EEG-fMRI & Language

Peter J. Molfese, Ph.D.
Section on Functional Imaging Methods, NIMH
July 28, 2017



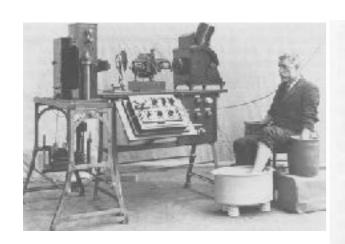


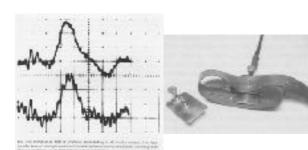
Purpose of this Talk

- Give an overview of EEG/ERP Techniques
 - Particularly some of the uses
- Admit fully that not a lot of studies have been done on simultaneous EEG-fMRI on Language
- Illustrate converging evidence from fMRI and EEG/ERP of language (reading)
- Highlight the importance of simultaneous studies moving forward



Historical Moment





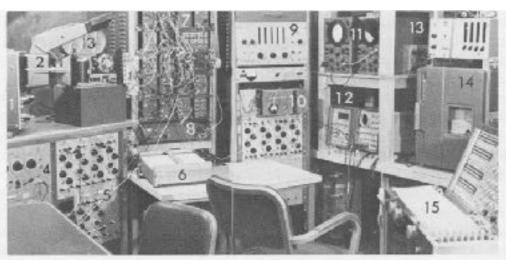


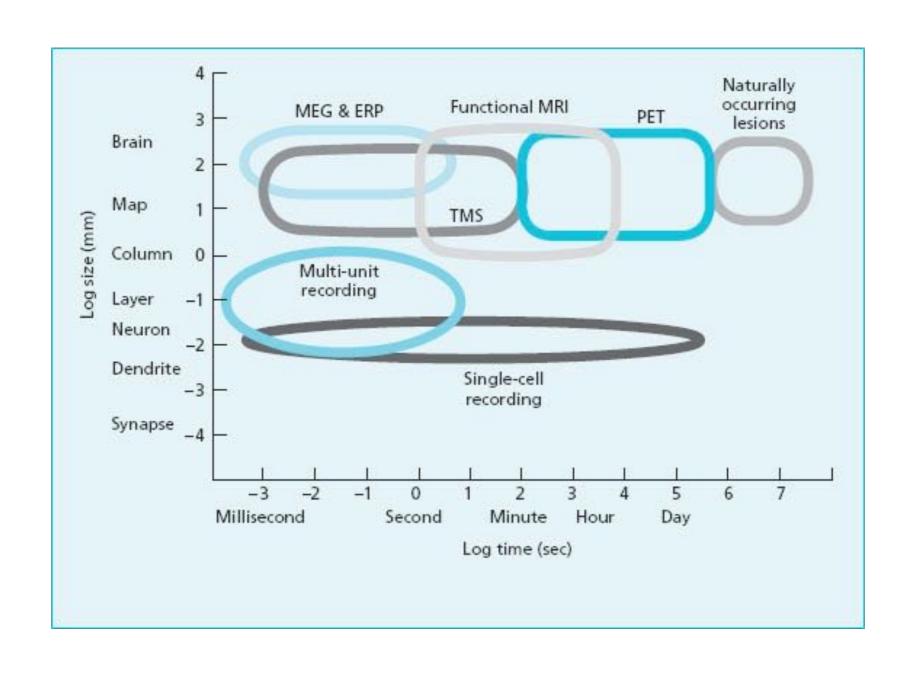
Fig. 1-21. A modern EEG and perceptual laboratory, equipped for EEG, average evoked potential, and CNV recording. 1. Tektronix 502 oscilloscope; 2. Tektronix Polaroid camera; 3. Grass Kymograph camera; 4. Grass physiological stimulators; 5. Tektronix power supply, waveform and pulse generators; 6. Moseley X-Y plotter; 7. Massey-Dickinson control and programming equipment; 8. Power supply; 9. Hewlett-Packard counter; 10. Tapereader; 11. Oscilloscope monitors; 12. Mnemotron computer of average transient (CAT) and accessories; 13. Preset controller and tape coder; 14. Ampex FR-1300 7-channel FM tape recorder; 15. Grass Model 6.8-channel EEG.



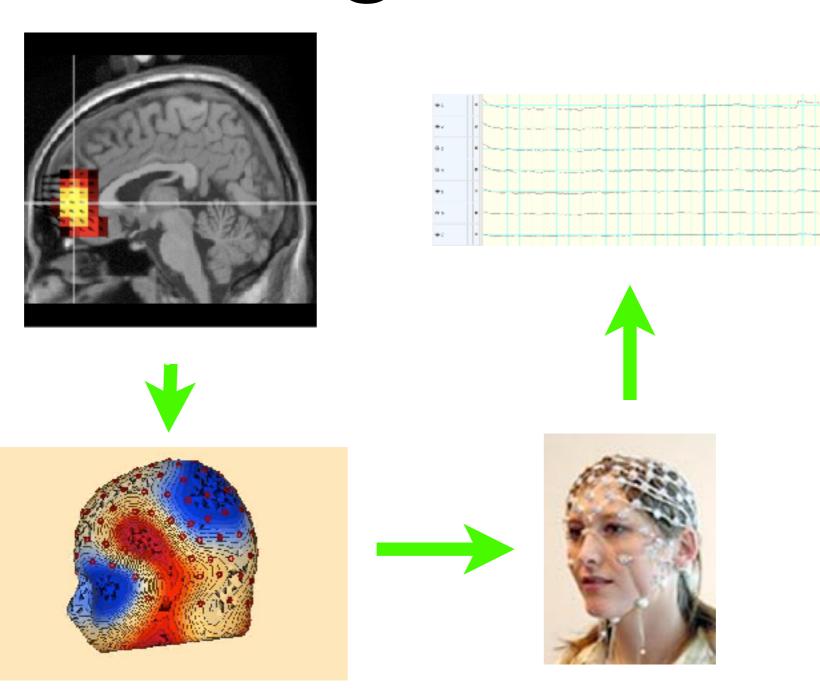
Hans Berger, 1924



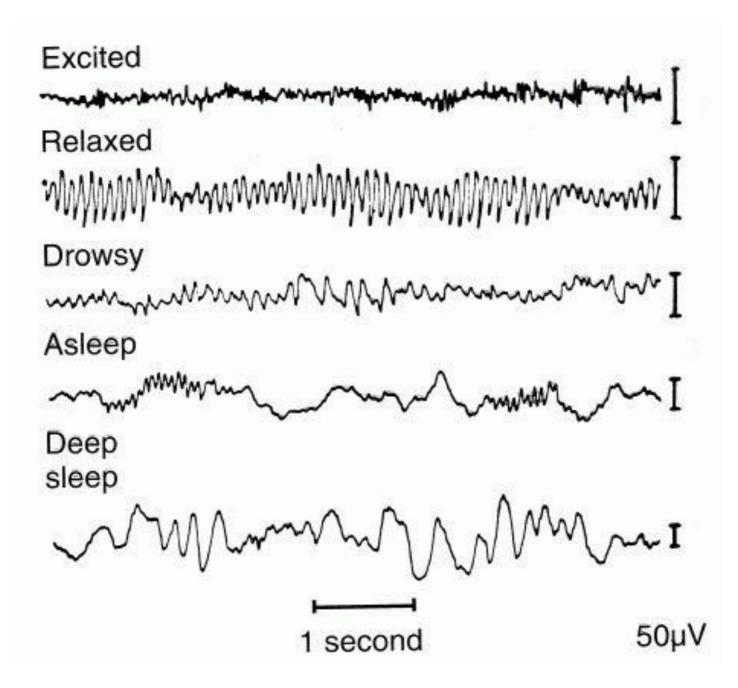
Temporal-Spatial Tradeoffs



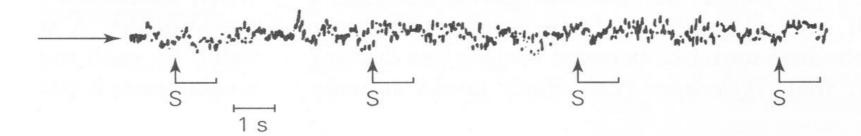
Neural Origins of EEG



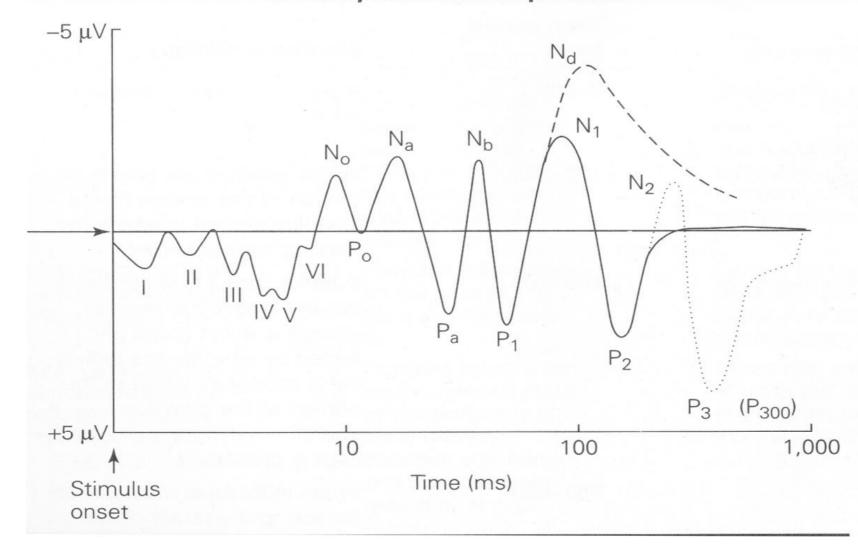
EEG Reflects Brain State



Ongoing EEG



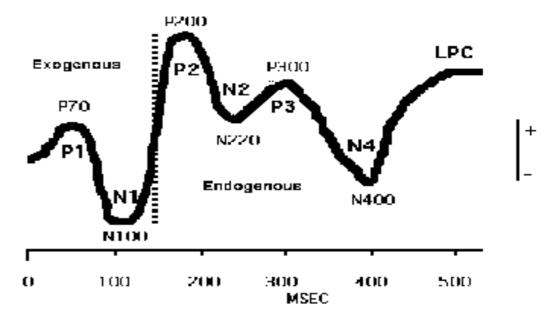
Auditory event-related potential

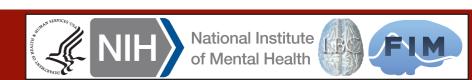


The Evoked Potential

- Event-Related Potential
- Portion of the ongoing EEG
- Time-Locked to Stimulus Onset
- Strong Temporal Information
- Comparability across the lifespan

Event Related Potential (ERP)

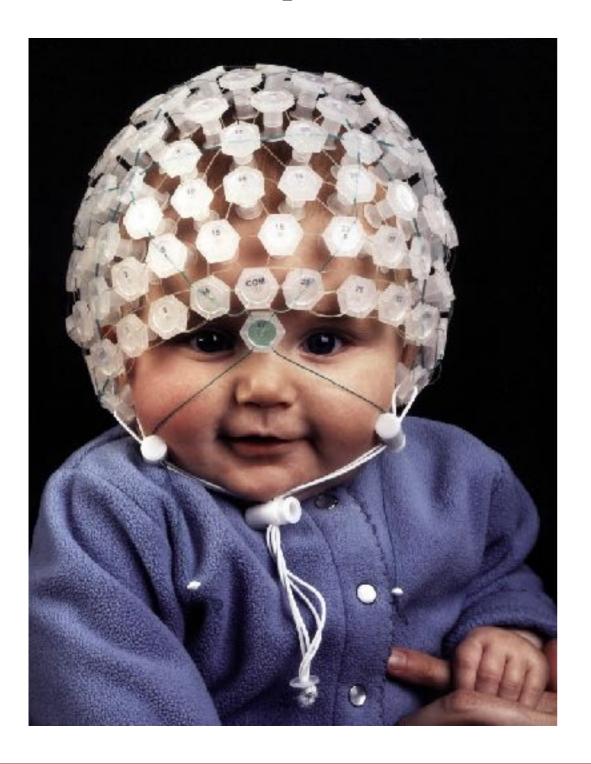


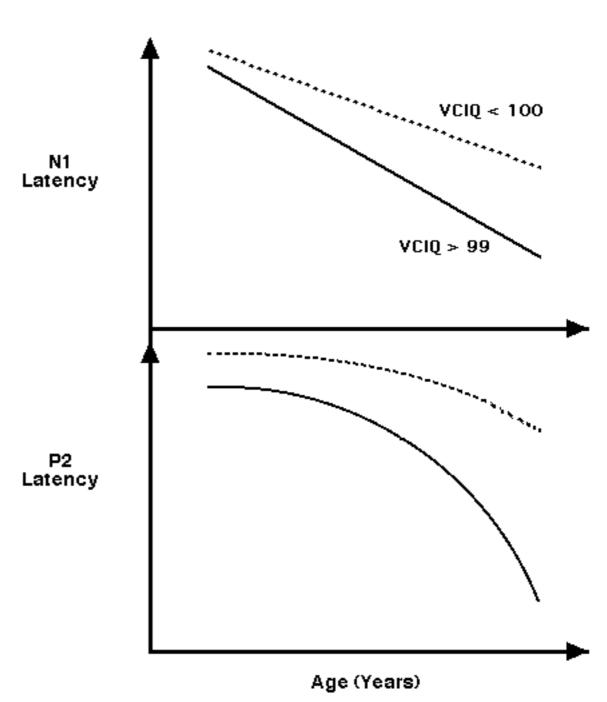


ERP Definition

 Sequence of overlapping components, each perhaps representing activity of different populations of nerve cells and each sometimes standing in different, perhaps orthogonal, relations to experimental variables (Donchin, 1979, p24).

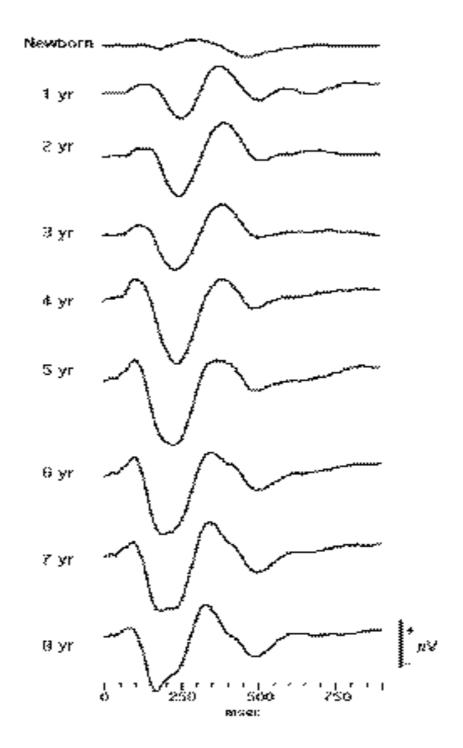
Comparable Across Lifespan







Developmental Changes

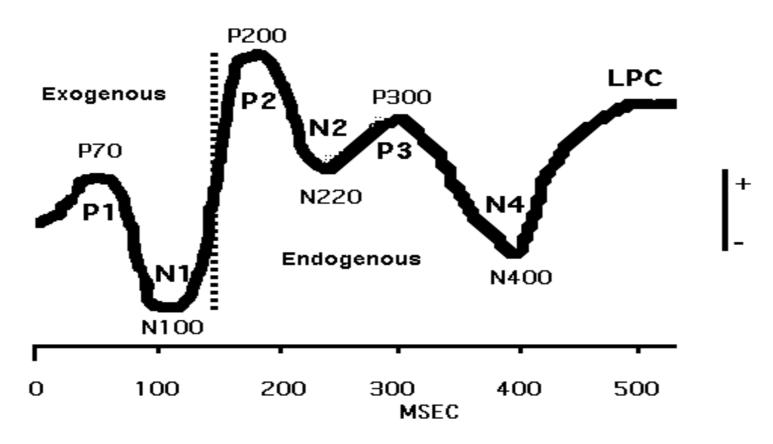


Infant ERP Testing

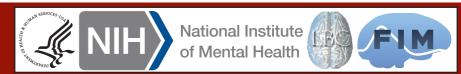


ERP Nomenclature

Event Related Potential (ERP)



Exogenous = influenced by external factors Endogenous = internal factors controlling



Peaks	Latency (ms)	Experiment Manipulation	Max Scalp Amplitude	Interpretation	Source
P1	50 Auditory	None Specific	Anterior	arousal level, suppression of unattended information	primary auditory cortex, superior temporal gyres, medial frontal
	100 Visual		Occipital		striate or extra-striate (posterior fusiform), posterior- parietal regions
N1	100 Auditory	None Specific	Temporal	selective filtering, basic stimulus characteristics, initial selection for later pattern recognition	Primary auditory cortex, superior temporal plane
	100-161 Visual		Central, Midline, Occipital		inferior occipital, occipito- temporal junction, inferior temporal lobe
P2	150-275 Auditory	None Specific	Central	selective attention, stimulus change, feature detection, short-term memory	primary auditory cortex, secondary auditory cortex
	200+ Visual		Occipital, Frontal		inferior occipital region



Peaks	Latency	Experimental Manipulation	Max Amplitude	Interpretation	Source
N2	200 ms Auditory	None Specific	vertex, pre- occipital, and frontal	Detect changes in attended stimuli	supratemporal, auditory cortex
	156-189 ms (N170)	Face and Object Recognition	inferior temporal	Facial and Object Expertise	Fusiform Gyrus, lateral occipital-temporal
	100-300 ms	Go/No-Go	frontal, central	inhibition	caudal and astral anterior cingulate
MMN	100-250 ms Auditory	Physically different stimuli	frontal, central	early sensory memory	temporal lobe, right superior temporal gyrus and plane
Р3	300 ms	oddball (P3b)	occipito-parietal	memory updating, stimulus discrimination	thalamus, hippocampus, superior temporal gyrus and junction
	300 ms	novel stimuli (P3a)	frontal	involuntary attention, inhibition	medial parietal lobe, left superior prefrontal cortex



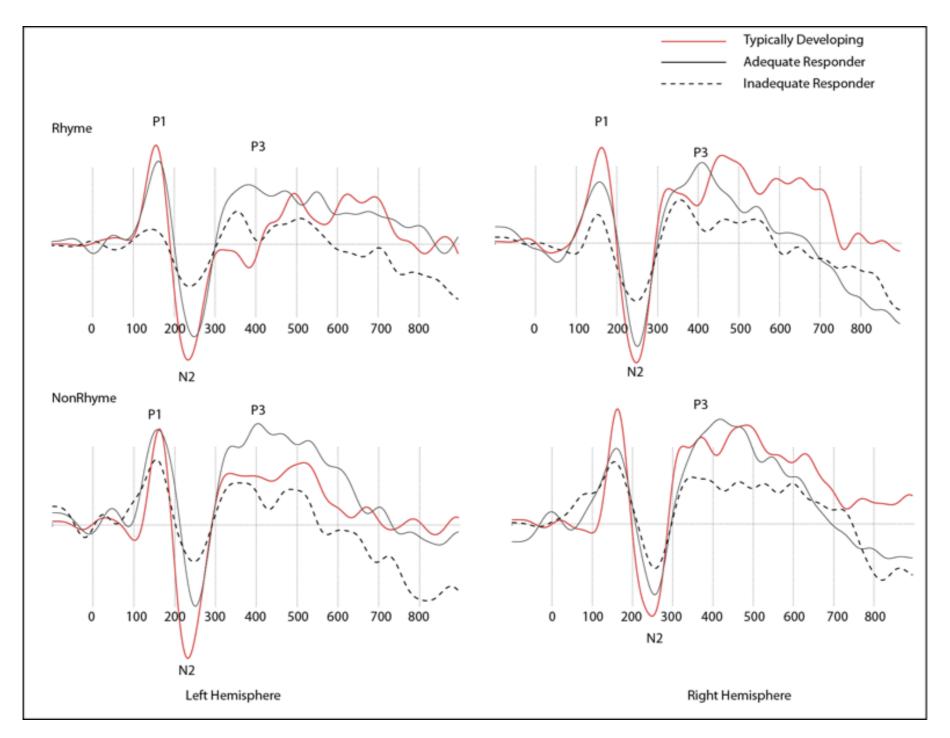
Peaks	Latency	Experimental Manipulation	Max Amplitude	Interpretation	Source
N400	475 ms Visual	Sentence Processing with	Parietal and Temporal	Higher-Order Language Processing	Parahippocampal, anterior fusiform, medial temporal, hippocampus, amygdala, lateral temporal regions
	525 ms Auditory	Semantic Violations			
P600	400-600 ms Memory	Learned vs. Novel	Left Temporal, Frontaocentral	Memory, Novel identification, can be cross-modal	Prefrontal, anterior temporal lobe, anterior cingulate, hippocampus, frontal and temporal cortex
	500 ms Language	Syntactic Violation	Frontocentral	Syntactic violation, phrase structure, subcategorization	Superior Parietal, Precuneus, Posterior Cingulate, Basal Ganglia



Peaks	Latency	Experimental Manipulation	Max Amplitude	Interpretation	Source
ERN	80-150 ms (Response Locked)	Forced Choice RT/ACC (e.g. Flanker)	Frontal & Central	Intent and Motivation	Anterior Cingulate Cortex, DLPFC
FRN	250-350 ms	Any reward feedback	Anterior Frontal Central	Expectation differs from feedback Feedback positive vs. negative	Anterior Cingulate Cortex
CNV	Pre-stimulus negative deflection	Go/No-Go Set ITIs	Vertex	Anticipation	Premotor (BA 6)

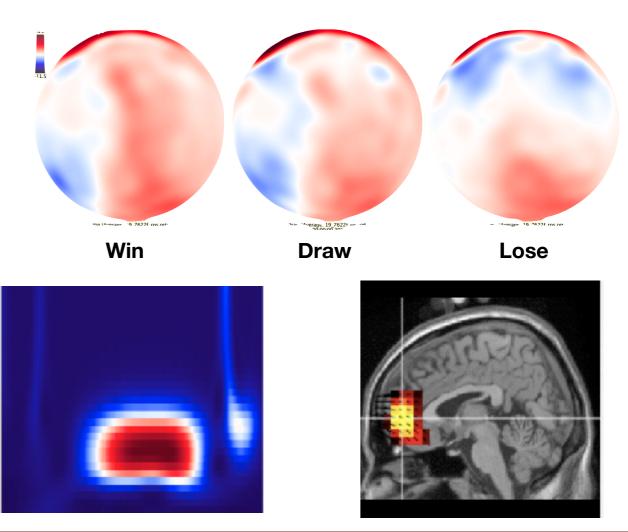


Amplitude and Latency



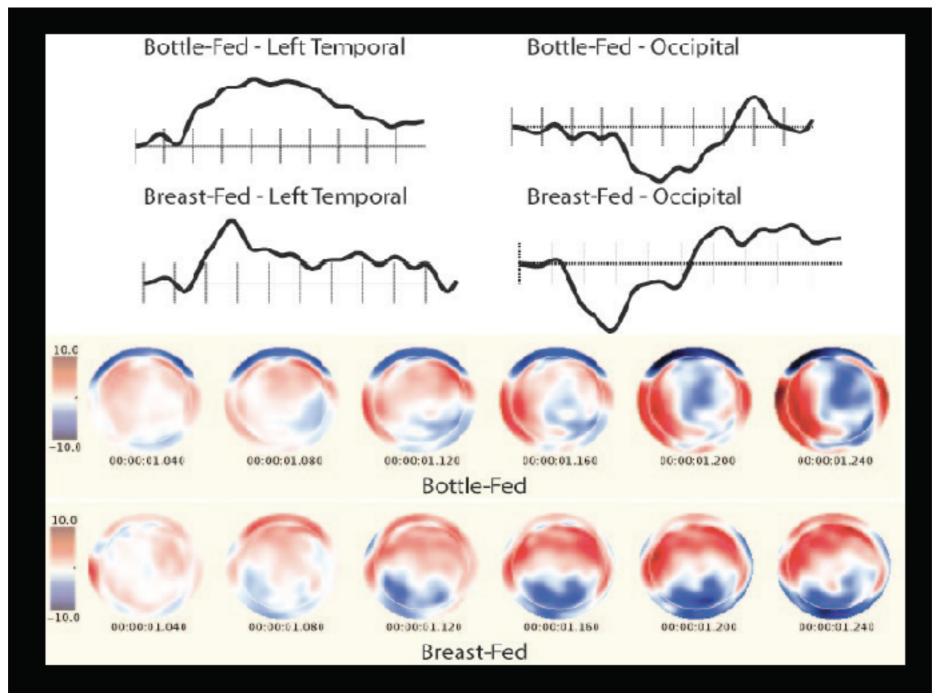
Multiple Metrics of Comparison

- In addition to peak amplitude & latency
 - Scalp Topography
 - Frequency Analysis
 - Wavelets
 - Source Analysis



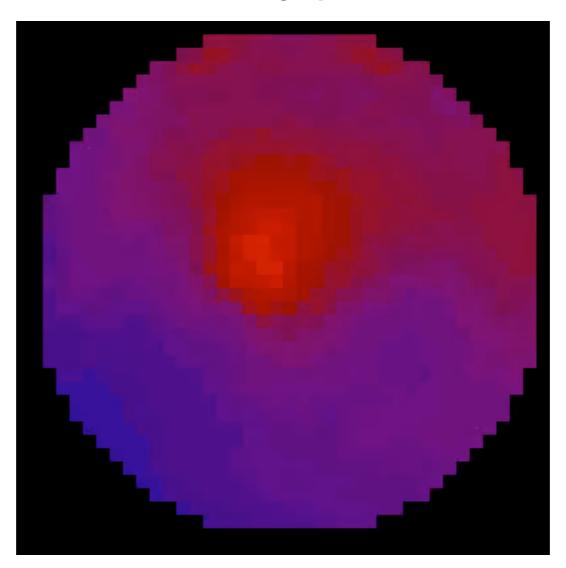


Link between amplitude and topography



Changing Scalp Topography

Front



Right

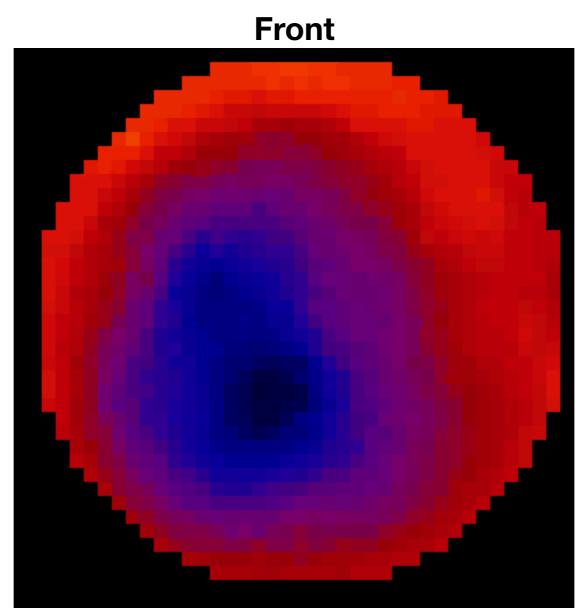
Infant Data

Back of Head



Left

Changing Scalp Topography



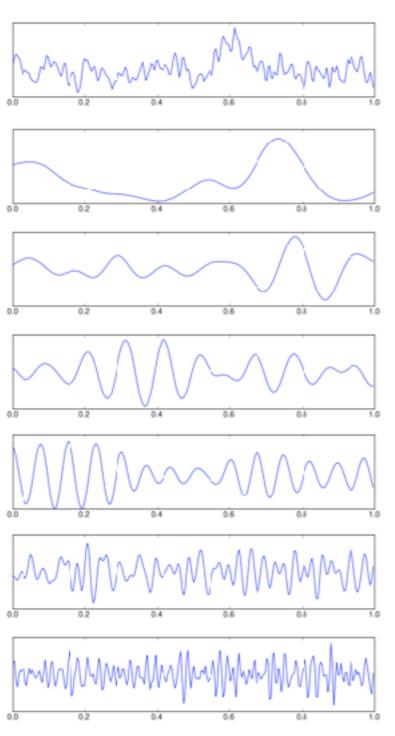
Right

Adult Data

Back of Head

Left

Frequency Analysis



Ongoing EEG

Delta 0-4 Hz

Theta 4-7 Hz

Alpha 8-11 Hz

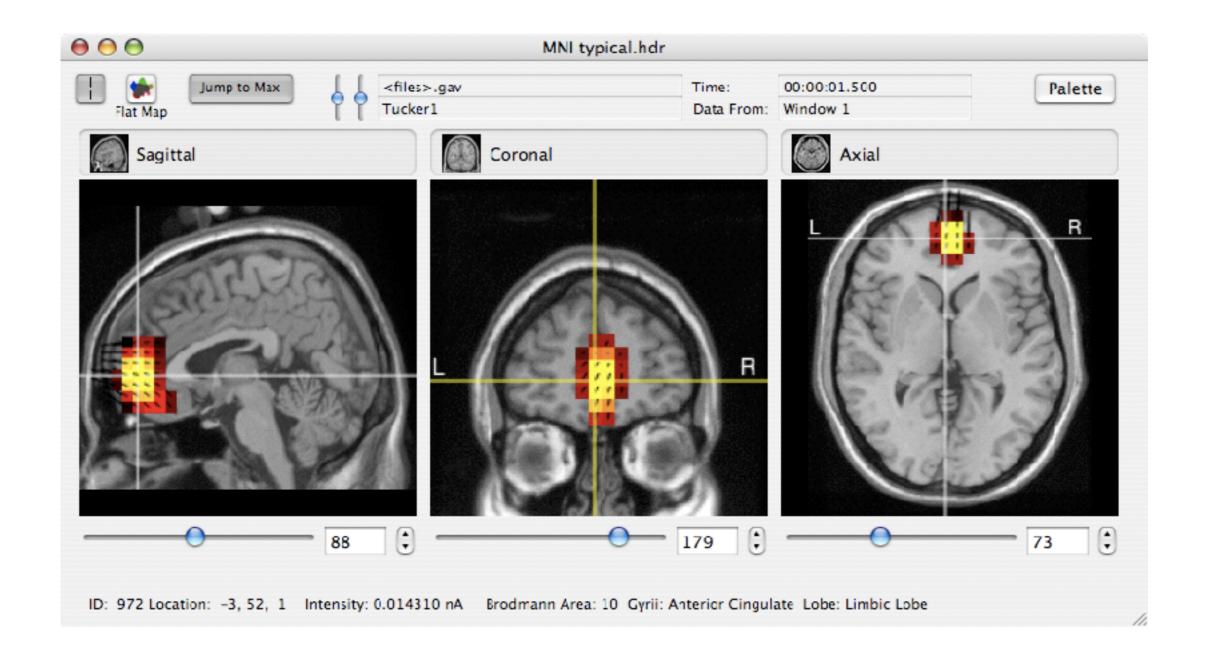
Sensory Motor Rhythm (SMR) 9-12 Hz

Beta 12-25 Hz

Gamma 26+ Hz



Source Analysis



Source Analysis Terminology

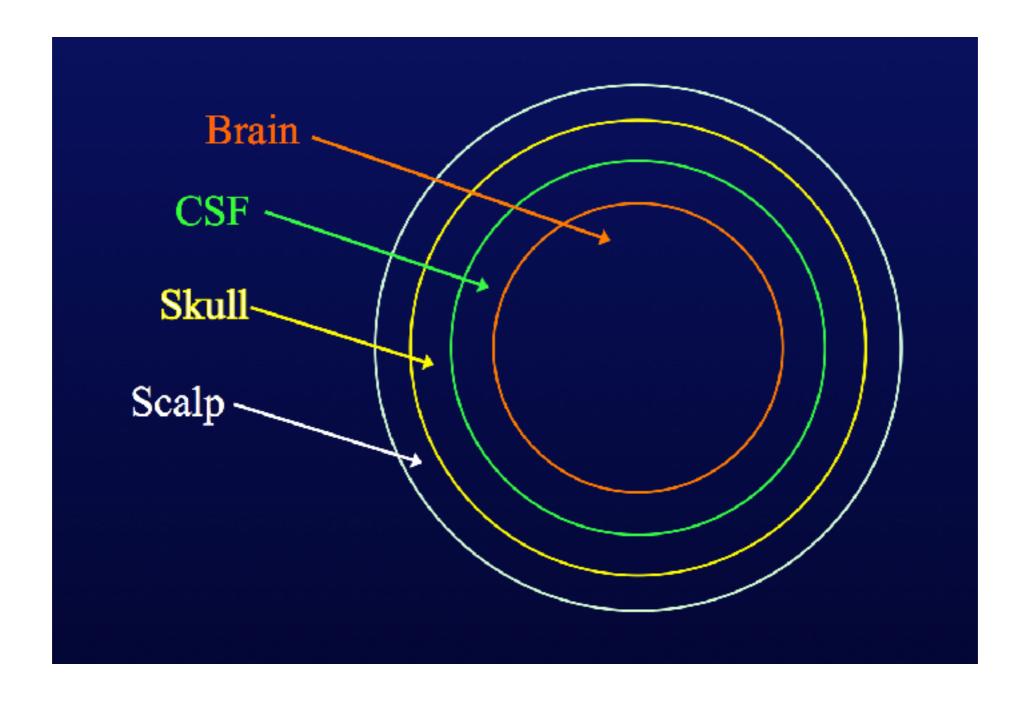
- Forward Problem: Given the sources and a propagation model, what does the scalp topography look like?
- Inverse Problem: Given scalp topography and propagation model, what does the scalp topography look like?

Best Practices in Source Analysis

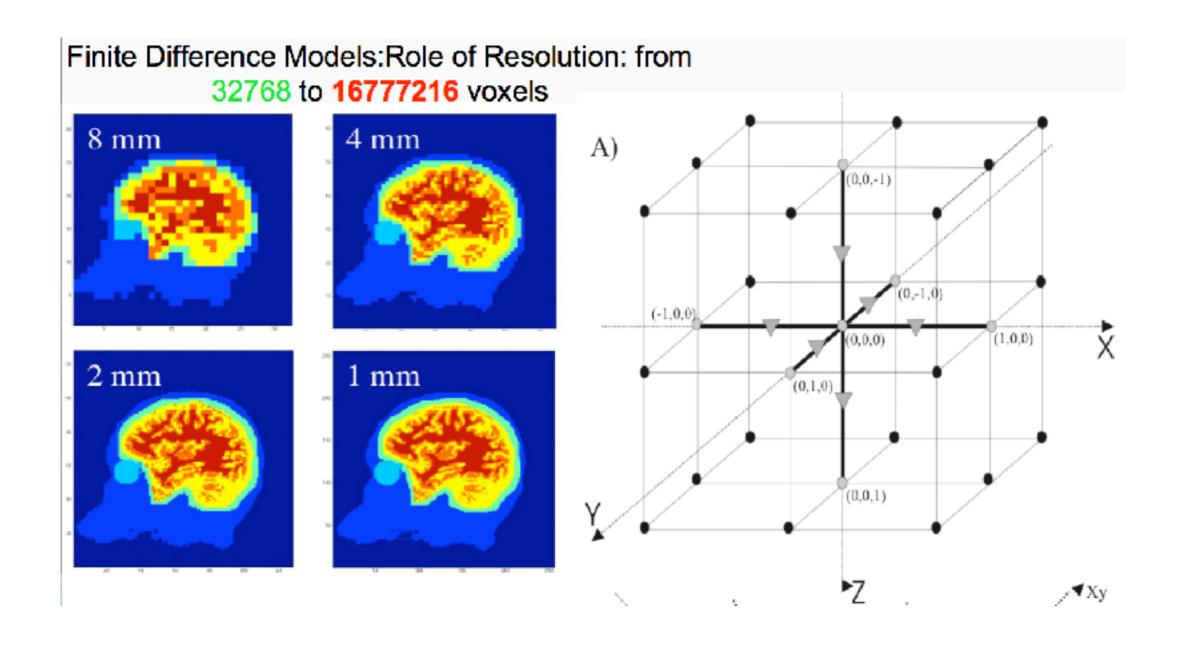
- Adequate Number of Sensors
- Accurate Head Model
- Know the Geometry
- Know the Sensor Locations
- Know Conductivity of the head
- Make due with some propagation model



Head Models

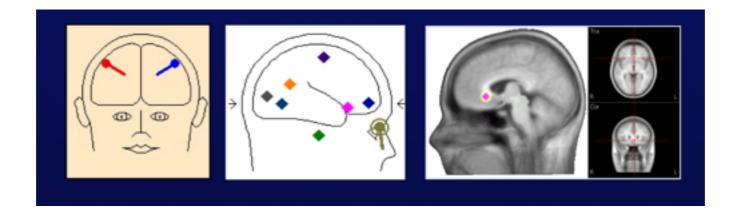


Head Models

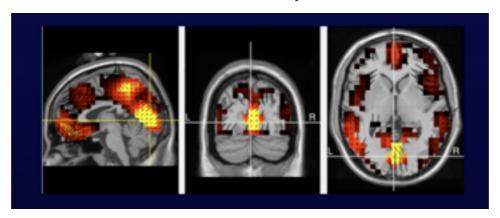


Types of Source Analysis

Equivalent Current Dipole (ECD)



Distributed Linear Inverse (LORETTA)



ECD

- Assumes a small number of sources can model surface topography
- Iteratively nonlinear search for location that accounts for largest proportion of variance in scalp topography
- Repeat and add more sources until percentage variance desired is accounted for
- Challenges:
 - High user involvement (low replication)
 - Have to balance time window, number of dipoles, and orientation of dipoles



Distributed Linear Inverse

- Often synonymous with Minimum Norm Solutions
- Linear mapping of scalp data to distributed sources within the brain
- All dipoles are active in the brain to different degrees
- Underdetermined



Distributed Linear Inverse

- No true/unique solution. Need to apply constraints:
 - Minimum Norm: smallest overall intensity of activation
 - Weighted Minimum Norm: Apply weights to compensate for superficial sources (smoothing)
 - Regularization reduce the effect of random noise



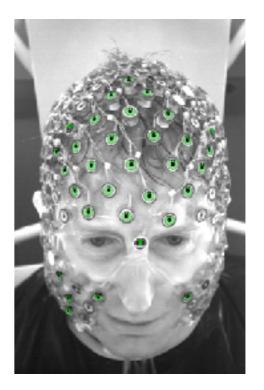
Options for Source Analysis

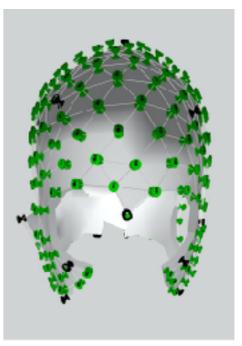
- Best: MRI + EEG Electrode Locations (GPS) + EEG
 - Individual Head Models
- Good: EEG Electrode Locations (GPS) + EEG
 - Template Head Models with custom sensor loc
- OK: EEG
 - Template Head Models only



Electrode Coordinates







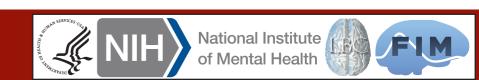


Solving Electrode Locations



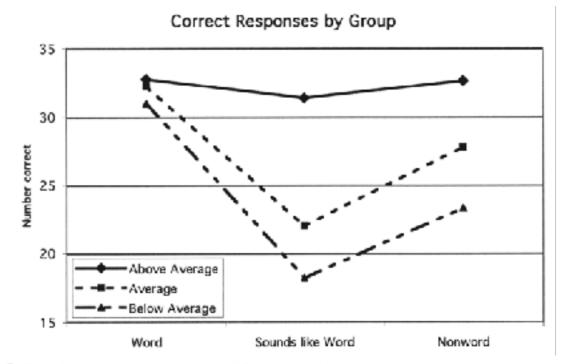
EEG and fMRI Similar Results

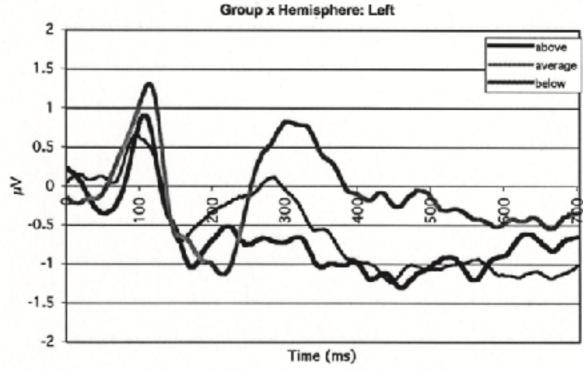
- Limited research covering language and simultaneous EEG-fMRI
- Extensive research in both fMRI and EEG showing
 - Differences between TD and RD/Dyslexia
 - Changes in brain with development of language
 - Changes in brain following reading intervention
- Converging findings



ERPs of Reading Groups

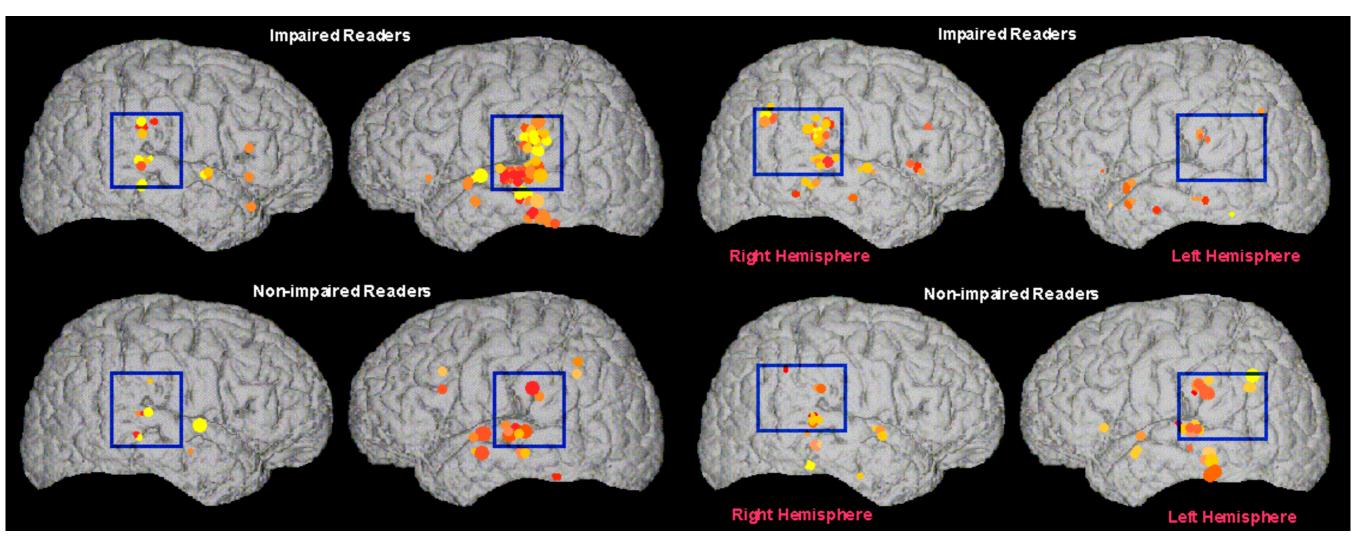
- N=27 children
- 9-12 years of age
- ERP amplitudes and peak latencies decreased as reading skills in- creased
- Hemisphere differences increased with higher reading skill levels
 - LH > RH is more proficient reading skill







MEG of Reading Groups

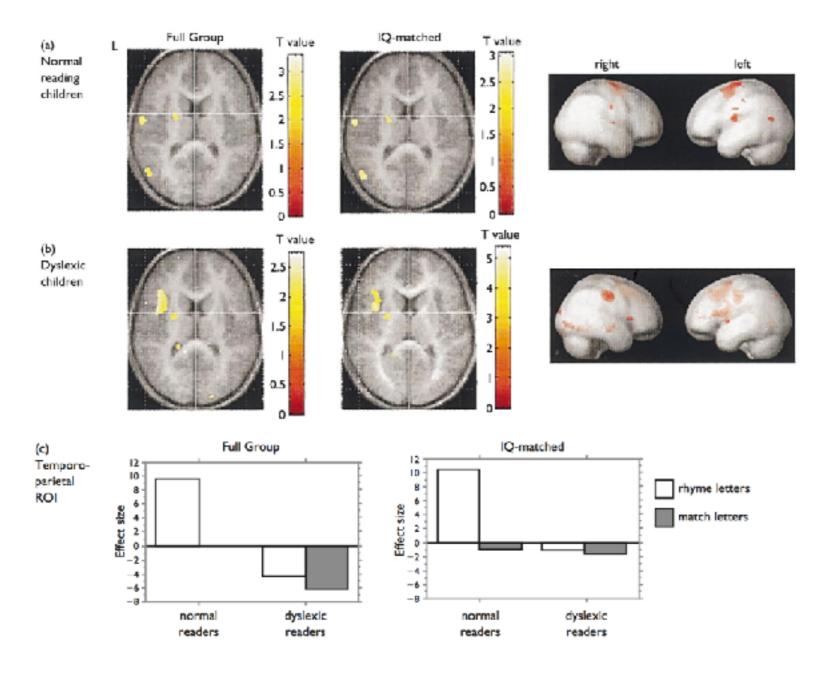


Listening to Words

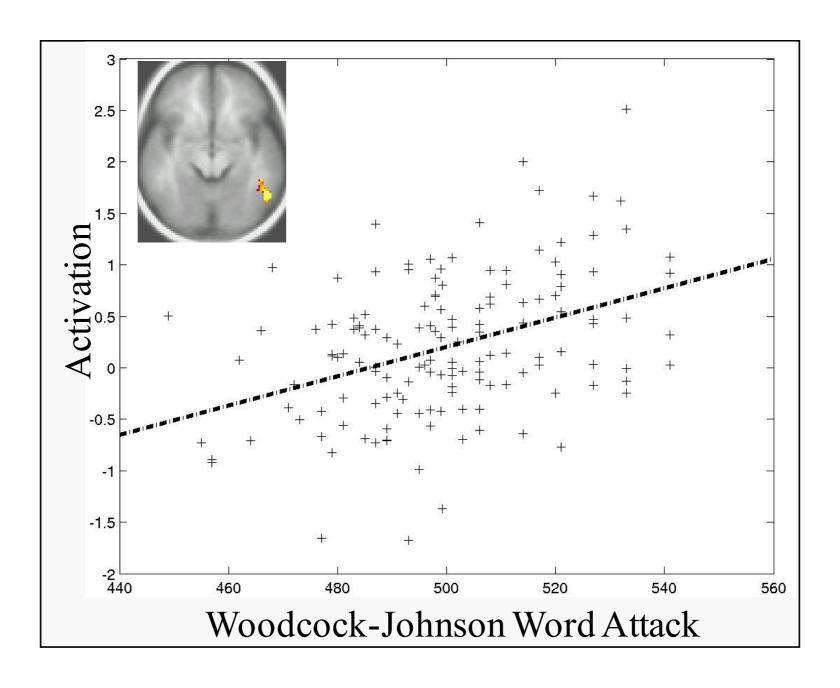
Reading Words



fMRI Differences RD and TD



Reading Skill & VWFA



Intervention Studies

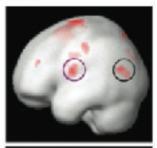
- Looking at changes in reading circuit associated with improvement in reading ability
- fMRI
- MEG
- ERP



fMRI Reading Intervention I

A Children with no remediation

Normal reading children while rhyming



Dyslexic reading children while rhyming before remediation



B Dyslexic children increases after remediation

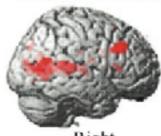




Fig. 1. Neural effects of remediation in children with developmental dyslexia. (A) Left hemisphere activations of control children and children with dyslexia are shown during rhyming (as compared with matching) letters (P < 0.025, 20-voxel threshold; ref. 12). (B) Brain areas that showed increased activity during phonological processing in the dyslexic group after remediation. Shown at P < 0.01, 20-voxel threshold. Black circles highlight left tempora parietal region, which is disrupted in children with dyslexia and affected by remediation. Purple circles highlight the left frontal region that is active in control children and is affected by remediation in children with dyslexia.

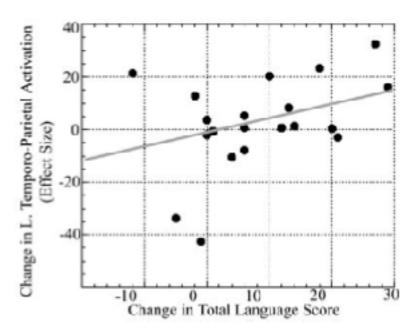
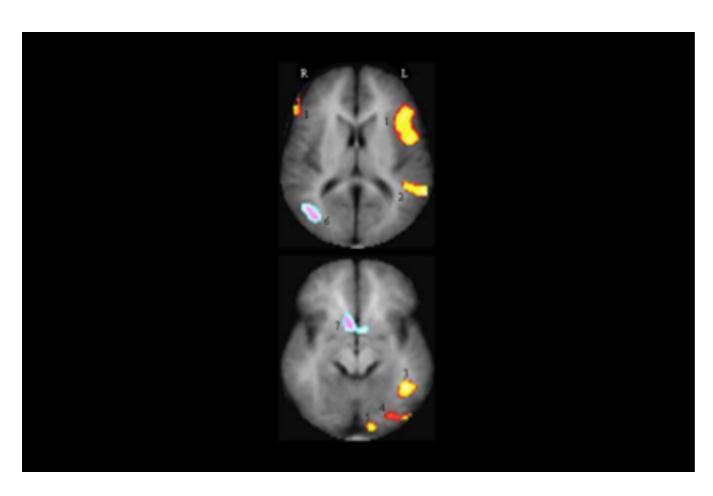
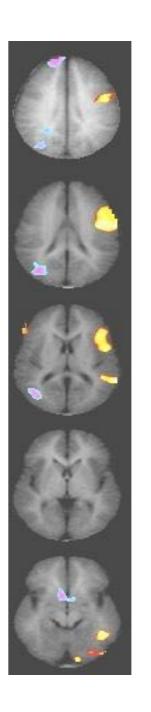


Fig. 2. Language improvement and increased brain function. Correlation between magnitude of change in left temporo-parietal ROI (BA 39) and improvement in oral language (r = 0.41, P = 0.03). Left temporo-parietal ROI encompassed brain areas that showed underactivation and increases after training in children with dyslexia. Change in effect size is on the vertical axis; change in total language score (CELF-3) is on the horizontal axis. Effect size is the weighted sum of parameter estimates from the multiple regression for rhyme vs. match contrast pre- and posttraining.

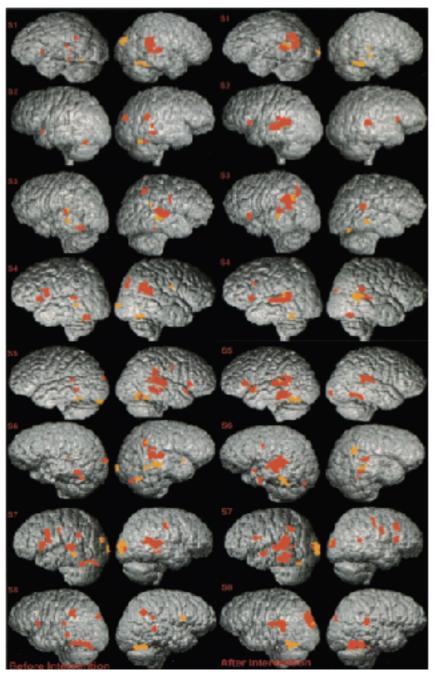
fMRI Reading Intervention II



Year 3 - Year 1



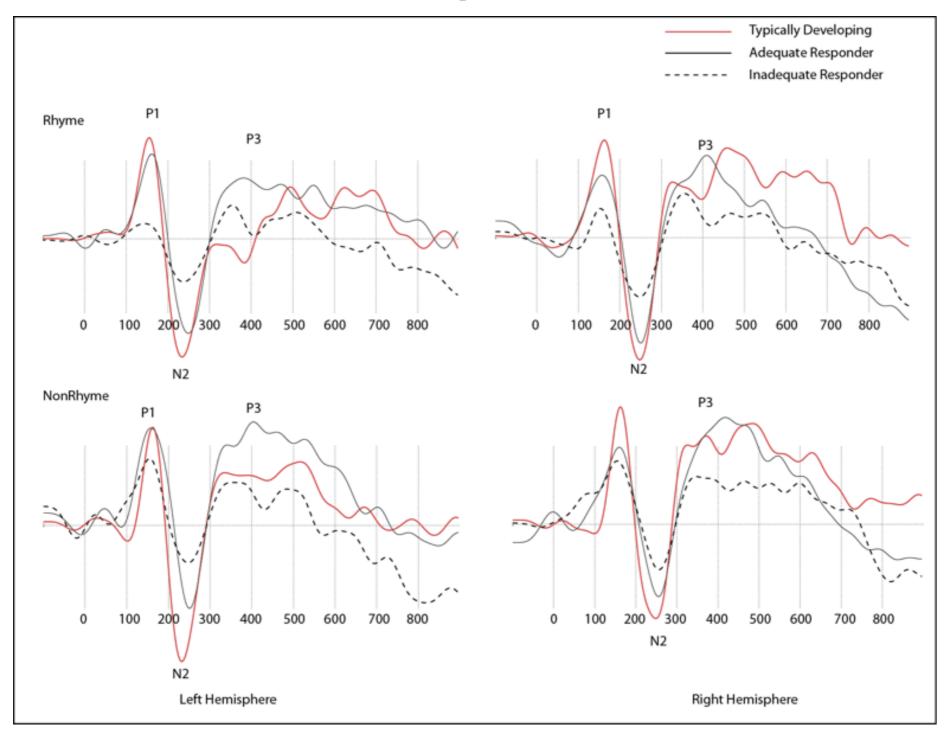
MEG Reading Intervention



Pre

Post

ERP Reading Intervention



Classifying Intervention

Discriminant Function - p<0.003

	TD	AR	IR
TD	5	3	
AR	2	6	
IR			7

	Reading >= GL	Reading < GL	
Reading >= GL	16		96%
Reading < GL	I	7	

General Themes

- Activation differences between good and poor readers
- Modulation of parts of the reading circuit
- Laterality differences
- Reading Intervention leads to "normalization" of the reading circuit and in some cases "compensatory" activation

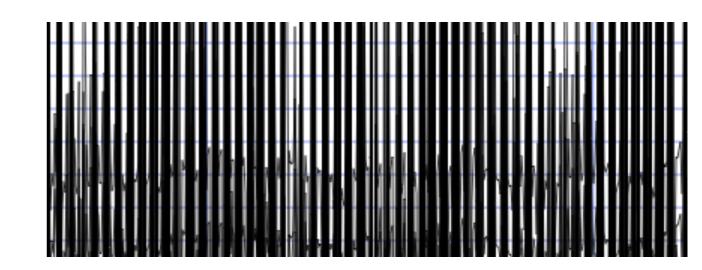
The Advantages to Simultaneous EEG-fMRI

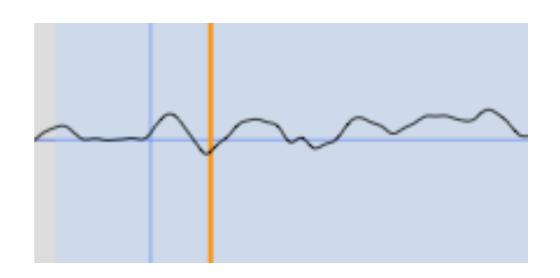
- Quite a bit of information can be gathered from separate sessions of EEG and MRI
- The real benefits are:
 - Same environment in both cases (inside an MRI)
 - Looking at individual trial modulation of either ERP/ BOLD
 - Looking at coherence and resting state relationship at the same time



Artifacts

- MR Artifacts
 - Caused by the gradients
- Ballistocardiogram (BCG)
 - Caused by movement of the electrode within the magnetic field

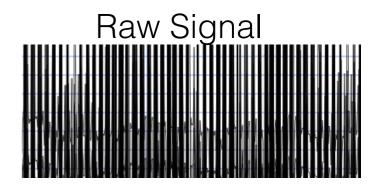




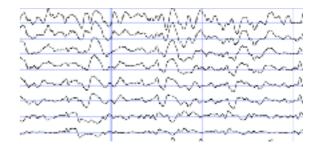


Artifact Removal

- Gradient Artifacts
 - Fairly straightforward to remove
 - Template subtraction
- BCG Artifacts
 - More prominent on the facial electrodes
 - Follows the heartbeat by ~250 ms
 - PCA to model the artifact, remove, reconstruct



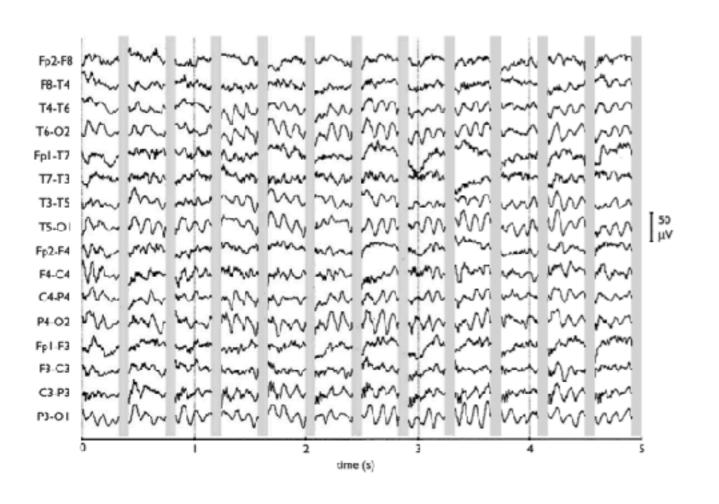
Artifact Removed

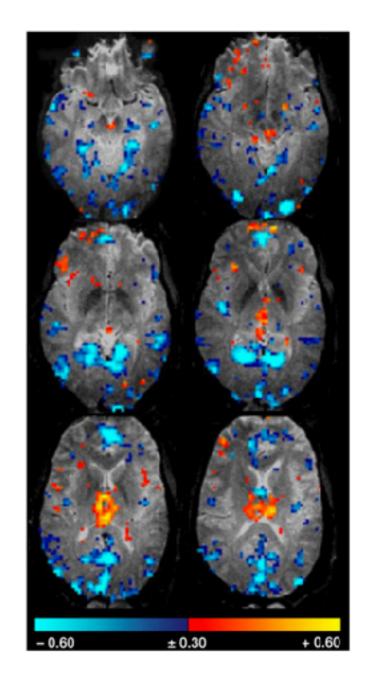




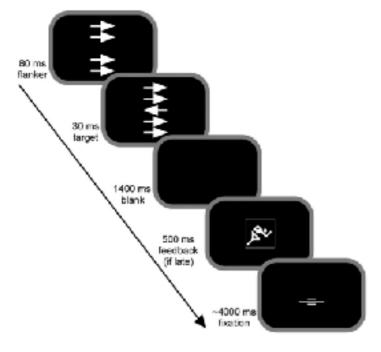


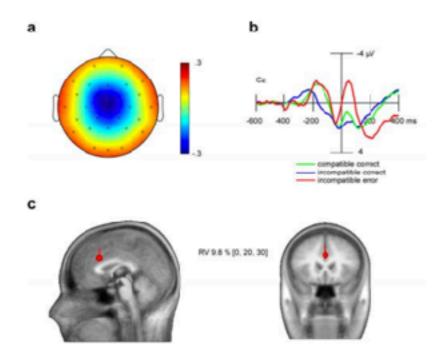
Alpha Rhythm

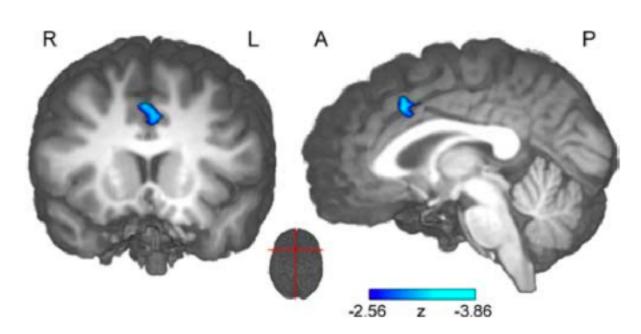


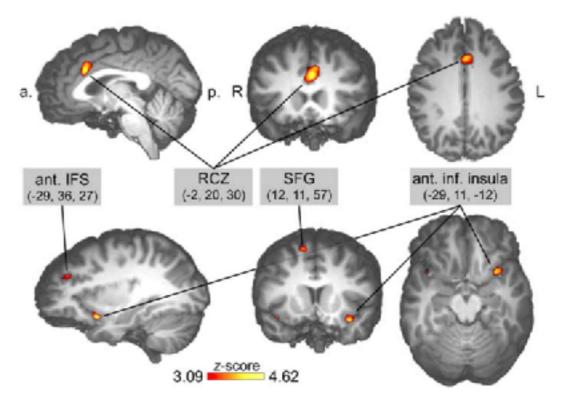


Flanker Example

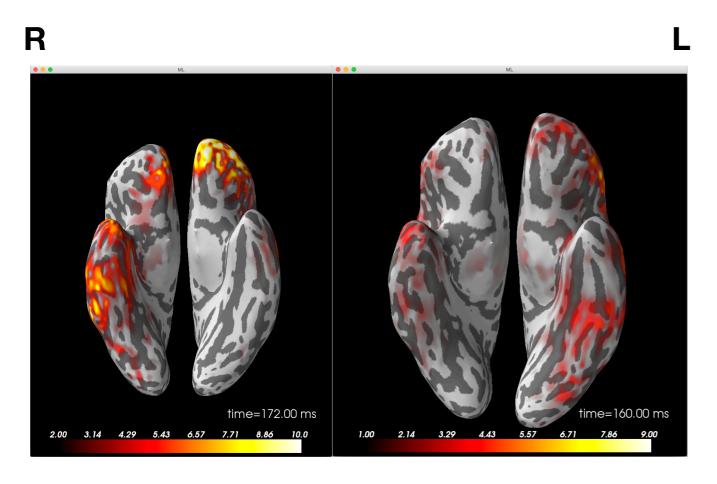








N170 Faces / Characters



Elit FFA 104,105,113,114

Characters
Faces

0.0 0.2 0.4 0.6 0.8

Faces Characters

Left FF

Questions?

Peter.Molfese@nih.gov

