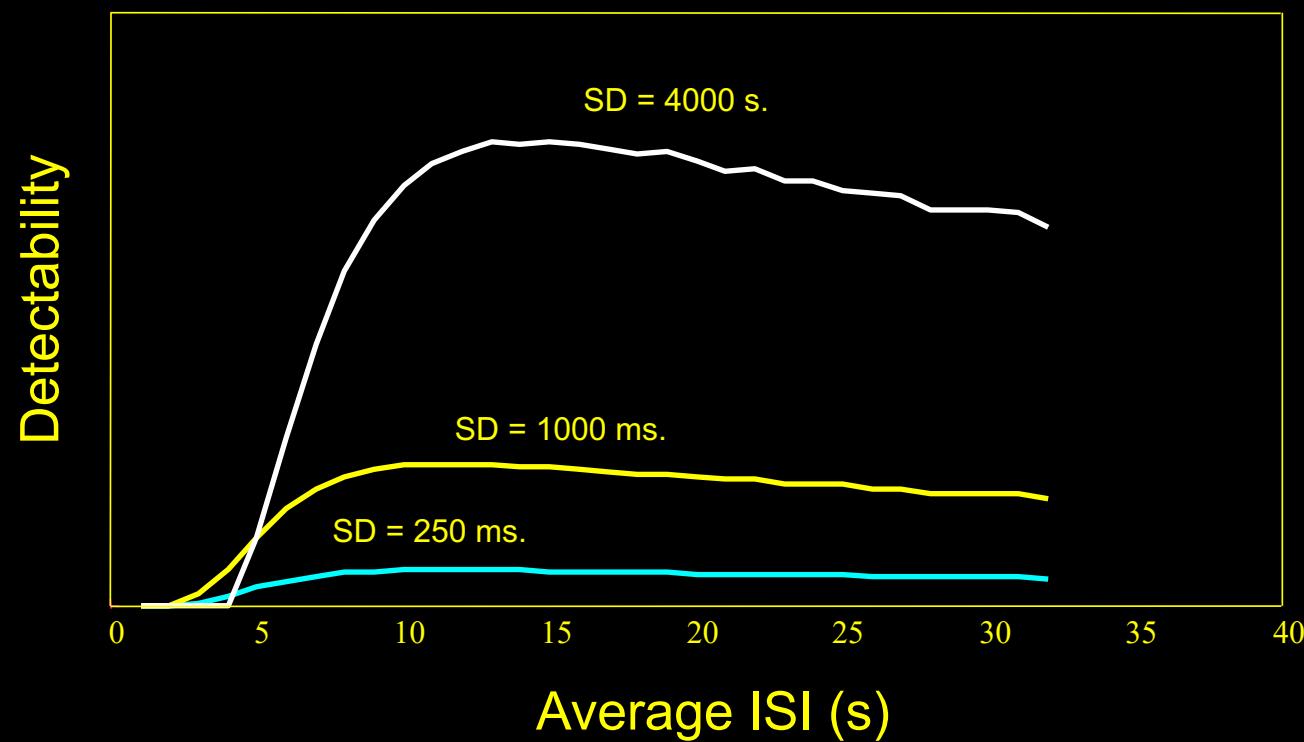
P. A. Bandettini, R. W. Cox. Functional contrast in constant interstimulus interval event - related fMRI: theory and experiment. *Magn. Reson. Med.* 43: 540-548 (2000).

Detectability – constant ISI

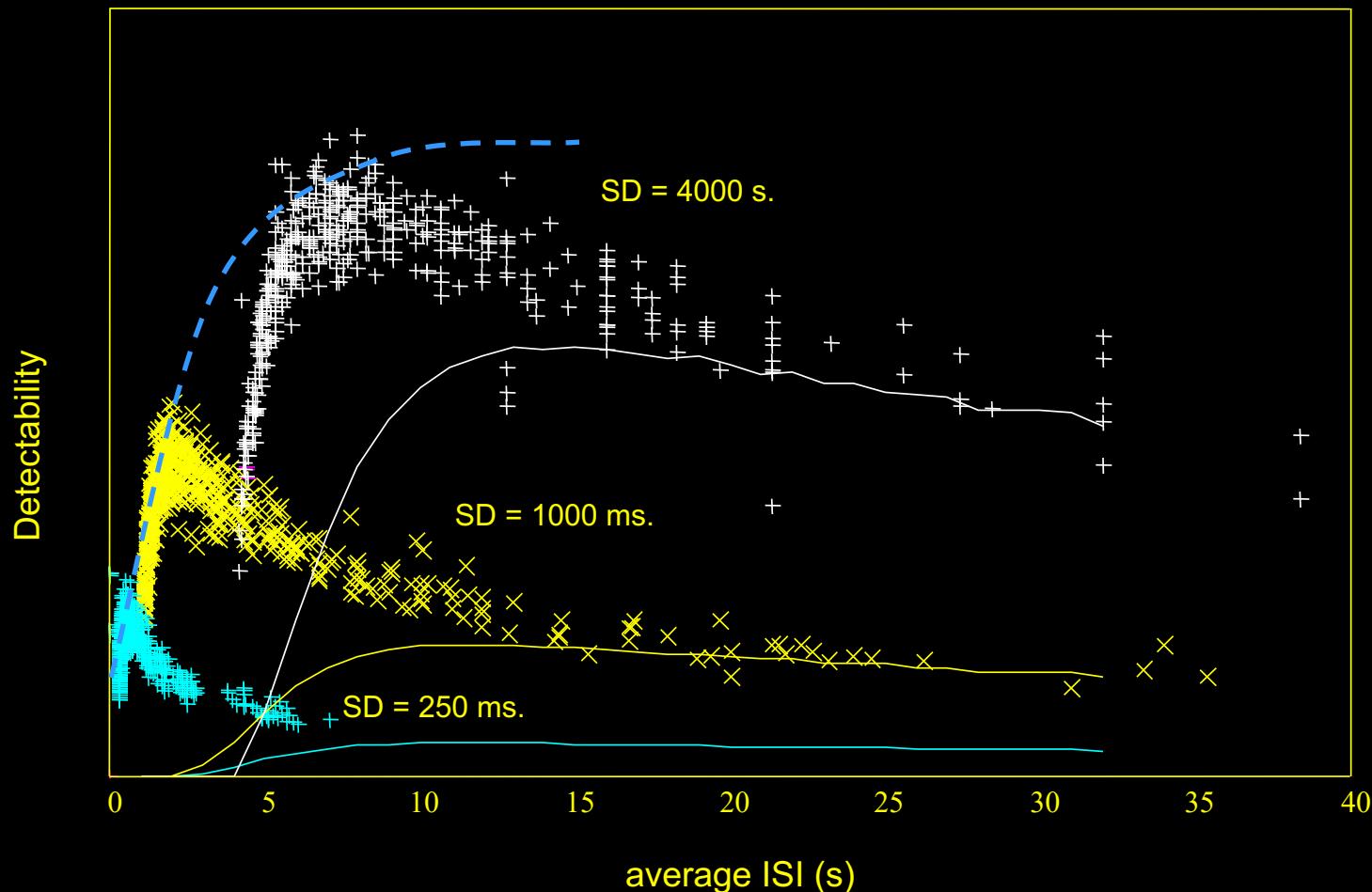
SD – stimulus duration



ISI – inter-stimulus interval

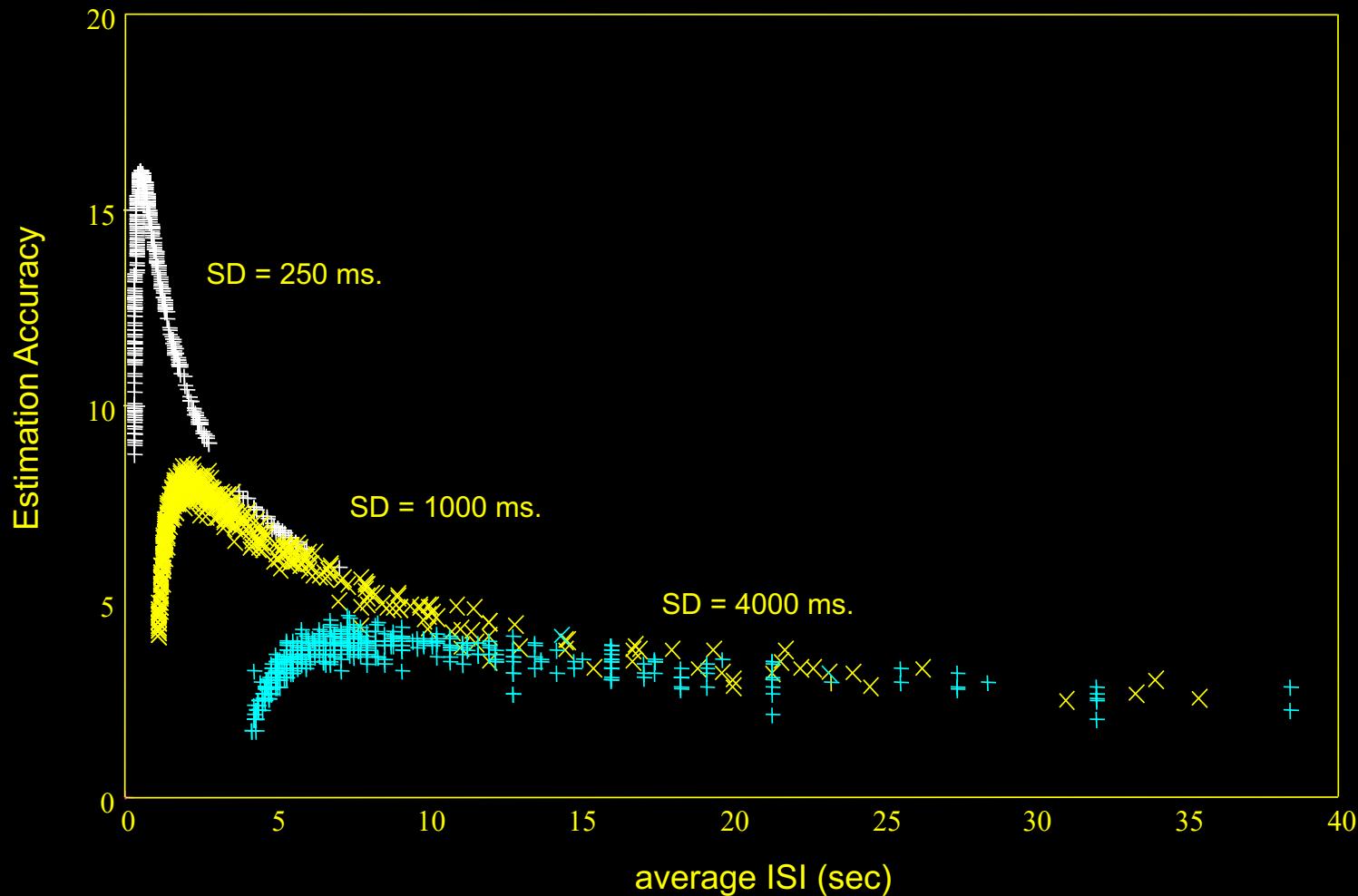


Detectability vs. Average ISI



R. M. Birn, R. W. Cox, P. A. Bandettini, Detection versus estimation in Event-Related fMRI: choosing the optimal stimulus timing. *NeuroImage* 15: 262-264, (2002).

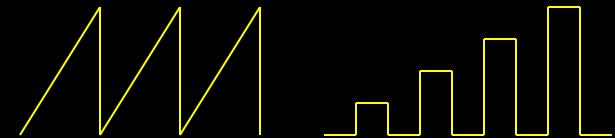
Estimation accuracy vs. average ISI



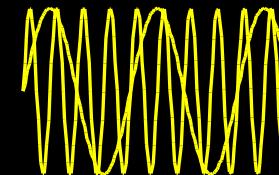
R. M. Birn, R. W. Cox, P. A. Bandettini, Detection versus estimation in Event-Related fMRI: choosing the optimal stimulus timing. *NeuroImage* 15: 262-264, (2002).

Neuronal Activation Input Strategies

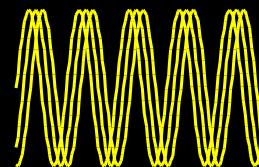
1. Block Design



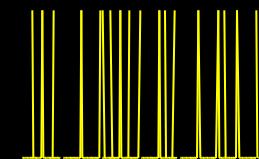
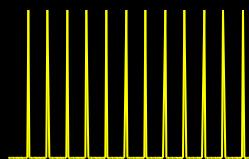
2. Parametric Design



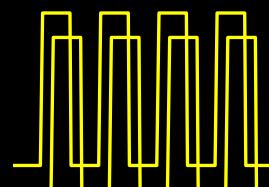
3. Frequency Encoding



4. Phase Encoding



5. Event Related

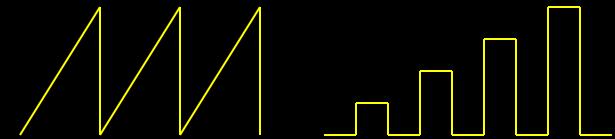


6. Orthogonal Design

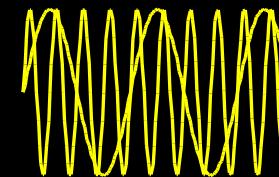
7. Free Behavior Design

Neuronal Activation Input Strategies

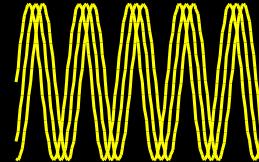
1. Block Design



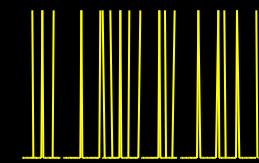
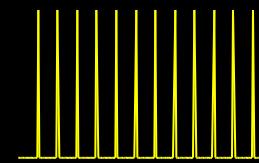
2. Parametric Design



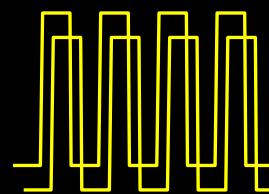
3. Frequency Encoding



4. Phase Encoding



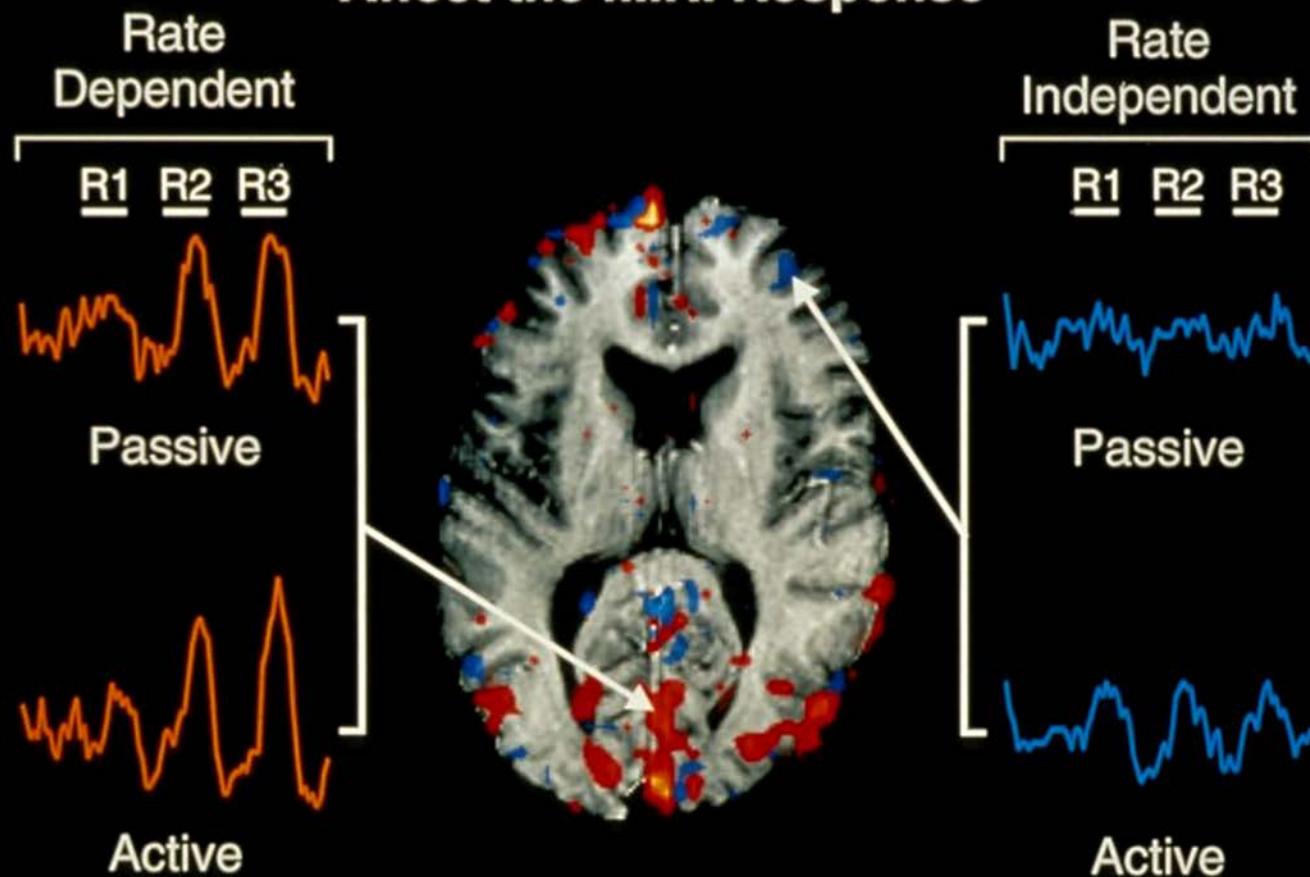
5. Event Related



6. Orthogonal Design

7. Free Behavior Design

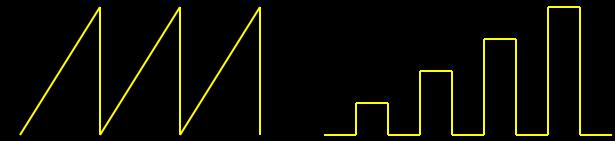
Both the Task and Presentation Rate Affect the fMRI Response



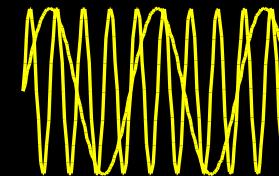
E. A. DeYoe, P. A. Bandettini, J. Nietz, D. Miller, P. Winas, Methods for functional magnetic resonance imaging (fMRI). *J. Neuroscience Methods* 54, 171-187 (1994).

Neuronal Activation Input Strategies

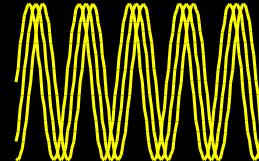
1. Block Design



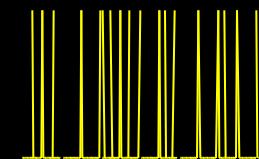
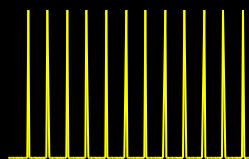
2. Parametric Design



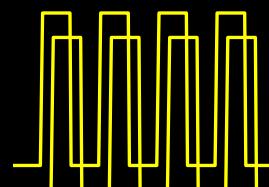
3. Frequency Encoding



4. Phase Encoding

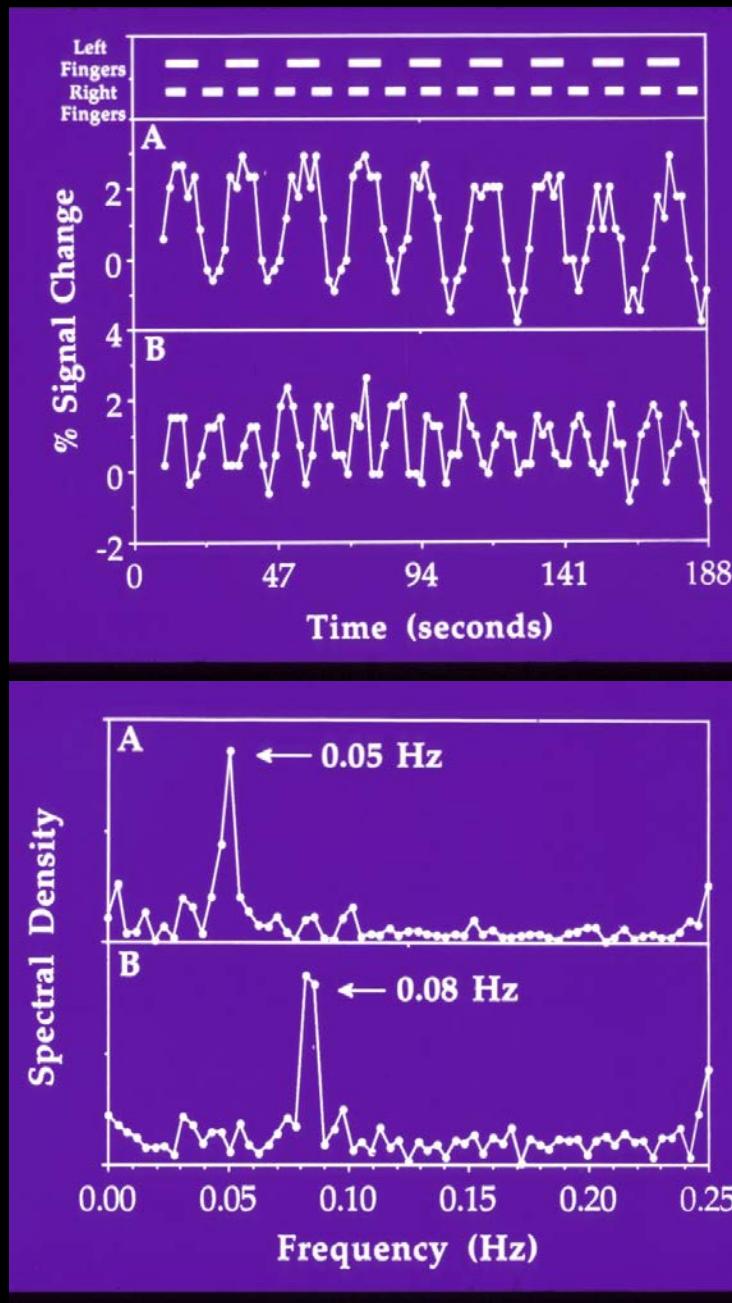
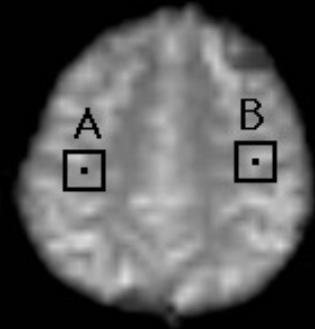


5. Event Related



6. Orthogonal Design

7. Free Behavior Design



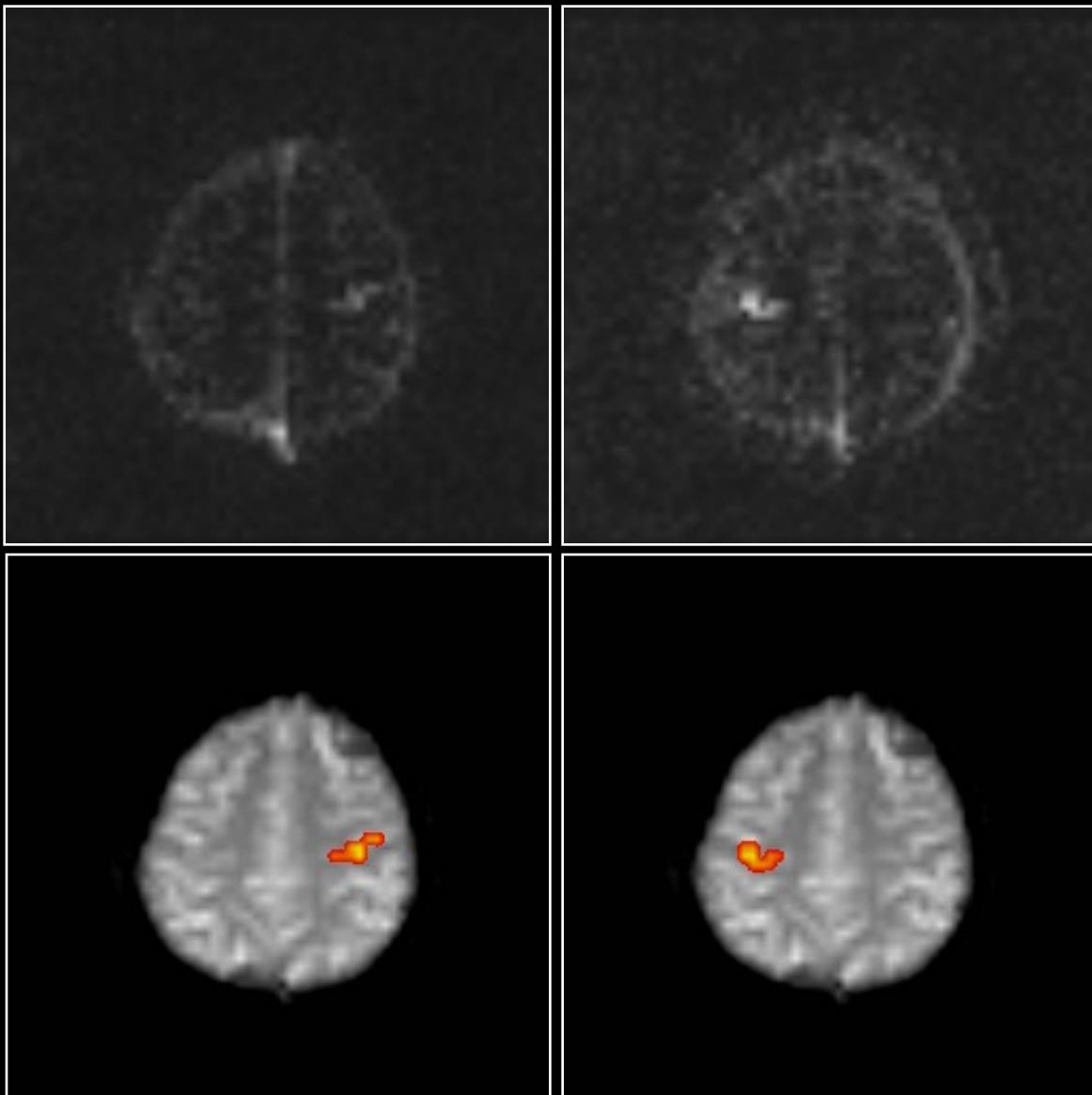
P. A. Bandettini, A. Jesmanowicz, E. C. Wong, J. S. Hyde, Processing strategies for time-course data sets in functional MRI of the human brain. *Magn. Reson. Med.* 30, 161-173 (1993).

0.08 Hz

0.05 Hz

**spectral
density**

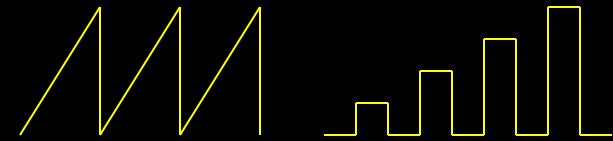
**c.c. > 0.5
with spectra**



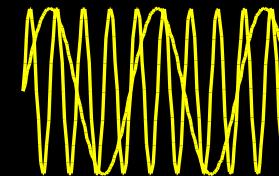
P. A. Bandettini, A. Jesmanowicz, E. C. Wong, J. S. Hyde, Processing strategies for time-course data sets in functional MRI of the human brain. *Magn. Reson. Med.* 30, 161-173 (1993).

Neuronal Activation Input Strategies

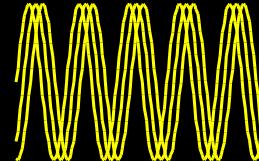
1. Block Design



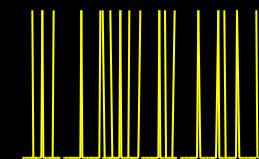
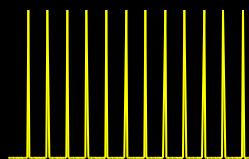
2. Parametric Design



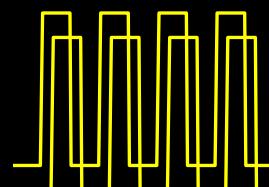
3. Frequency Encoding



4. Phase Encoding



5. Event Related

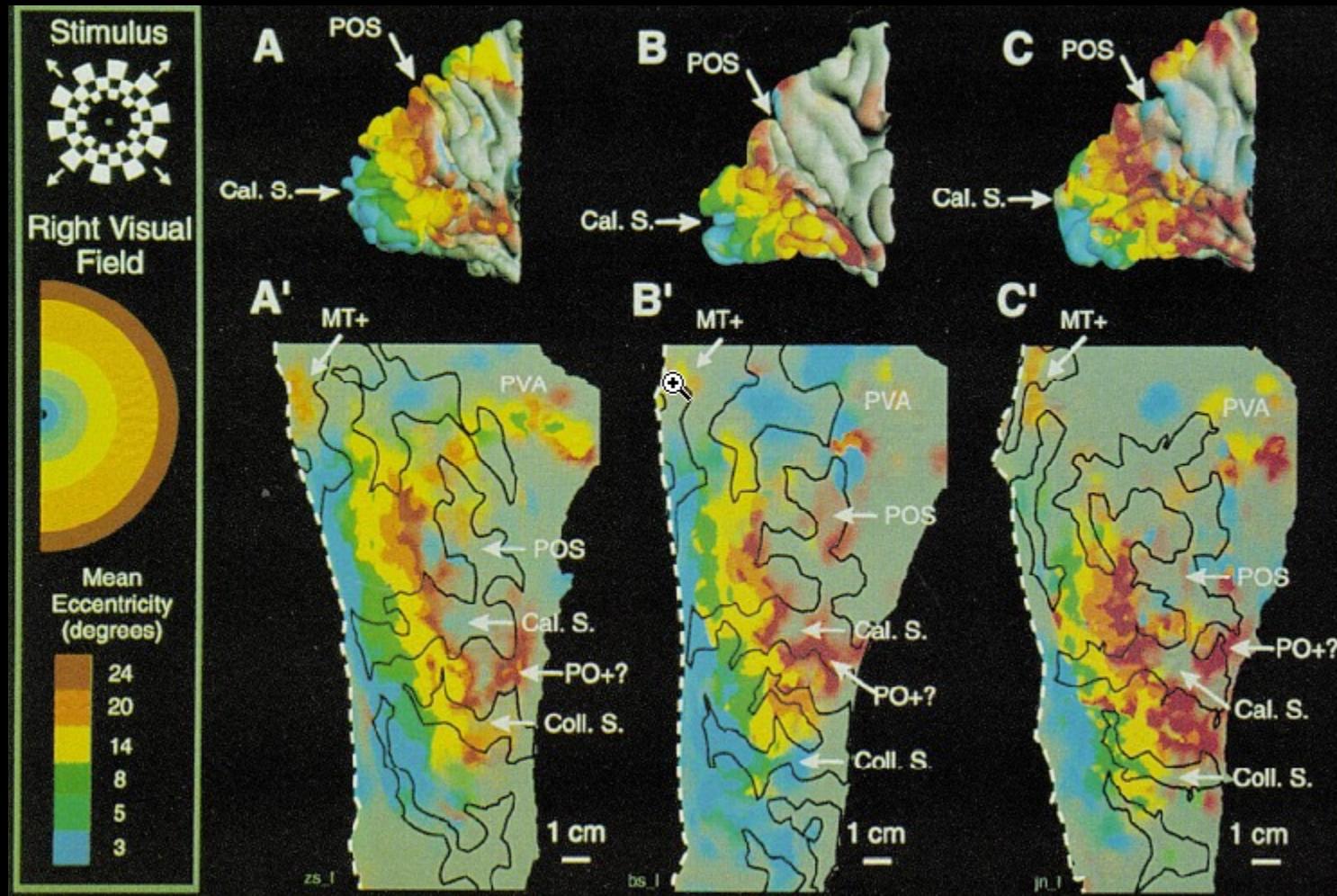


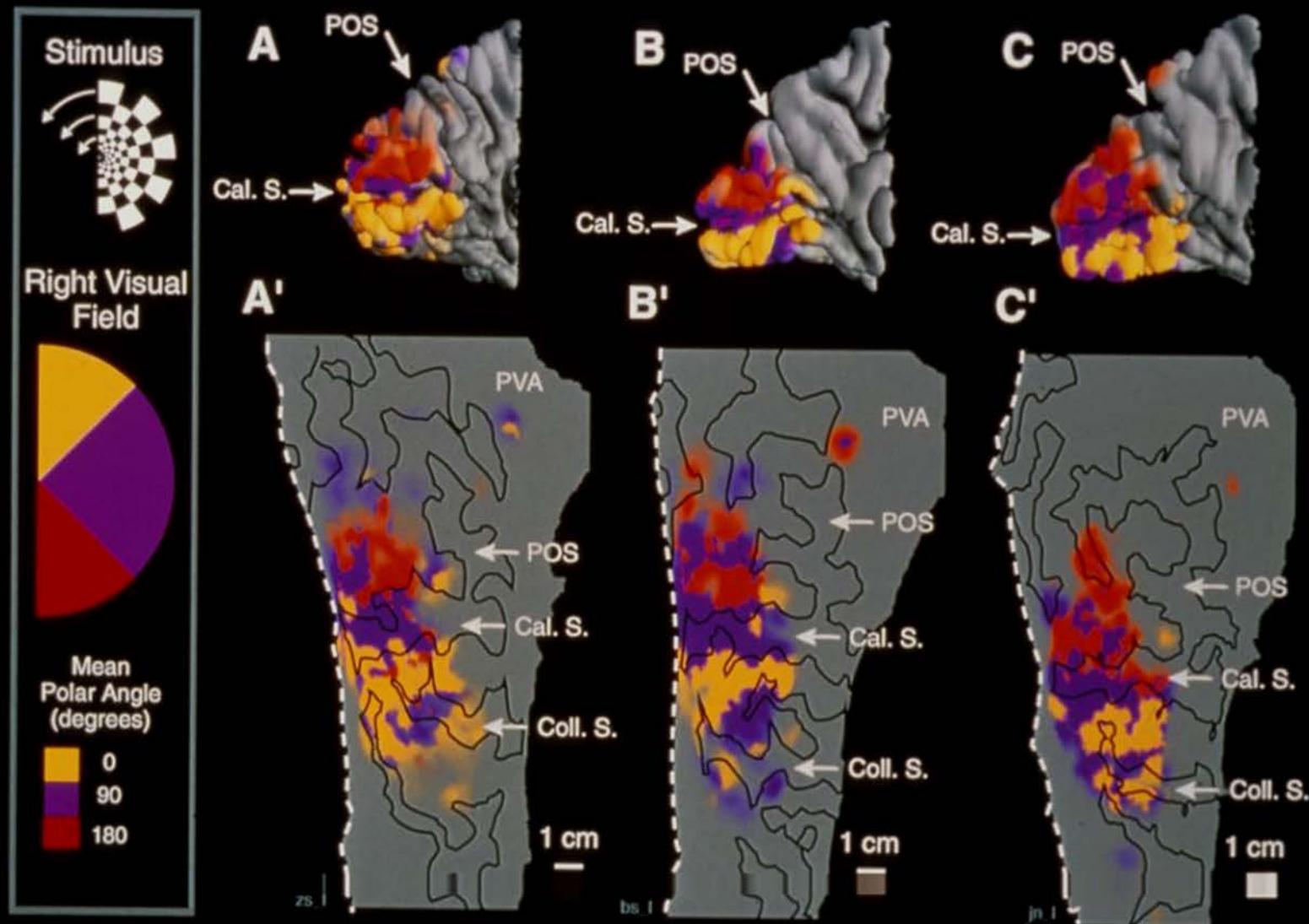
6. Orthogonal Design

7. Free Behavior Design

Mapping striate and extrastriate visual areas in human cerebral cortex

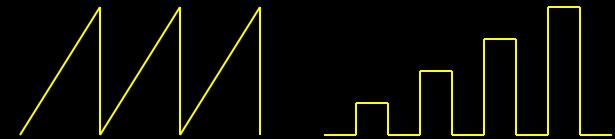
EDGAR A. DEYOE*, GEORGE J. CARMAN†, PETER BANDETTINI‡, SETH GLICKMAN*, JON WIESER*, ROBERT COX§,
DAVID MILLER¶, AND JAY NEITZ*



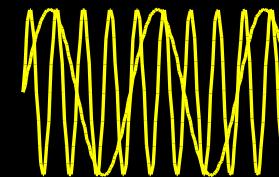


Neuronal Activation Input Strategies

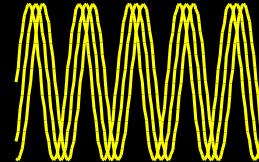
1. Block Design



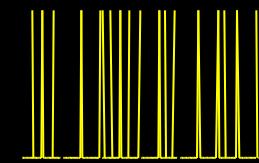
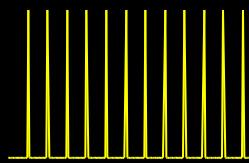
2. Parametric Design



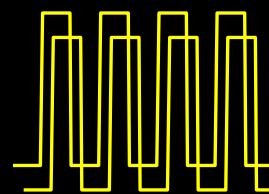
3. Frequency Encoding



4. Phase Encoding



5. Event Related



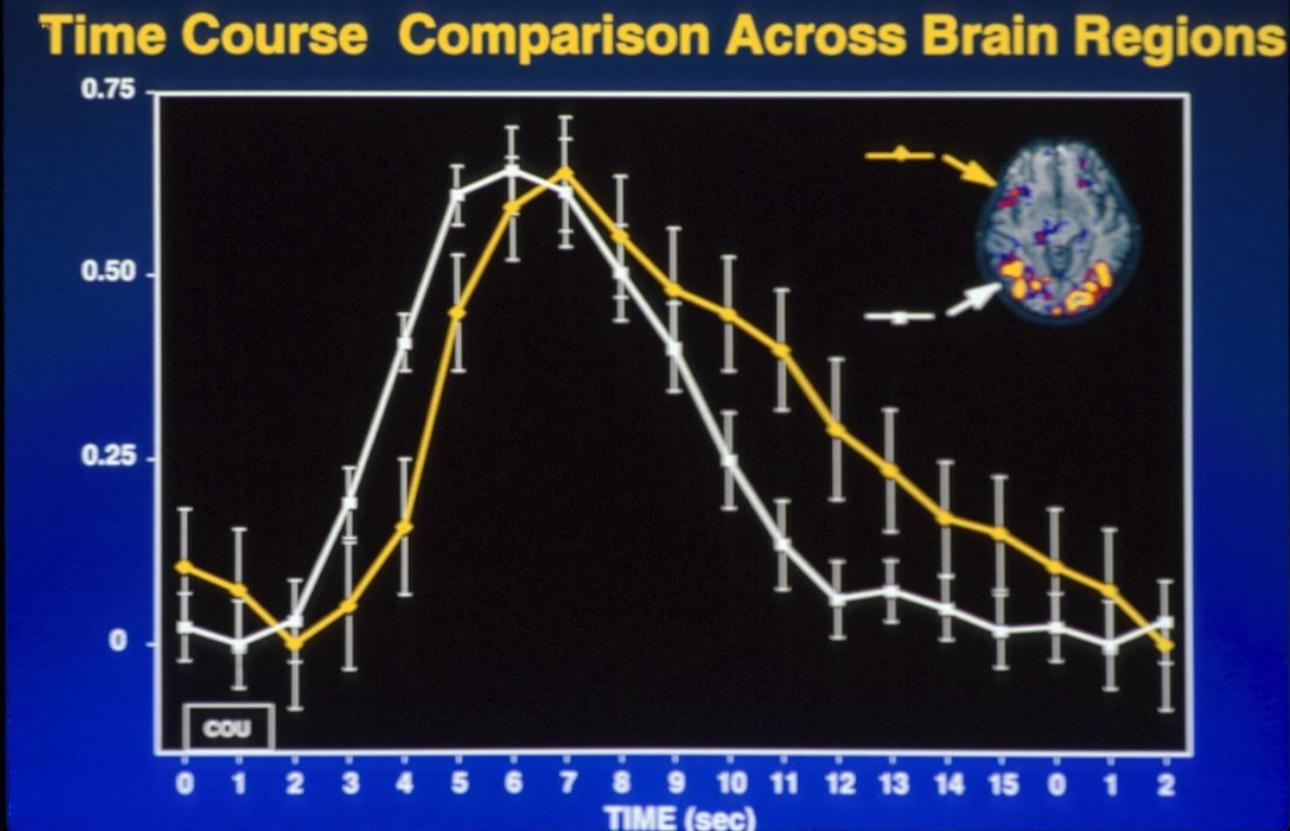
6. Orthogonal Design

7. Free Behavior Design

Detection of cortical activation during averaged single trials of a cognitive task using functional magnetic resonance imaging

(neuroimaging/single trial/language/prefrontal)

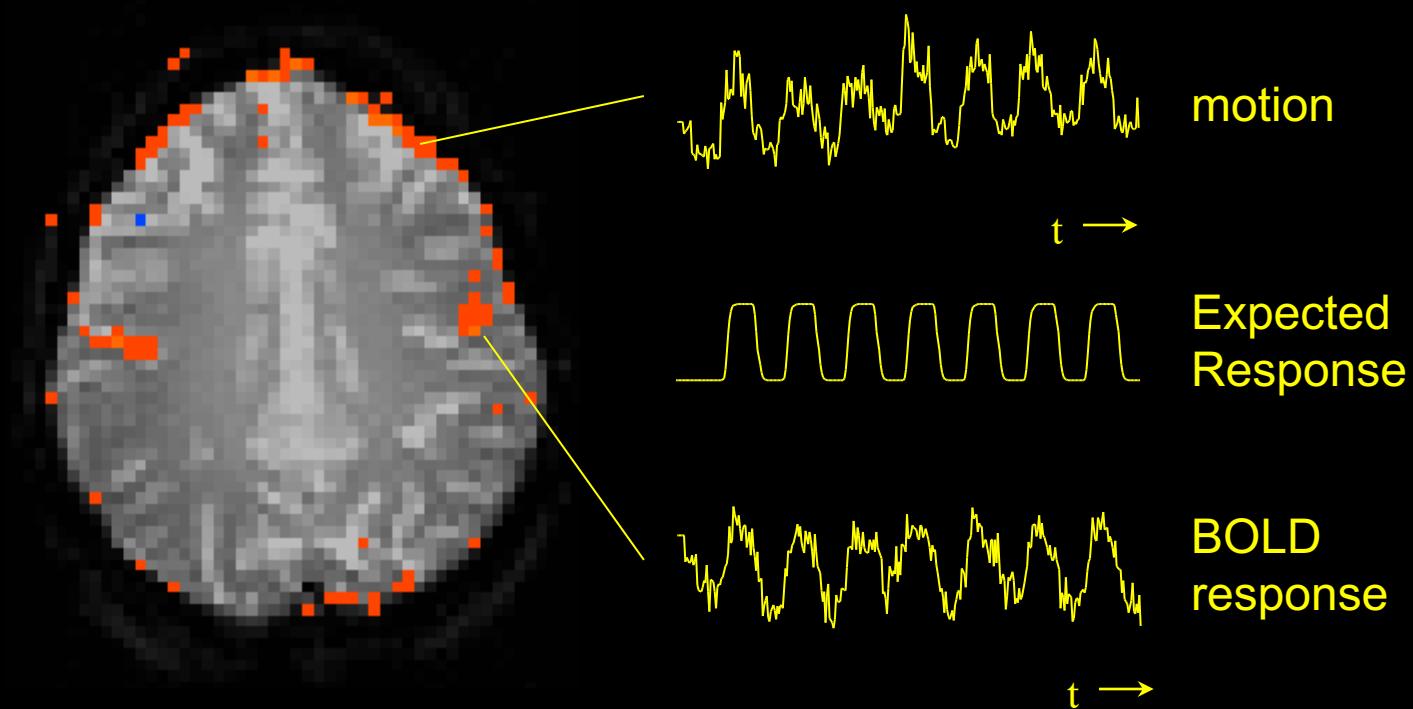
RANDY L. BUCKNER^{†‡§¶||}, PETER A. BANDETTINI^{†‡}, KATHLEEN M. O'CRAVEN^{†||}, ROBERT L. SAVOY^{†||},
STEVEN E. PETERSEN^{*++††}, MARCUS E. RAICHLE^{§++††}, AND BRUCE R. ROSEN^{†‡}



Event Related Advantages

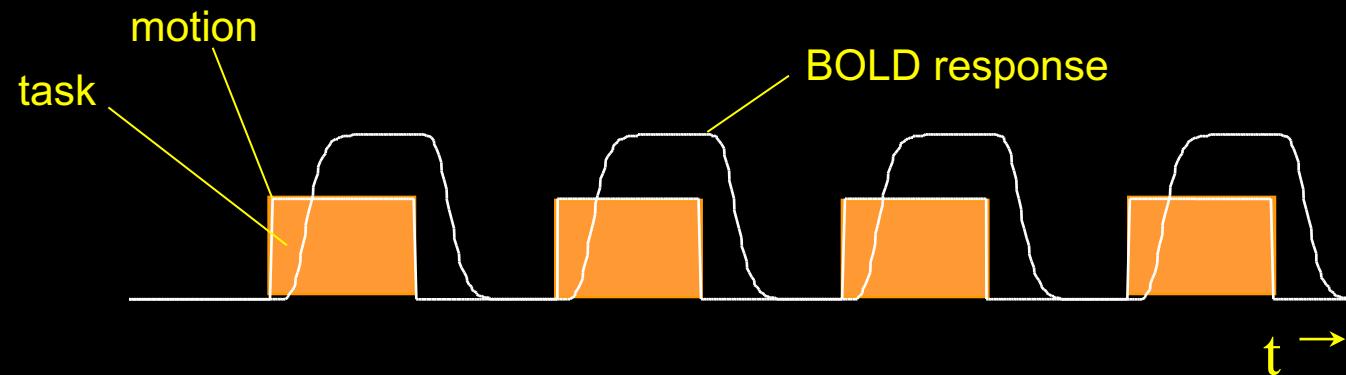
- Task Randomization
- Post acquisition, Performance-based, data binning
- Natural presentation
- Reduction of habituation effects
- Overt responses
- Reduction of scanner noise effects
- More precise estimation of hemodynamic responses

Speaking - Blocked Trial

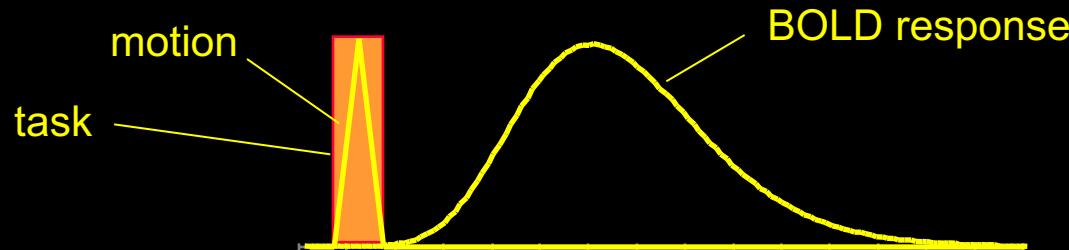


fMRI during tasks that involve brief motion

Blocked Design

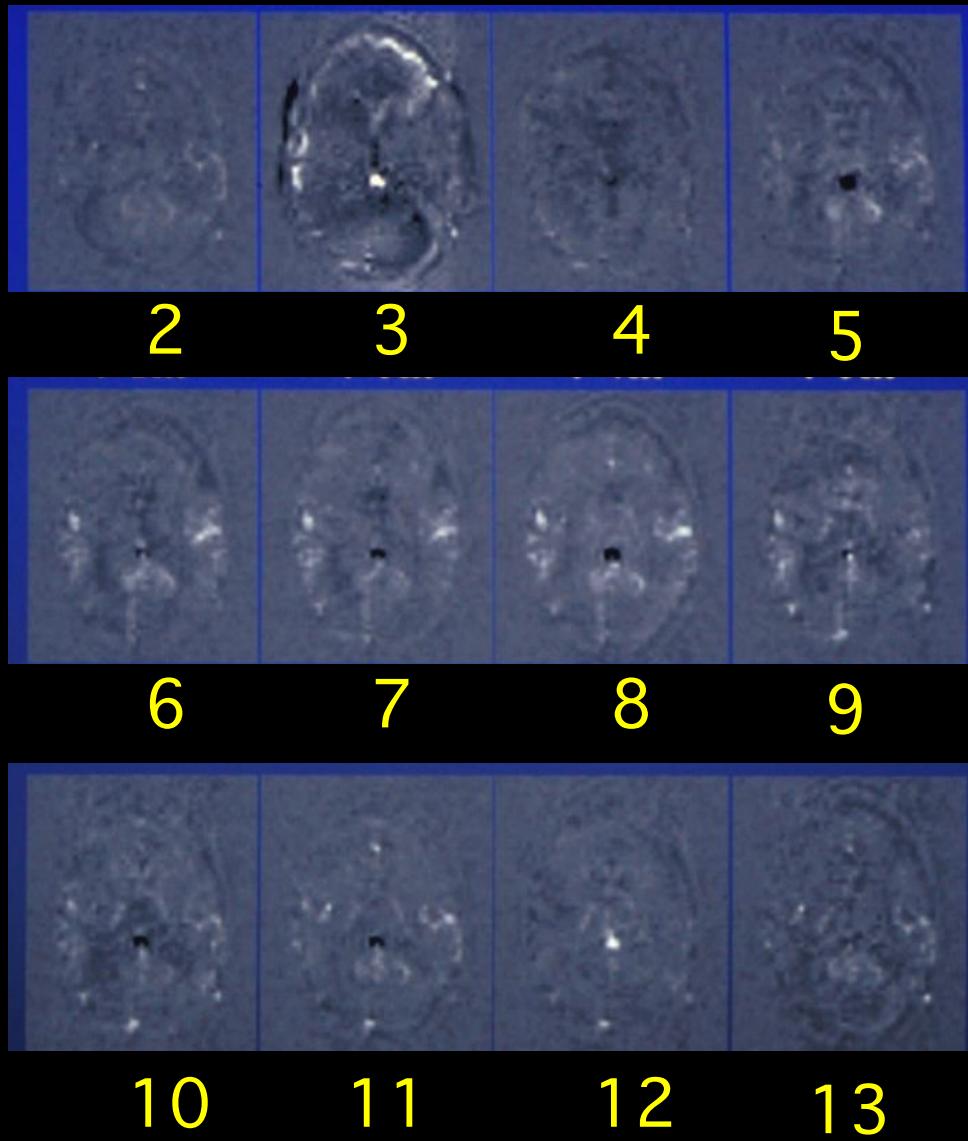


Event-Related Design



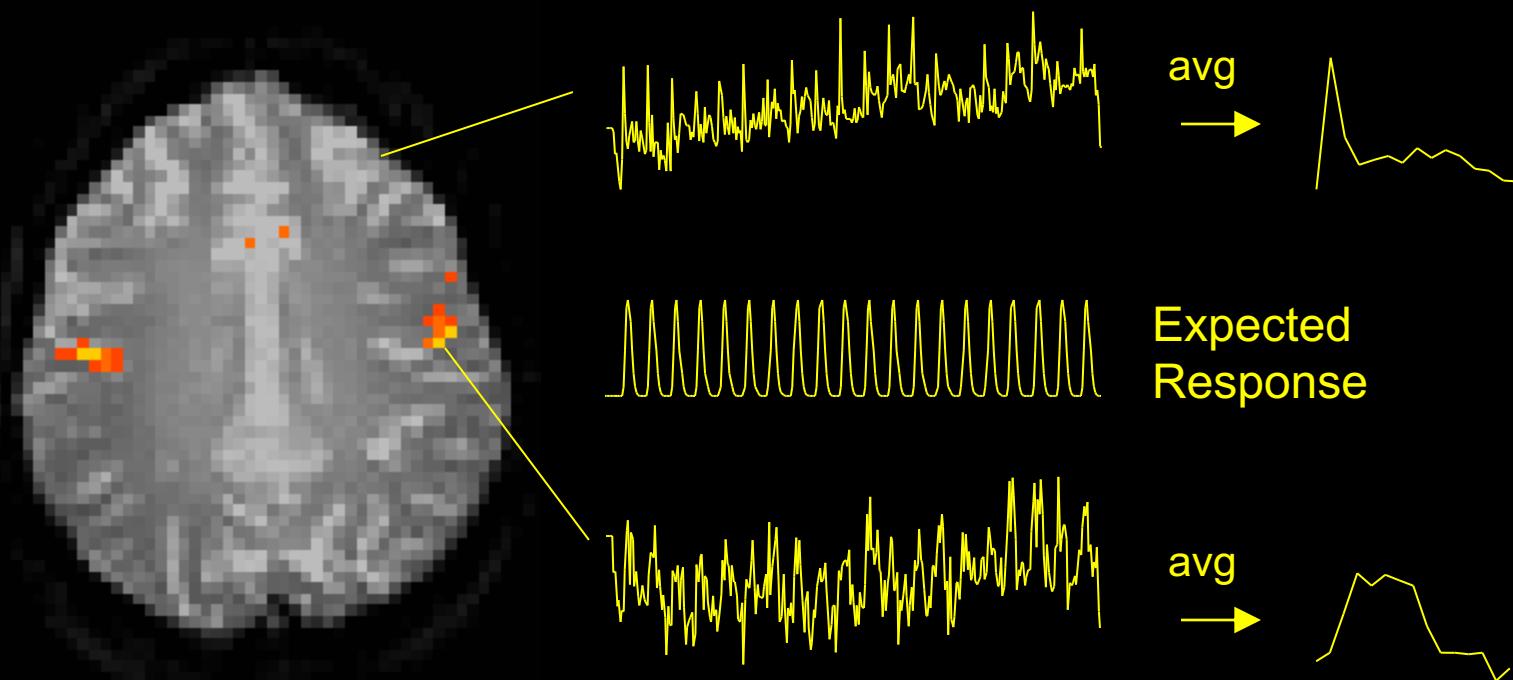
R. M. Birn, P. A. Bandettini, R. W. Cox, R. Shaker, Event - related fMRI of tasks involving brief motion. *Human Brain Mapping* 7: 106-114 (1999).

Overt Word Production



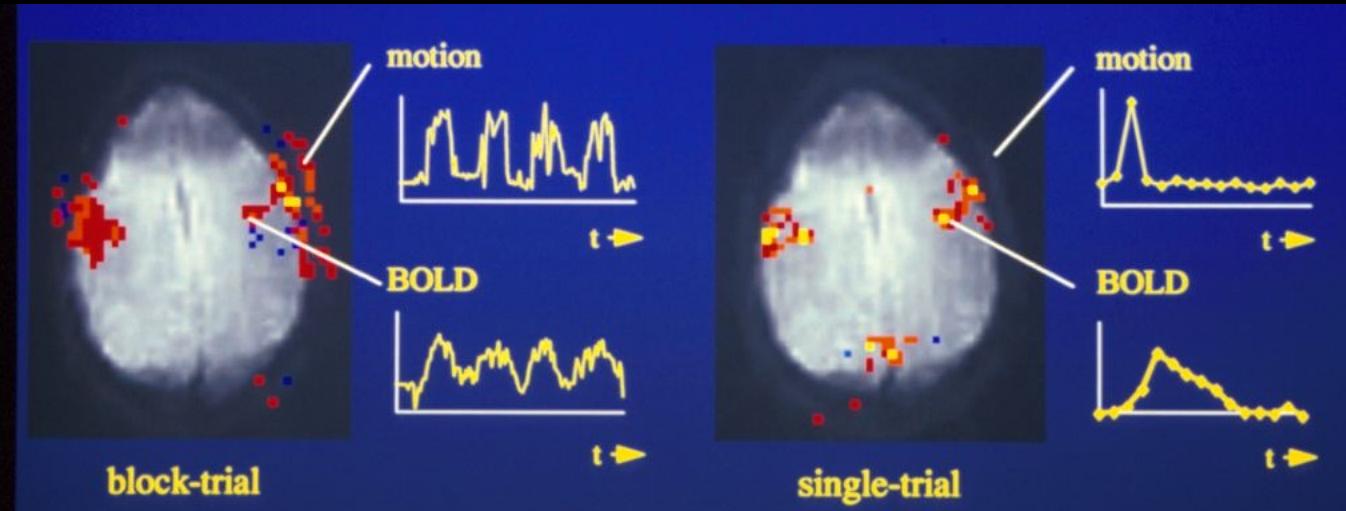
R. M. Birn, P. A. Bandettini, R. W. Cox, R. Shaker, Event - related fMRI of tasks involving brief motion. *Human Brain Mapping* 7: 106-114 (1999).

Speaking - ER-fMRI

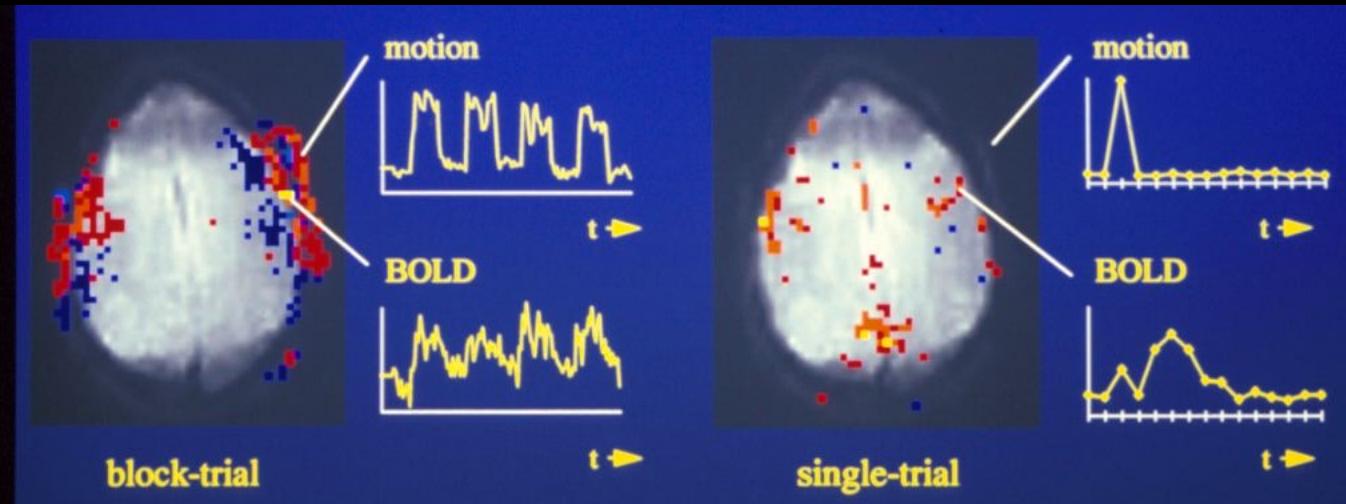


R. M. Birn, P. A. Bandettini, R. W. Cox, R. Shaker, Event - related fMRI of tasks involving brief motion. *Human Brain Mapping* 7: 106-114 (1999).

Tongue Movement

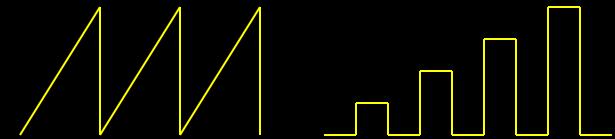


Jaw Clenching

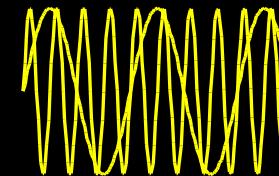


Neuronal Activation Input Strategies

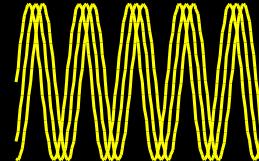
1. Block Design



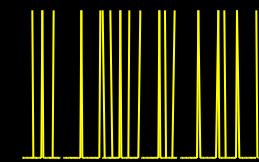
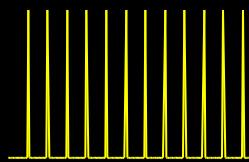
2. Parametric Design



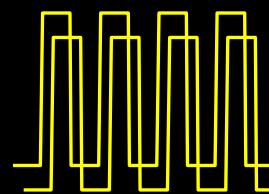
3. Frequency Encoding



4. Phase Encoding



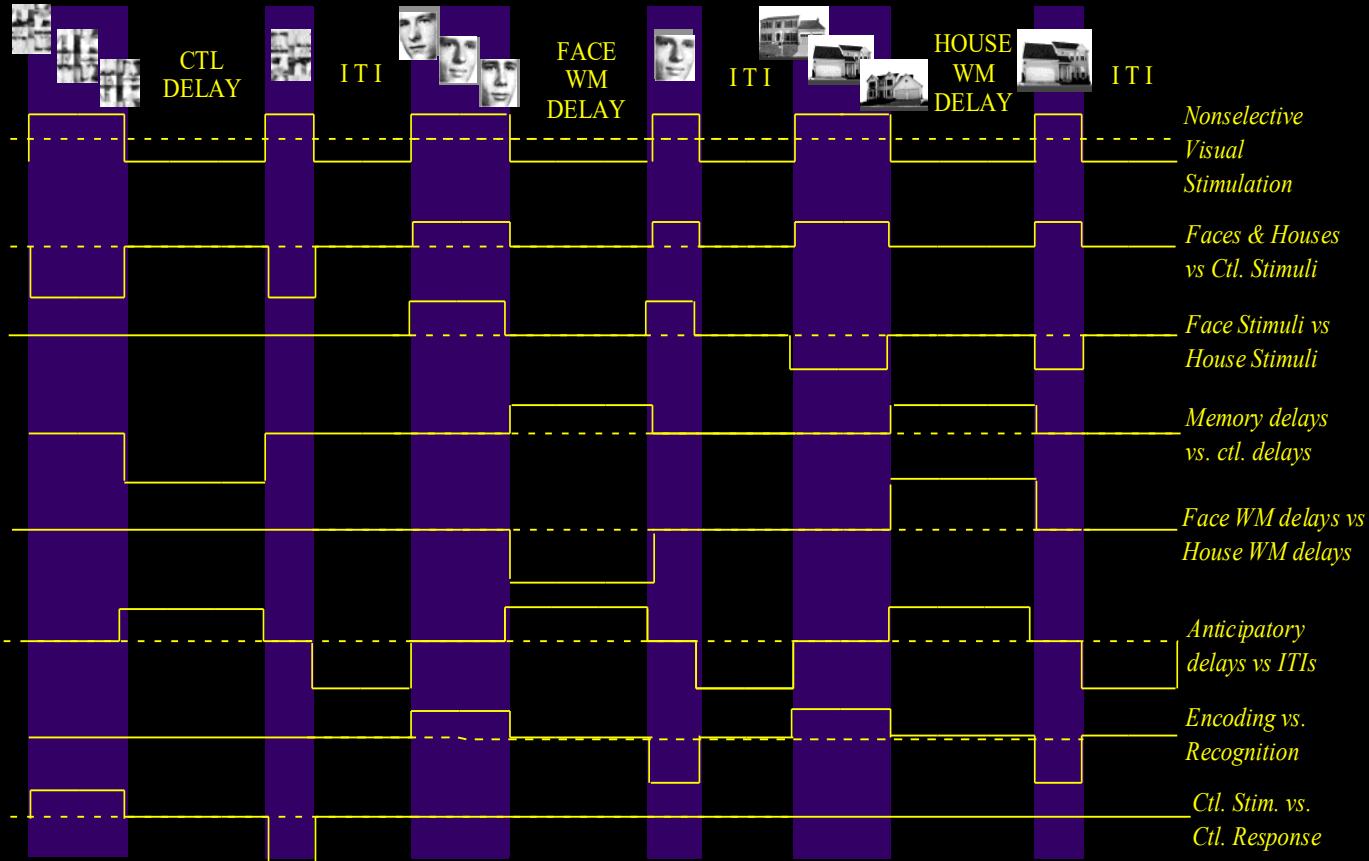
5. Event Related



6. Orthogonal Design

7. Free Behavior Design

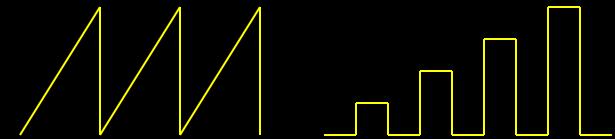
Example of a Set of Orthogonal Contrasts for Multiple Regression



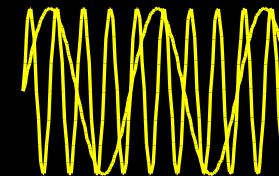
Courtney, S. M., L. G. Ungerleider, et al. (1997). "Transient and sustained activity in a distributed neural system for human working memory." *Nature* 386(6625): 608-11.

Neuronal Activation Input Strategies

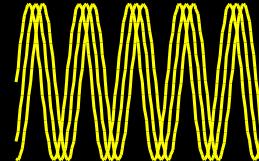
1. Block Design



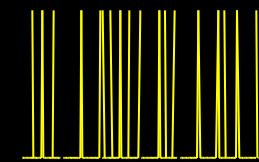
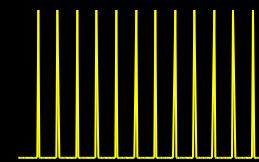
2. Parametric Design



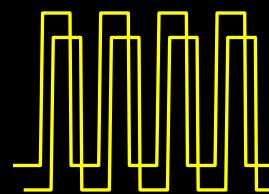
3. Frequency Encoding



4. Phase Encoding



5. Event Related



6. Orthogonal Design

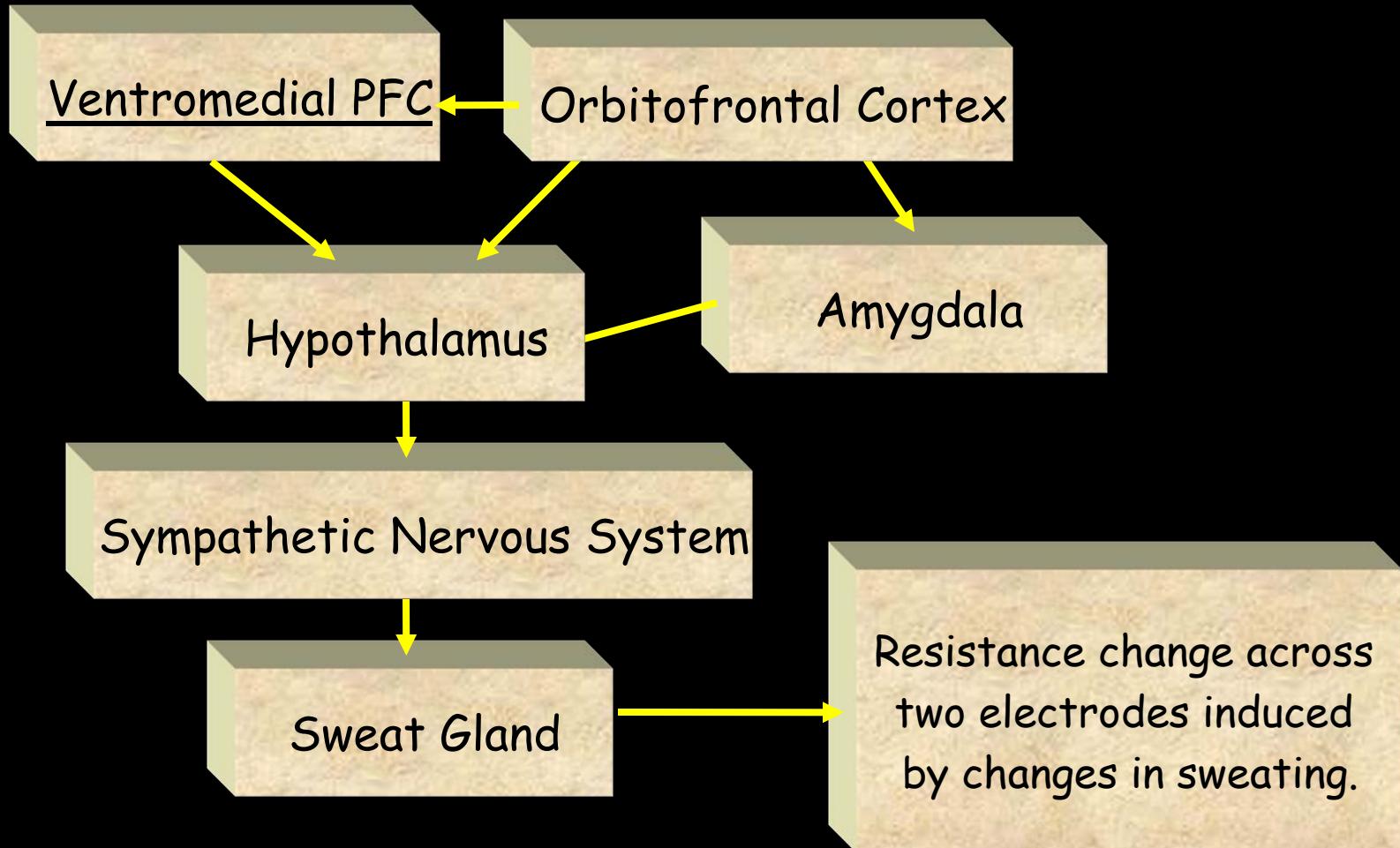
7. Free Behavior Design

Free Behavior Design

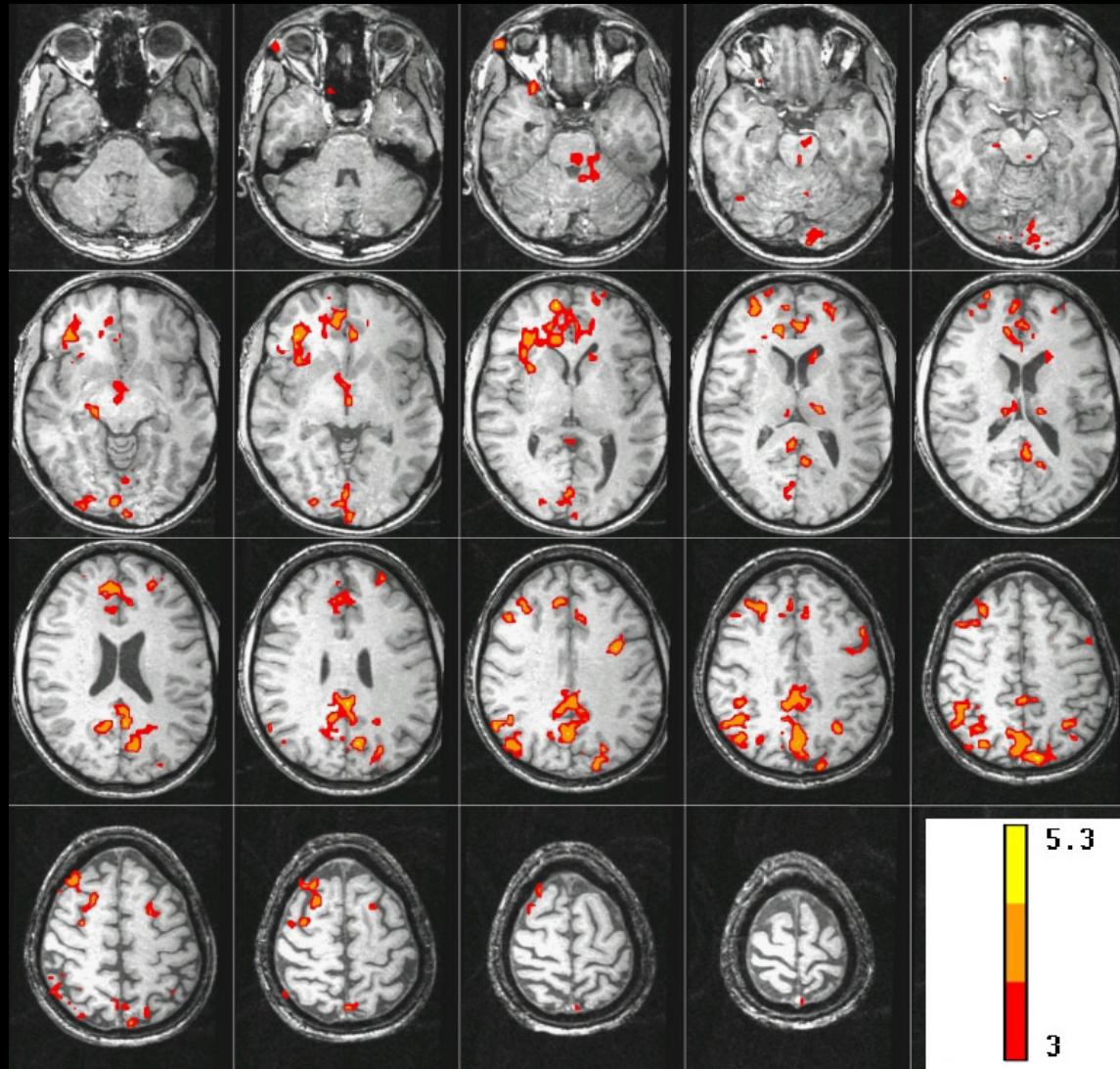
Use a continuous measure as a reference function:

- Task performance
- Skin Conductance
- Heart, respiration rate..
- Eye position
- EEG

The Skin Conductance Response (SCR)

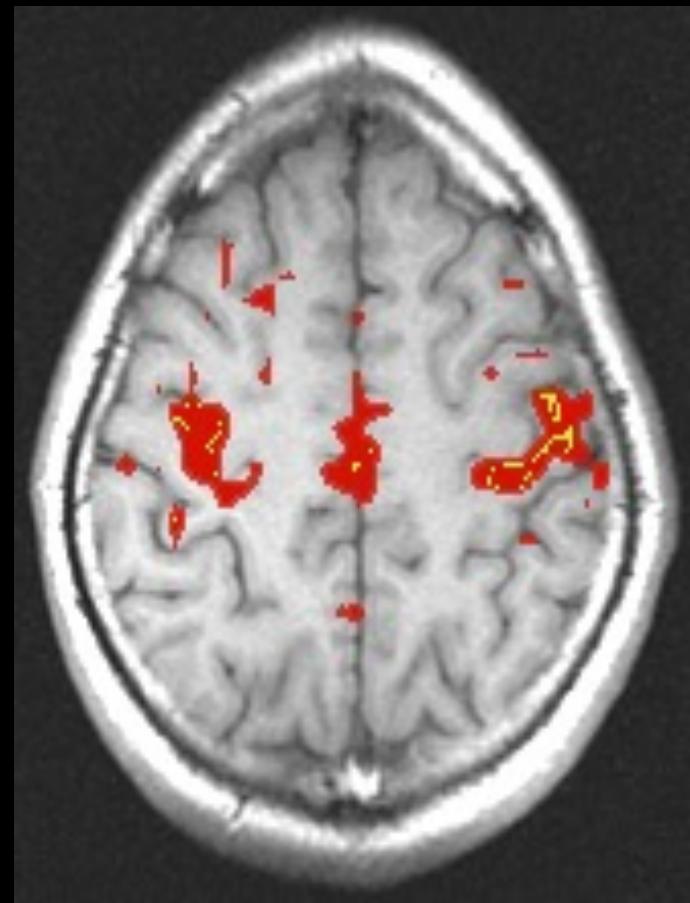
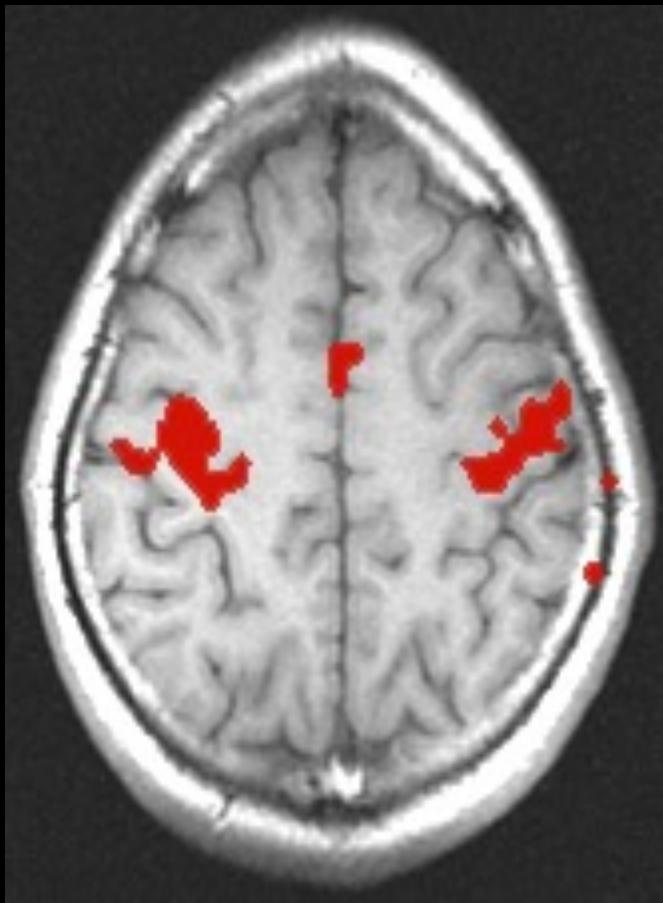


Brain activity correlated with SCR during “Rest”



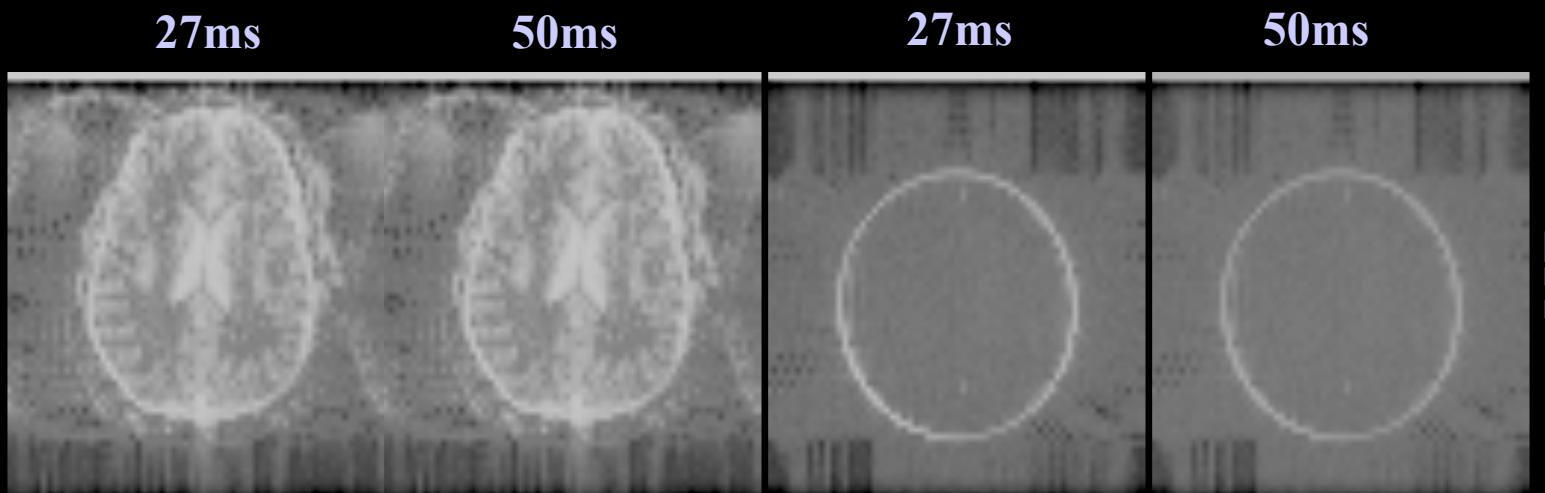
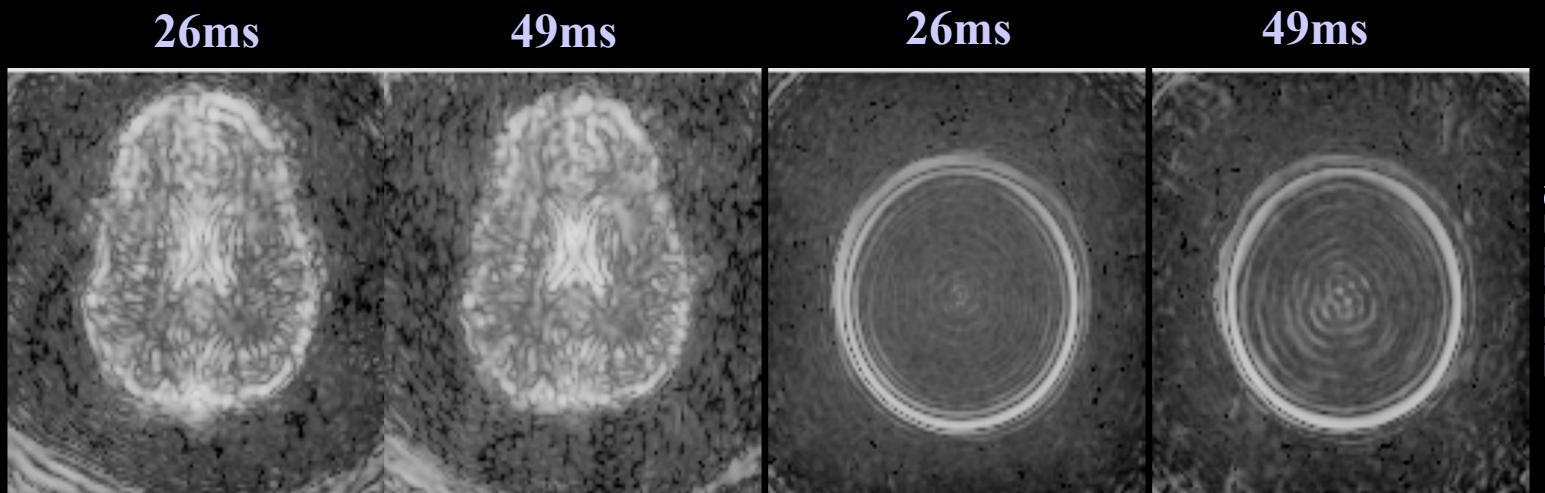
Patterson et al. (submitted)

Resting Hemodynamic Autocorrelations



B. Biswal *et al.*, MRM, 34:537 (1995)

Temporal vs. Spatial SNR- 3T



SPIRAL

EPI

0.25 Hz Breathing at 3T

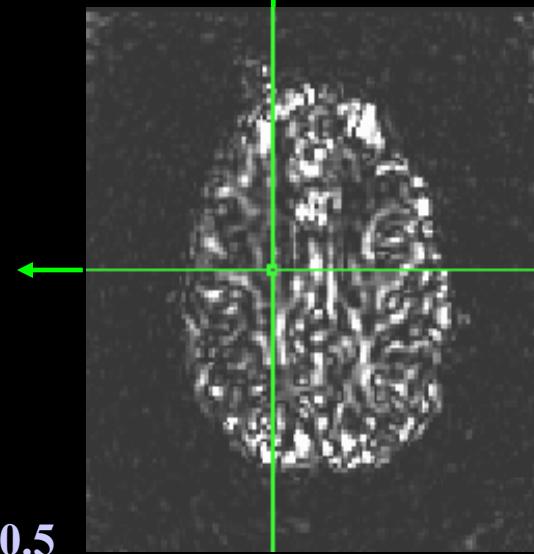
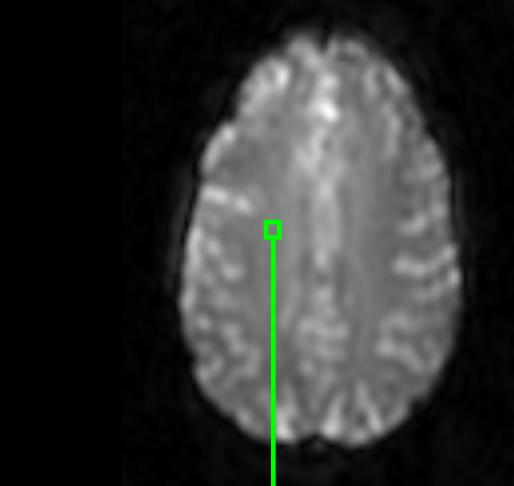
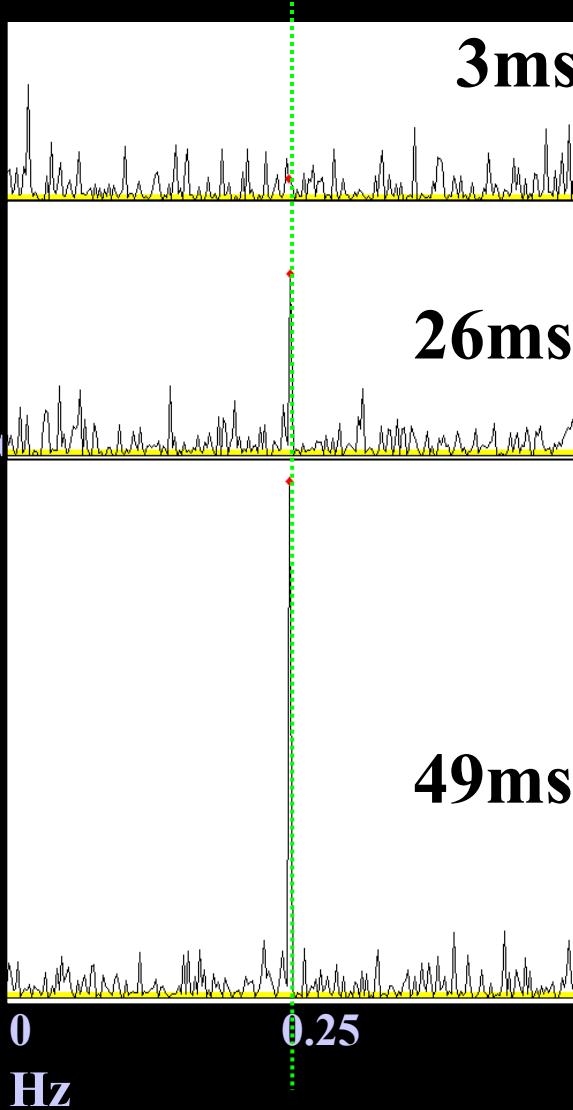


Image Respiration map

0.68 Hz Cardiac rate at 3T

Power Spectra

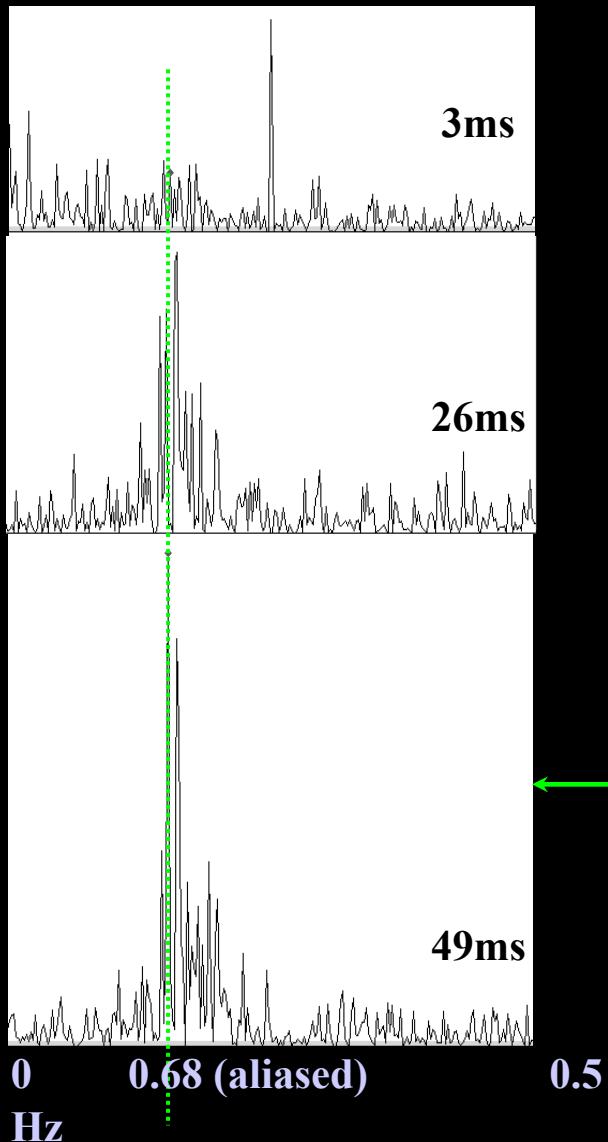
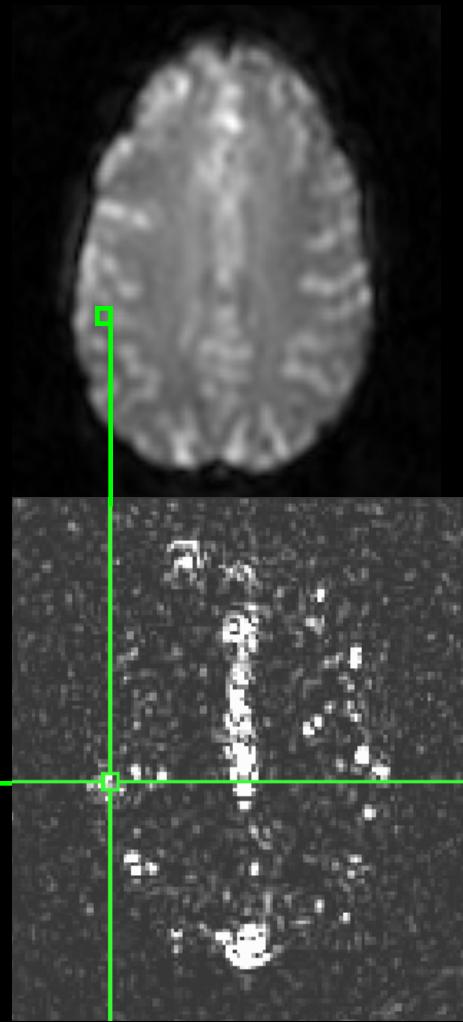
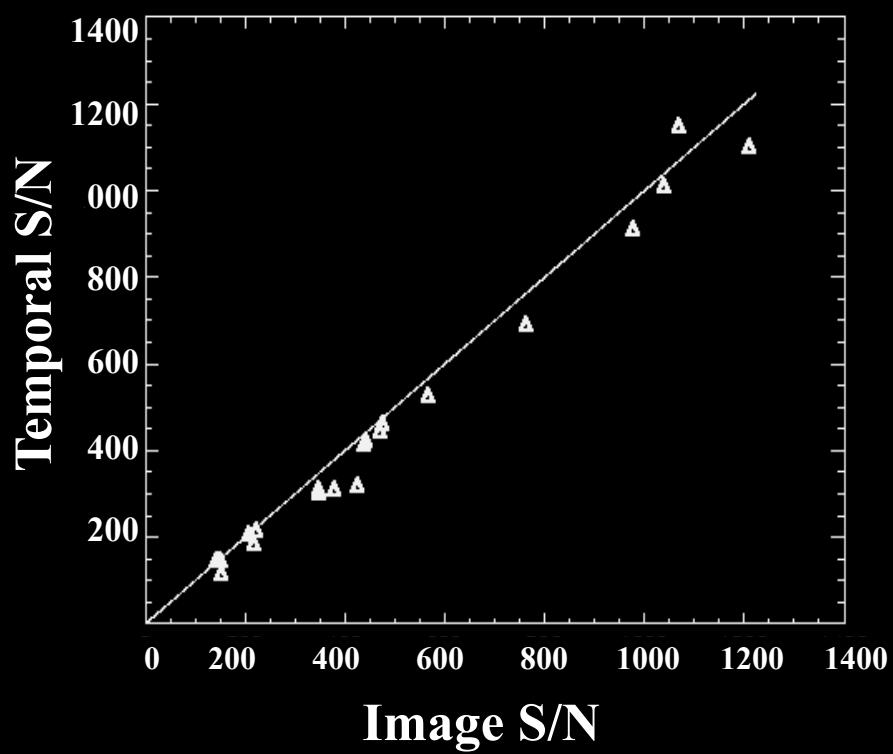


Image
Cardiac map

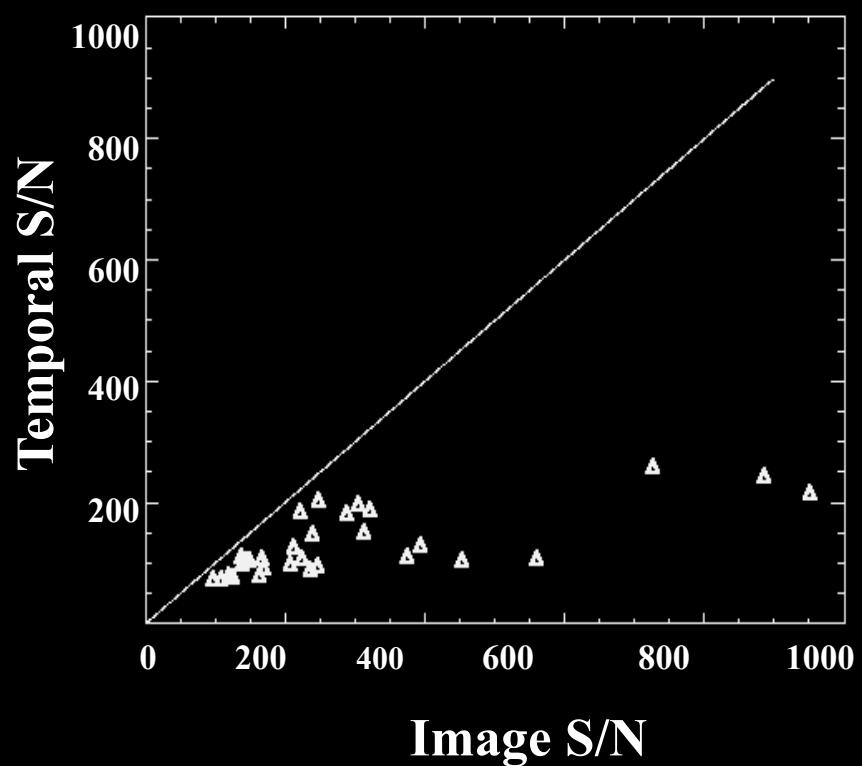


Temporal S/N vs. Image S/N

PHANTOMS

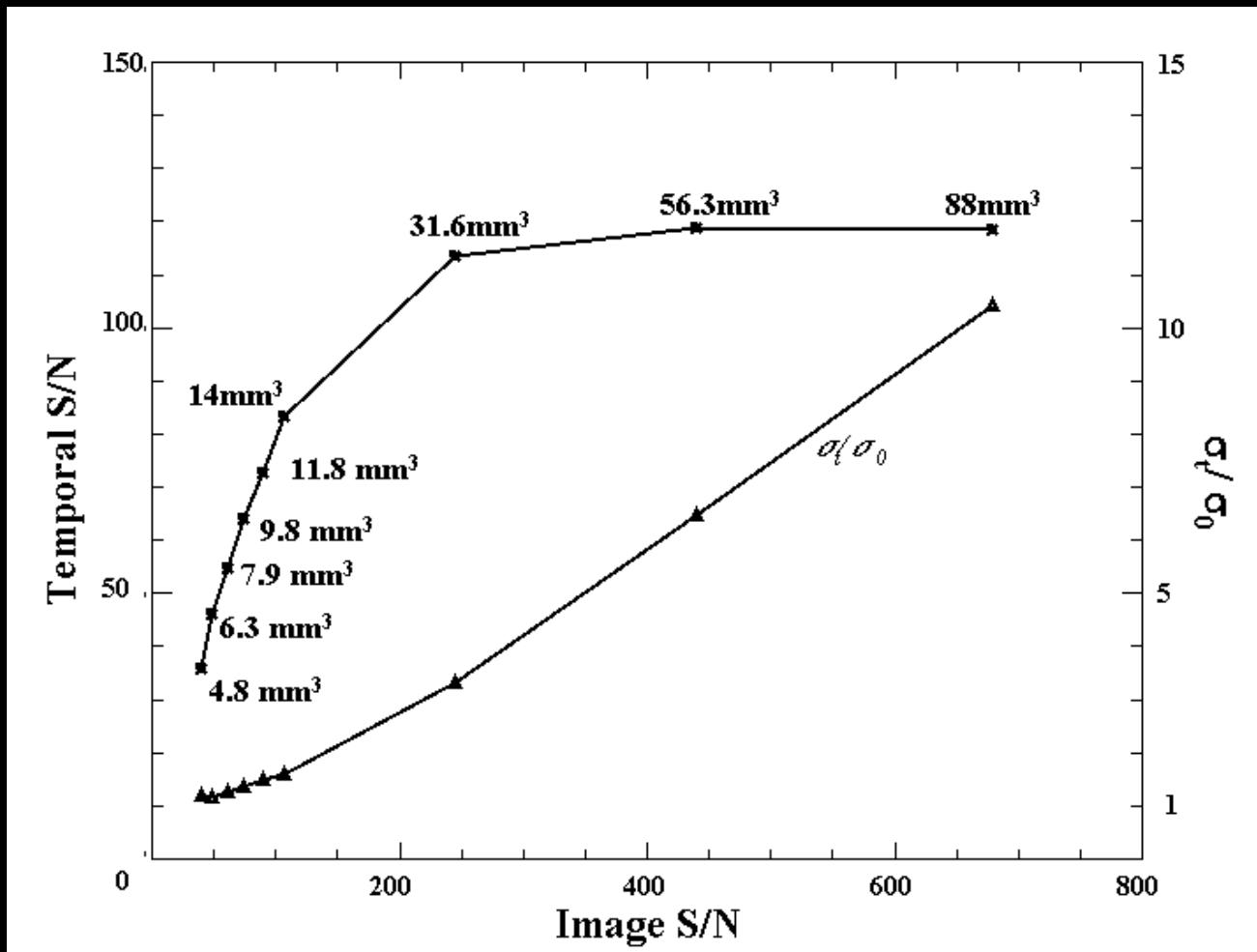


SUBJECTS



N. Petridou

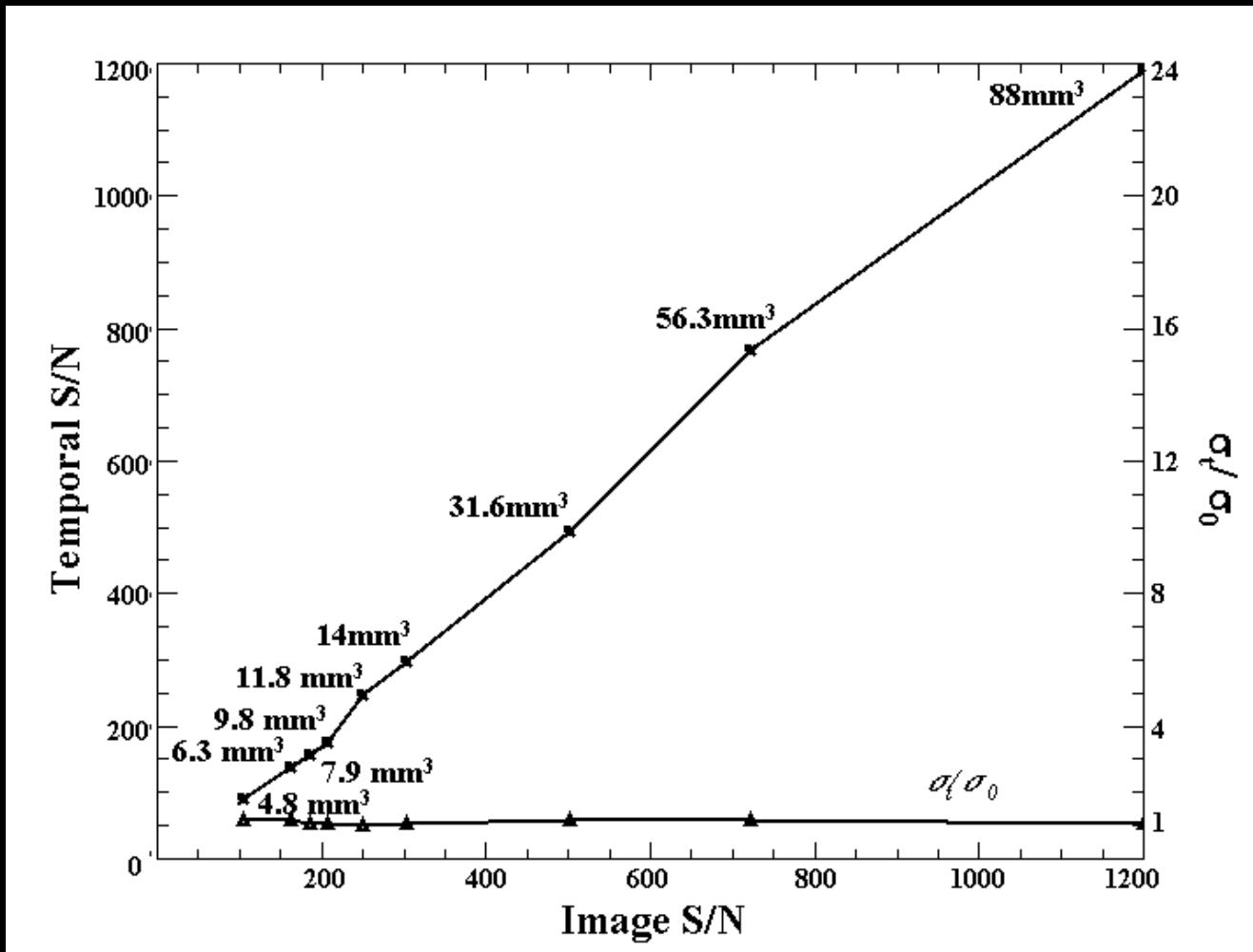
Temporal vs. Image S/N Optimal Resolution Study



Human data

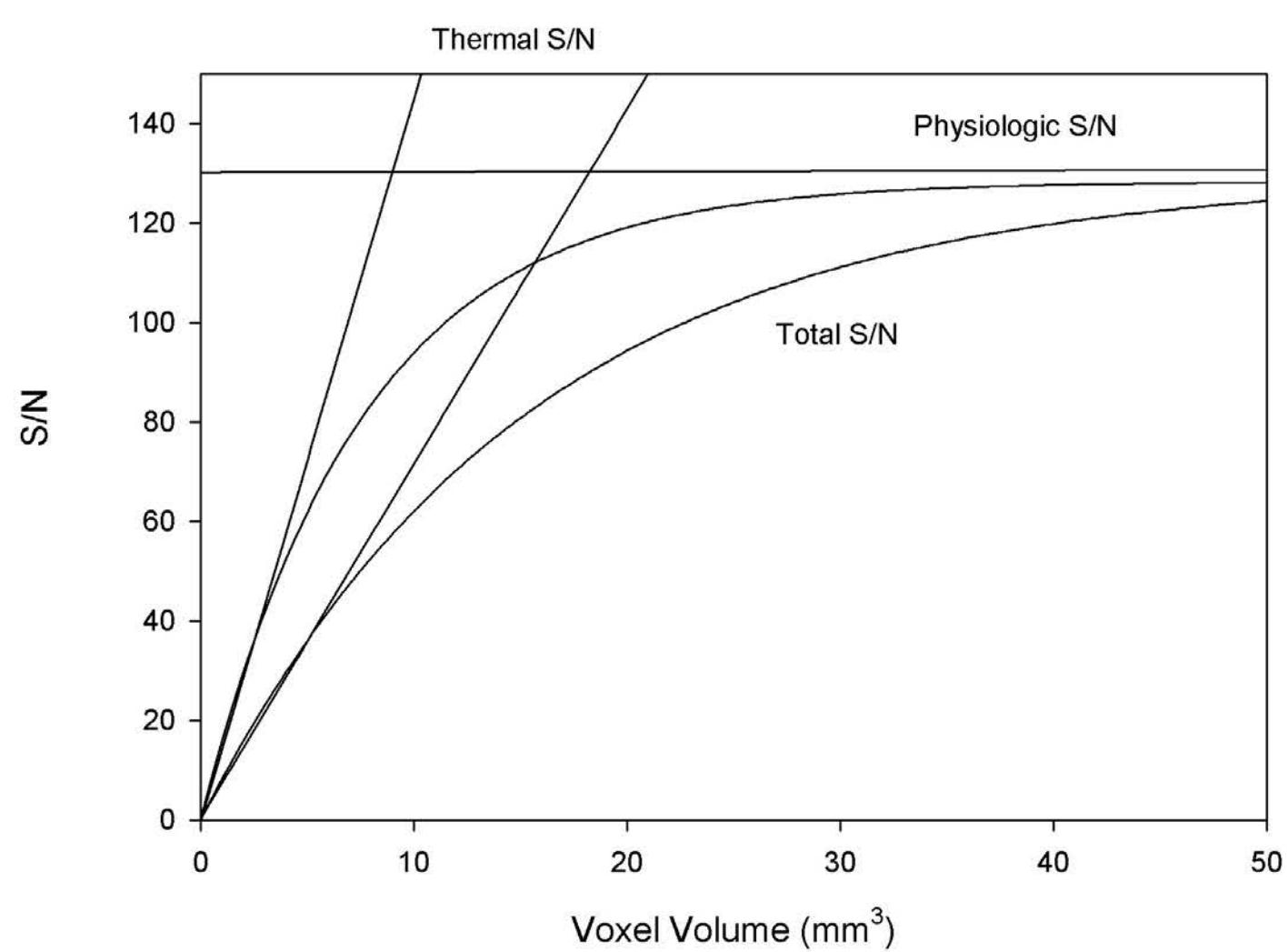
Petridou et al

Temporal vs. Image S/N Optimal Resolution Study

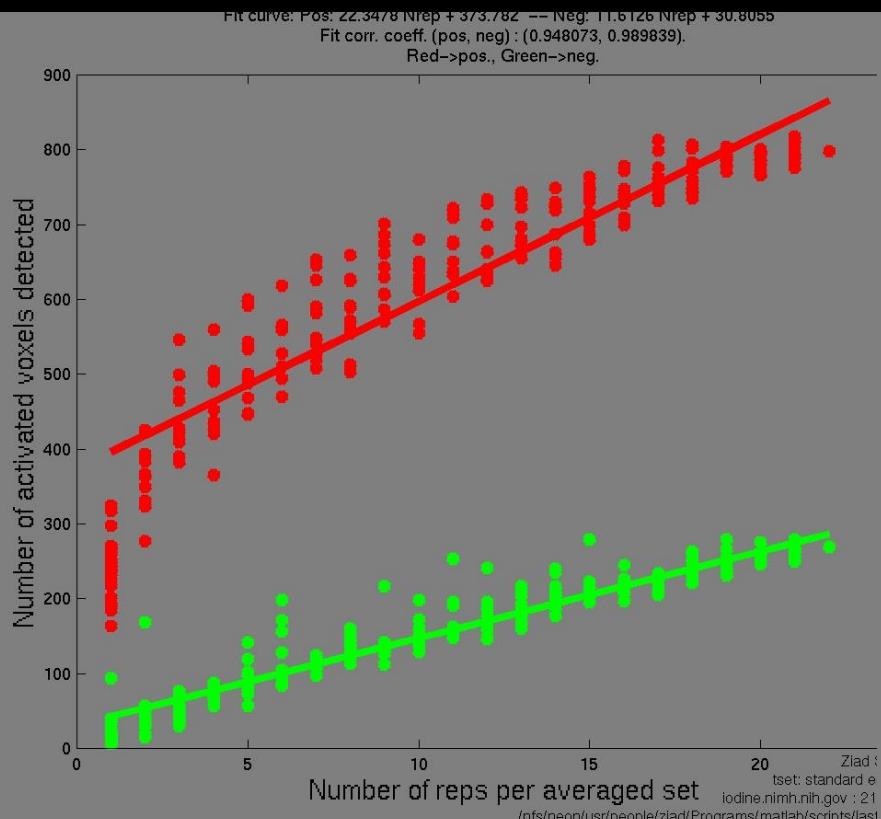


Phantom data

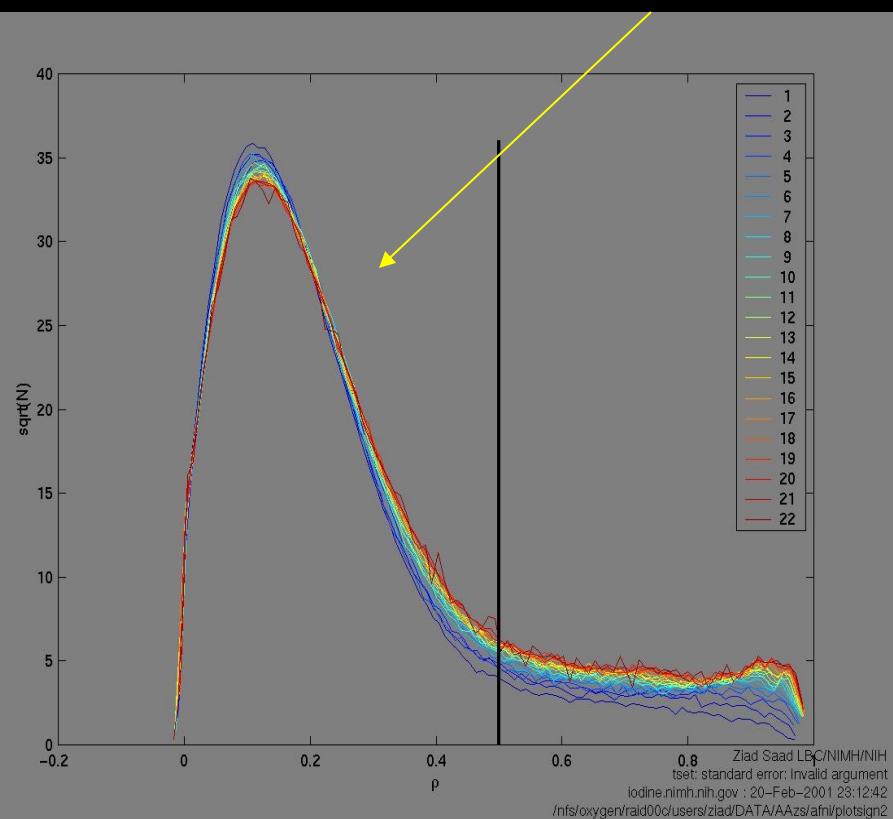
Petridou et al



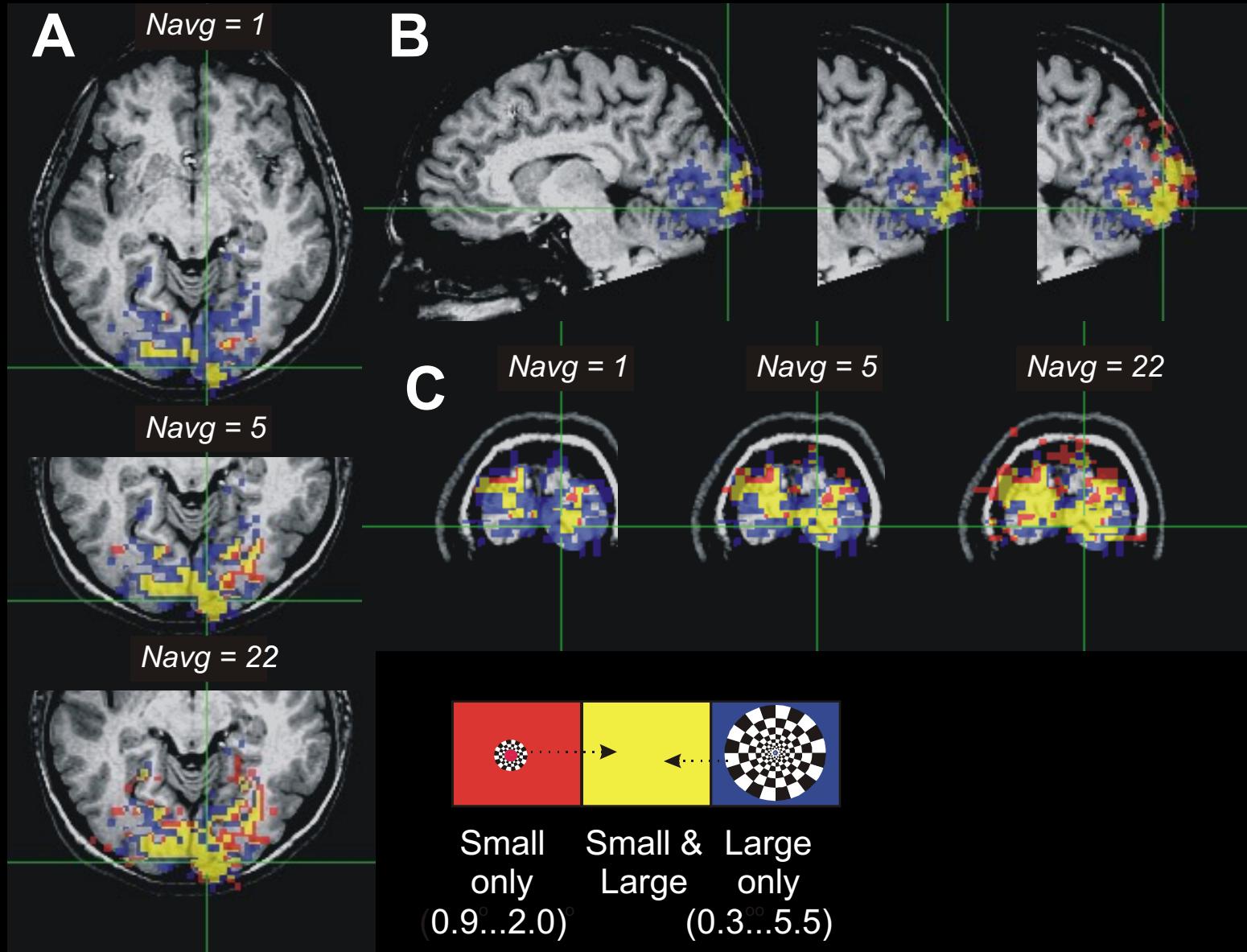
Continuously Growing Activation Area



CC Histogram Inflection Point



Ziad Saad, et al (Submitted)



Motion

Recognize?

- Edge effects
- Shorter signal change latencies
- Unusually high signal changes
- External measuring devices

Correct?

- Image registration algorithms
- Orthogonalize to motion-related function (*cardiac, respiration, movement*)
- Navigator echo for k-space alignment
(for multishot techniques)
- Re-do scan

Bypass?

- Paradigm timing strategies..
- Gating (with T1-correction)

Suppress?

- Flatten image contrast
- Physical restraint
- Averaging, smoothing

Refinements

BOLD Contrast Interpretation

Dynamics, Paradigm Design and Processing

Applications

Technology

MRI	EPI	1.5T,3T, 4T	EPI on Clin. Syst.	Diff. tensor	Mg ⁺	7T
		Local Human Head Gradient Coils		Real time fMRI	Venography	
	ASL	Spiral EPI	Nav. pulses	Quant. ASL	Z-shim	SENSE
	BOLD		Multi-shot fMRI	Dynamic IV volume	Simultaneous ASL and BOLD	Baseline Susceptibility
						Current Imaging?

Methodology

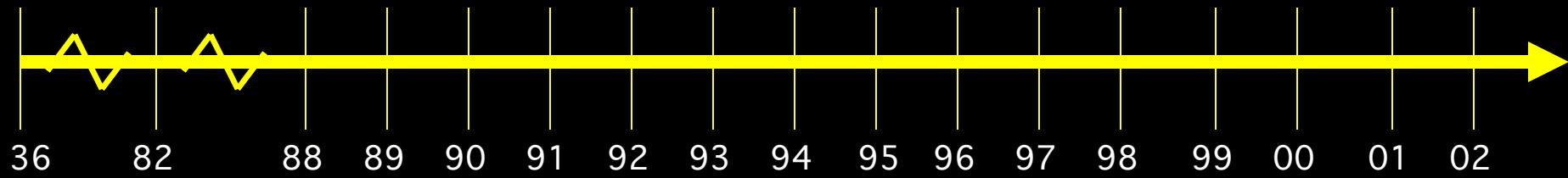
IVIM	Baseline Volume	Correlation Analysis	Motion Correction	CO ₂ Calibration
		Parametric Design		Multi-Modal Mapping
	Linear Regression	Surface Mapping		Free-behavior Designs
		Phase Mapping		
		Event-related	Mental Chronometry	Deconvolution

Interpretation

Blood T2	BOLD models	PET correlation		
	B ₀ dep.	IV vs EV	ASL vs. BOLD	
		Pre-undershoot	PSF of BOLD	
	TE dep	Resolution Dep.		Extended Stim.
		Post-undershoot		
Hemoglobin	SE vs. GE	CO ₂ effect	Linearity	Metab. Correlation
		NIRS Correlation	Fluctuations	Optical Im. Correlation
	Veins	Inflow	Balloon Model	Electophys. correlation

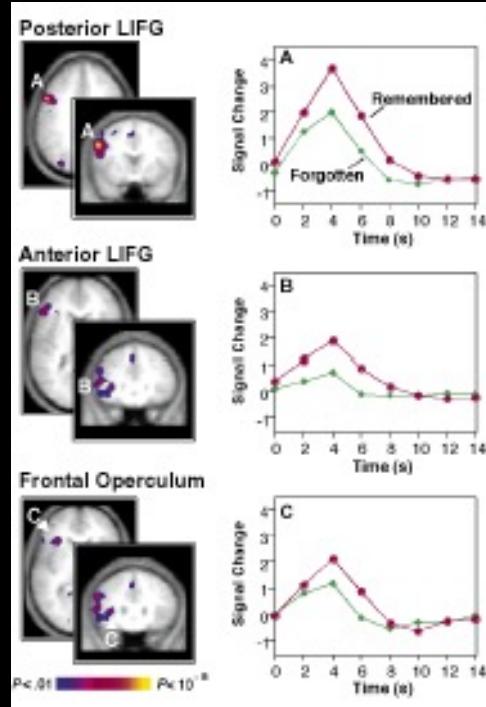
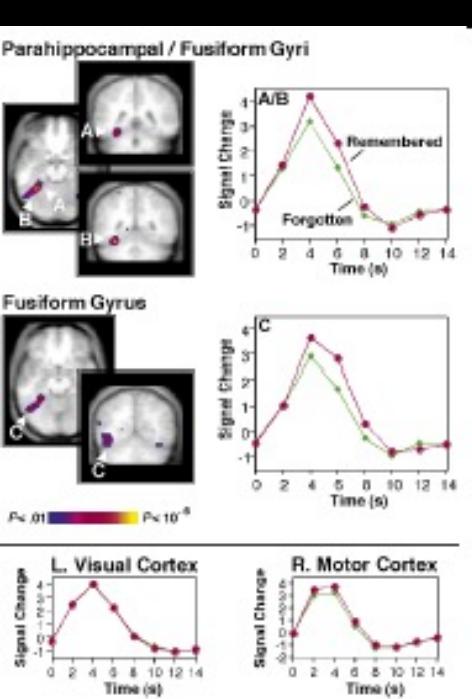
Applications

Volume - Stroke	Complex motor			
	Language	Imagery	Memory	Emotion
	Motor learning	Children	Tumor vasc.	Drug effects
	BOLD -V1, M1, A1	Presurgical	Attention	Ocular Dominance
Δ Volume-V1	V1, V2..mapping	Priming/Learning	Clinical Populations	
		Plasticity	Face recognition	Performance prediction



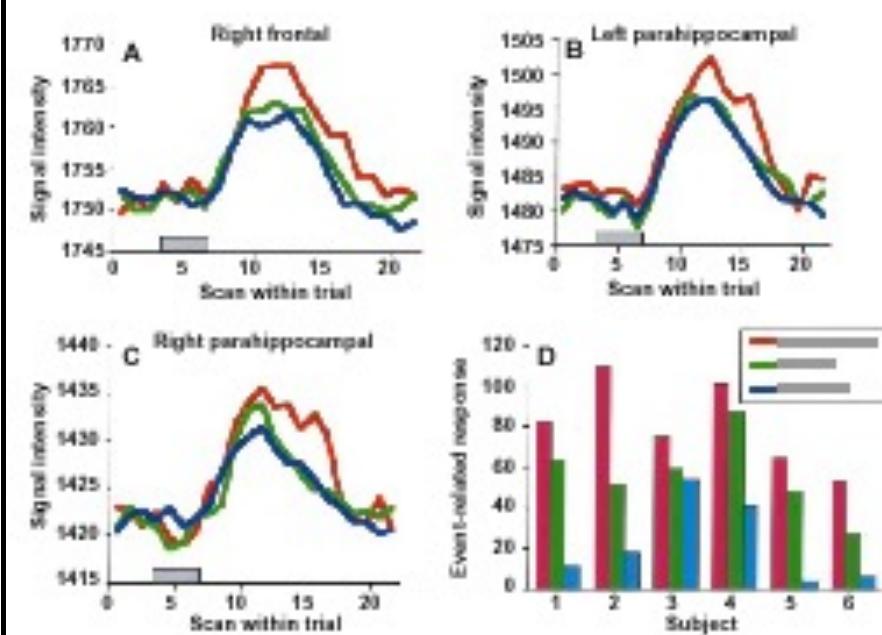
Building Memories: Remembering and Forgetting of Verbal Experiences as Predicted by Brain Activity

Anthony D. Wagner,* Daniel L. Schacter, Michael Rotte,†
Wilma Koutstaal, Anat Maril, Anders M. Dale, Bruce R. Rosen,
Randy L. Buckner



Making Memories: Brain Activity that Predicts How Well Visual Experience Will Be Remembered

James B. Brewer,* Zuo Zhao, John E. Desmond, Gary H. Glover,
John D. E. Gabrieli



Technology

MRI	EPI	1.5T,3T, 4T	EPI on Clin. Syst. Nav. pulses	Diff. tensor	Mg ⁺	7T Venography SENSE
		Local Human Head Gradient Coils		Real time fMRI		
		ASL		Quant. ASL	Z-shim	
		BOLD		Dynamic IV volume		
				Multi-shot fMRI	Simultaneous ASL and BOLD	Baseline Susceptibility Current Imaging?

Methodology

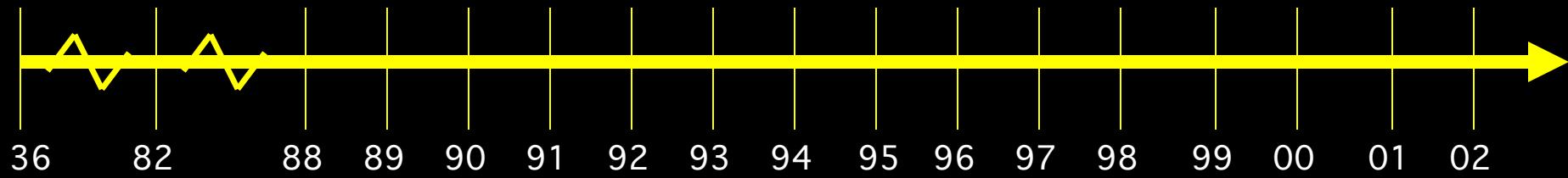
IVIM	Baseline Volume	Correlation Analysis	CO ₂ Calibration
		Motion Correction	
		Parametric Design	Multi-Modal Mapping
		Surface Mapping	
		Phase Mapping	Free-behavior Designs
		Linear Regression	Mental Chronometry
		Event-related	Deconvolution

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		Post-undershoot	Linearity Metab. Correlation
		SE vs. GE	CO ₂ effect
		NIRS Correlation	Fluctuations Optical Im. Correlation
		Veins	Inflow Balloon Model Electophys. correlation

Applications

Volume - Stroke	△ Volume-V1	Complex motor	Memory	Emotion
		Language		
		Imagery		
		Motor learning		
		Children		
		Tumor vasc.		Drug effects
		Presurgical		Ocular Dominance
		Attention		
		Priming/Learning		Clinical Populations
		Plasticity		Performance prediction
		Face recognition		



Δ Neuronal Activity

Number of Neurons
Local Field Potential
Spiking Coherence
Spiking Rate

Δ Metabolism

Aerobic Metabolism

Anaerobic Metabolism

Δ Hemodynamics

Blood Volume

Deoxygenated Blood

Flow Velocity

Oxygenated Blood

Perfusion

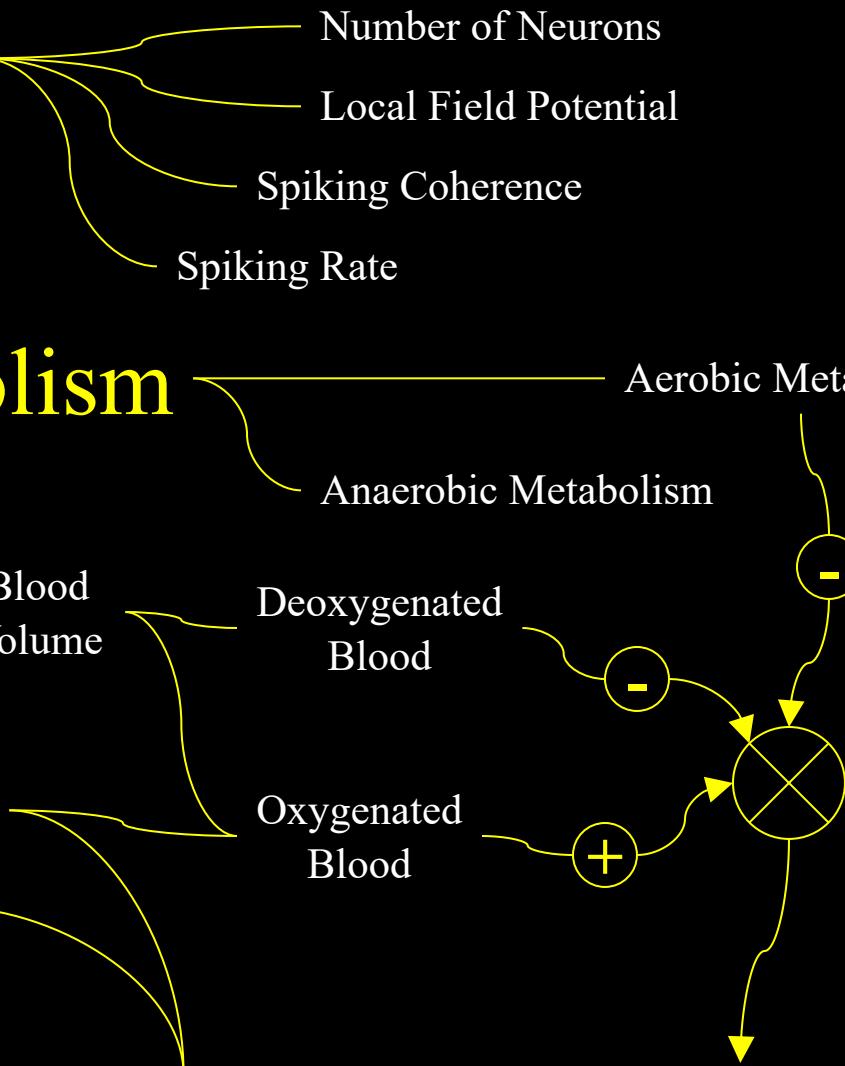
Δ BOLD Contrast

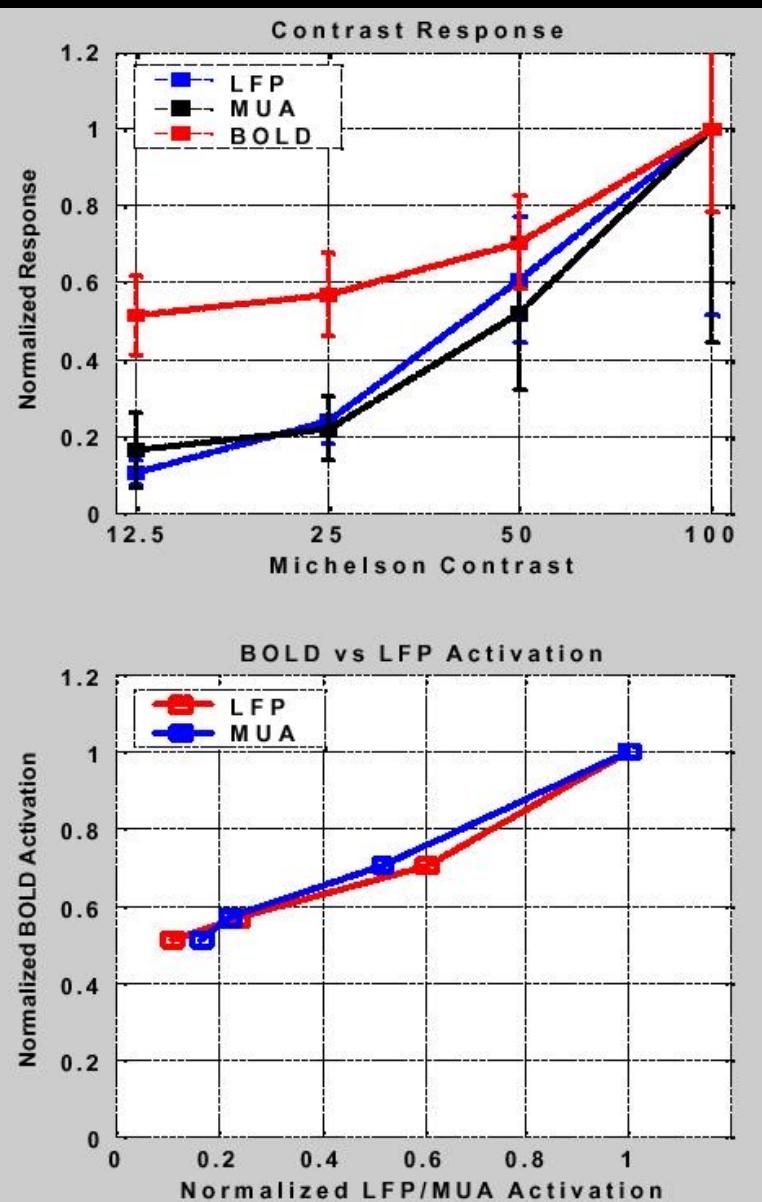
Δ Perfusion Contrast

Δ Inflow Contrast

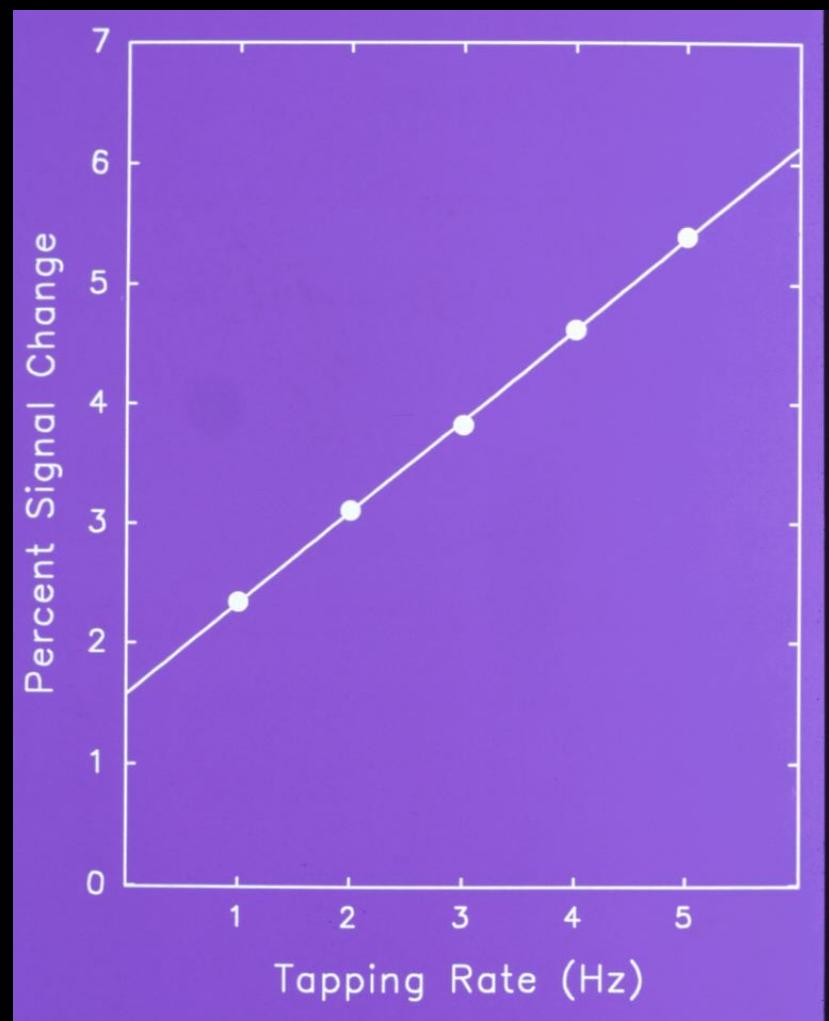
MRI Pulse Sequence

Δ Deoxy-Hb





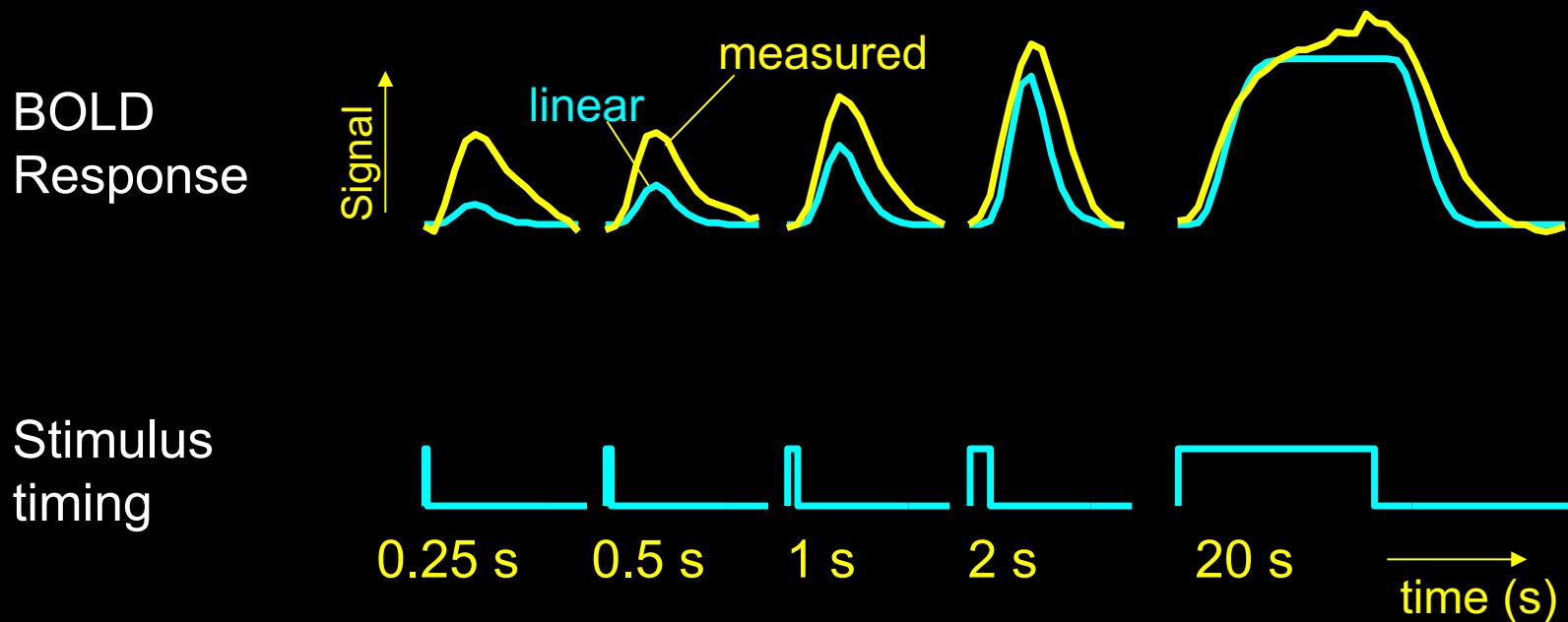
Logothetis et al. (2001) “Neurophysiological investigation of the basis of the fMRI signal” Nature, 412, 150-157



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.

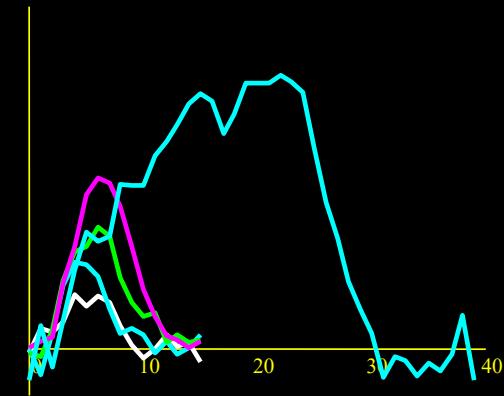
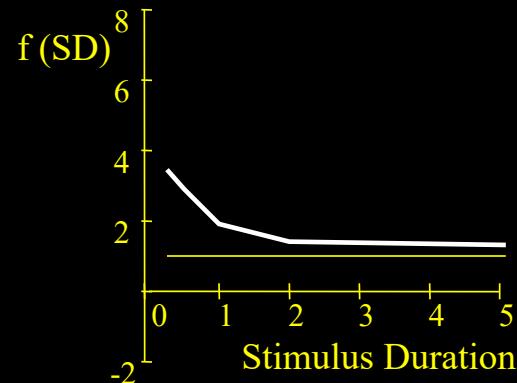
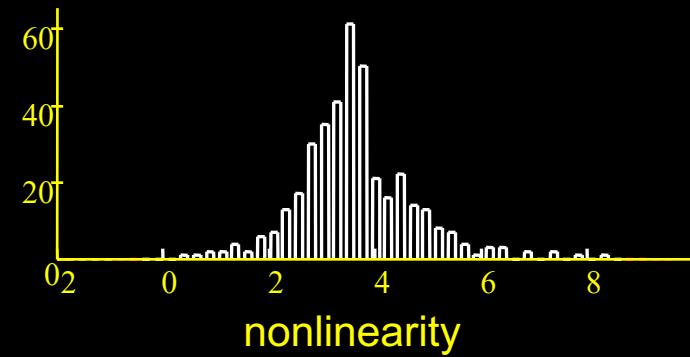
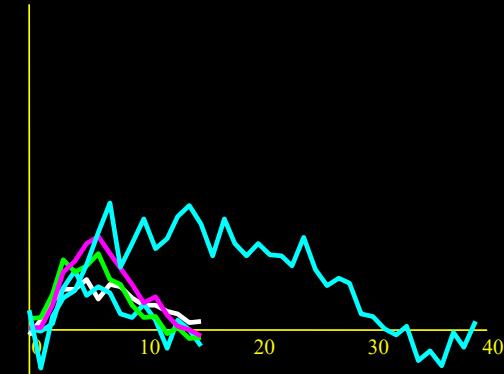
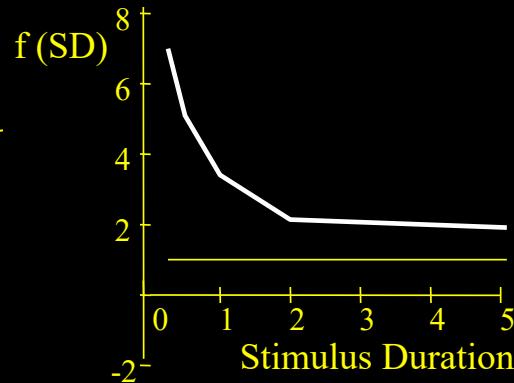
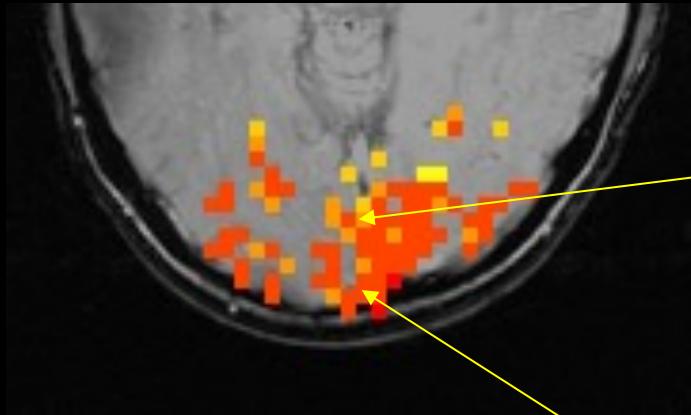
Dynamic Nonlinearity Assessment

Different stimulus “ON” periods



Brief stimuli produce larger responses than expected

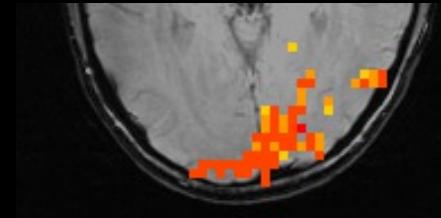
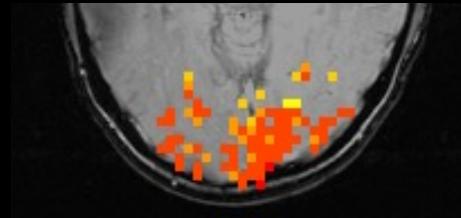
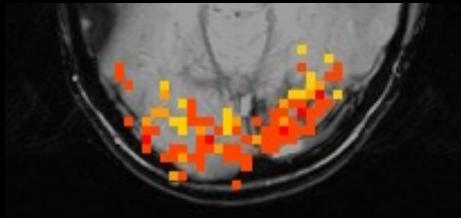
Spatial Heterogeneity of BOLD Nonlinearity



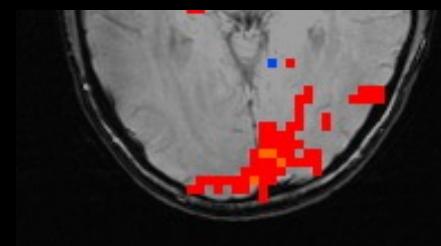
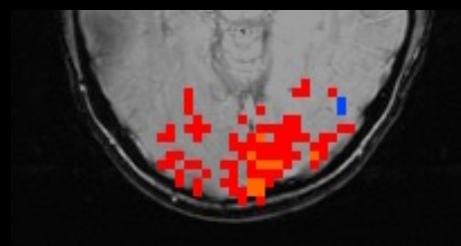
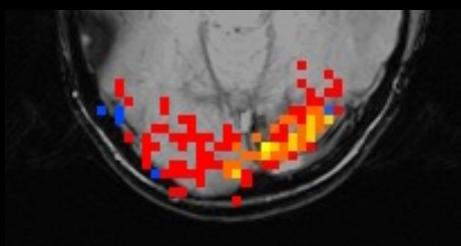
R. M. Birn, Z. Saad, P. A. Bandettini, (2001) "Spatial heterogeneity of the nonlinear dynamics in the fMRI BOLD response." *NeuroImage*, 14: 817-826.

Results – visual task

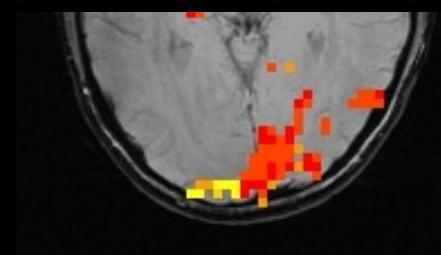
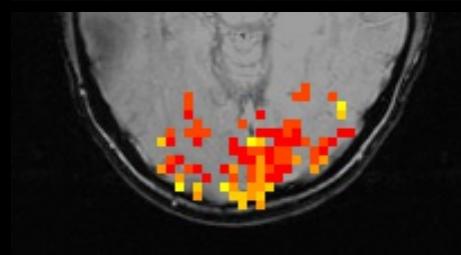
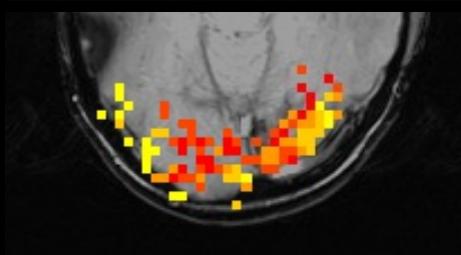
Nonlinearity



Magnitude

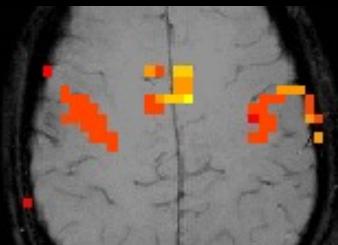


Latency

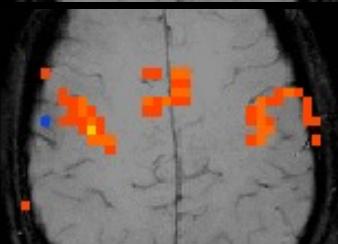


Results – motor task

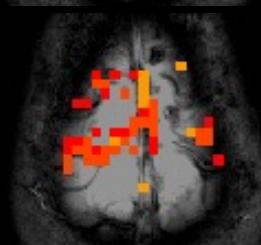
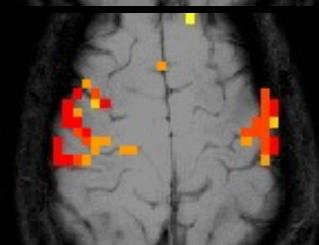
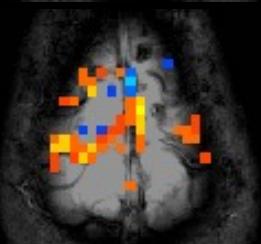
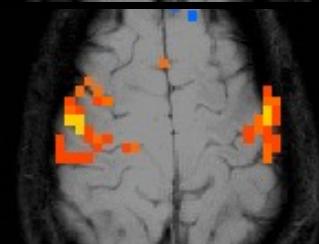
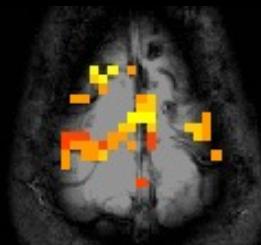
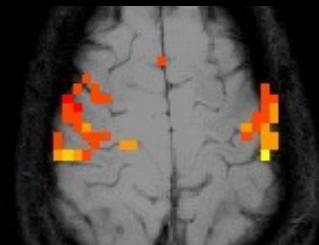
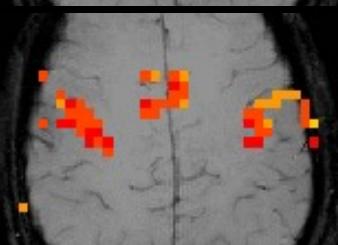
Nonlinearity



Magnitude

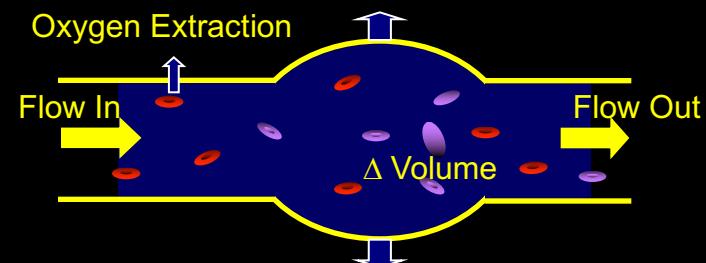
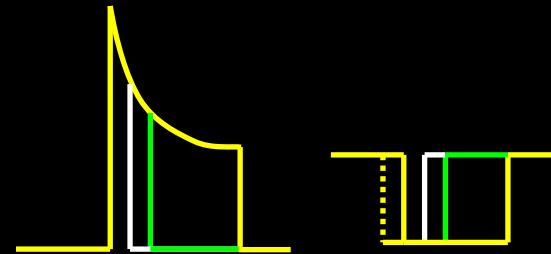
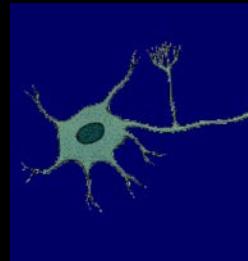


Latency



Sources of this Nonlinearity

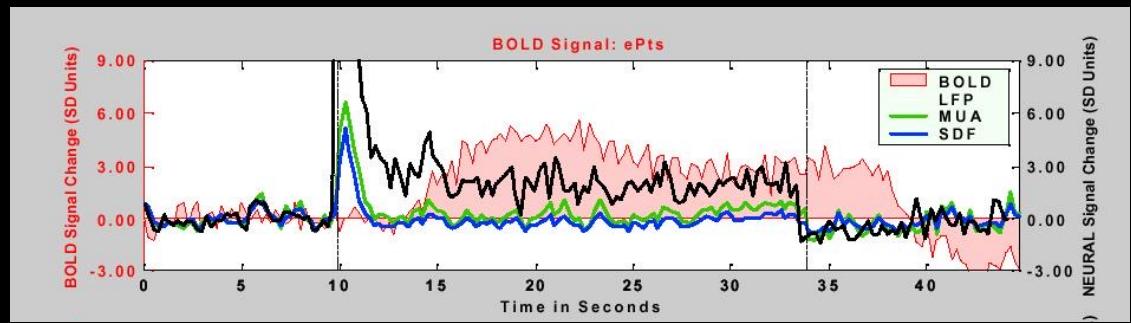
- Neuronal
- Hemodynamic
 - Oxygen extraction
 - Blood volume dynamics



BOLD Correlation with Neuronal Activity

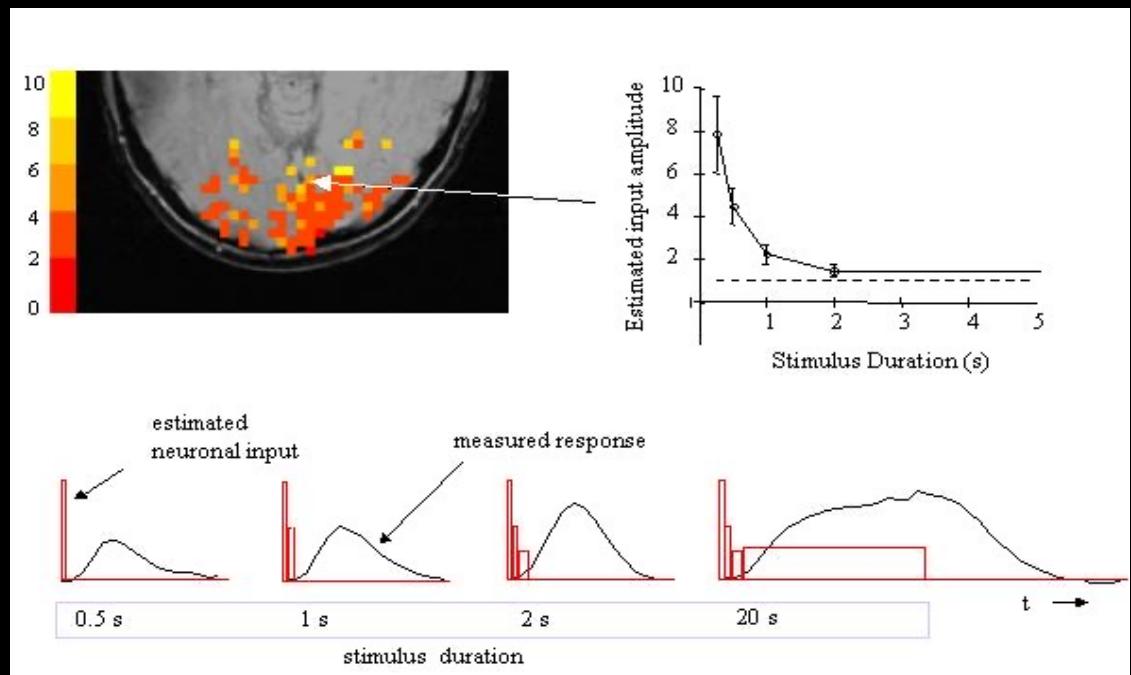
Logothetis et al. (2001)

“Neurophysiological investigation
of the basis of the fMRI signal”
Nature, 412, 150-157.



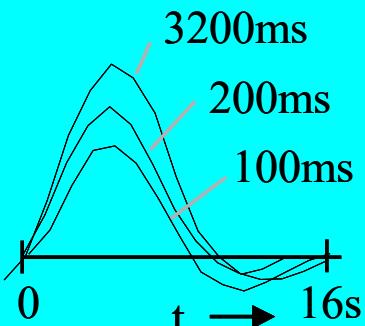
P. A. Bandettini and L. G.

Ungerleider, (2001) “From neuron
to BOLD: new connections.”
Nature Neuroscience, 4: 864-866.

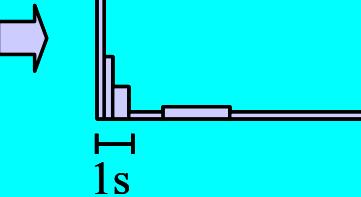


Stationary grating

BOLD response

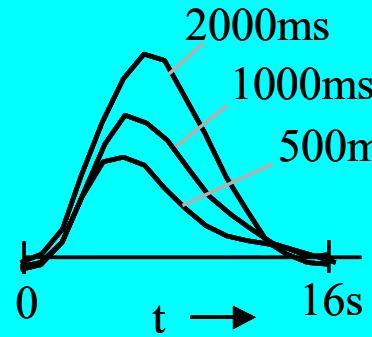


Estimated
Neuronal Input

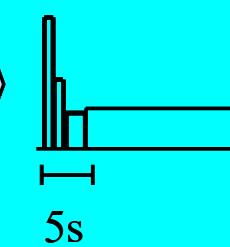


Contrast-reversing checkerboard

BOLD response



Estimated
Neuronal Input



Varying “ON” and “OFF” periods

- *Rapid event-related design with varying ISI*



8% ON



25% ON

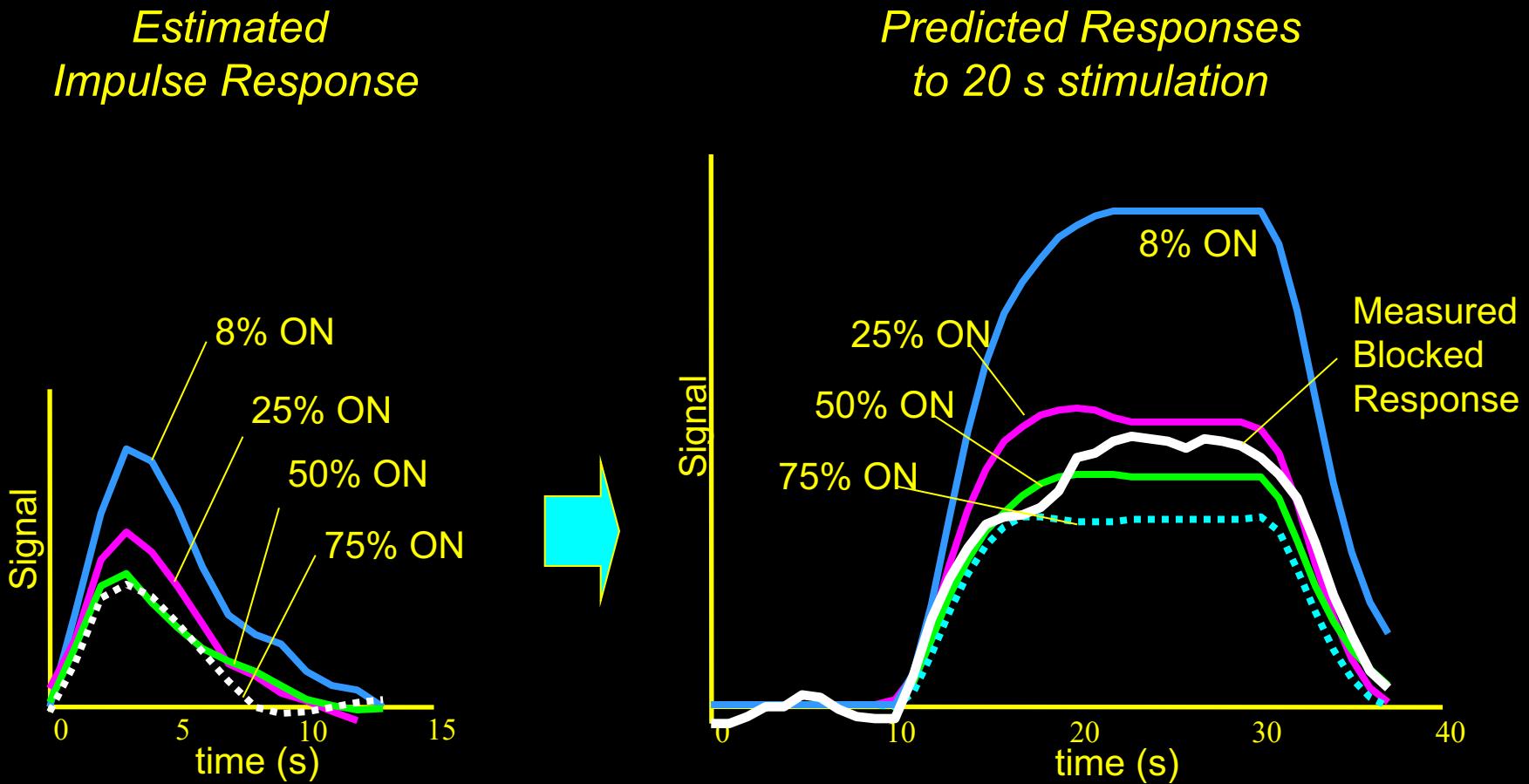


50% ON

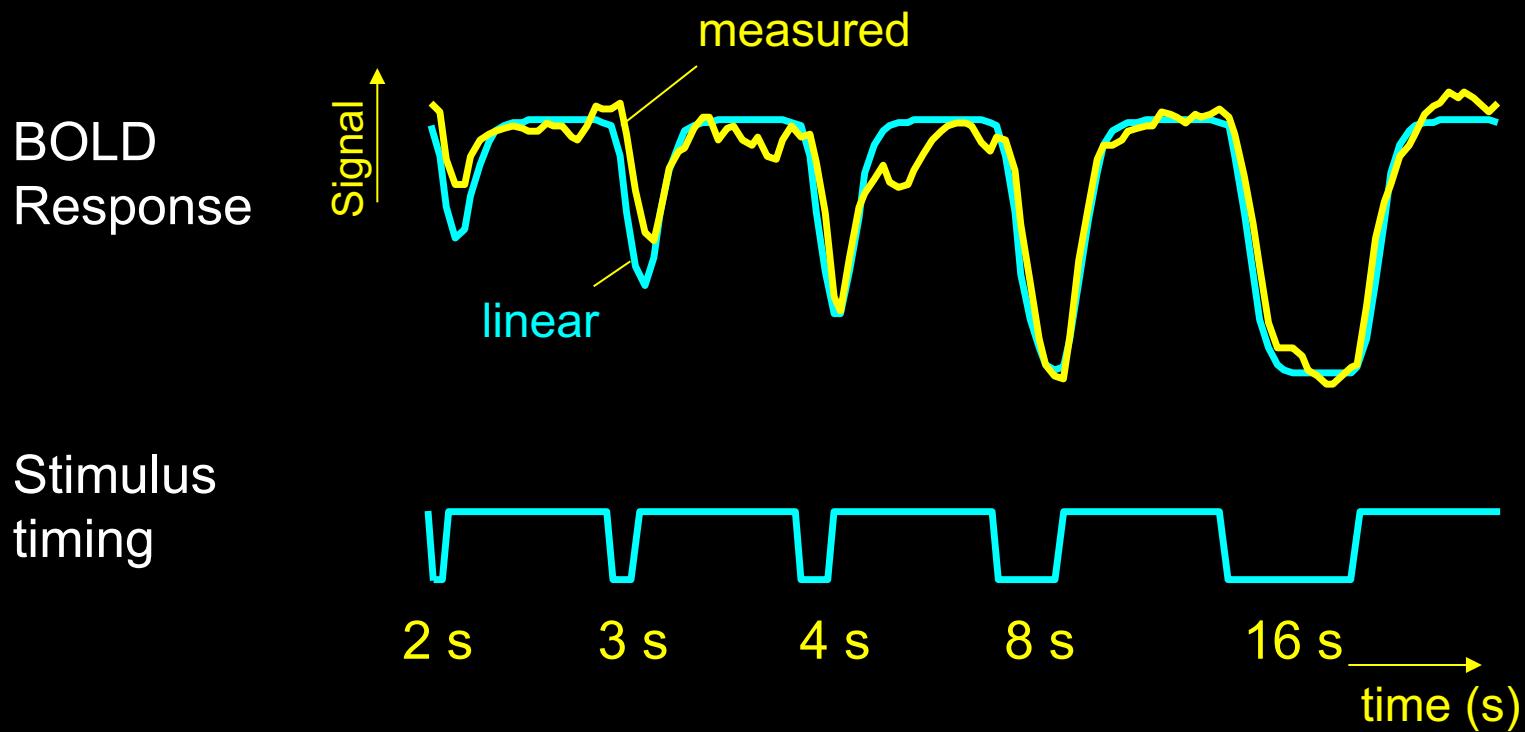


75% ON

Varying “ON” and “OFF” periods



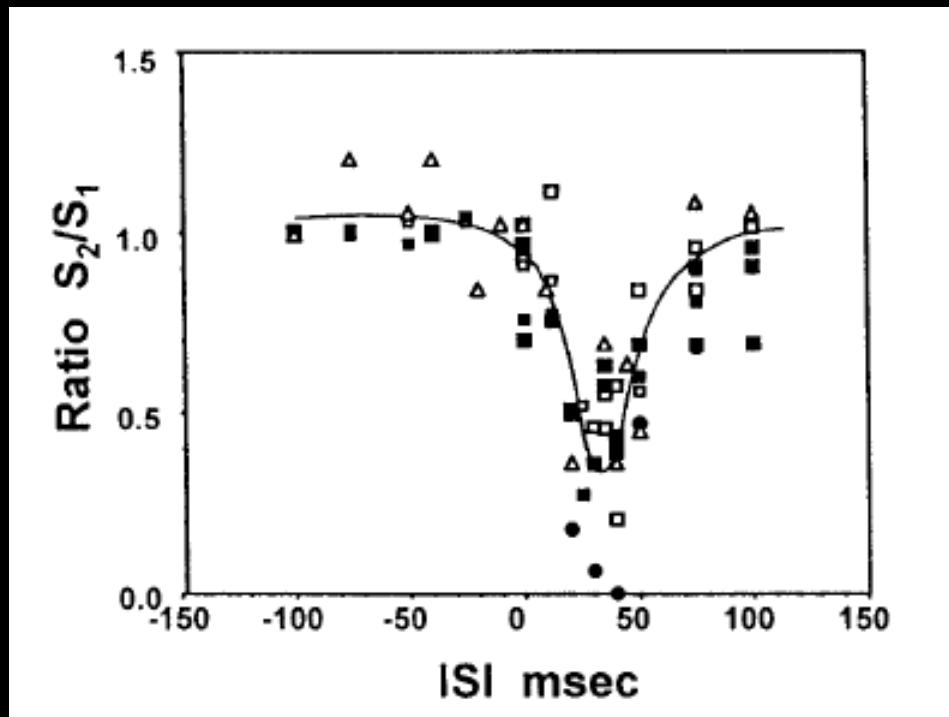
Different stimulus “OFF” periods



Brief stimulus OFF periods produce smaller decreases than expected

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

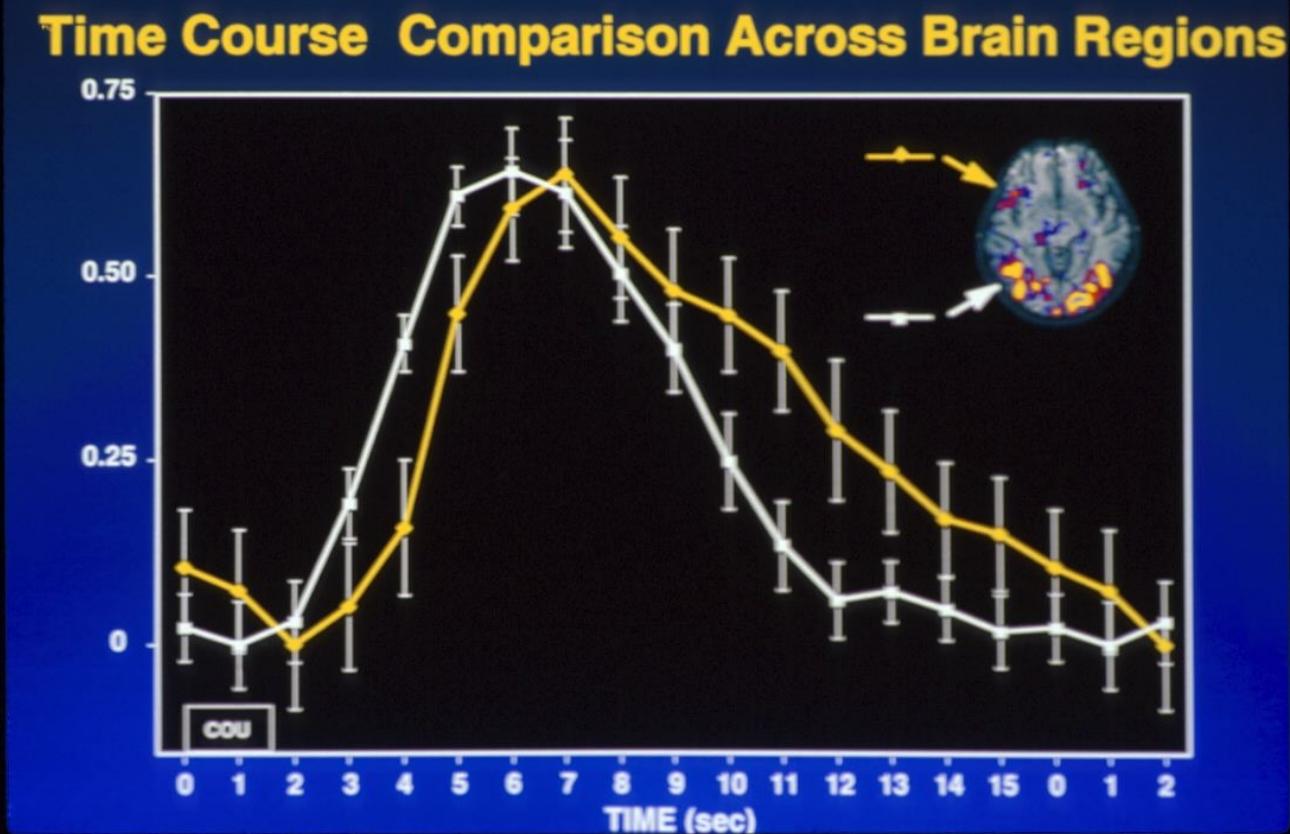
Seiji Ogawa^{†‡}, Tso-Ming Lee[†], Ray Stepnoski[†], Wei Chen[§], Xiao-Hong Zhu[§], and Kamil Ugurbil[§]



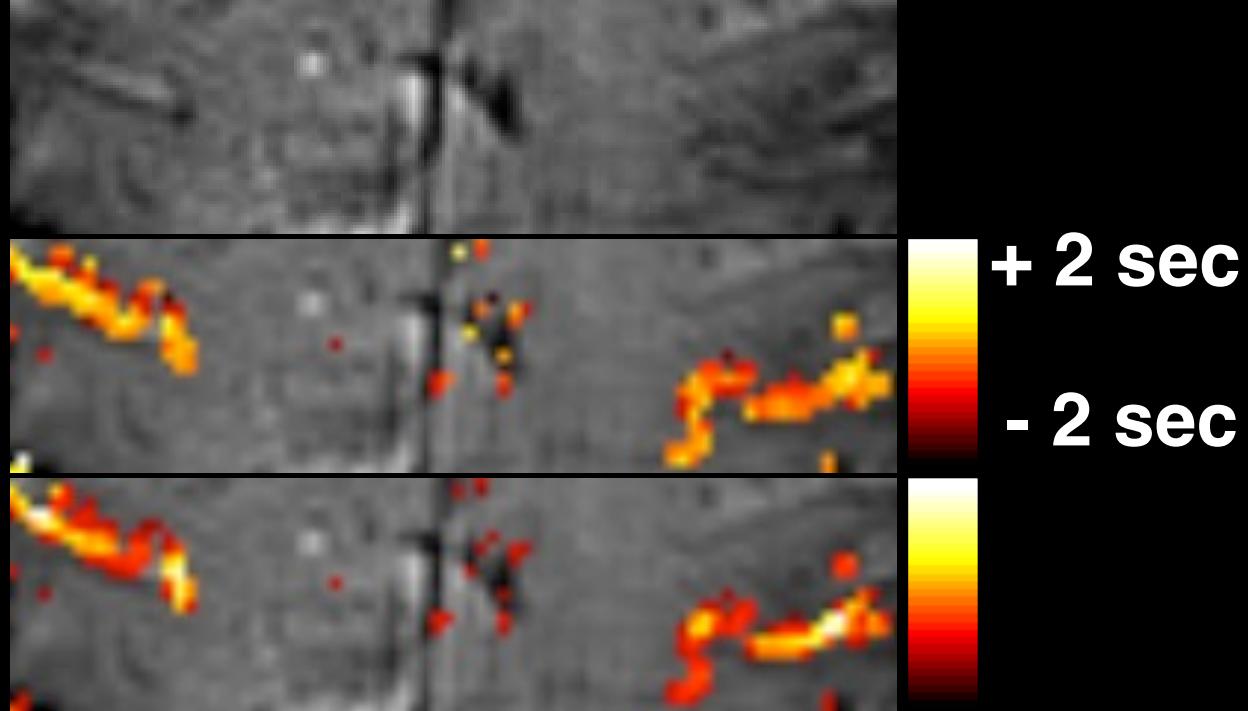
Detection of cortical activation during averaged single trials of a cognitive task using functional magnetic resonance imaging

(neuroimaging/single trial/language/prefrontal)

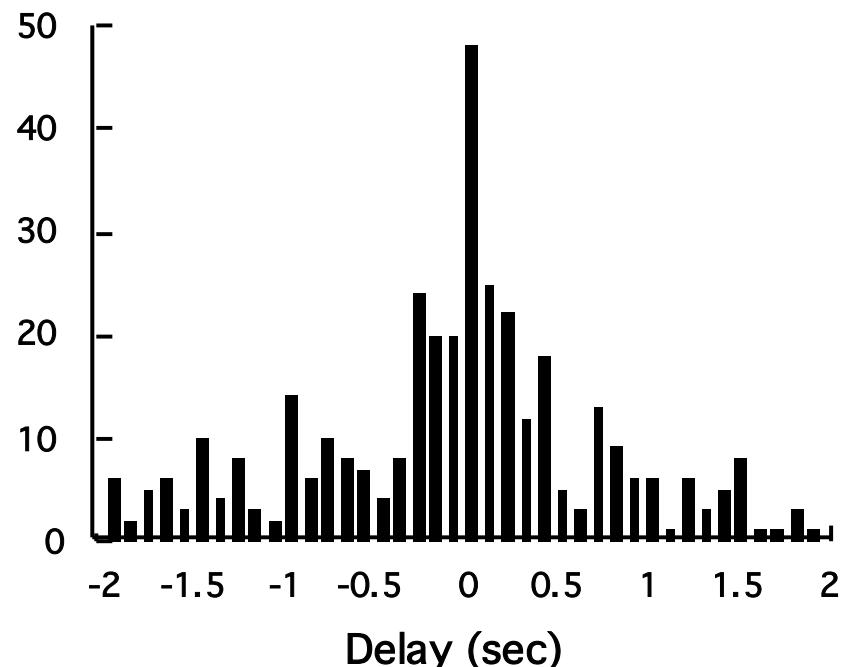
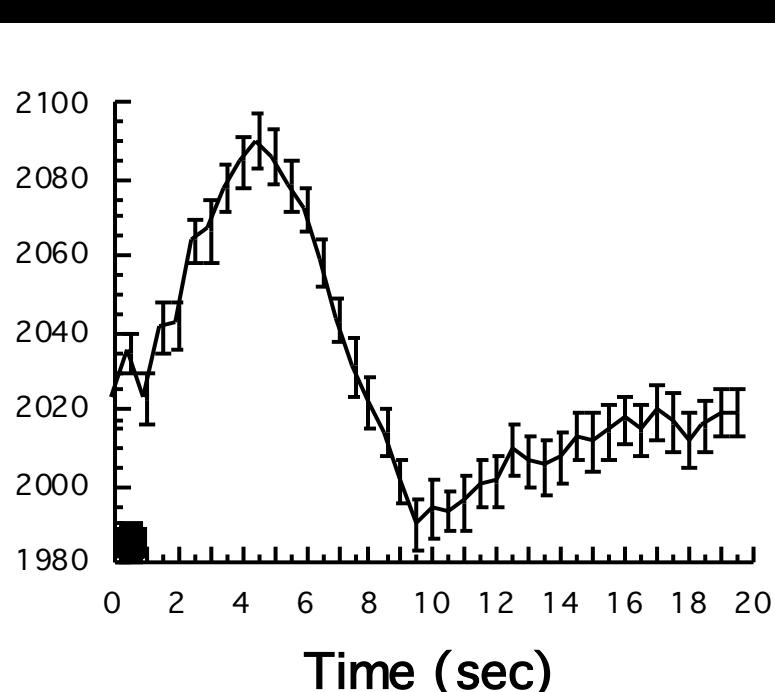
RANDY L. BUCKNER^{†‡§¶||}, PETER A. BANDETTINI^{†‡}, KATHLEEN M. O'CRAVEN^{†||}, ROBERT L. SAVOY^{†||},
STEVEN E. PETERSEN^{*++††}, MARCUS E. RAICHLE^{§++††}, AND BRUCE R. ROSEN^{†‡}



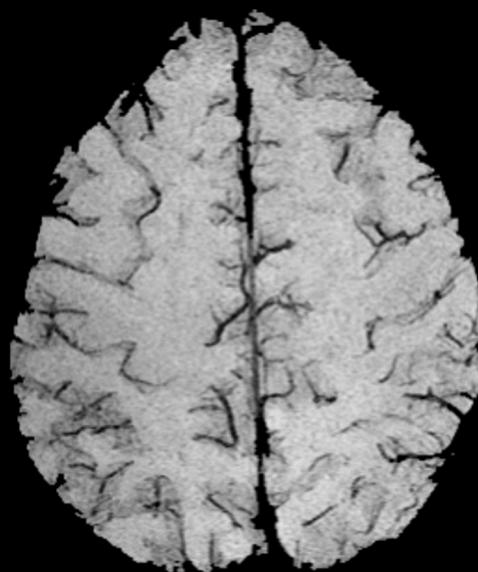
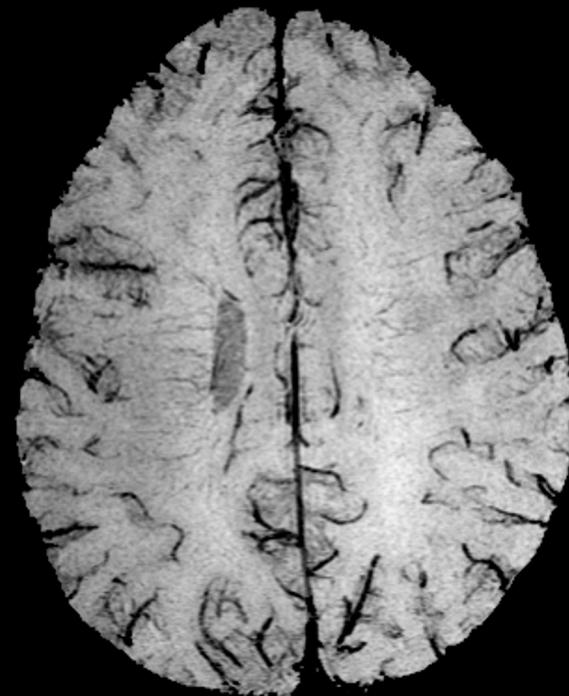
Latency

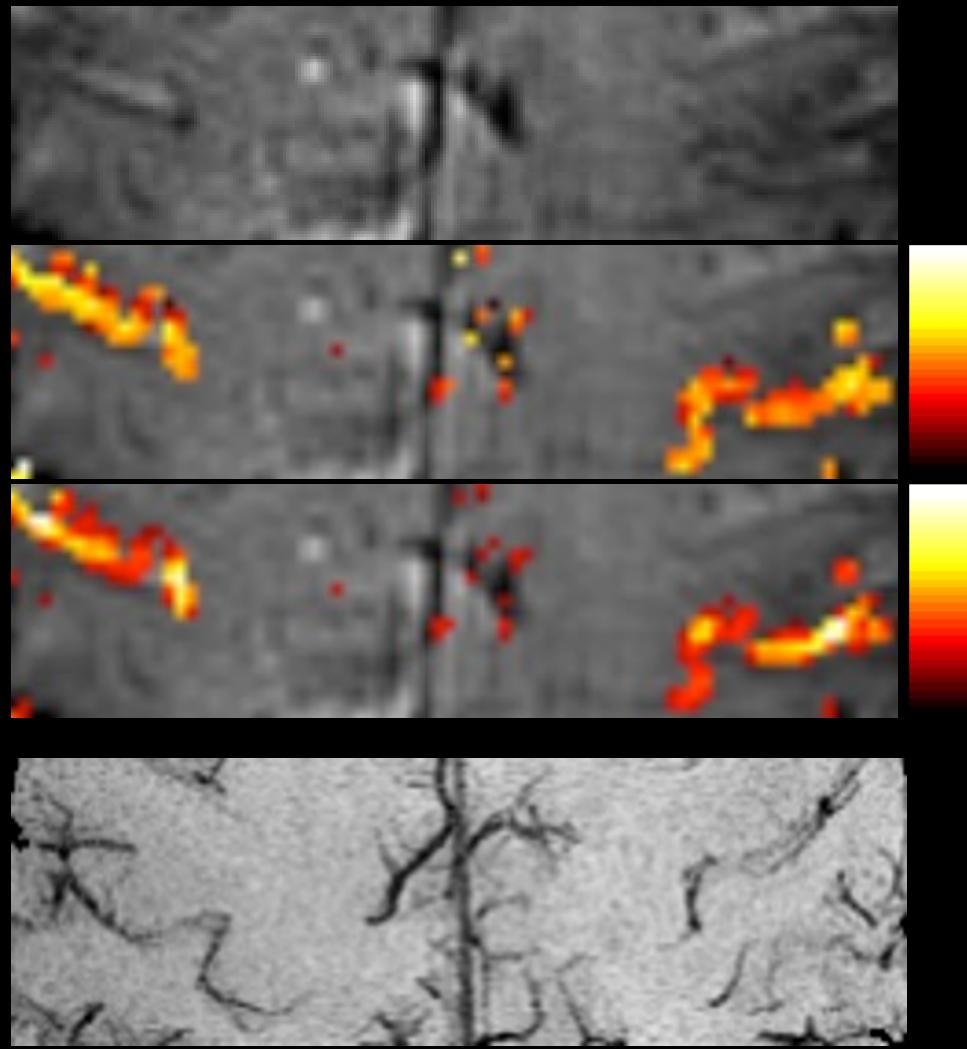


Magnitude



Venograms (3T)

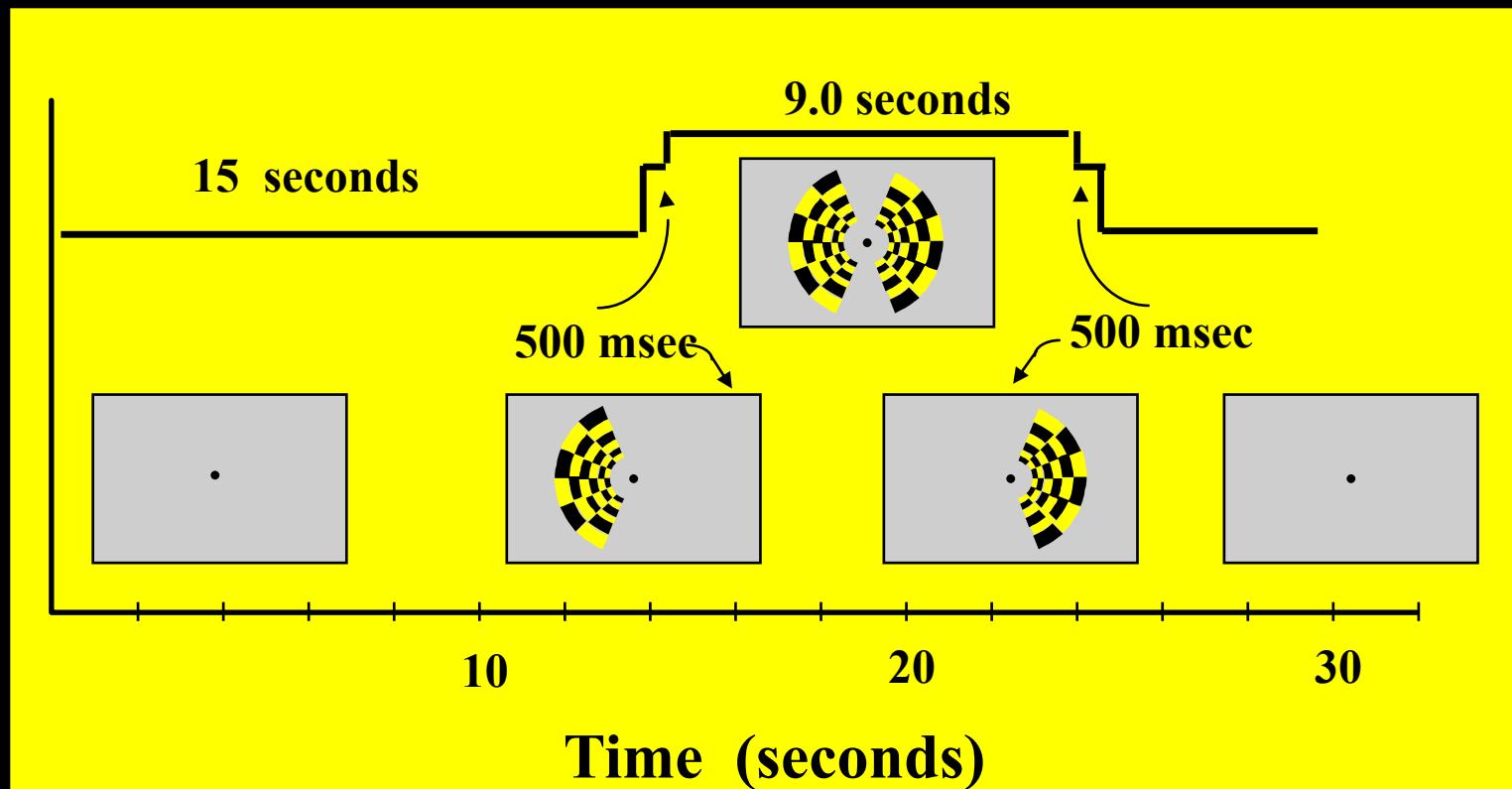


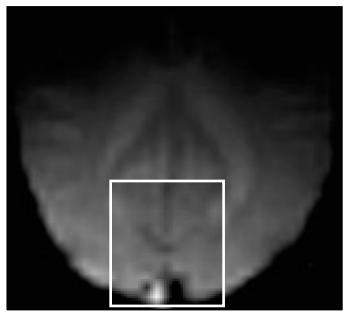


Hemi-Field Experiment

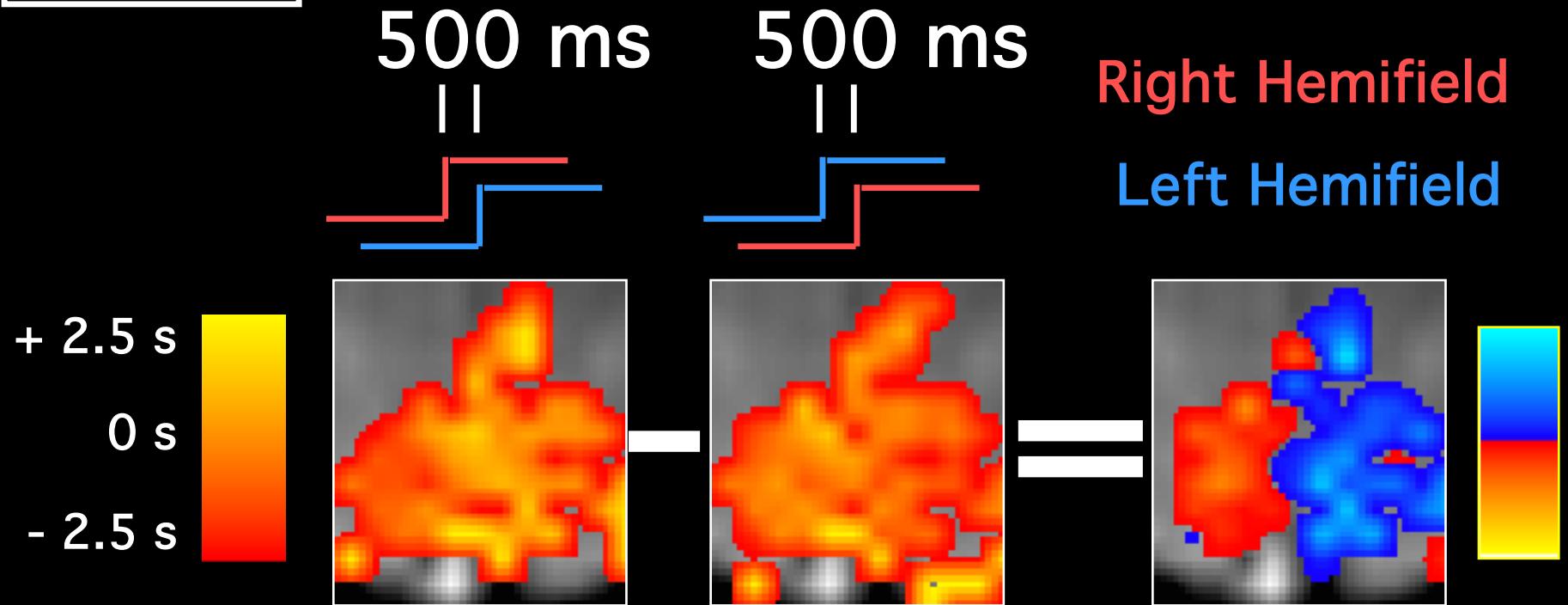
**Left
Hemisphere**

**Right
Hemisphere**

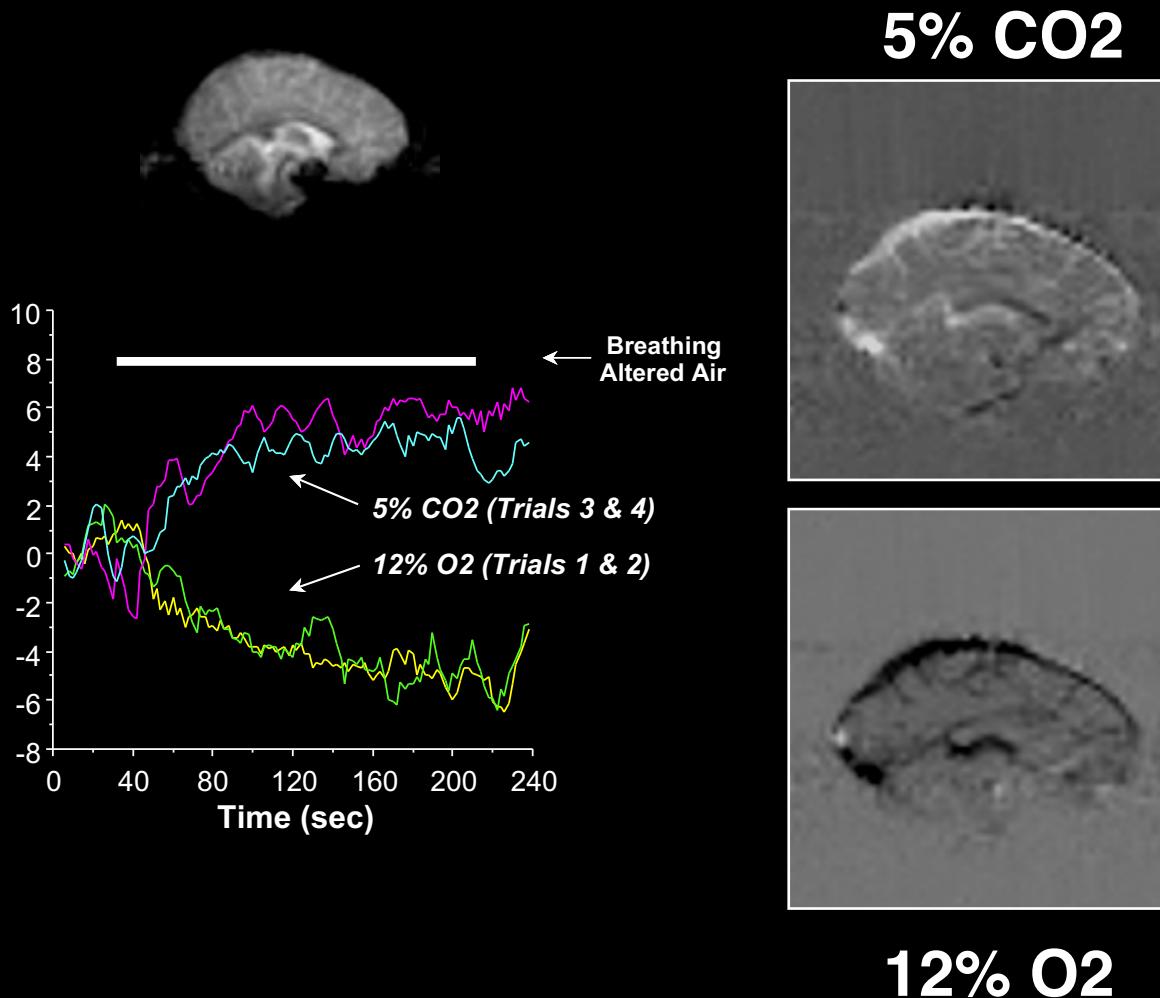




Calibration Techniques.....



Hemodynamic Stress Calibration

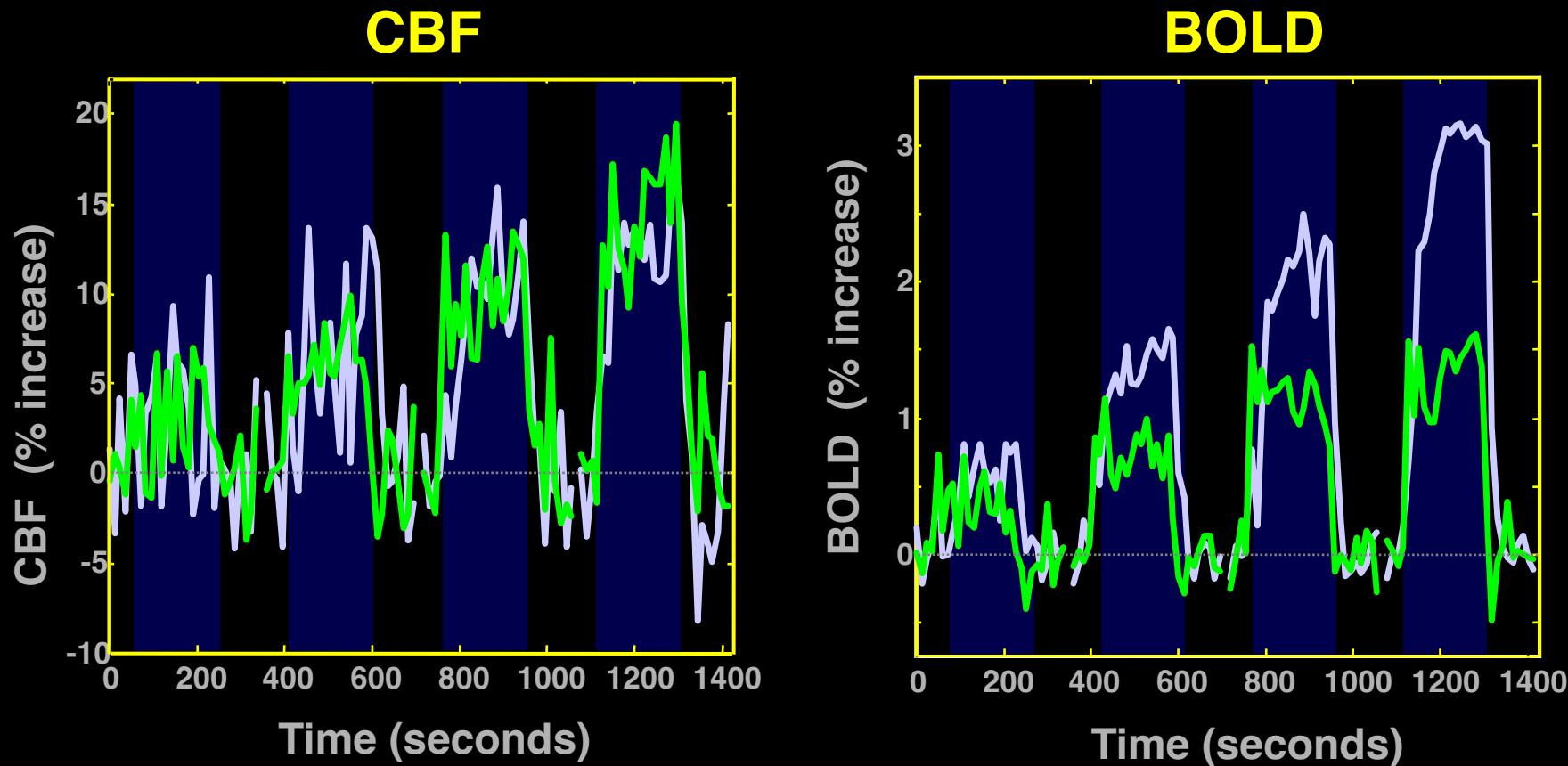


P. A. Bandettini, E. C. Wong, A hypercapnia - based normalization method for improved spatial localization of human brain activation with fMRI. *NMR in Biomedicine* 10, 197-203 (1997).

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*

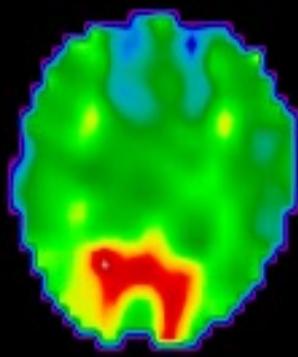
*Room WB325, McConnell Brain Imaging Centre, Montreal Neurological Institute, Quebec, Canada H3A 2B4; and [‡]Nuclear Magnetic Resonance Center, Massachusetts General Hospital, Building 149, 13th Street, Charlestown, MA 02129



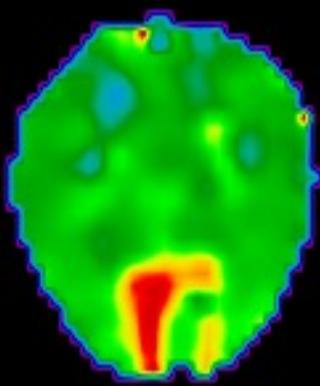
Simultaneous Perfusion and BOLD imaging during
graded visual activation and hypercapnia

N=12

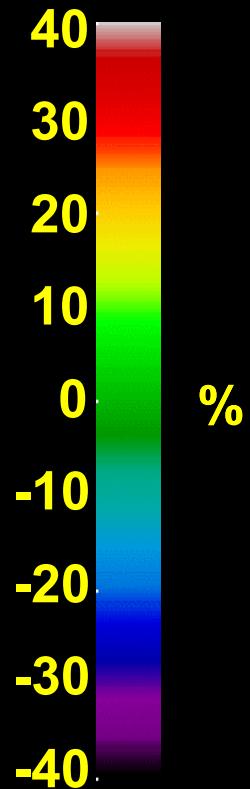
Computed CMRO₂ Changes



Subject 1

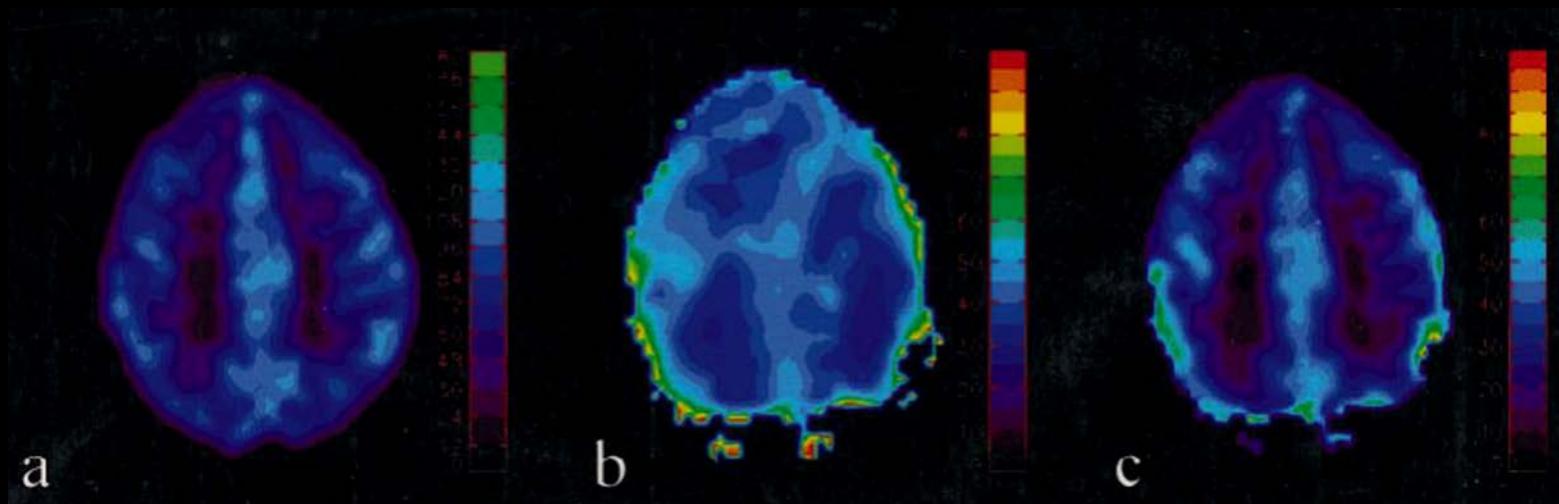


Subject 2



Quantitative measurements of cerebral metabolic rate of oxygen utilization using MRI: a volunteer study

Hongyu An,¹ Weili Lin,^{2*} Azim Celik³ and Yueh Z. Lee²

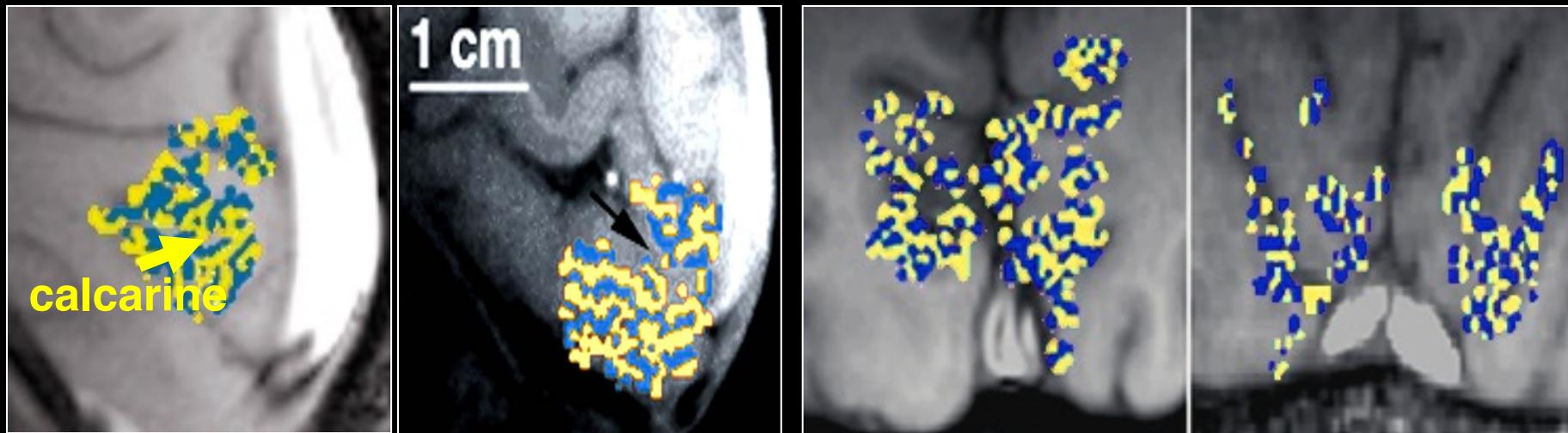


CBF

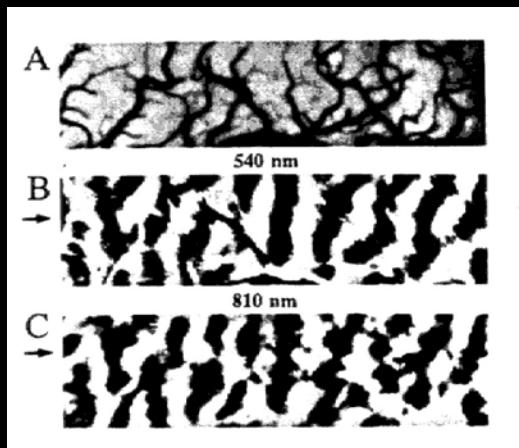
OEF

CMRO₂

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.

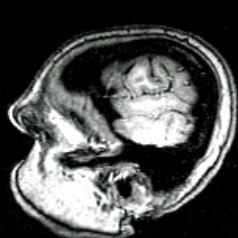
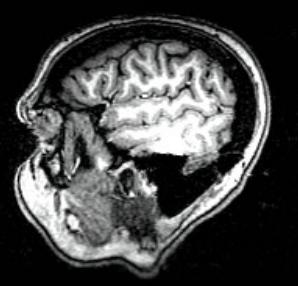
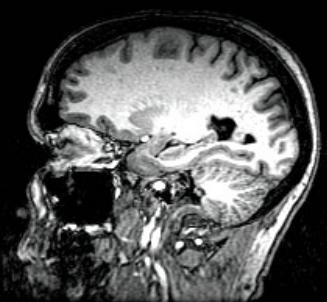
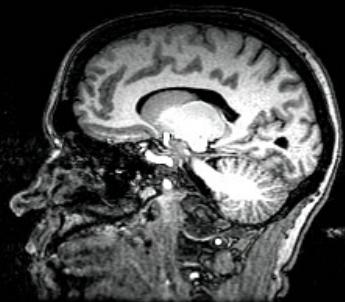
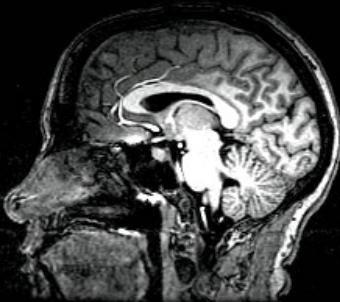
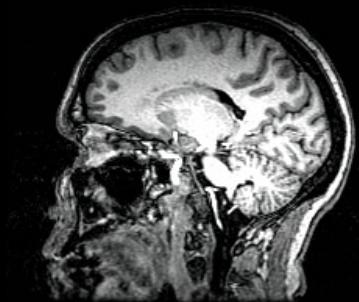


Optical Imaging

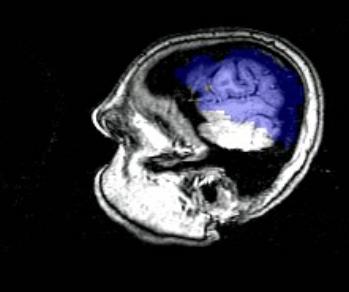
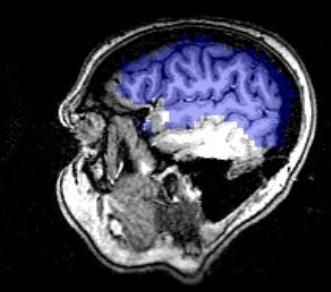
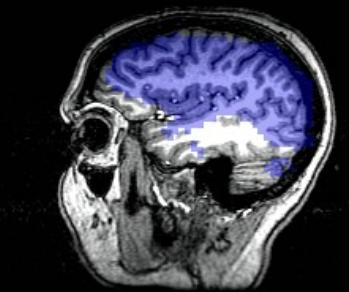
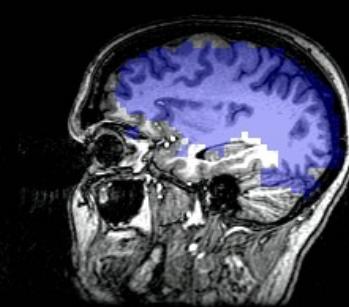
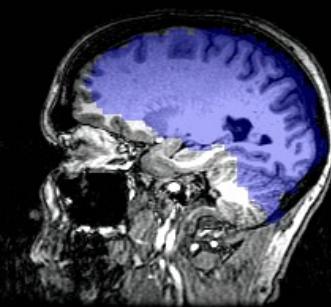
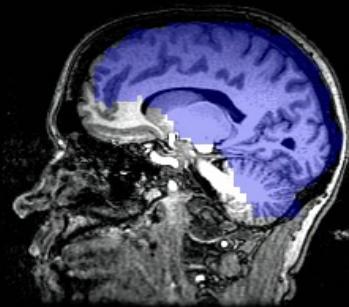
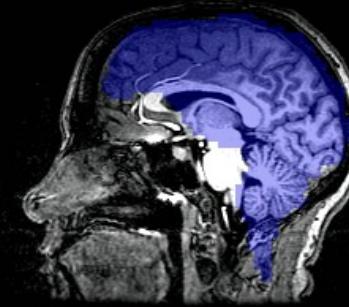
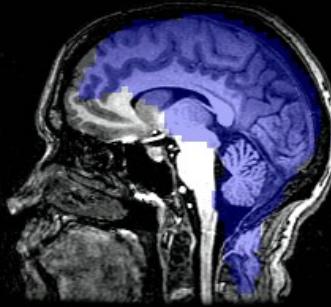
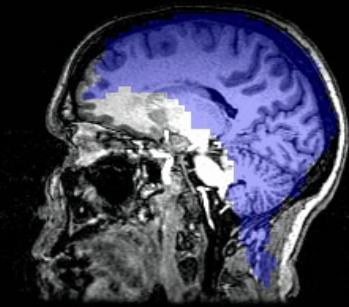
R. D. Frostig et. al, PNAS 87: 6082-6086, (1990).

Future....

- Shimming
- Acoustic Noise
- Multishot Techniques
- Increased Gradient Performance
- Higher Field Strengths
- Surface Coil Arrays
- Calibration / Quantification
- Embedded Functional Contrast
- Noise / Fluctuations
- Direct Neuronal Current Imaging
- Clinical Populations
- Neuronal, Vascular, and Metabolic Information

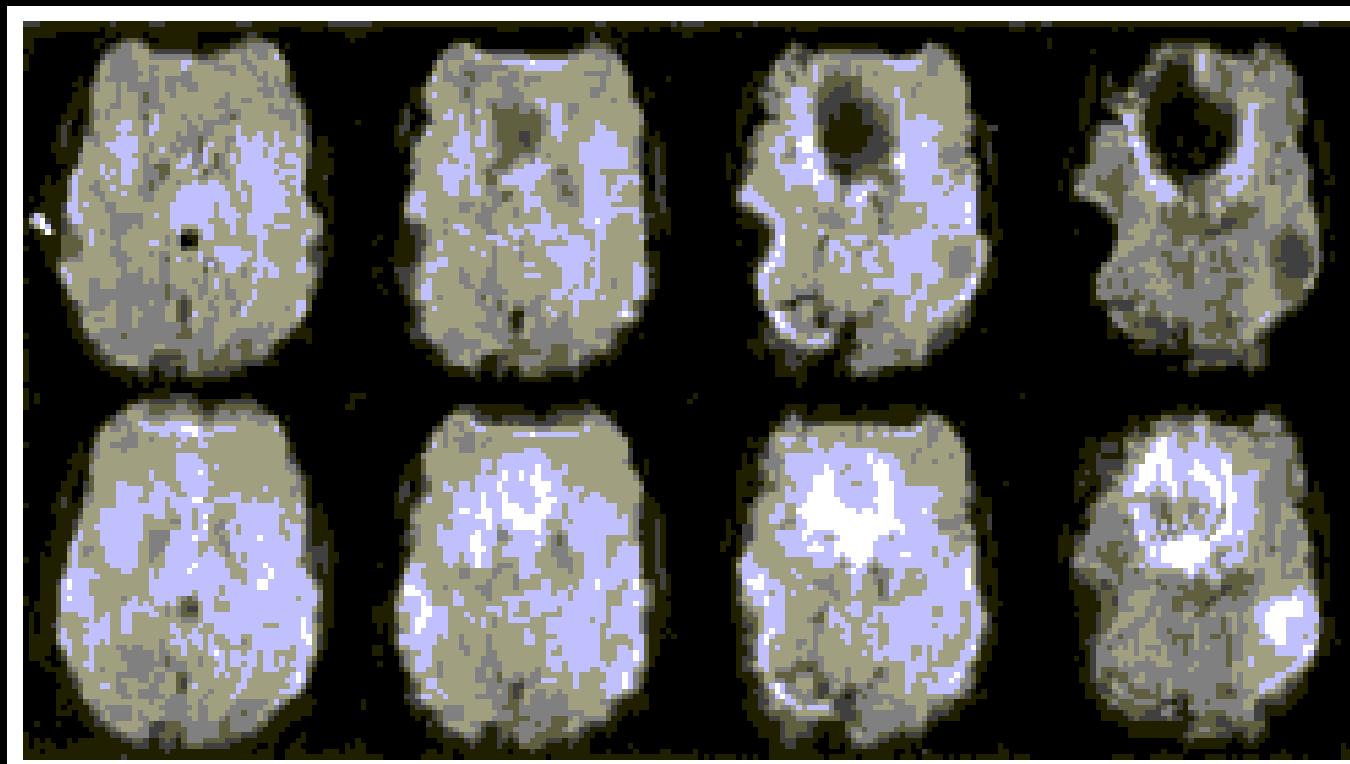






3D z-Shim Method for Reduction of Susceptibility Effects in BOLD fMRI

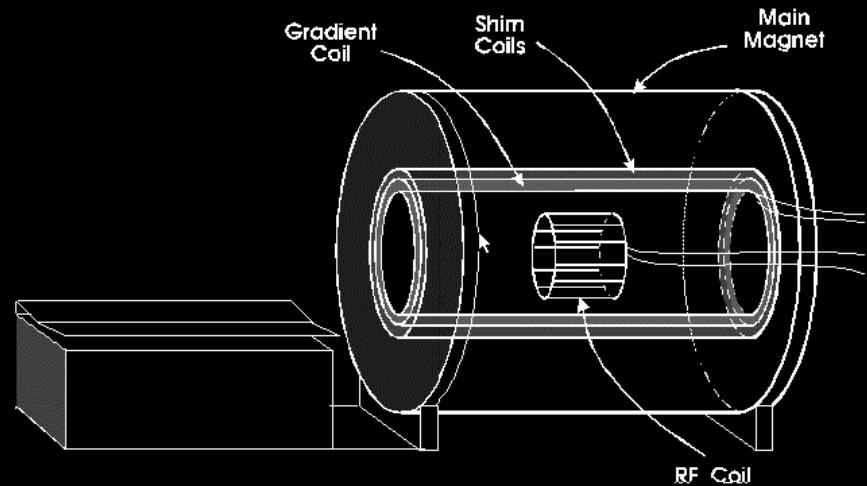
Gary H. Glover*



2 G/cm, 350 T/m/s



4 G/cm, 150 T/m/s



10 G/cm, 1000 T/m/s



Diffusion imaging
Faster imaging
Higher resolution

Neuronal Current Imaging

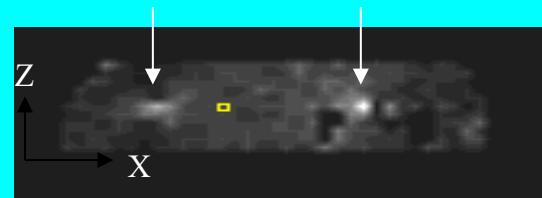
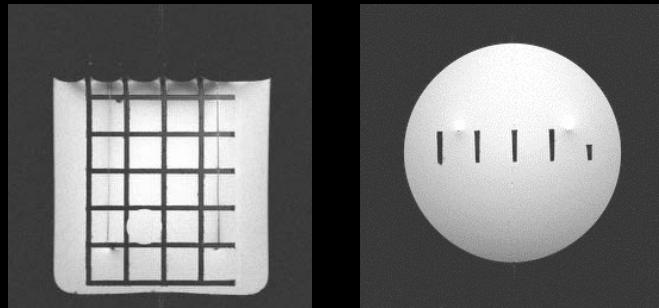
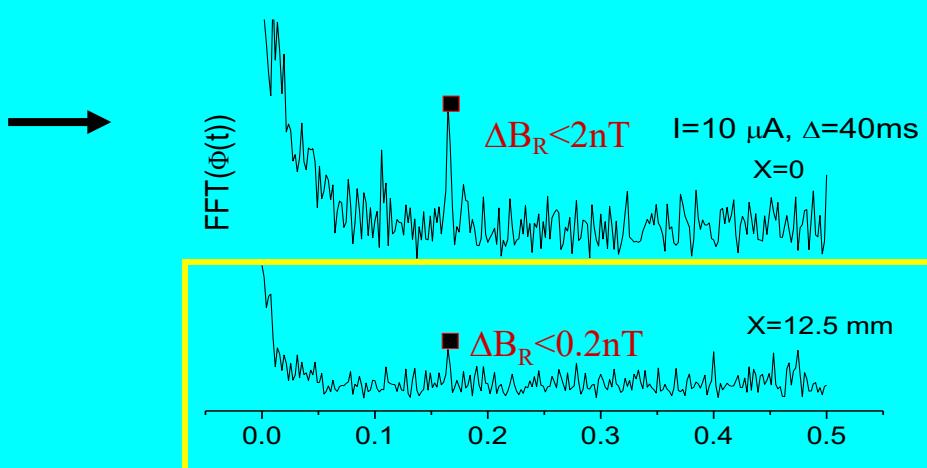


Figure 1



FIM Unit & FMRI Core Facility

Director:

Peter Bandettini

Staff Scientists:

Sean Marrett

Jerzy Bodurka

Frank Ye

Wen-Ming Luh

Computer Specialist:

Adam Thomas

Post Docs:

Rasmus Birn

Hauke Heekeren

David Knight

Patrick Bellgowan

Ziad Saad

Graduate Student:

Natalia Petridou

Post-Back. IRTA Students:

Elisa Kapler

August Tuan

Dan Kelley

Visiting Fellows:

Sergio Casciaro

Marta Maierov

Guosheng Ding

Clinical Fellow:

James Patterson

Psychologist:

Julie Frost

Summer Students:

Hannah Chang

Courtney Kemps

Douglass Ruff

Carla Wettig

Kang-Xing Jin

Program Assistant:

Kay Kuhns

Scanning Technologists:

Karen Bove-Bettis

Paula Rowser