

fMRI: Past, Present, and Future

Peter A. Bandettini, Ph.D

Unit on Functional Imaging Methods
&
3T Neuroimaging Core Facility

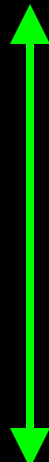
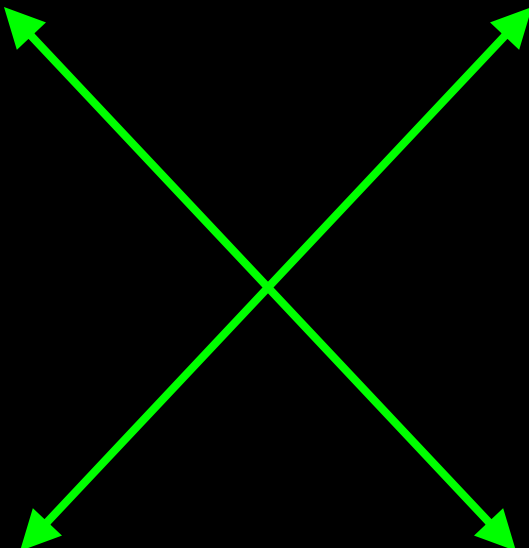
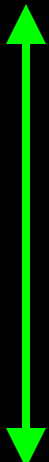
Laboratory of Brain and Cognition
National Institute of Mental Health



Technology



Methodology



Interpretation



Applications

Technology

Methodology

Engineers

Statisticians

Physicists

Mathematicians

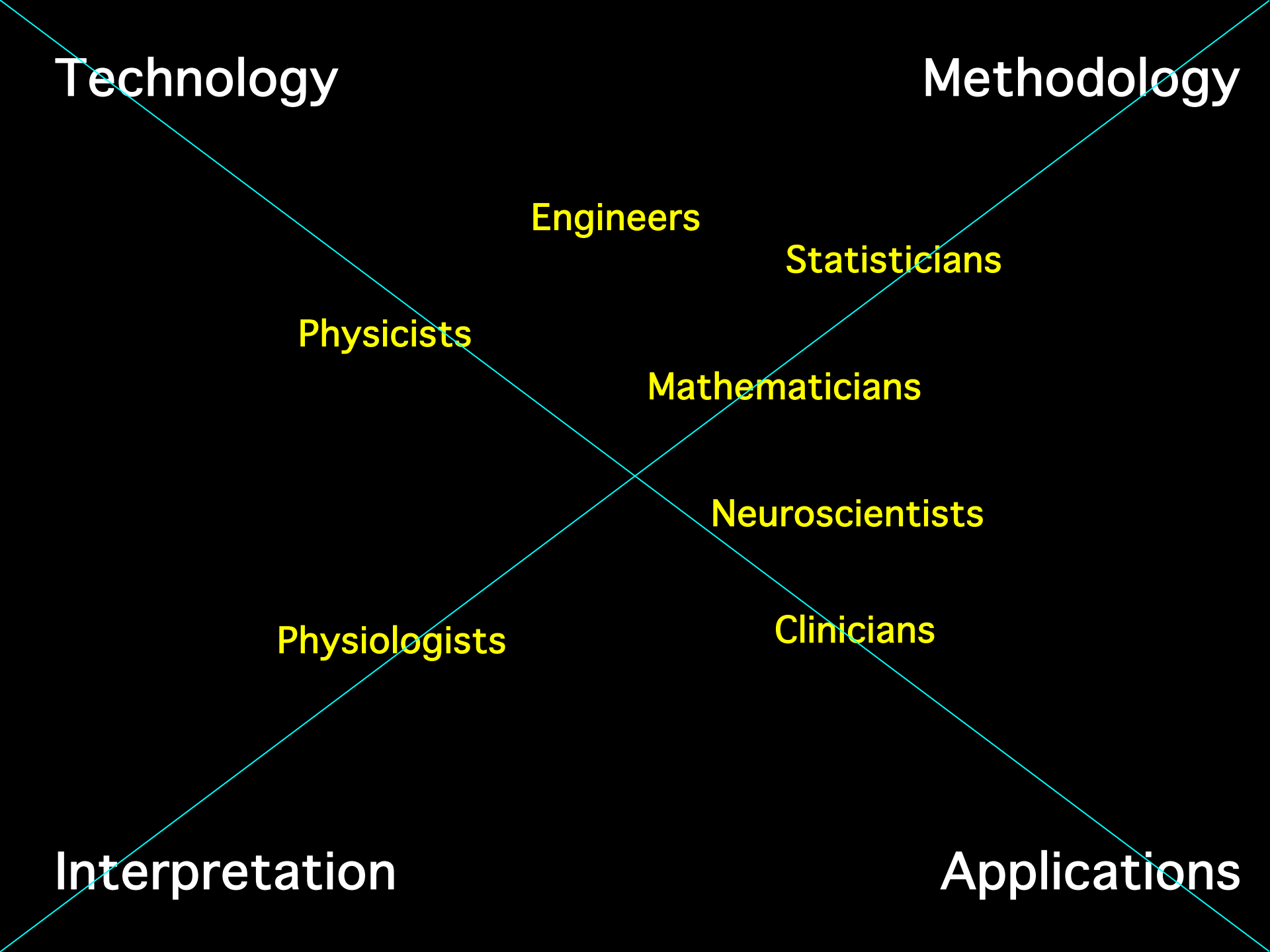
Neuroscientists

Physiologists

Clinicians

Interpretation

Applications



Technology

MRI

1.5T,3T, 4T

EPI

Local Human Head Gradient Coils

ASL

BOLD

EPI on Clin. Syst.

Nav. pulses

Spiral EPI

Multi-shot fMRI

Diff. tensor

Real time fMRI

Quant. ASL

Dynamic IV volume

Simultaneous ASL and BOLD

Mg⁺

Venography

Z-shim

Baseline Susceptibility

7T

SENSE

Current Imaging?

Methodology

Baseline Volume

IVIM

Correlation Analysis

Parametric Design

Surface Mapping

Phase Mapping

Linear Regression

Event-related

Motion Correction

Multi-Modal Mapping

Free-behavior Designs

Mental Chronometry

Deconvolution

CO₂ Calibration

Interpretation

Blood T2

Hemoglobin

BOLD models

B₀ dep.

TE dep

SE vs. GE

NIRS Correlation

Veins

PET correlation

IV vs EV

Pre-undershoot

Resolution Dep.

Post-undershoot

CO₂ effect

NIRS Correlation

Inflow

ASL vs. BOLD

PSF of BOLD

Extended Stim.

Linearity

Fluctuations

Balloon Model

Metab. Correlation

Optical Im. Correlation

Electrophys. correlation

Applications

Complex motor Language

Imagery

Memory

Emotion

Motor learning

Children

Tumor vasc.

Drug effects

BOLD -V1, M1, A1

Presurgical

Attention

Ocular Dominance

Volume - Stroke

V1, V2..mapping

Priming/Learning

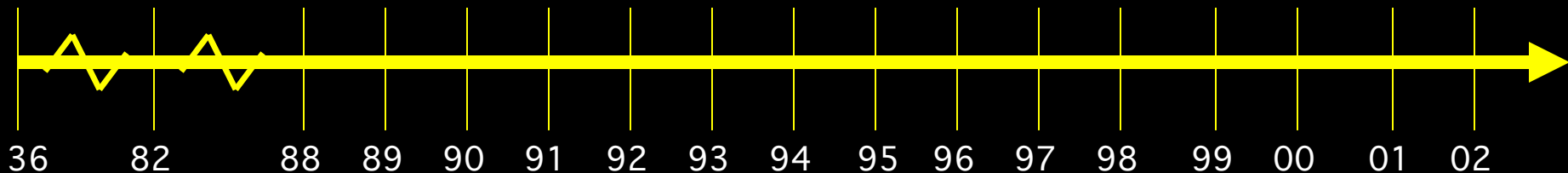
Clinical Populations

Δ Volume-V1

Plasticity

Face recognition

Performance prediction



Technology

MRI

EPI 1.5T,3T, 4T EPI on Clin. Syst. Diff. tensor Mg⁺ 7T

Local Human Head Gradient Coils Nav. pulses Real time fMRI Venography SENSE

ASL Spiral EPI Quant. ASL Dynamic IV volume Z-shim Baseline Susceptibility

BOLD Multi-shot fMRI Simultaneous ASL and BOLD Current Imaging?

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Correlation Analysis Motion Correction CO₂ Calibration

Parametric Design Surface Mapping Multi-Modal Mapping

Phase Mapping Free-behavior Designs

Linear Regression Mental Chronometry

Event-related Deconvolution

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Blood T2

Hemoglobin

BOLD models PET correlation

B₀ dep. IV vs EV ASL vs. BOLD

TE dep. Resolution Dep. Pre-undershoot PSF of BOLD

Post-undershoot Extended Stim. Metab. Correlation

SE vs. GE CO₂ effect Linearity

NIRS Correlation Fluctuations Optical Im. Correlation

Veins Inflow Balloon Model Electrophys. correlation

Applications

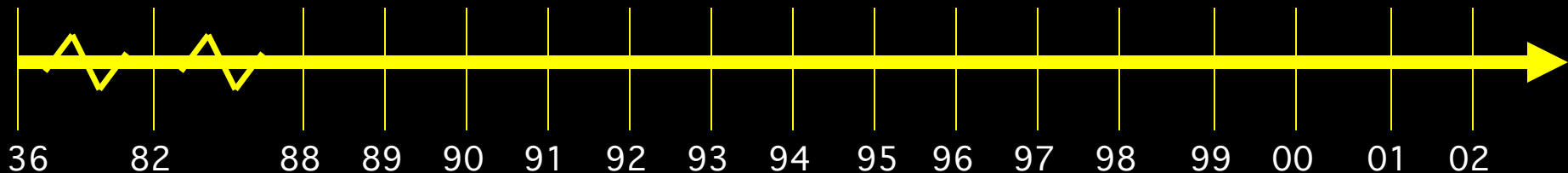
Complex motor Language Imagery Memory Emotion

Motor learning Children Tumor vasc. Drug effects

BOLD -V1, M1, A1 Presurgical Attention Ocular Dominance

Volume - Stroke V1, V2..mapping Priming/Learning Clinical Populations

Δ Volume-V1 Plasticity Face recognition Performance prediction





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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 ASL
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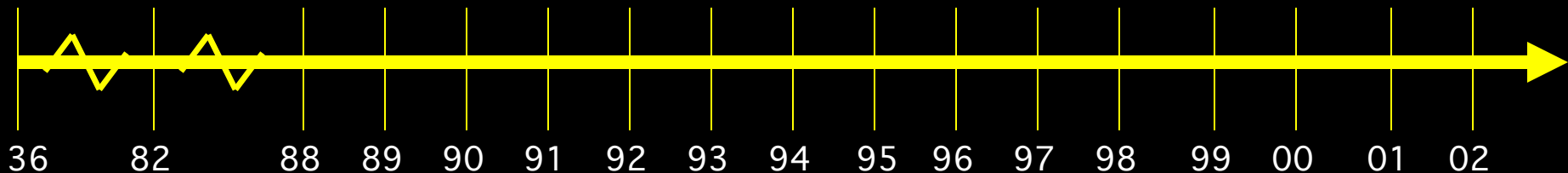
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$$f_0 = \gamma B_0$$

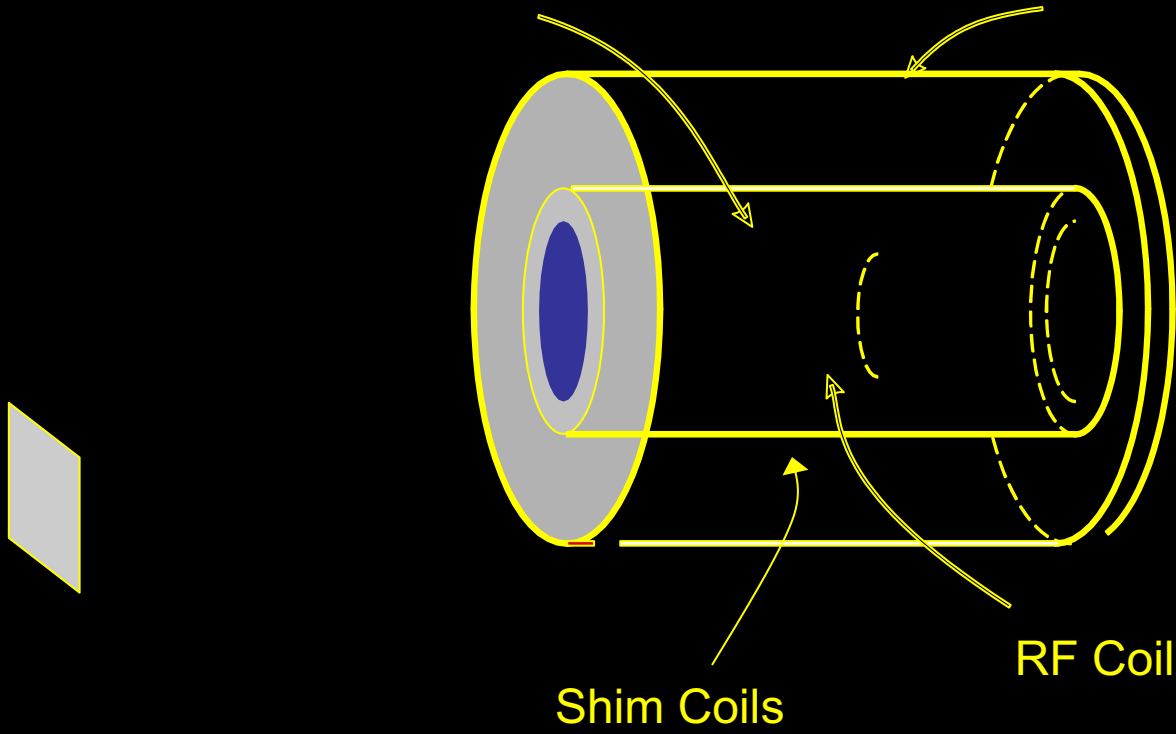
f_0 = Larmor frequency

γ = gyromagnetic ratio (42.6 MHz/Tesla)

B_0 = magnetic field strength (Tesla)

Gradient coil

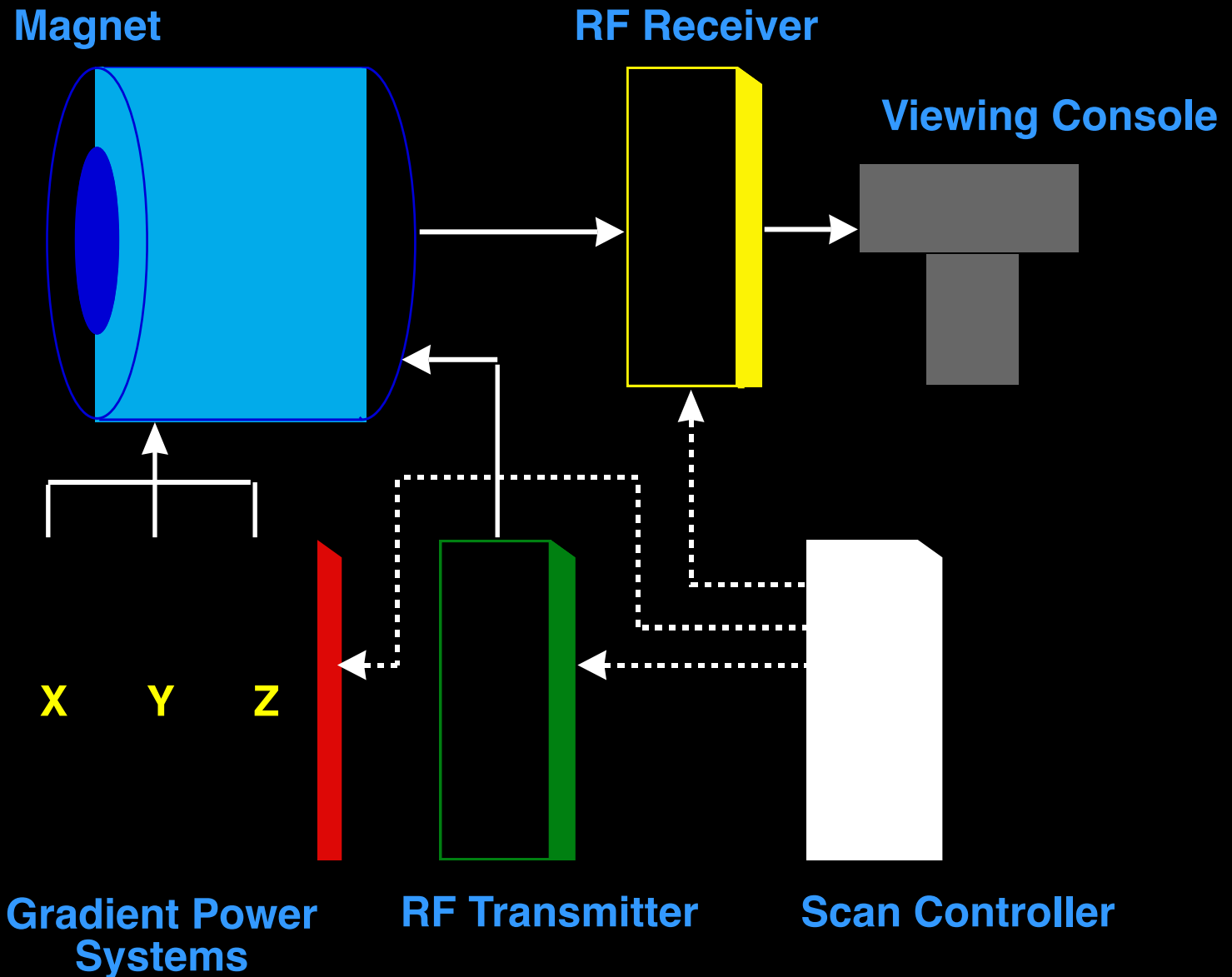
Main Magnet



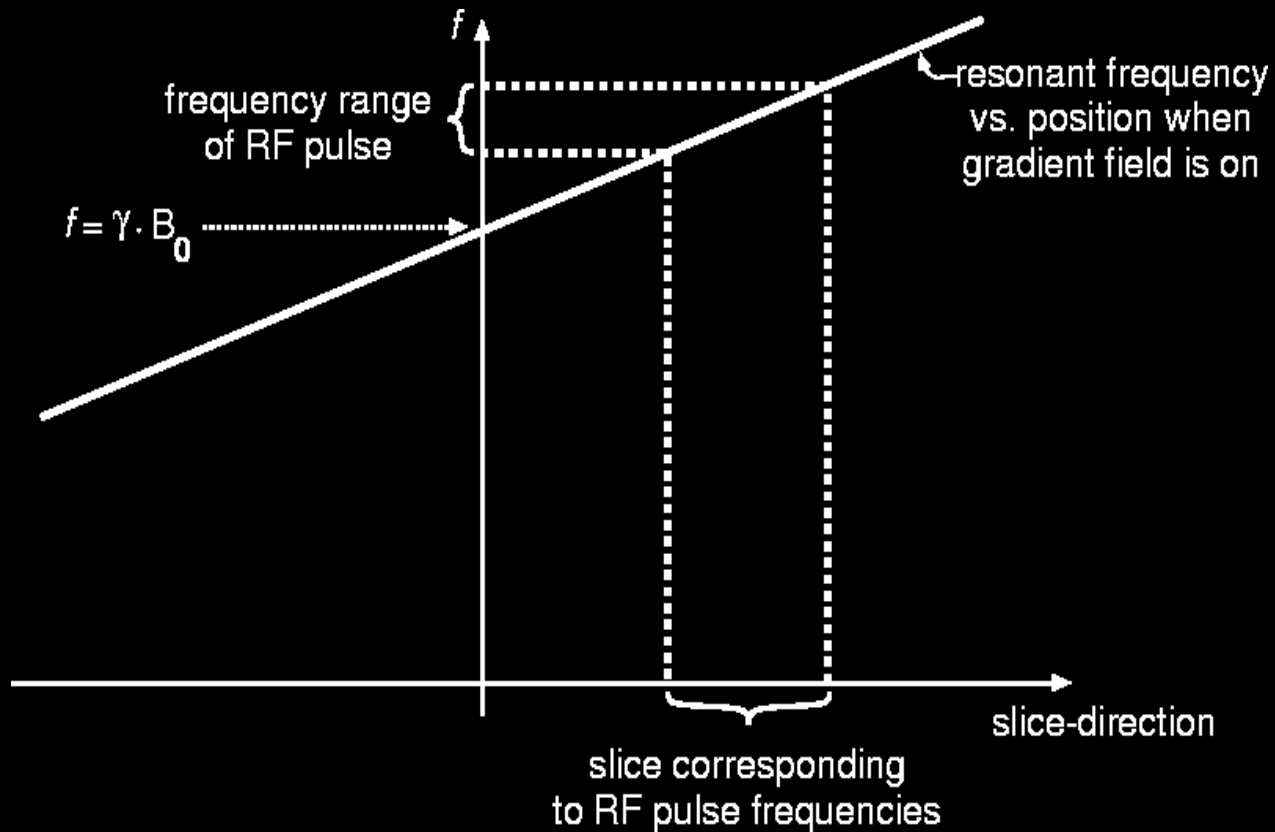
Shim Coils

RF Coil

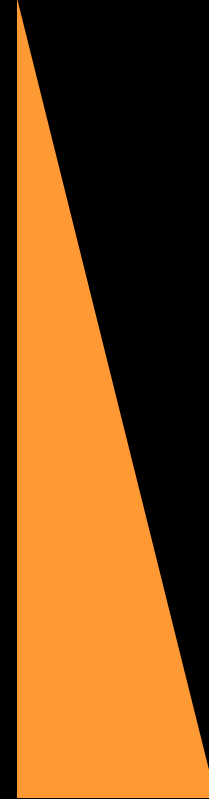
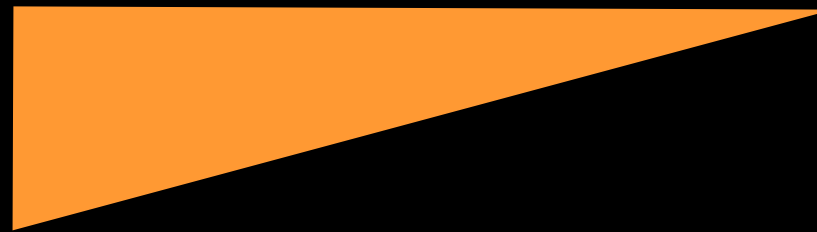
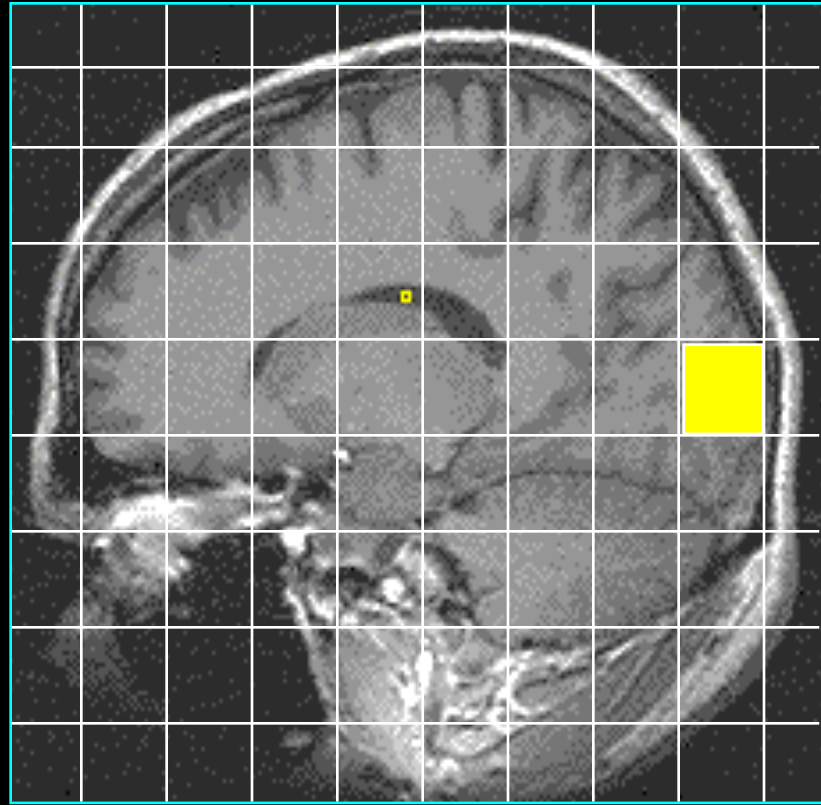
Imaging System Components

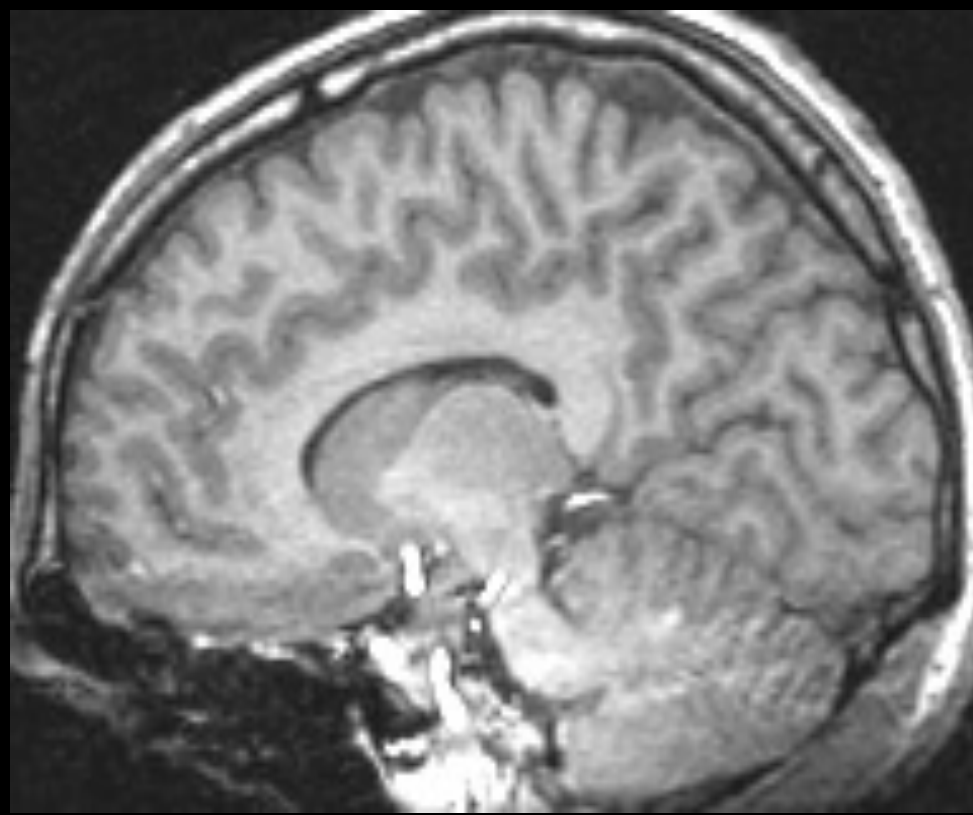
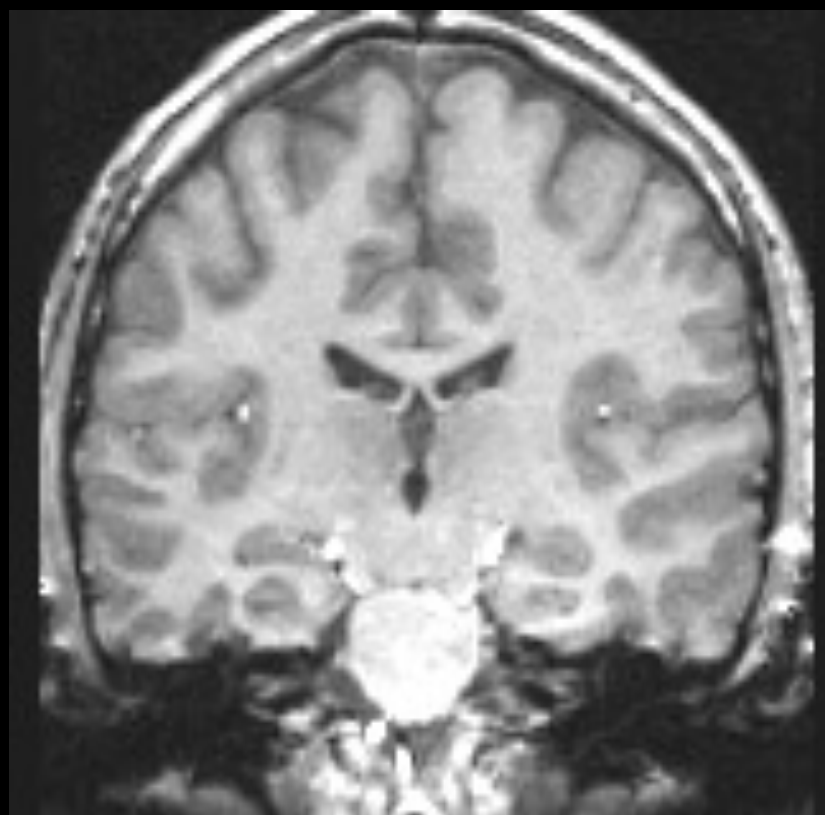


Slice Selection



In – Plane Spatial Localization





Source of Anatomical Contrast: MRI Parameters

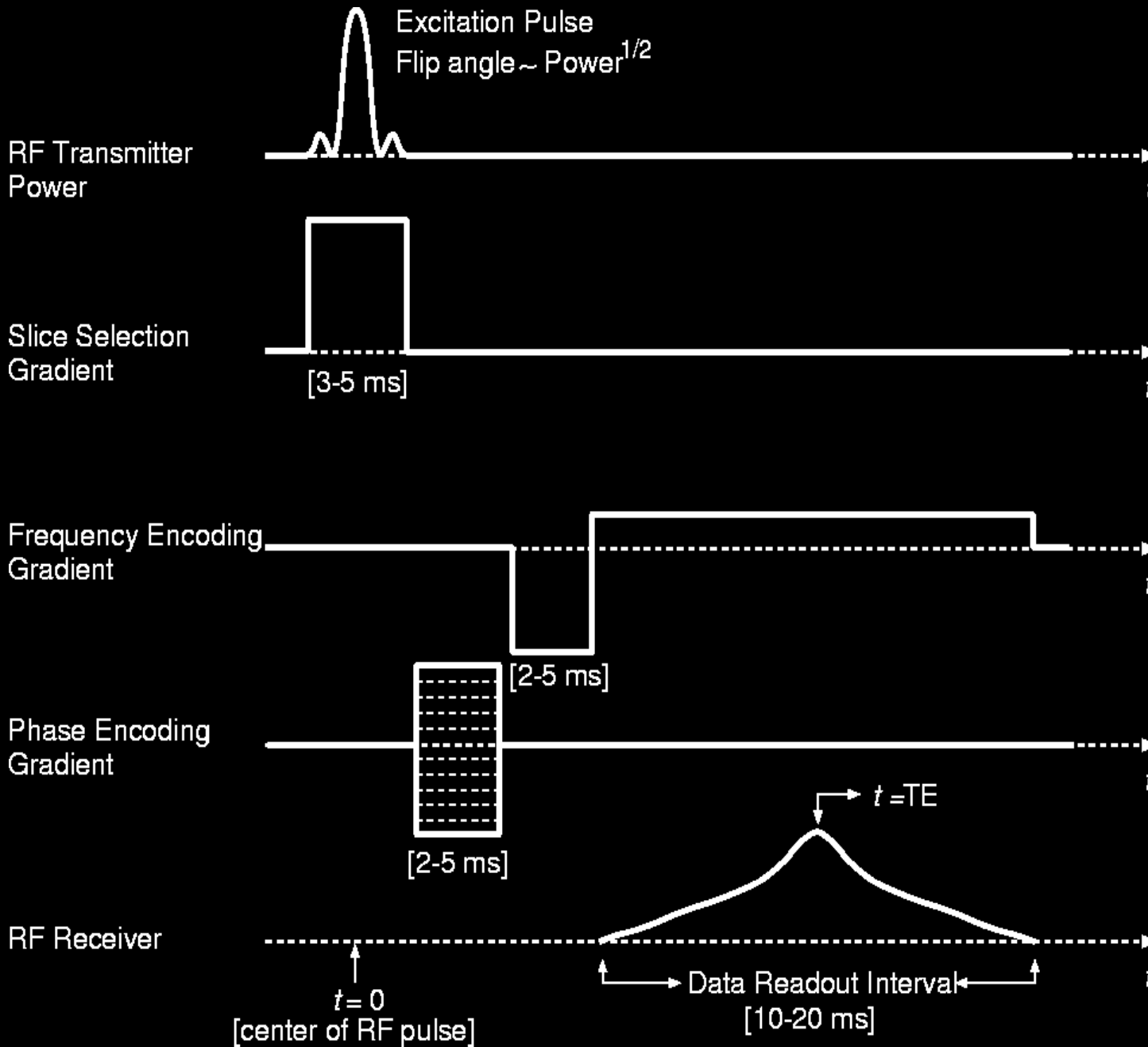


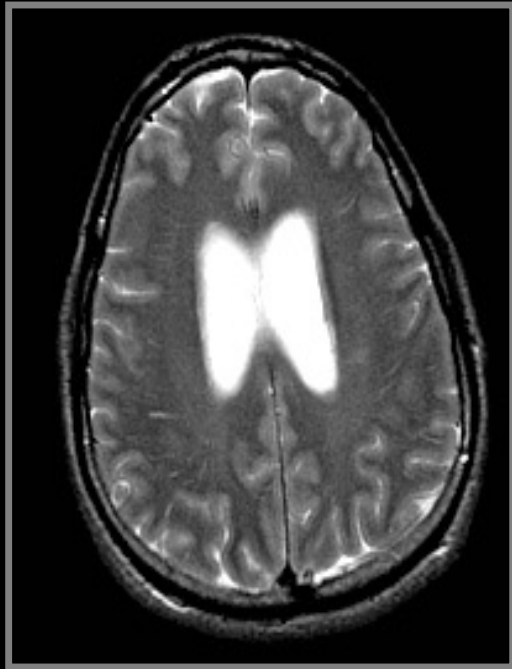
- Spin (Proton) Density
- T1 Relaxation Time
- T2, T2* Relaxation Times

MRI Pulse Sequence for Gradient Echo Imaging

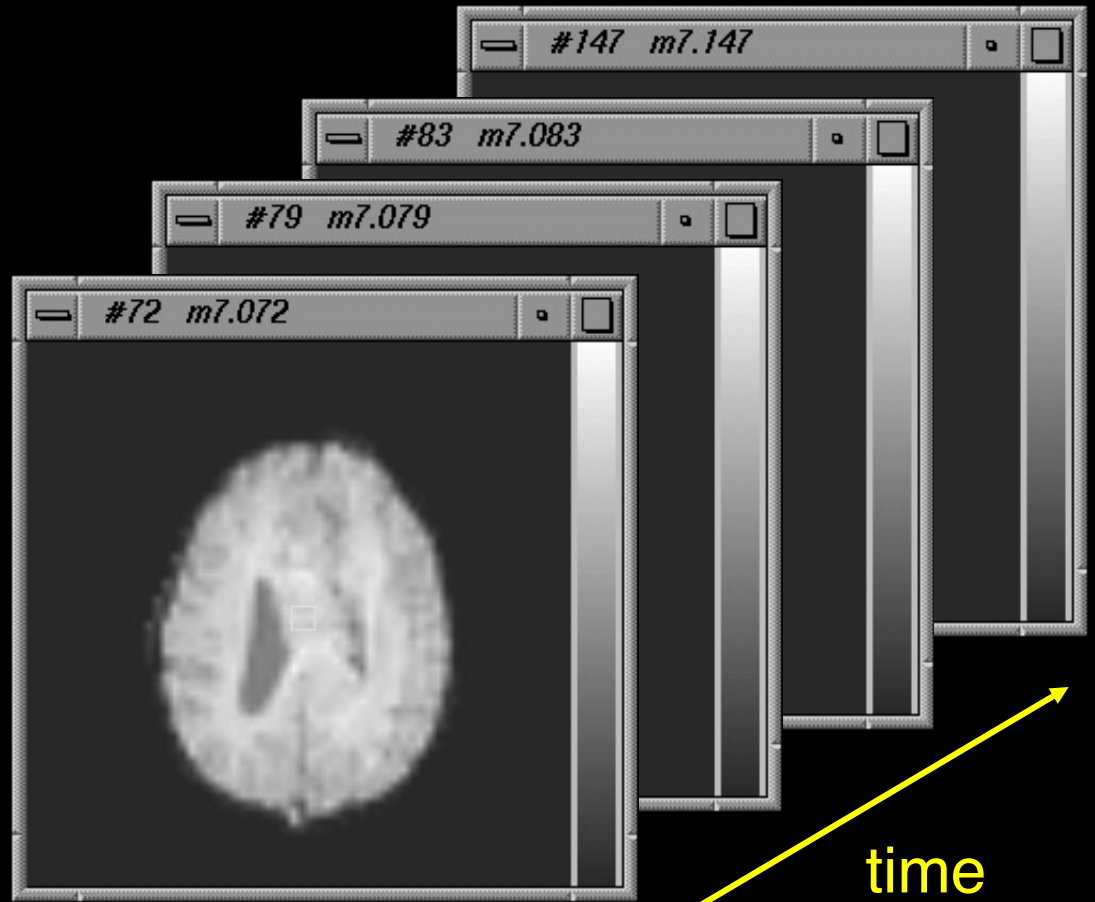
Illustrates sequence of events during scanning

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



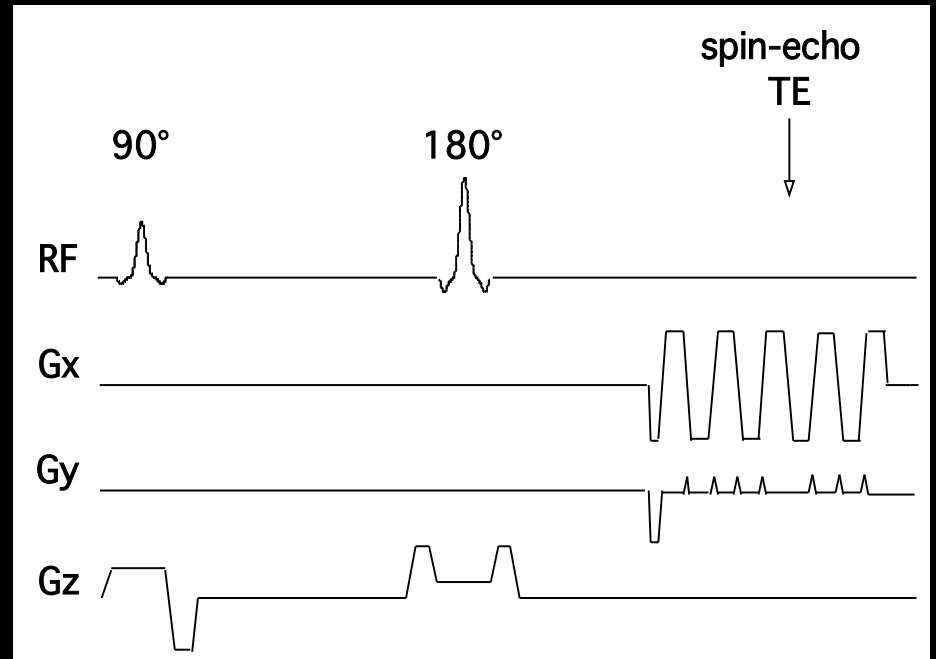
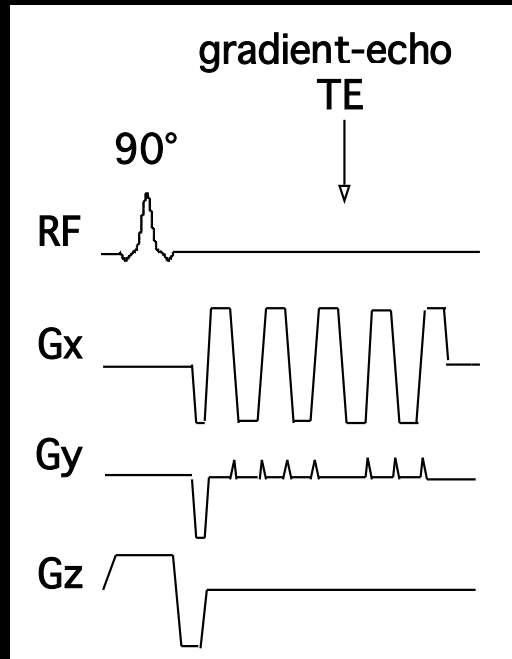


Anatomic

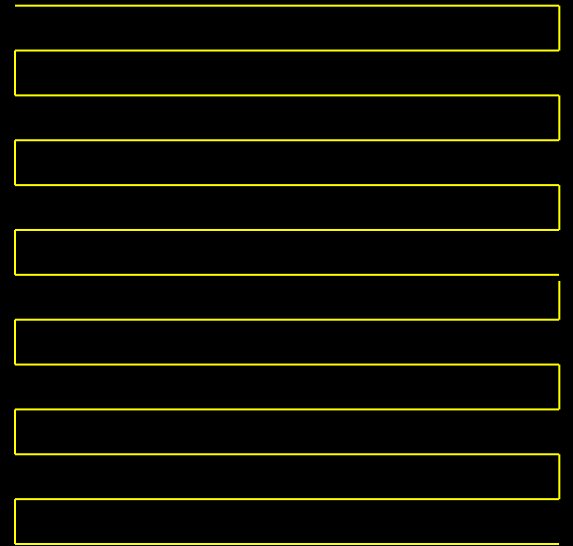
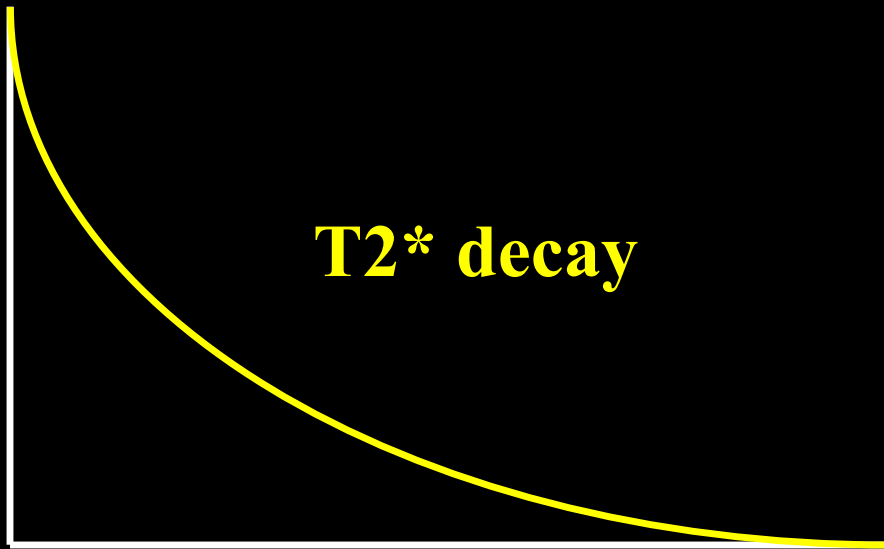


Functional

Echo-Planar Imaging



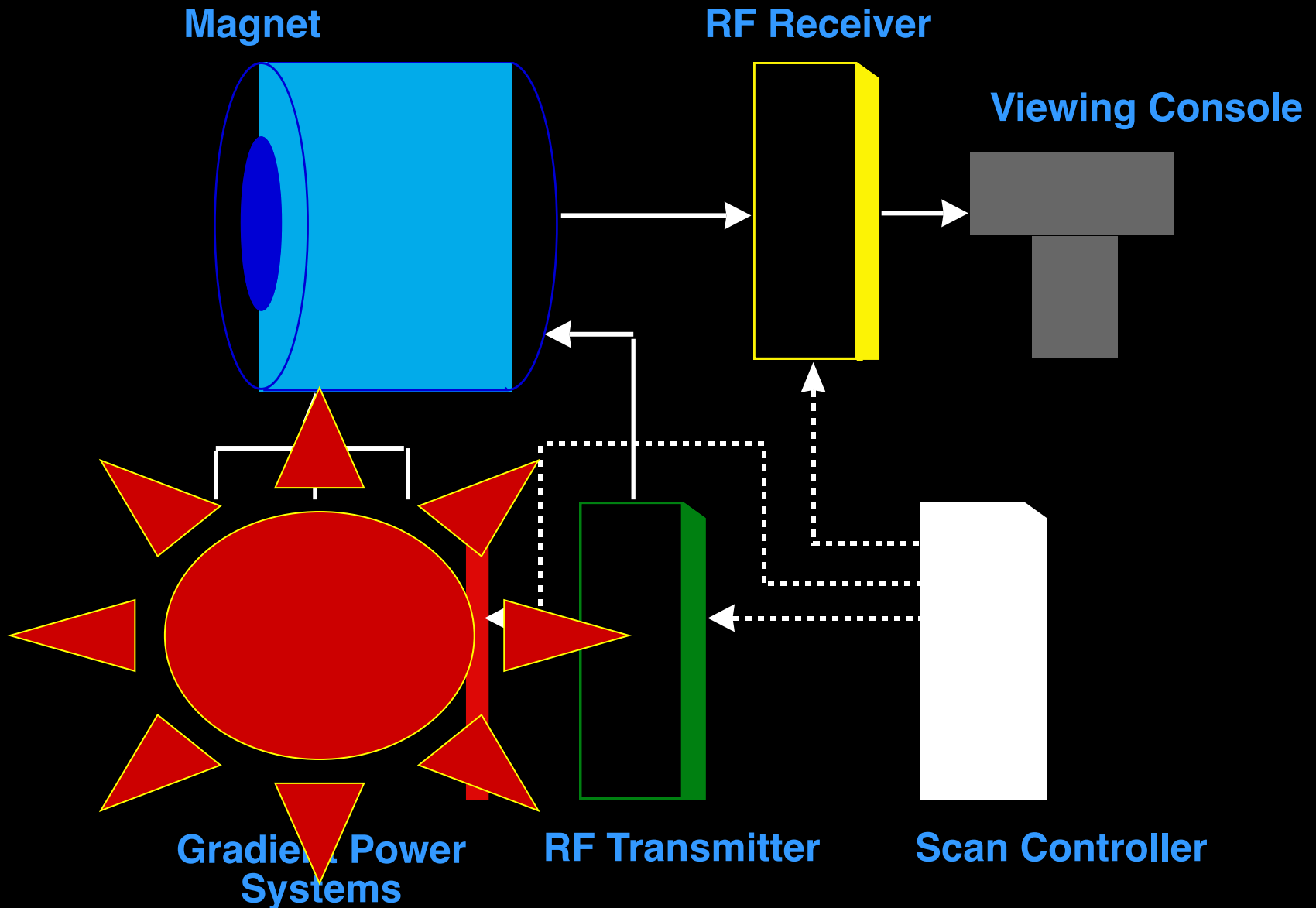
Single Shot Imaging



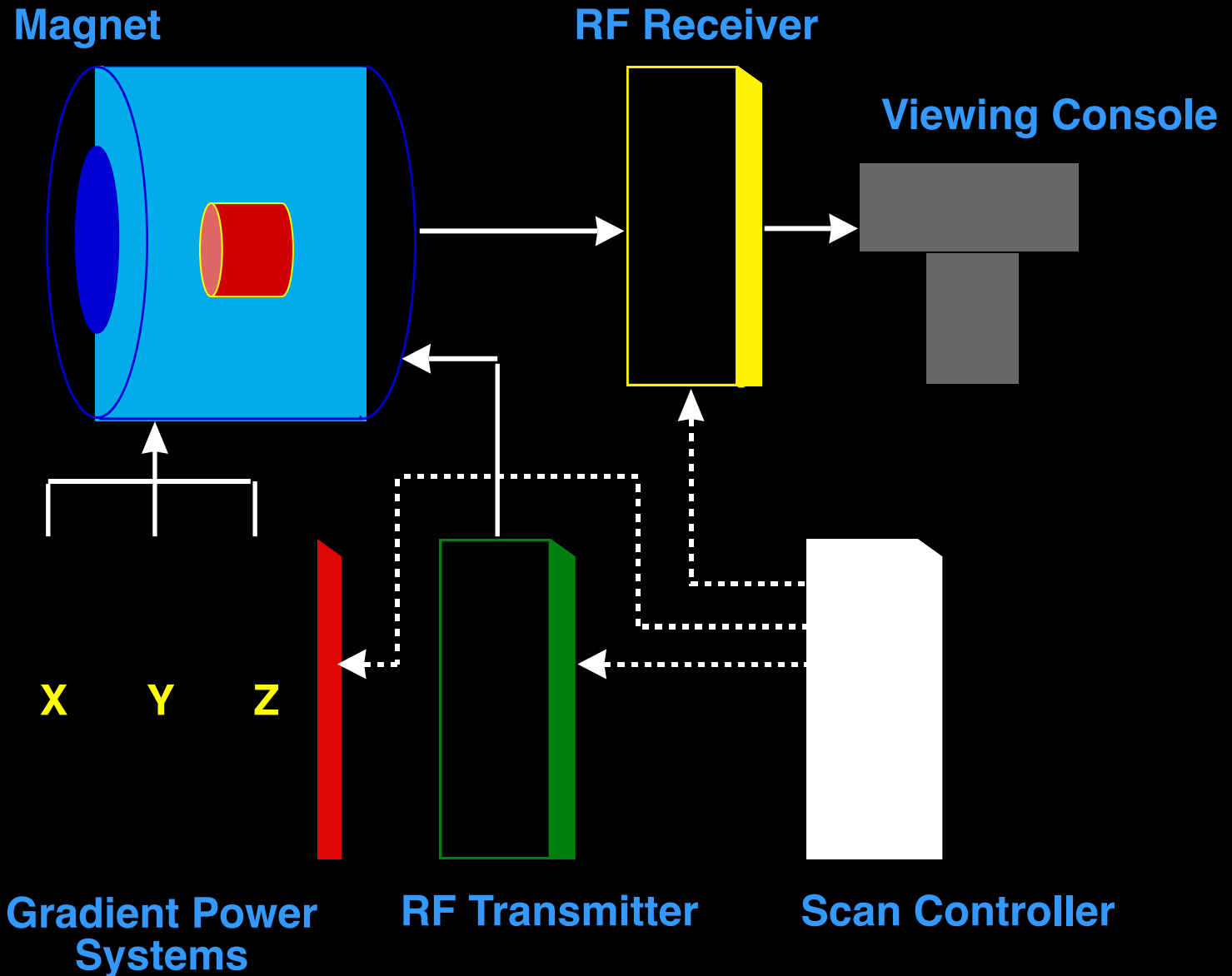
EPI Readout Window

≈ 20 to 40 ms

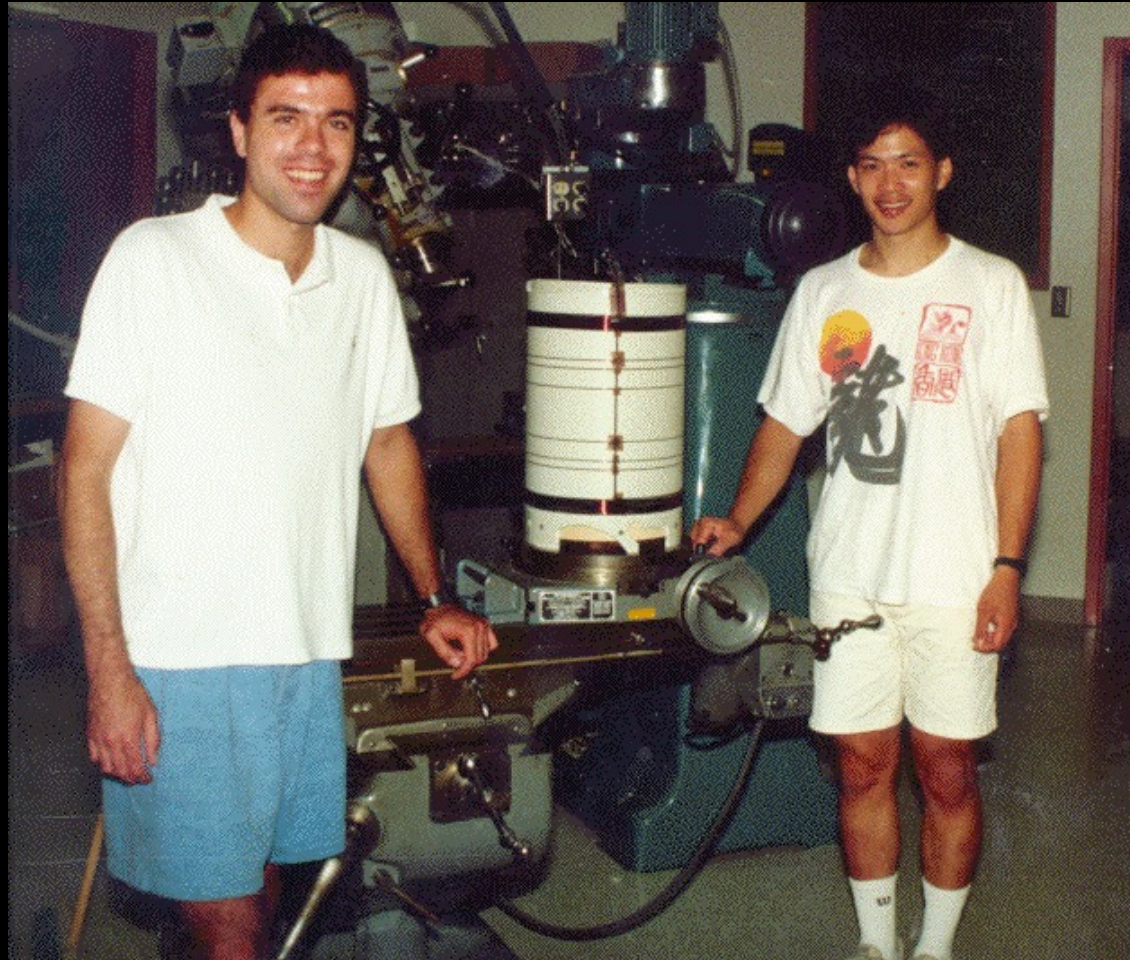
Imaging System Components



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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Local Human Head Gradient Coils
1.5T,3T, 4T
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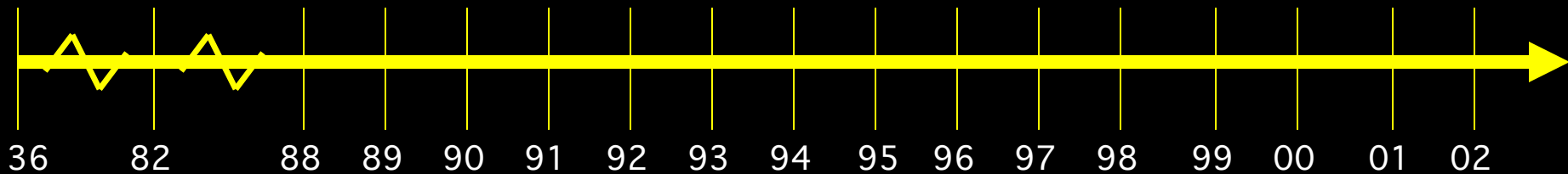
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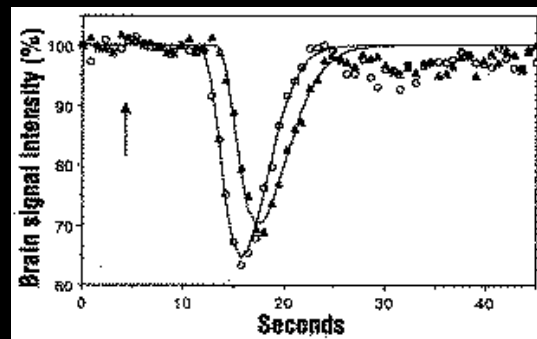
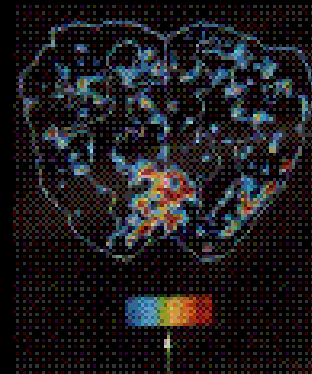
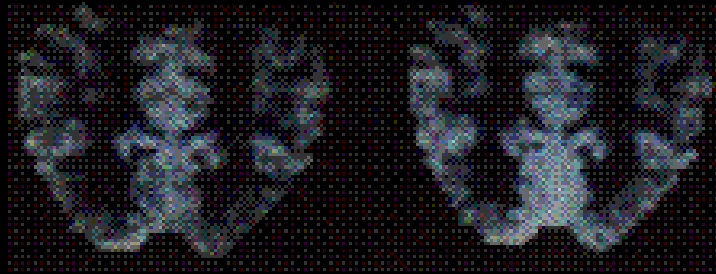


Blood Volume Imaging

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

Resting

Active

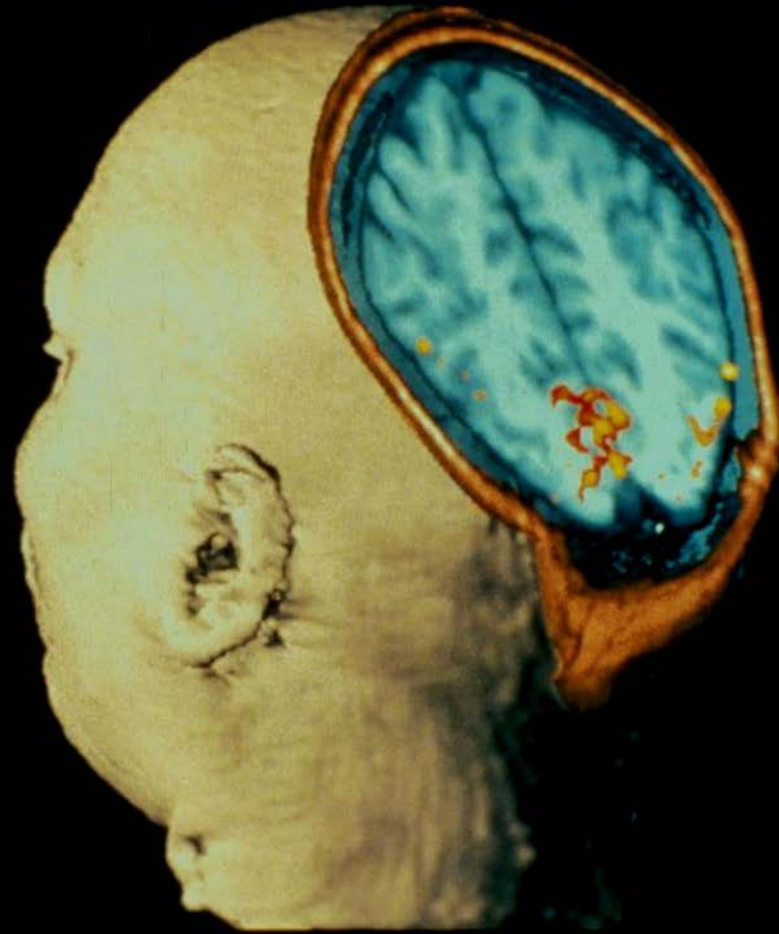


Blood Volume

**Photic
Stimulation**

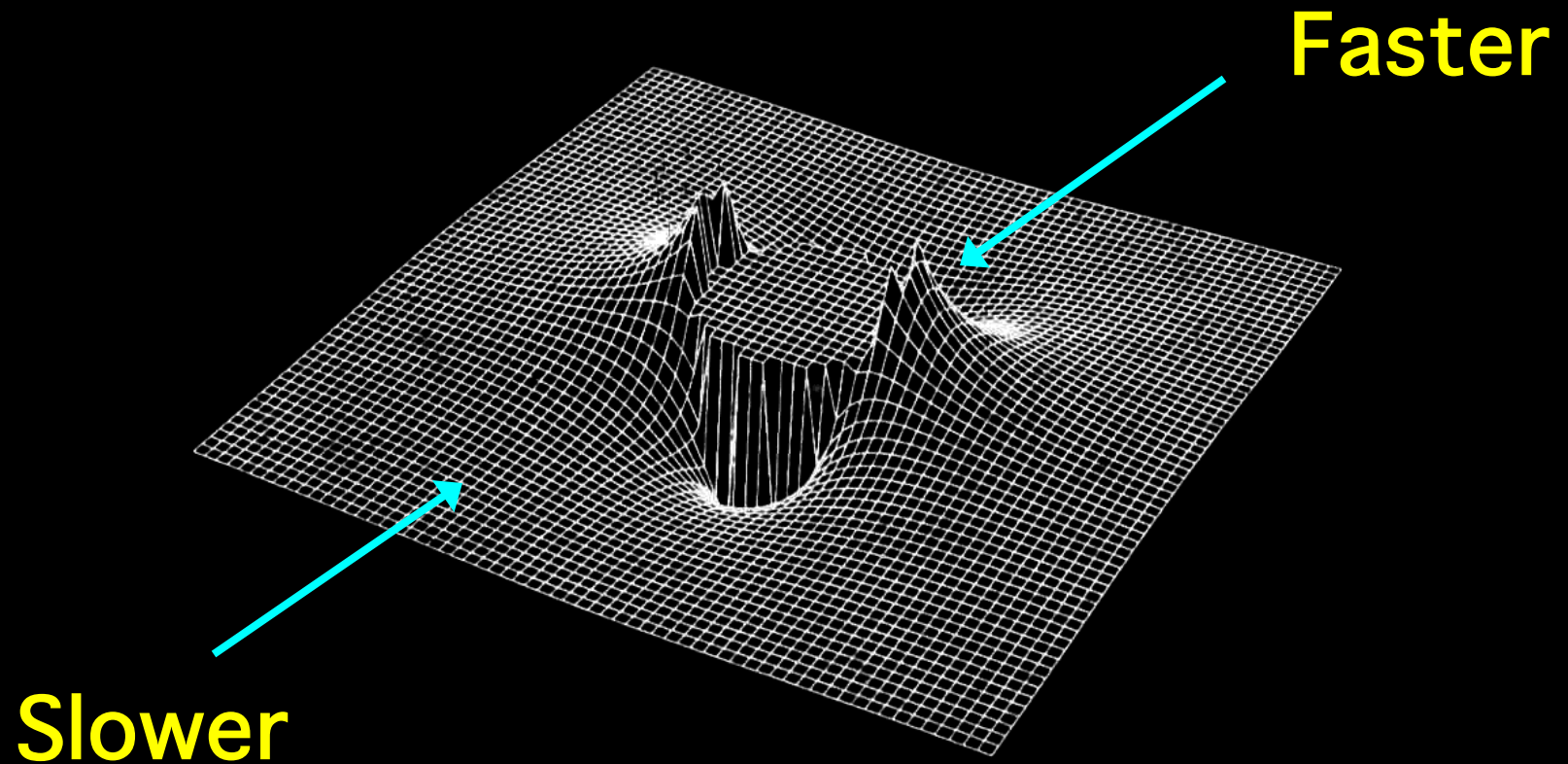
**MRI Image showing
activation of the
Visual Cortex**

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



Susceptibility Contrast

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



Alternating Left and Right Finger Tapping



~ 1992

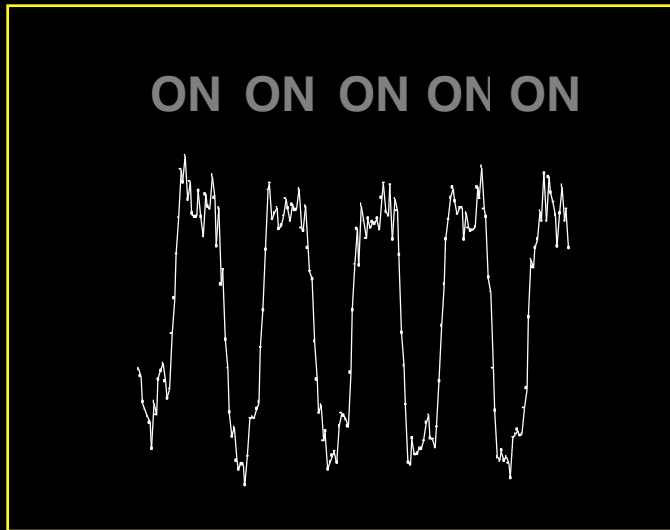
K. K. Kwong, et al, (1992) “Dynamic magnetic resonance imaging of human brain activity during primary sensory stimulation.” Proc. Natl. Acad. Sci. USA. 89, 5675-5679.

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.

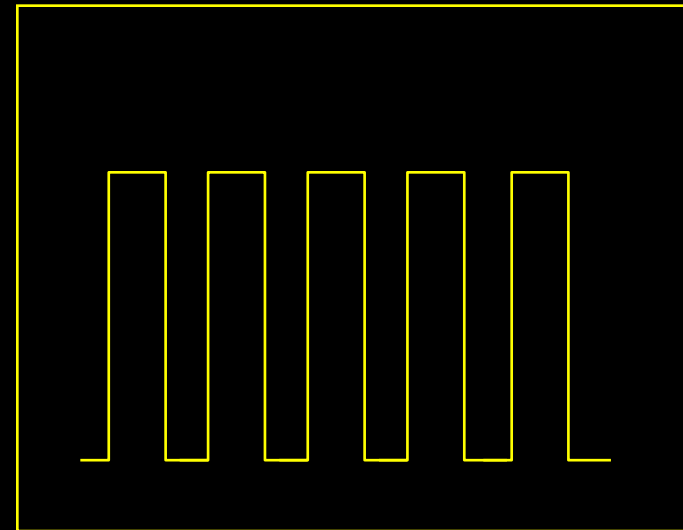
Blamire, A. M., et al. (1992). “Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging.” Proc. Natl. Acad. Sci. USA 89: 11069-11073.

Creating a Functional Image



Signal Time Course

X



Reference Function

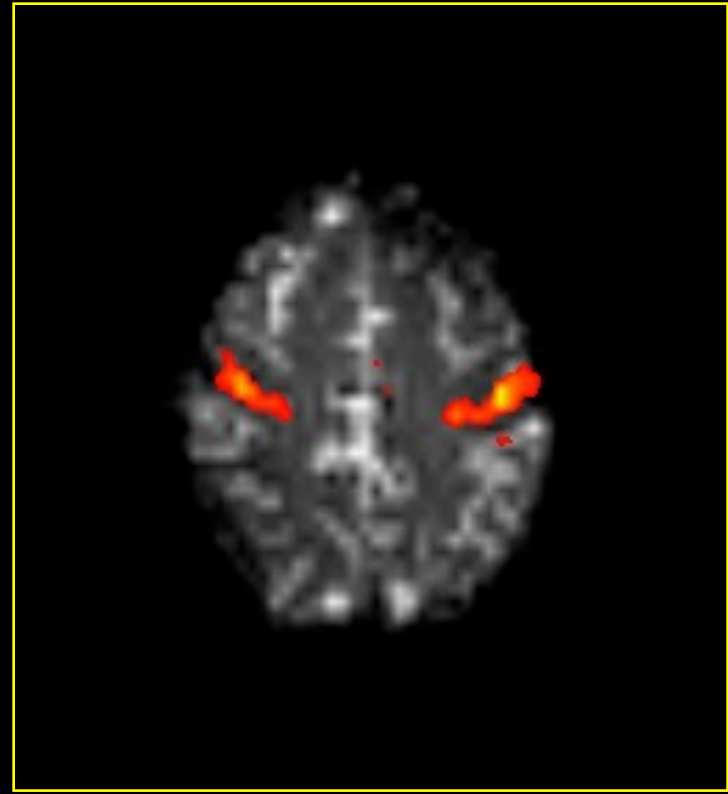
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P. A. Bandettini, A. Jesmanowicz, E. C. Wong, J. S. Hyde, Processing strategies for time-course data sets in functional MRI of the human brain. *Magn. Reson. Med.* **30**, 161-173 (1993).



Cross Correlation Image



Cross Correlation Image
Anatomical Image

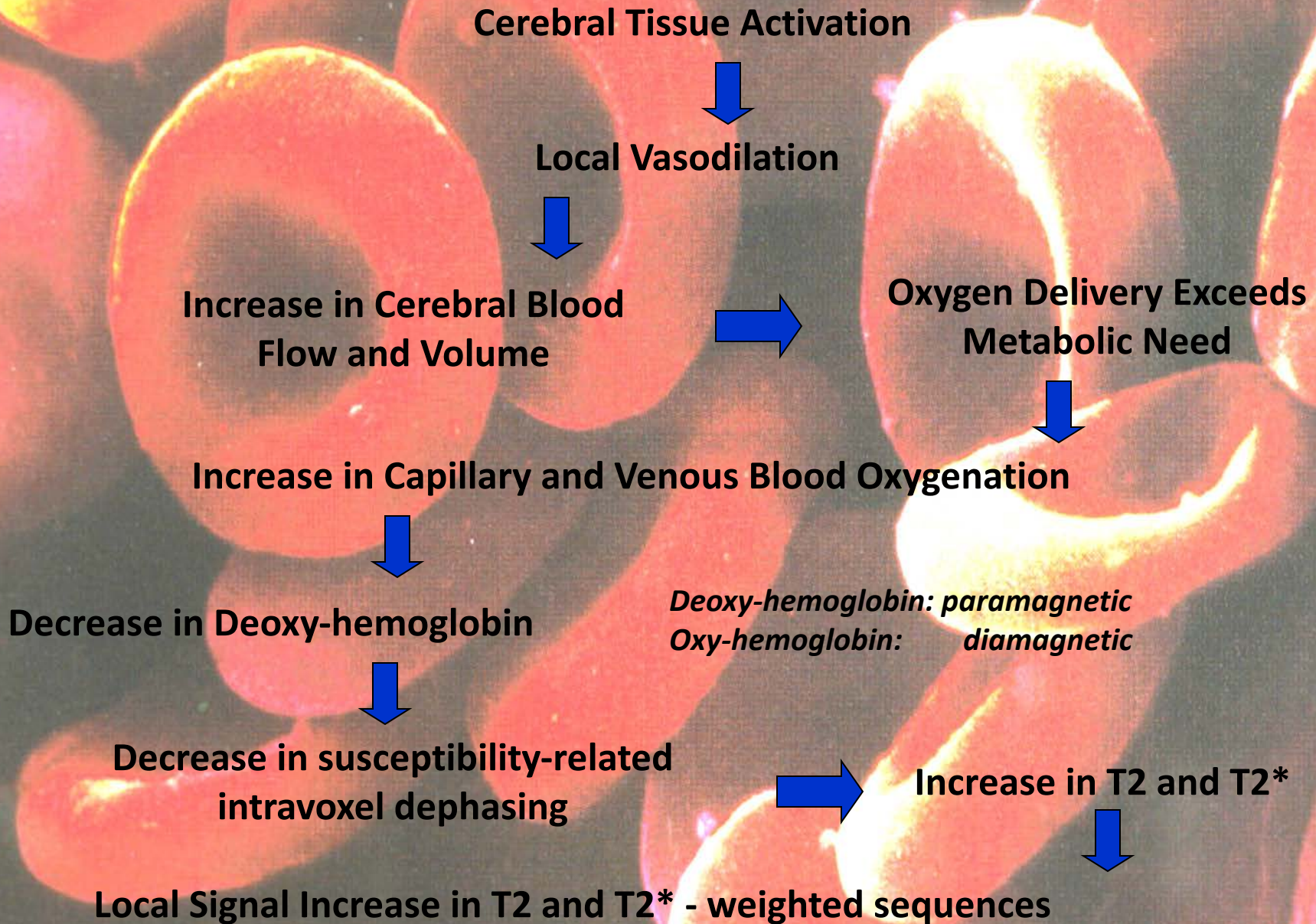
P. A. Bandettini, A. Jesmanowicz, E. C. Wong, J. S. Hyde, Processing strategies for time-course data sets in functional MRI of the human brain. *Magn. Reson. Med.* 30, 161-173 (1993).

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



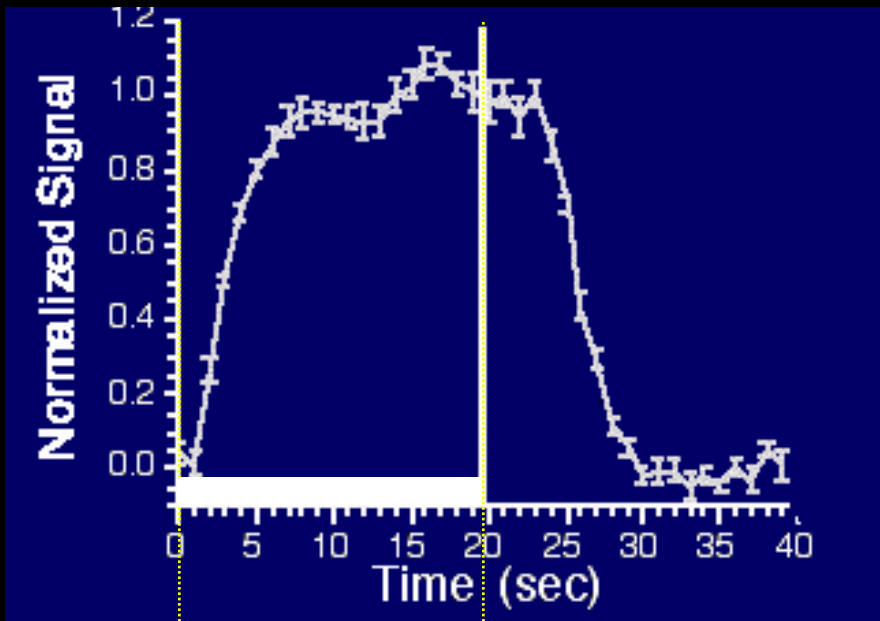
*Quality of results and importance of the findings depends on
type of question asked, experimental method, and analysis method...*

BOLD Contrast in the Detection of Neuronal Activity

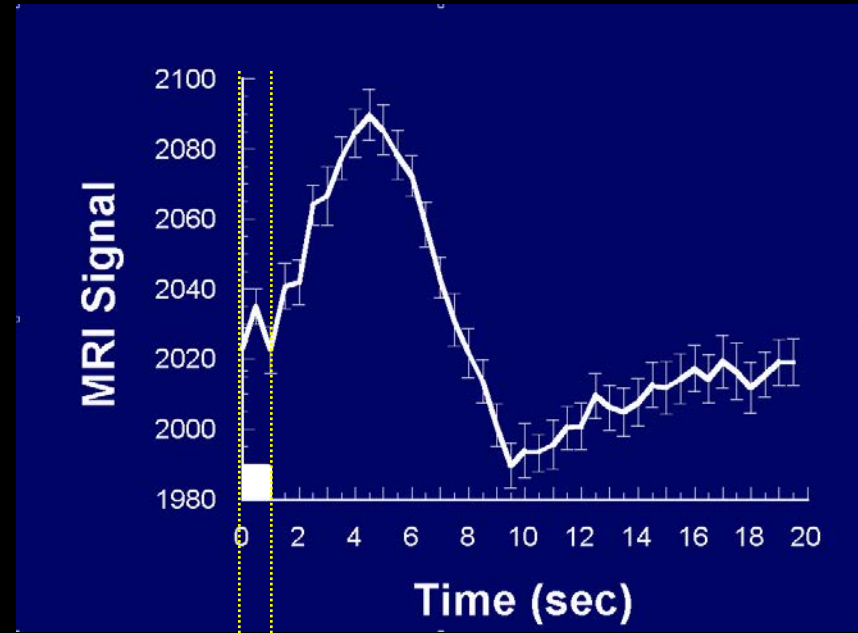


The BOLD Signal

Blood Oxygenation Level Dependent (BOLD) signal changes

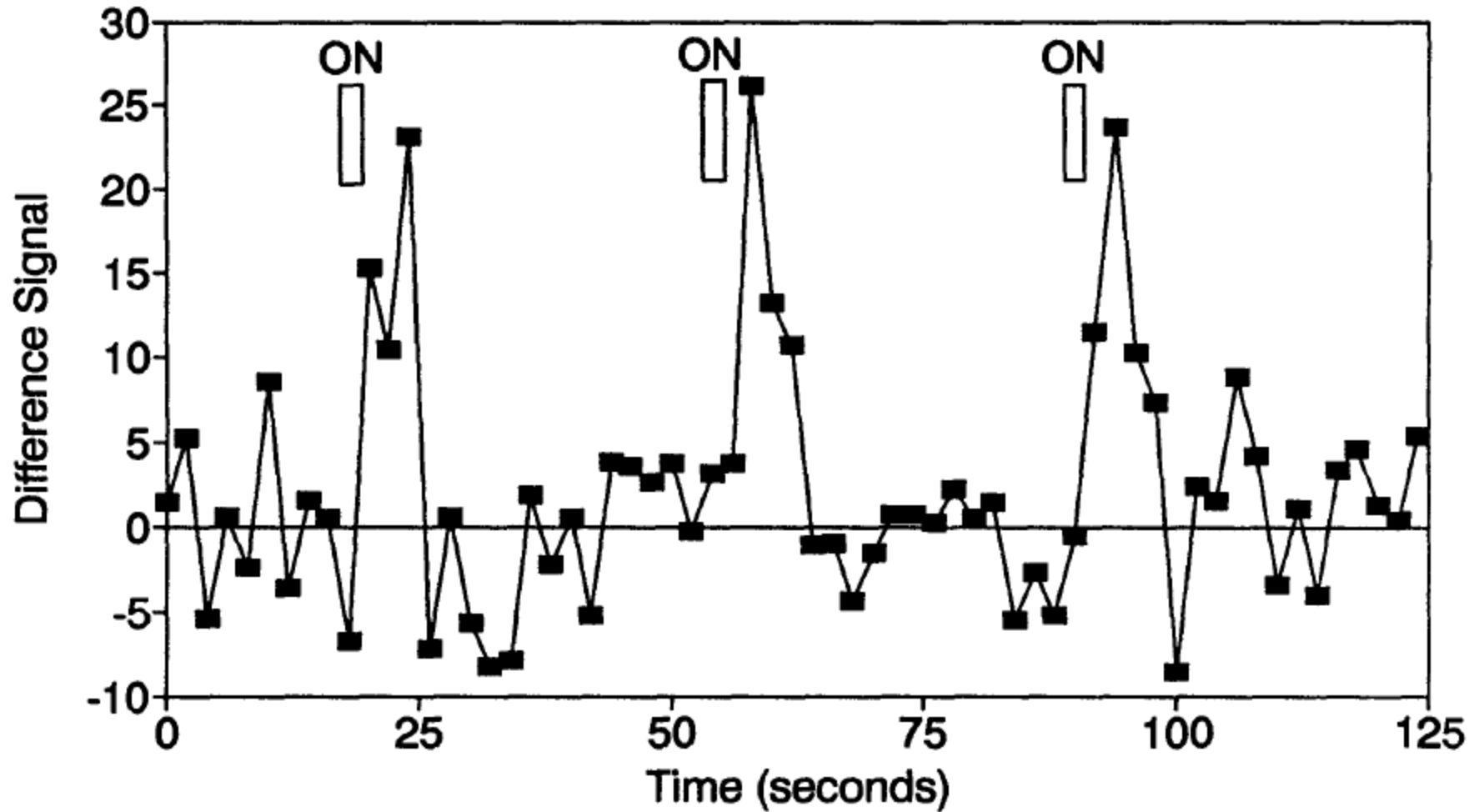


task



task

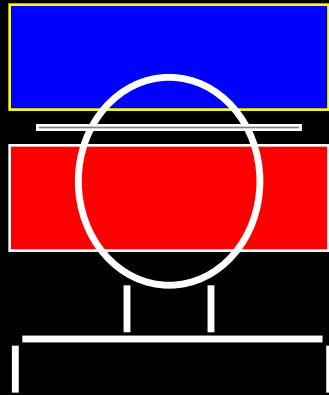
First Event-related fMRI Results



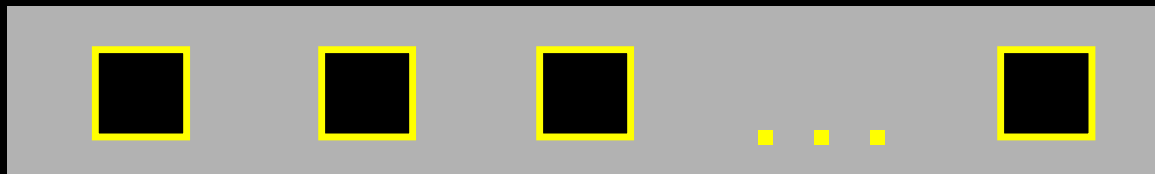
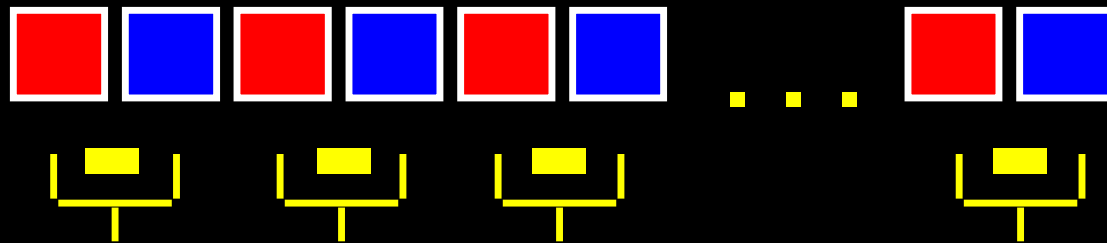
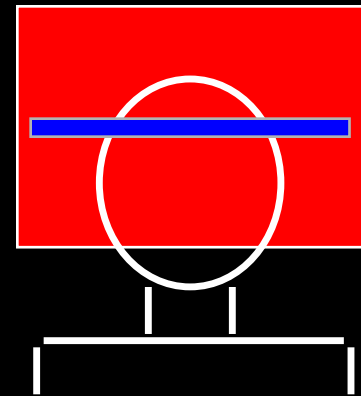
Blamire, A. M., et al. (1992). "Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging." *Proc. Natl. Acad. Sci. USA* 89: 11069-11073.

Blood Perfusion

EPISTAR



FAIR



**Perfusion
Time Series**

TI (ms)

FAIR

EPISTAR

200

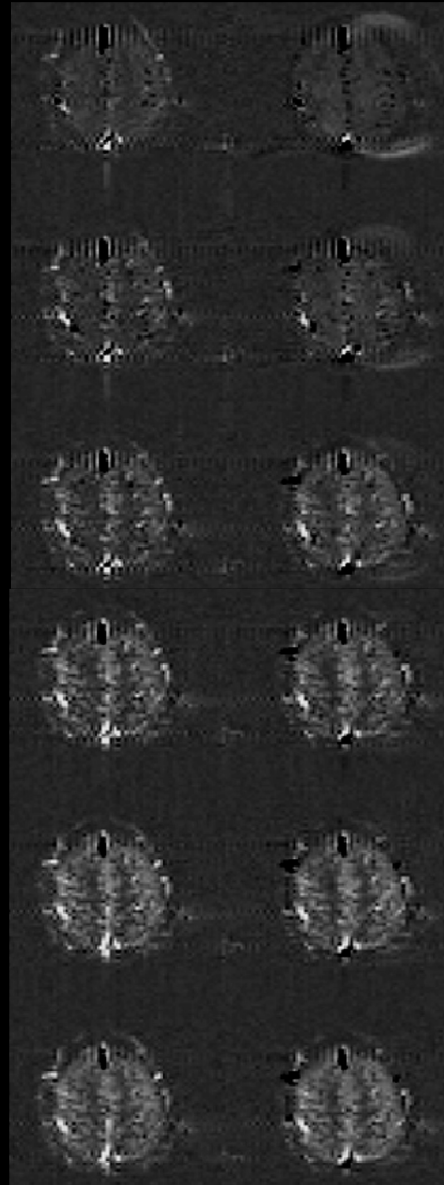
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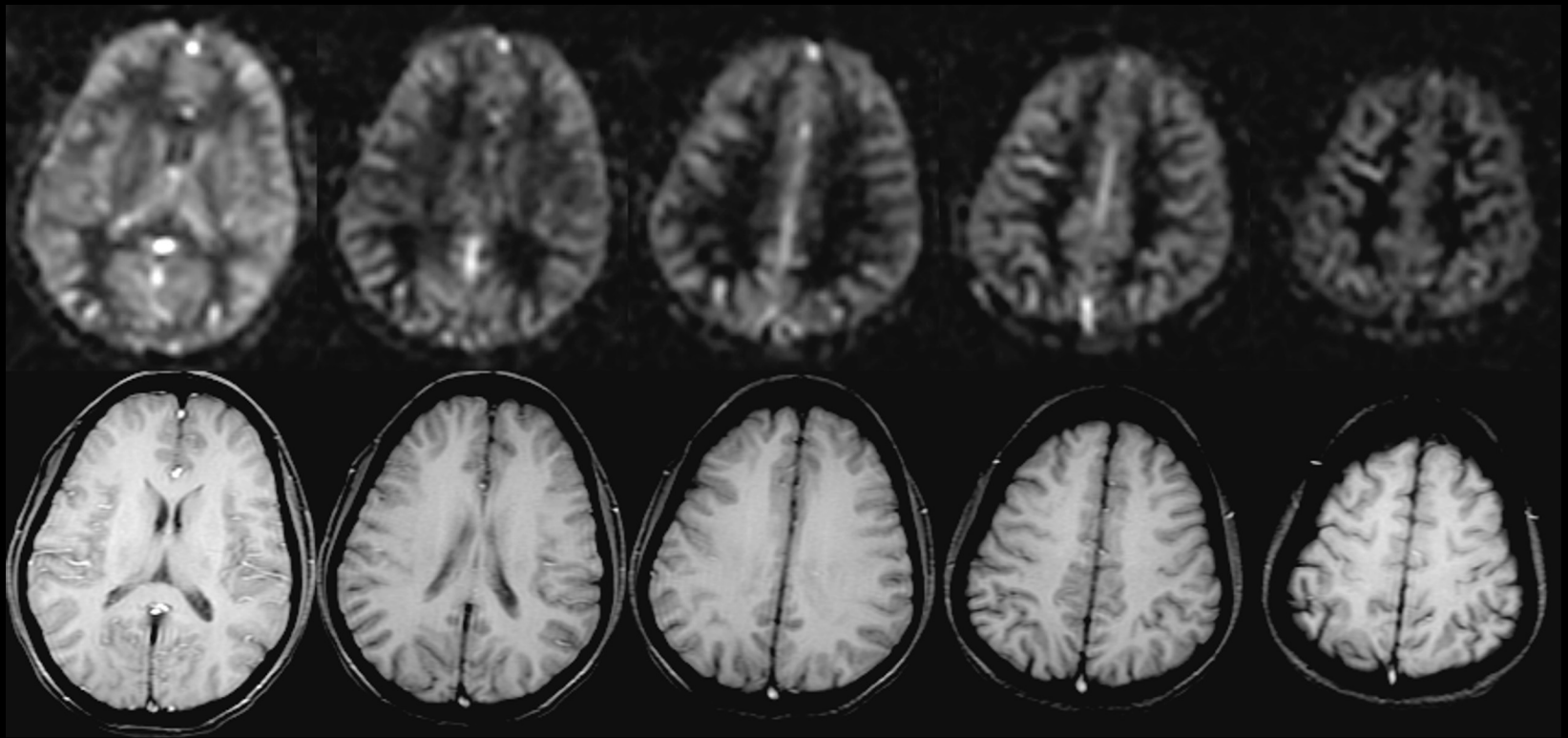
600

800

1000

1200





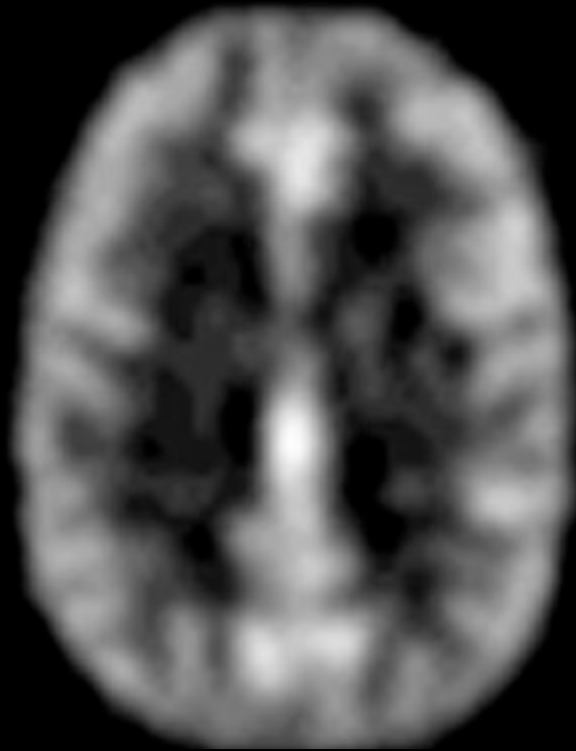
Williams, D. S., Detre, J. A., Leigh, J. S. & Koretsky, A. S. (1992) "Magnetic resonance imaging of perfusion using spin-inversion of arterial water." *Proc. Natl. Acad. Sci. USA* 89, 212-216.

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.

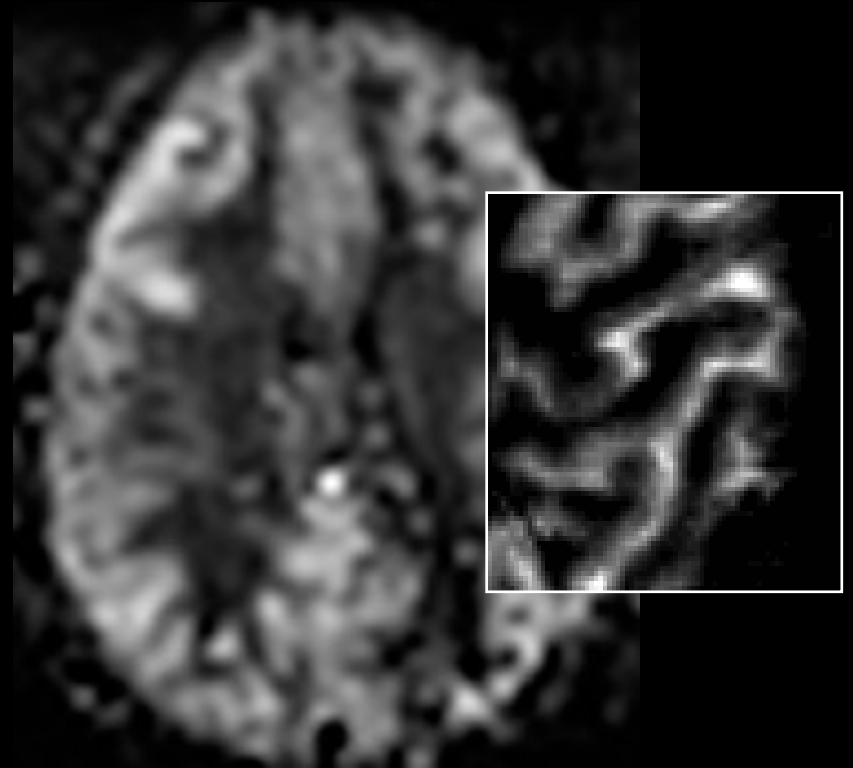
Kim, S.-G. (1995) "Quantification of relative cerebral blood flow change by flow-sensitive alternating inversion recovery (FAIR) technique: application to functional mapping." *Magn. Reson. Med.* 34, 293-301.

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: H₂¹⁵O



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

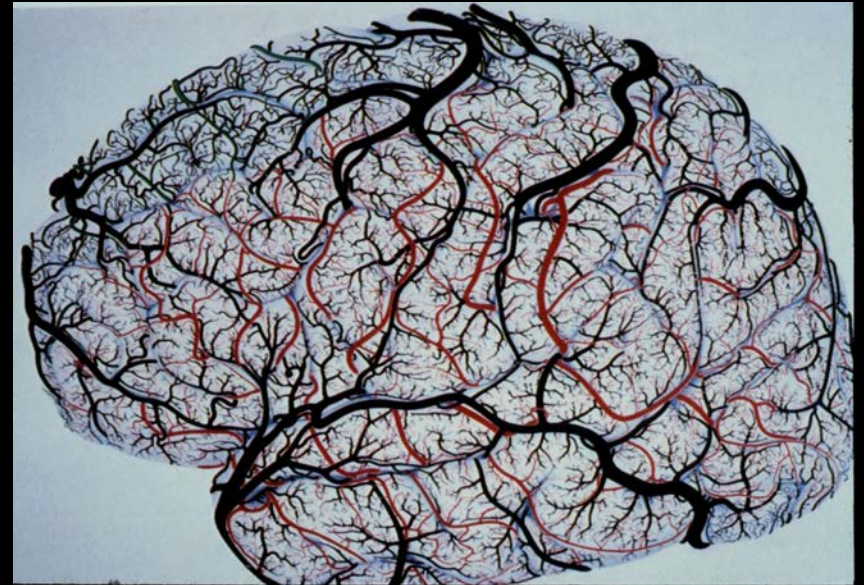
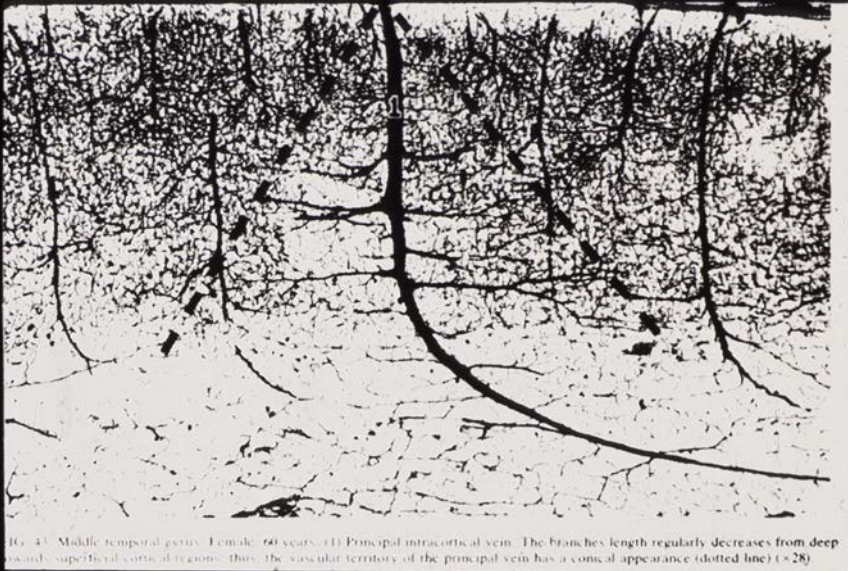
BOLD Contrast Interpretation

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.



Neuronal
Activation



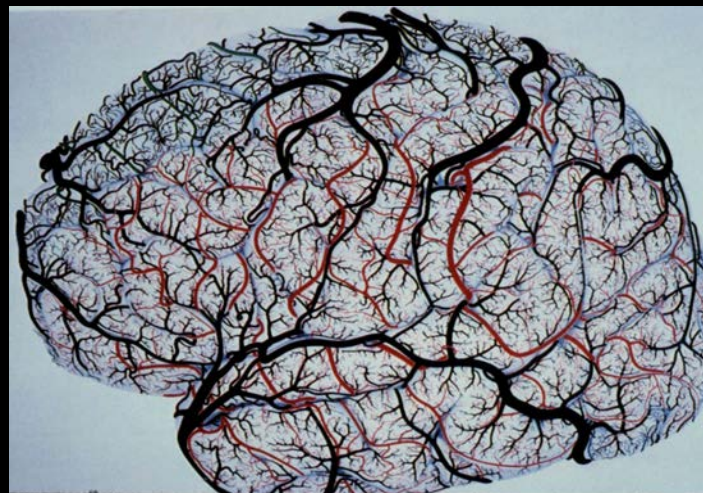
Measured
Signal

Hemodynamics

?

?

?



Noise

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V1, V2..mapping

Priming/Learning

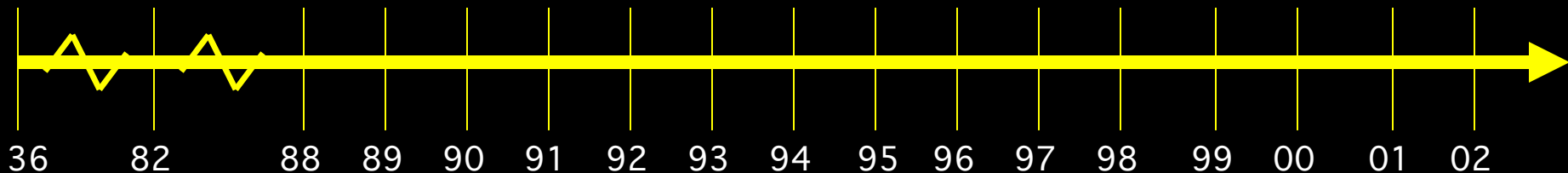
Clinical Populations

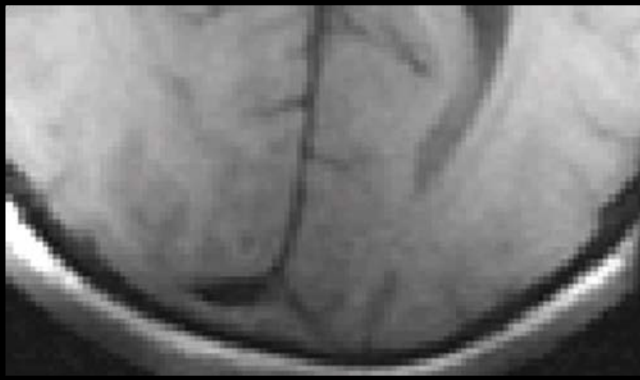
Δ Volume-V1

Plasticity

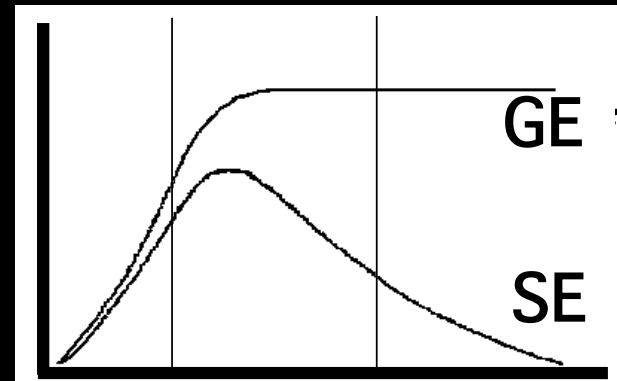
Face recognition

Performance prediction





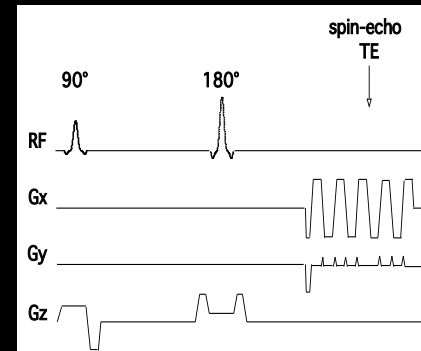
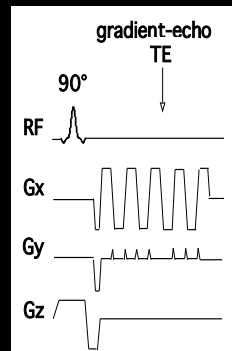
Contrast



2.5 to 3 μm 3 to 15 μm 15 to ∞ μm

compartment size

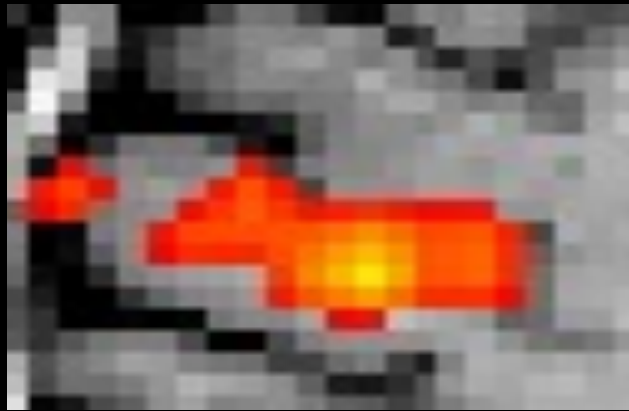
Gradient - Echo



Spin - Echo

T1 - weighted

Flow weighted



T2* weighted

BOLD weighted



T1 and T2* weighted

Flow and BOLD weighted



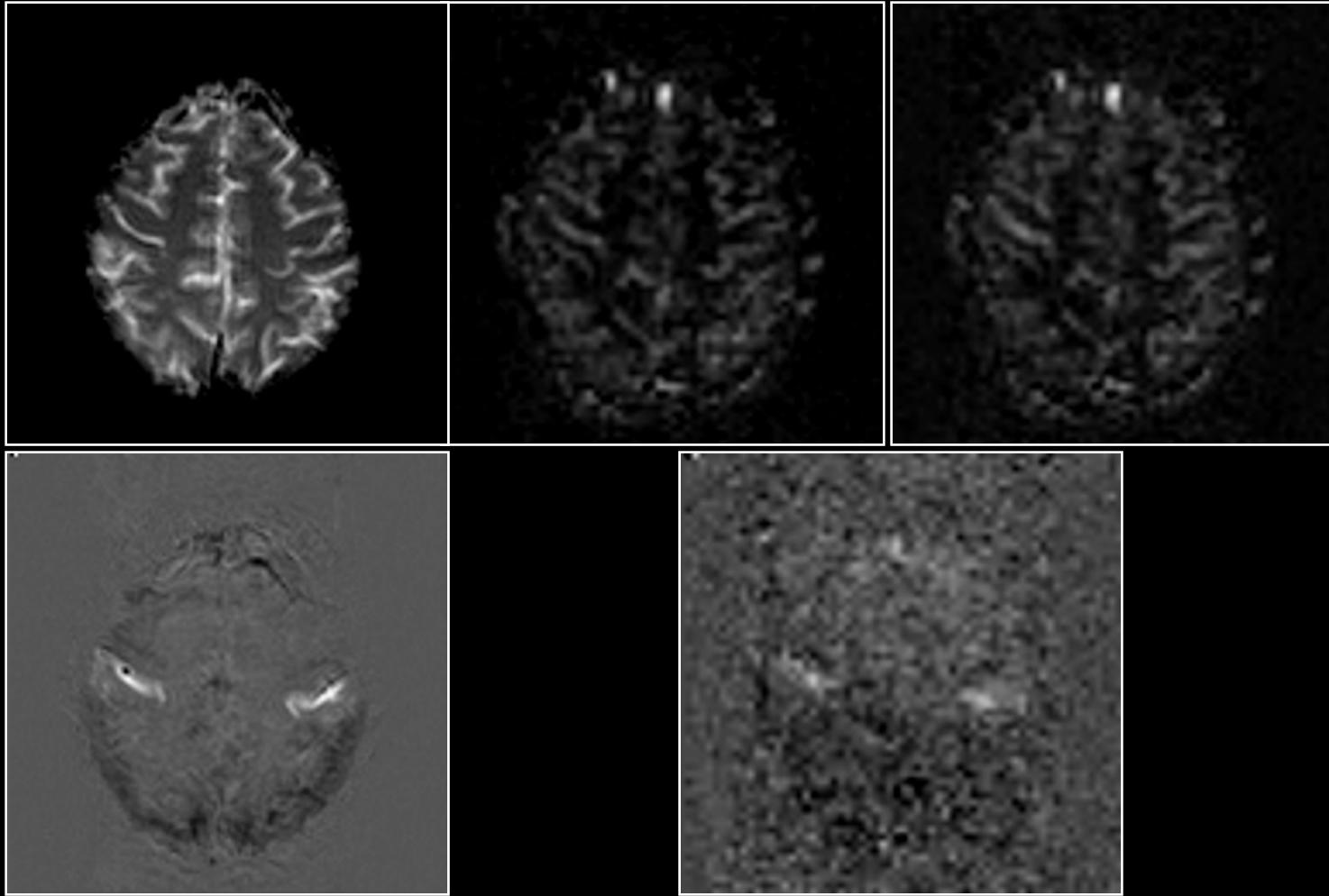
P. A. Bandettini, E. C. Wong, Echo - planar magnetic resonance imaging of human brain activation, in "Echo Planar Imaging: Theory, Technique, and Application" (F. Schmitt, M. Stehling, R. Turner, Eds.), p.493-530, Springer - Verlag, Berlin, 1997

Perfusion

BOLD

Rest

Activation

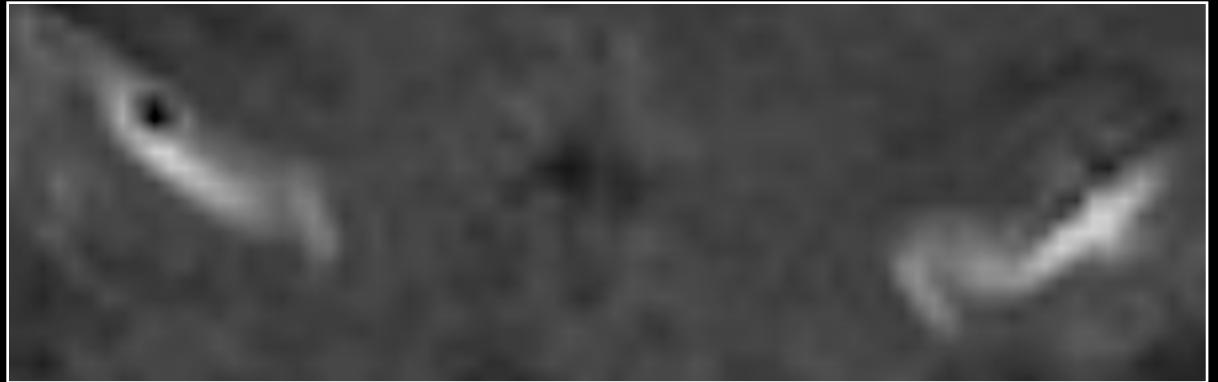


P. A. Bandettini, E. C. Wong, Magnetic resonance imaging of human brain function: principles, practicalities, and possibilities, *in* "Neurosurgery Clinics of North America: Functional Imaging" (M. Haglund, Ed.), p.345-371, W. B. Saunders Co., 1997.

Anatomy



BOLD



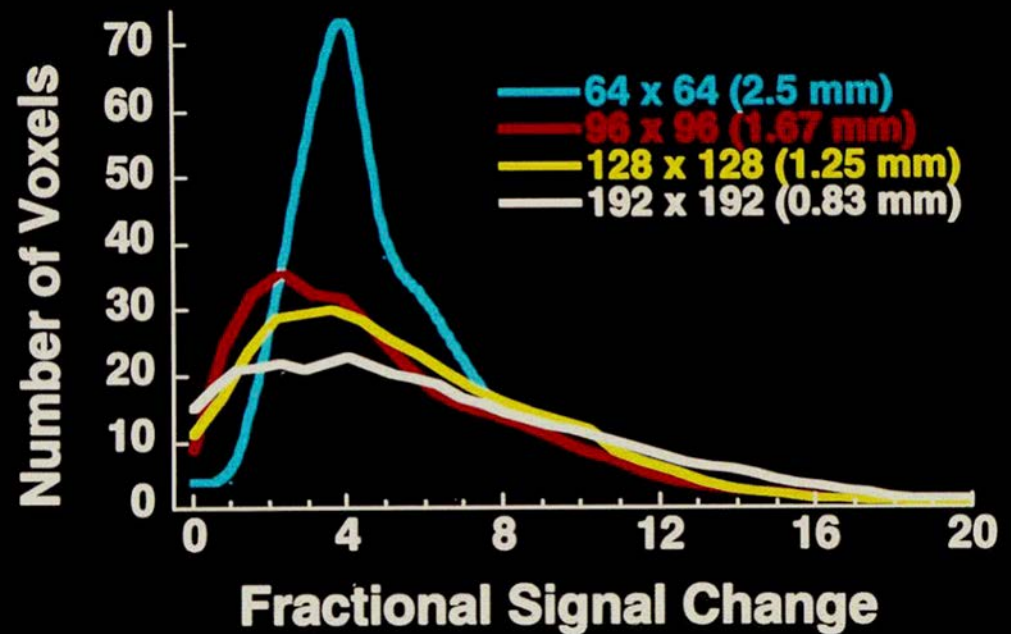
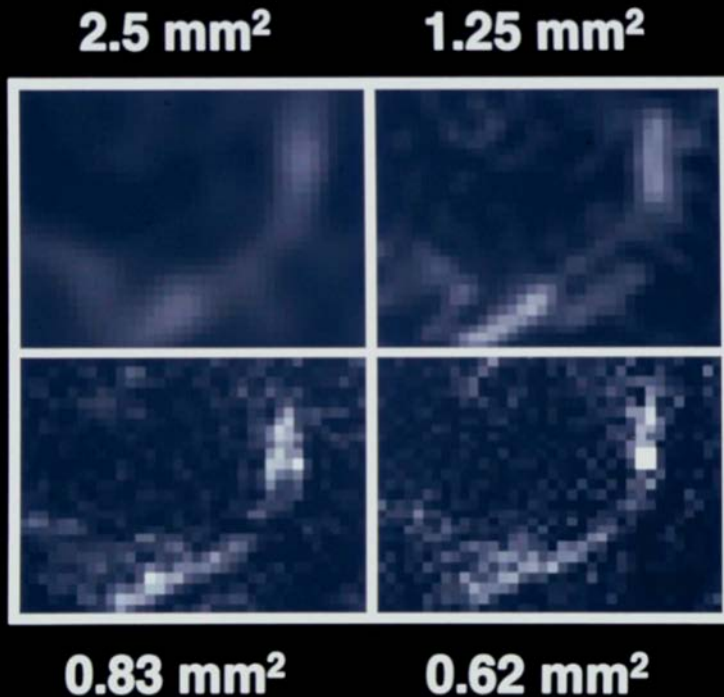
Perfusion



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)

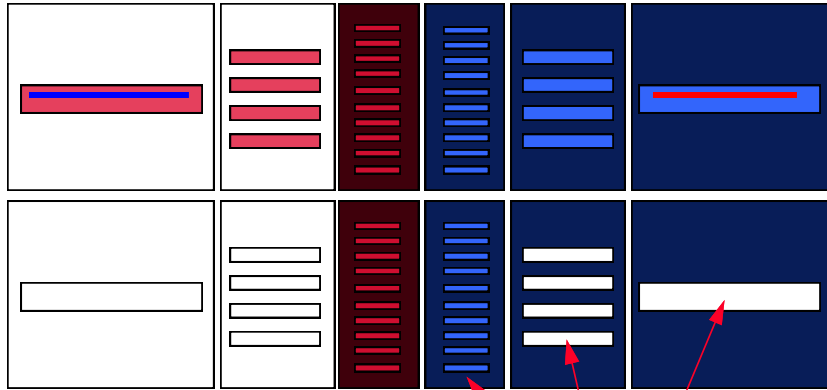
Perfusion

BOLD

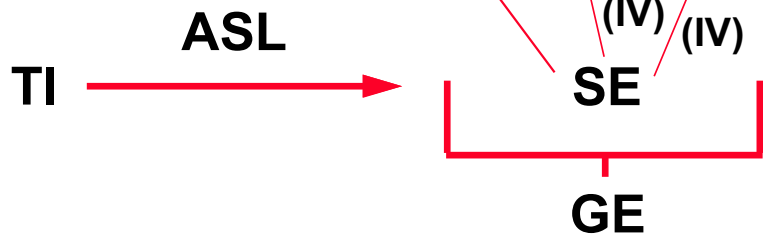
Venous inflow
(for ASL, w/ no VN)

No
Velocity
Nulling

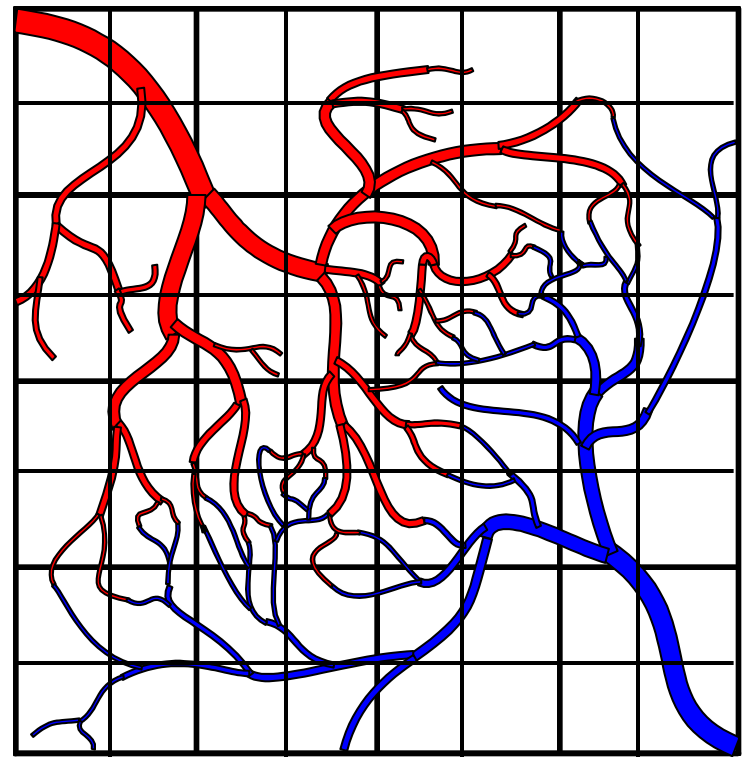
Velocity
Nulling



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. Diff. tensor Mg⁺ 7T

Local Human Head Gradient Coils Nav. pulses Real time fMRI Venography SENSE

ASL Spiral EPI Quant. ASL Dynamic IV volume Z-shim Baseline Susceptibility

BOLD Multi-shot fMRI Simultaneous ASL and BOLD Current Imaging?

Methodology

Baseline Volume

IVIM

Correlation Analysis Motion Correction CO₂ Calibration

Parametric Design Multi-Modal Mapping

Surface Mapping Free-behavior Designs

Phase Mapping Mental Chronometry

Linear Regression Deconvolution

Event-related

Interpretation

Blood T2

Hemoglobin

BOLD models PET correlation

B₀ dep. IV vs EV ASL vs. BOLD

TE dep Resolution Dep. Pre-undershoot PSF of BOLD

Post-undershoot Extended Stim.

SE vs. GE CO₂ effect Linearity Metab. Correlation

NIRS Correlation Fluctuations Optical Im. Correlation

Veins Inflow Balloon Model Electrophys. correlation

Applications

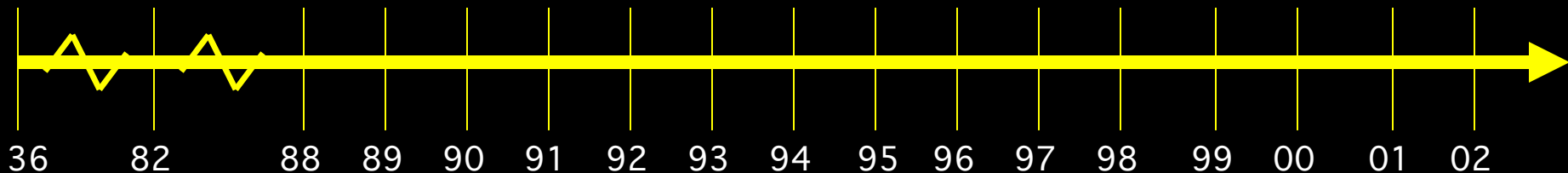
Complex motor Language Imagery Memory Emotion

Motor learning Children Tumor vasc. Drug effects

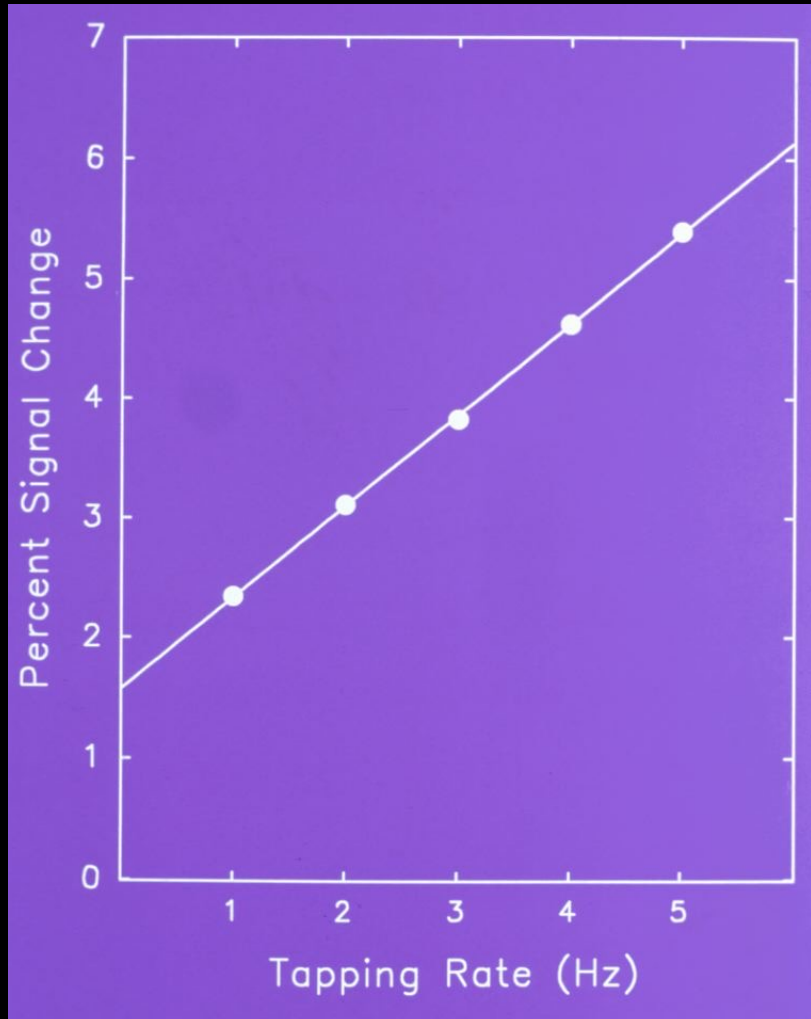
BOLD -V1, M1, A1 Presurgical Attention Ocular Dominance

Volume - Stroke V1, V2..mapping Priming/Learning Clinical Populations

Δ Volume-V1 Plasticity Face recognition Performance prediction

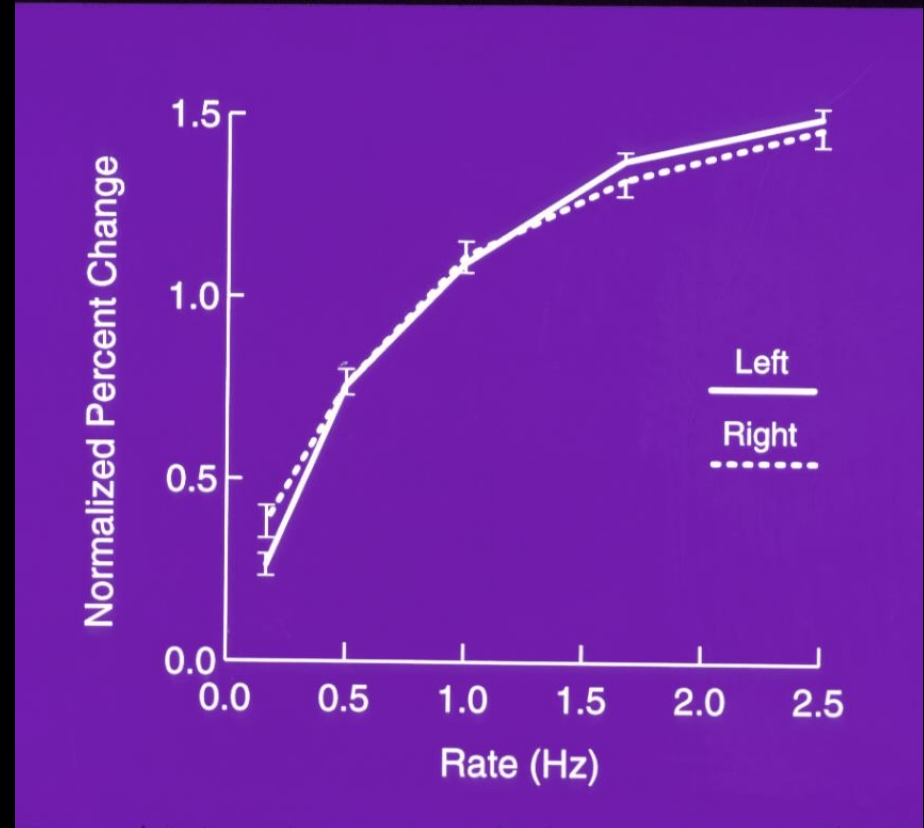


Motor Cortex



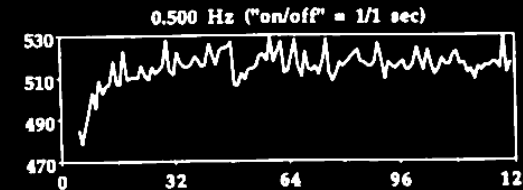
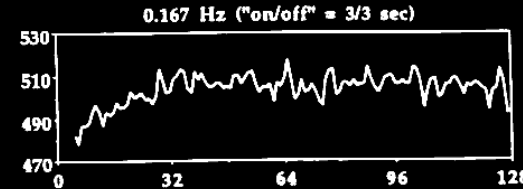
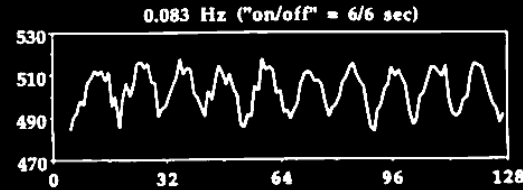
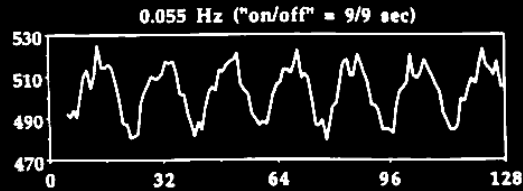
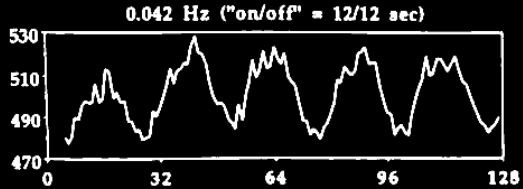
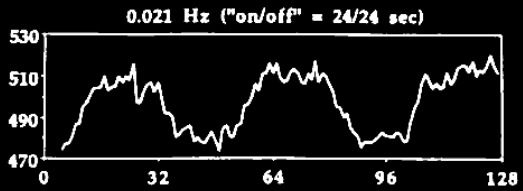
S. M. Rao et al, (1996) "Relationship between finger movement rate and functional magnetic resonance signal change in human primary motor cortex." *J. Cereb. Blood Flow and Met.* 16, 1250-1254.

Auditory Cortex

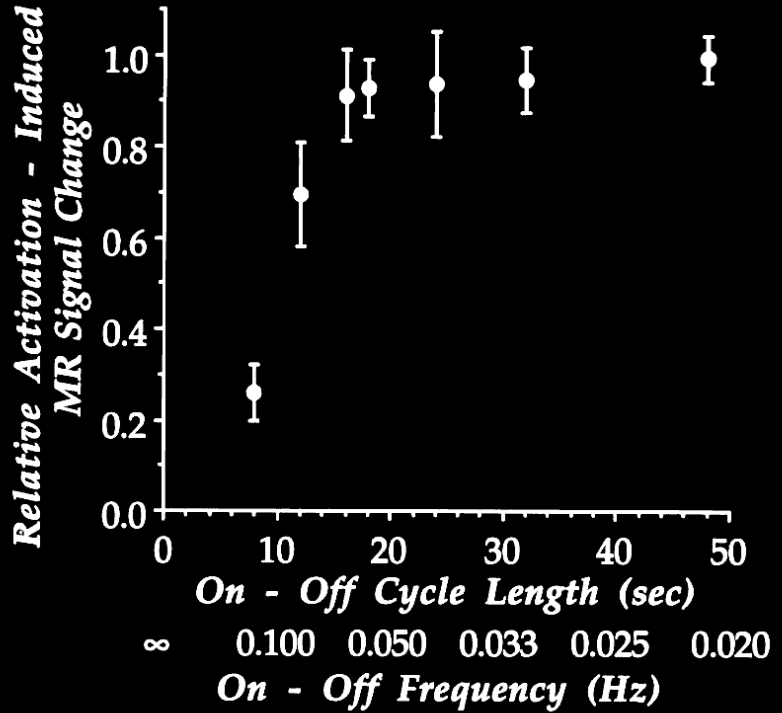


J. R. Binder, et al, (1994). "Effects of stimulus rate on signal response during functional magnetic resonance imaging of auditory cortex." *Cogn. Brain Res.* 2, 31-38

MRI Signal

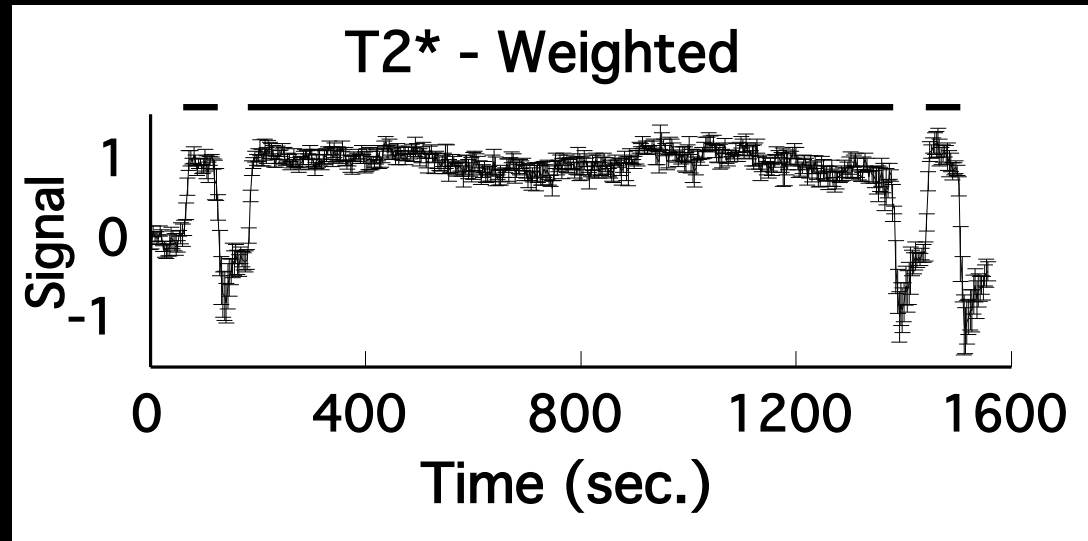


Time (seconds)

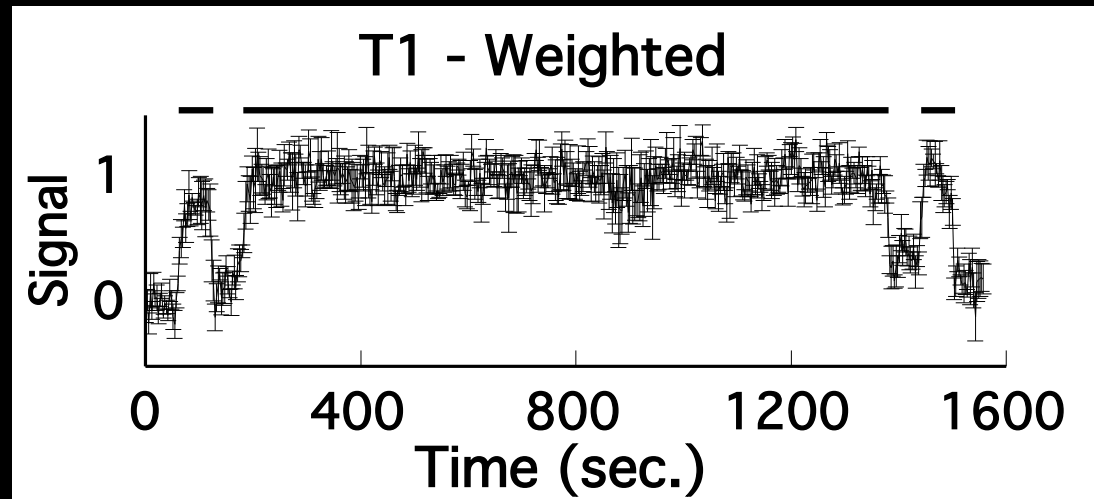


P. A. Bandettini, Functional MRI temporal resolution in "Functional MRI" (C. Moonen, and P. Bandettini., Eds.), p. 205-220, Springer - Verlag, 1999.

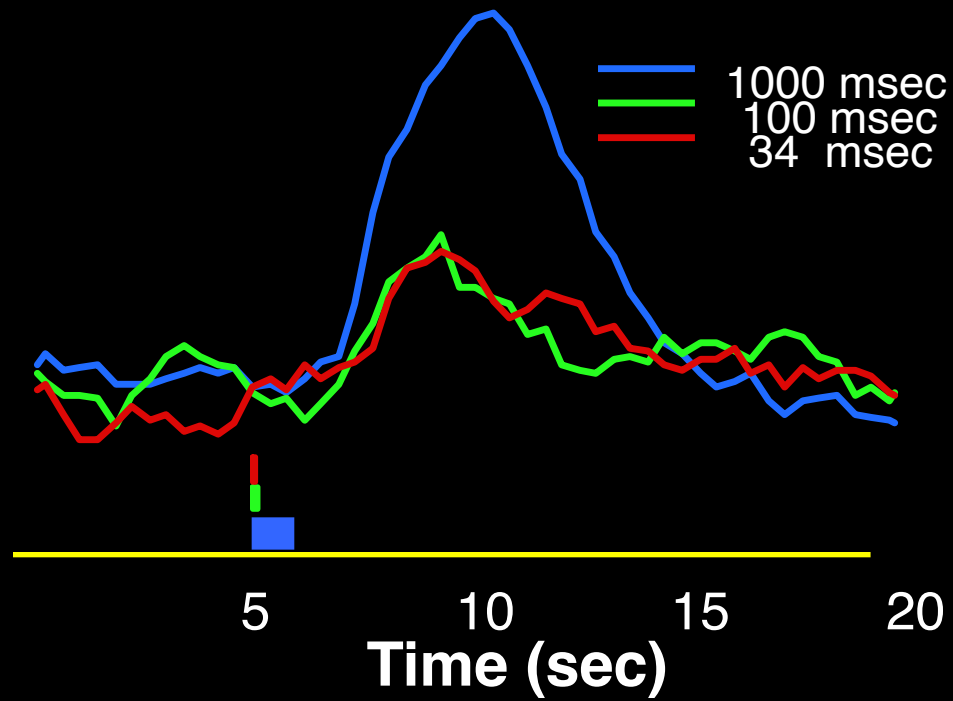
BOLD



Flow

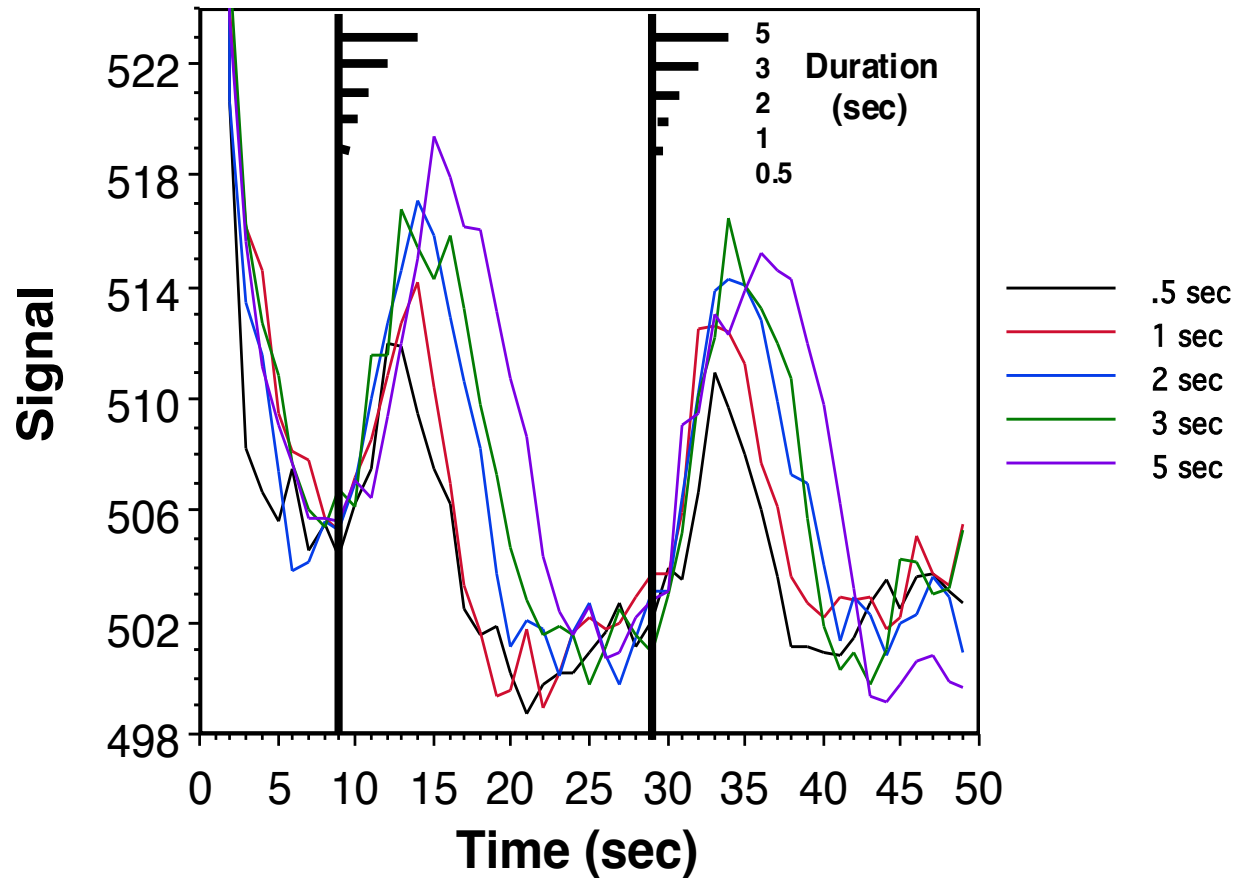


P. A. Bandettini, K. K. Kwong, T. L. Davis, R. B. H. Tootell, E. C. Wong, P. T. Fox, J. W. Belliveau, R. M. Weisskoff, B. R. Rosen, (1997). "Characterization of cerebral blood oxygenation and flow changes during prolonged brain activation." *Human Brain Mapping* 5, 93-109.



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], 3rd Proc. Soc. Magn. Reson., Nice, p. 450. (1995).

Motor Cortex

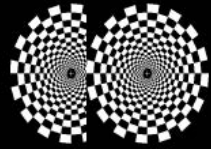


Bandettini, et al., The functional dynamics of blood oxygenation level contrast in the motor cortex, 12'th Proc. Soc. Magn. Reson. Med., New York, p. 1382. (1993).



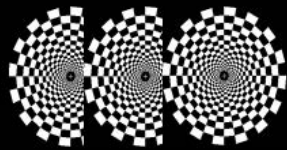
0 sec

20 sec



0 sec 2 sec

20 sec



0 sec 2 sec 4 sec

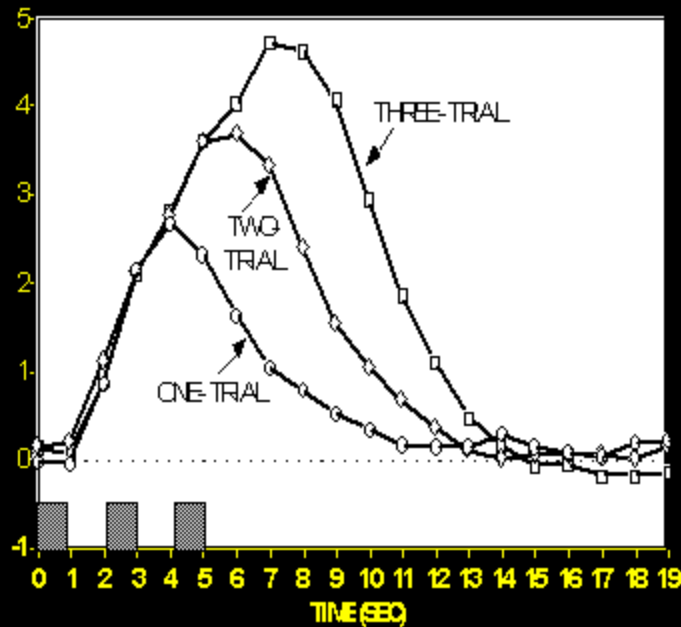
20 sec

♦ Human Brain Mapping 5:329-340(1997) ♦

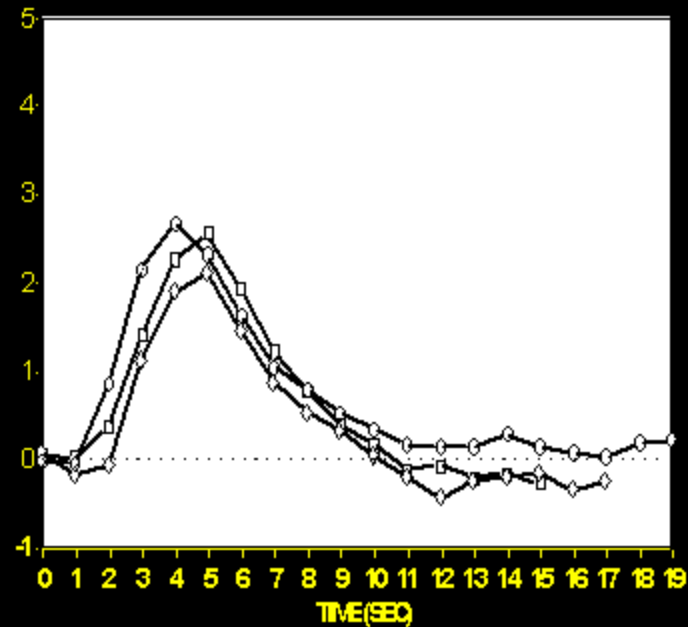
Selective Averaging of Rapidly Presented Individual Trials Using fMRI

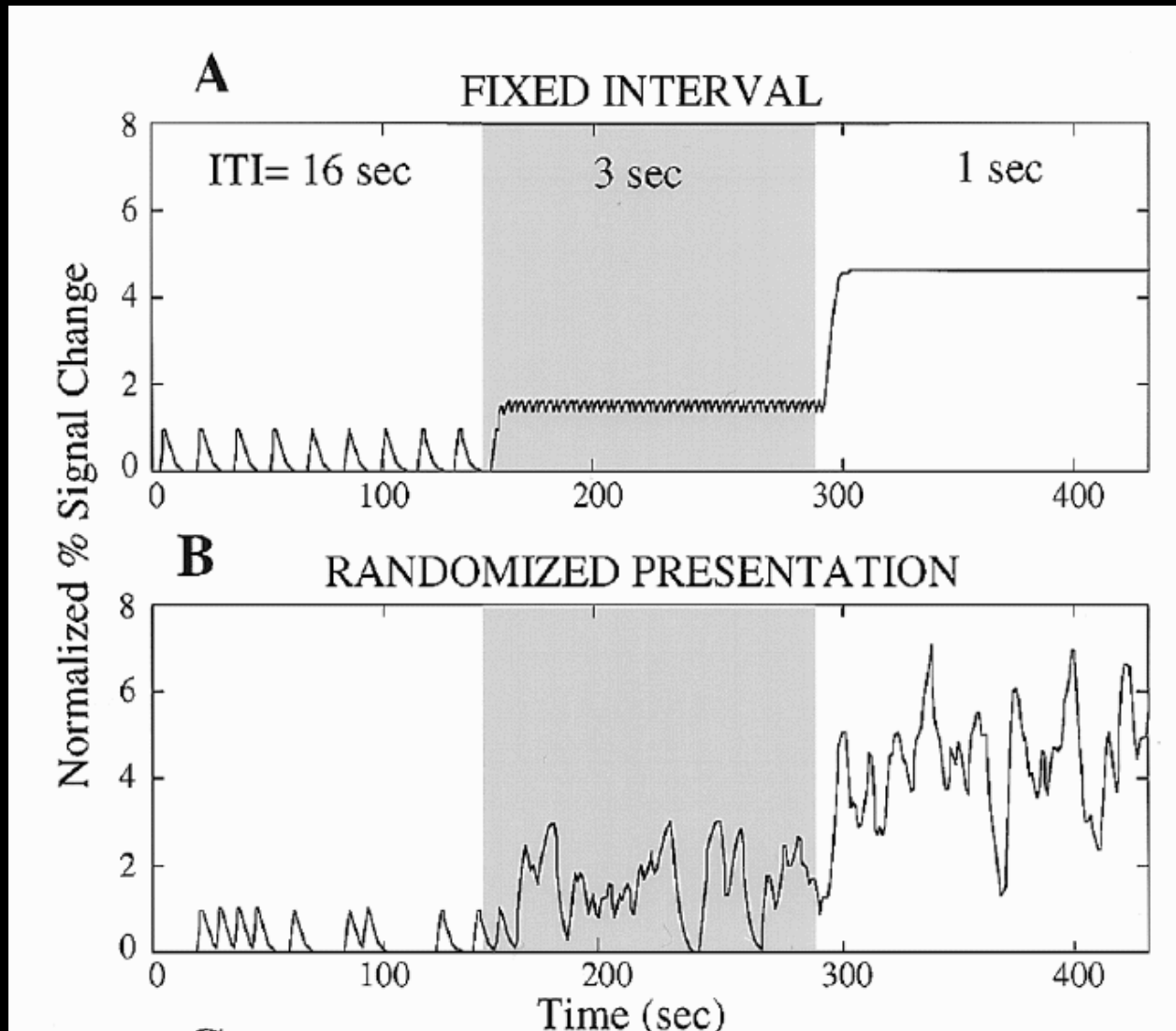
Anders M. Dale* and Randy L. Buckner

RAW DATA



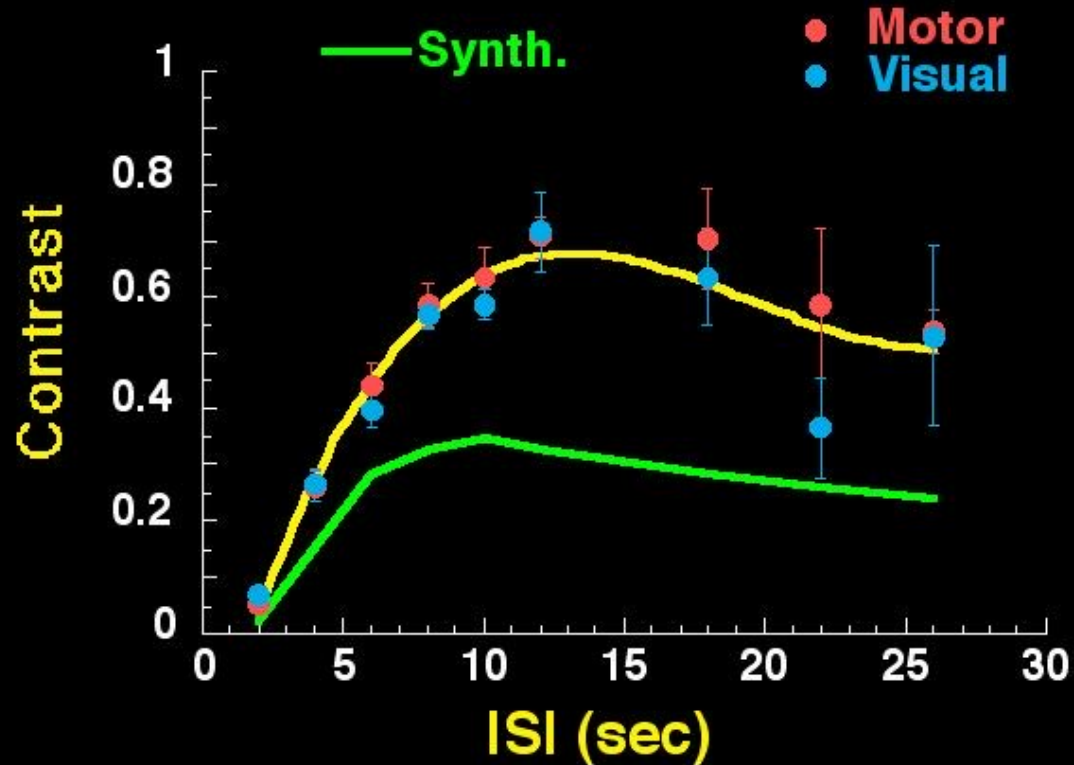
ESTIMATED RESPONSES





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

Functional Contrast



(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

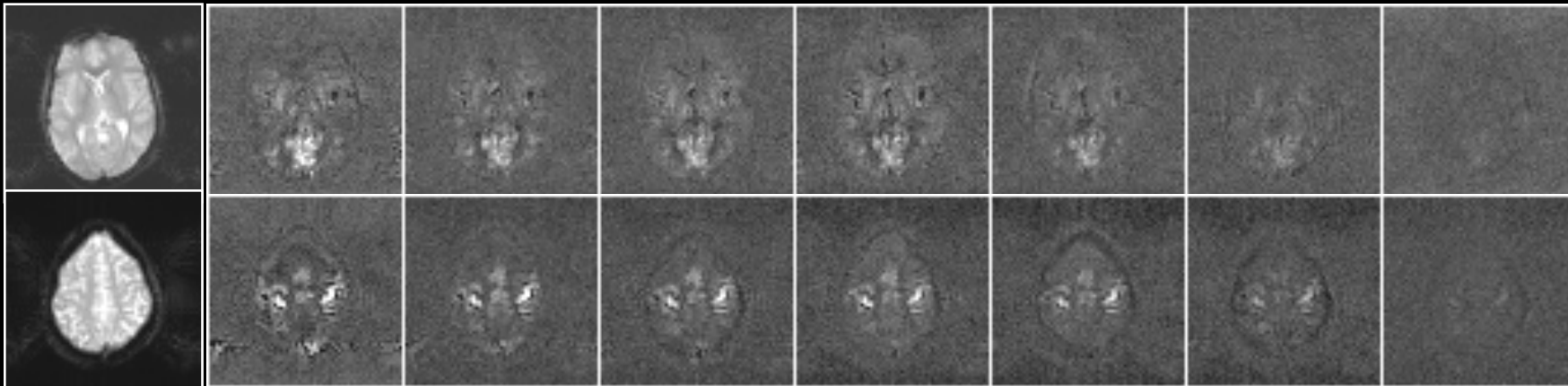
8, 2

6, 2

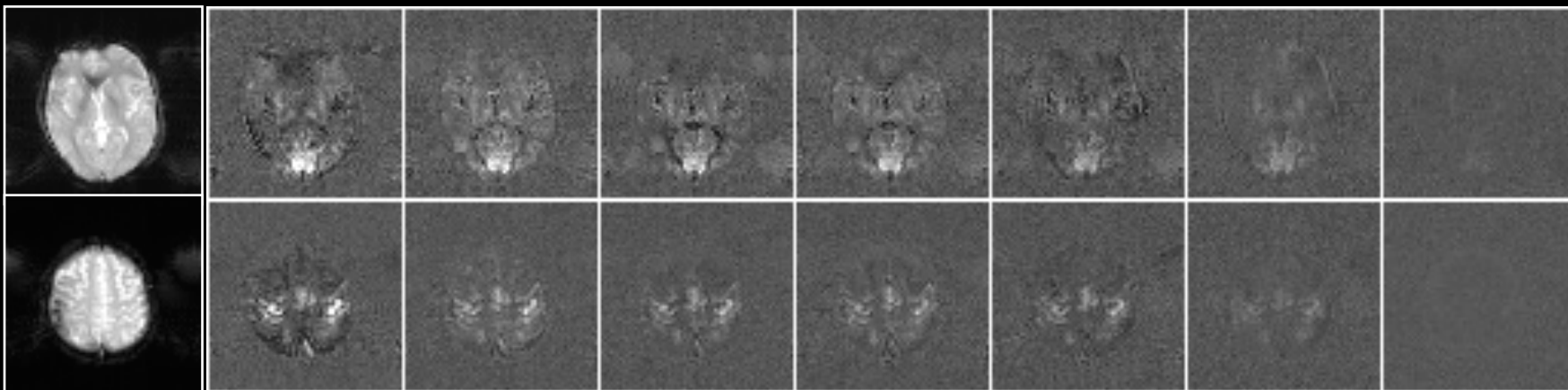
4, 2

2, 2

S1



S2

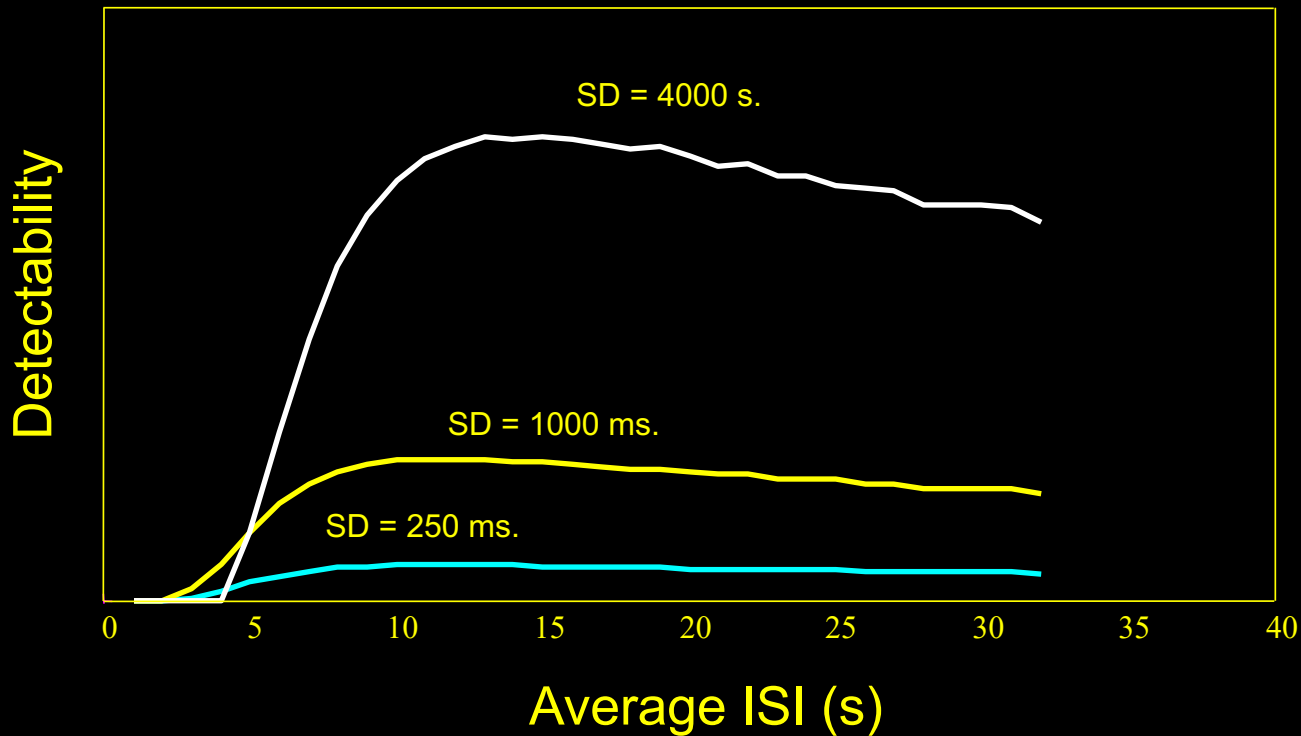
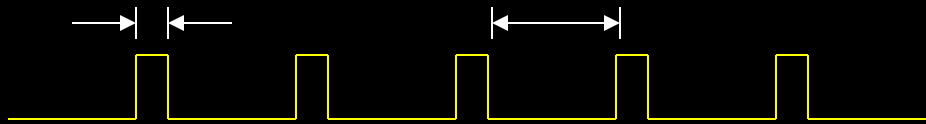


P. A. Bandettini, R. W. Cox. Functional contrast in constant interstimulus interval event - related fMRI: theory and experiment. *Magn. Reson. Med.* 43: 540-548 (2000).

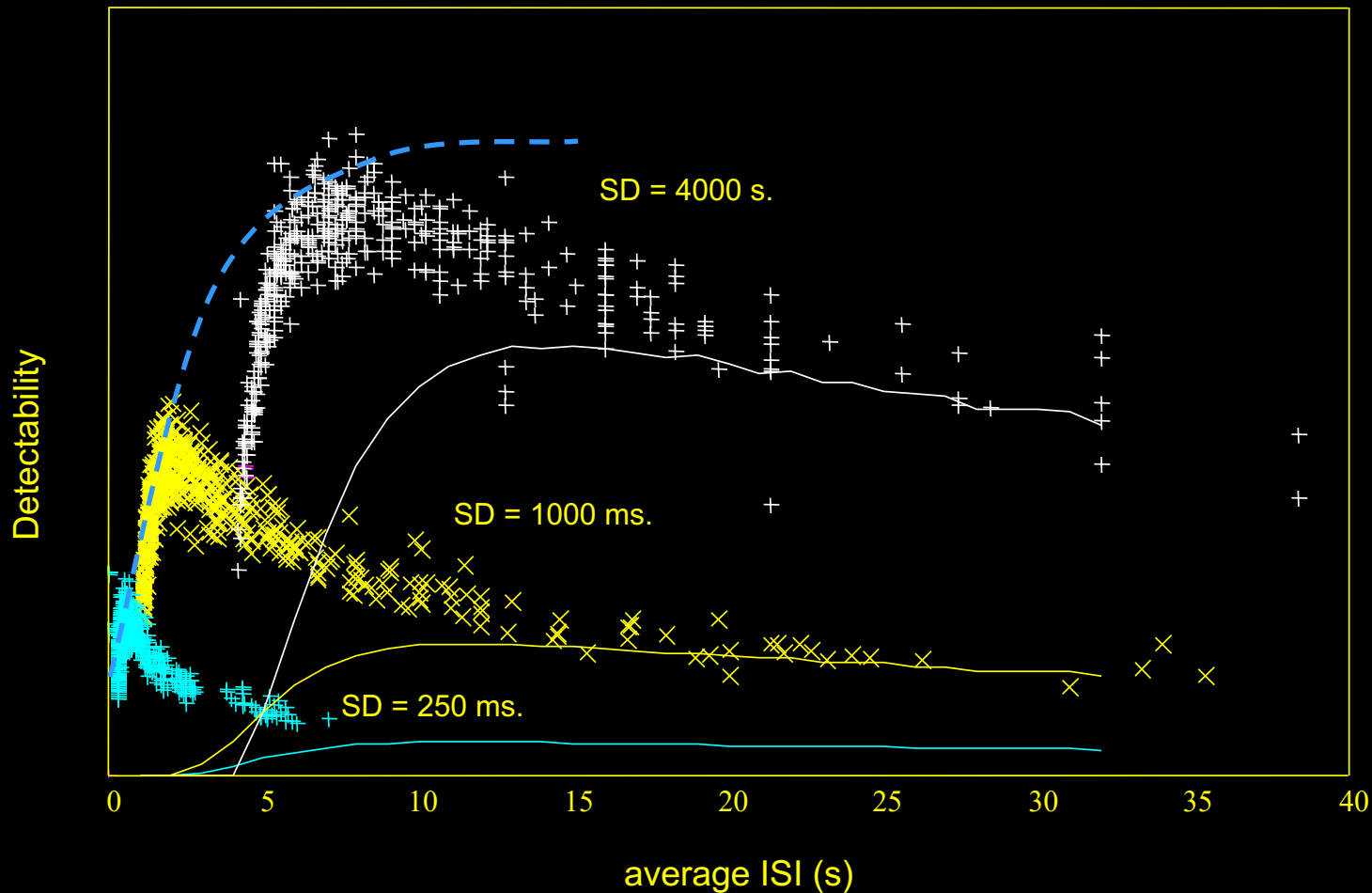
Detectability – constant ISI

SD – stimulus duration

ISI – inter-stimulus interval

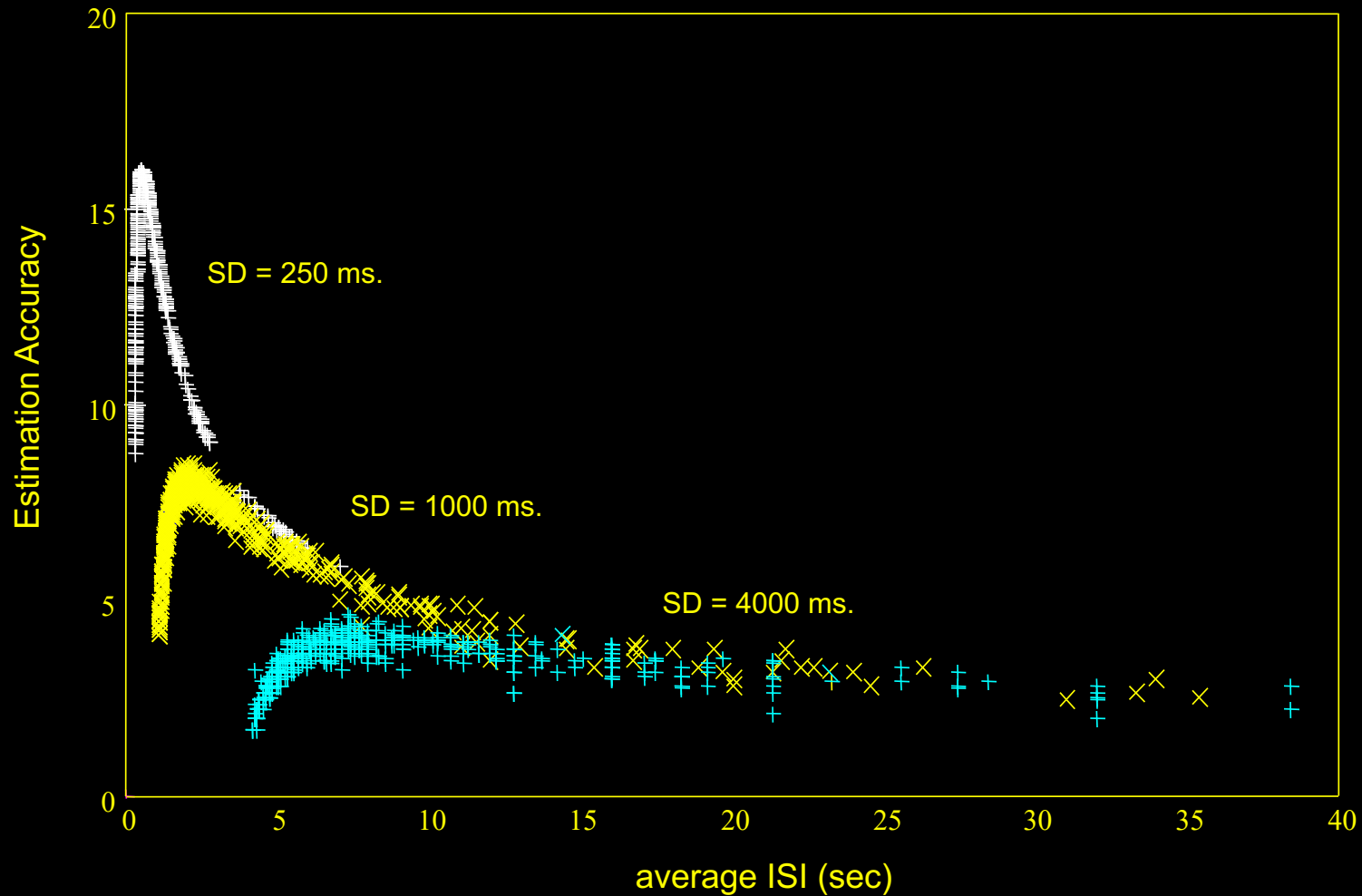


Detectability vs. Average ISI



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

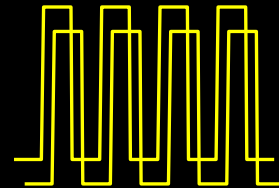
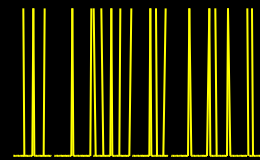
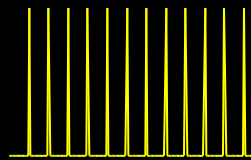
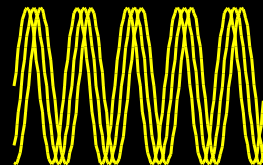
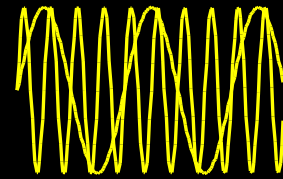
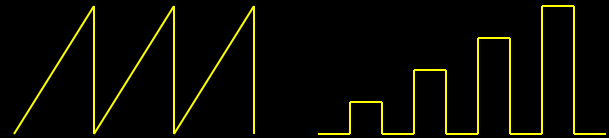
3. Frequency Encoding

4. Phase Encoding

5. Event Related

6. Orthogonal Design

7. Free Behavior Design



Neuronal Activation Input Strategies

1. Block Design

2. Parametric Design

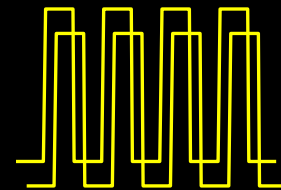
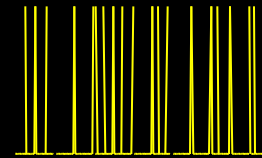
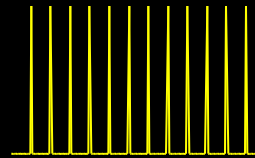
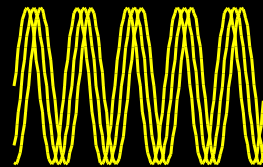
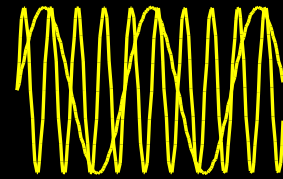
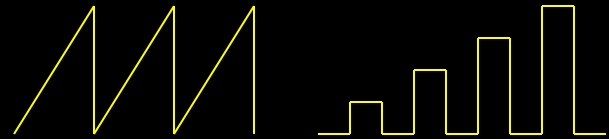
3. Frequency Encoding

4. Phase Encoding

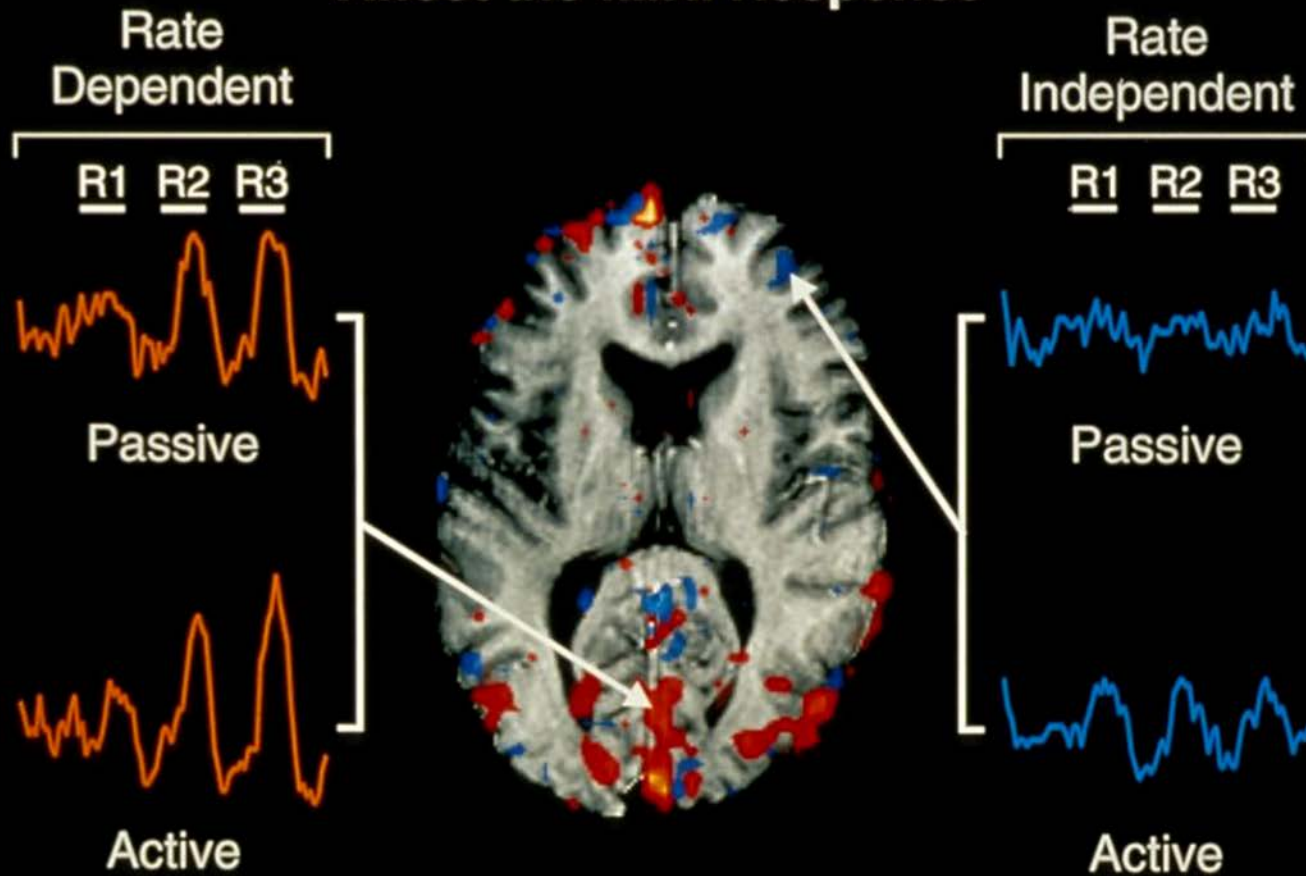
5. Event Related

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

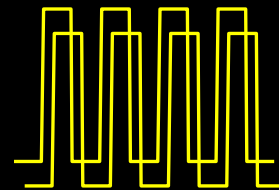
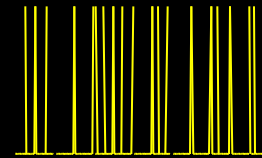
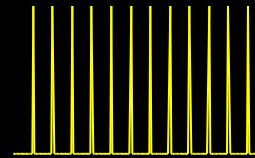
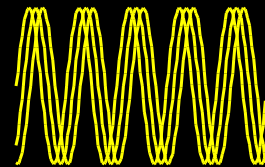
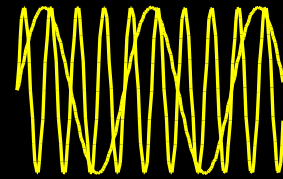
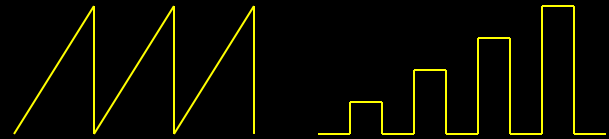
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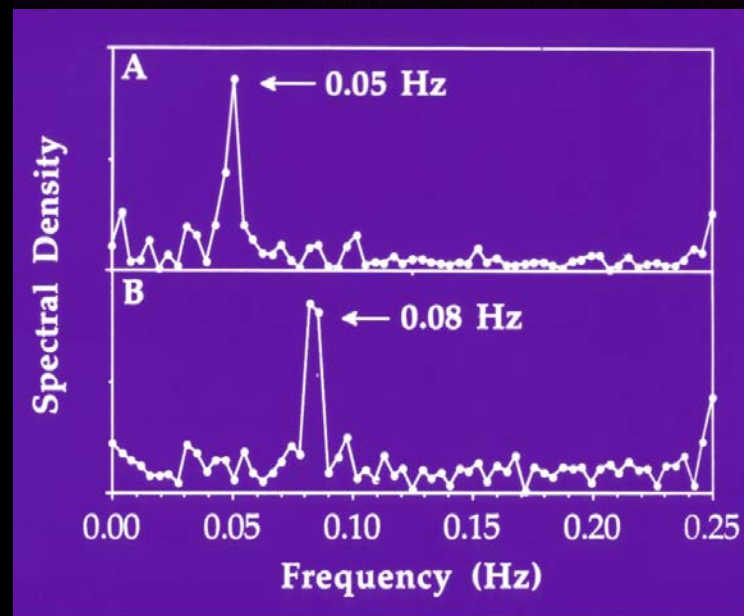
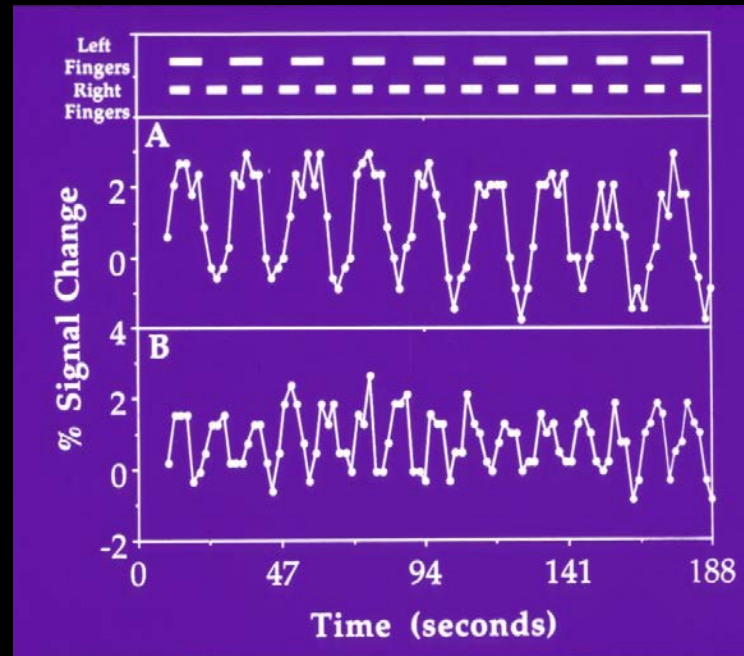
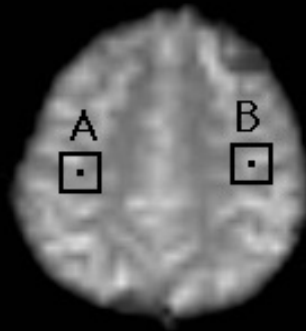
4. Phase Encoding

5. Event Related

6. Orthogonal Design

7. Free Behavior Design



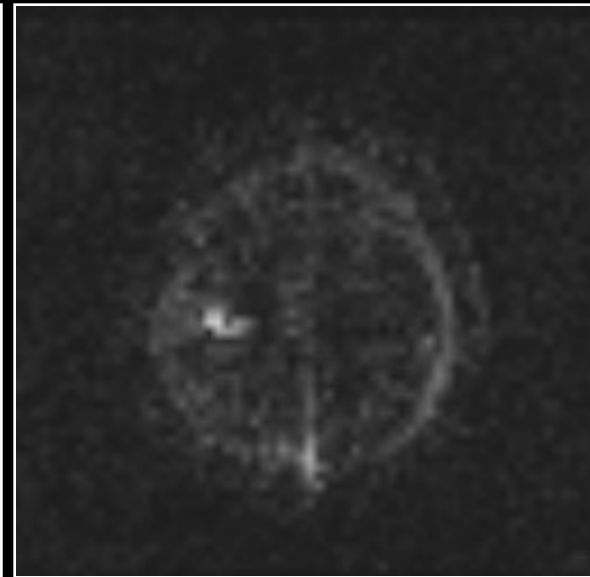
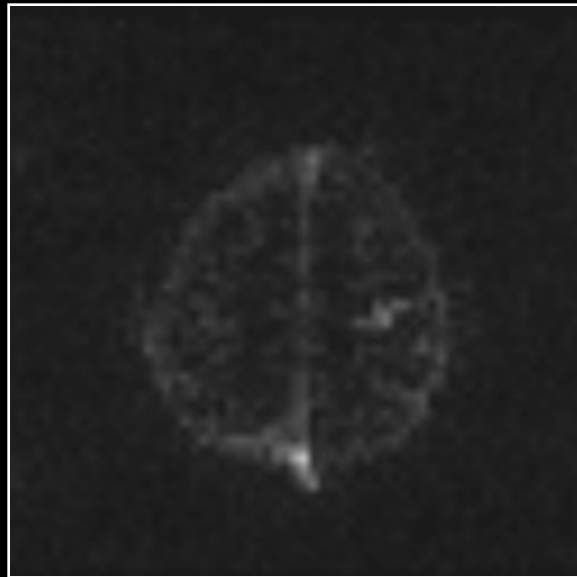


P. A. Bandettini, A. Jesmanowicz, E. C. Wong, J. S. Hyde, Processing strategies for time-course data sets in functional MRI of the human brain. *Magn. Reson. Med.* 30, 161-173 (1993).

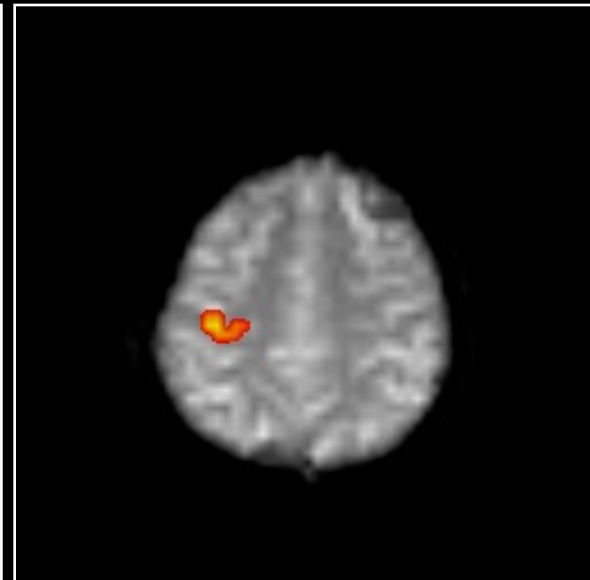
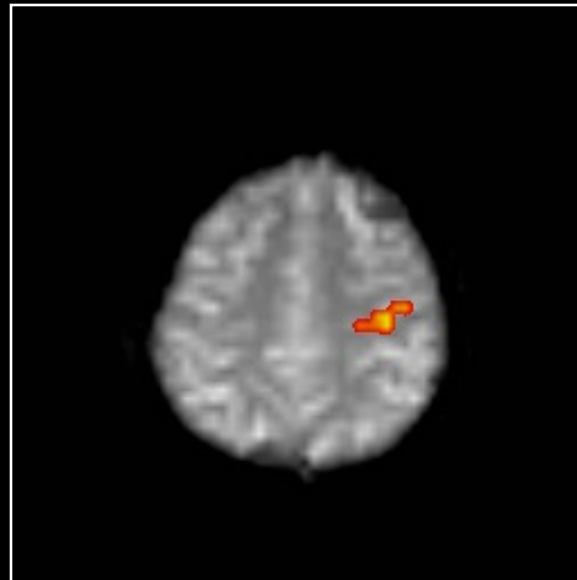
0.08 Hz

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

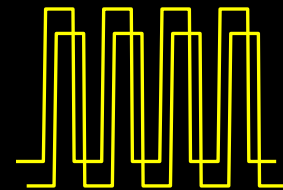
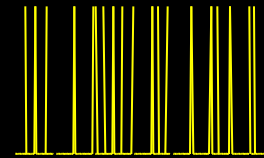
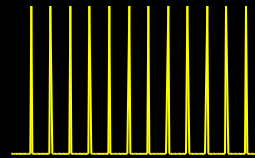
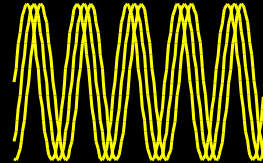
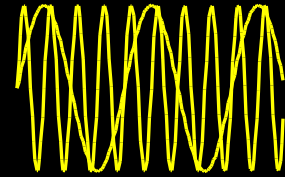
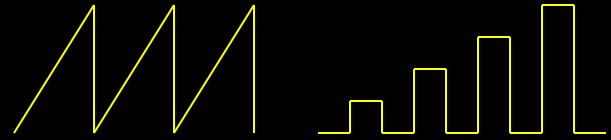
3. Frequency Encoding

4. Phase Encoding

5. Event Related

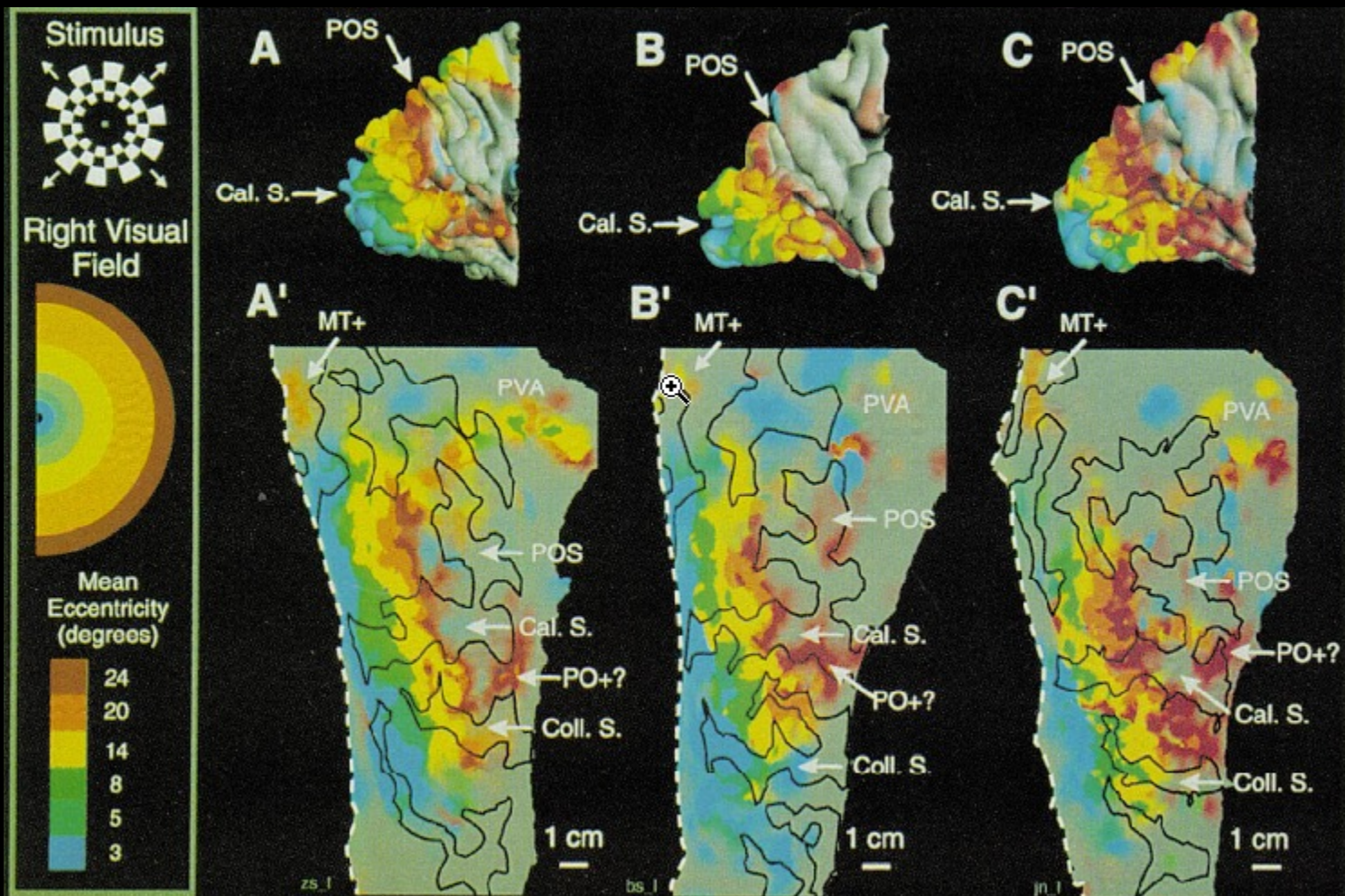
6. Orthogonal Design

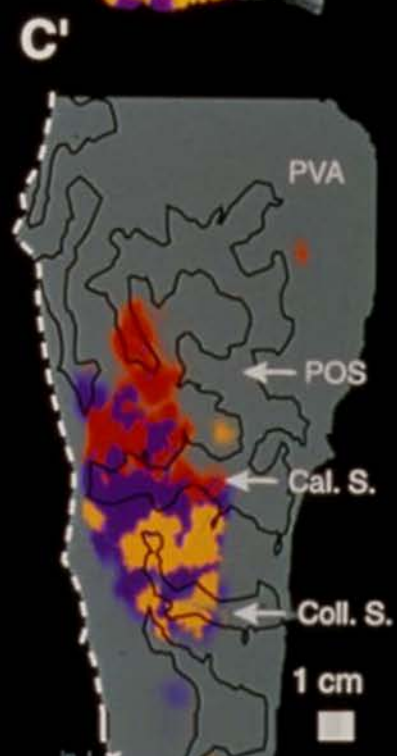
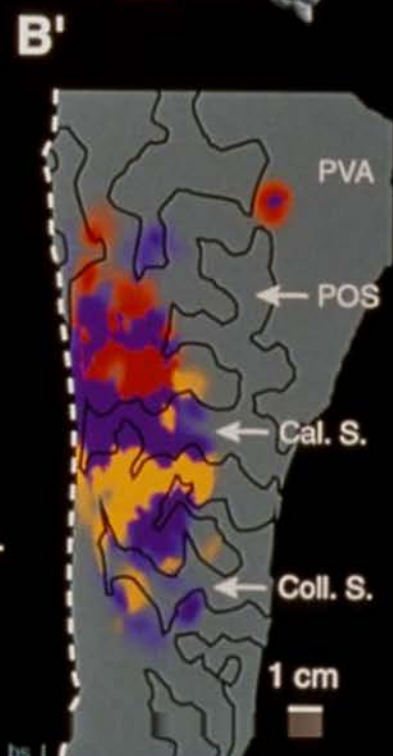
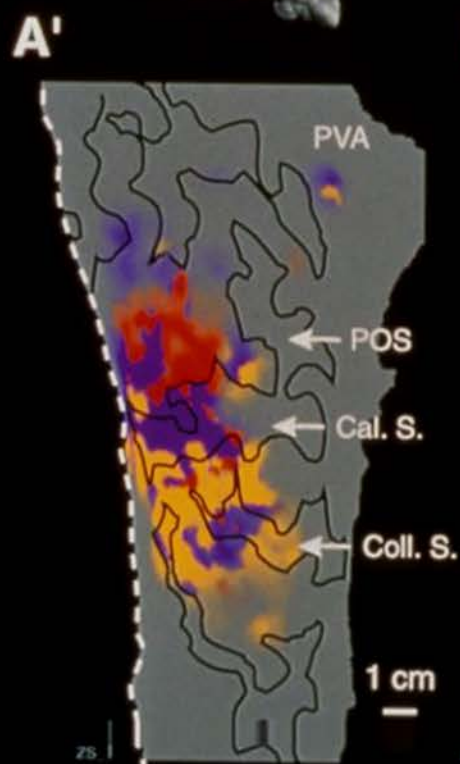
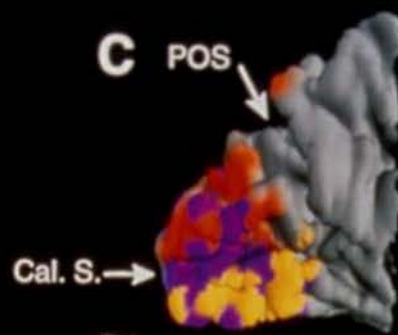
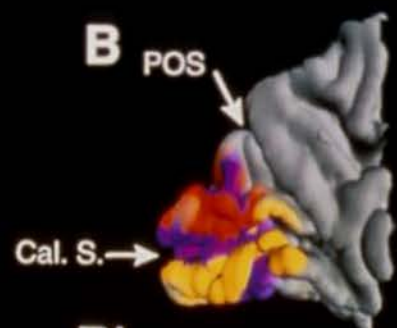
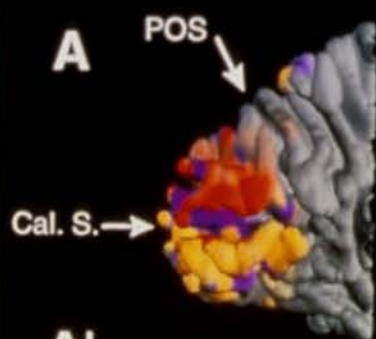
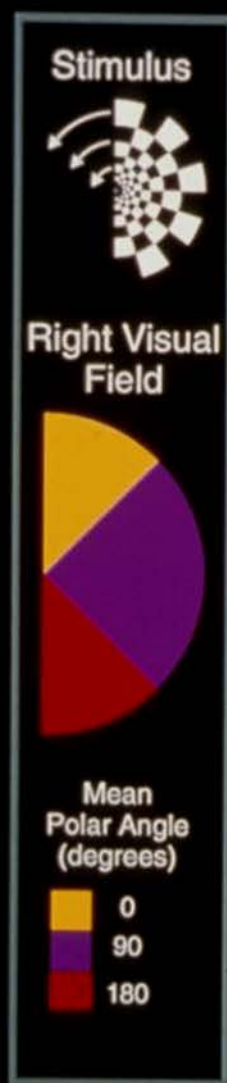
7. Free Behavior Design



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

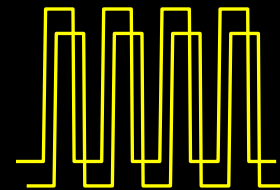
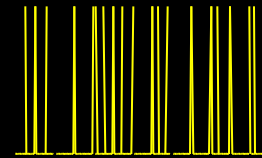
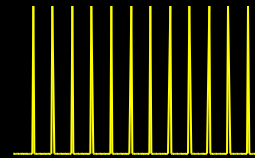
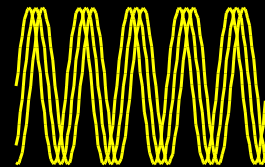
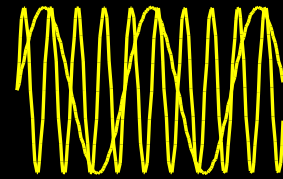
3. Frequency Encoding

4. Phase Encoding

5. Event Related

6. Orthogonal Design

7. Free Behavior Design

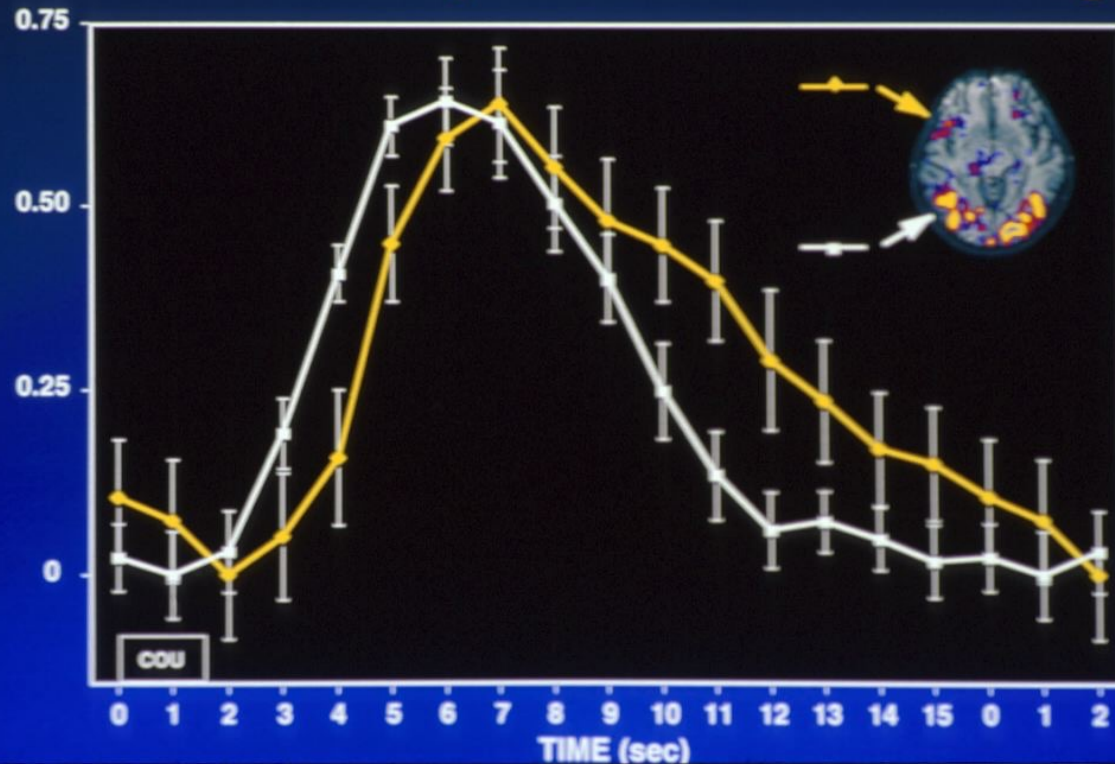


Detection of cortical activation during averaged single trials of a cognitive task using functional magnetic resonance imaging

(neuroimaging/single trial/language/prefrontal)

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

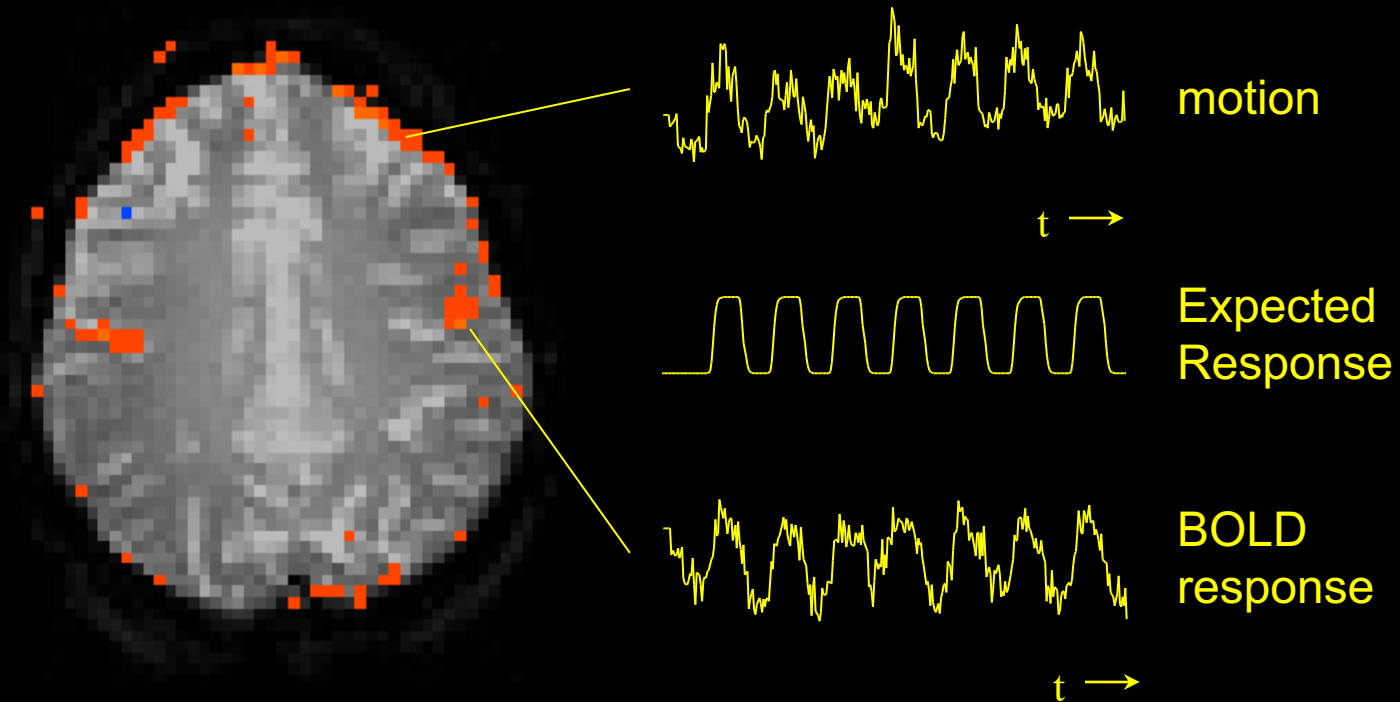
Time Course Comparison Across Brain Regions



Event Related Advantages

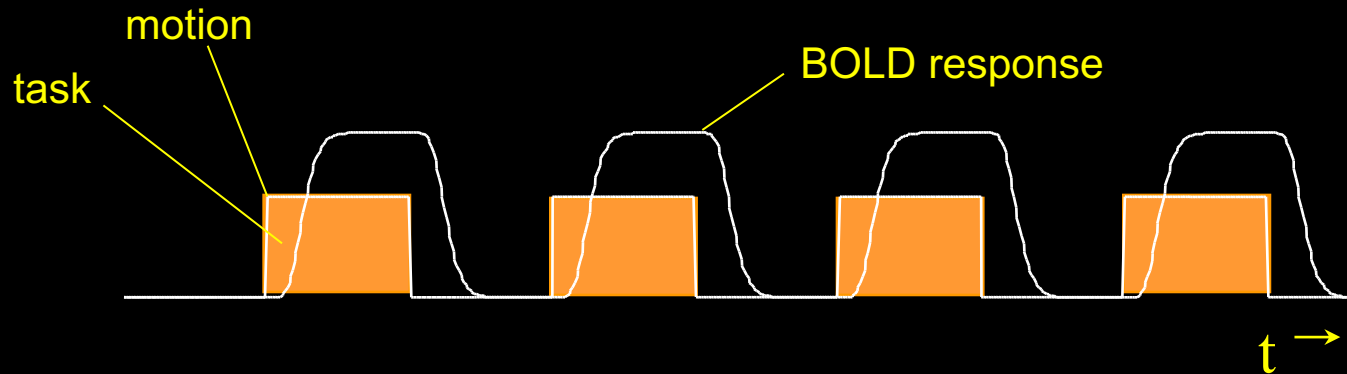
- Task Randomization
- Post acquisition, Performance-based, data binning
- Natural presentation
- Reduction of habituation effects
- Overt responses
- Reduction of scanner noise effects
- More precise estimation of hemodynamic responses

Speaking - Blocked Trial

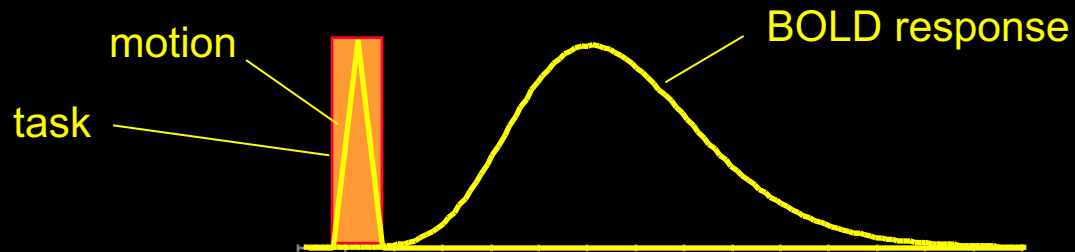


fMRI during tasks that involve brief motion

Blocked Design

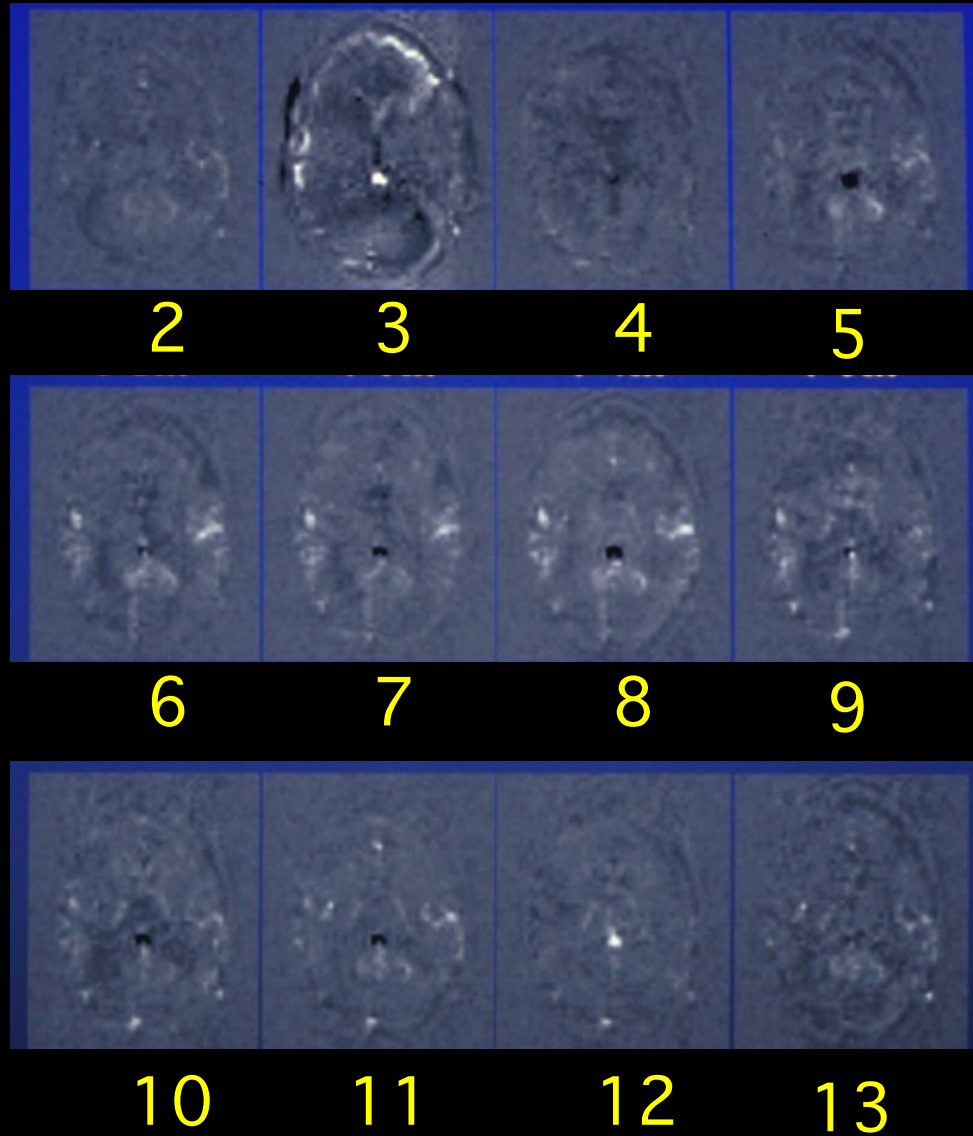


Event-Related Design



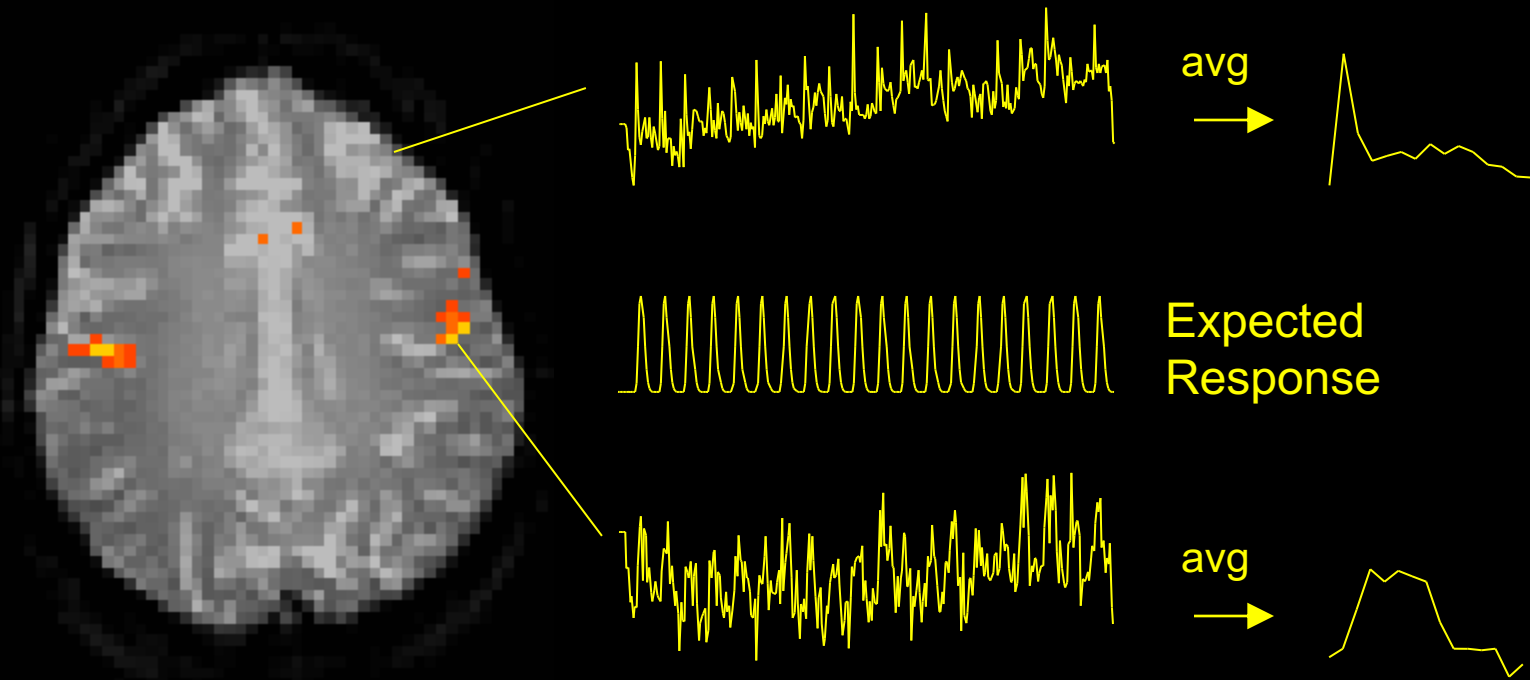
R. M. Birn, P. A. Bandettini, R. W. Cox, R. Shaker, Event - related fMRI of tasks involving brief motion. *Human Brain Mapping* 7: 106-114 (1999).

Overt Word Production



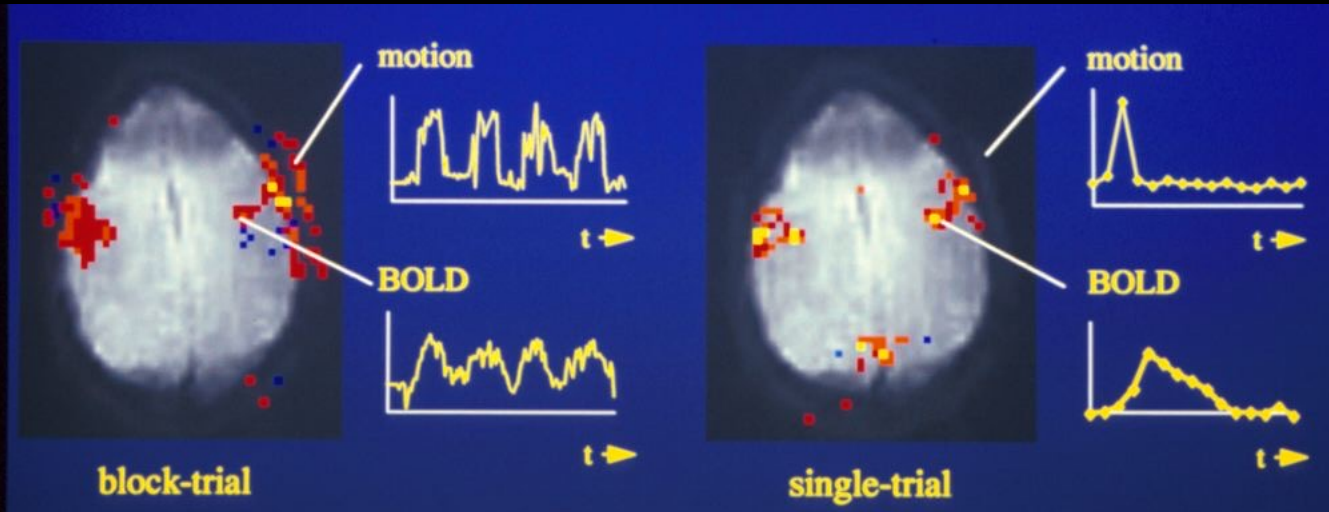
R. M. Birn, P. A. Bandettini, R. W. Cox, R. Shaker, Event - related fMRI of tasks involving brief motion. *Human Brain Mapping* 7: 106-114 (1999).

Speaking - ER-fMRI

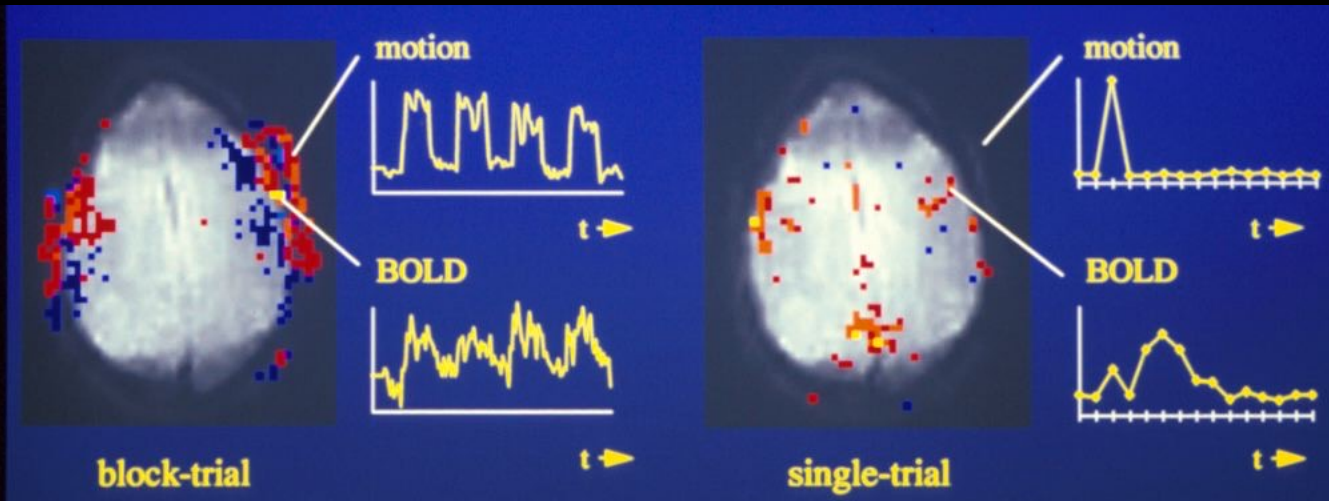


R. M. Birn, P. A. Bandettini, R. W. Cox, R. Shaker, Event - related fMRI of tasks involving brief motion. *Human Brain Mapping* 7: 106-114 (1999).

Tongue Movement



Jaw Clenching



Neuronal Activation Input Strategies

1. Block Design

2. Parametric Design

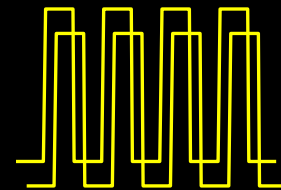
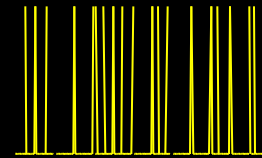
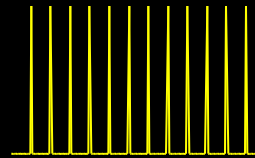
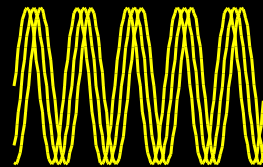
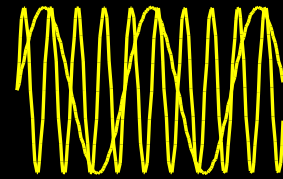
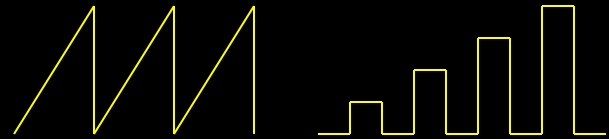
3. Frequency Encoding

4. Phase Encoding

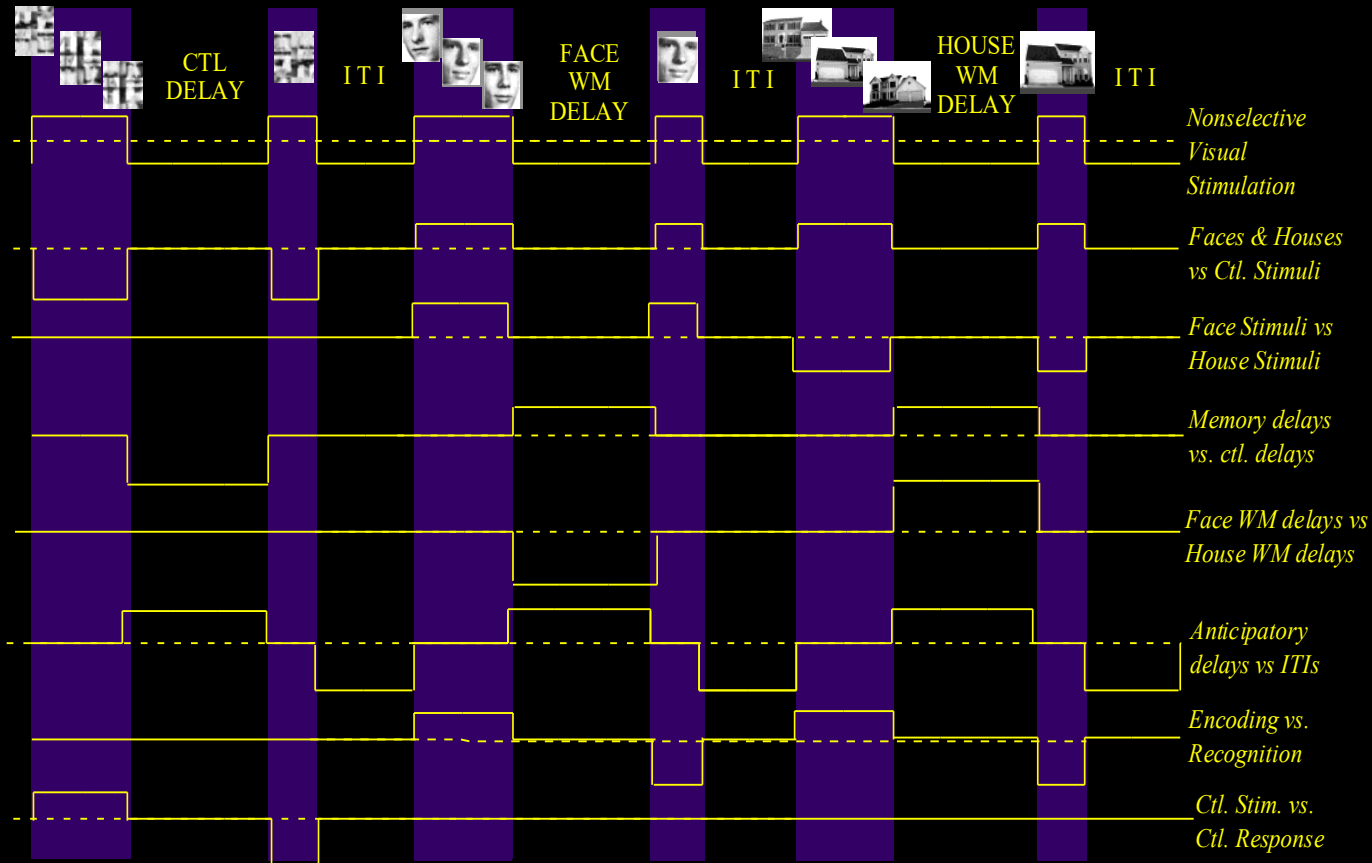
5. Event Related

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

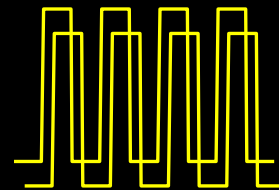
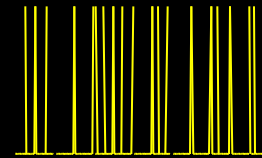
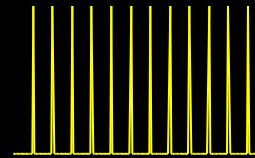
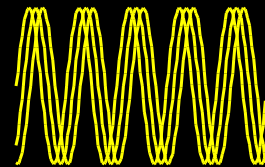
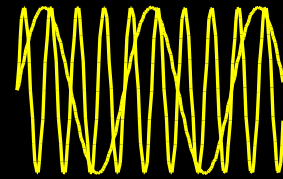
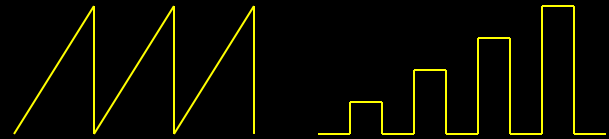
3. Frequency Encoding

4. Phase Encoding

5. Event Related

6. Orthogonal Design

7. Free Behavior Design

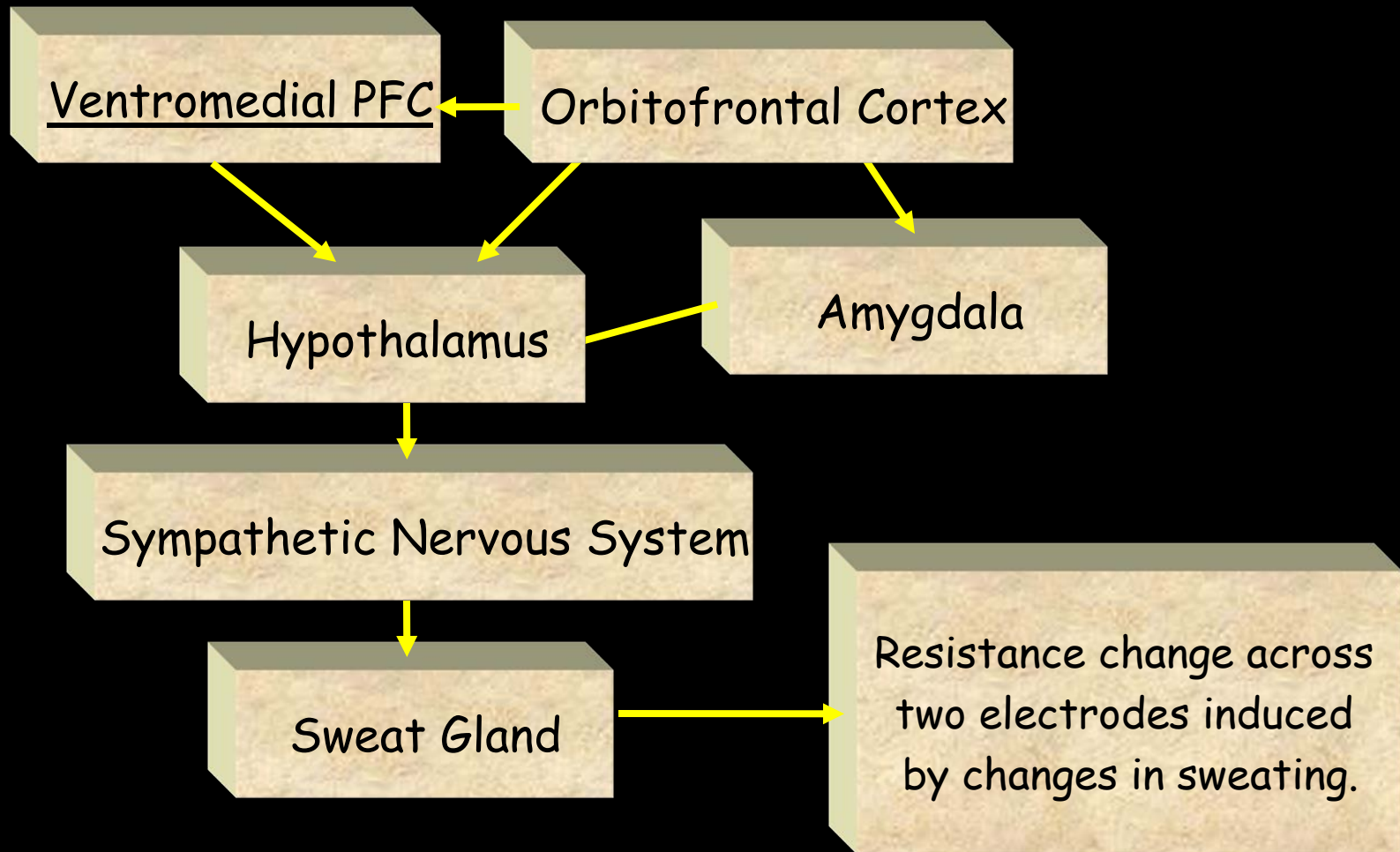


Free Behavior Design

Use a continuous measure as a reference function:

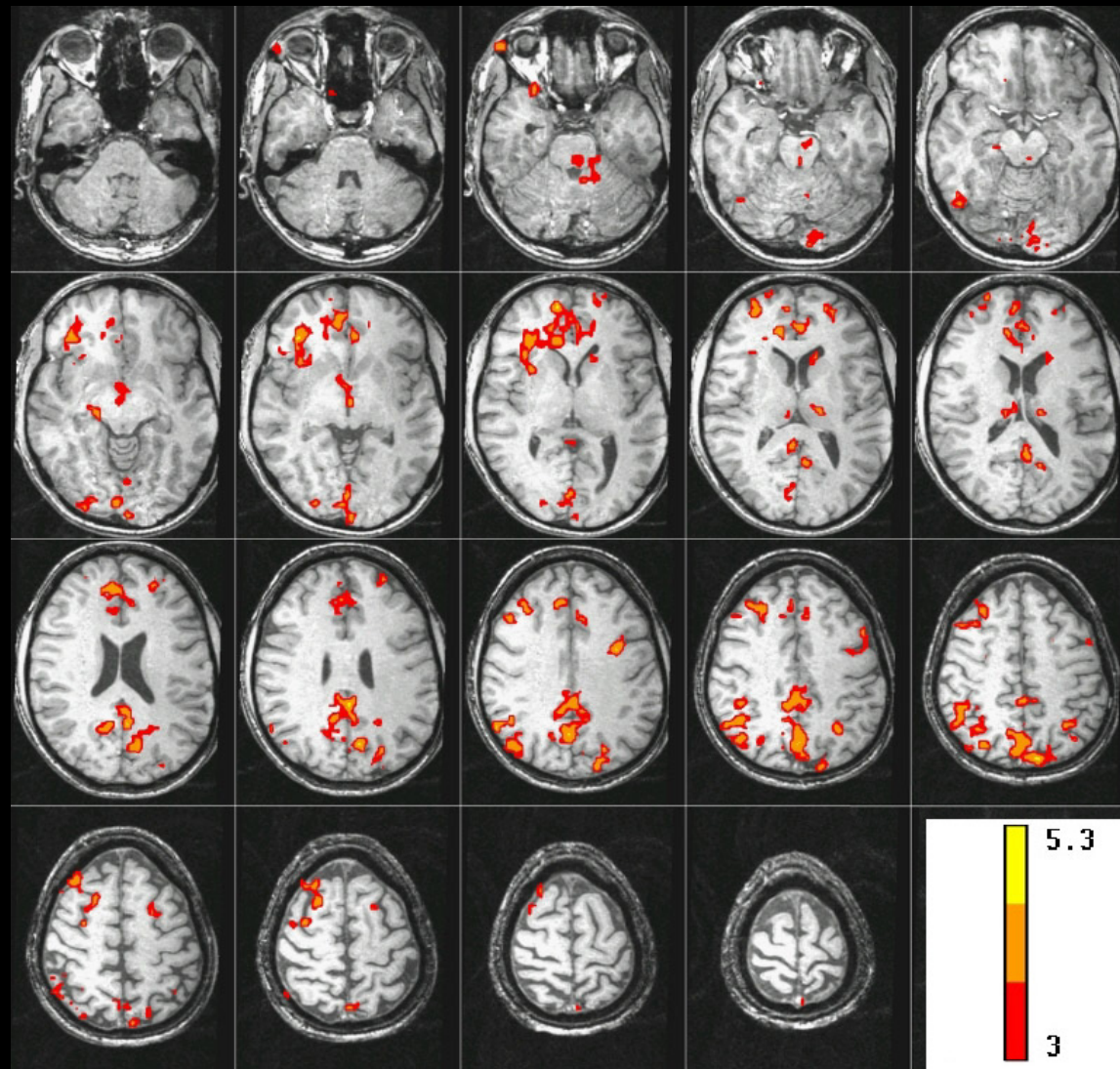
- Task performance
- Skin Conductance
- Heart, respiration rate..
- Eye position
- EEG

The Skin Conductance Response (SCR)



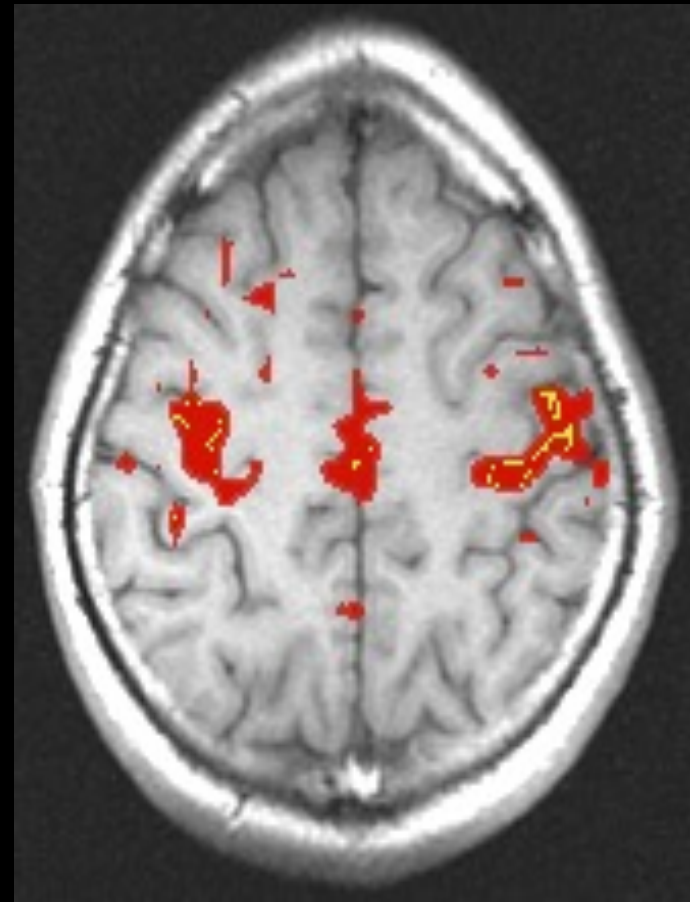
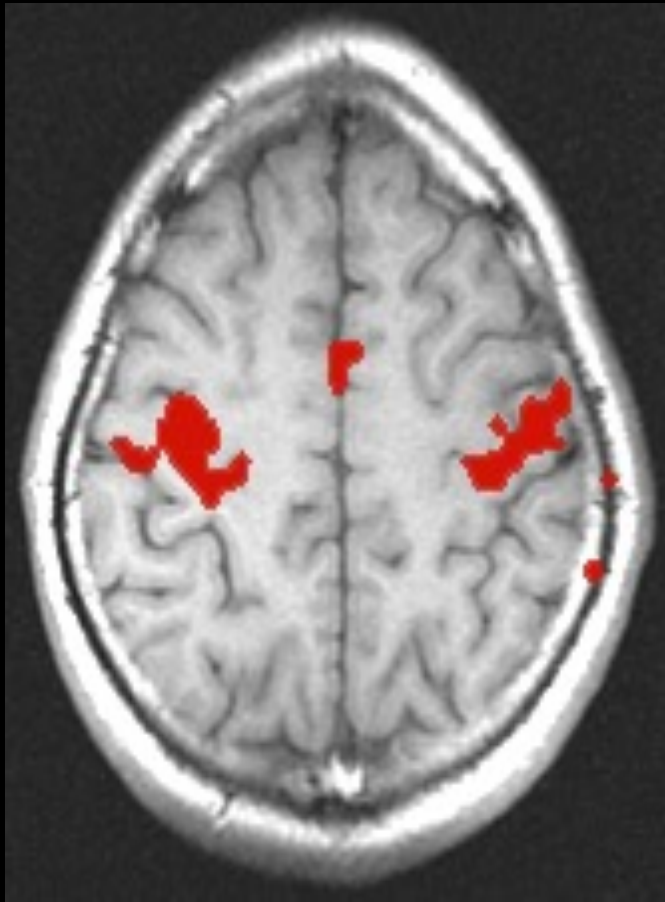
Patterson et al. (submitted)

Brain activity correlated with SCR during “Rest”

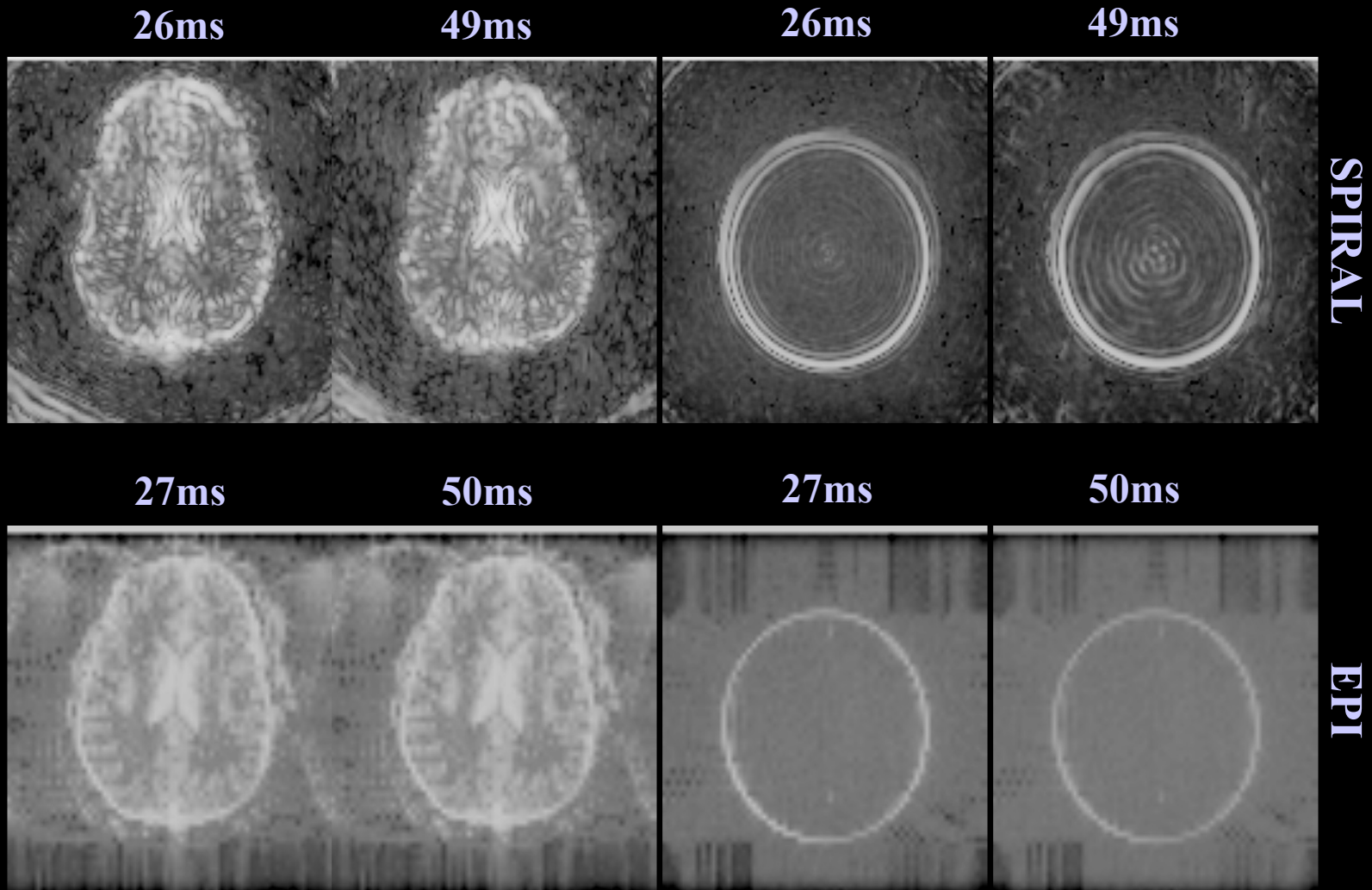


Patterson et al. (submitted)

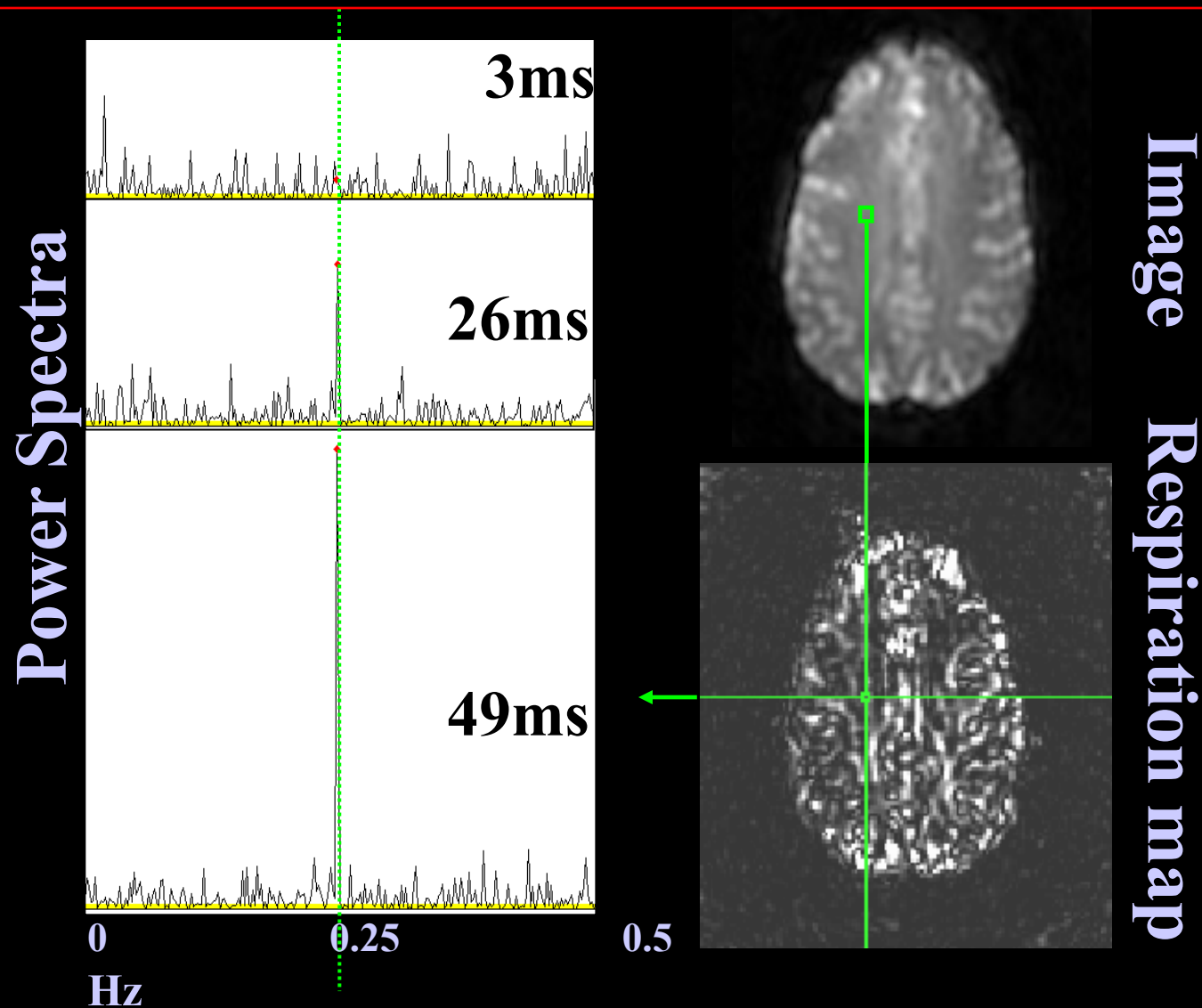
Resting Hemodynamic Autocorrelations



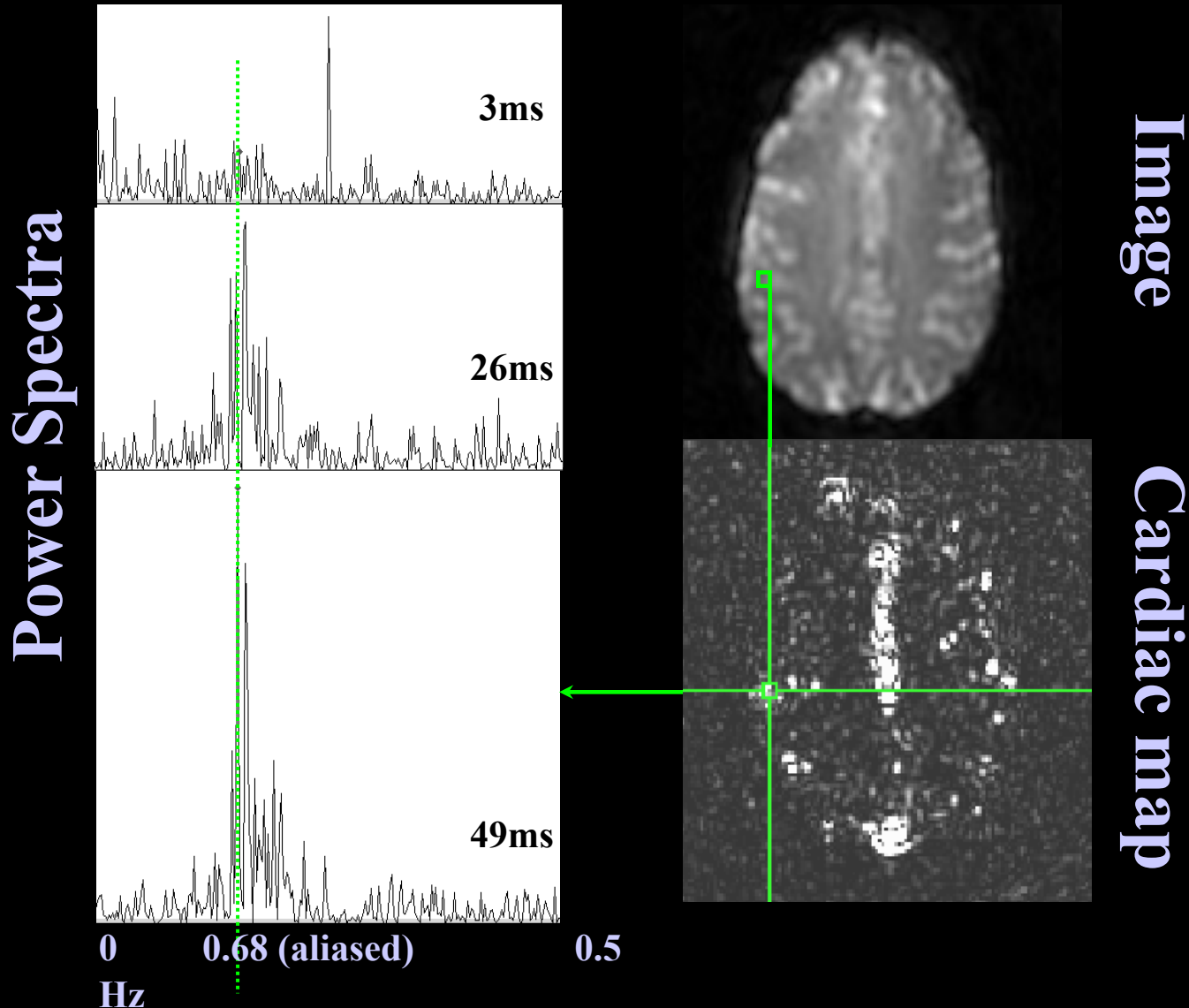
Temporal vs. Spatial SNR- 3T



0.25 Hz Breathing at 3T

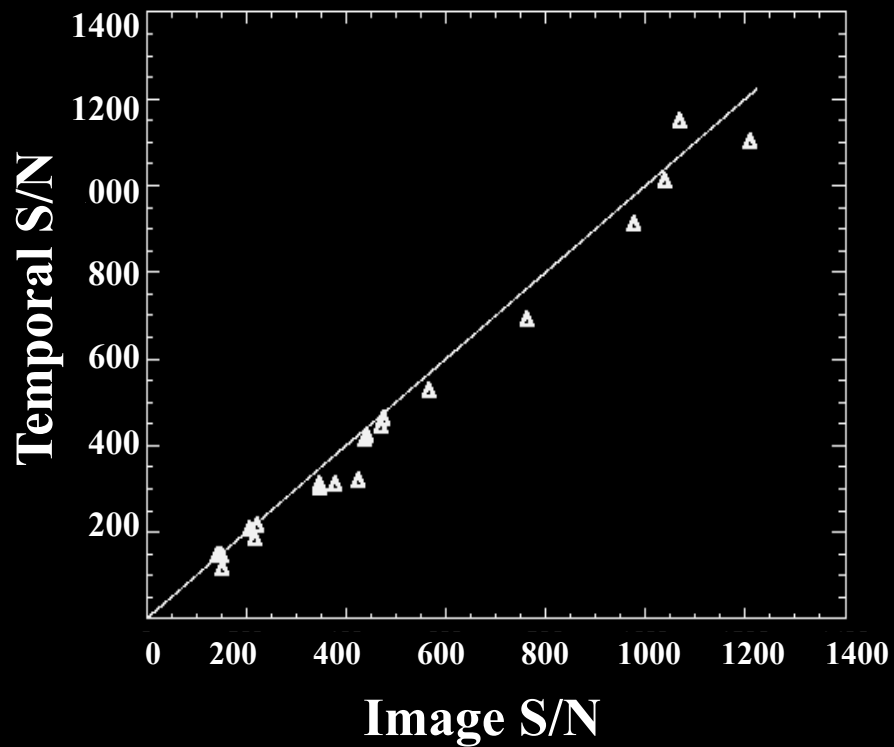


0.68 Hz Cardiac rate at 3T

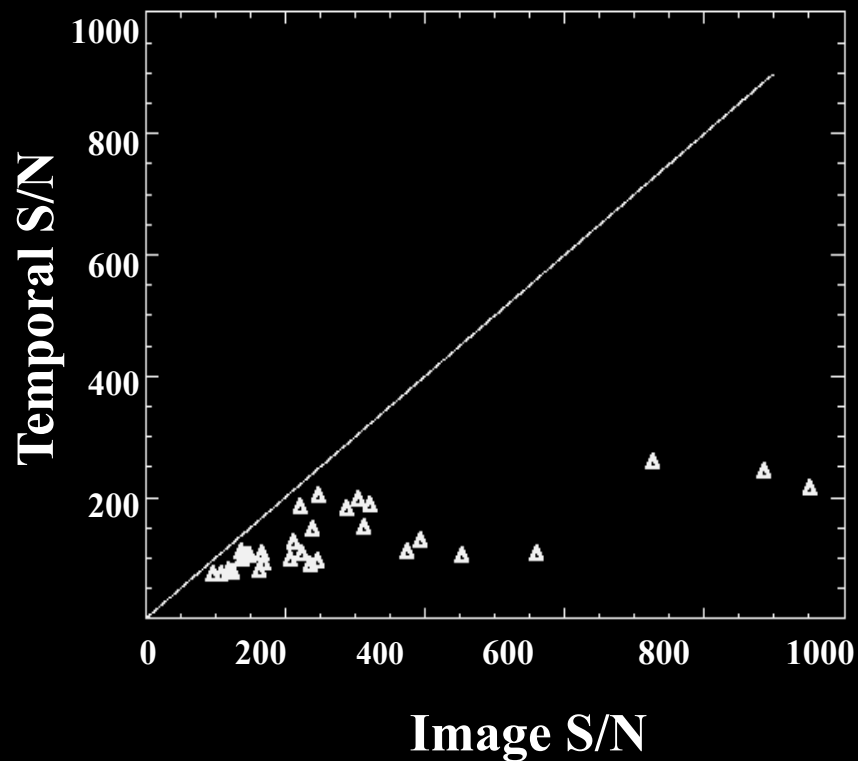


Temporal S/N vs. Image S/N

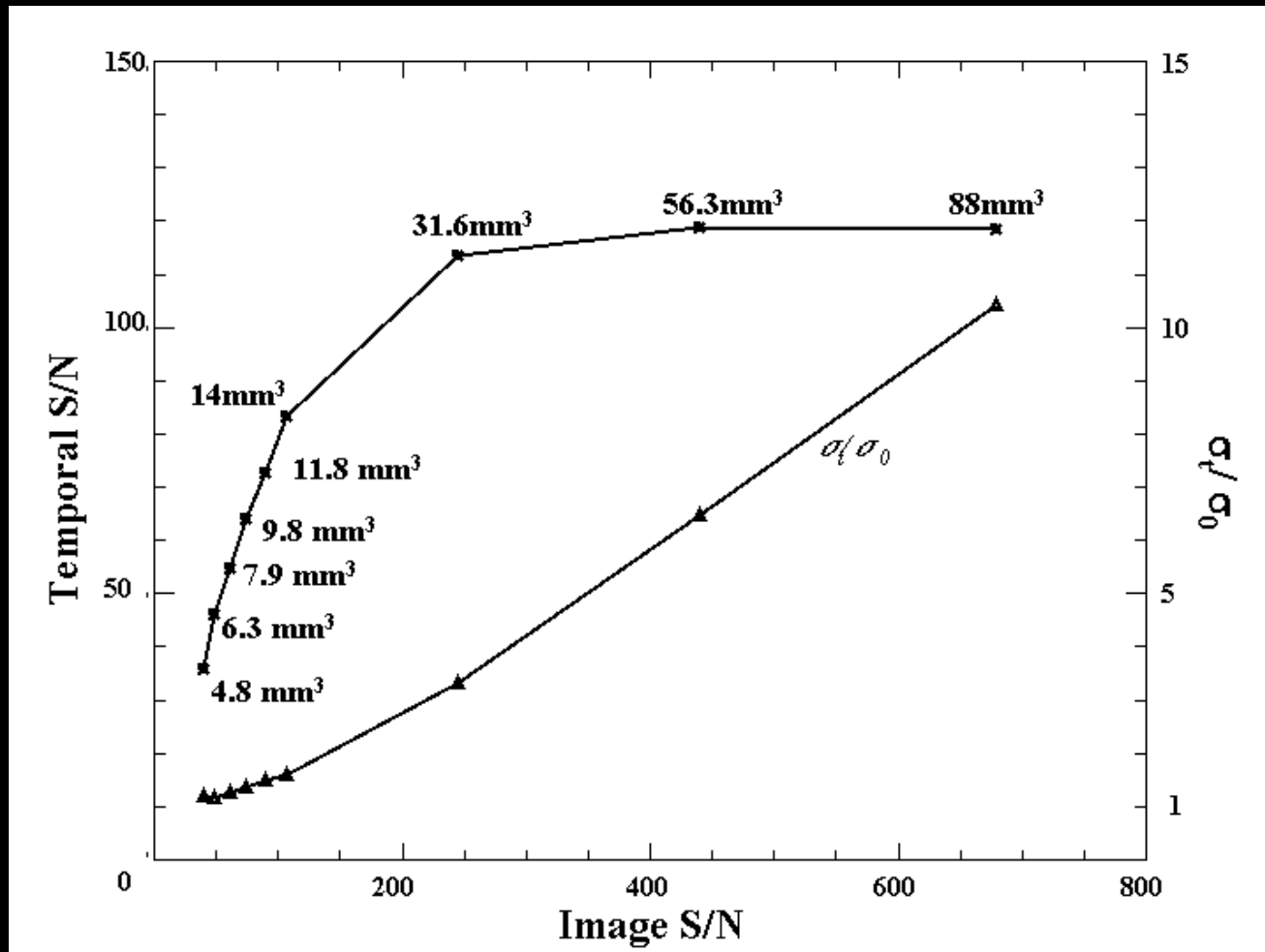
PHANTOMS



SUBJECTS



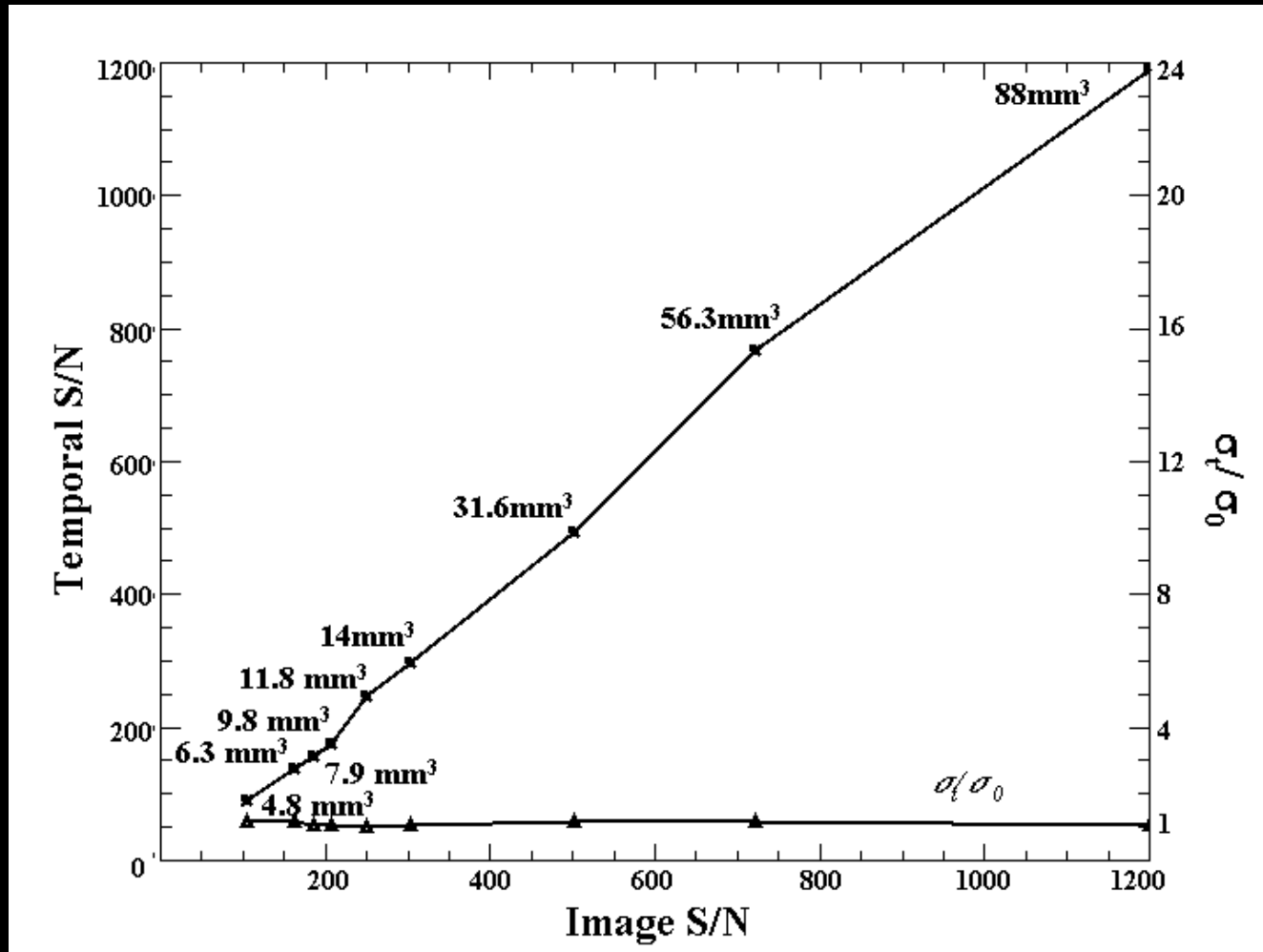
Temporal vs. Image S/N Optimal Resolution Study



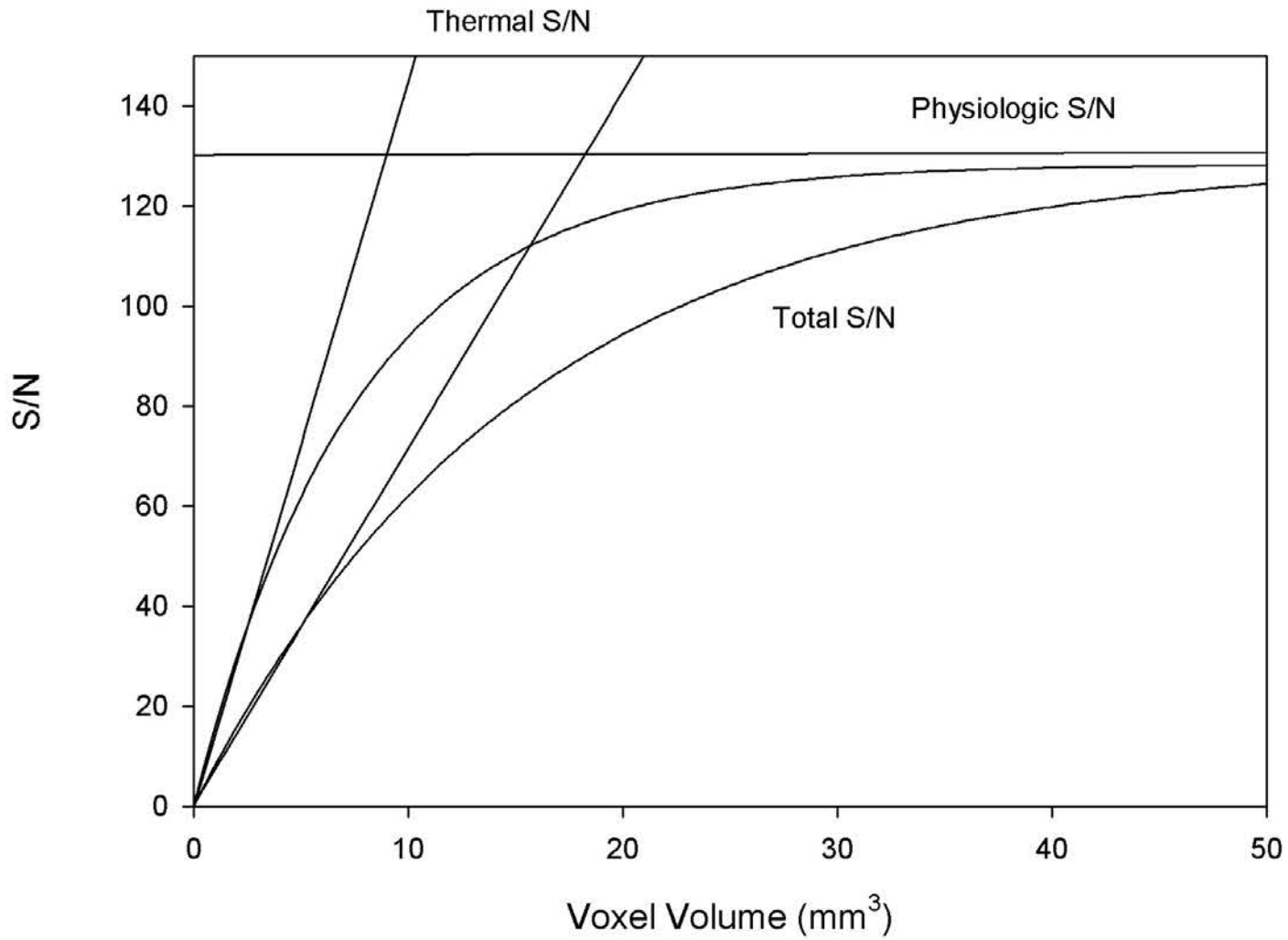
Human data

Petridou et al

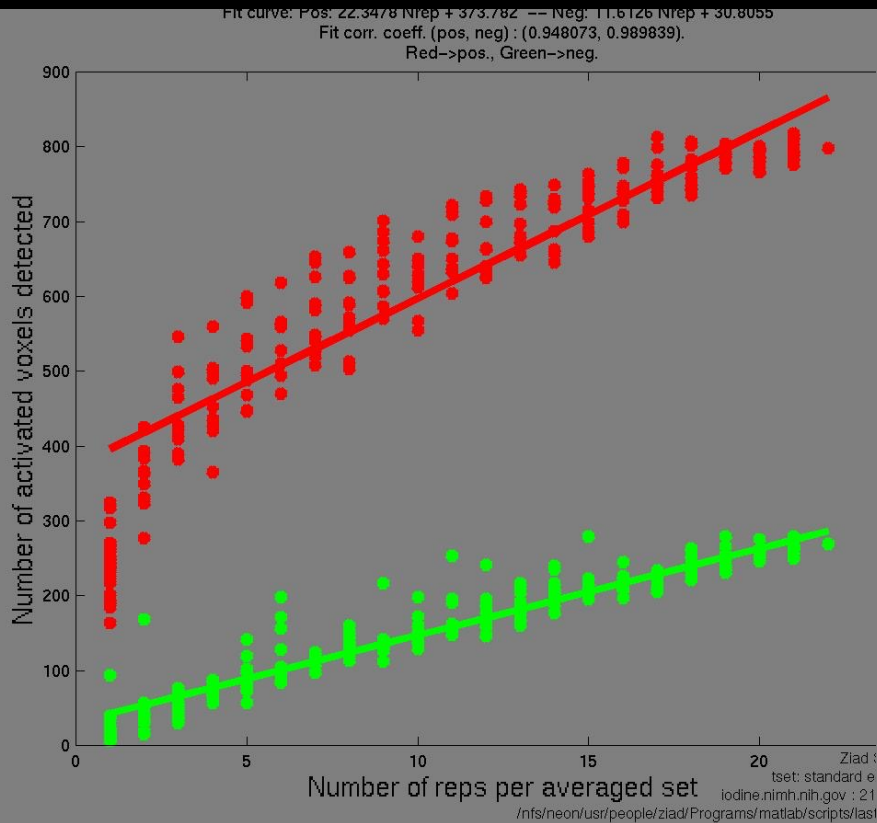
Temporal vs. Image S/N Optimal Resolution Study



Phantom data

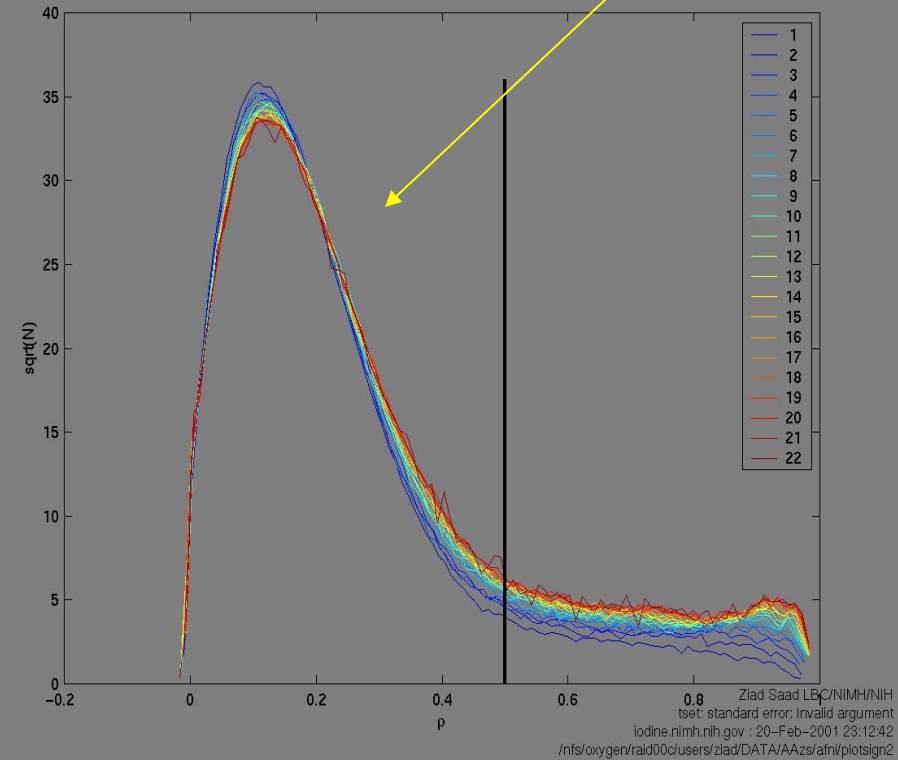


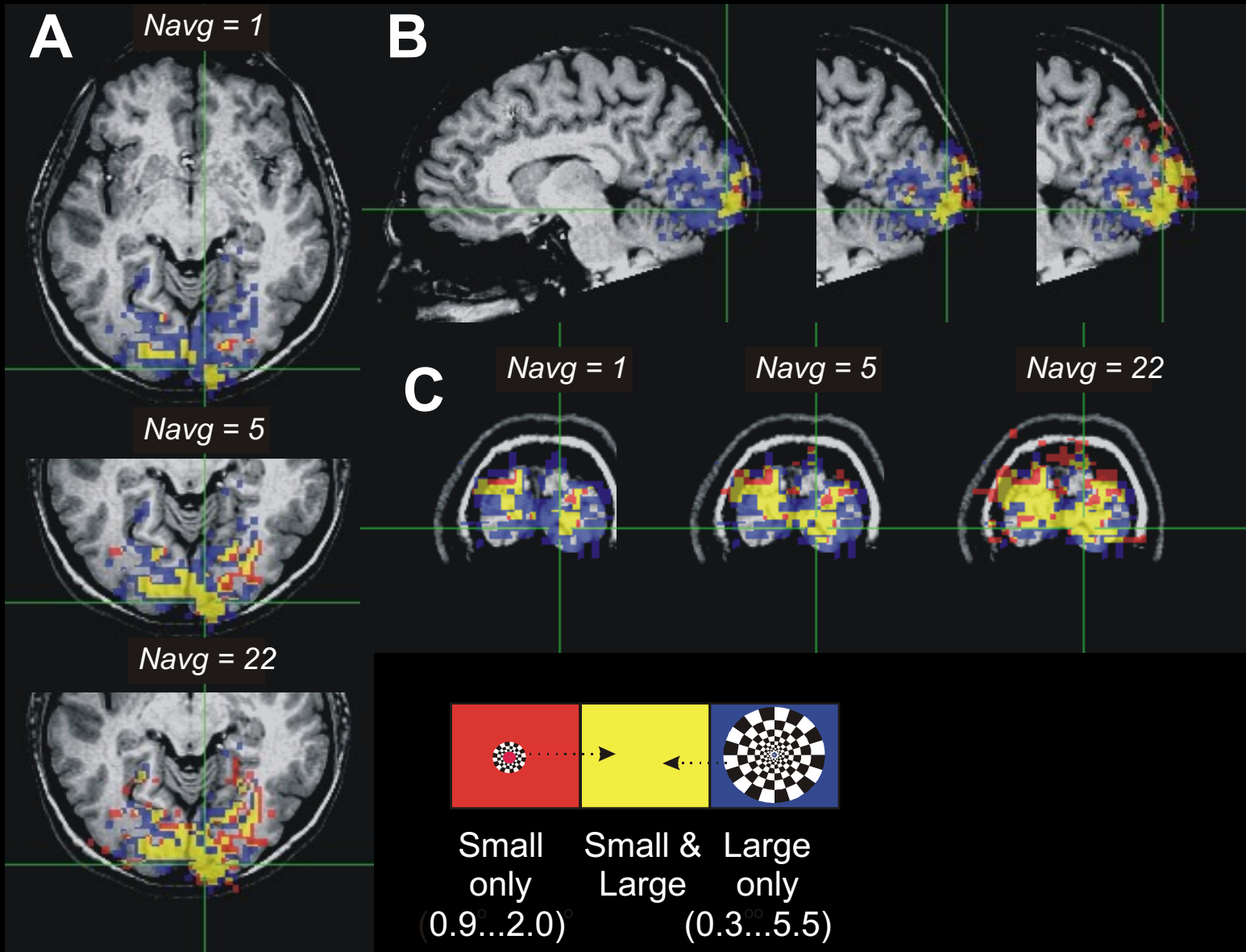
Continuously Growing Activation Area



CC Histogram

Inflection Point





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

Methodology

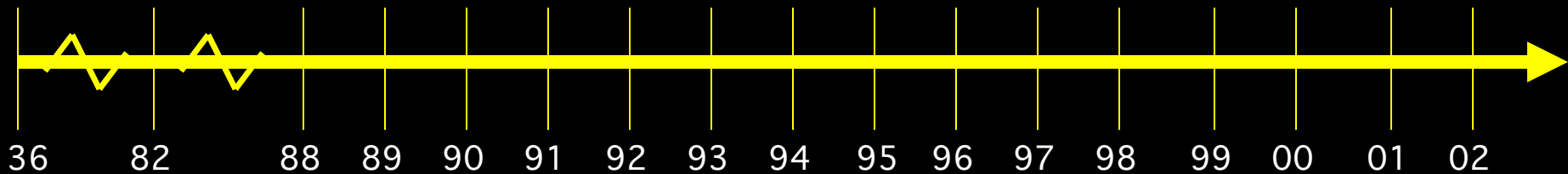
Baseline Volume
 IVIM
 Correlation Analysis
 Parametric Design
 Surface Mapping
 Phase Mapping
 Linear Regression
 Event-related
 Motion Correction
 Multi-Modal Mapping
 Free-behavior Designs
 Mental Chronometry
 Deconvolution
 CO₂ Calibration

Interpretation

Blood T2
 Hemoglobin
 BOLD models
 B₀ dep.
 TE dep
 SE vs. GE
 NIRS Correlation
 Veins
 PET correlation
 IV vs EV
 Pre-undershoot
 Resolution Dep.
 Post-undershoot
 CO₂ effect
 Inflow
 ASL vs. BOLD
 PSF of BOLD
 Extended Stim.
 Linearity
 Fluctuations
 Balloon Model
 Metab. Correlation
 Optical Im. Correlation
 Electrophys. correlation

Applications

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

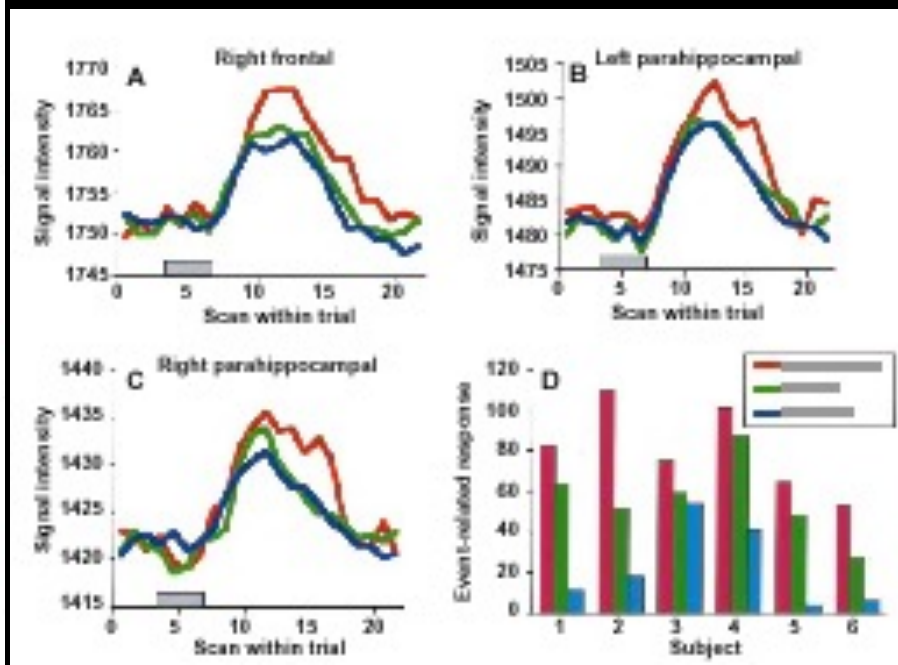
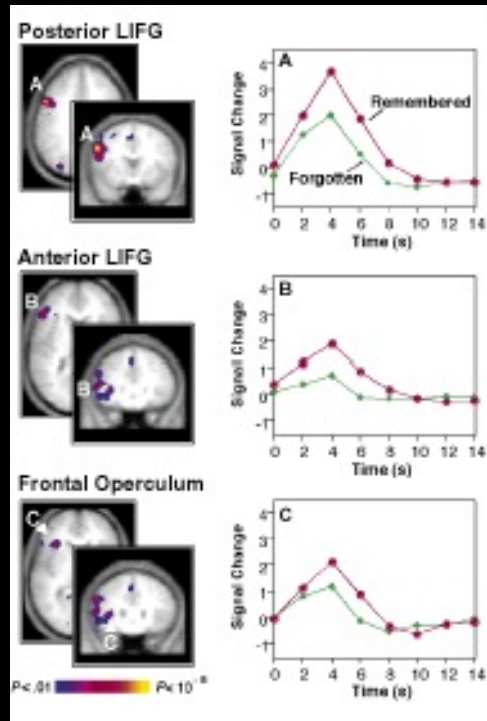
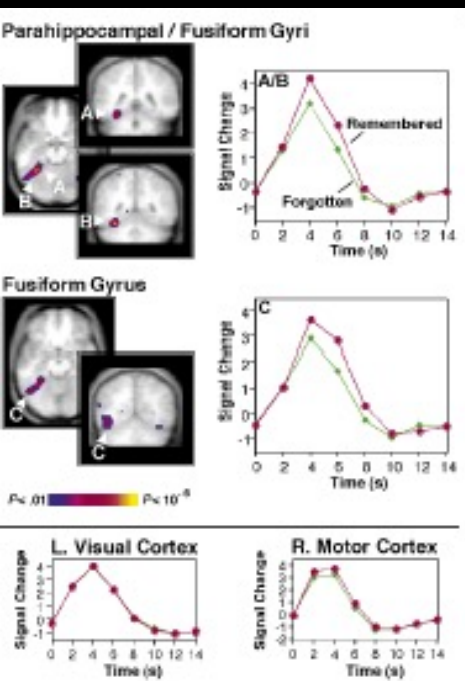


Building Memories: Remembering and Forgetting of Verbal Experiences as Predicted by Brain Activity

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



Science, Vol 281, August 1998

Technology

MRI

1.5T,3T, 4T

EPI

Local Human Head Gradient Coils

ASL

BOLD

EPI on Clin. Syst.

Nav. pulses

Spiral EPI

Multi-shot fMRI

Diff. tensor

Real time fMRI

Quant. ASL

Dynamic IV volume

Simultaneous ASL and BOLD

Mg⁺

Venography

Z-shim

Baseline Susceptibility

7T

SENSE

Current Imaging?

Methodology

Baseline Volume

IVIM

Correlation Analysis

Parametric Design

Surface Mapping

Phase Mapping

Linear Regression

Event-related

Motion Correction

Multi-Modal Mapping

Free-behavior Designs

Mental Chronometry

Deconvolution

CO₂ Calibration

Interpretation

Blood T2

Hemoglobin

BOLD models

B₀ dep.

TE dep

SE vs. GE

NIRS Correlation

Veins

PET correlation

IV vs EV

Pre-undershoot

Resolution Dep.

Post-undershoot

CO₂ effect

NIRS Correlation

Inflow

PSF of BOLD

Extended Stim.

Linearity

Fluctuations

Balloon Model

Metab. Correlation

Optical Im. Correlation

Electrophys. correlation

Applications

Complex motor Language

Imagery

Memory

Emotion

Motor learning

Children

Tumor vasc.

Drug effects

BOLD -V1, M1, A1

Presurgical

Attention

Ocular Dominance

Volume - Stroke

V1, V2..mapping

Priming/Learning

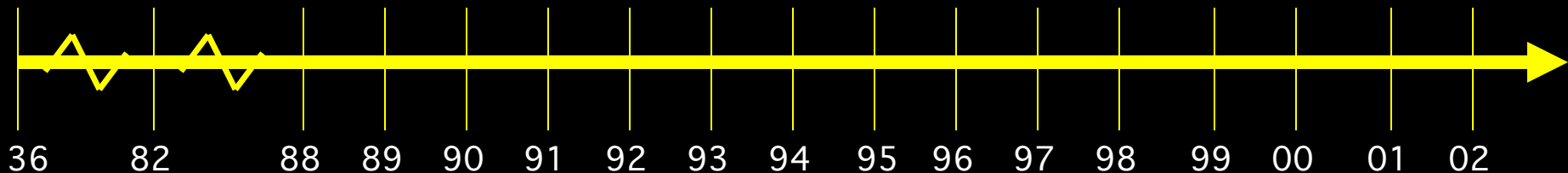
Clinical Populations

Performance prediction

Δ Volume-V1

Plasticity

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

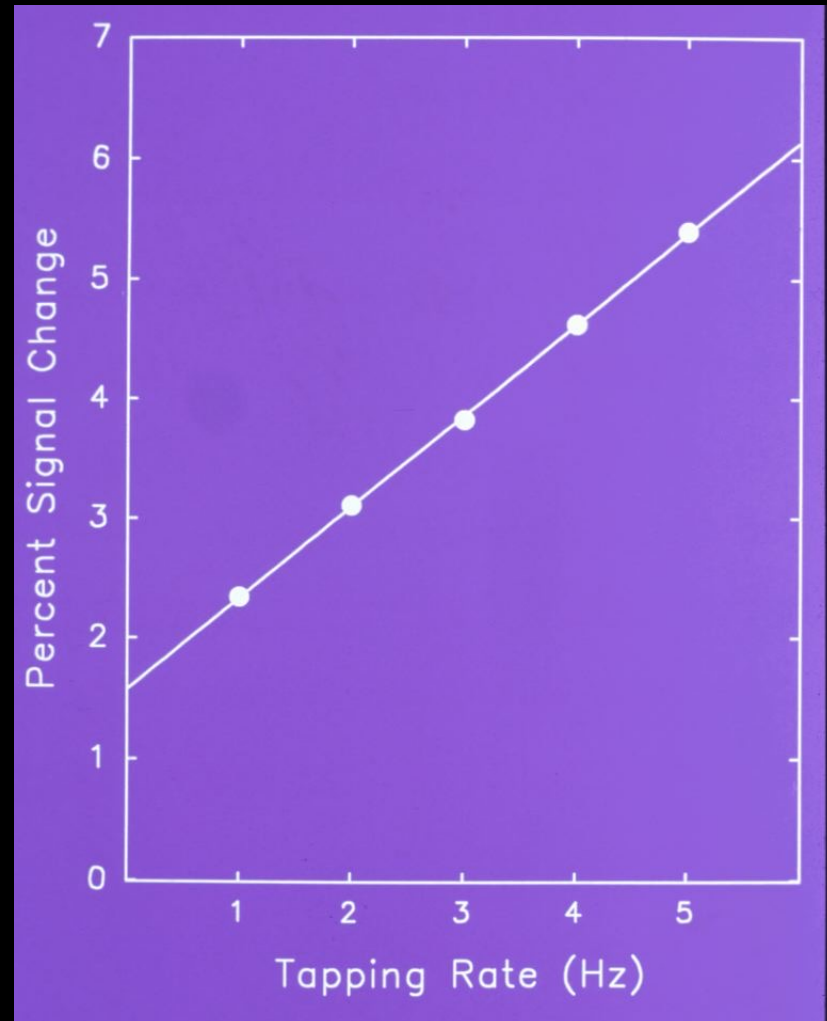
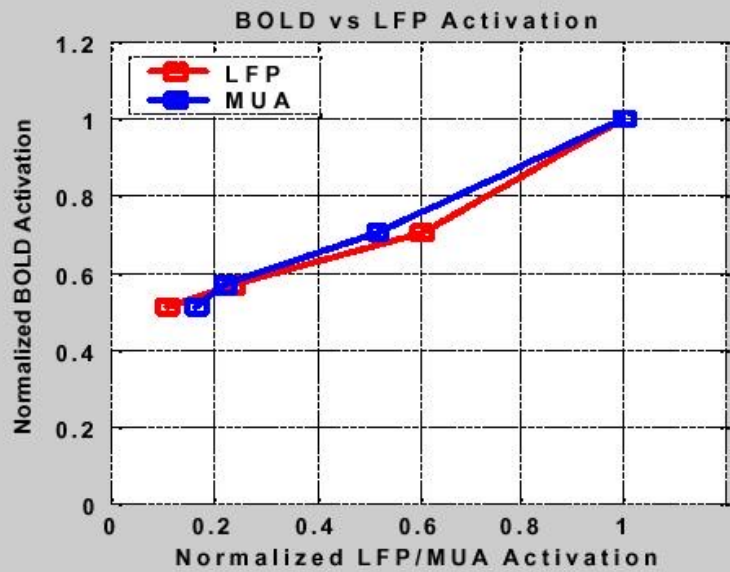
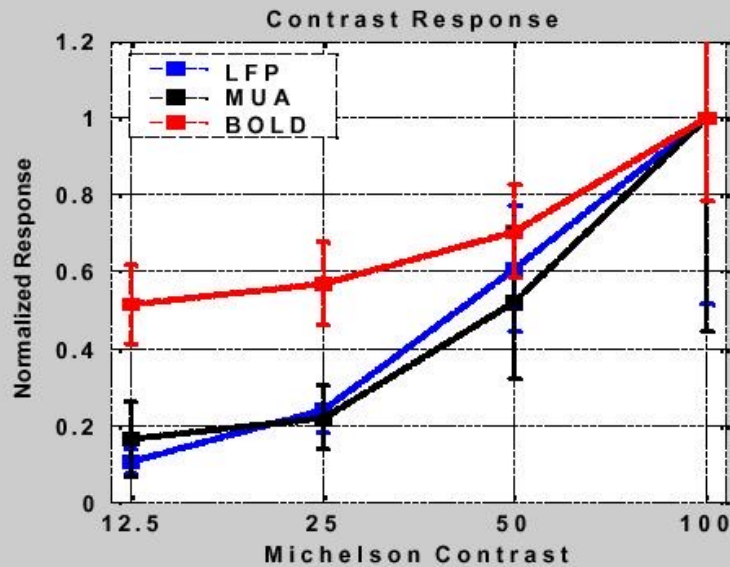
Δ Perfusion Contrast

Δ Inflow Contrast

MRI Pulse Sequence

Δ Deoxy-Hb



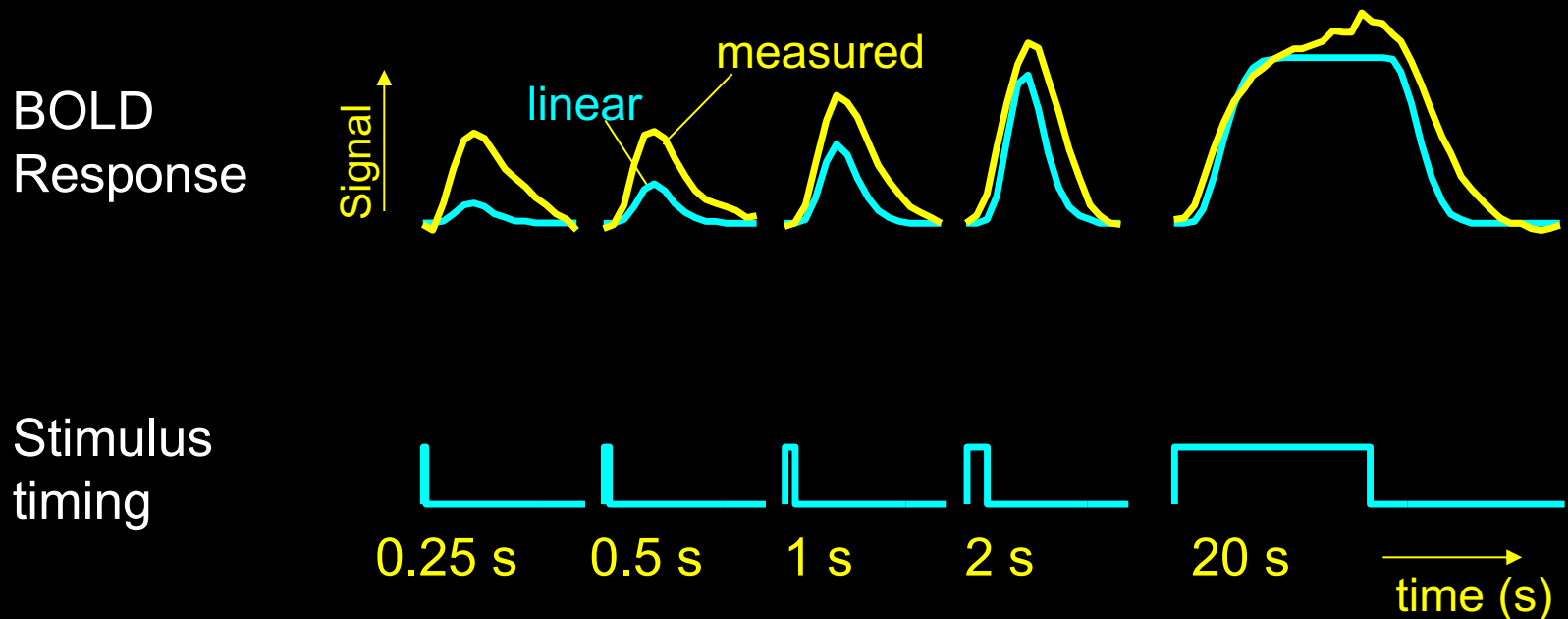


Logothetis et al. (2001) “Neurophysiological investigation of the basis of the fMRI signal” *Nature*, 412, 150-157

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.

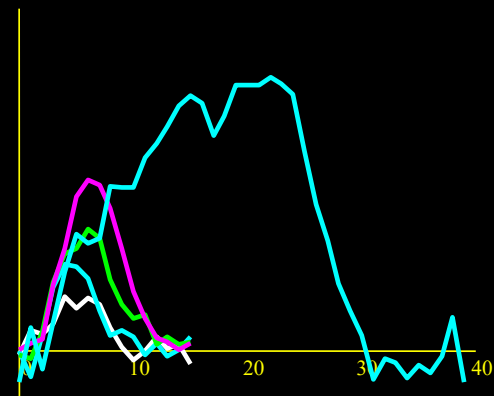
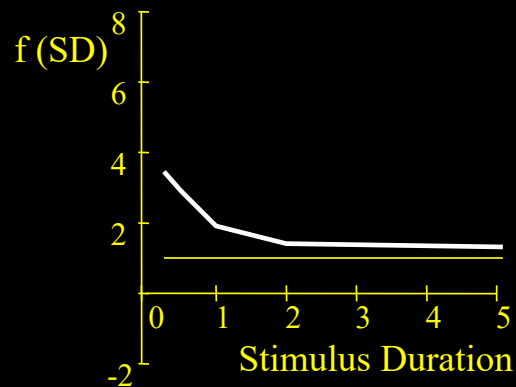
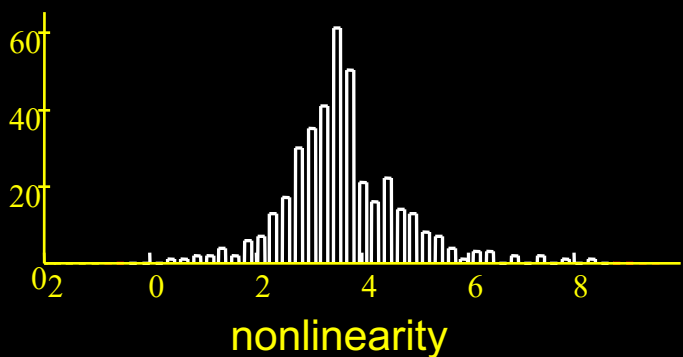
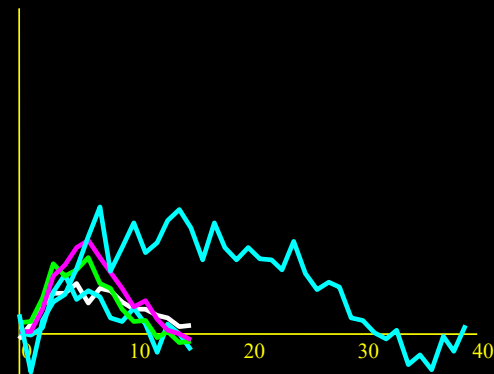
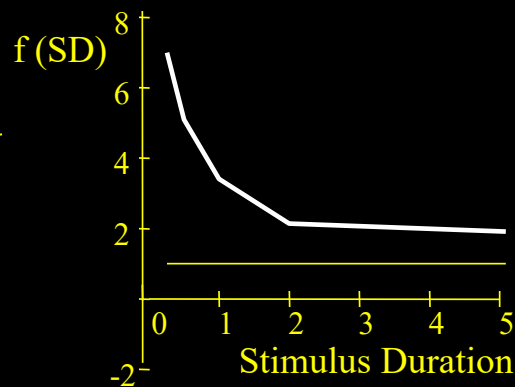
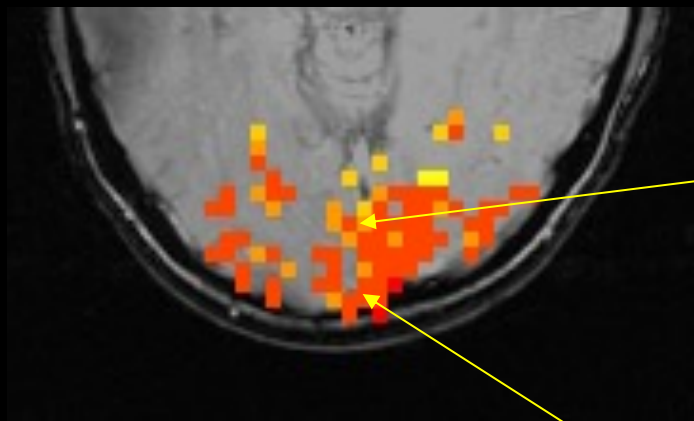
Dynamic Nonlinearity Assessment

Different stimulus “ON” periods



Brief stimuli produce larger responses than expected

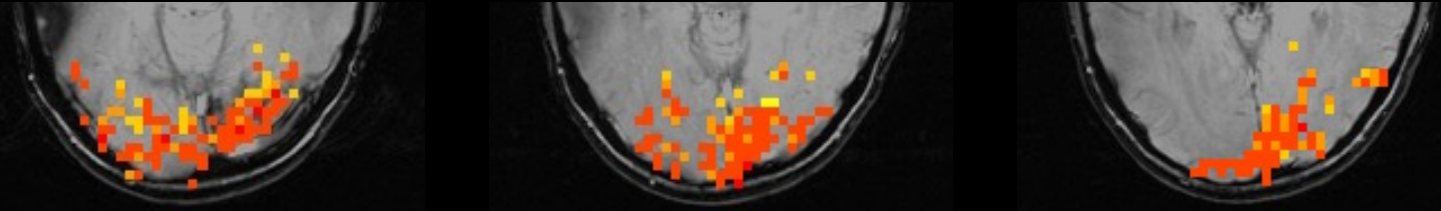
Spatial Heterogeneity of BOLD Nonlinearity



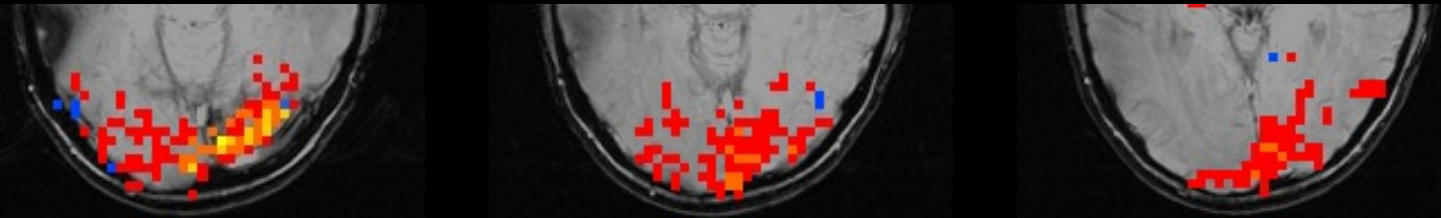
R. M. Birn, Z. Saad, P. A. Bandettini, (2001) "Spatial heterogeneity of the nonlinear dynamics in the fMRI BOLD response." *NeuroImage*, 14: 817-826.

Results – visual task

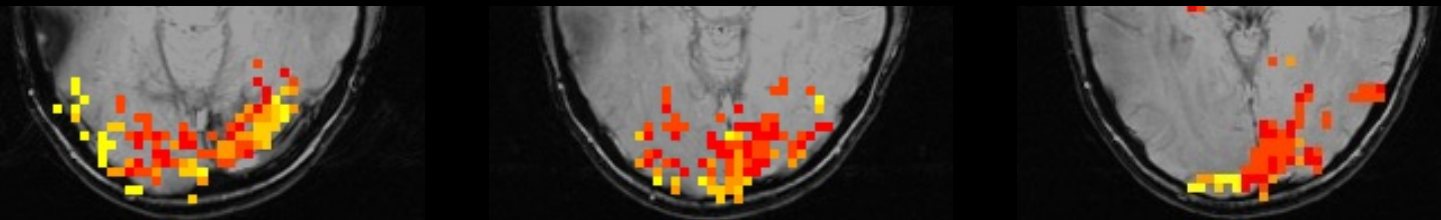
Nonlinearity



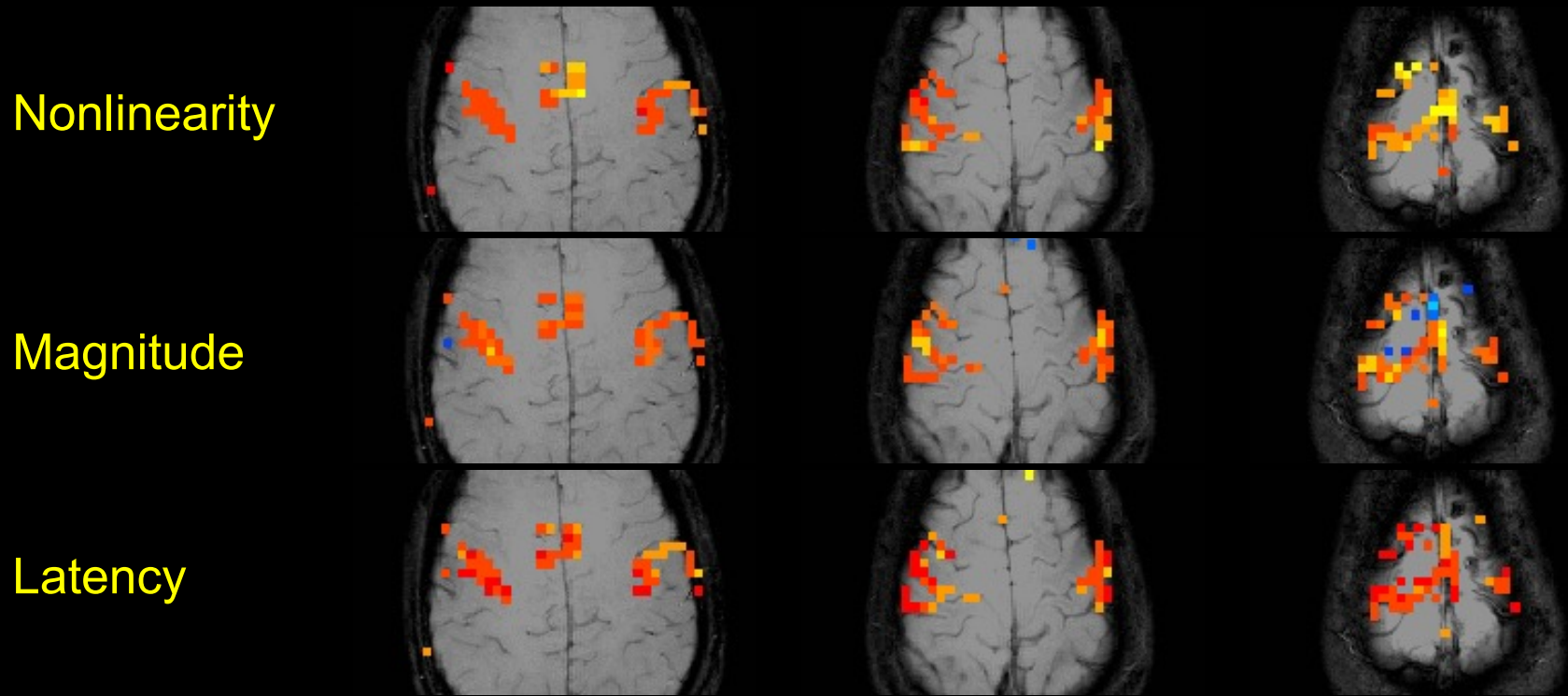
Magnitude



Latency



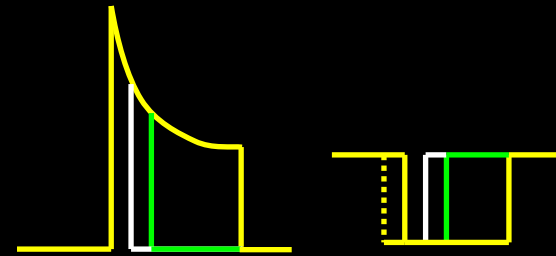
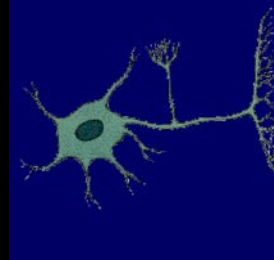
Results – motor task



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

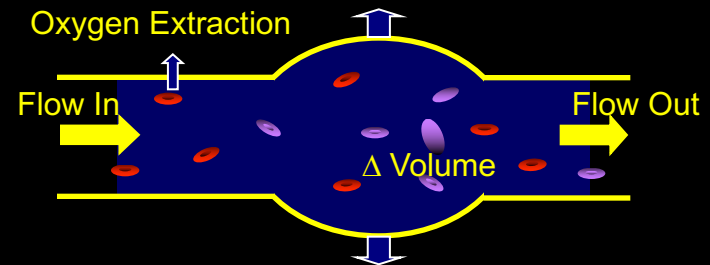
Sources of this Nonlinearity

- Neuronal



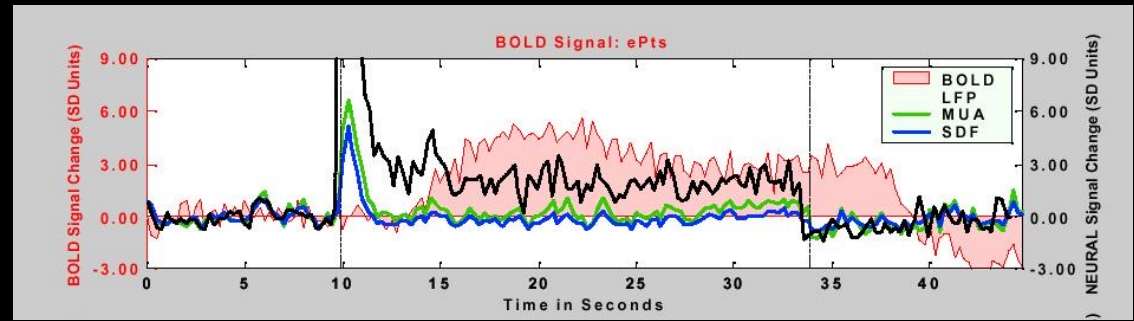
- Hemodynamic

- Oxygen extraction
- Blood volume dynamics

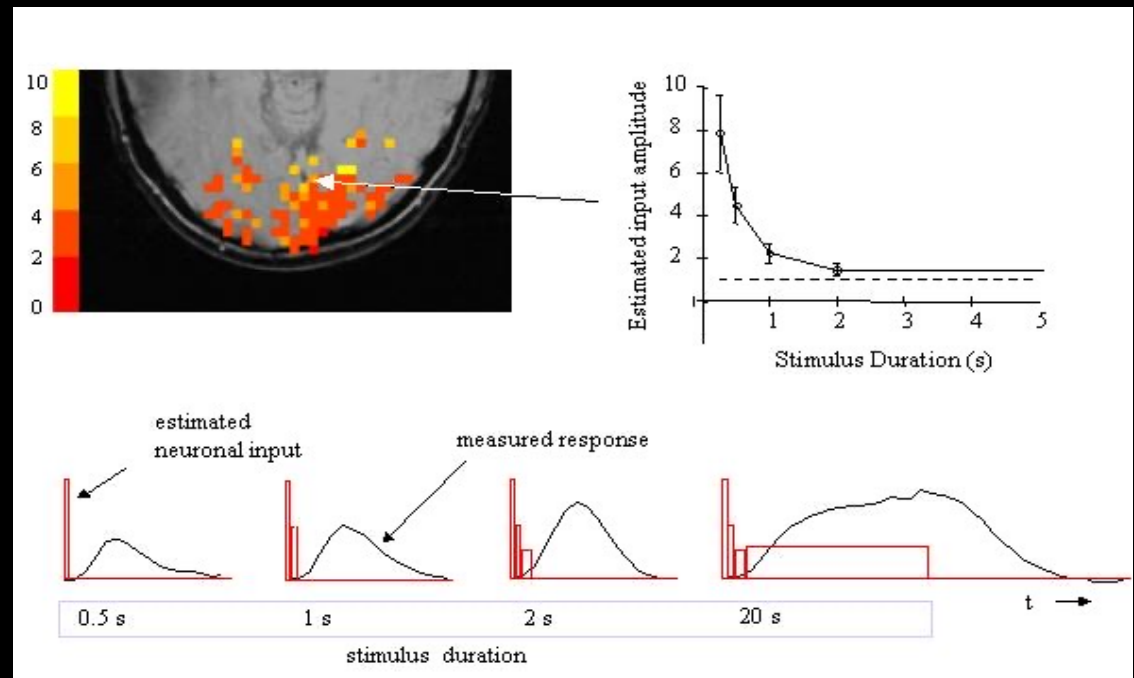


BOLD Correlation with Neuronal Activity

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

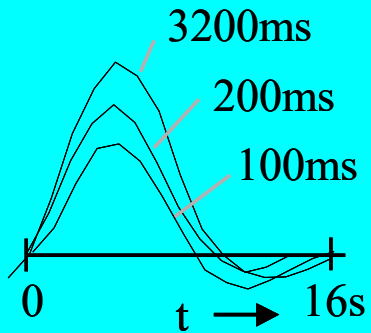


P. A. Bandettini and L. G. Ungerleider, (2001) “From neuron
to BOLD: new connections.”
Nature Neuroscience, 4: 864-866.

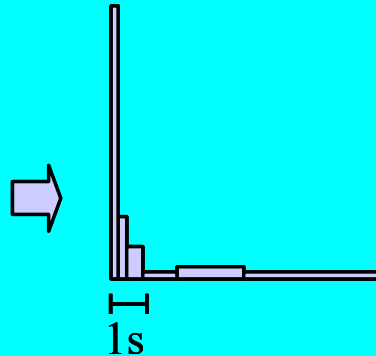


Stationary grating

BOLD response

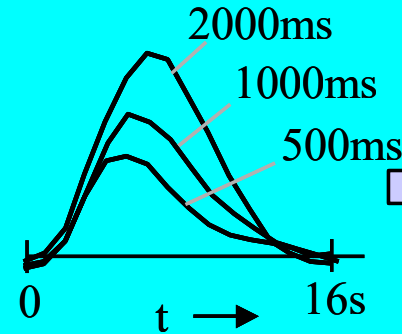


Estimated
Neuronal Input

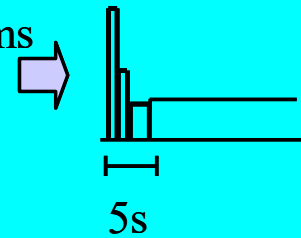


Contrast-reversing checkerboard

BOLD response



Estimated
Neuronal Input



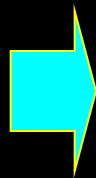
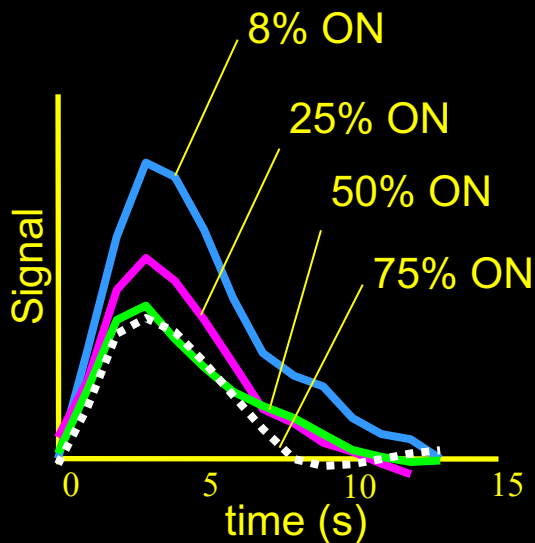
Varying “ON” and “OFF” periods

- *Rapid event-related design with varying ISI*

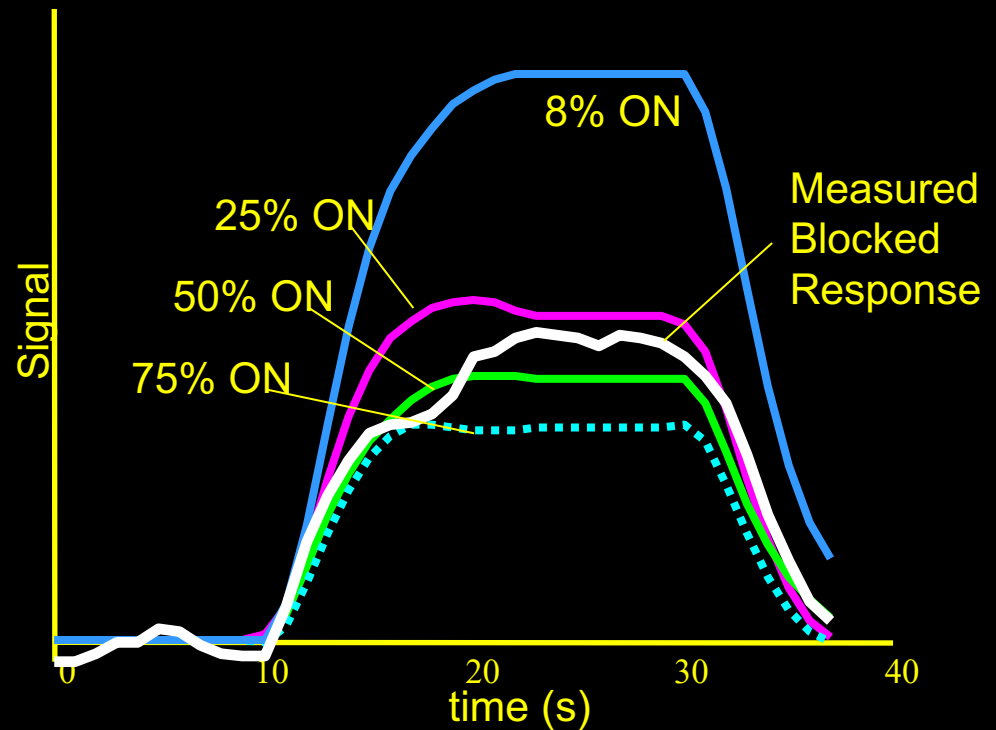


Varying “ON” and “OFF” periods

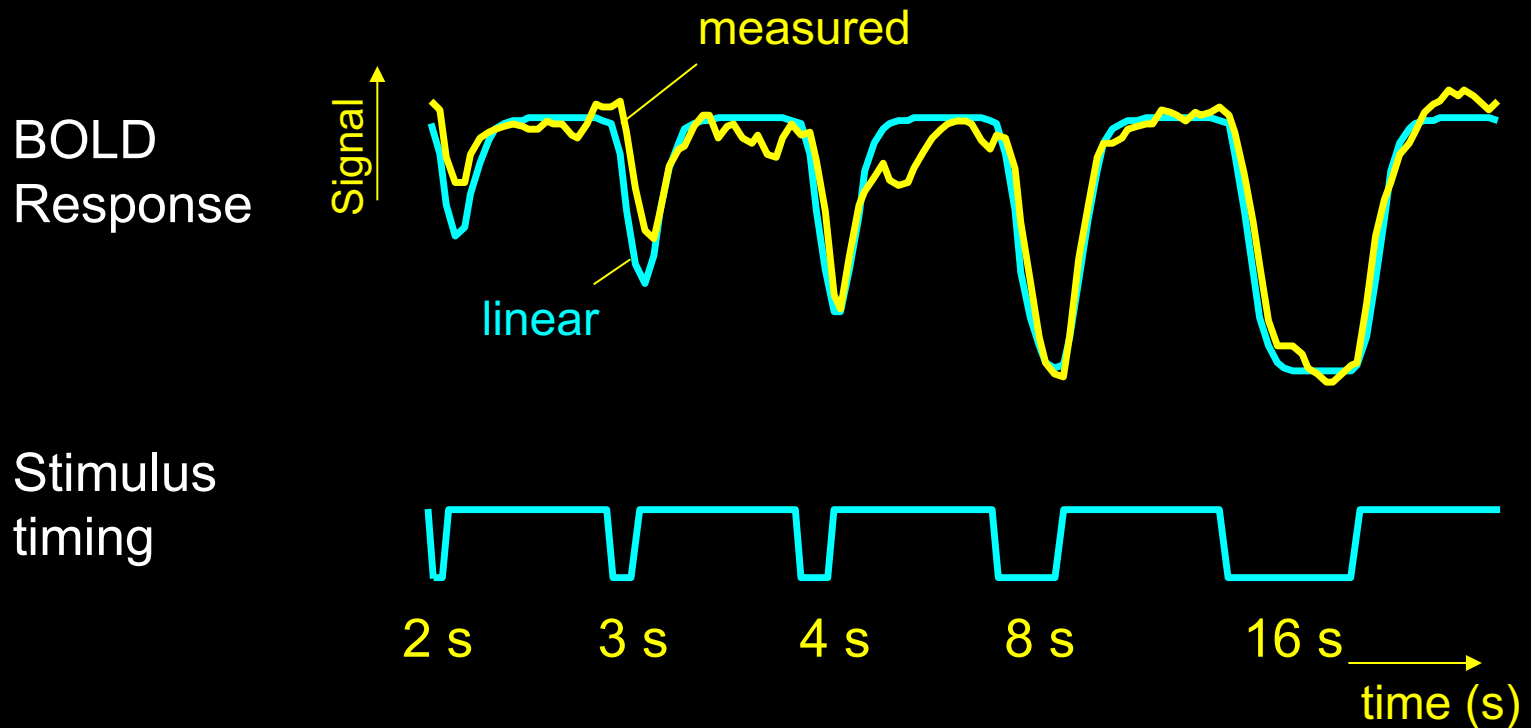
Estimated Impulse Response



Predicted Responses to 20 s stimulation



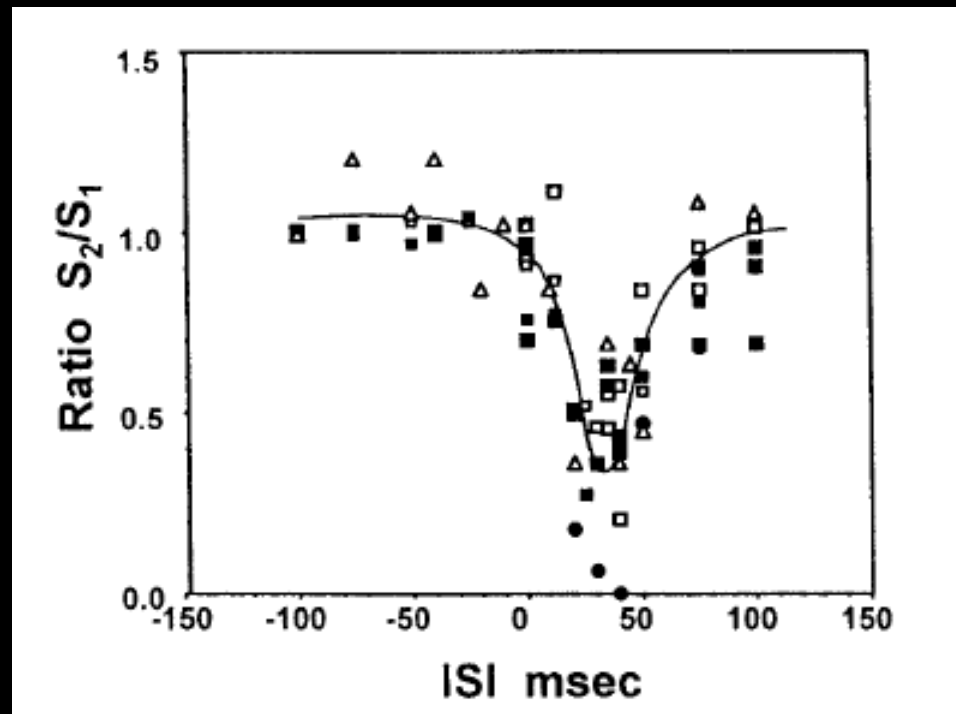
Different stimulus “OFF” periods



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[§]

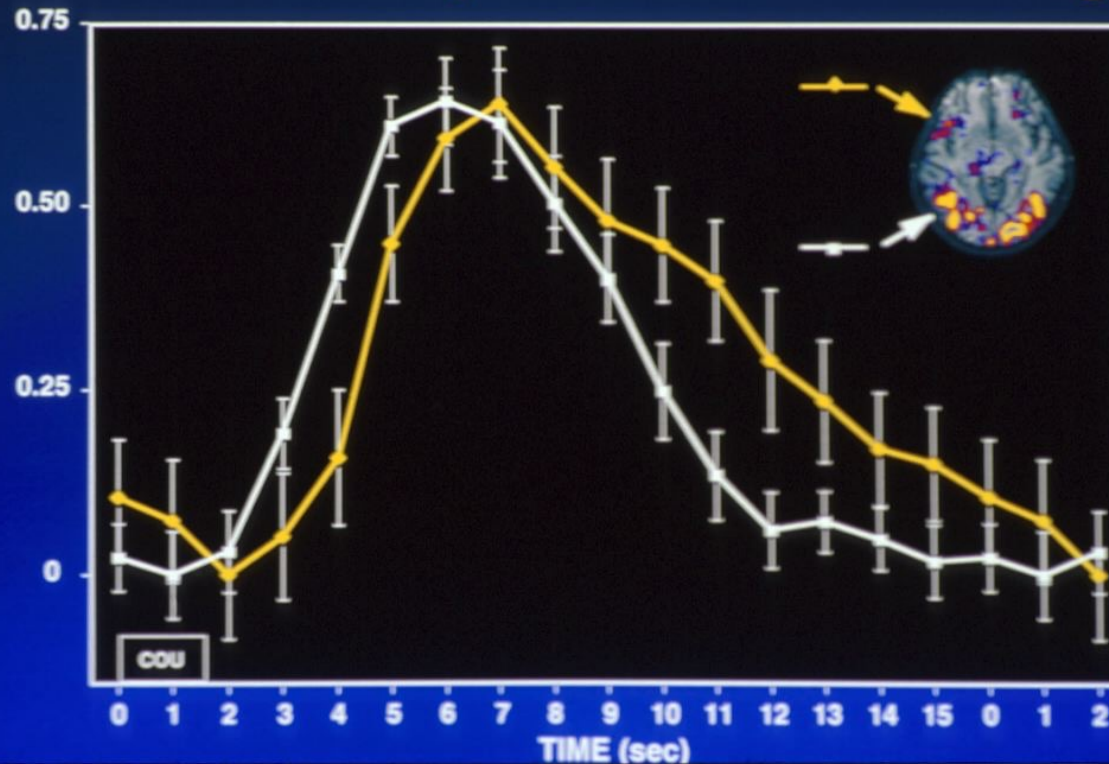


Detection of cortical activation during averaged single trials of a cognitive task using functional magnetic resonance imaging

(neuroimaging/single trial/language/prefrontal)

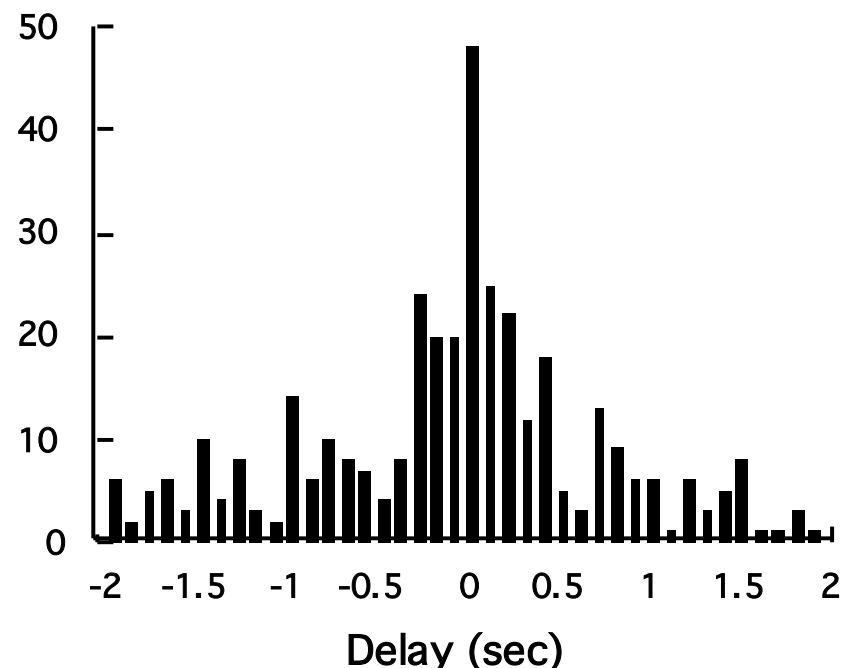
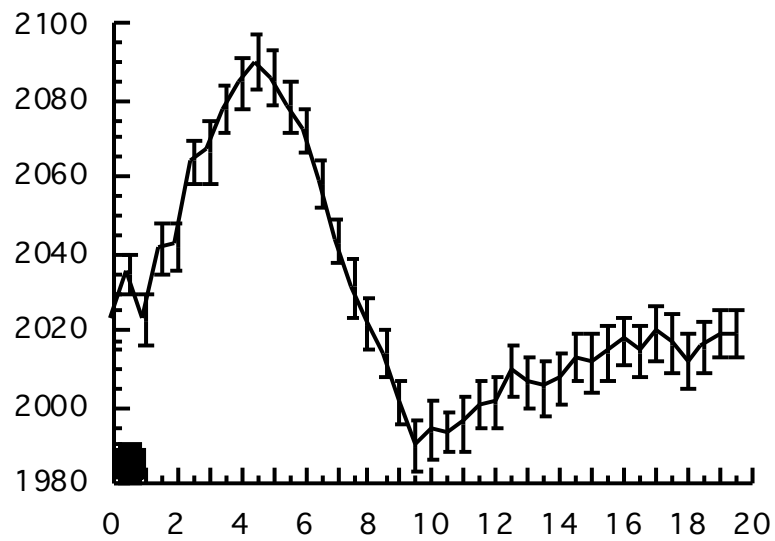
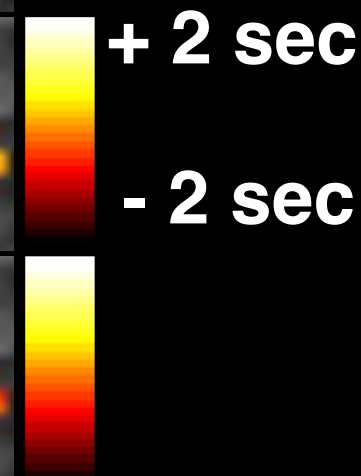
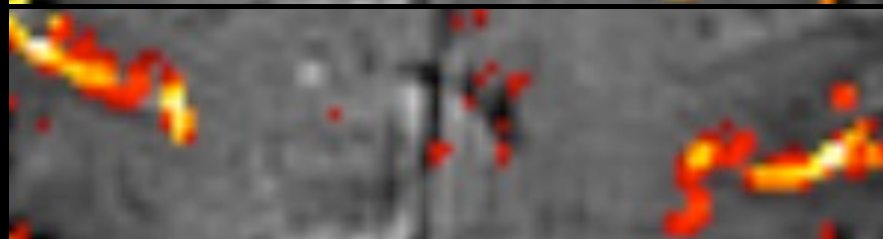
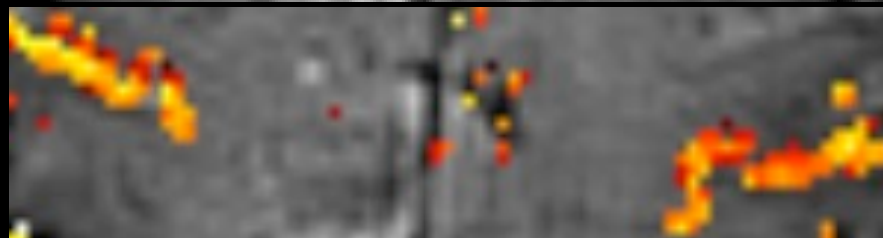
RANDY L. BUCKNER^{†‡§¶}, PETER A. BANDETTINI^{†‡}, KATHLEEN M. O' CRAVEN^{†||}, ROBERT L. SAVOY^{†||},
STEVEN E. PETERSEN^{**††}, MARCUS E. RAICHEL^{§**††}, AND BRUCE R. ROSEN^{†‡}

Time Course Comparison Across Brain Regions

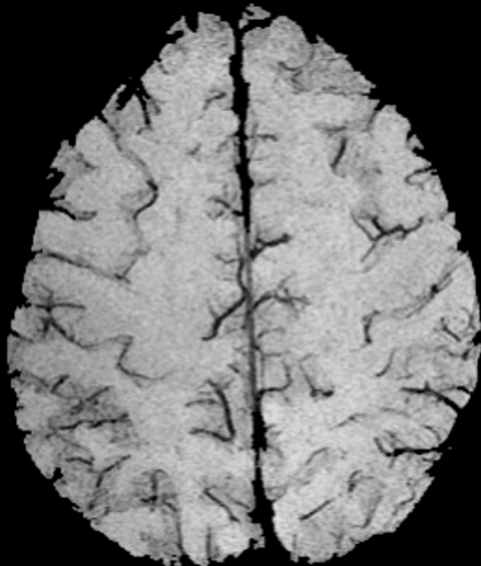
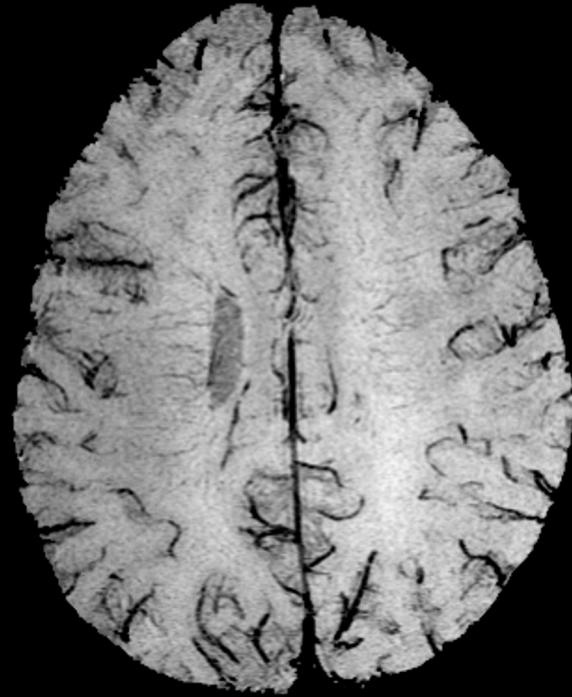


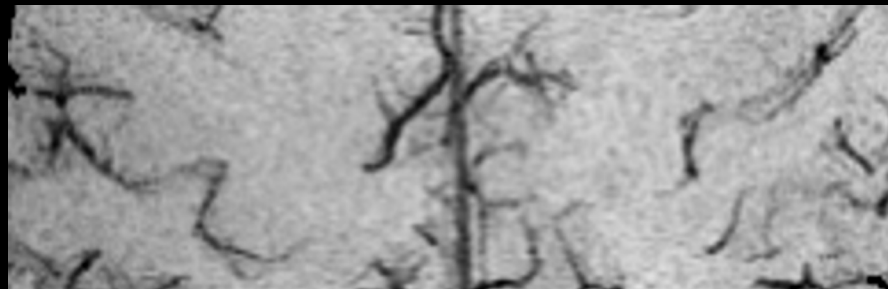
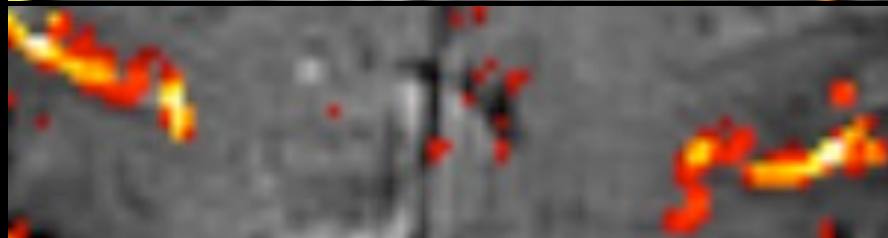
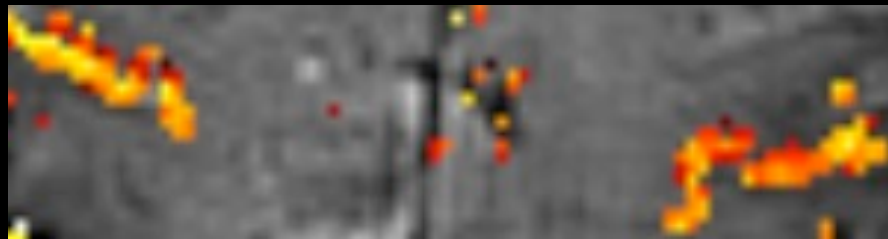
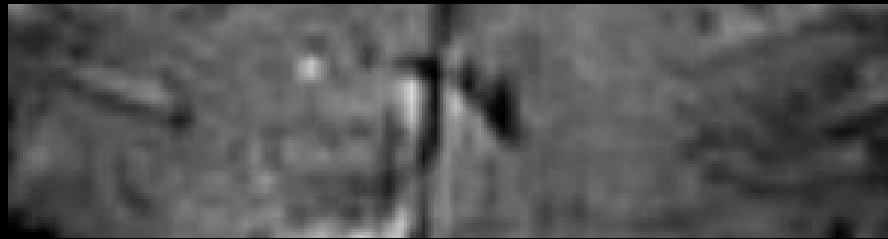
Latency

Magnitude



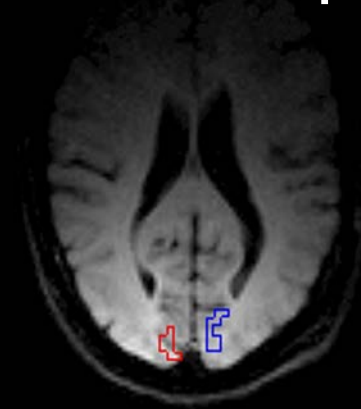
Venograms (3T)



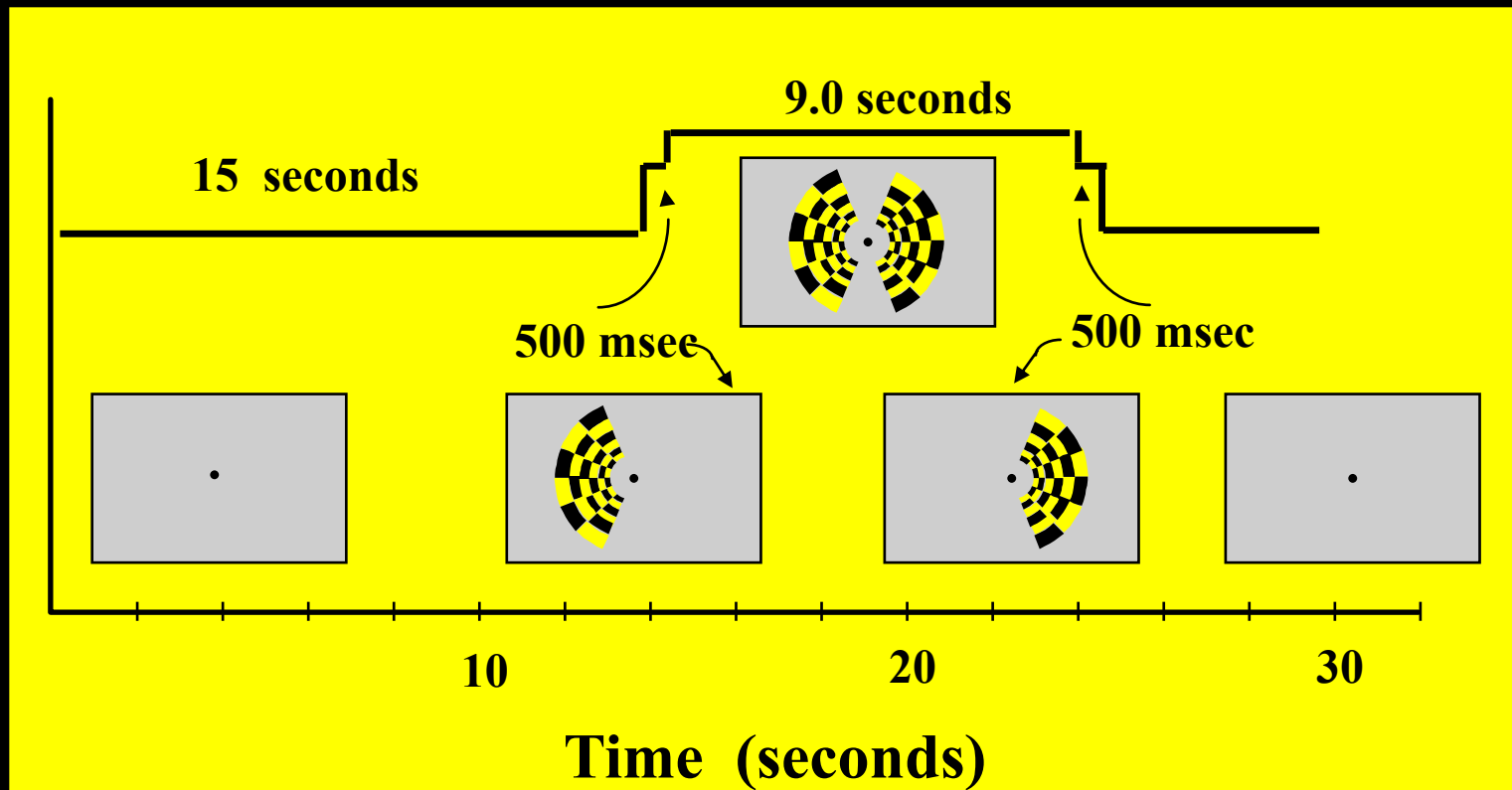


Hemi-Field Experiment

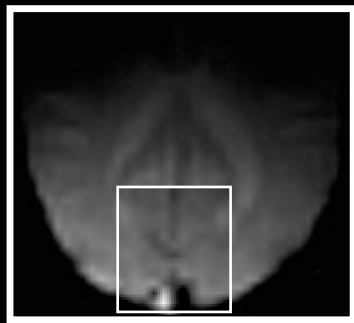
Left Hemisphere



Right Hemisphere

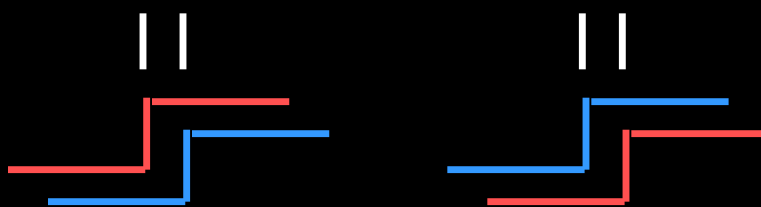


Calibration Techniques.....



500 ms

500 ms



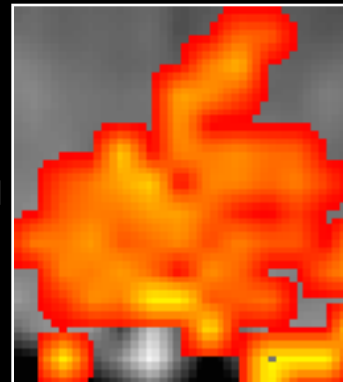
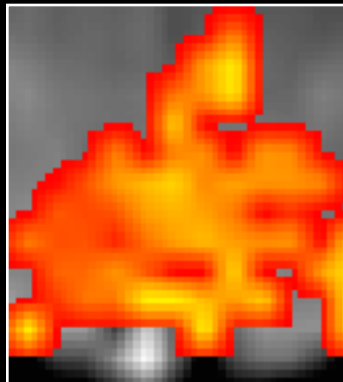
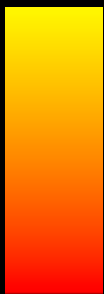
Right Hemifield

Left Hemifield

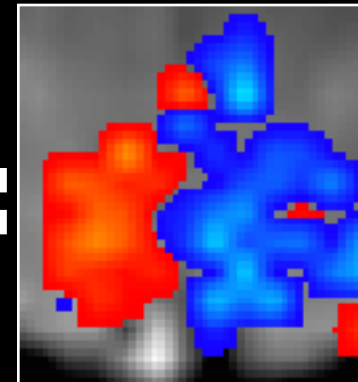
+ 2.5 s

0 s

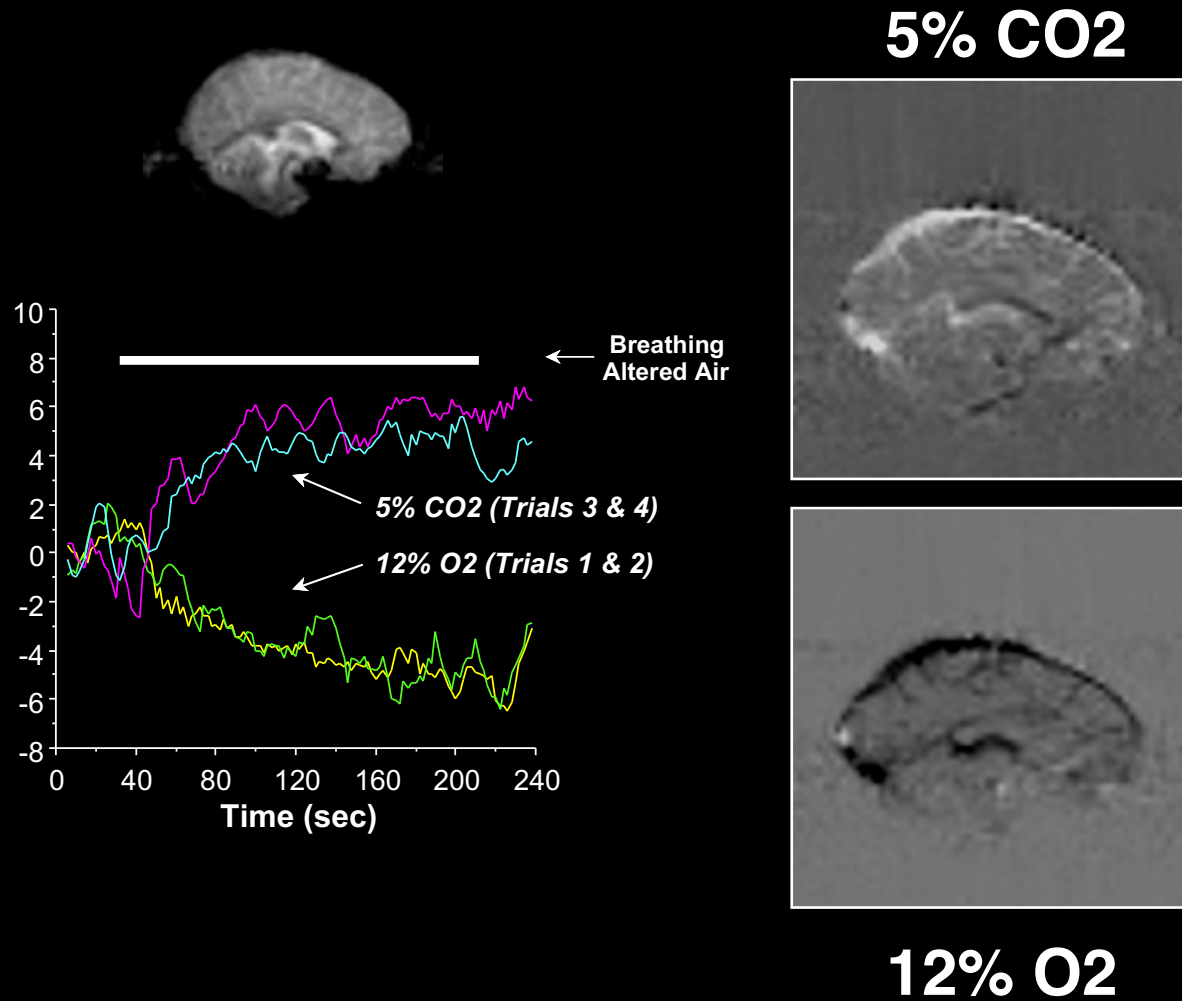
- 2.5 s



=



Hemodynamic Stress Calibration



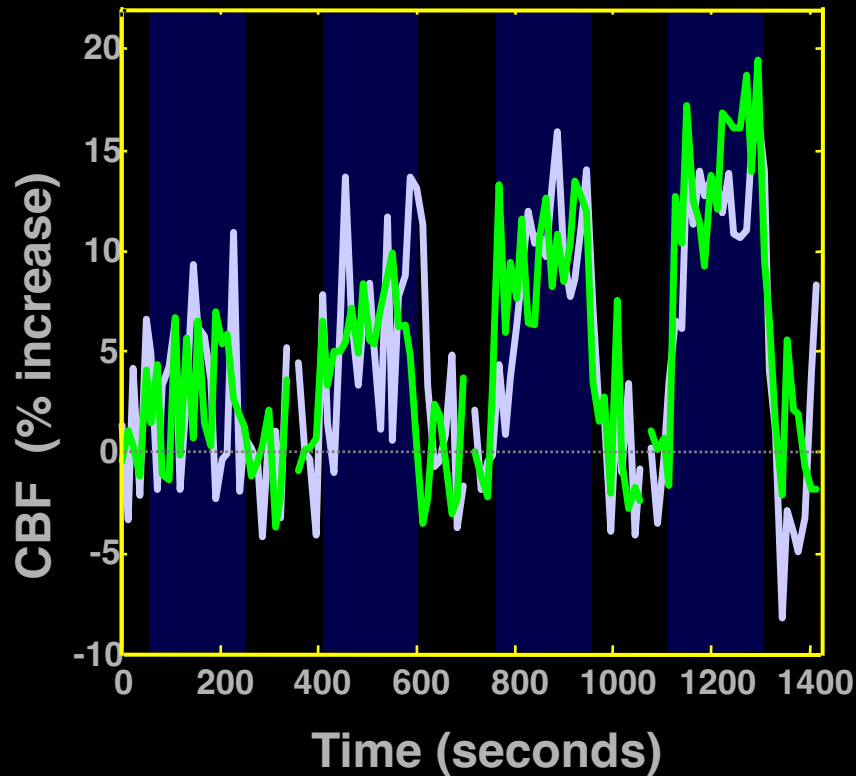
P. A. Bandettini, E. C. Wong, A hypercapnia - based normalization method for improved spatial localization of human brain activation with fMRI. *NMR in Biomedicine* 10, 197-203 (1997).

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

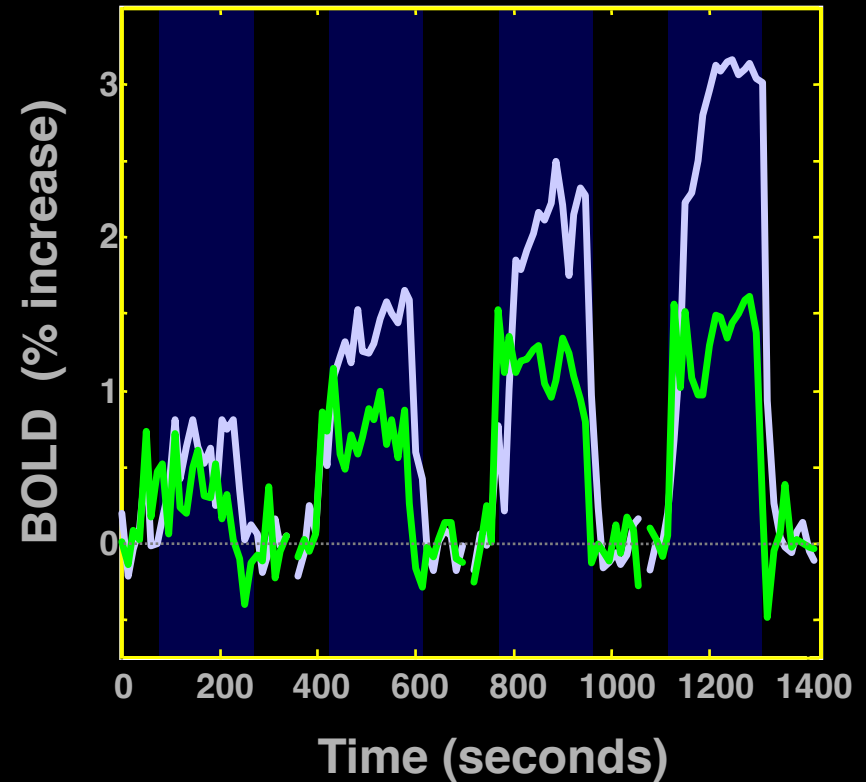
RICHARD D. HOGE^{*†}, JEFF ATKINSON^{*}, BRAD GILL^{*}, GÉRARD R. CRELIER^{*}, SEAN MARRETT[‡], AND G. BRUCE PIKE^{*}

^{*}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

CBF



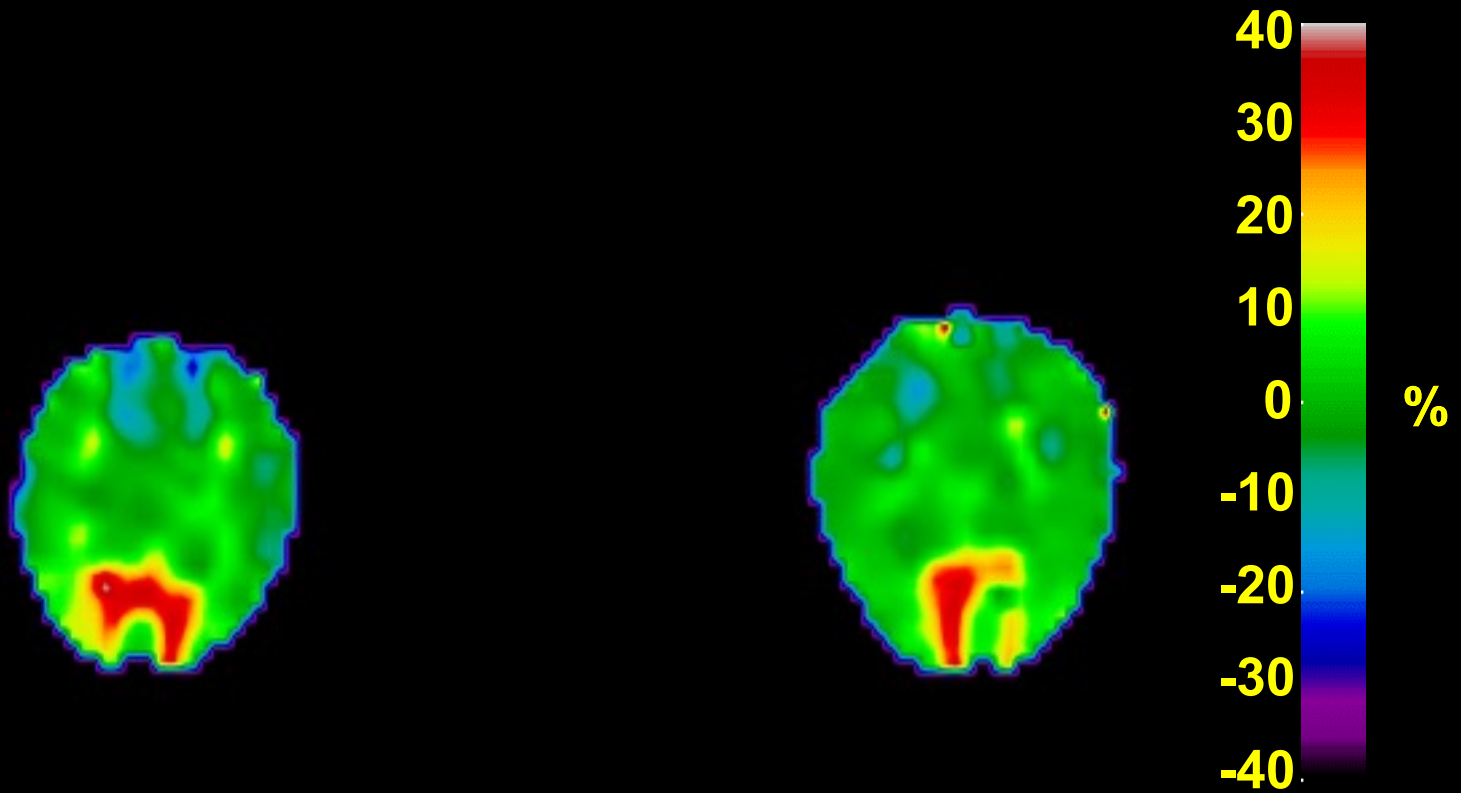
BOLD



Simultaneous Perfusion and BOLD imaging during
graded visual activation and hypercapnia

N=12

Computed CMRO₂ Changes

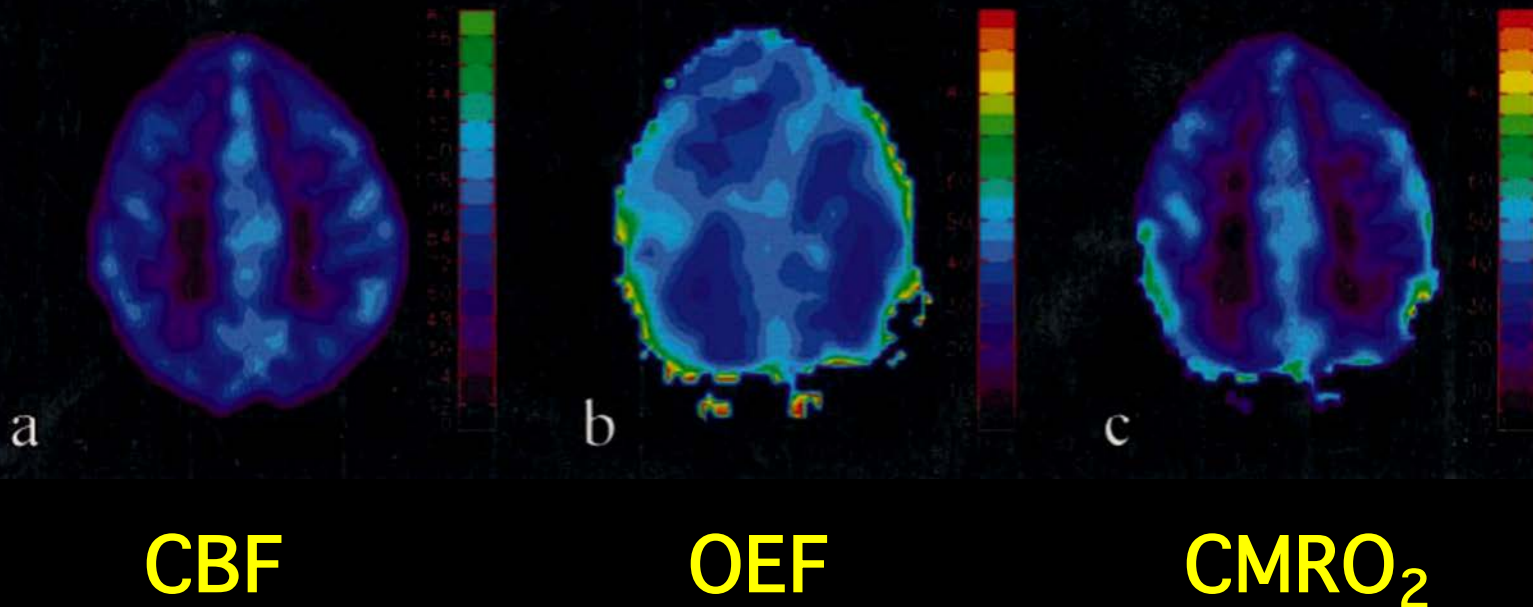


Subject 1

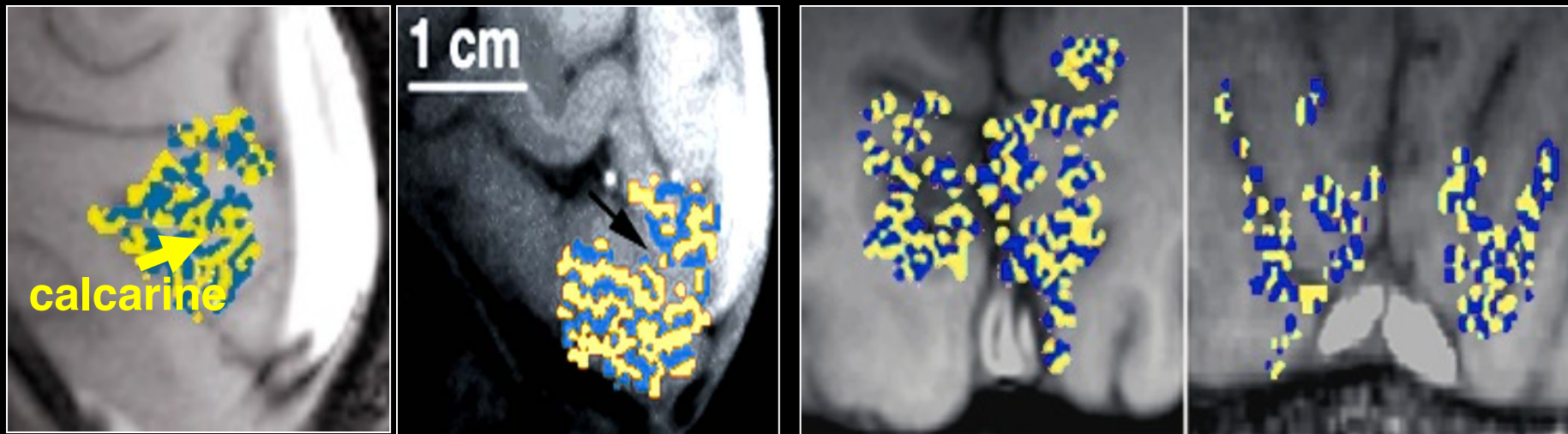
Subject 2

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

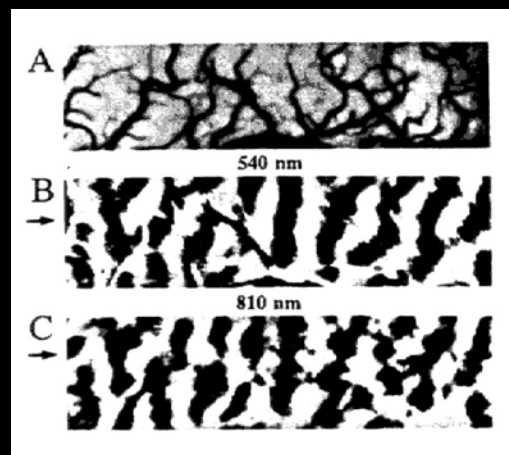
Hongyu An,¹ Weili Lin,^{2*} Azim Celik³ and Yueh Z. Lee²



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.

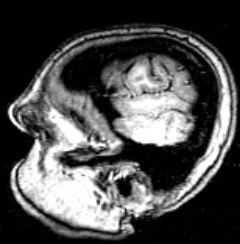
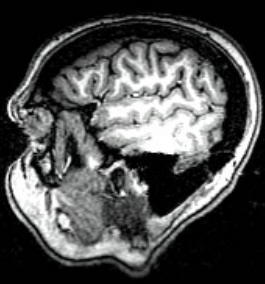
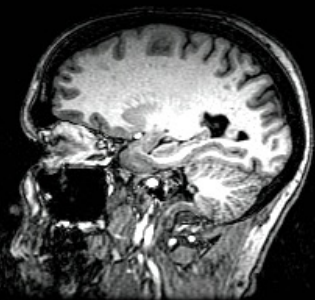
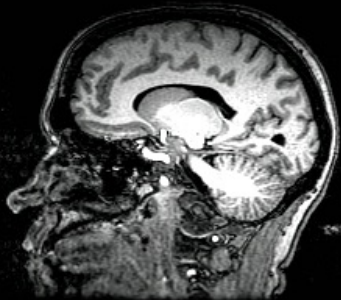
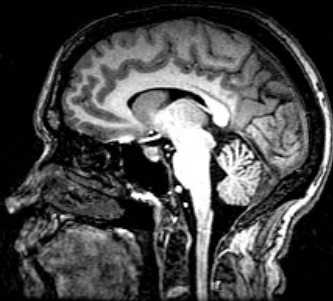
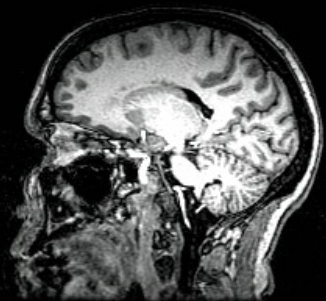


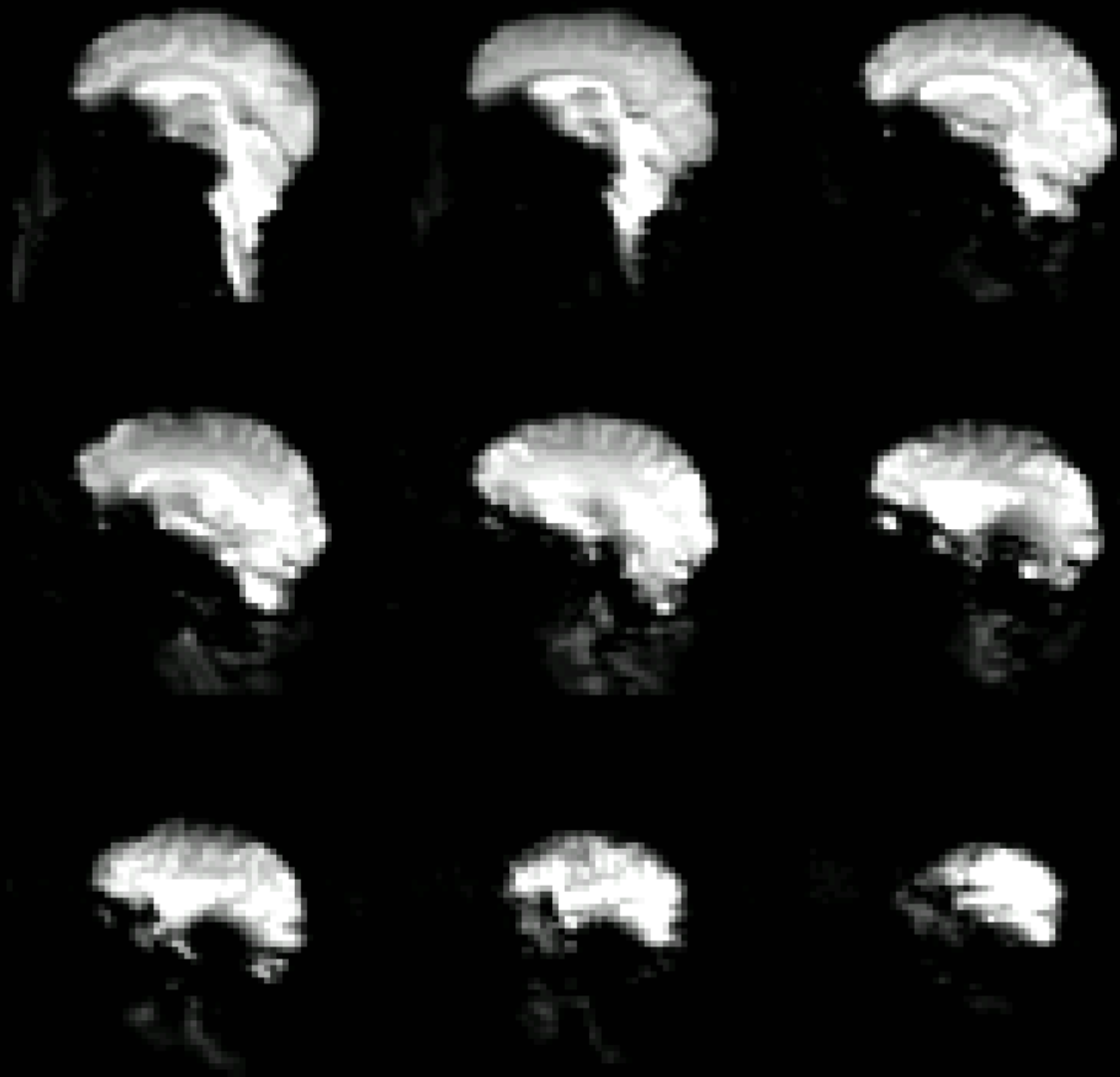
Optical Imaging

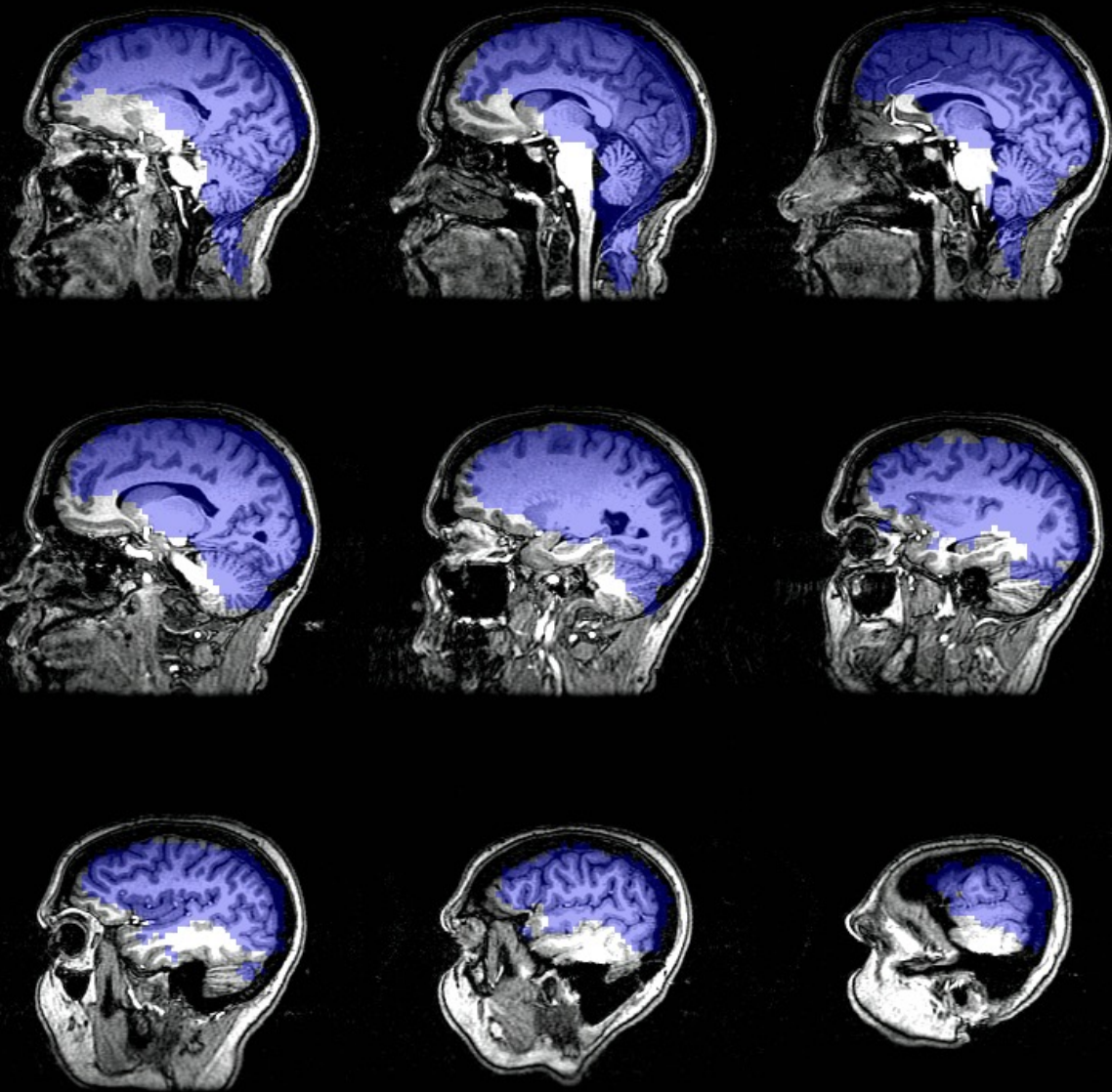
R. D. Frostig et. al, PNAS 87: 6082-6086, (1990).

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

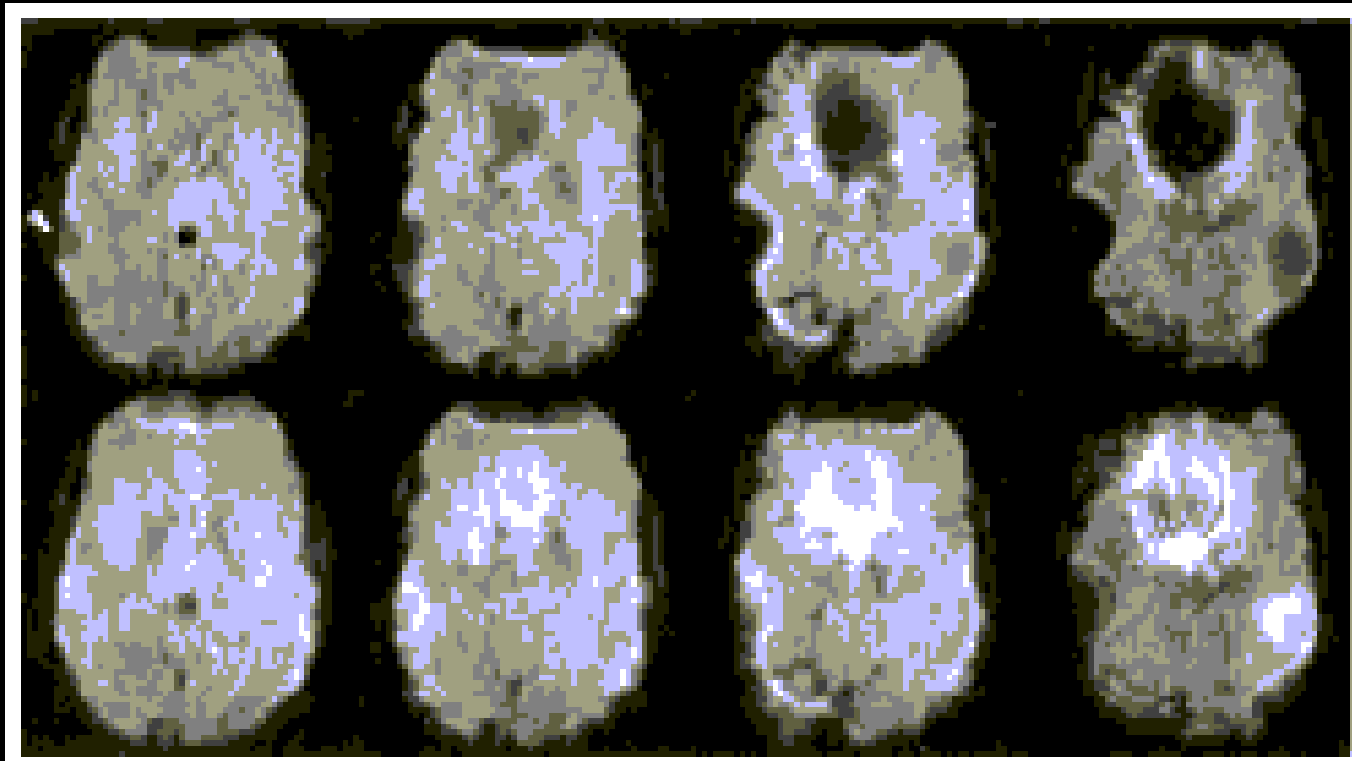






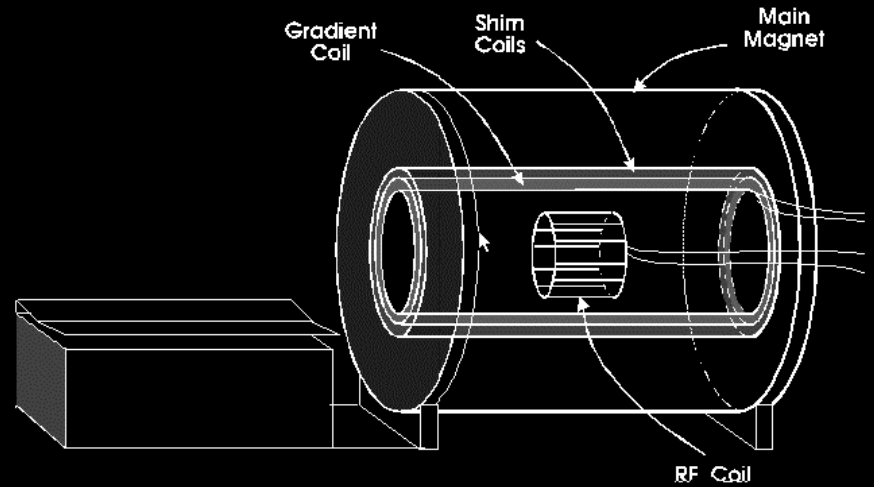
3D z-Shim Method for Reduction of Susceptibility Effects in BOLD fMRI

Gary H. Glover*



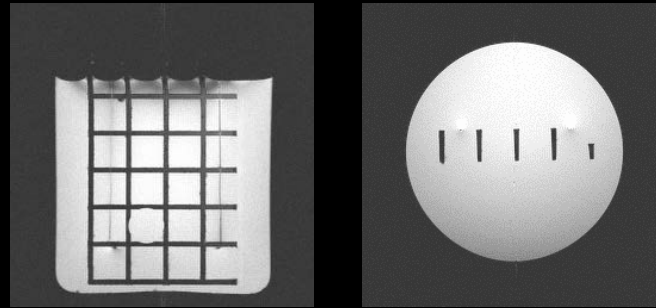
2 G/cm, 350 T/m/s

4 G/cm, 150 T/m/s



10 G/cm, 1000 T/m/s

Diffusion imaging
Faster imaging
Higher resolution



Neuronal Current Imaging

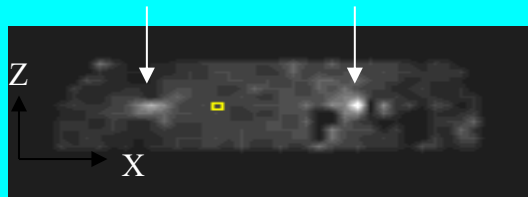
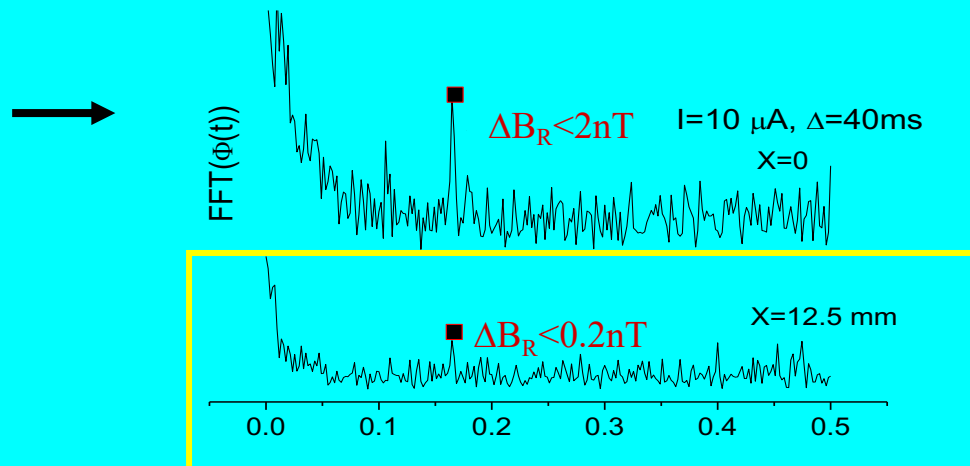


Figure 1



J. Bodurka, P. A. Bandettini. Toward direct mapping of neuronal activity: MRI detection of ultra weak transient magnetic field changes, Magn. Reson. Med. .

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 Maieron

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