

Multimodal Neuroimaging Overview

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NIH Summer Neuroimaging Course

July 18, 2024





cmn.nimh.nih.gov



Simultaneous EEG-fMRI

- Equipment
- Experiment Design
- Analysis
- Interpretation / Writeup

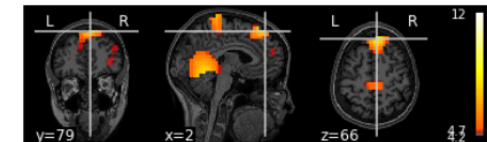
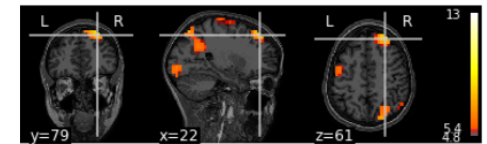


Workshops

- 2019 - Multimodal Neuroimaging
- 2021 - Naturalistic Stimuli & Individual Differences
- 2022 - Simultaneous PET-MRI
- 2024 - Eye Tracking
- 202X - Layer fMRI & Other Modalities

Multi-Modal

- Combine Data from multiple modalities
 - EEG, fMRI, DTI, EyeTracking, Behavior
- Refer folks to expert cores
 - AFNI, MEG, PET



Why This Talk?

13	Thursday	7/18/24	2:00PM	FAES B1C207	Multimodal Neuroimaging Overview	PDF	Pete Molfese
14	Tuesday	7/23/24	2:00PM	FAES B1C209	Electroencephalography (EEG) with and without simultaneous fMRI	PDF	Pete Molfese

- Two talks with potentially overlap
- Today: Mostly Theory
- Next week: More applications / Video Demos
 - How to do EEG/EEG-MRI at the NIH

Overview

- What is Multimodal Neuroimaging?
 - Definition
 - Motivation
- Doing this well is hard
- Doing this well is important

Multimodal Neuroimaging

- Any neuroimaging that uses multiple (complimentary) methods

- fMRI + EEG*



- MEG + EEG*

- fMRI + MRI



- fMRI + DTI

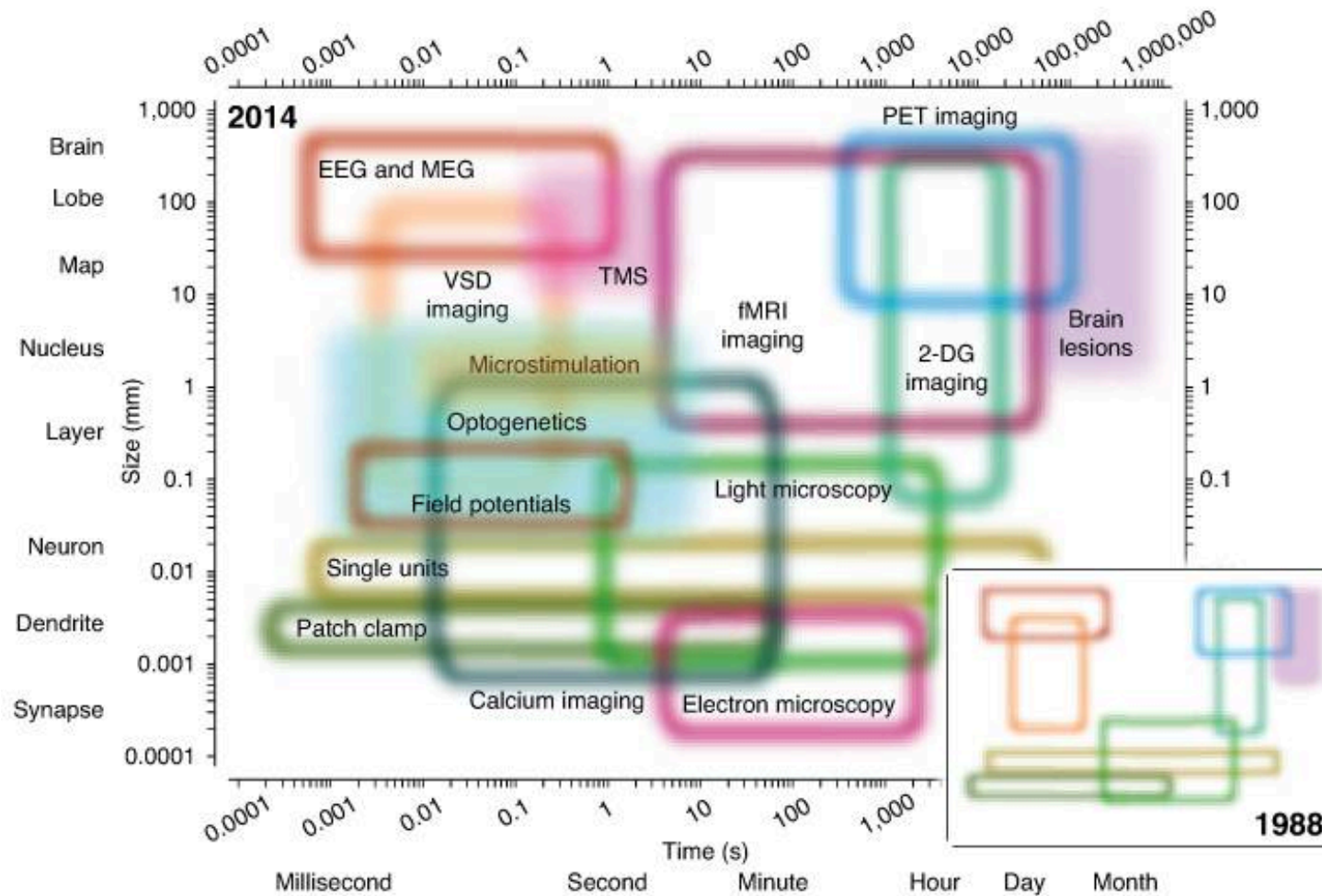
- fMRI + MEG

- tDCS/tACS/TMS + EEG*

- TMS + fMRI*

- Etc...

Temporal-Spatial Tradeoffs



Multimodal Possibilities

	EEG	MEG	sMRI	fMRI	DTI	MRS	PET	TMS/ tXCS
EEG	Black							
MEG	Green	Black						
sMRI		Red	Black					
fMRI	Green	Red		Black				
DTI	Red	Red			Black			
MRS		Red			Red	Black		
PET	Green	Red		Green	Red	Red	Black	
TMS/ tXCS	Green	Green			Red	Red	Red	Black



Possible



Not Currently Possible

Doesn't Have To Be Simultaneous

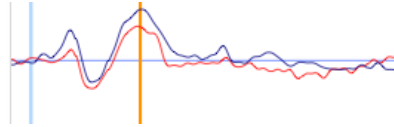
	EEG	MEG	sMRI	fMRI	DTI	MRS	PET	TMS/ tXCS
EEG	Black							
MEG	Pink	Black						
sMRI	Pink	Pink	Black					
fMRI	Pink	Pink	Pink	Black				
DTI	Pink	Pink	Pink	Pink	Black			
MRS	Pink	Pink	Pink	Pink	Pink	Black		
PET	Pink	Pink	Pink	Pink	Pink	Pink	Black	
TMS/ tXCS	Pink	Pink	Pink	Pink	Pink	Pink	Pink	Black

 Possible As Separate To Combine Data

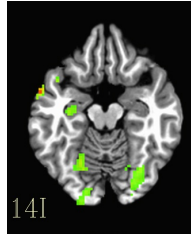
Tradeoffs

Information Tradeoffs

- “When in the brain”



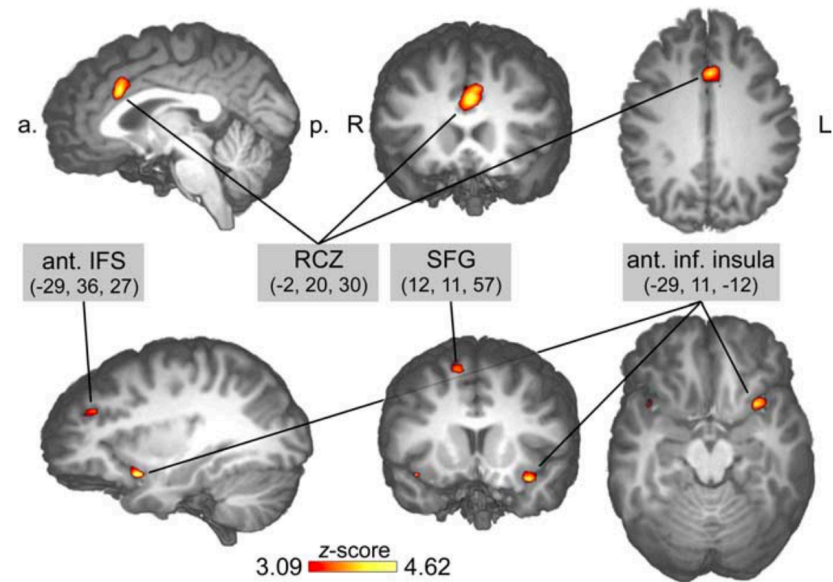
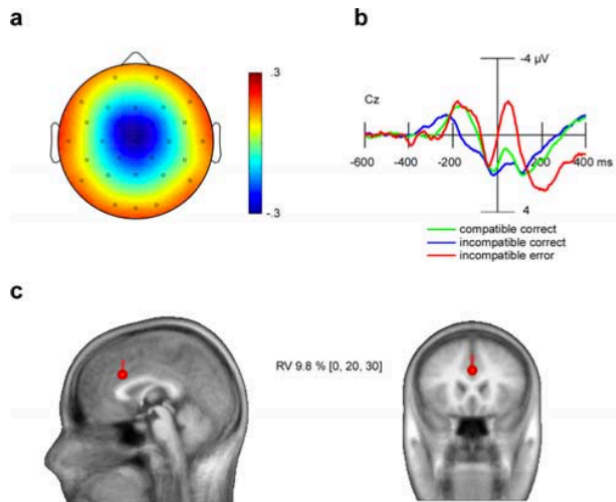
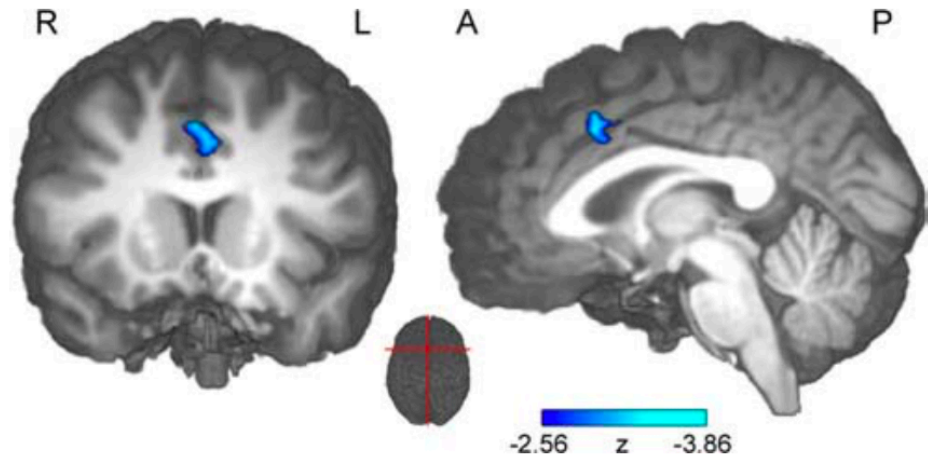
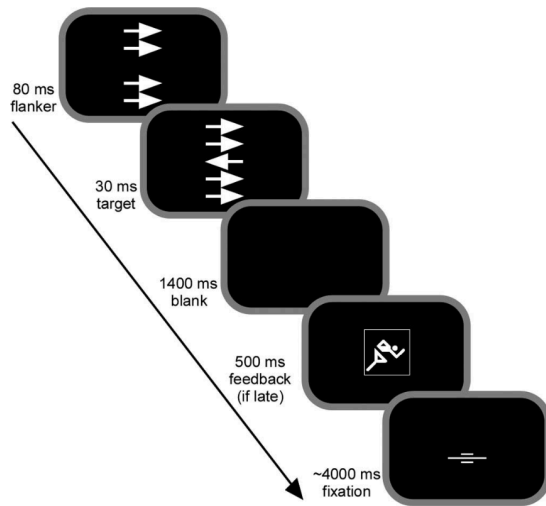
- “Where in the brain”



- “What happens to a process when X area is disrupted?”
 - Is there a fallback network?
- “What happens to a process when disrupted at Y time?”
 - How does the brain compensate?



EEG-fMRI Example

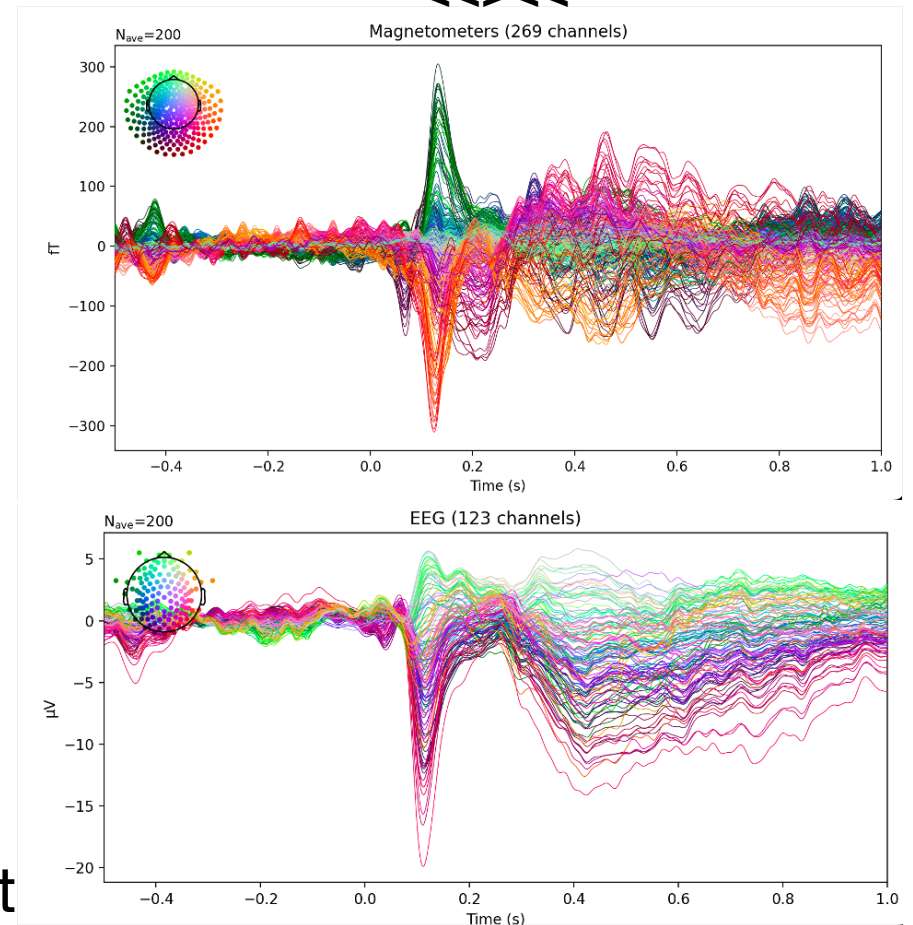


EEG-MEG

- EEG & MEG have different sensitivity profiles
 - EEG picks up on “all” sources (including radial), lower SNR, source “smearing”
 - MEG most sensitive to tangential sources, higher SNR
 - These complimentary profiles allow improved source localization accuracy (Sharon, et al. 2007; Aydin et al. 2015)

Flanker Task

<<><<



Courtesy Lucrezia Liuzzi (SDAN)

fMRI + Diffusion

- Can use fMRI activation (resting state or task) as seeds for diffusion tensor imaging
- Memory scores correlated with global and local “efficiency” in DMN
- Anxiety negatively correlated

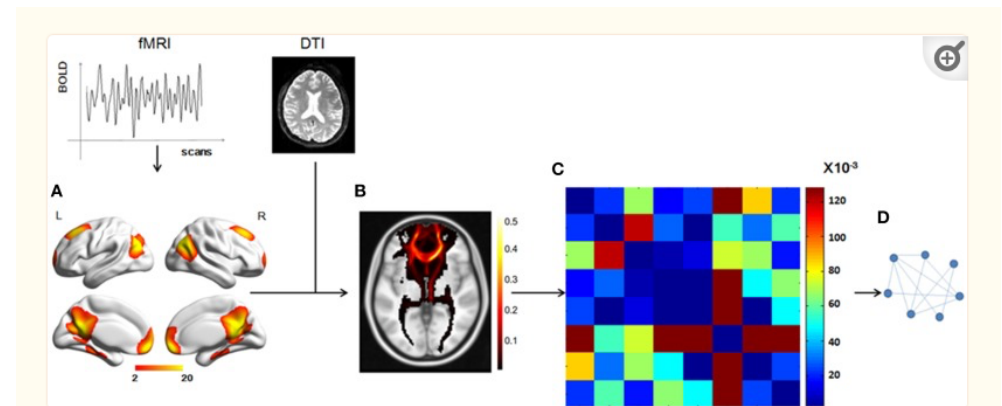
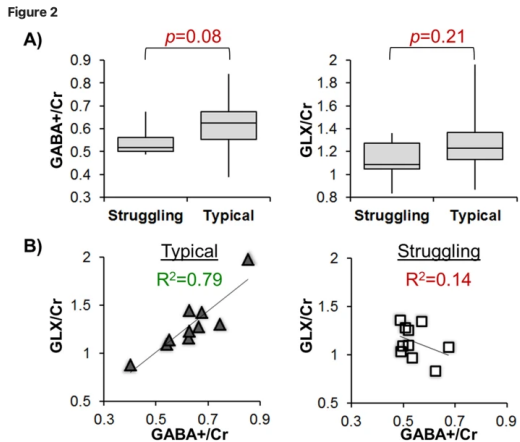


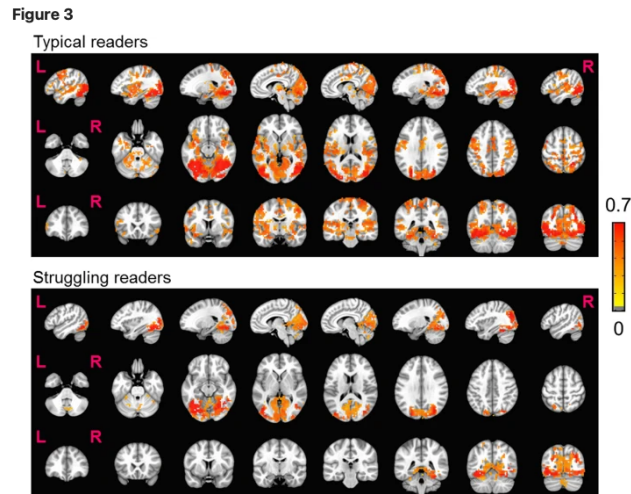
Figure 1

Total workflow before the statistical analysis. (A) The DMN mask was extracted from fMRI data using group ICA, with a threshold of $z = 2$. (B) A probabilistic fiber tractography method was used to track the fibers from each ROI and the connectivity probability between any two ROIs was calculated to get the matrix (C). (D) A graph theory method, in which the nodes represented the ROIs and the weighted edges reflected the connectivity probabilities between any two nodes, was used to analyze the DMN structural network.

fMRI + MRS



The age-corrected neurotransmitter concentrations for both typical and struggling reader groups. (A) Though not significant, the struggling readers have lower GABA+/Cr and GLX/Cr concentrations in the frontal regions. (B) The typical readers show a strong relationship between GABA+/Cr and GLX/Cr, whereas struggling readers may have a neurotransmitter imbalance in their frontal system.



Functional connectivity maps in typical and struggling readers ($p=0.001$, cluster size 100, FWE corrected) when seeded from L-FG. Color bar indicates the Z(CC). Note: L=left, R=right.

- Typical and struggling adult readers
- GABA in L IFG predicts real word reading behavior, but only if there is significant rsFC between fusiform and higher-order reading areas
- rsFC between L Fusiform and other language systems predicts oral reading of real words, irrespective of GABA

Technique Tradeoffs

	Pro	Con
EEG	High Temporal Resolution Comparable Across Lifespan Inexpensive	Low (Source) Spatial Resolution Source Analysis Difficult
MEG	High Temporal Resolution Less Difficult Source Analysis	Expensive Difficult to use with Children Limited Signal from Deep Tissue
fMRI	Excellent Spatial Resolution	Expensive Difficult to use with Children Low Temporal Resolution (HRF)

Doing Multimodal is Hard

Design



- Knowledge of the domain or topic
- Knowledge of component parts (e.g. MRI + EEG)
- Technical Knowledge to carry out some portion of the experiment

Data Analysis



Interpretation



More Design Questions

- Do you have the equipment?
- Do you have the expertise to operate the equipment?
- Is it worthwhile?
 - More data isn't always better.
- Can you do this well?

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Interpretation



Doing Multimodal is Hard

Design



- Knowledge of the domain or topic
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- What's the value added?
- Does it have to be simultaneous?

Data Analysis



Interpretation



Simultaneous? Really?

- Quite a bit of information can be gathered from separate sessions of EEG and MRI
- The benefits of simultaneous 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

Doing Multimodal is Hard

Design



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- Important choice points: Simultaneous vs. Separate
- What's the value added?

Data Analysis



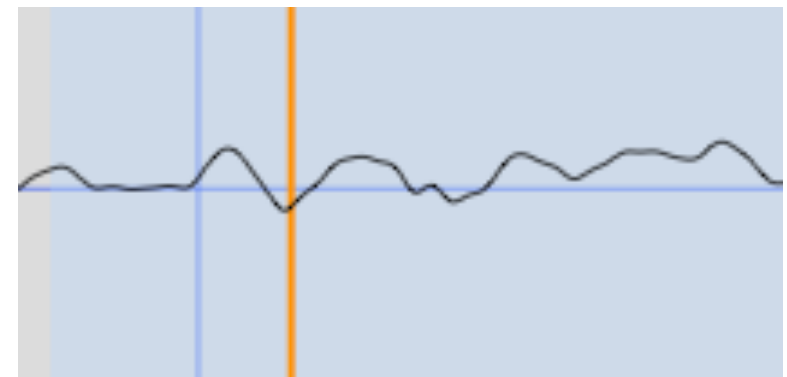
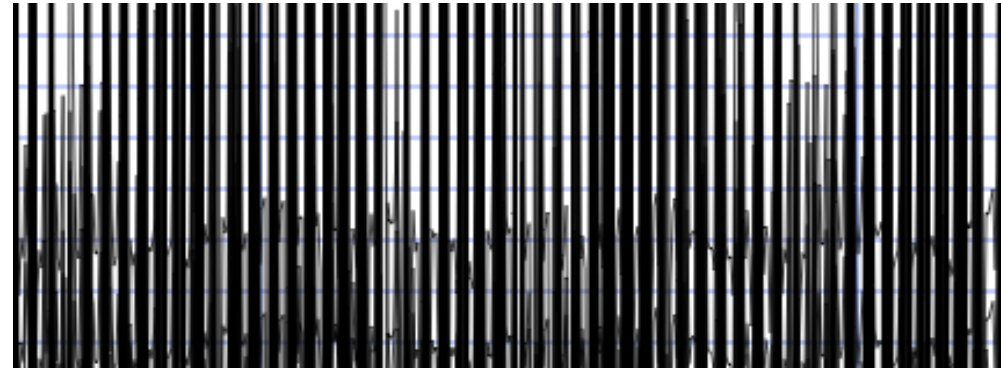
- Are there artifacts from multimodal imaging?
 - Can you fix them?

Interpretation



Artifacts

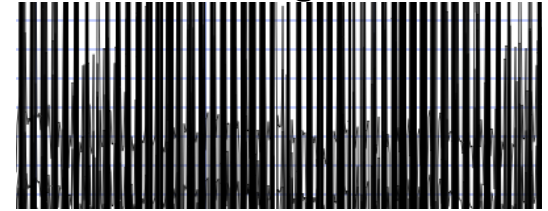
- MR Artifacts
 - Caused by the gradients
- Ballistocardiogram (BCG)
 - Caused by movement of the electrode within the magnetic field



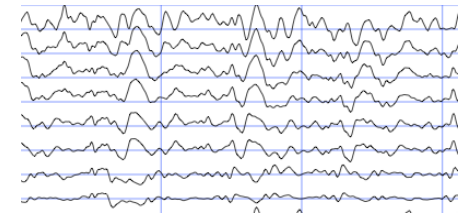
Artifact Removal

- 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

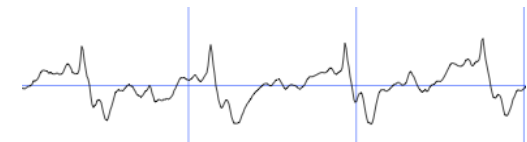
Raw Signal



Artifact Removed



BCG Detection



Doing Multimodal is Hard

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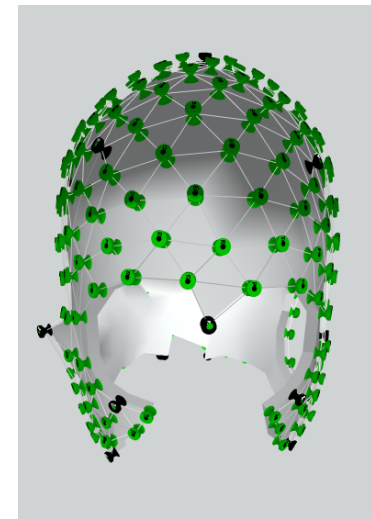
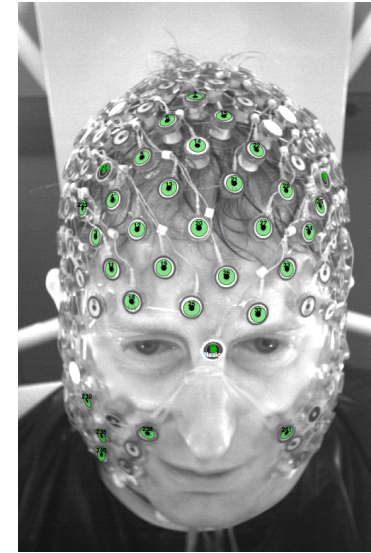


- Are there artifacts from multimodal imaging?
 - Can you fix them?
- If applicable, how will you combine the data?
 - Co-register data across modalities
 - Head Models / Single Subject vs. Group
 - EEG Sources

Interpretation

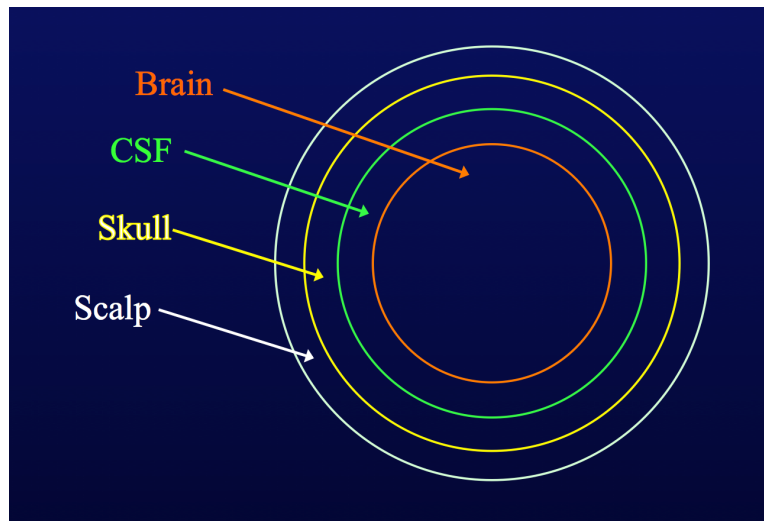


Electrode Coordinates



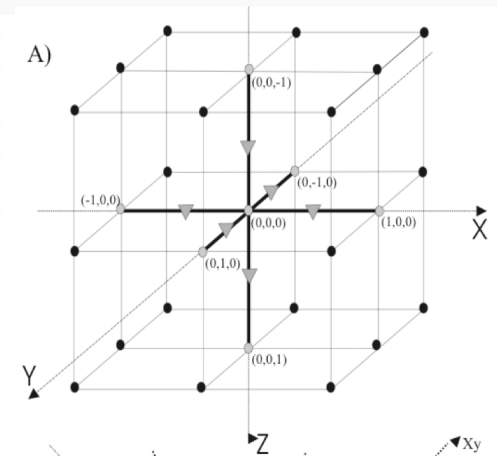
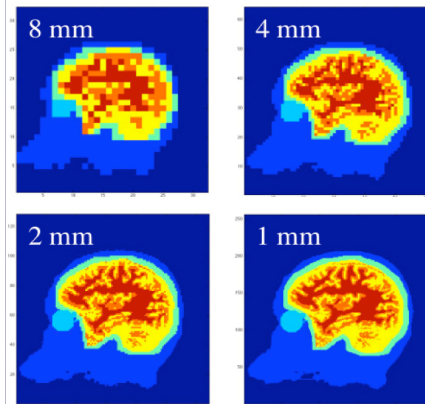
Head Models

Shell Models



Finite Difference/Element Models

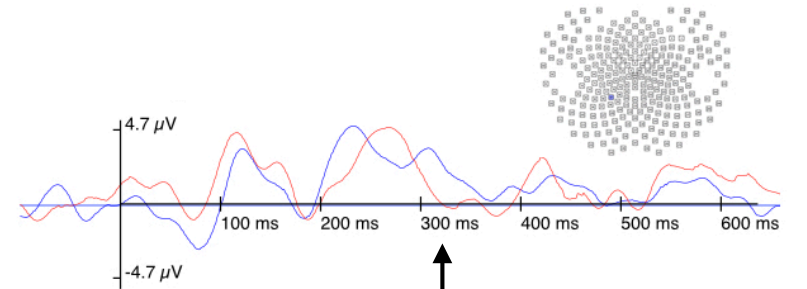
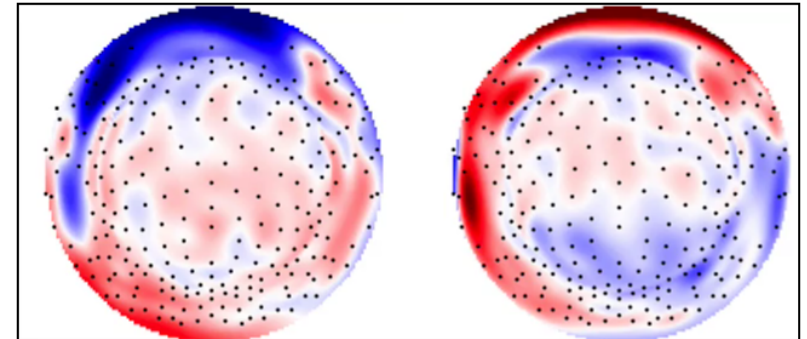
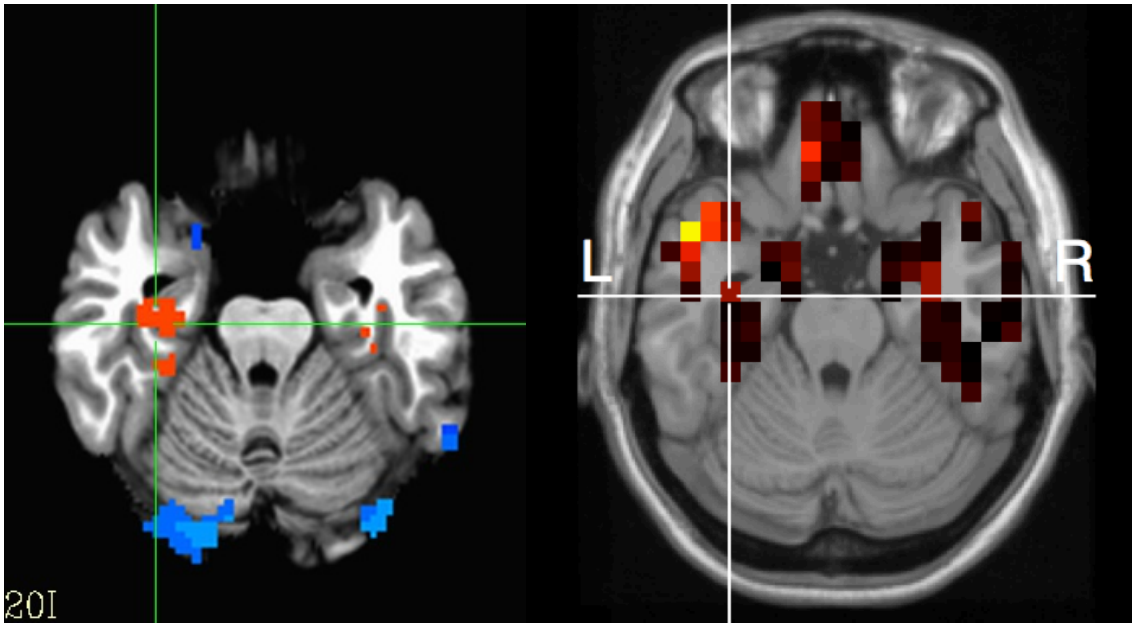
Finite Difference Models: Role of Resolution: from
32768 to 1677216 voxels



Difficulty Combining Very Different Measures

fMRI

EEG



Parahippocampal Activation to P300

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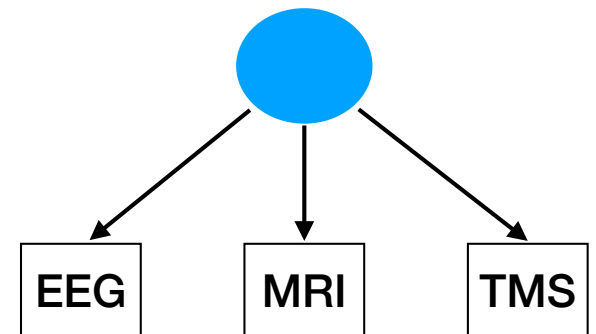
- How to interpret findings?
 - Do the analyses of your data let you make certain generalizations?
- What does it really mean?

Good News

- We're making great progress on integrating Multi-modal techniques
 - Simultaneous methods have improved noise and artifact reduction
 - Non-concurrent methods continue to improve integration and comparison of results

Multimodal is Important

- Provides (hopefully) complimentary and (possibly) overlapping information
- May facilitate better studies of individual differences
 - May be used as a control within subject!
 - Allow for creation of multi-measure latent variable models
- Improve reliability/replication



Open Questions

- Design: What Questions best lend themselves to multi-modal imaging?

- Analysis: How should analyses be standardized to best make use of multimodal imaging?

- Interpretation: What will be the scientific/statistical level of rigor to support multimodal imaging conclusions

Questions?

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