

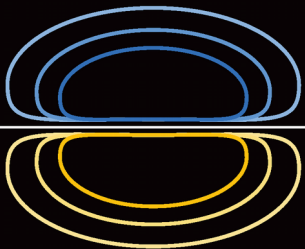


**Karolinska  
Institutet**

Seminar on:

Open data – Standardized analysis  
pipeline for  
Magnetoencephalography (MEG)

By Lau M. Andersen




NatMEG

The Swedish National Facility for Magnetoencephalography



# Group Analysis in MNE-Python of Evoked Responses from a Tactile Stimulation Paradigm: A Pipeline for Reproducibility at Every Step of Processing, Going from Individual Sensor Space Representations to an across-Group Source Space Representation

 Lau M. Andersen\*

NatMEG, Department of Clinical Neuroscience, Karolinska Institutet, Stockholm, Sweden



Open-source Python software for exploring, visualizing, and analyzing human neurophysiological data: MEG, EEG, sEEG, ECoG, and more.



PROTOCOLS ARTICLE

Front. Neurosci., 01 May 2018 | <https://doi.org/10.3389/fnins.2018.00261>



# Group Analysis in FieldTrip of Time-Frequency Responses: A Pipeline for Reproducibility at Every Step of Processing, Going From Individual Sensor Space Representations to an Across-Group Source Space Representation

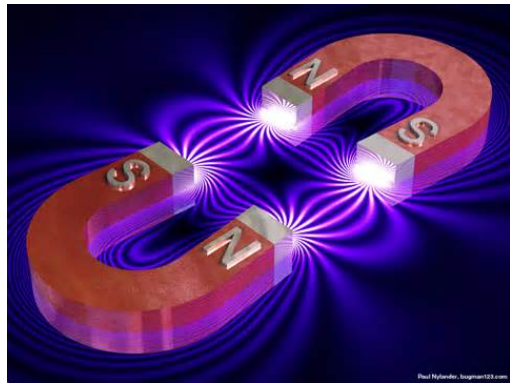
 Lau M. Andersen\*

NatMEG, Department of Clinical Neuroscience, Karolinska Institutet, Stockholm, Sweden

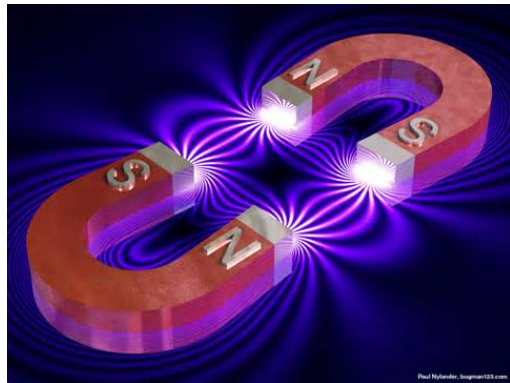


# Magneto·encephalo·graphy

# Magneto-encephalo-graphy



# Magneto·encephalo·graphy



*Writing out the magnetic fields of the brain*

# Goals for today

- 1) (*What is MEG?*) You are to be able to recognize MEG signals
- 2) (*How to analyse MEG*) You are to be able to find relevant resources for group level analysis, so you can get started yourself
- 3) (*How to decide about MEG*) You are to know about statistical tools for answering questions about MEG

*What is MEG?*

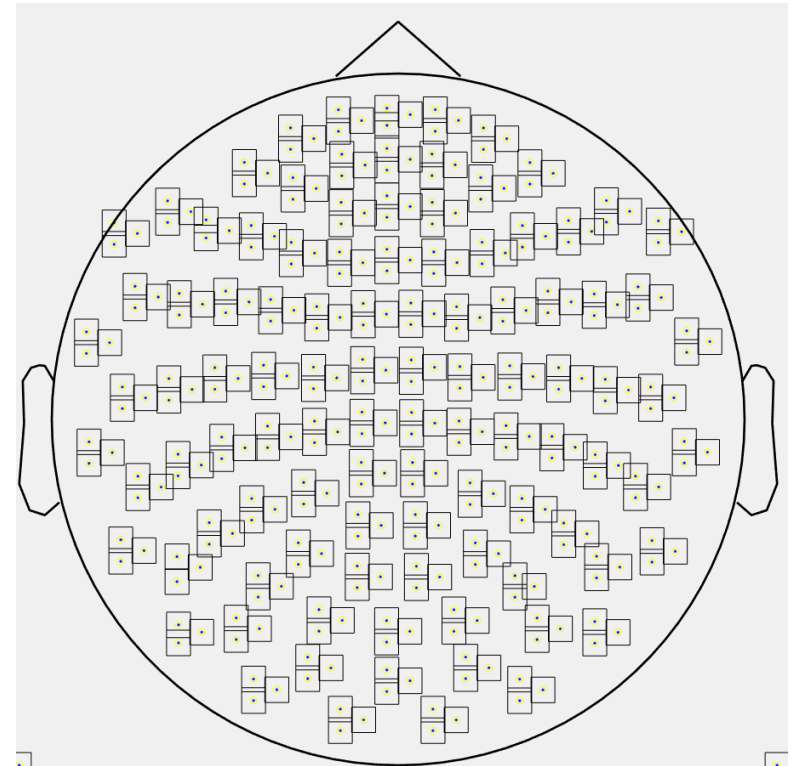
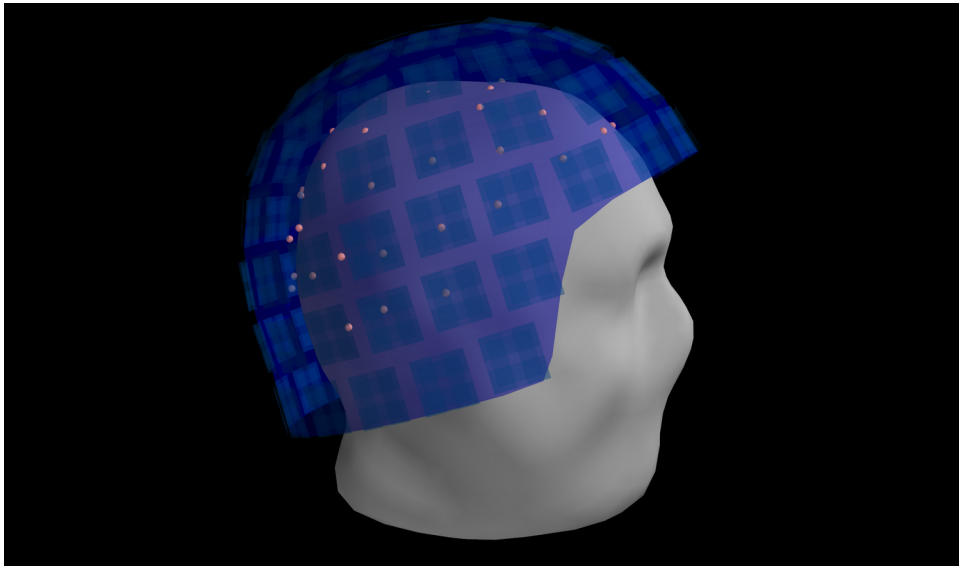
# How do we record it?



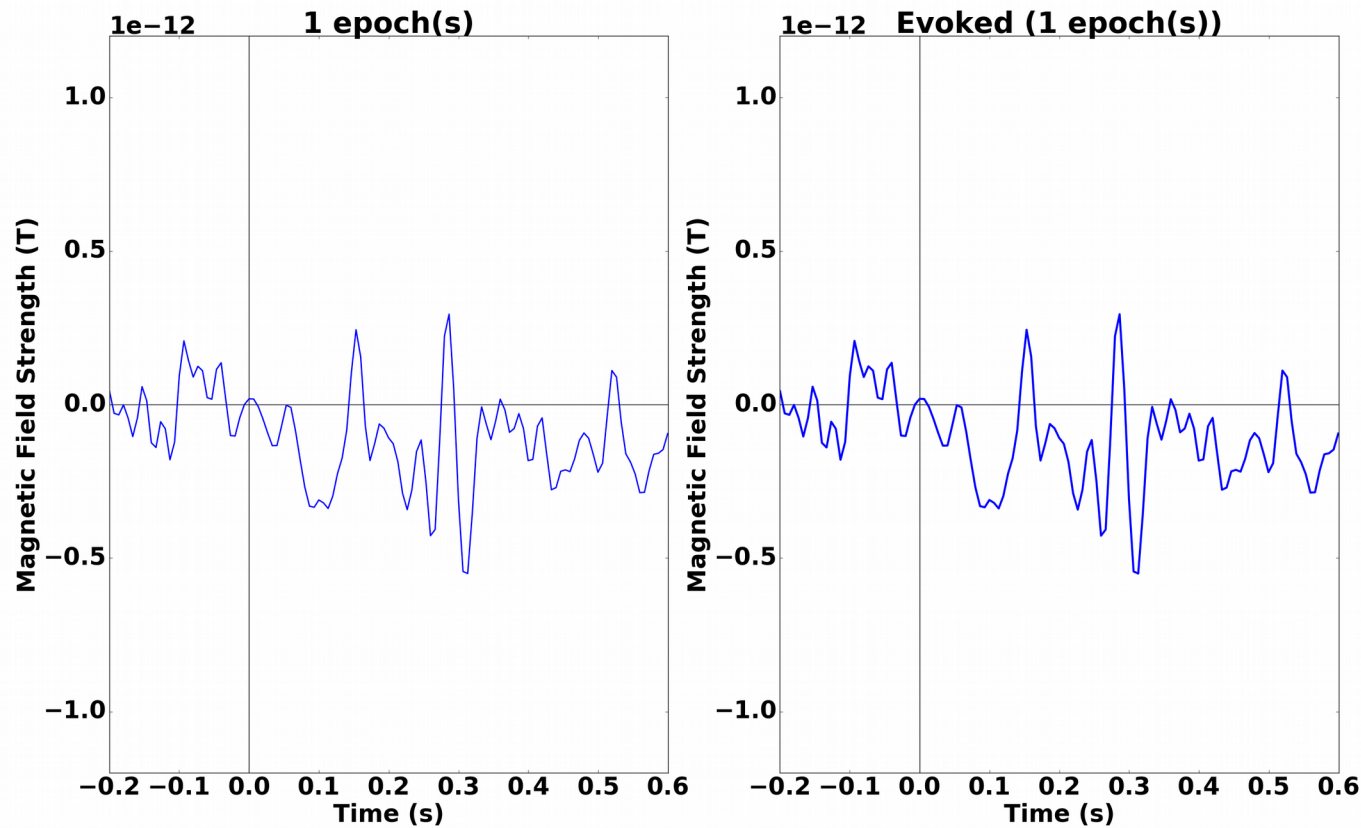


# What is MEG?

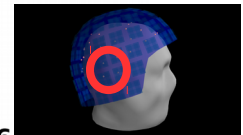
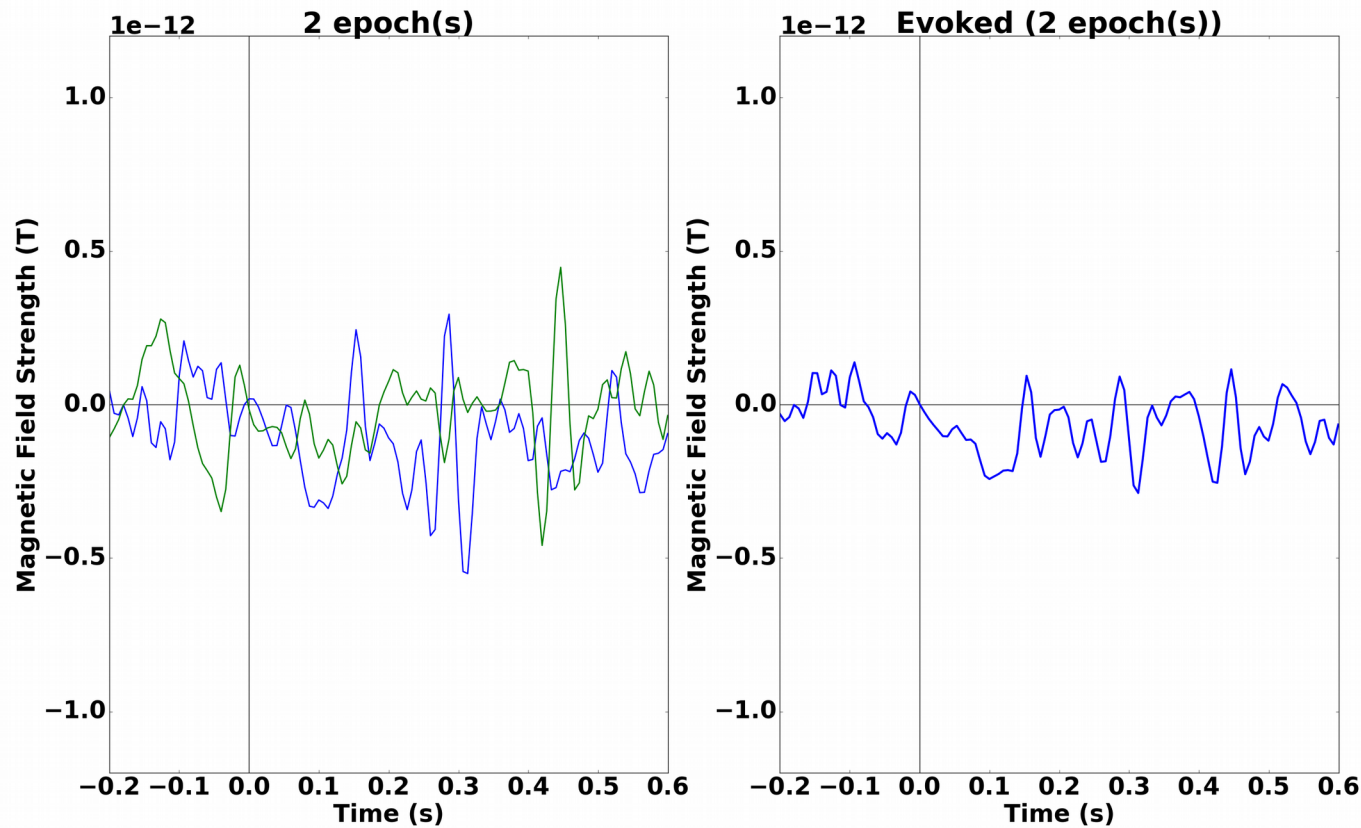
## How do we record it?



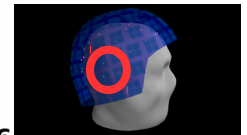
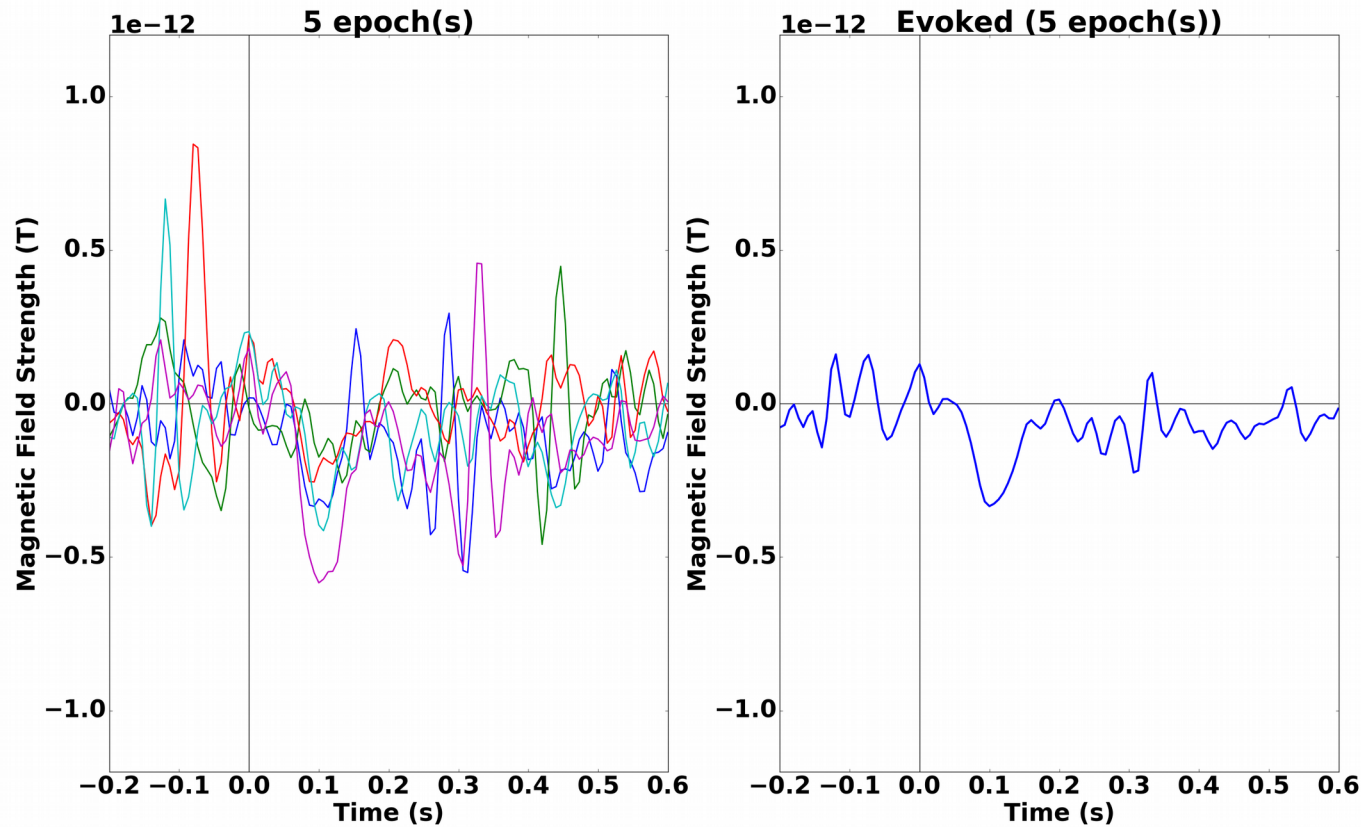
# EPOCHS AND EVOKED



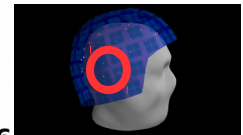
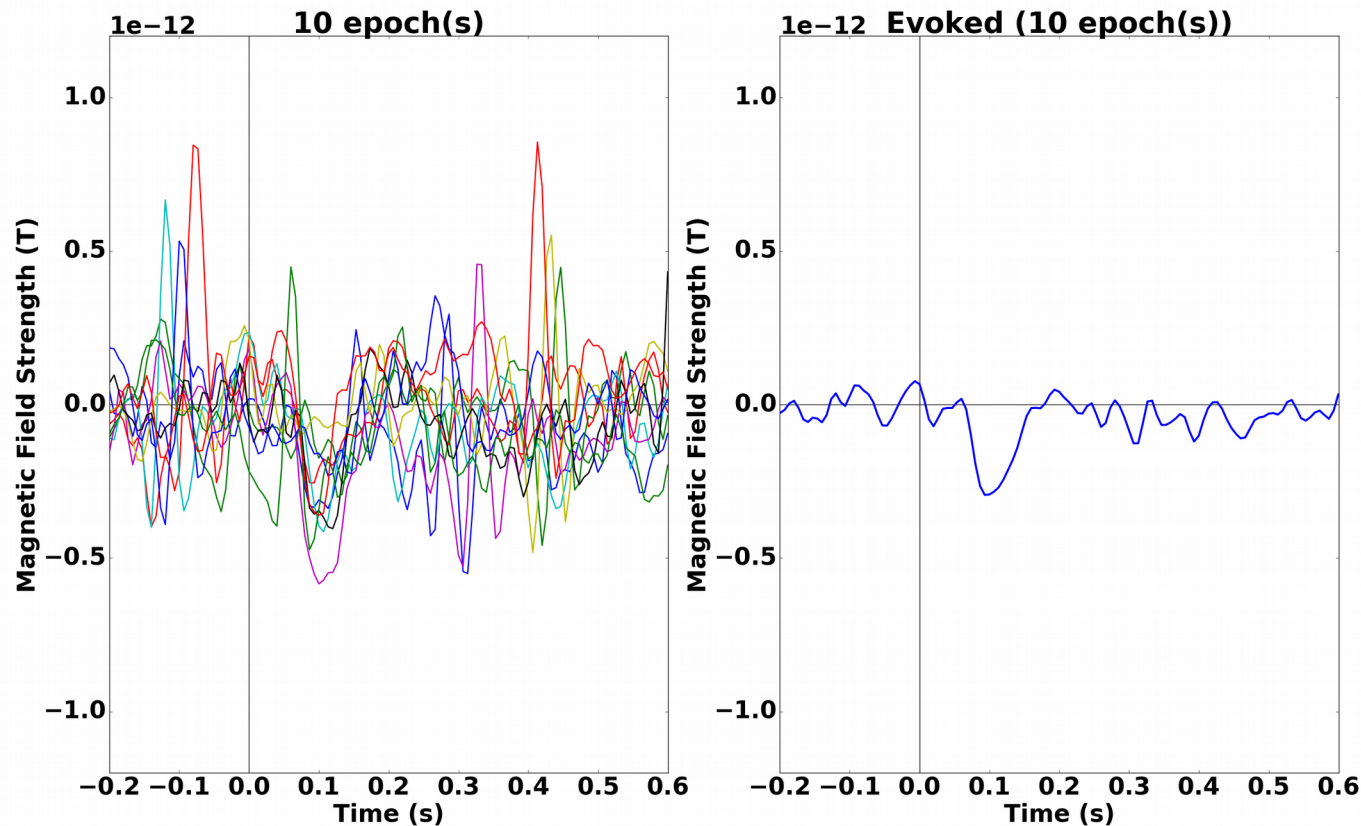
# EPOCHS AND EVOKED



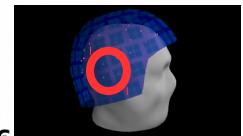
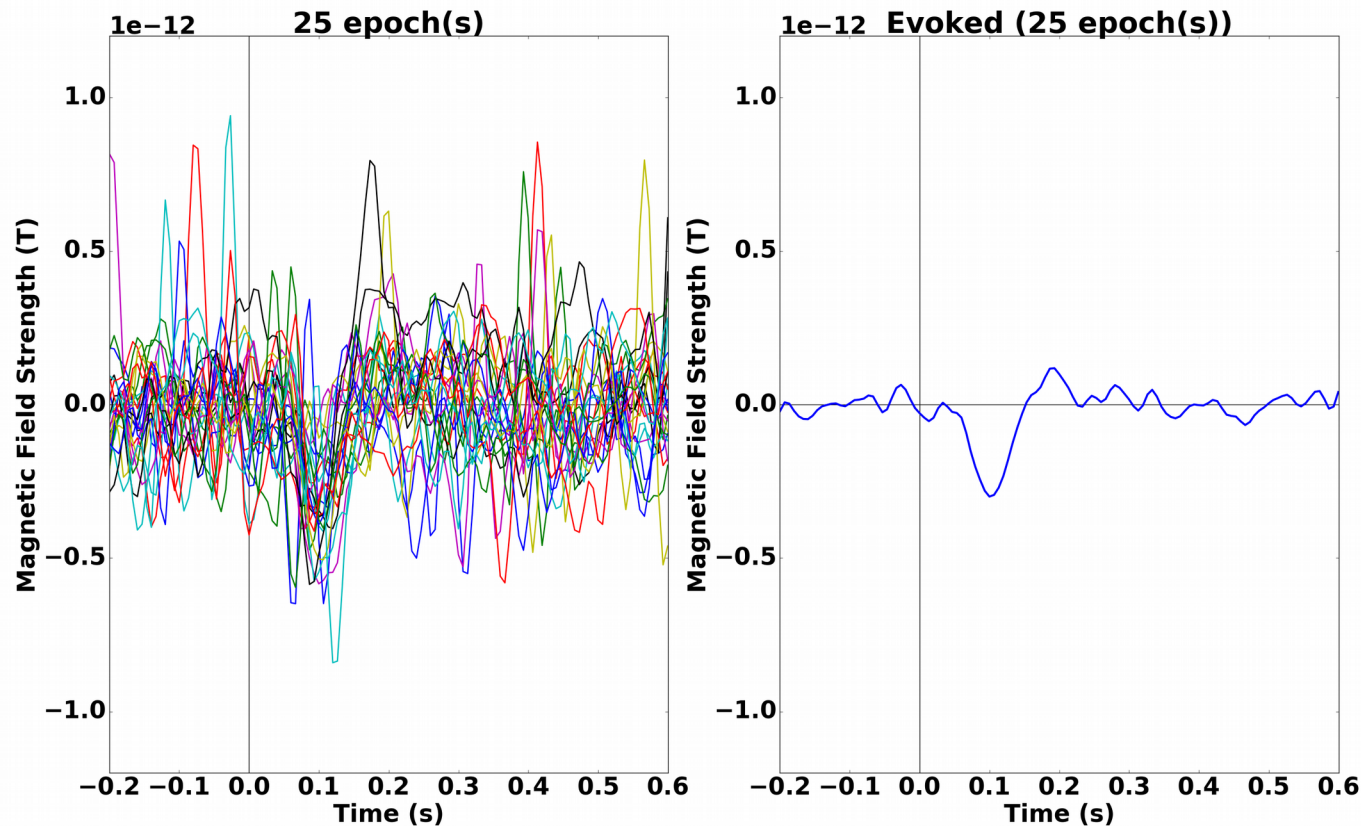
# EPOCHS AND EVOKED



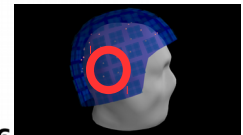
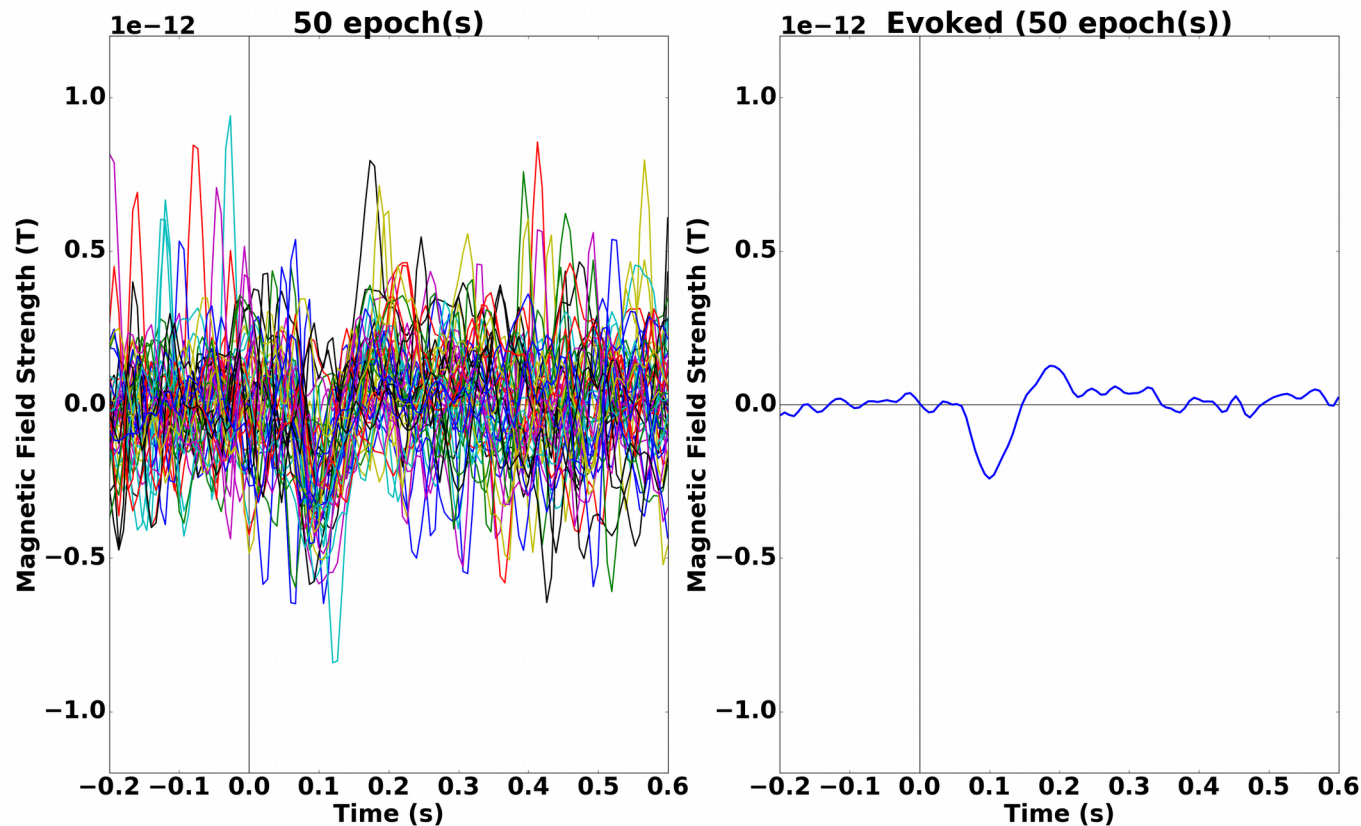
# EPOCHS AND EVOKED



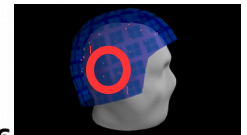
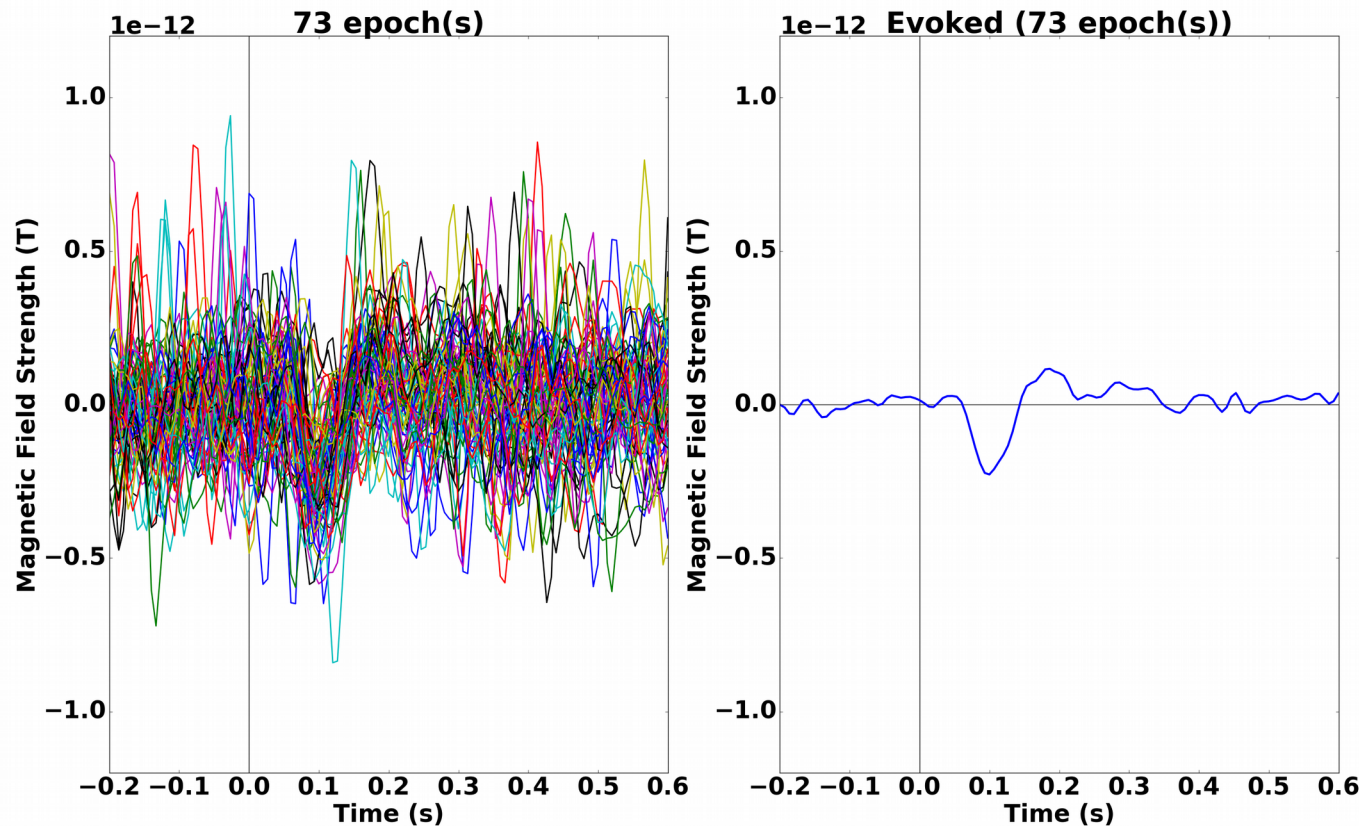
# EPOCHS AND EVOKED



# EPOCHS AND EVOKED

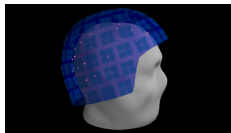
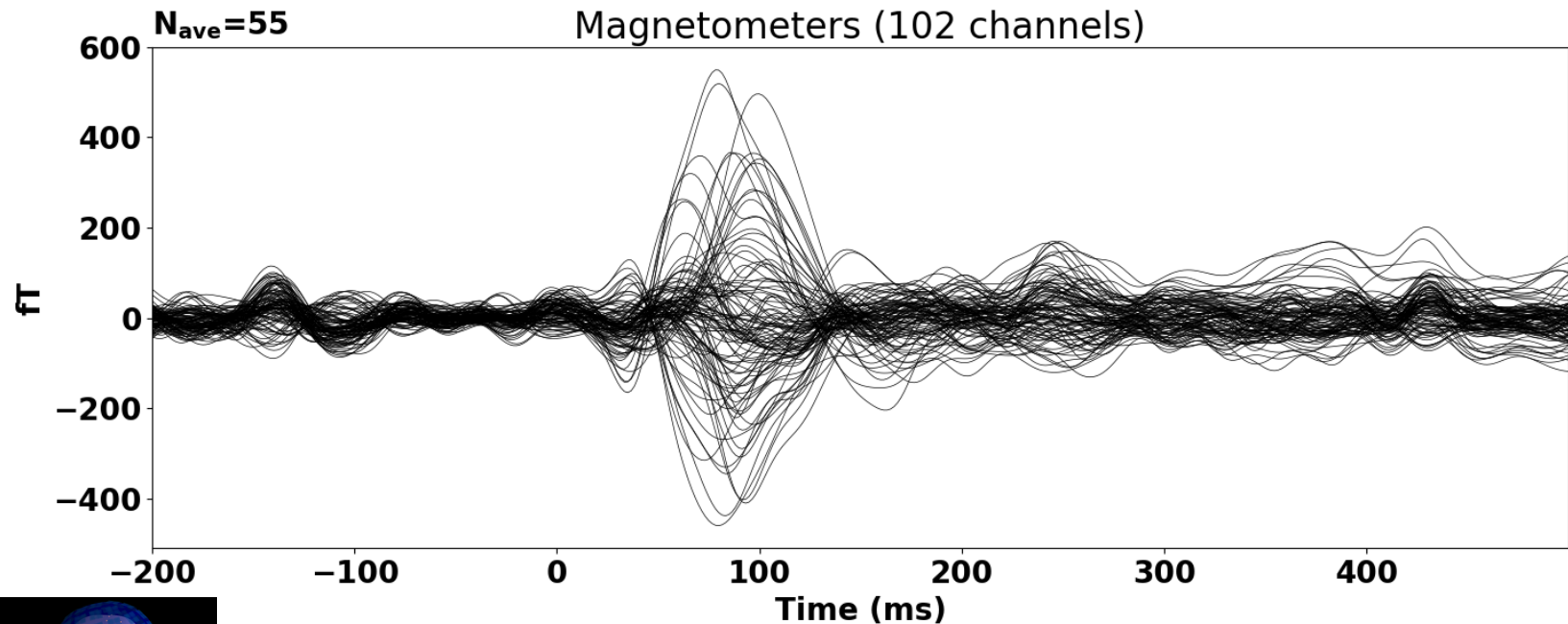


# EPOCHS AND EVOKED

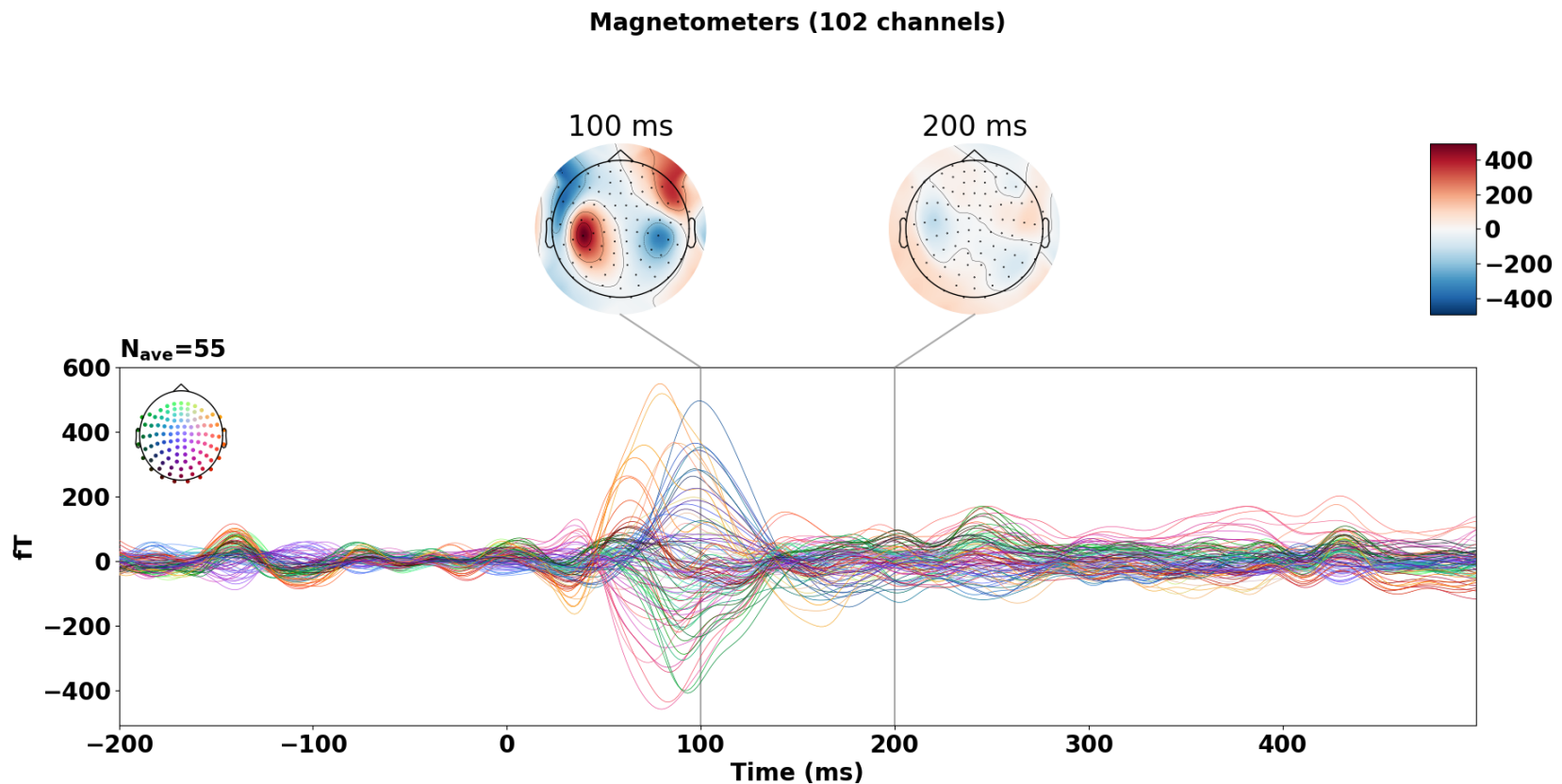




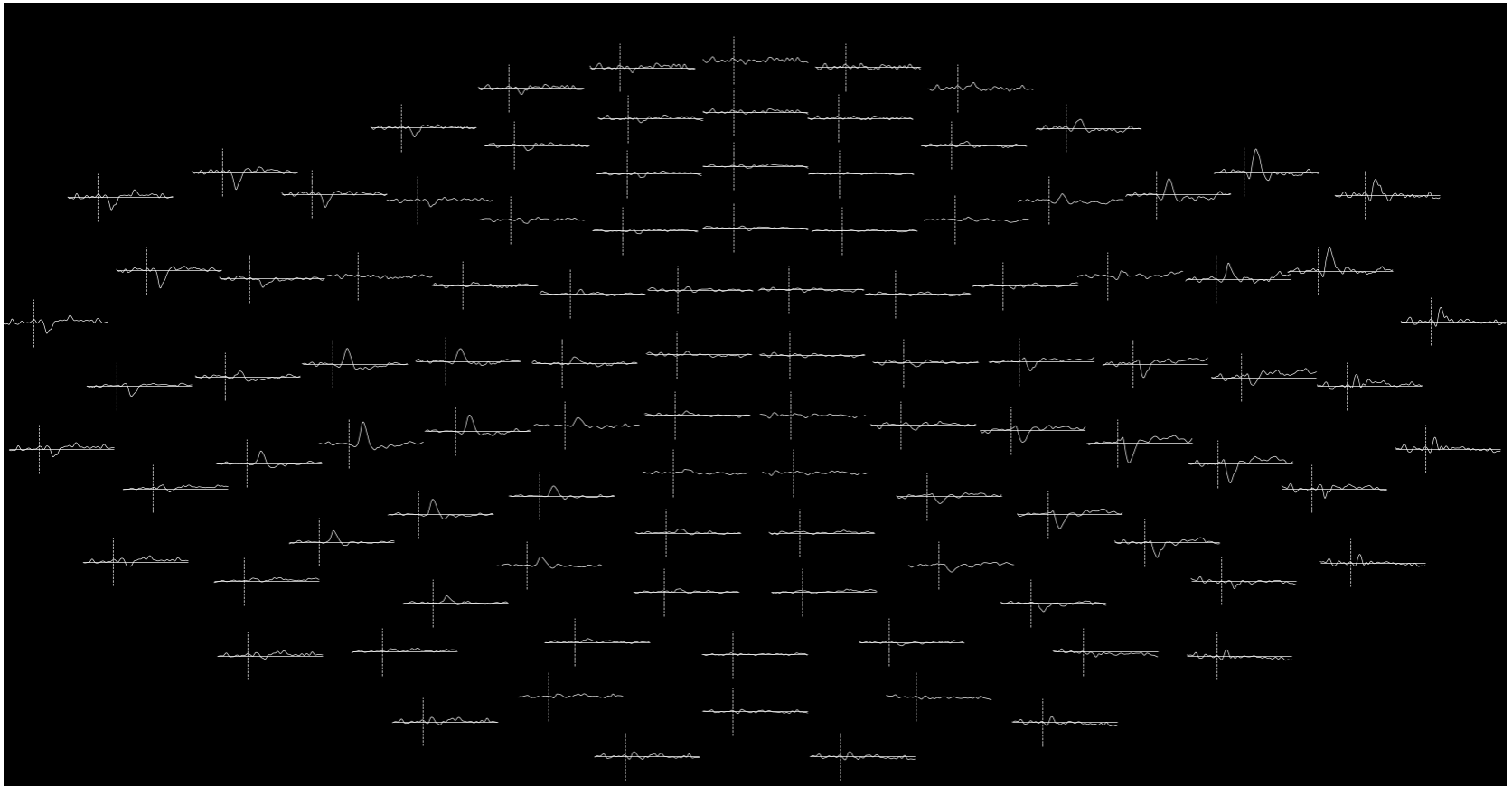
# What does the signal look like?



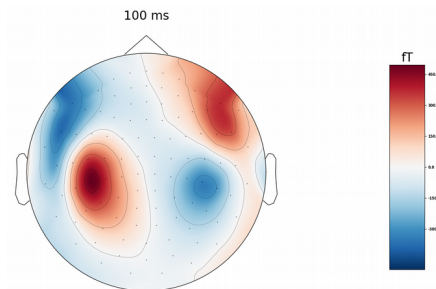
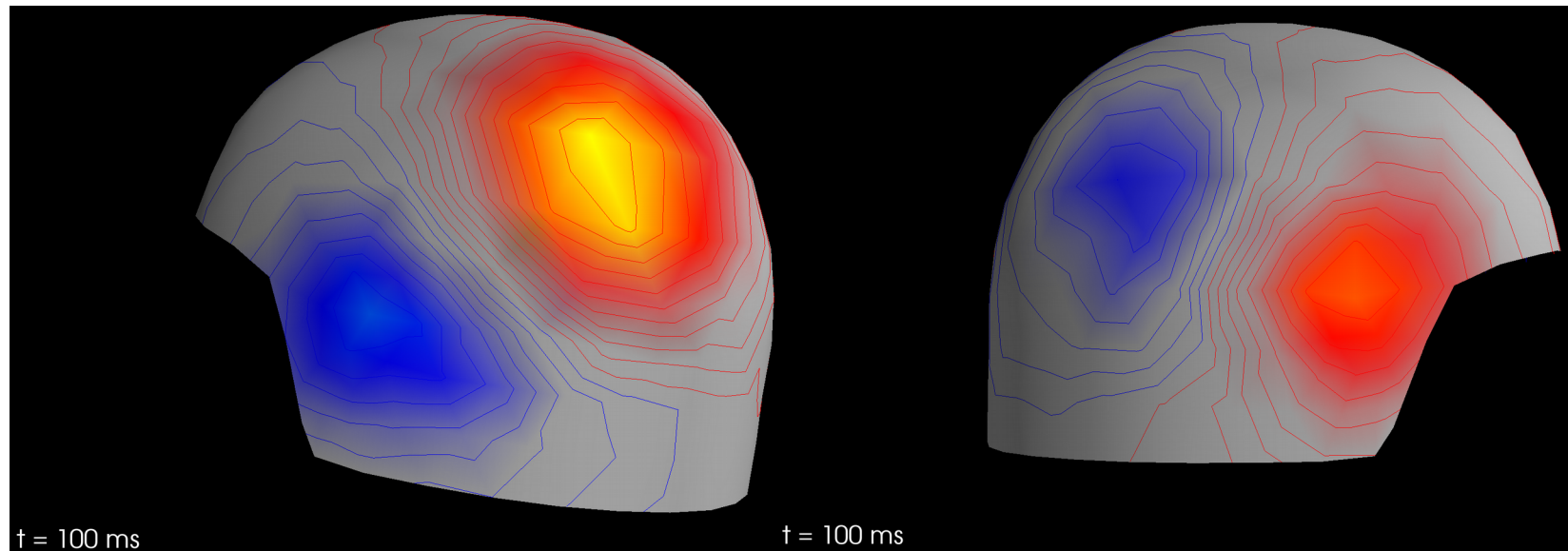
# What does the signal look like?



# What does the signal look like?

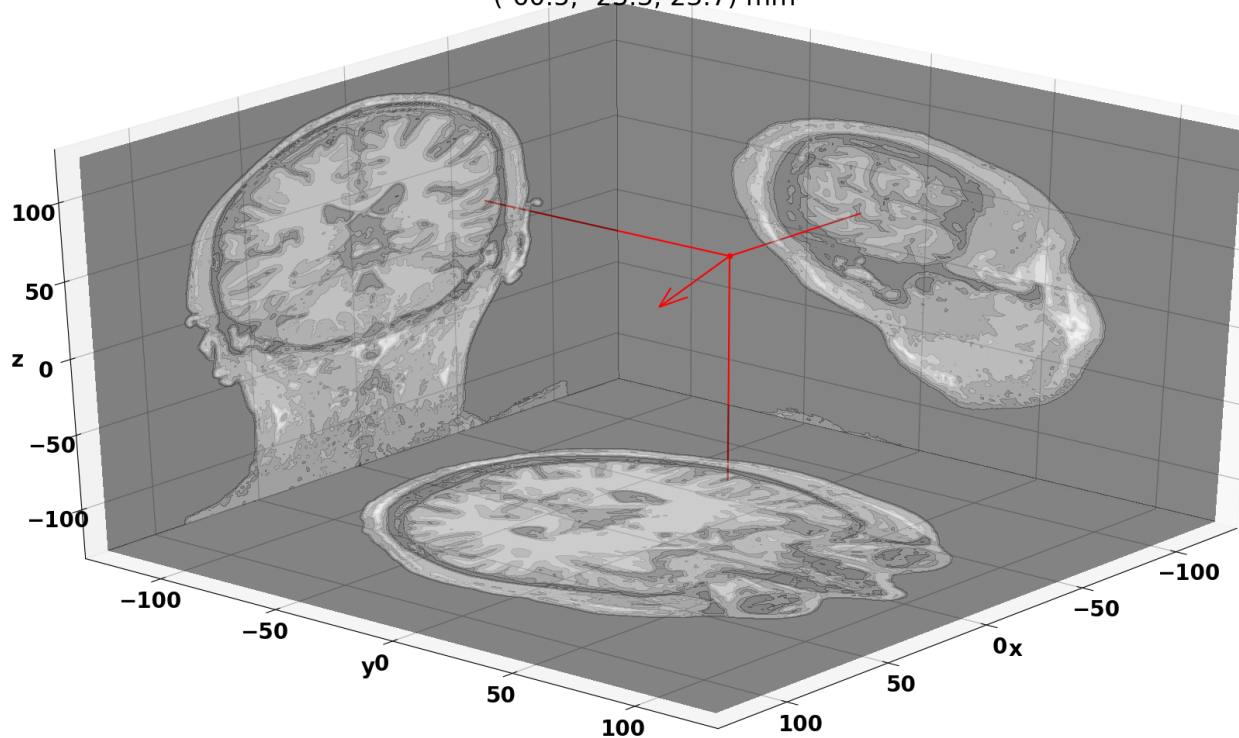


# What does the signal look like?



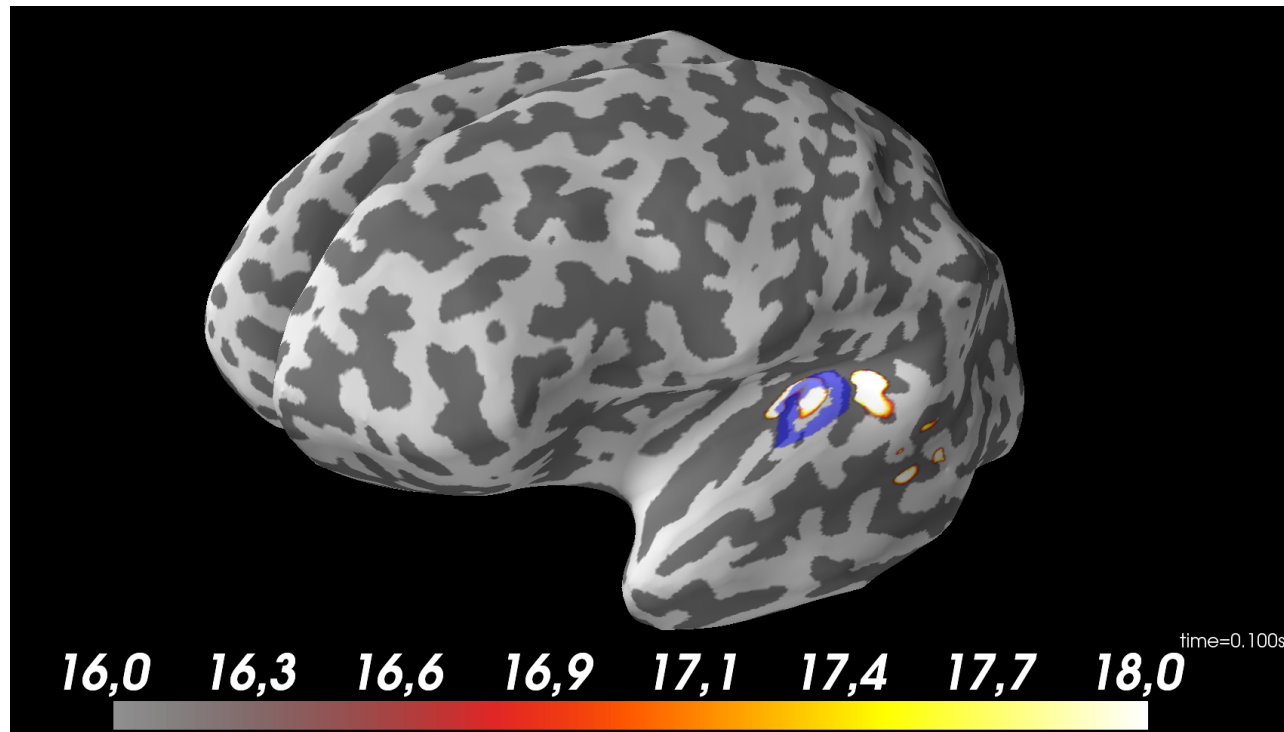
# What is its origin?

Dipole #1 / 1 @ 0.100s, GOF: 43.6%, 27.5nAm  
(-60.5, -23.3, 23.7) mm



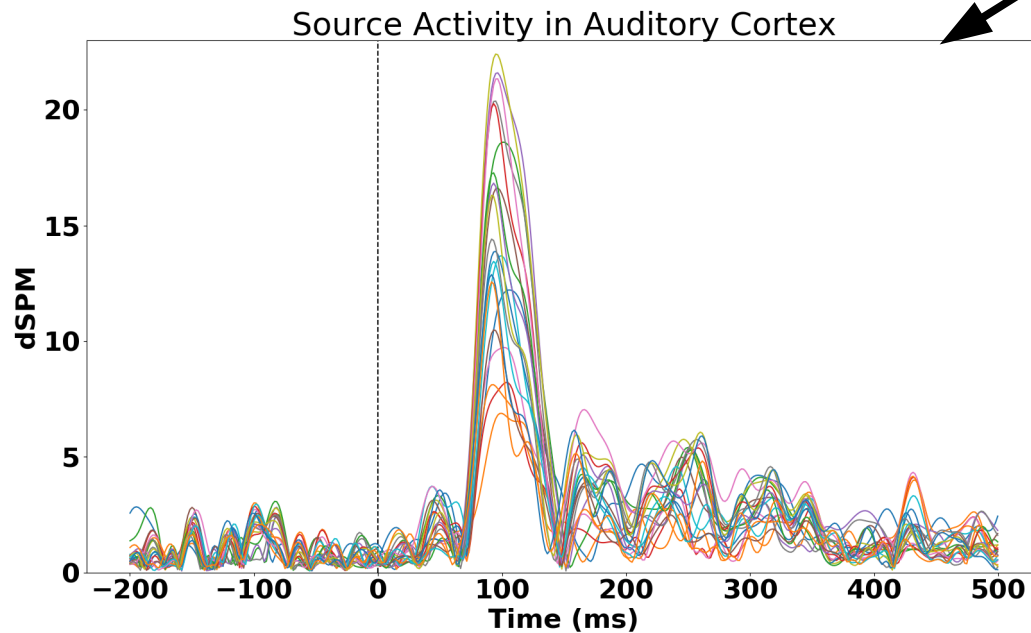
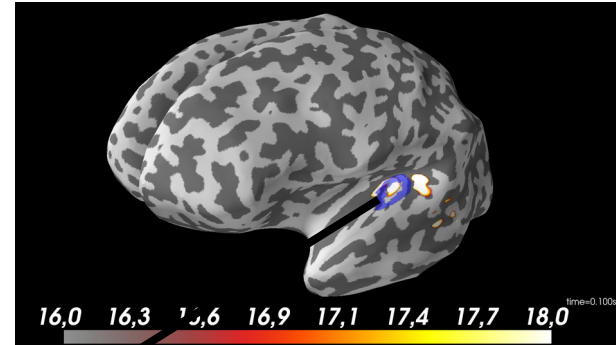
*What is MEG?*

# What is its origin?

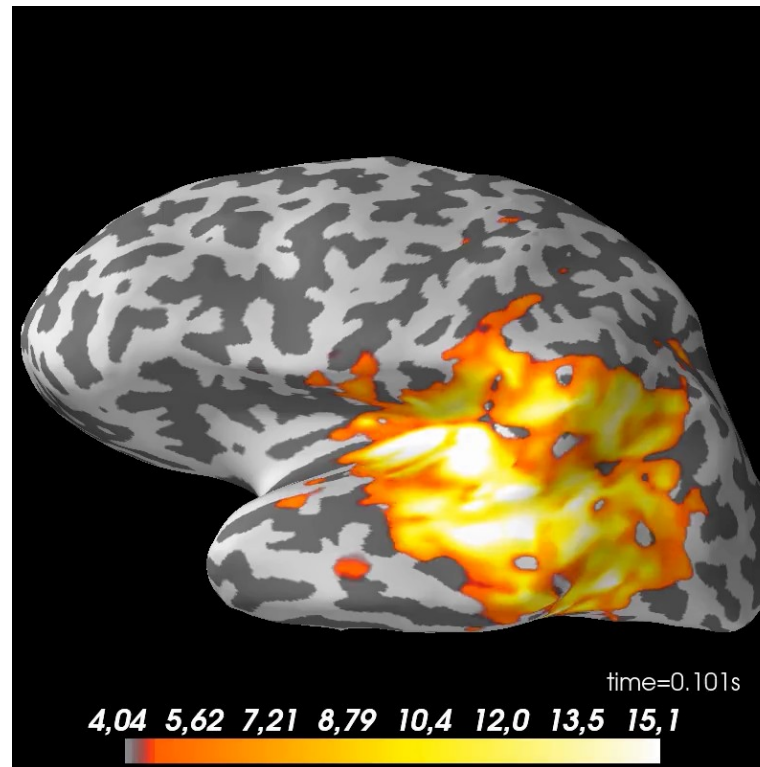


What is MEG?

# What is its origin?



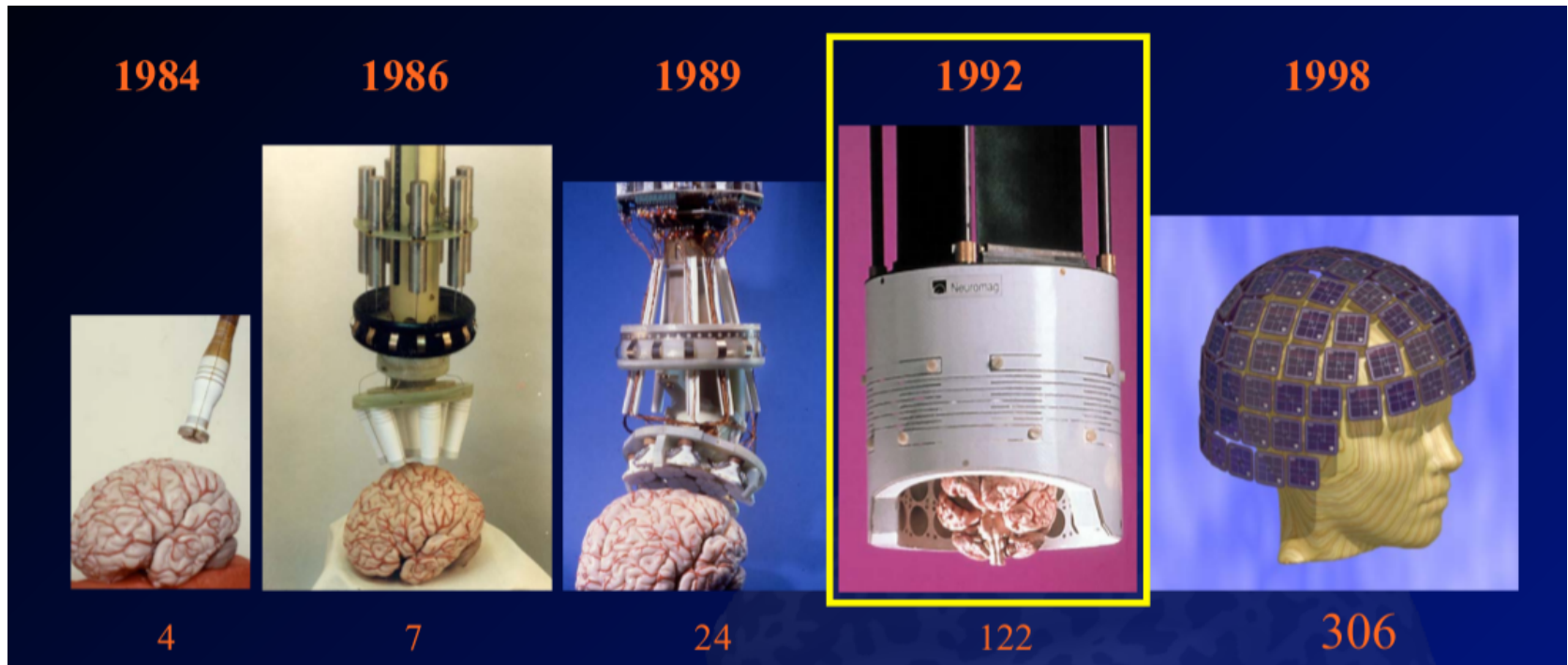
# What is its origin? (video)





# What is MEG?

## Some history



Credit: Matti Hämäläinen

# Some history

2016 – what existed?

- Single subject tutorials
- Recommendations paper

## Single subject tutorials

### Compare evoked responses for different conditions

In this example, an Epochs object for visual and auditory responses is created. Both conditions are then accessed by their respective names to create a sensor layout plot of the related evoked responses.

```
# Authors: Denis Engemann <denis.engemann@gmail.com>
#          Alexandre Gramfort <alexandre.gramfort@telecom-paristech.fr>

# License: BSD (3-clause)

import matplotlib.pyplot as plt
import mne

from mne.viz import plot_evoked_topo
from mne.datasets import sample

print(__doc__)

data_path = sample.data_path()
```

#### Set parameters

```
raw_fname = data_path + '/MEG/sample/sample_audvis_filt-0-40_raw.fif'
event_fname = data_path + '/MEG/sample/sample_audvis_filt-0-40_raw-eve.fif'
event_id = 1
tmin = -0.2
tmax = 0.5

# Setup for reading the raw data
raw = mne.io.read_raw_fif(raw_fname)
events = mne.read_events(event_fname)

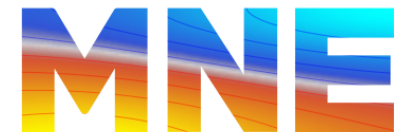
# Set up pick list: MEG + STI 014 - bad channels (modify to your needs)
include = [] # or stim channels ['STI 014']
# bad channels in raw.info['bads'] will be automatically excluded

# Set up amplitude-peak rejection values for MEG channels
reject = dict(grad=4000e-13, mag=4e-12)

# pick MEG channels
picks = mne.pick_types(raw.info, meg=True, eeg=False, stim=False, eog=True,
                       include=include, exclude='bads')
```

Download Python source  
code:  
plot\_topo\_compare\_conditions  
.py

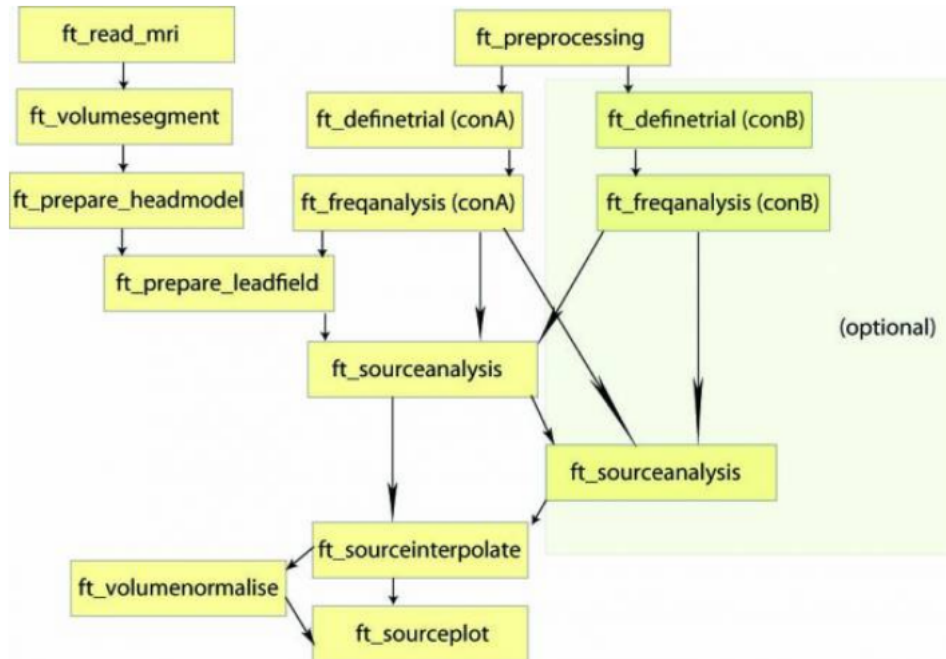
Download Jupyter  
notebook:  
plot\_topo\_compare\_conditions  
.ipynb



MEG + EEG ANALYSIS & VISUALIZATION

Open-source Python software for exploring, visualizing, and analyzing human neurophysiological data: MEG, EEG, sEEG, ECoG, and more.

## Single subject tutorials

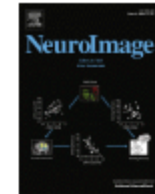


# Recommendations paper



NeuroImage

Volume 65, 15 January 2013, Pages 349-363



Comments and Controversies

## Good practice for conducting and reporting MEG research

Joachim Gross <sup>a</sup>  , Sylvain Baillet <sup>b</sup>, Gareth R. Barnes <sup>c</sup>, Richard N. Henson <sup>d</sup>, Arjan Hillebrand <sup>e</sup>, Ole Jensen <sup>f</sup>, Karim Jerbi <sup>g</sup>, Vladimir Litvak <sup>c</sup>, Burkhard Maess <sup>h</sup>, Robert Oostenveld <sup>f</sup>, Lauri Parkkonen <sup>i, j</sup>, Jason R. Taylor <sup>d</sup>, Virginie van Wassenhove <sup>k, l, m</sup>, Michael Wibral <sup>n</sup>, Jan-Mathijs Schoffelen <sup>f, o</sup>

 **Show more**

<https://doi.org/10.1016/j.neuroimage.2012.10.001>

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# Some history

2016 – what was missing?

- Group level tutorials
- Open data on which to apply them

# *How to analyse MEG?*



# How to analyse MEG?

Research Topic

## From raw MEG/EEG to publication: how to perform MEG/EEG group analysis with free academic software.

Comment

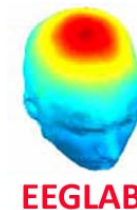
0

f 4    t 28    g+ 1    in 2    38

Submission closed.

Overview    **19** Articles    **72** Authors    Impact    Comments

VIEWS  
**42,827**




source Python software for exploring, visualizing, and analyzing neurophysiological data: MEG, EEG, sEEG, ECoG, and more.



SCIENTIFIC DATA 

Comment | [OPEN](#) | Published: 19 June 2018

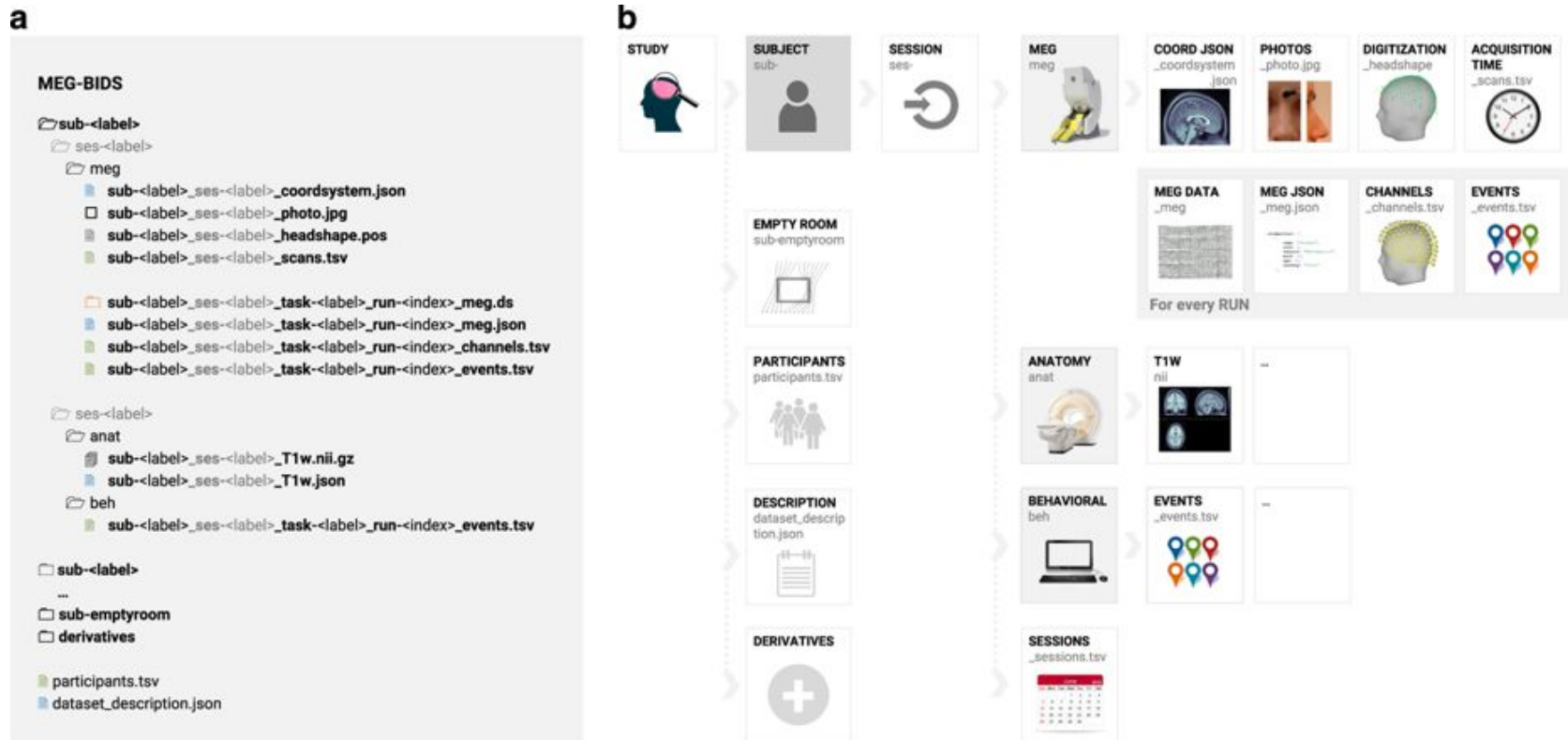
## MEG-BIDS, the brain imaging data structure extended to magnetoencephalography

Guiomar Niso, Krzysztof J. Gorgolewski, Elizabeth Bock, Teon L. Brooks, Guillaume Flandin, Alexandre Gramfort, Richard N. Henson, Mainak Jas, Vladimir Litvak, Jeremy T. Moreau, Robert Oostenveld, Jan-Mathijs Schoffelen, Francois Tadel, Joseph Wexler & Sylvain Baillet 

*Scientific Data* **5**, Article number: 180110 (2018) | [Download Citation](#) 

# How to analyse MEG?

# MEG-BIDS



Niso et al. 2018, *Scientific data*

# DATA REPOSITORIES

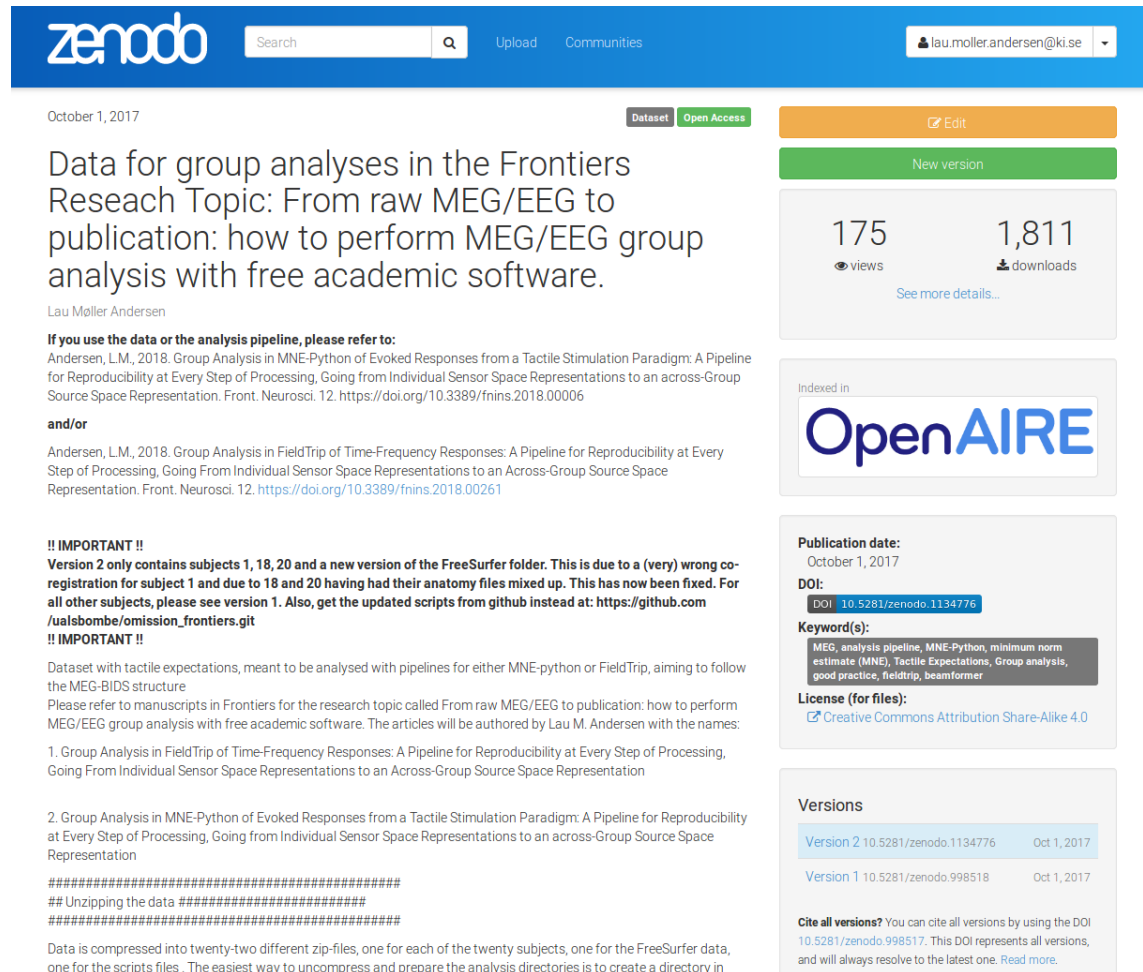


**OpenNEURO**<sup>BETA</sup>

A free and open platform for analyzing  
and sharing neuroimaging data

# How to analyse MEG?

# DATA



The screenshot shows a Zenodo dataset page for a research topic. The page includes a search bar, user profile, and various action buttons like 'Edit' and 'New version'. It displays statistics such as 175 views and 1,811 downloads. The main title is 'Data for group analyses in the Frontiers Reseach Topic: From raw MEG/EEG to publication: how to perform MEG/EEG group analysis with free academic software.' The author is Lau Møller Andersen. The page contains two important notices regarding subject registration and script updates. It also lists two publications related to the dataset and provides a list of versions with their respective DOIs and dates. A license of Creative Commons Attribution Share-Alike 4.0 is mentioned.

zenodo Search Upload Communities lau.moller.andersen@ki.se

October 1, 2017 Dataset Open Access Edit

## Data for group analyses in the Frontiers Reseach Topic: From raw MEG/EEG to publication: how to perform MEG/EEG group analysis with free academic software.

Lau Møller Andersen

**If you use the data or the analysis pipeline, please refer to:**  
Andersen, L.M., 2018. Group Analysis in MNE-Python of Evoked Responses from a Tactile Stimulation Paradigm: A Pipeline for Reproducibility at Every Step of Processing, Going from Individual Sensor Space Representations to an across-Group Source Space Representation. *Front. Neurosci.* 12. <https://doi.org/10.3389/fnins.2018.00006>  
**and/or**  
Andersen, L.M., 2018. Group Analysis in FieldTrip of Time-Frequency Responses: A Pipeline for Reproducibility at Every Step of Processing, Going From Individual Sensor Space Representations to an Across-Group Source Space Representation. *Front. Neurosci.* 12. <https://doi.org/10.3389/fnins.2018.00261>

**!! IMPORTANT !!**  
**Version 2 only contains subjects 1, 18, 20 and a new version of the FreeSurfer folder. This is due to a (very) wrong co-registration for subject 1 and due to 18 and 20 having had their anatomy files mixed up. This has now been fixed. For all other subjects, please see version 1. Also, get the updated scripts from github instead at: [https://github.com/ualsbombe/omission\\_frontiers.git](https://github.com/ualsbombe/omission_frontiers.git)**  
**!! IMPORTANT !!**

Dataset with tactile expectations, meant to be analysed with pipelines for either MNE-python or FieldTrip, aiming to follow the MEG-BIDS structure  
Please refer to manuscripts in Frontiers for the research topic called From raw MEG/EEG to publication: how to perform MEG/EEG group analysis with free academic software. The articles will be authored by Lau M. Andersen with the names:

1. Group Analysis in FieldTrip of Time-Frequency Responses: A Pipeline for Reproducibility at Every Step of Processing, Going From Individual Sensor Space Representations to an Across-Group Source Space Representation
2. Group Analysis in MNE-Python of Evoked Responses from a Tactile Stimulation Paradigm: A Pipeline for Reproducibility at Every Step of Processing, Going from Individual Sensor Space Representations to an across-Group Source Space Representation

```
#####  
## Unzipping the data #####  
#####
```

Data is compressed into twenty-two different zip-files, one for each of the twenty subjects, one for the FreeSurfer data, one for the scripts files . The easiest way to uncompress and prepare the analysis directories is to create a directory in

175 views 1,811 downloads See more details...

Indexed in OpenAIRE

**Publication date:** October 1, 2017  
**DOI:** DOI 10.5281/zenodo.1134776  
**Keyword(s):** MEG, analysis pipeline, MNE-Python, minimum norm estimate (MNE), Tactile Expectations, Group analysis, good practice, fieldtrip, beamformer  
**License (for files):** Creative Commons Attribution Share-Alike 4.0

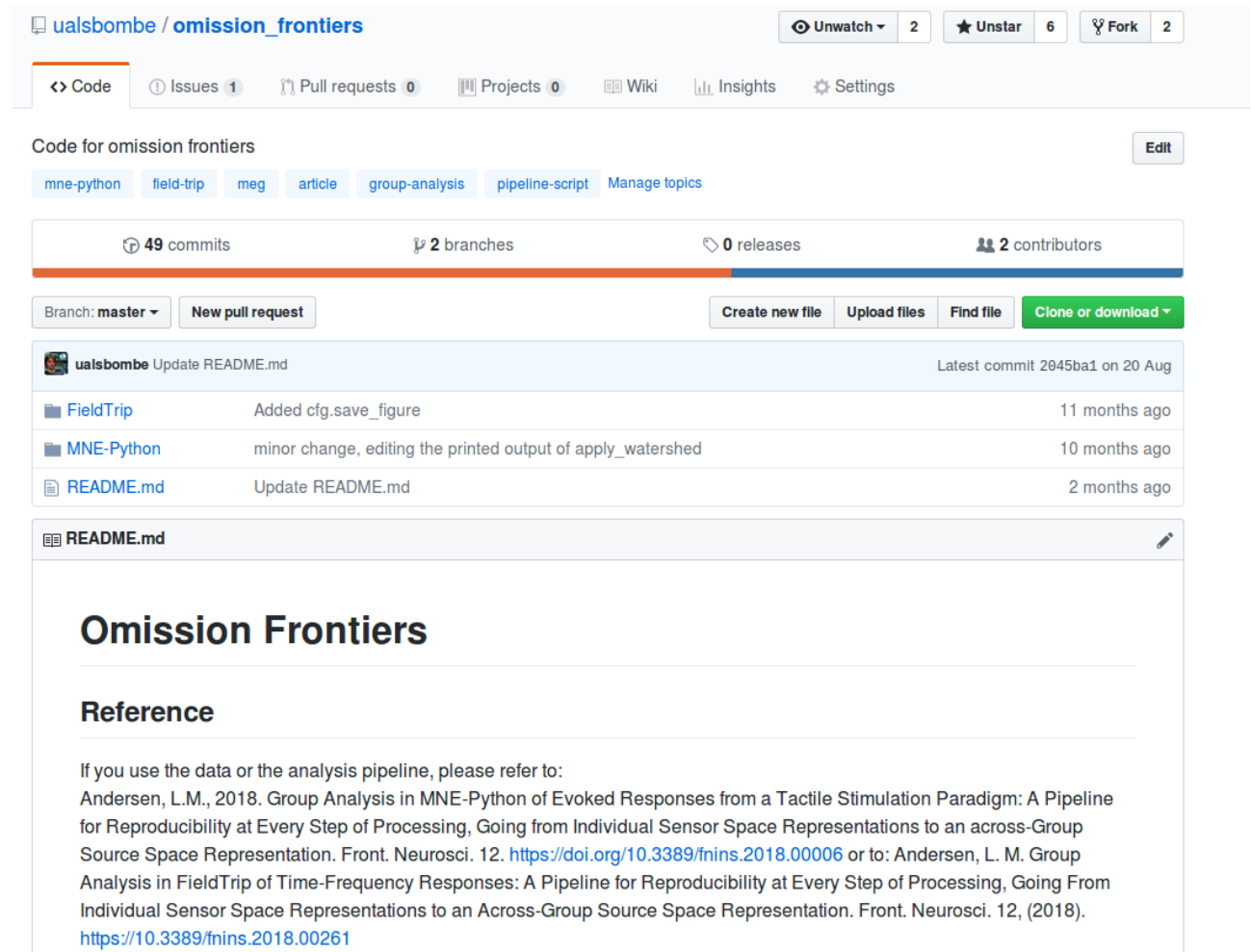
**Versions**

Version	DOI	Date
Version 2	10.5281/zenodo.1134776	Oct 1, 2017
Version 1	10.5281/zenodo.998518	Oct 1, 2017

**Cite all versions?** You can cite all versions by using the DOI 10.5281/zenodo.998517. This DOI represents all versions, and will always resolve to the latest one. [Read more.](#)

# How to analyse MEG?

## CODE



The screenshot shows a GitHub repository page for 'omission\_frontiers' by user 'ualsbombe'. The repository has 49 commits, 2 branches, 0 releases, and 2 contributors. The current branch is 'master'. The repository contains several files: 'FieldTrip', 'MNE-Python', and 'README.md'. The 'README.md' file is selected and its content is displayed below. The content of the README includes a title 'Omission Frontiers', a 'Reference' section, and a paragraph of text providing information about the data and analysis pipeline, along with two URLs for further reading.

ualsbombe / omission\_frontiers

Unwatch 2 Unstar 6 Fork 2

<> Code Issues 1 Pull requests 0 Projects 0 Wiki Insights Settings

Code for omission frontiers Edit

mne-python field-trip meg article group-analysis pipeline-script Manage topics

49 commits 2 branches 0 releases 2 contributors

Branch: master New pull request Create new file Upload files Find file Clone or download

ualsbombe Update README.md Latest commit 2045ba1 on 20 Aug

- FieldTrip Added cfg.save\_figure 11 months ago
- MNE-Python minor change, editing the printed output of apply\_watershed 10 months ago
- README.md Update README.md 2 months ago

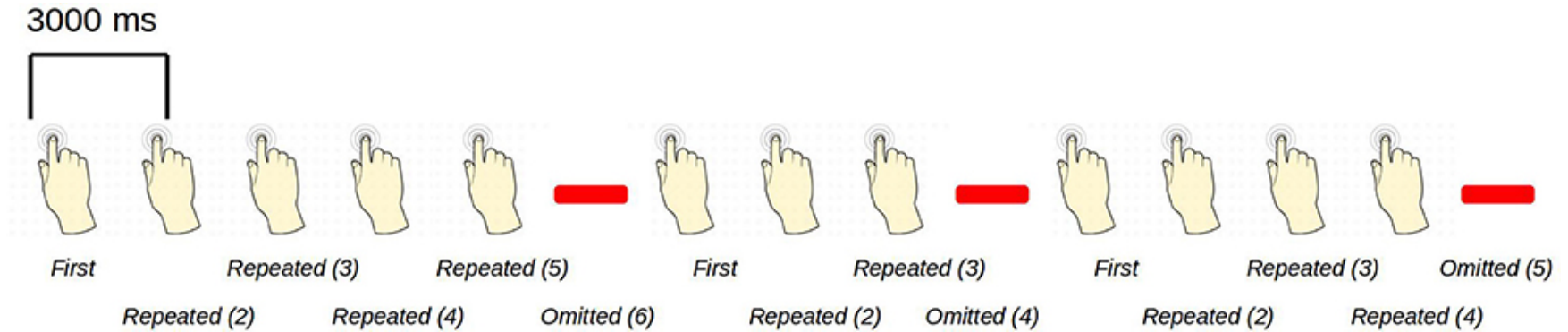
README.md

### Omission Frontiers

#### Reference

If you use the data or the analysis pipeline, please refer to:  
Andersen, L.M., 2018. Group Analysis in MNE-Python of Evoked Responses from a Tactile Stimulation Paradigm: A Pipeline for Reproducibility at Every Step of Processing, Going from Individual Sensor Space Representations to an across-Group Source Space Representation. *Front. Neurosci.* 12. <https://doi.org/10.3389/fnins.2018.00006> or to: Andersen, L. M. Group Analysis in FieldTrip of Time-Frequency Responses: A Pipeline for Reproducibility at Every Step of Processing, Going From Individual Sensor Space Representations to an Across-Group Source Space Representation. *Front. Neurosci.* 12, (2018). <https://10.3389/fnins.2018.00261>

## PARADIGM



Andersen 2018, *Frontiers in Neuroscience*

# QUESTION

Do *Stimulations* give rise to more activity than *Non-Stimulations* in the primary somatosensory cortex?

# How to analyse MEG?



NeuroImage

Volume 184, 1 January 2019, Pages 78-89



## Somatosensory responses to nothing: An MEG study of expectations during omission of tactile stimulations

Lau M. Andersen  , Daniel Lundqvist

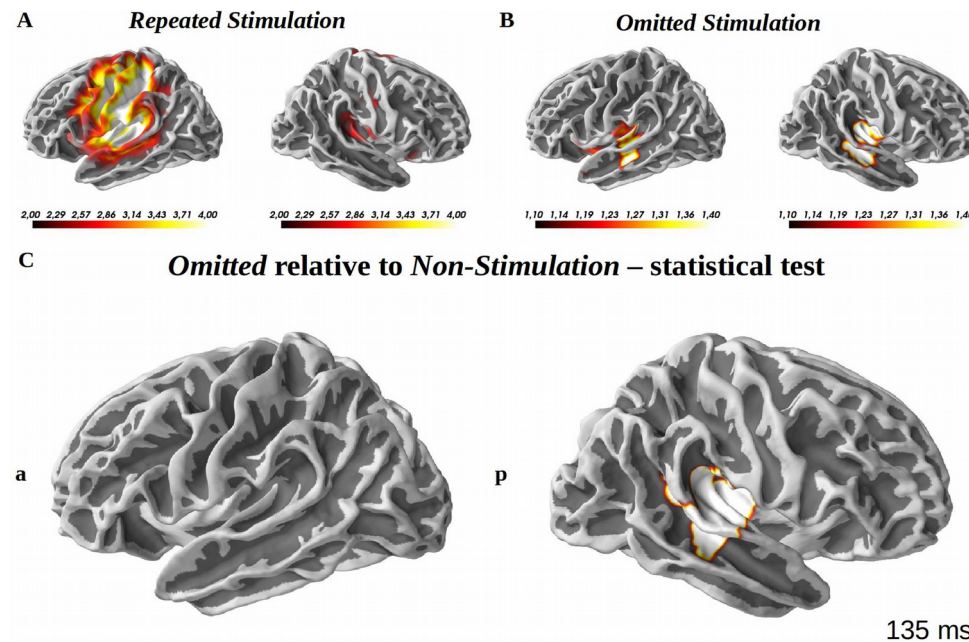
 [Show more](#)

<https://doi.org/10.1016/j.neuroimage.2018.09.014>

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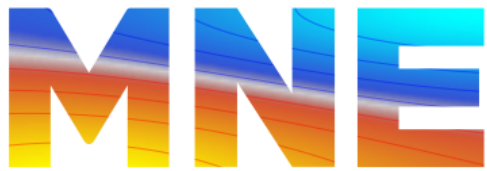
## SUBJECTS

```
subjects = [  
    'sub-01',  
    'sub-02',  
    'sub-03',  
    'sub-04',  
    'sub-05',  
    'sub-06',  
    'sub-07',  
    'sub-08',  
    'sub-09',  
    'sub-10',  
    'sub-11',  
    'sub-12',  
    'sub-13',  
    'sub-14',  
    'sub-15',  
    'sub-16',  
    'sub-17',  
    'sub-18',  
    'sub-19',  
    'sub-20'  
]
```

# 4 GUIDING PRINCIPLES FOR A PIPELINE

1. Should divide output files into folders belonging to the respective subjects and recordings
2. Should apply operations across a group of subjects
3. Should make it possible to (re)start the analysis at any intermediate point by saving output for each intermediate point
4. Should allow for plotting the results in a way that allows for changing the figures in a principled, but flexible manner

## TWO PIPELINES



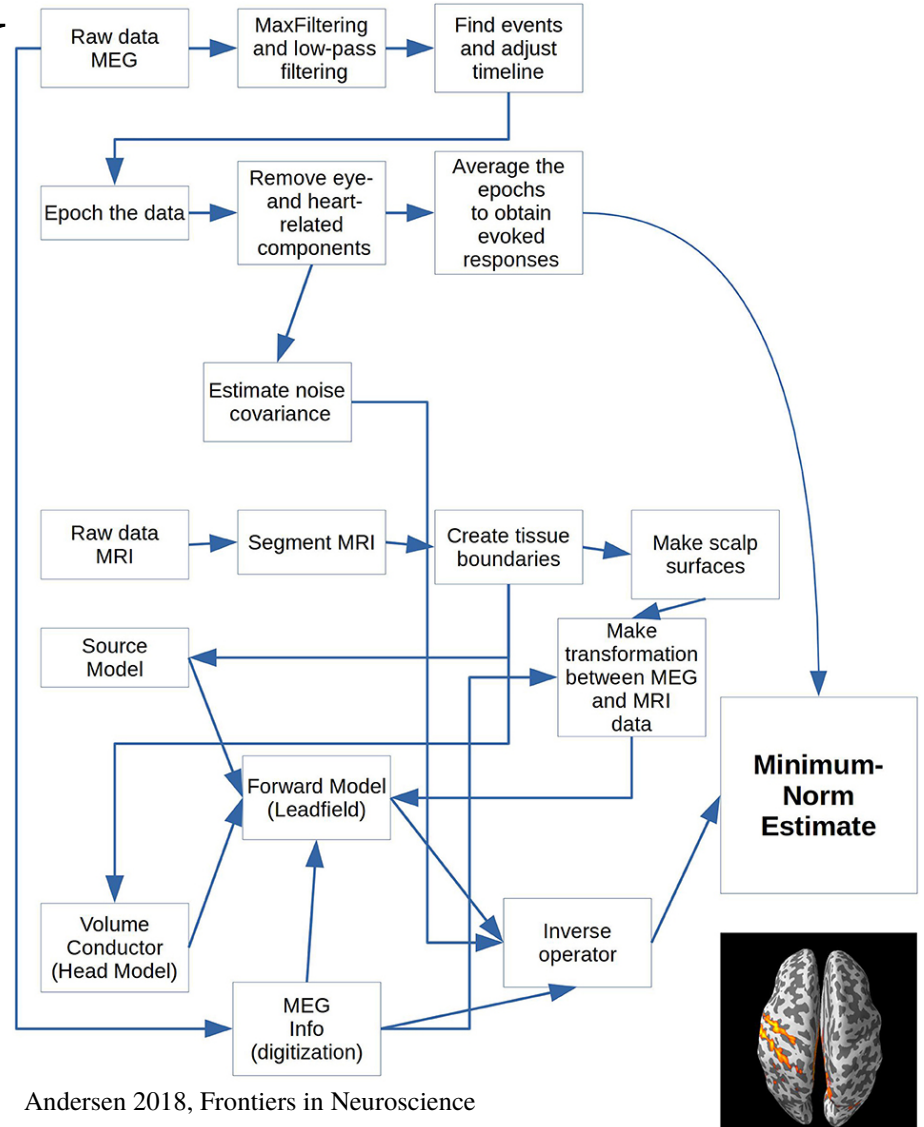
MEG + EEG ANALYSIS & VISUALIZATION

Open-source Python software for exploring, visualizing, and analyzing human neurophysiological data: MEG, EEG, sEEG, ECoG, and more.



# How to analyse MEG?

# MNE-PYTHON PIPELINE



Andersen 2018, Frontiers in Neuroscience

# How to analyse MEG?

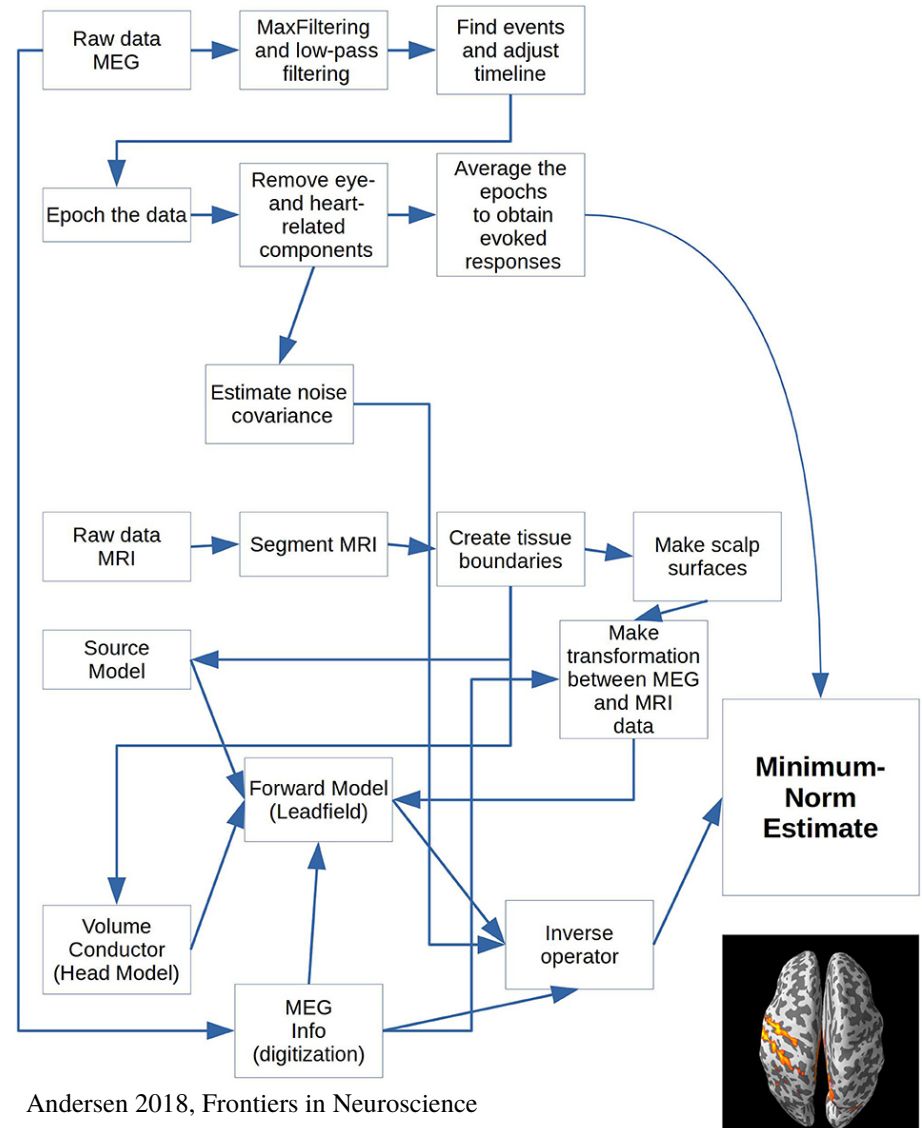
## OPERATIONS

### ## sensor space operations

```
filter_raw=0,  
find_events=0,  
epoch_raw=0,  
run_ica=0,  
apply_ica=0,  
get_evoked=0,
```

### ## source space operations

```
import_mri=0,  
segment_mri=0, # long process (>6 h)  
apply_watershed=0,  
make_source_space=0,  
make_dense_scalp_surfaces=0,  
make_bem_solutions=0,  
create_forward_solution=0,  
estimate_noise_covariance=0,  
create_inverse_operator=0,  
source_estimate=0,  
morph_to_fsaverage=0,
```



Andersen 2018, Frontiers in Neuroscience

## OPERATIONS

```
## sensor space operations
filter_raw=0,
find_events=0,
epoch_raw=1,
run_ica=0,
apply_ica=0,
get_evoked=0,

def epoch_raw(name, save_dir, lowpass, event_id, tmin, tmax,
              baseline, reject, bad_channels, decim, overwrite):

    epochs_name = name + filter_string(lowpass) + '-epo.fif'
    epochs_path = join(save_dir, epochs_name)
    if overwrite or not isfile(epochs_path):

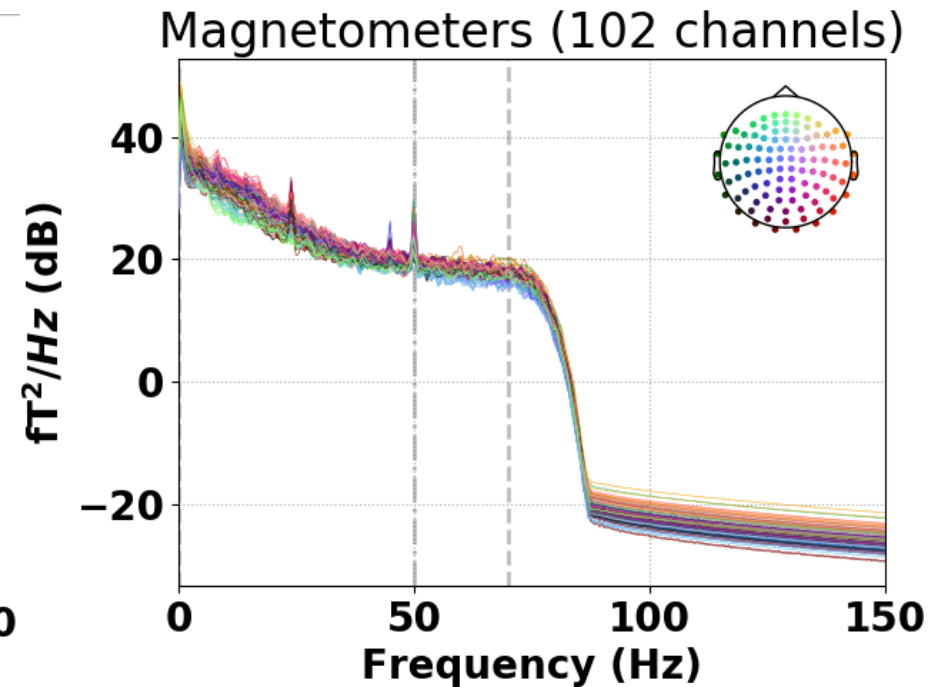
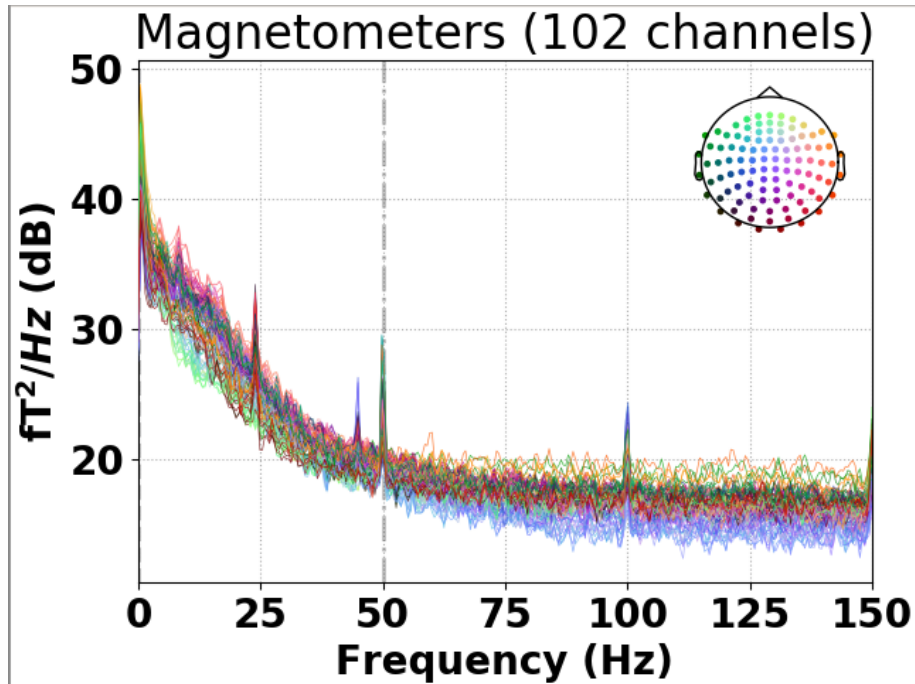
        events = io.read_events(name, save_dir)
        raw = io.read_filtered(name, save_dir, lowpass,)
        raw.info['bads'] = bad_channels
        picks = mne.pick_types(raw.info, meg=True, eog=True, ecg=True,
                               exclude='bads')

        epochs = mne.Epochs(raw, events, event_id, tmin, tmax, baseline,
                             reject=reject, preload=True, picks=picks,
                             decim=decim)

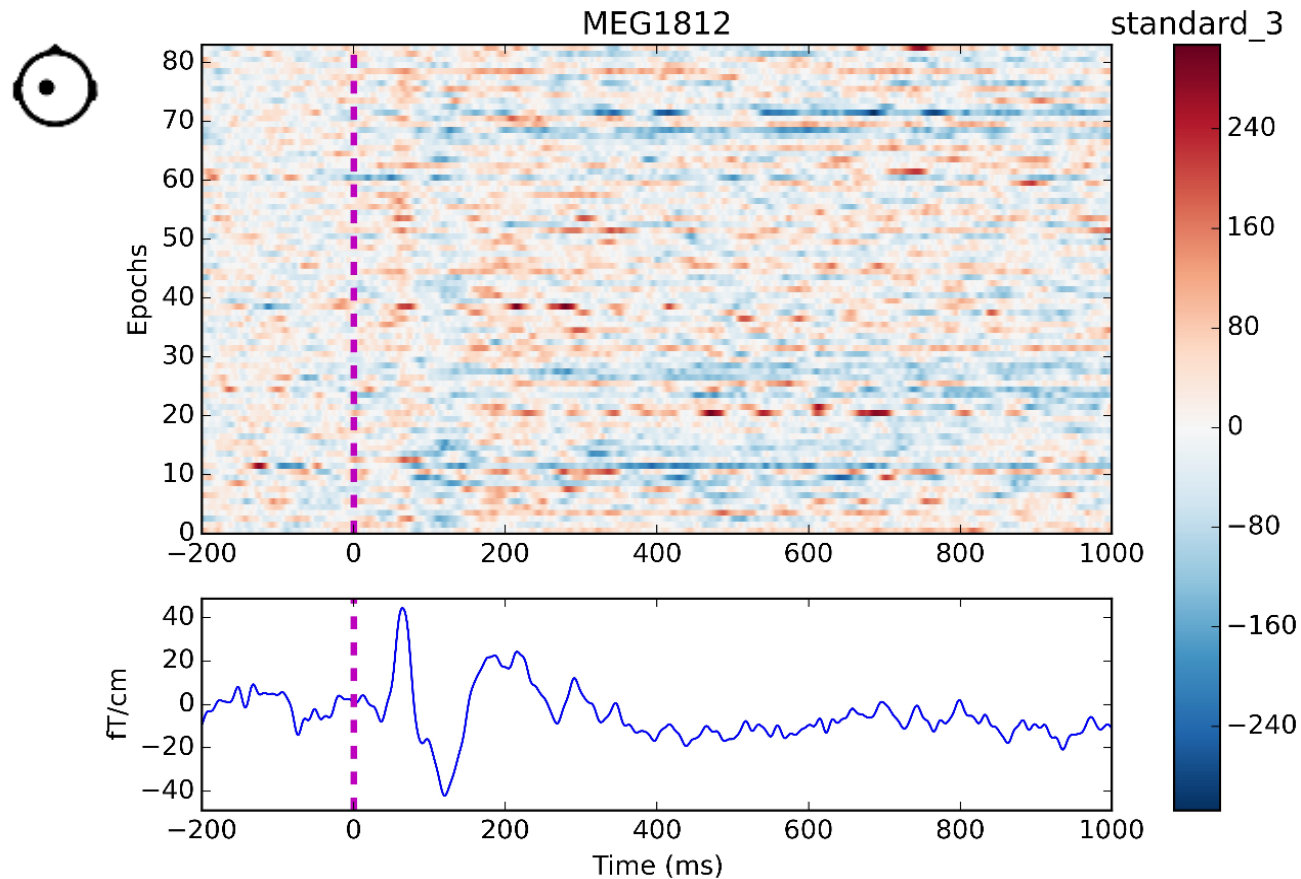
        epochs.save(epochs_path)

    else:
        print('epochs file: ' + epochs_path + ' already exists')
```

# OPERATIONS – FILTERING

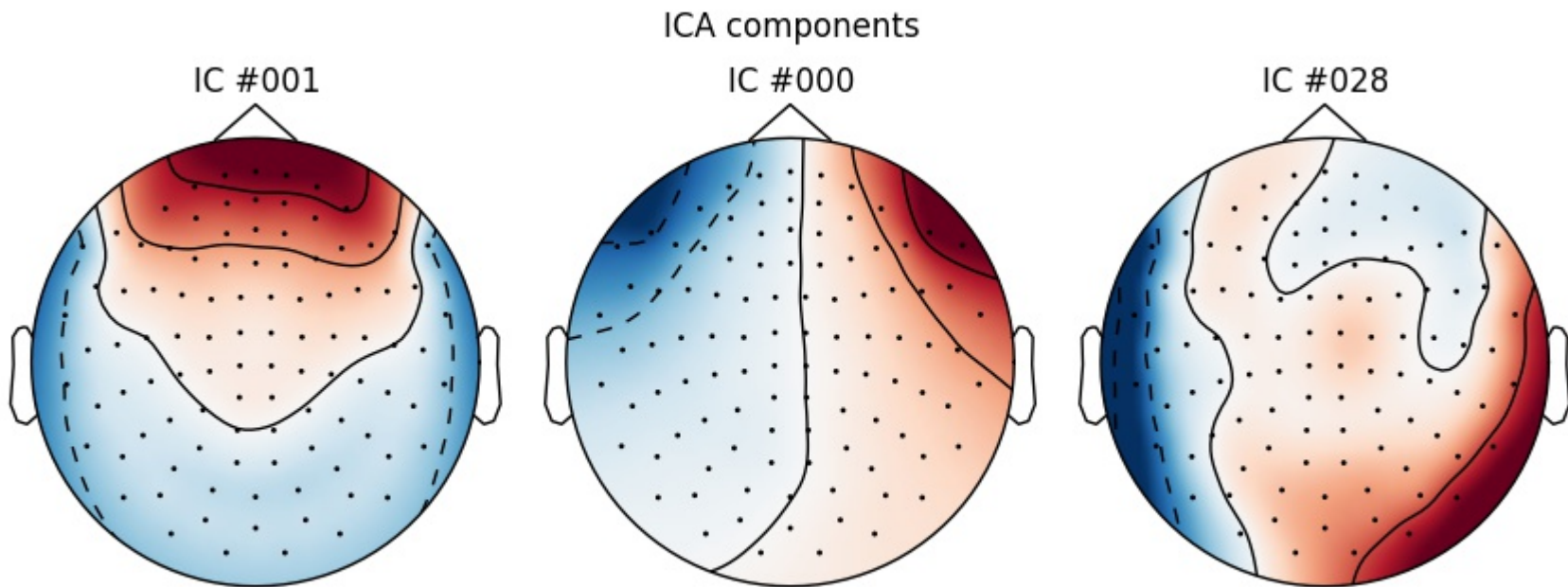


# OPERATIONS – EPOCHING



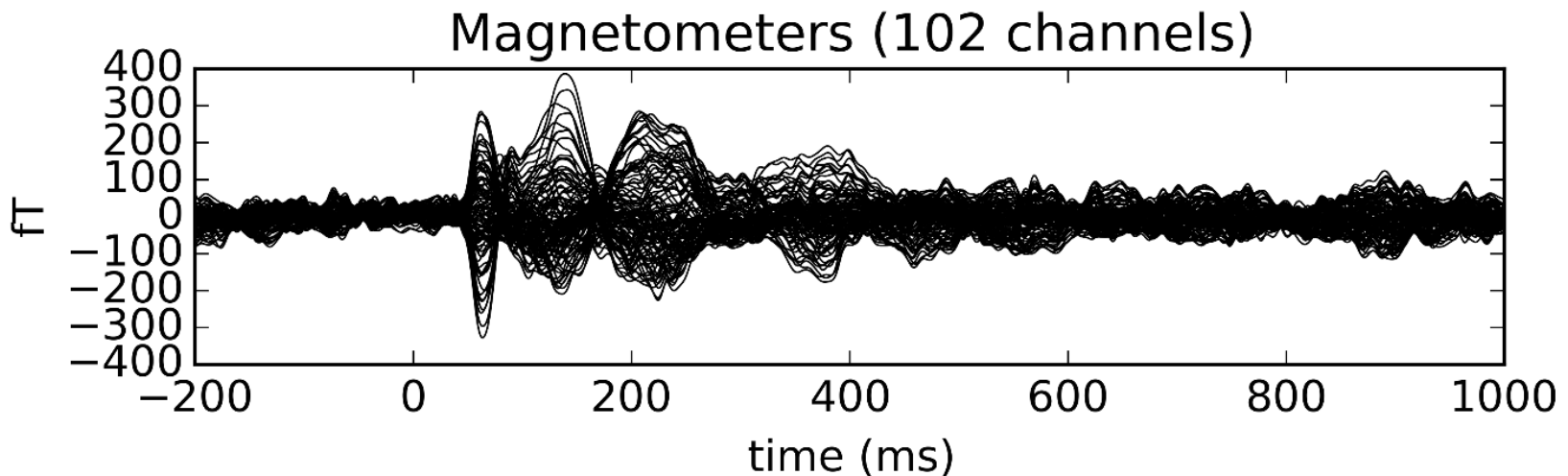


# OPERATIONS – ICA



PHYSIOLOGICAL NOISE, *EYE BLINKS* AND *HEART BEATS*

# OPERATIONS – EVOKEDS



THESE ARE THE ONES WE WANT TO SOURCE RECONSTRUCT

# How to analyse MEG?

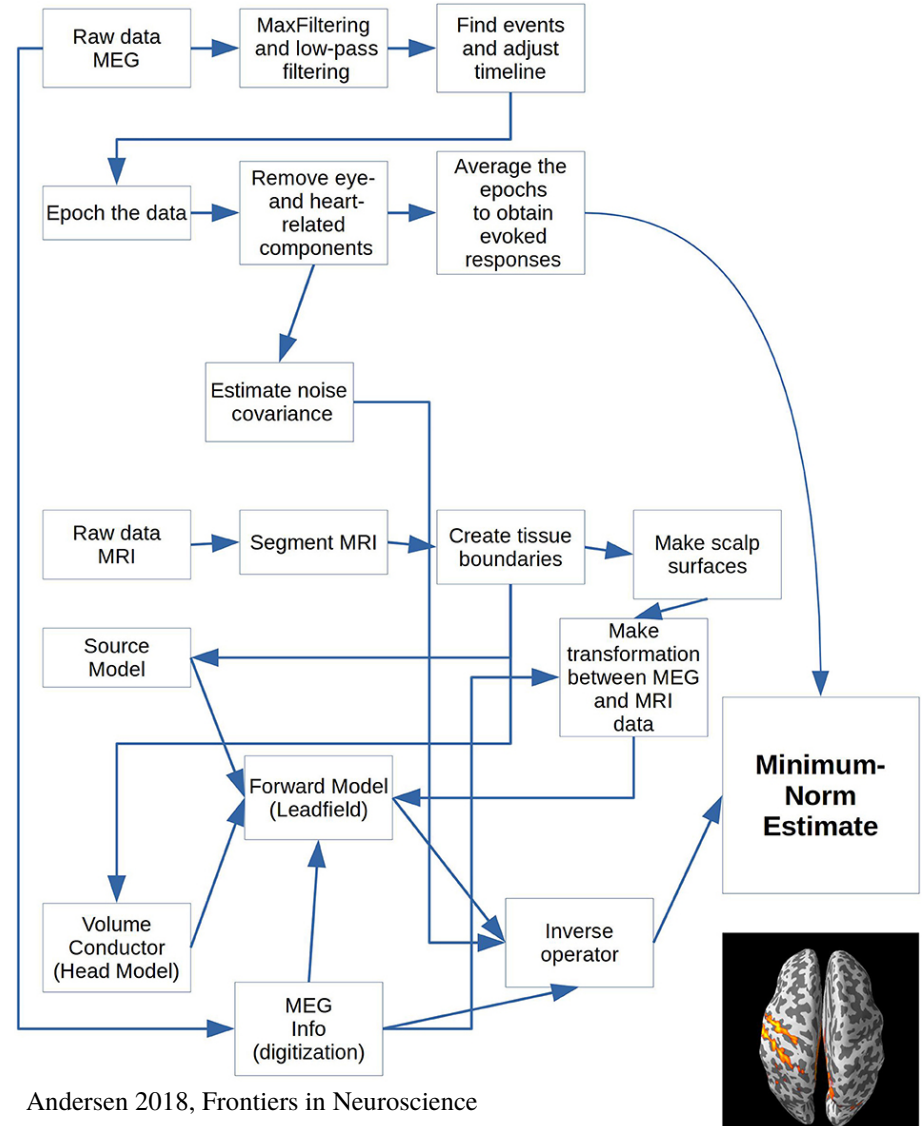
## OPERATIONS

### ## sensor space operations

```
filter_raw=0,  
find_events=0,  
epoch_raw=0,  
run_ica=0,  
apply_ica=0,  
get_evoked=0,
```

### ## source space operations

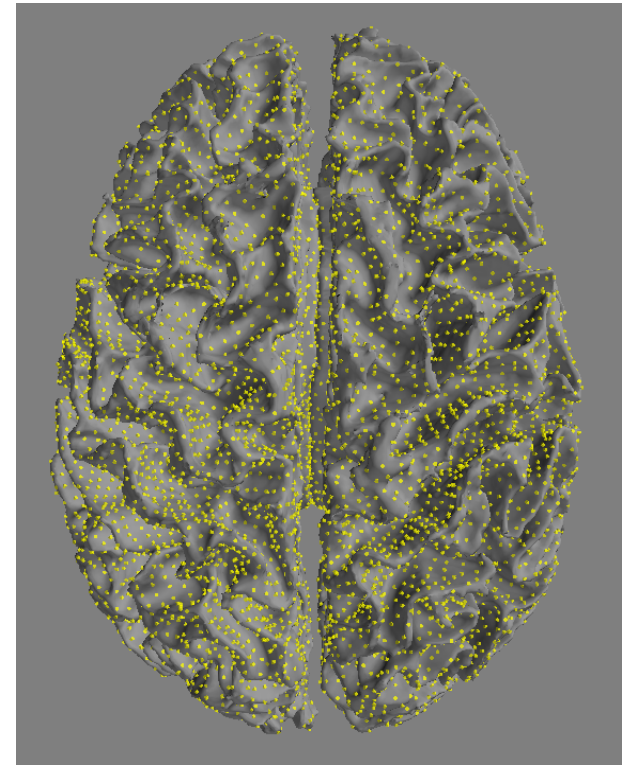
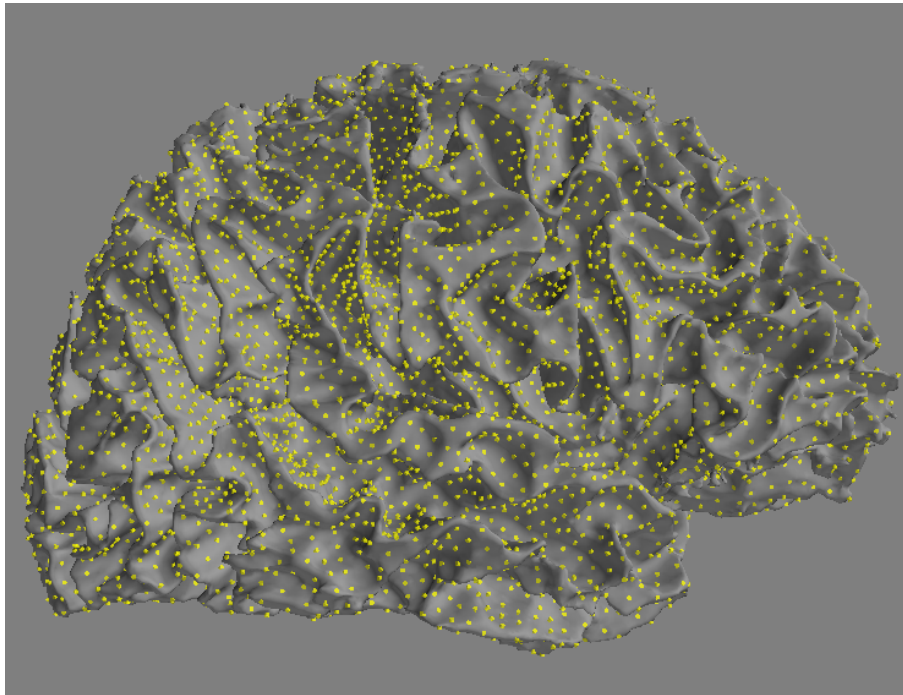
```
import_mri=0,  
segment_mri=0, # long process (>6 h)  
apply_watershed=0,  
make_source_space=0,  
make_dense_scalp_surfaces=0,  
make_bem_solutions=0,  
create_forward_solution=0,  
estimate_noise_covariance=0,  
create_inverse_operator=0,  
source_estimate=0,  
morph_to_fsaverage=0,
```



Andersen 2018, Frontiers in Neuroscience

*How to analyse MEG?*

# OPERATIONS – MAKE SOURCE SPACE



THE POSITIONS OF THE SOURCES

*How to analyse MEG?*

# OPERATIONS – CO- REGISTER MEG AND MRI

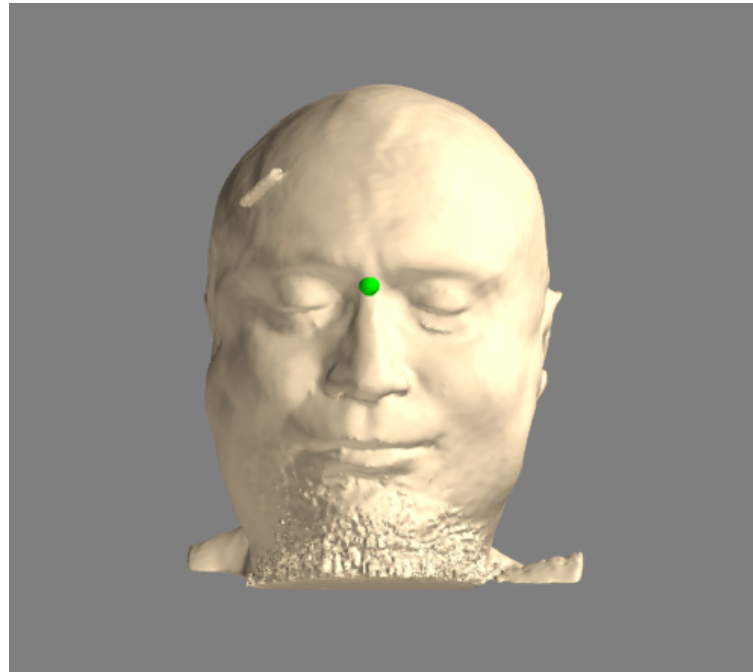


CONSTRUCT A HEAD MODEL BASED ON THE MRI

*How to analyse MEG?*

# OPERATIONS – CO- REGISTER MEG AND MRI

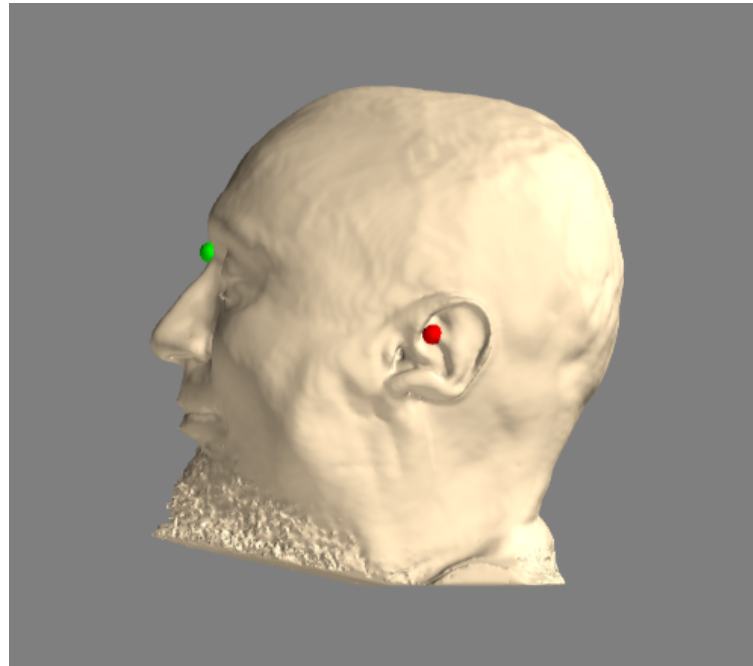
1) NASION



PLOT IN FIDUCIAL POINTS

# OPERATIONS – CO- REGISTER MEG AND MRI

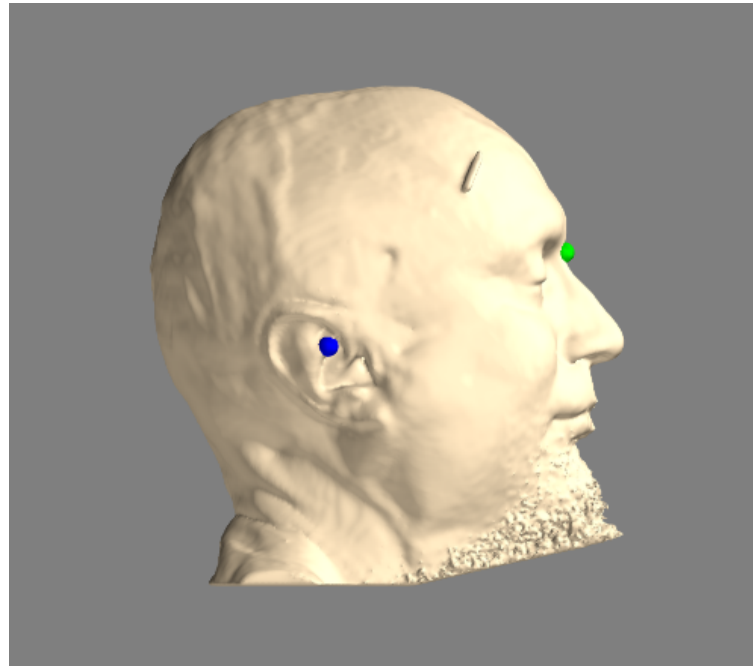
- 1) NASION
- 2) LPA



PLOT IN FIDUCIAL POINTS

# OPERATIONS – CO-REGISTER MEG AND MRI

- 1) NASION
- 2) LPA
- 3) RPA

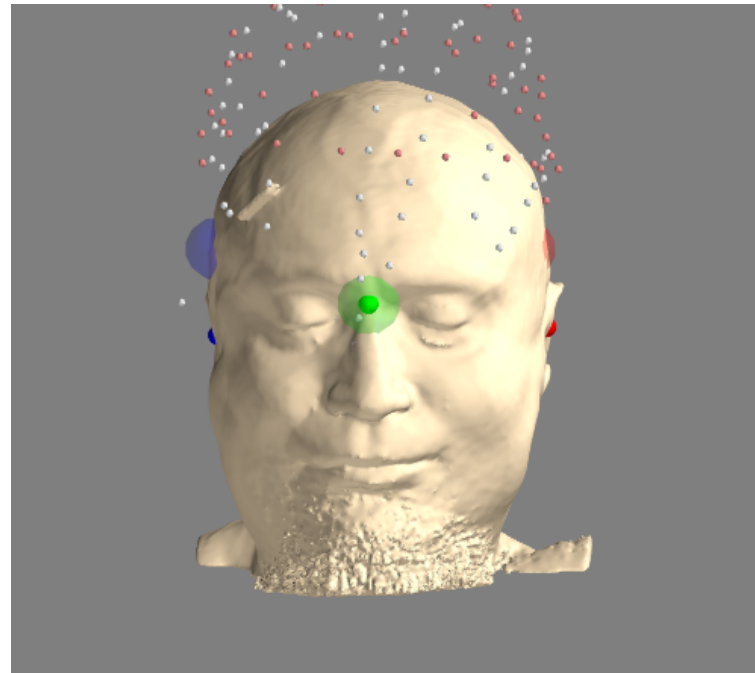


PLOT IN FIDUCIAL POINTS



# OPERATIONS – CO-REGISTER MEG AND MRI

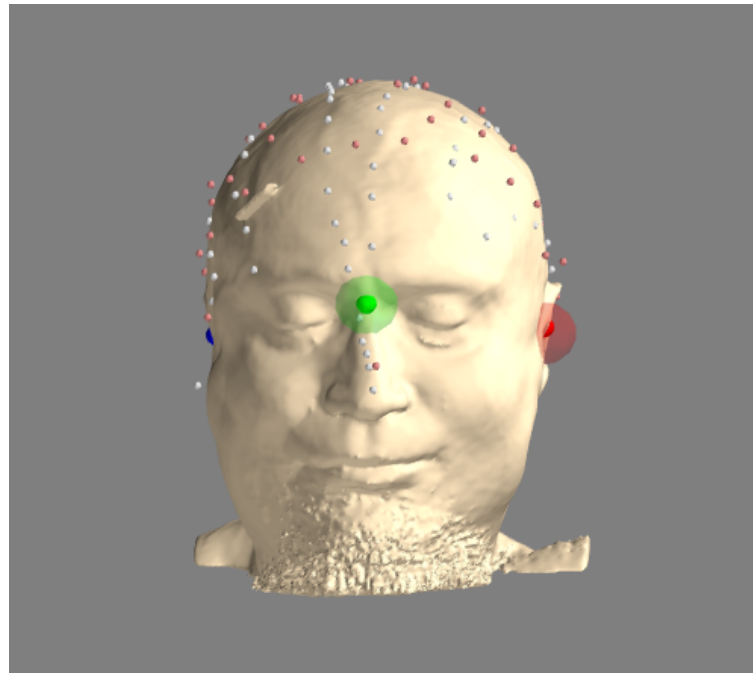
- 1) NASION
- 2) LPA
- 3) RPA



PLOT IN EXTRA HEAD POINTS

# OPERATIONS – CO-REGISTER MEG AND MRI

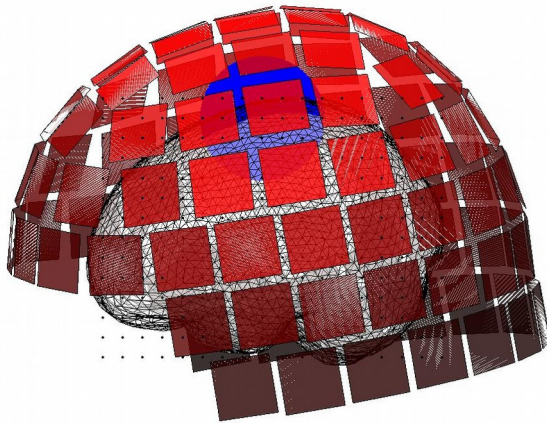
- 1) NASION
- 2) LPA
- 3) RPA



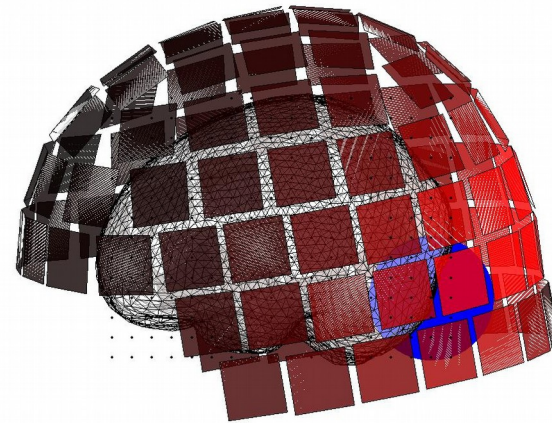
MINIMIZE DISTANCE BETWEEN HEAD POINTS AND HEAD MODEL

# OPERATIONS – FORWARD MODEL

"Tactile source"



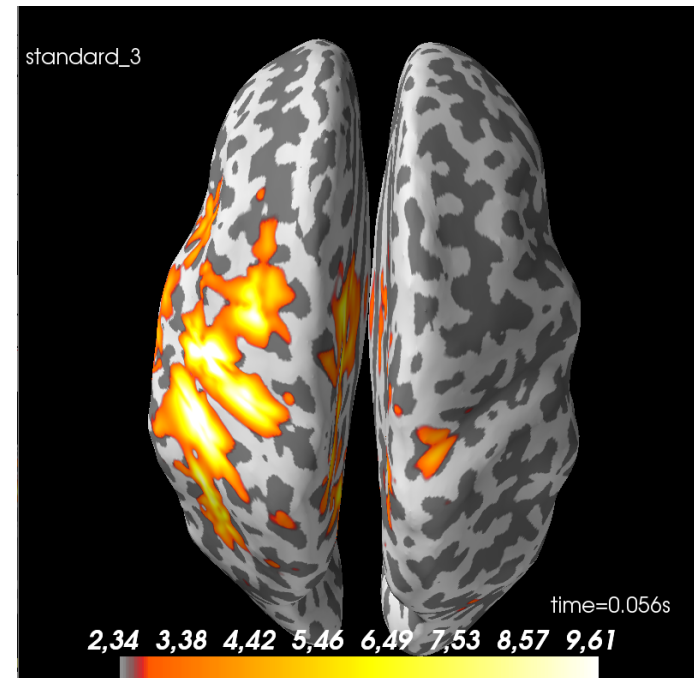
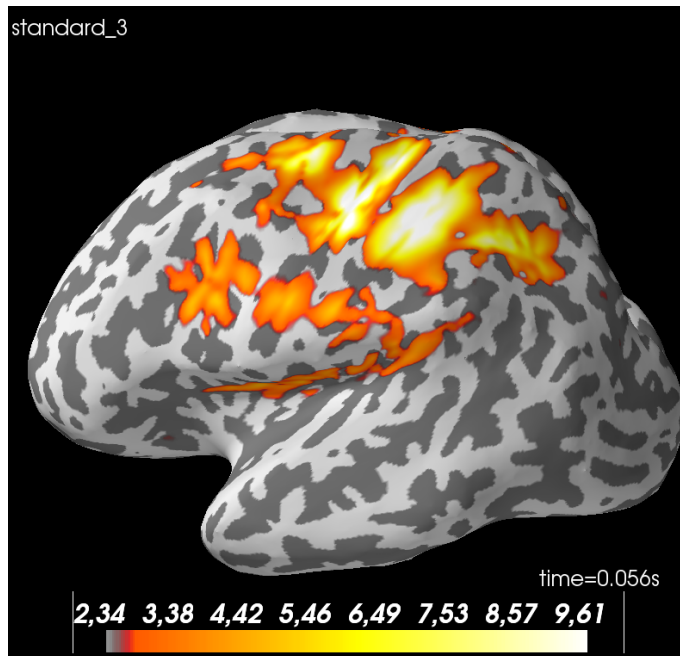
"Occipital" source



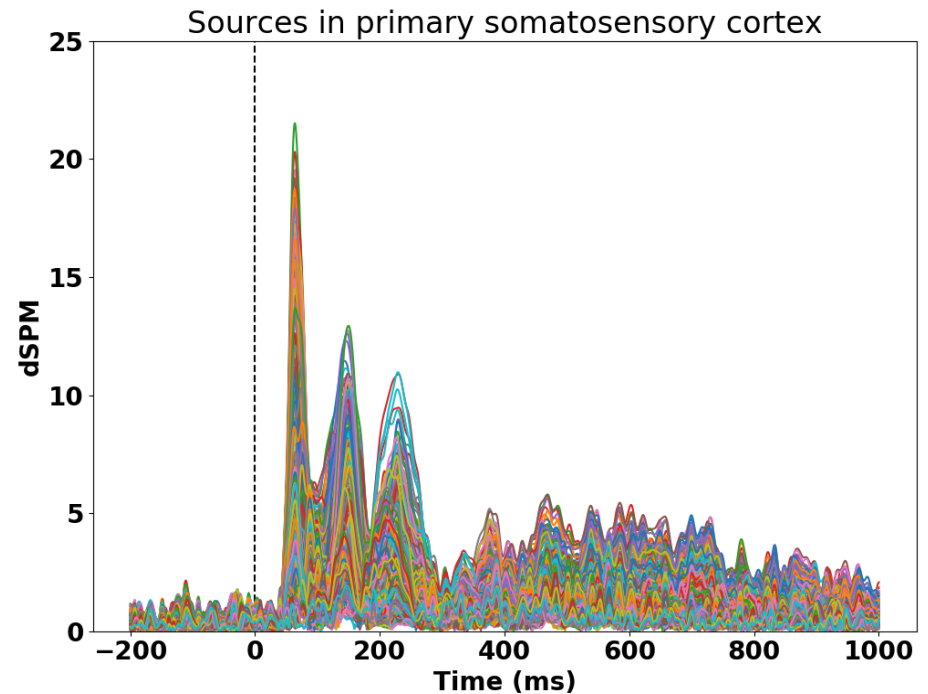
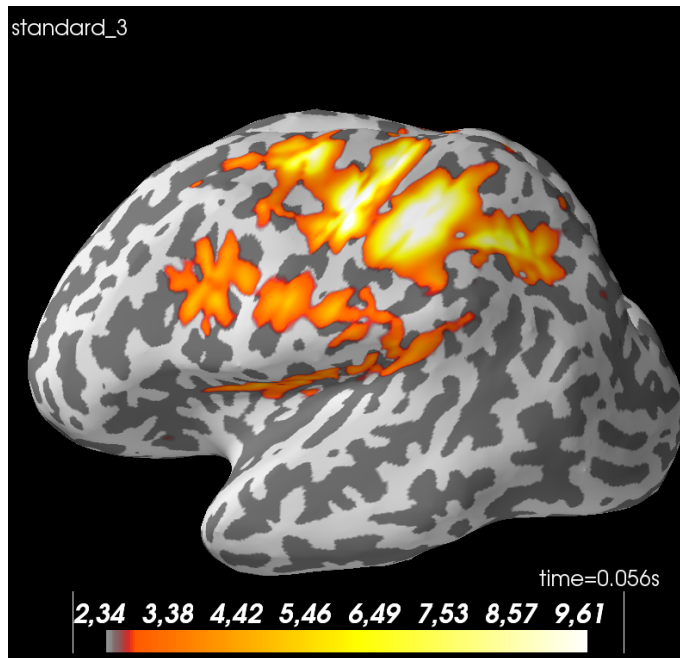
THE LINK BETWEEN SOURCES AND CHANNELS

*How to analyse MEG?*

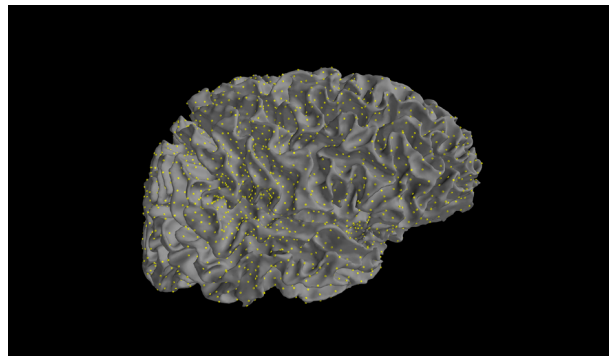
# OPERATIONS – SOURCE RECONSTRUCTION



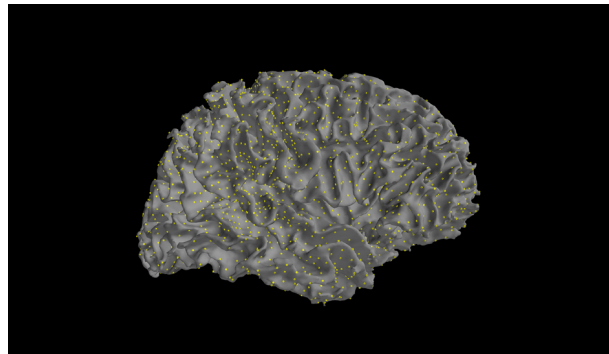
# OPERATIONS – SOURCE RECONSTRUCTION



# OPERATIONS – MORPH TO TEMPLATE



$n_1$

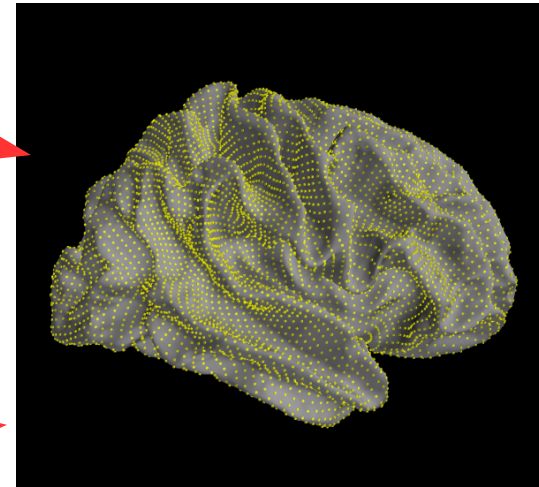


$n_2$



...

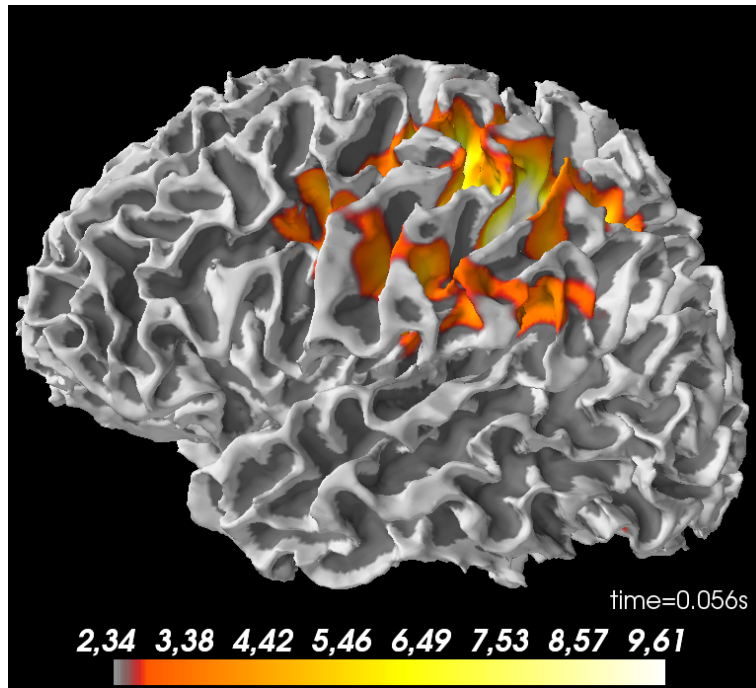
$n_{20}$



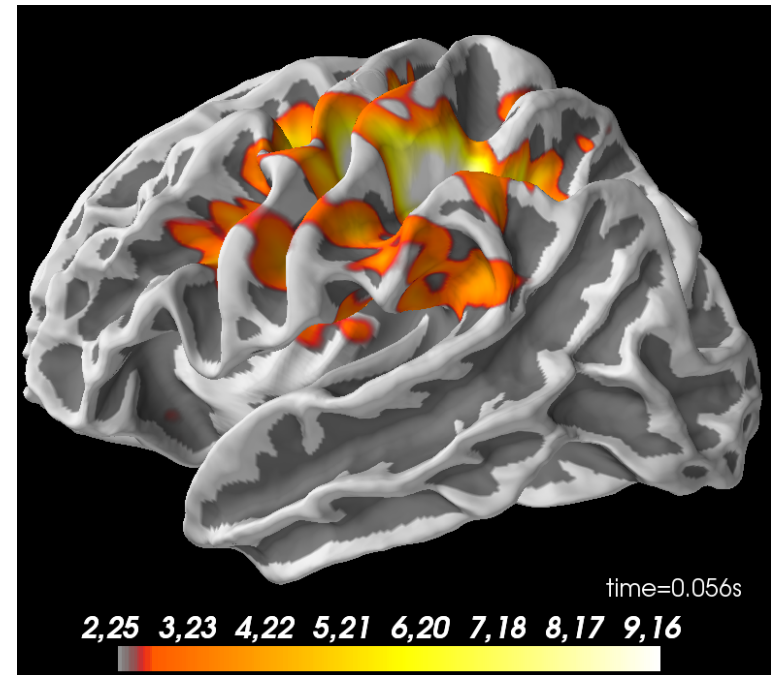
*fsaverage* template from the  
FreeSurfer suite, but many other  
anatomies exist

*How to analyse MEG?*

# OPERATIONS – MORPH TO TEMPLATE

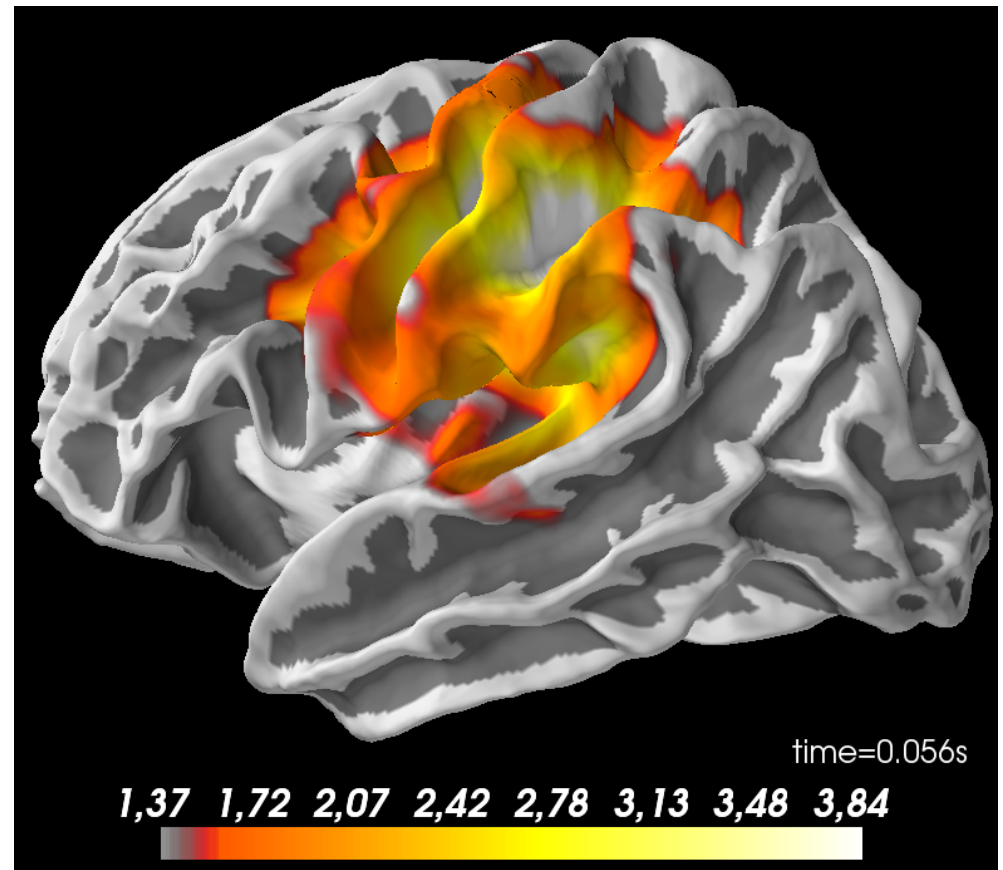


INDIVIDUAL'S ANATOMY



FS AVERAGE TEMPLATE

# OPERATIONS – GRAND AVERAGE

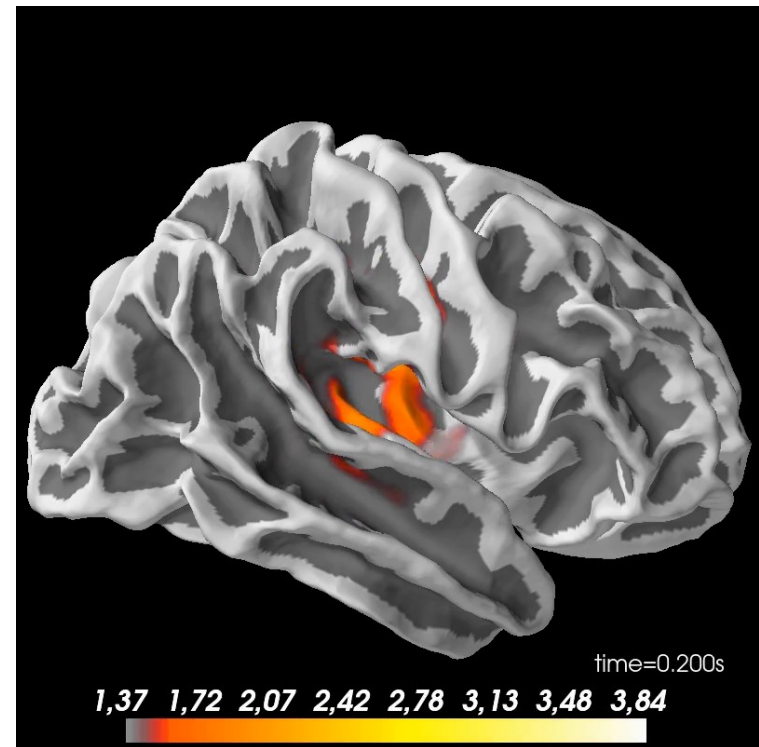
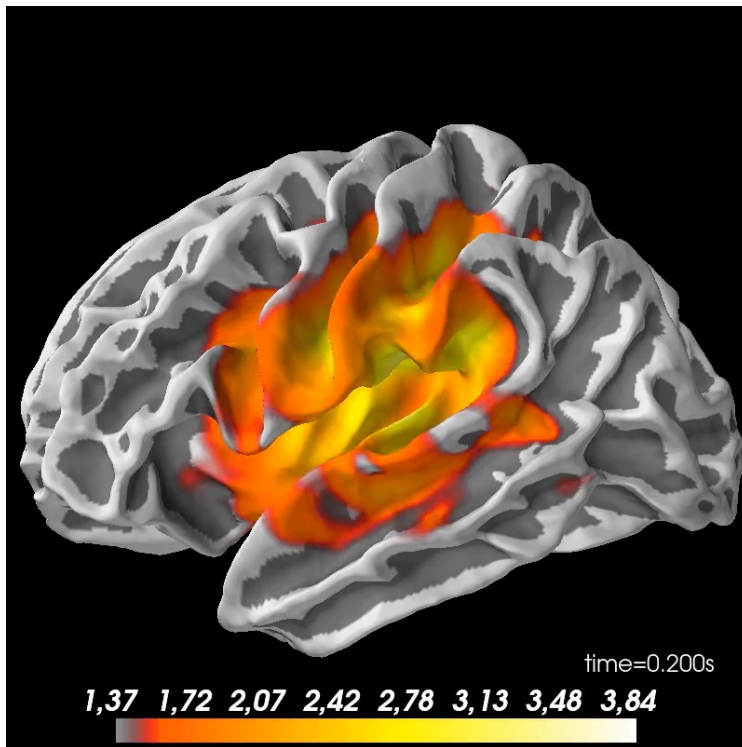


FSAVERAGE TEMPLATE



*How to analyse MEG?*

# OPERATIONS – GRAND AVERAGE VIDEOS



# OPERATIONS – STATISTICS

Do *Stimulations* give rise to more activity than *Non-Stimulations* in the primary somatosensory cortex?

## How do we decide?

# How do we decide?

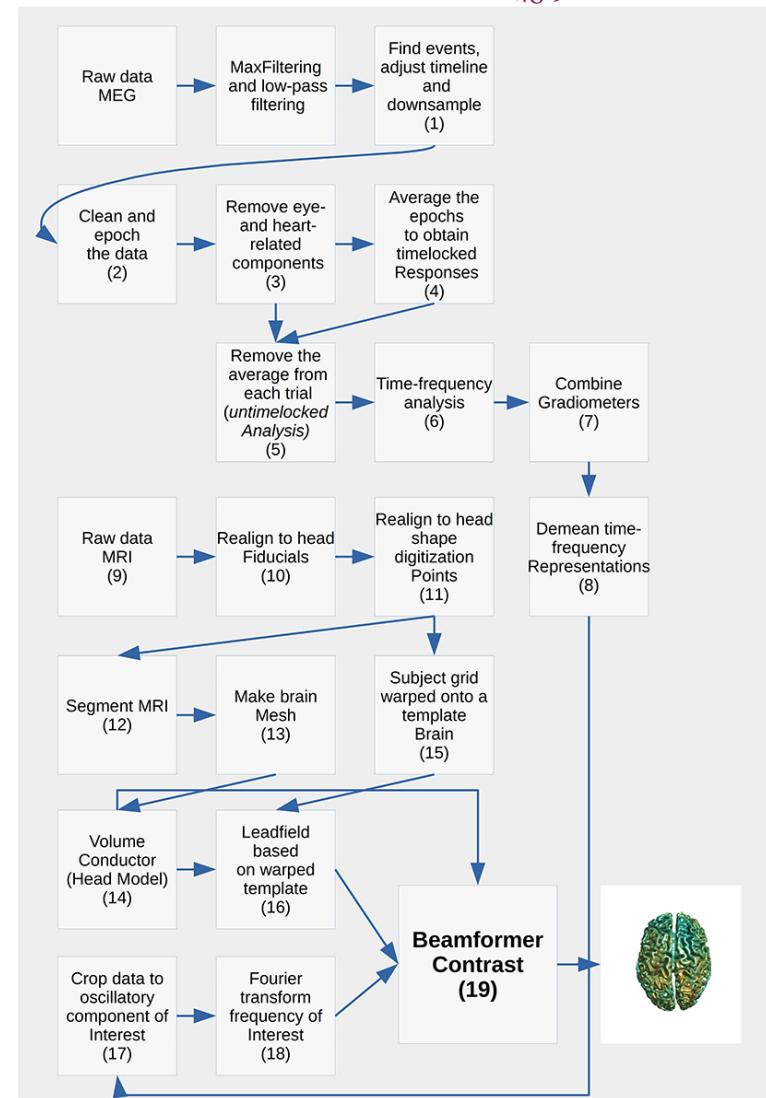
We'll postpone this question until we have gone through the next analysis pipeline

# How to analyse MEG?

# FIELDTRIP PIPELINE

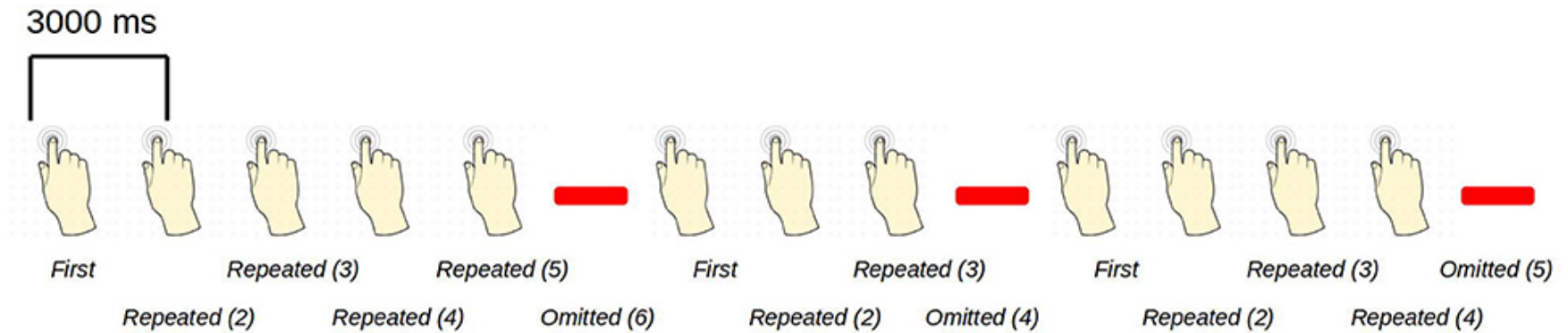
Argument	Purpose
<code>subjects</code>	Subject IDs indicating the directory name of the subject
<code>data_dir</code>	Whether data is MEG or MRI data
<code>function_name</code>	The function that should be applied to all subjects
<code>cfg</code>	Configuration structure, as known from FieldTrip
<code>output</code>	A cell array of name(s) of the output file(s)
<code>input</code>	A cell array of name(s) of the input file(s)
<code>figures_dir</code>	Where figures should be stored (leave empty, [], if no figures are produced)
<code>overwrite</code>	Whether existing output files should be overwritten

Arguments for the function `loop_through_subjects`, which structures input and output of all operations done on single subjects.



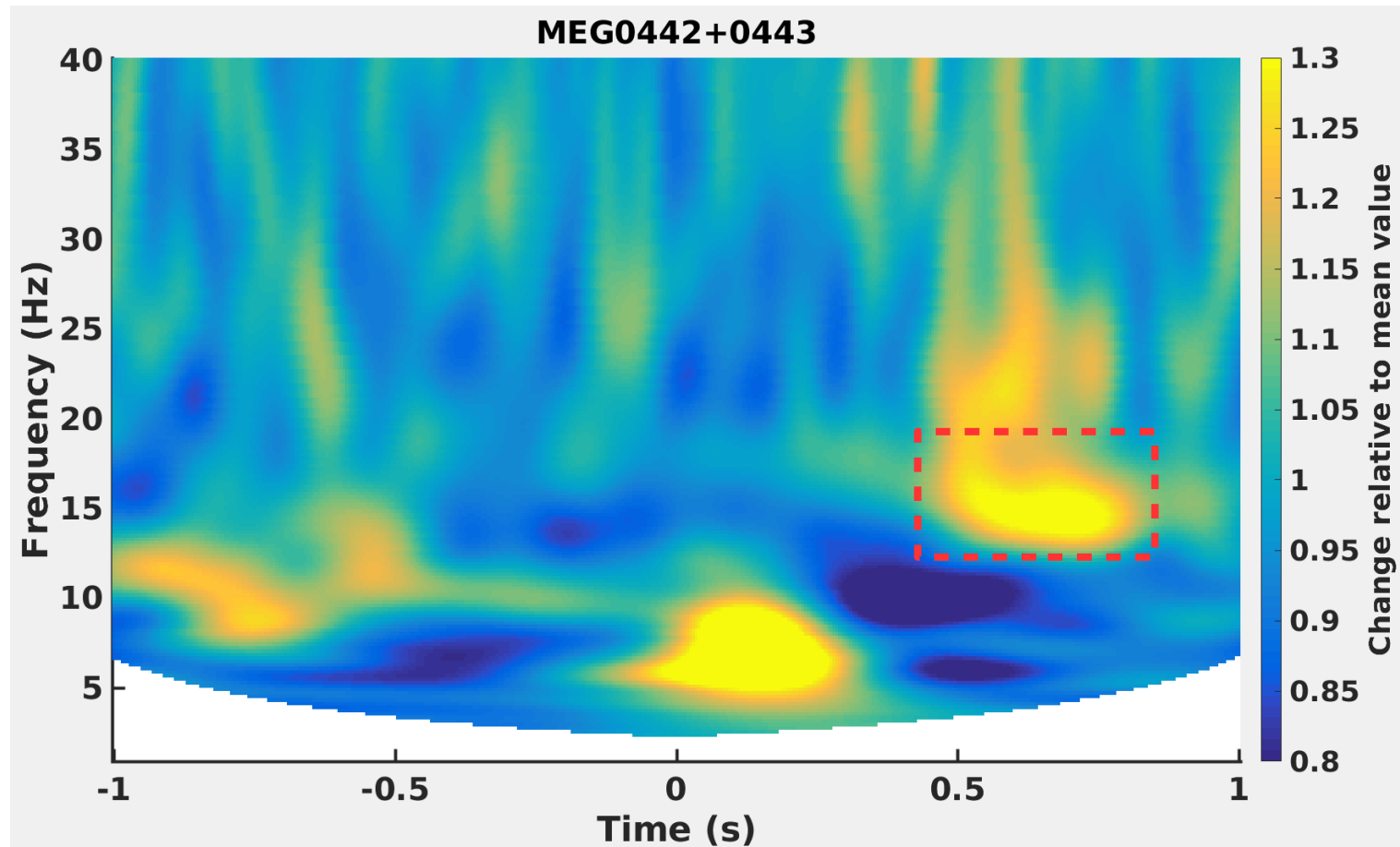
Andersen 2018, Frontiers in Neuroscience

## SAME PARADIGM



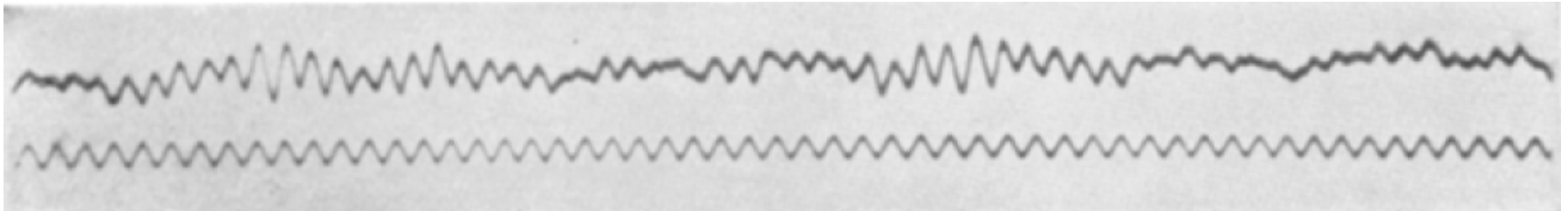
Andersen 2018, *Frontiers in Neuroscience*

# BETA REBOUND



# OSCILLATIONS

Berger 1929  
*Archives für Psychiatrie*



First EEG recorded:

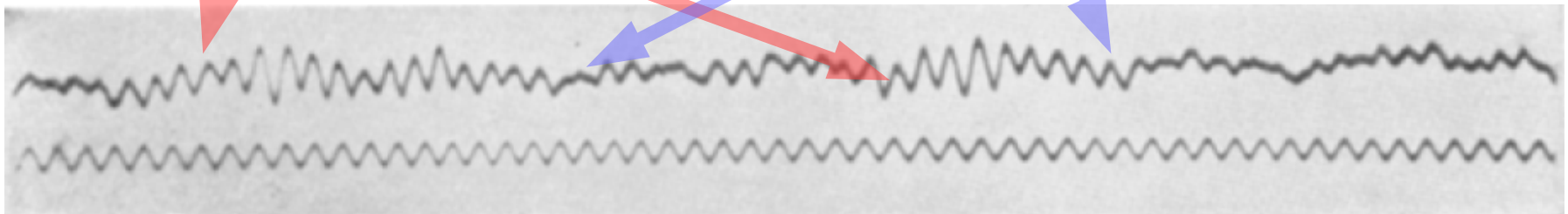
Upper row: EEG

Lower row: 10 Hz (alpha) rhythm

# OSCILLATIONS

**Subject closed his eyes**

**Subject opened his eyes**



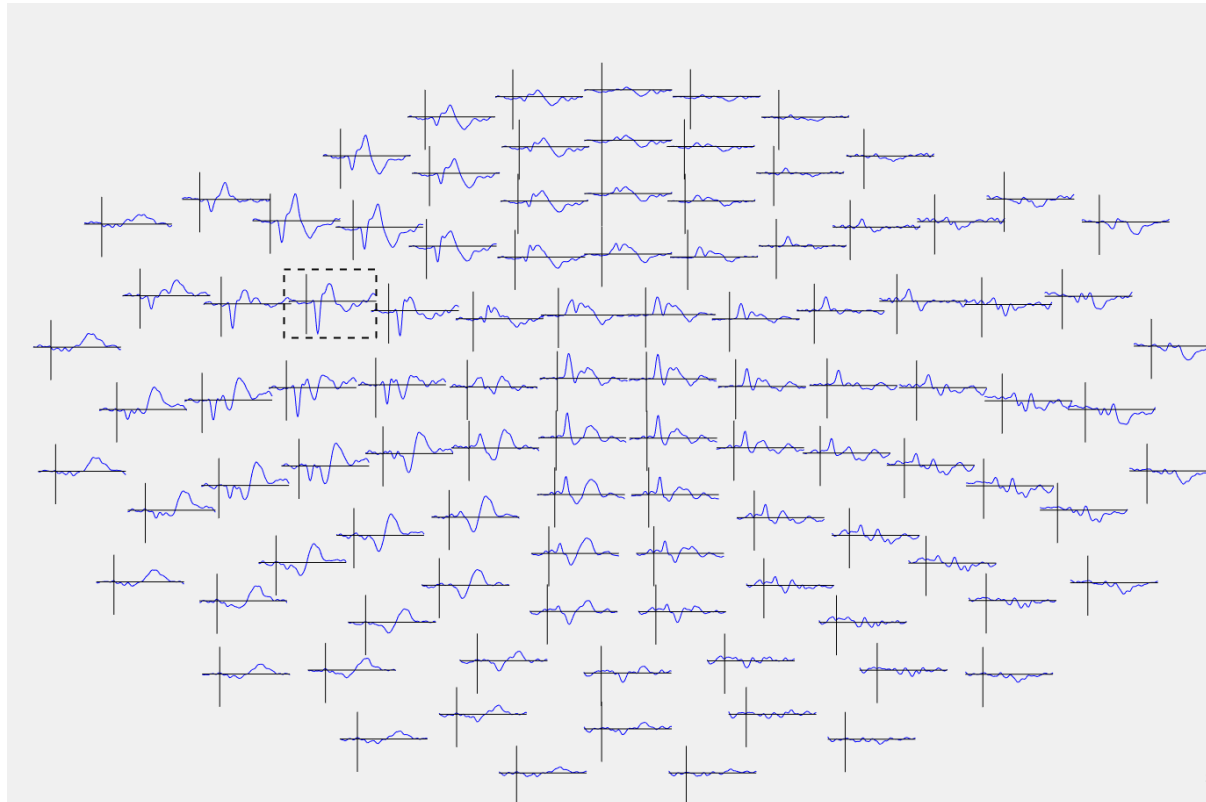
First EEG recorded:

Upper row: EEG

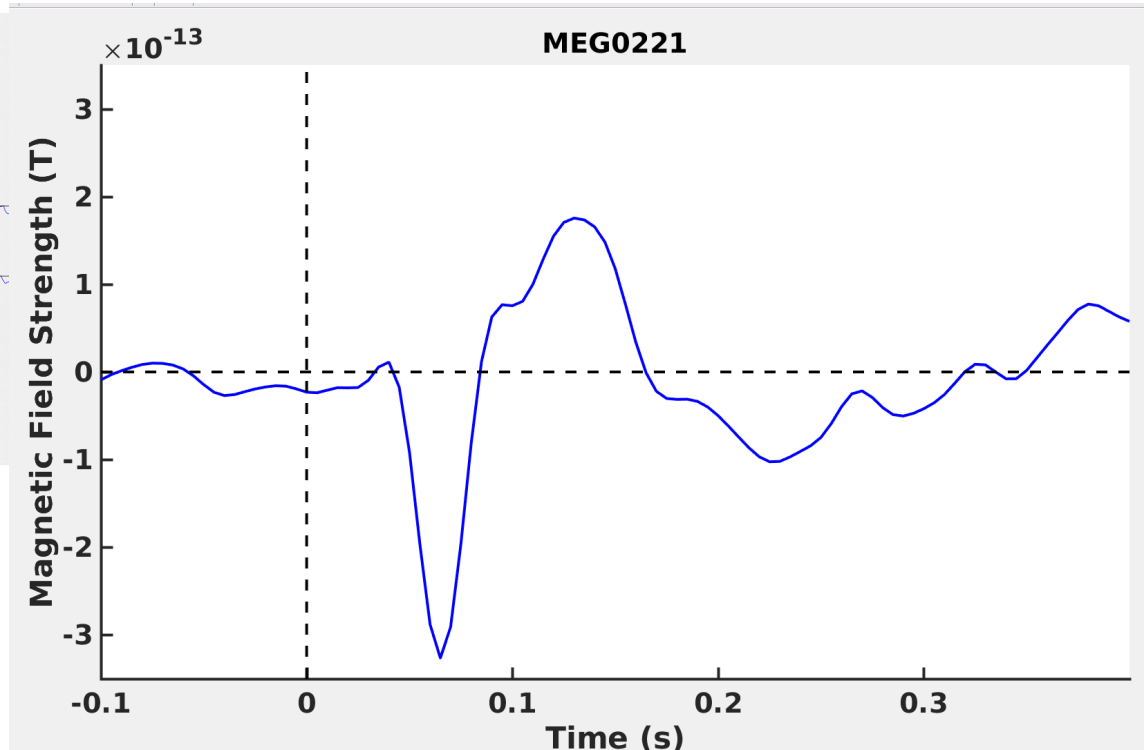
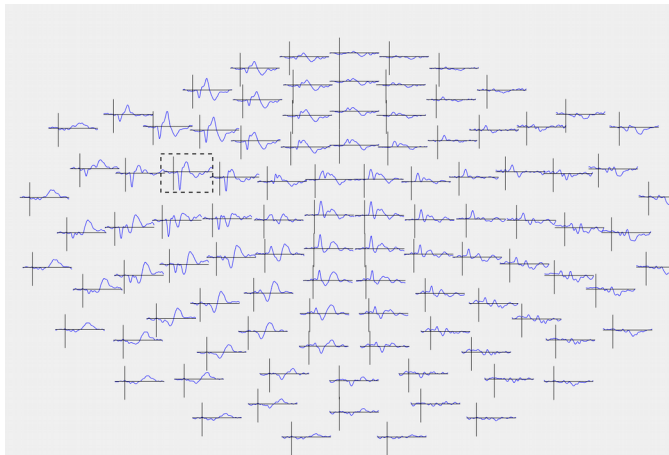
Lower row: 10 Hz (alpha) rhythm



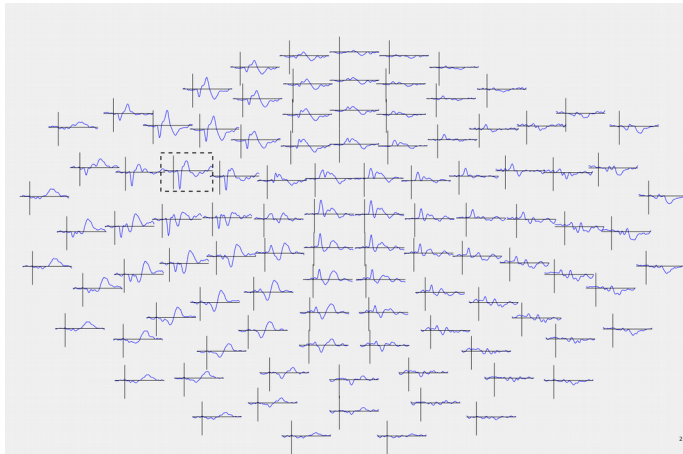
# OPERATIONS – TIMELOCKEDS (EVOKED)



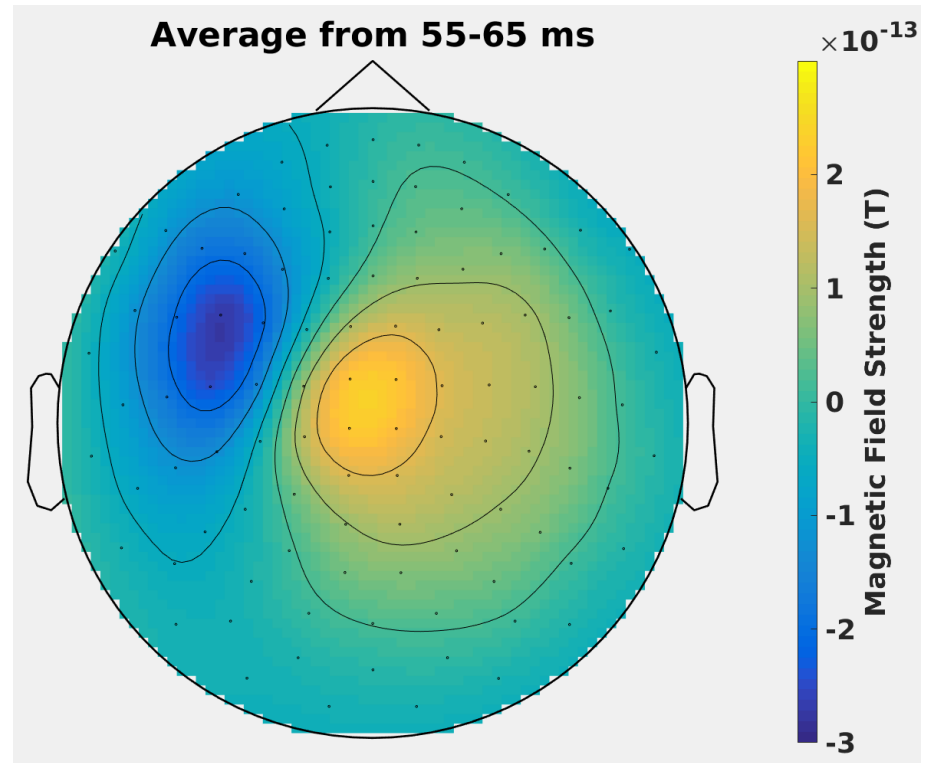
# OPERATIONS – TIMELOCKEDS (EVOKED)



# OPERATIONS – TIMELOCKEDS (EVOKED)

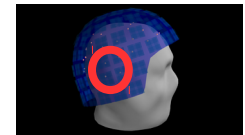
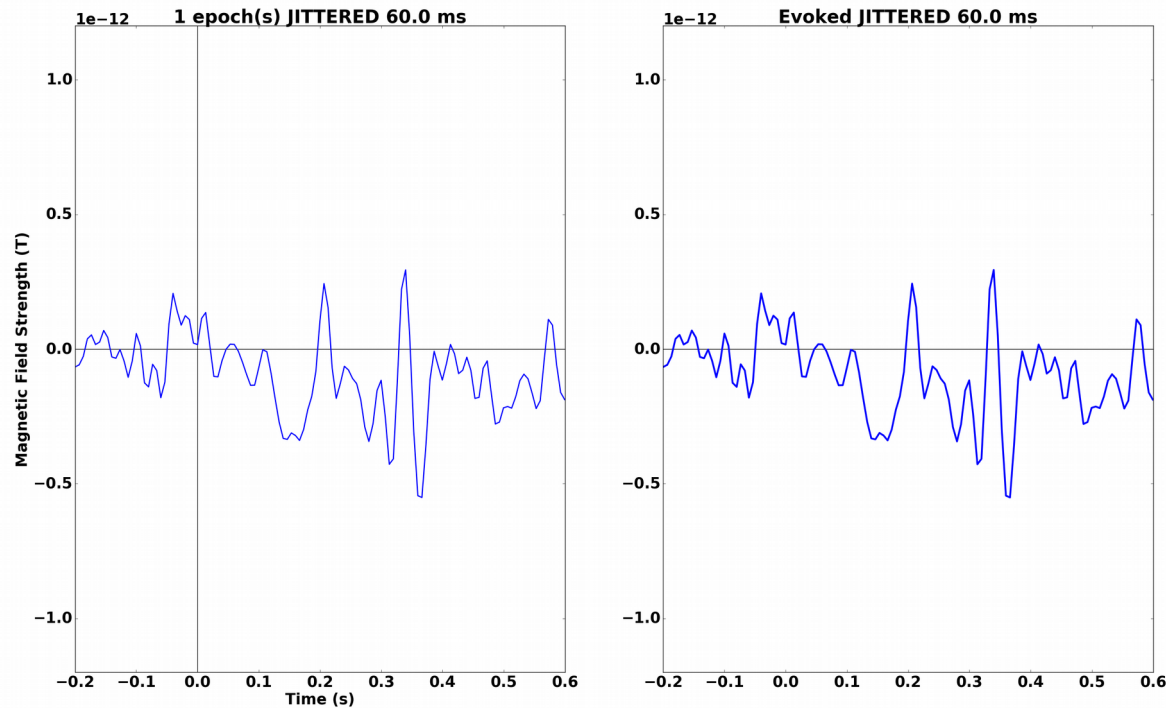


WE SEE NO OSCILLATIONS



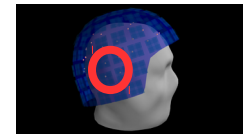
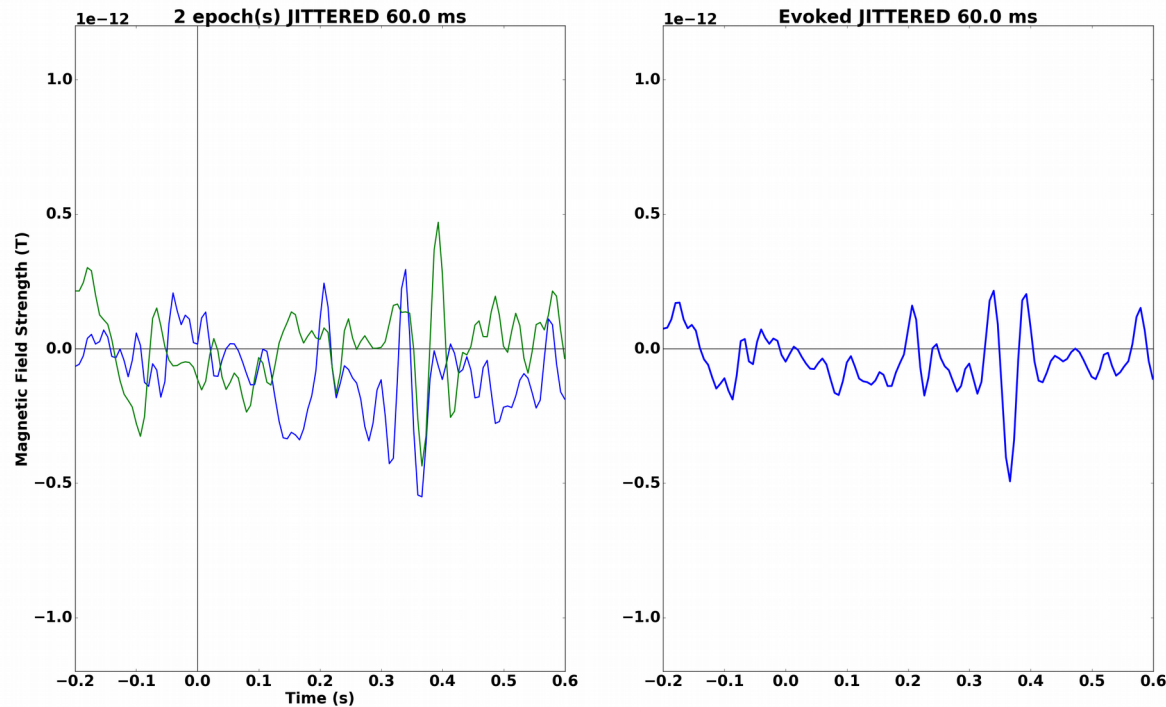
What is MEG?

# EPOCHS AND EVOKED (JITTERED 60 MS)



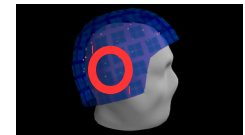
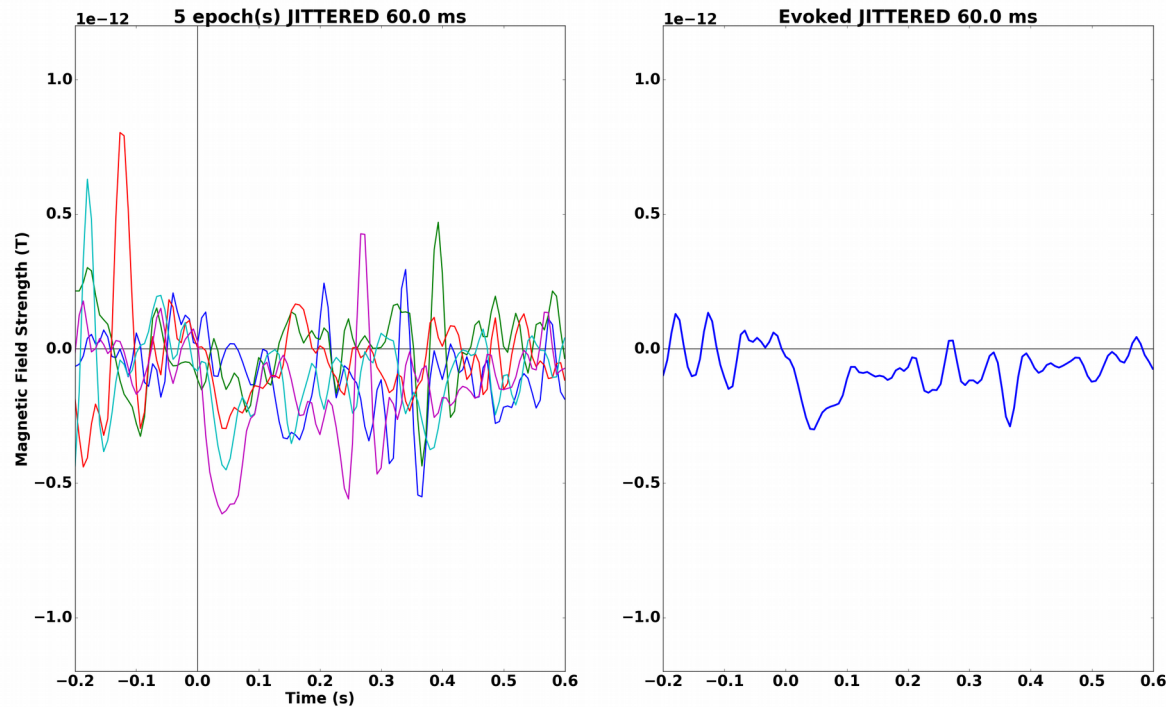
What is MEG?

# EPOCHS AND EVOKED (JITTERED 60 MS)



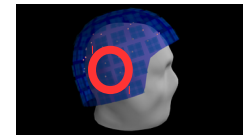
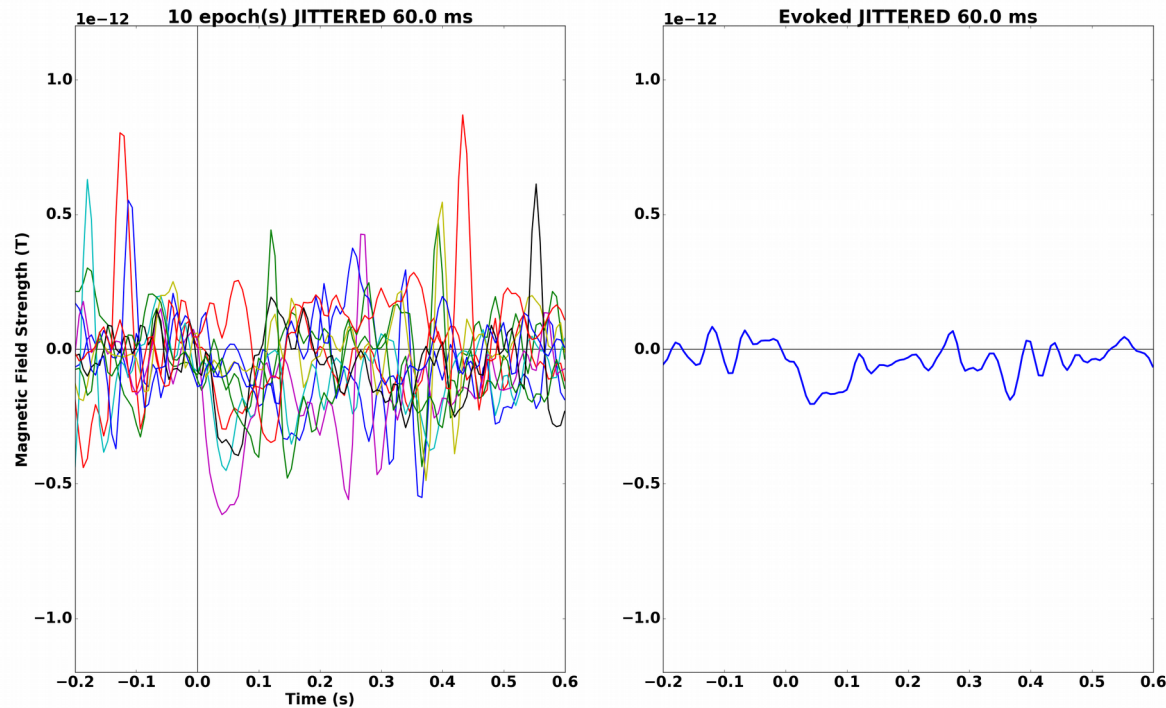
What is MEG?

# EPOCHS AND EVOKED (JITTERED 60 MS)



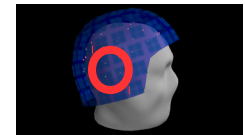
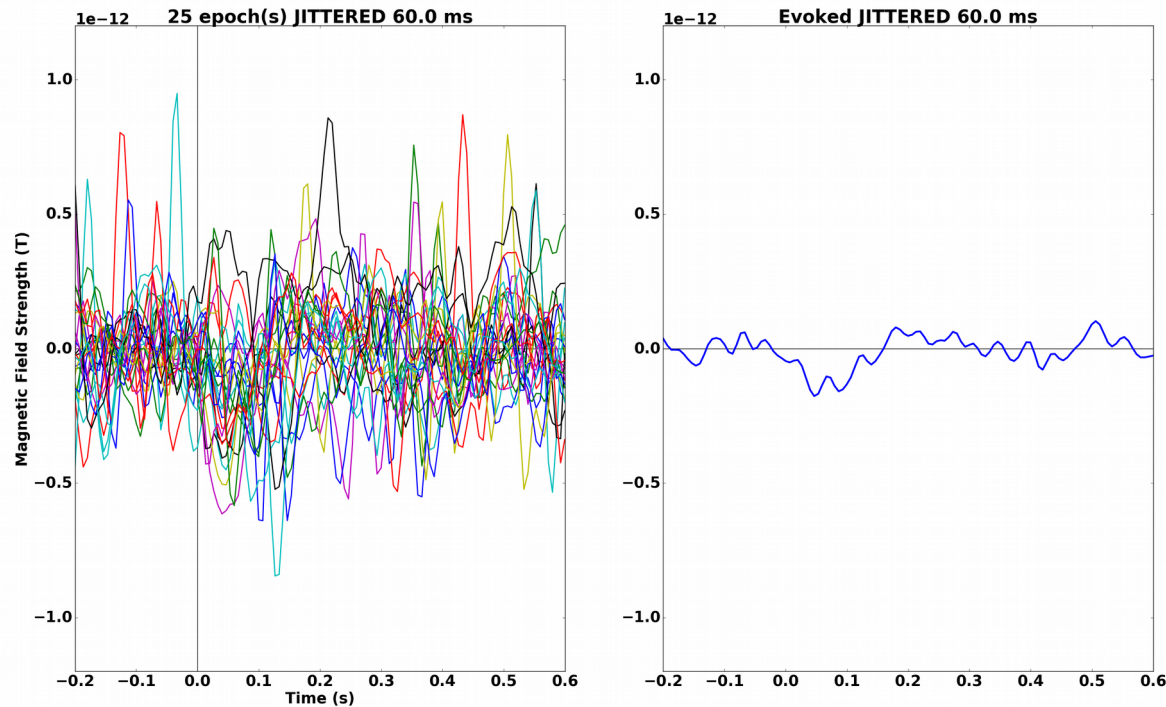
What is MEG?

# EPOCHS AND EVOKED (JITTERED 60 MS)



What is MEG?

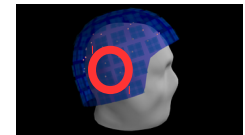
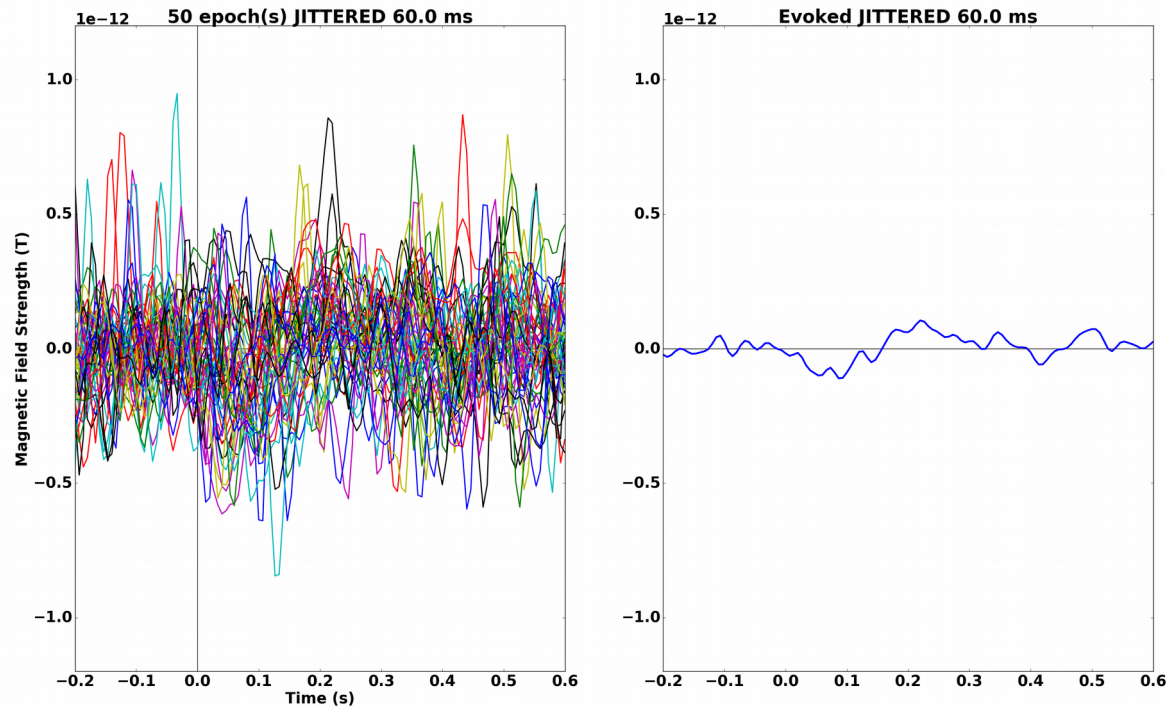
# EPOCHS AND EVOKED (JITTERED 60 MS)





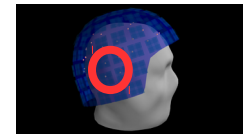
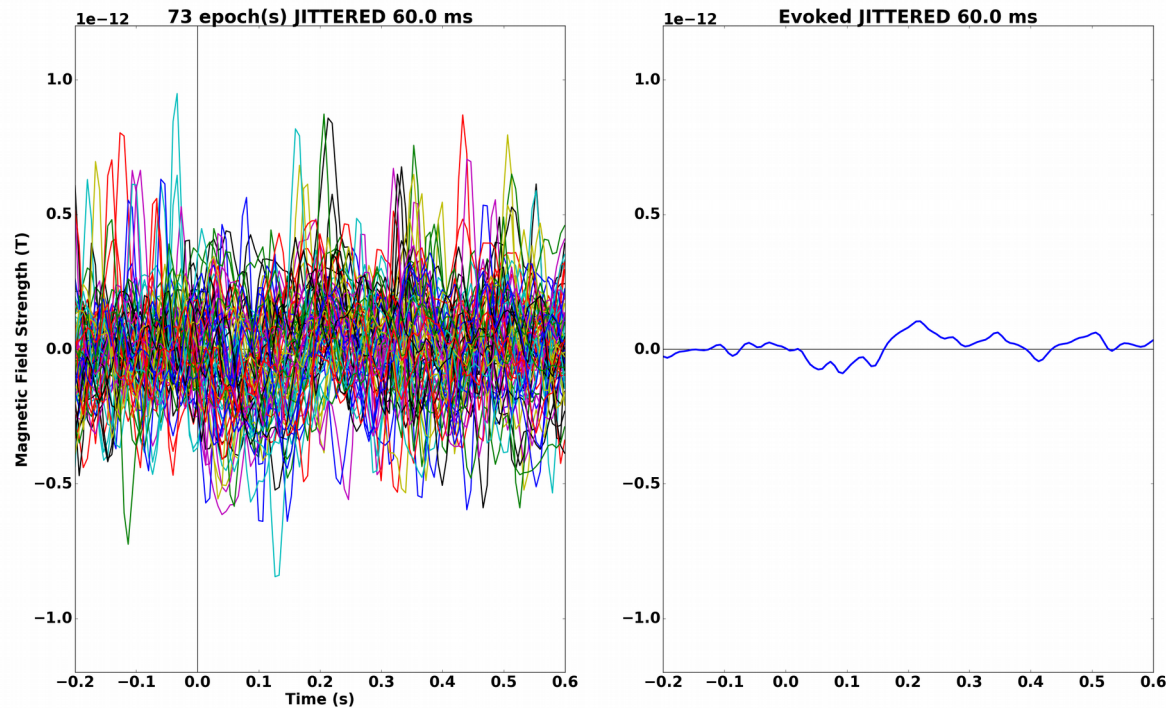
What is MEG?

# EPOCHS AND EVOKED (JITTERED 60 MS)

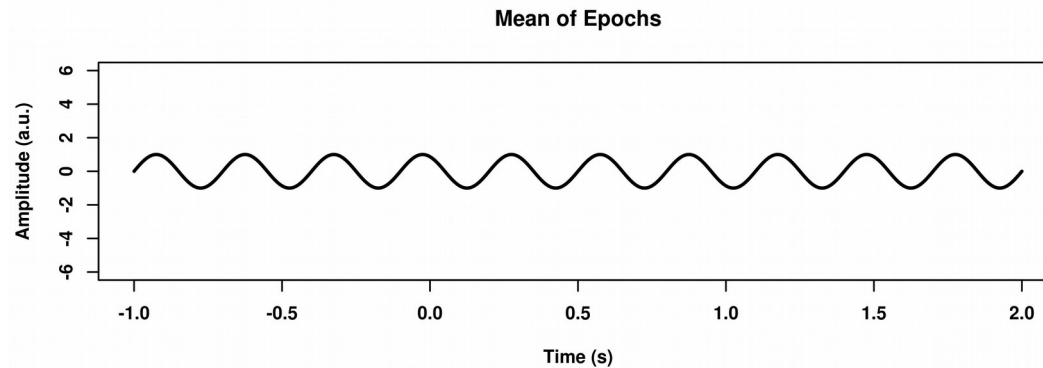
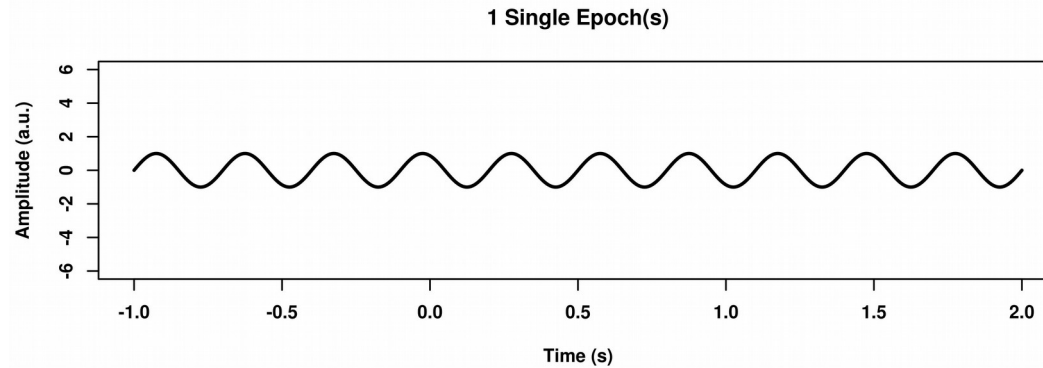


What is MEG?

# EPOCHS AND EVOKED (JITTERED 60 MS)

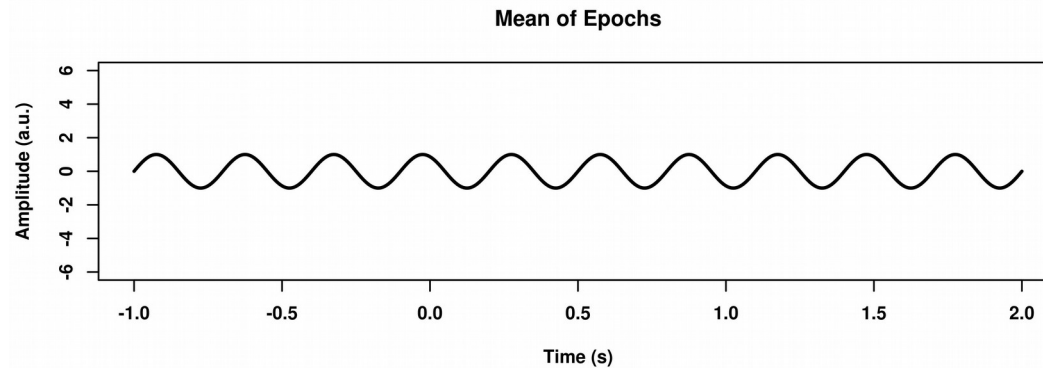
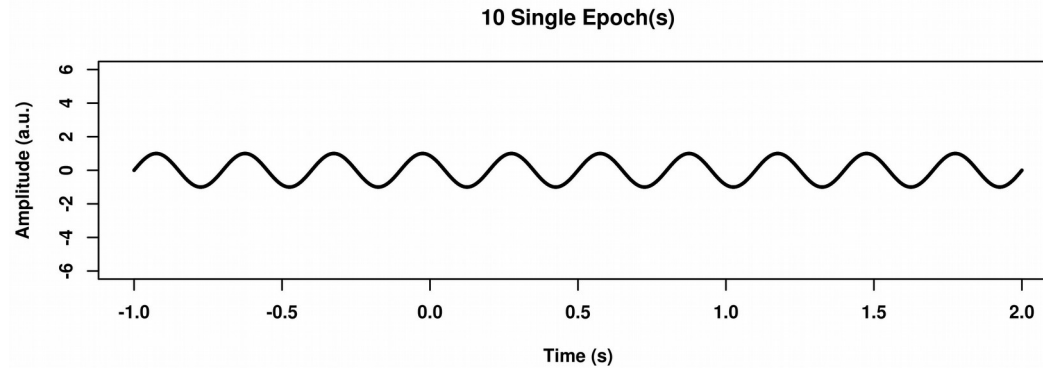


# IN-PHASE OR OUT-OF-PHASE



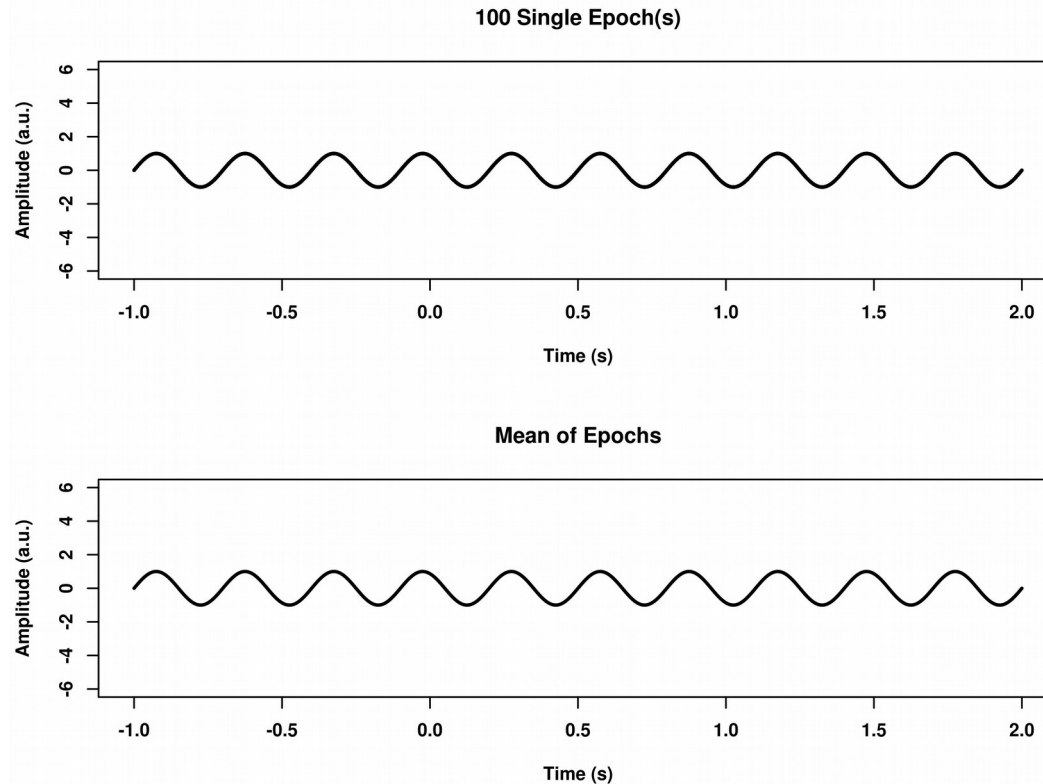
IN PHASE, IDEAL,  
Real Mean = 1.0

# IN-PHASE OR OUT-OF-PHASE



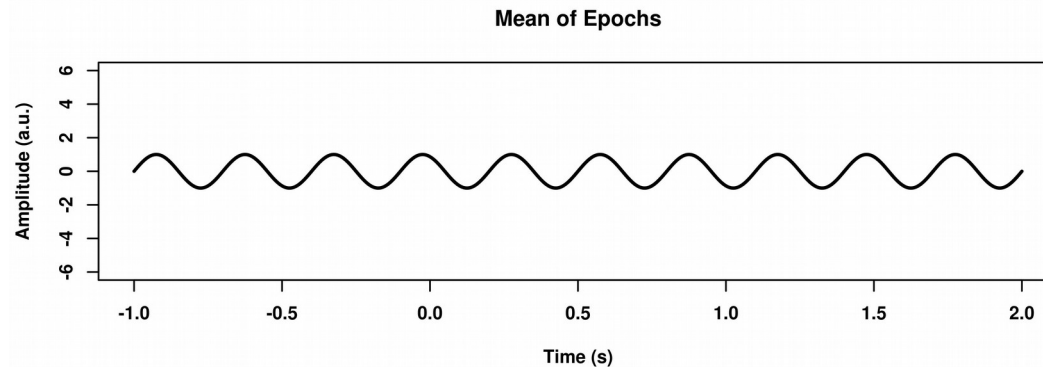
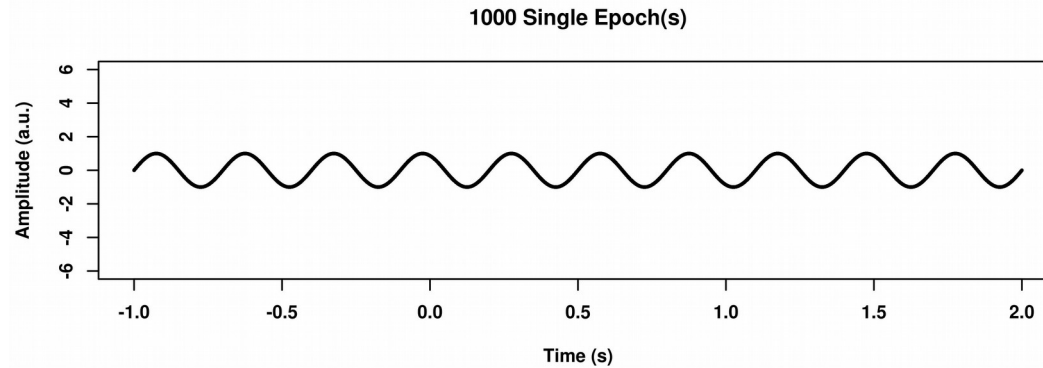
IN PHASE, IDEAL,  
Real Mean = 1.0

# IN-PHASE OR OUT-OF-PHASE



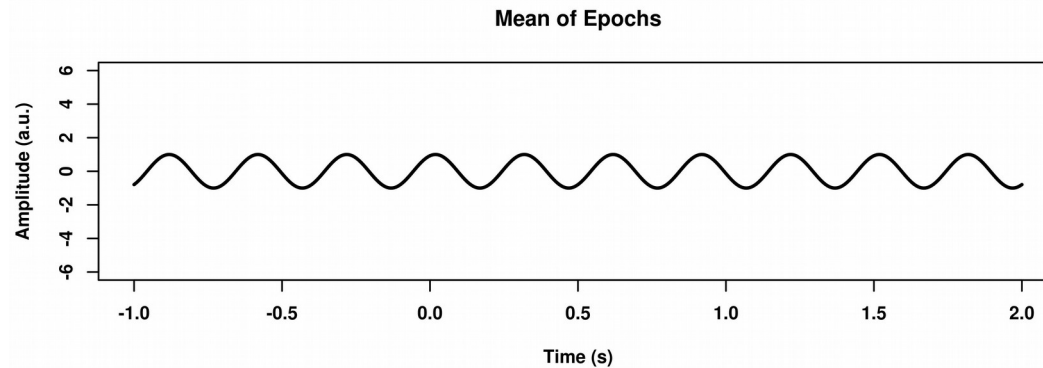
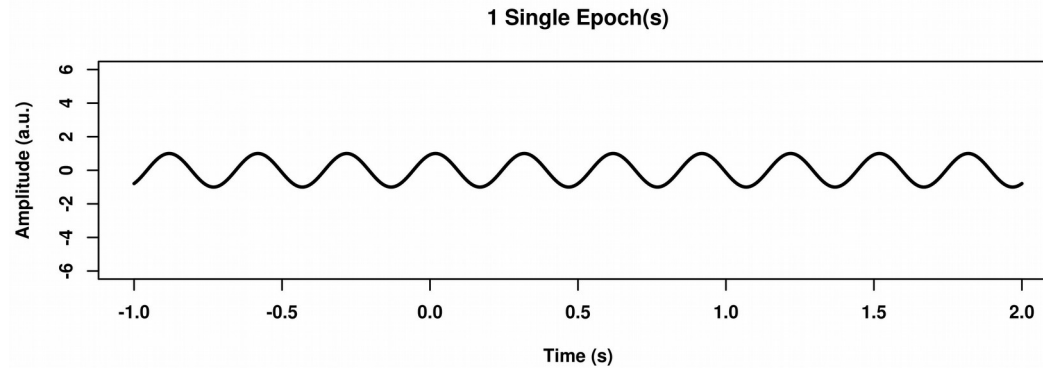
IN PHASE, IDEAL,  
Real Mean = 1.0

# IN-PHASE OR OUT-OF-PHASE



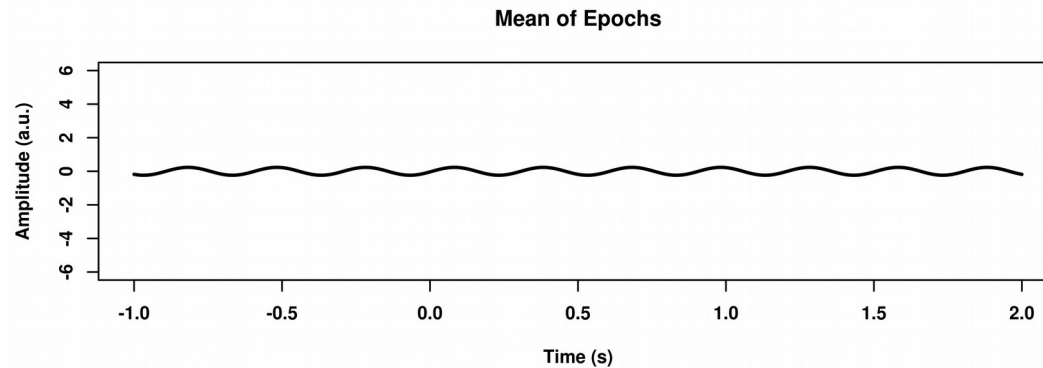
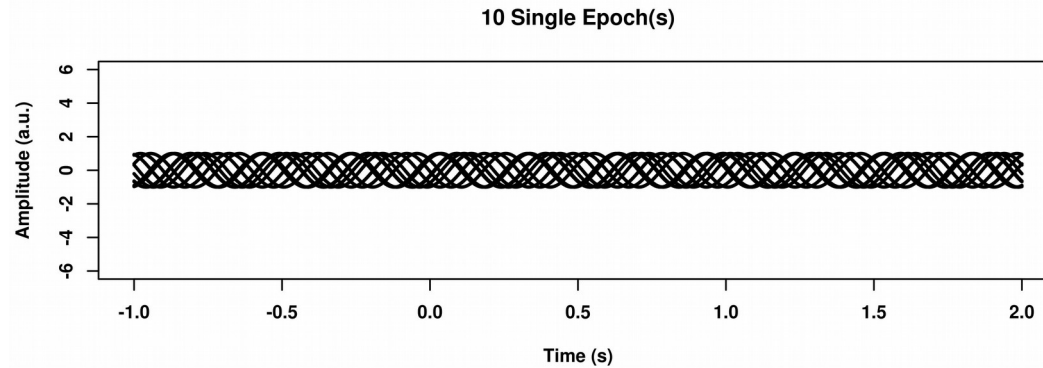
IN PHASE, IDEAL,  
Real Mean = 1.0

# IN-PHASE OR OUT-OF-PHASE



OUT OF PHASE,  
IDEAL, Real Mean = 1.0

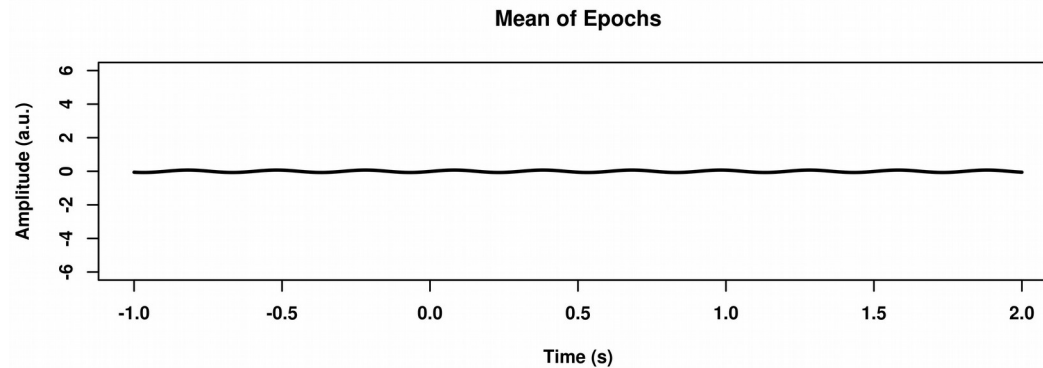
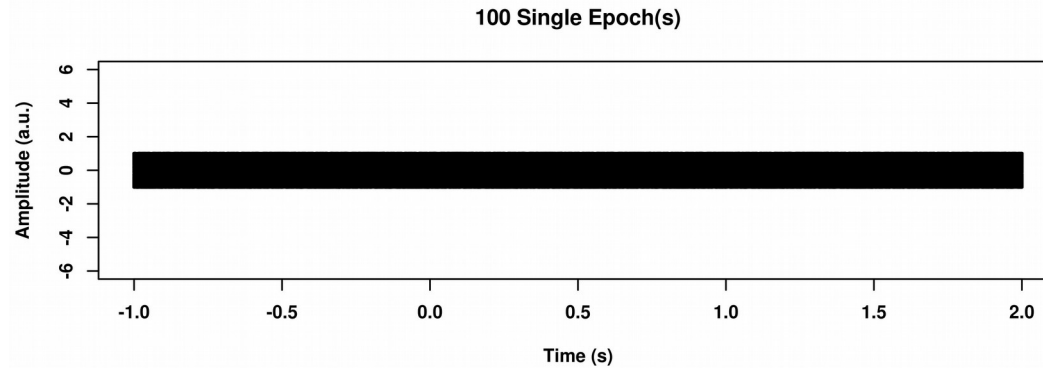
# IN-PHASE OR OUT-OF-PHASE



OUT OF PHASE,  
IDEAL, Real Mean = 1.0

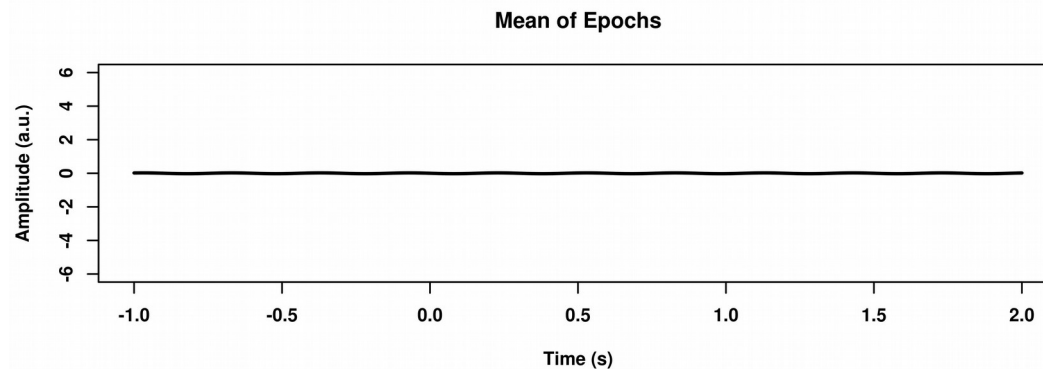
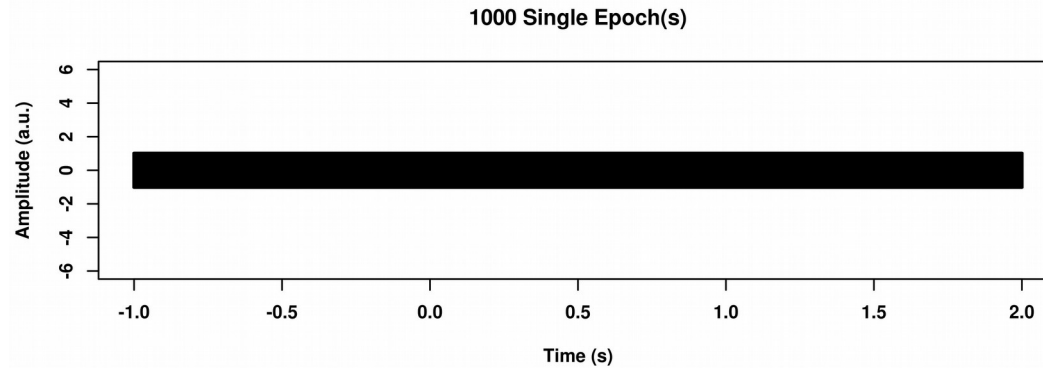


# IN-PHASE OR OUT-OF-PHASE



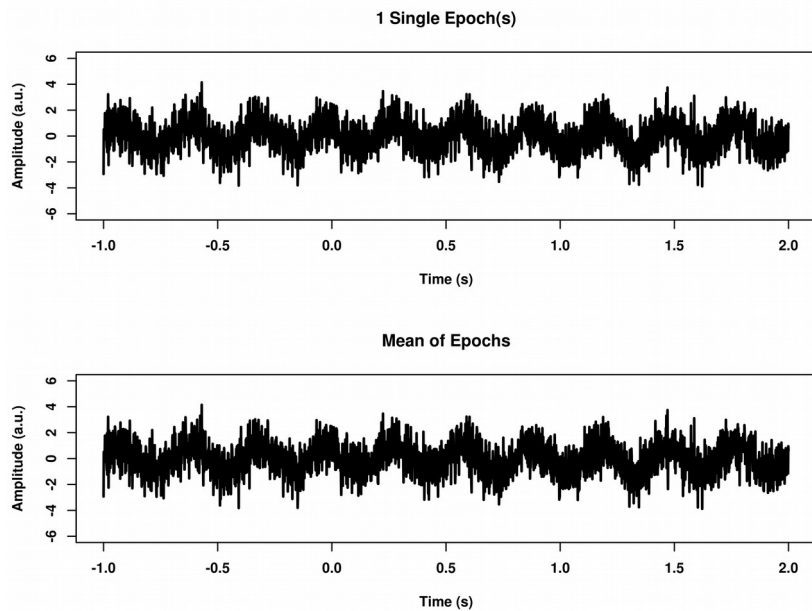
OUT OF PHASE,  
IDEAL, Real Mean = 1.0

# IN-PHASE OR OUT-OF-PHASE



OUT OF PHASE,  
IDEAL, Real Mean = 1.0

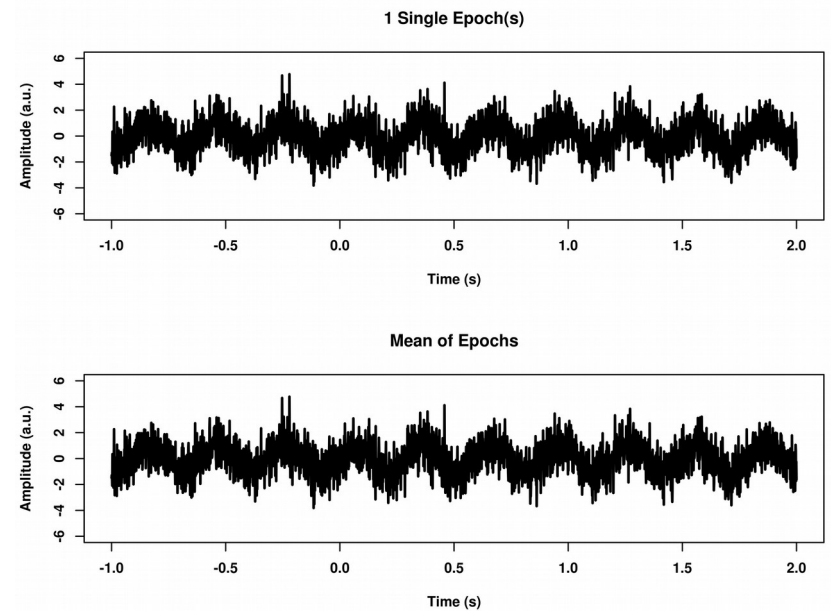
# IN-PHASE OR OUT-OF-PHASE WITH NOISE



IN PHASE

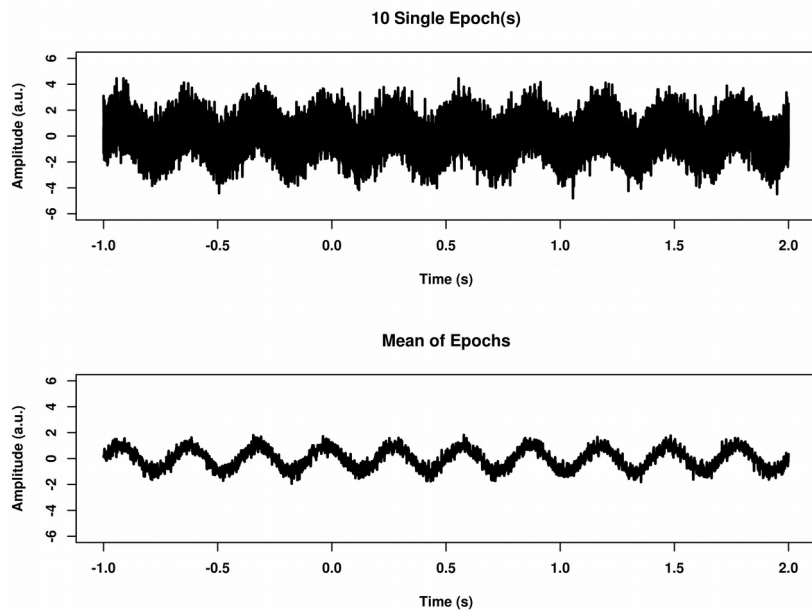
More realistic

Real Mean = 1.0



OUT OF PHASE

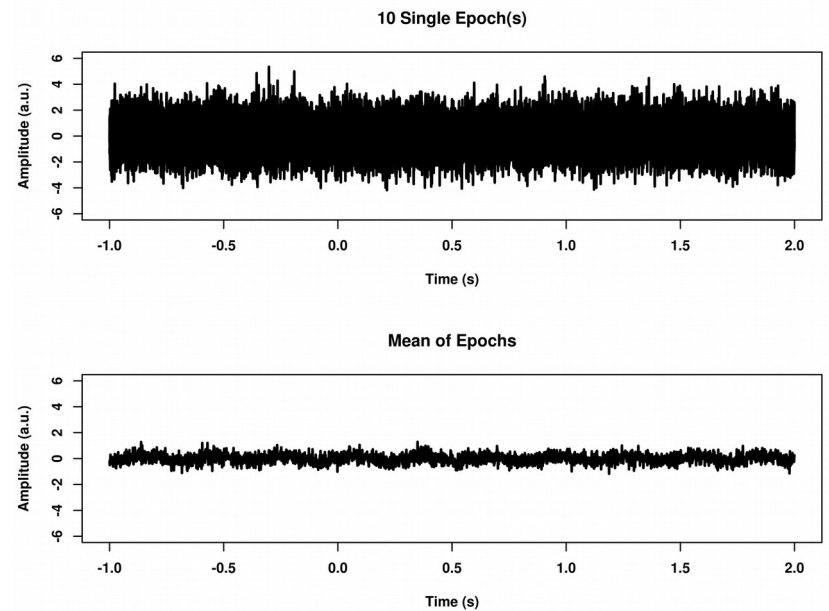
# IN-PHASE OR OUT-OF-PHASE WITH NOISE



IN PHASE

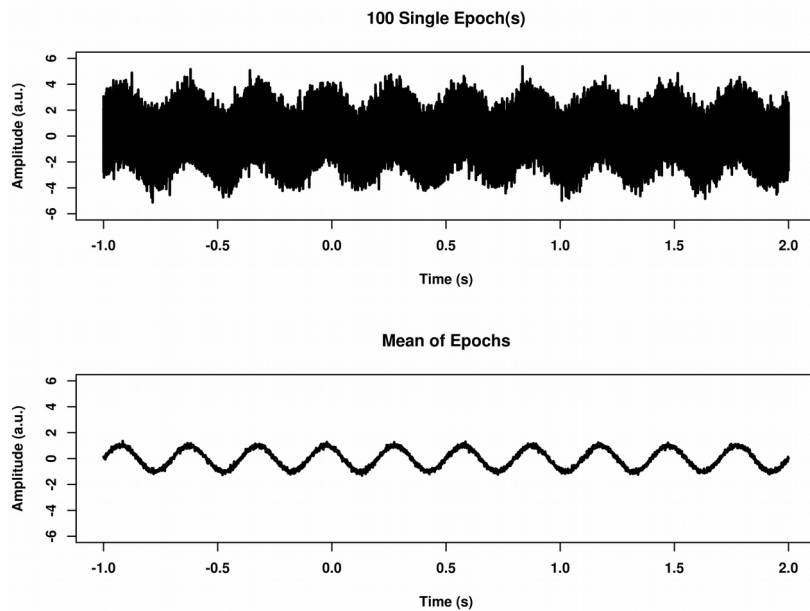
More realistic

Real Mean = 1.0



OUT OF PHASE

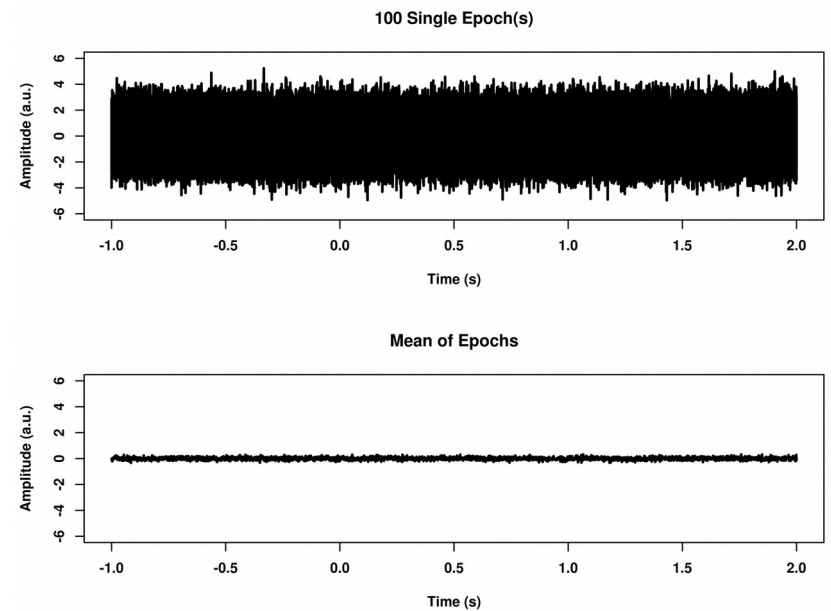
# IN-PHASE OR OUT-OF-PHASE WITH NOISE



IN PHASE

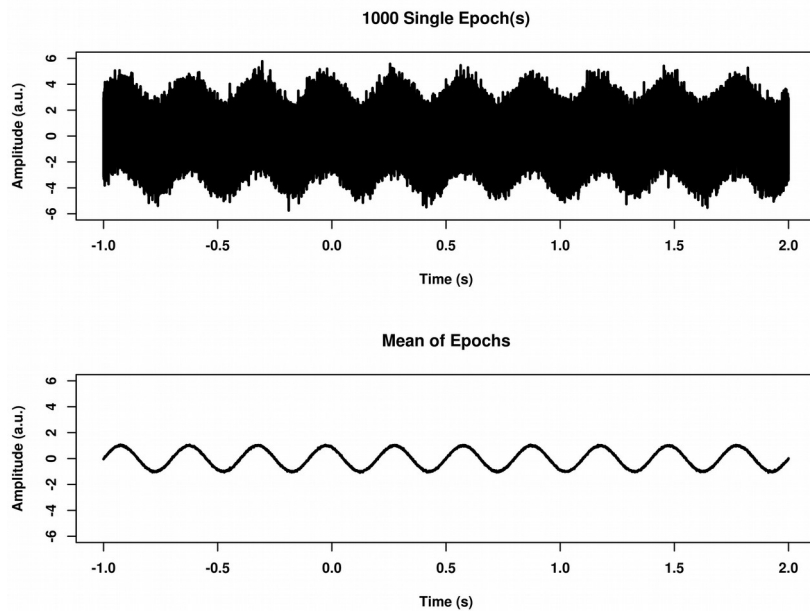
More realistic

Real Mean = 1.0



OUT OF PHASE

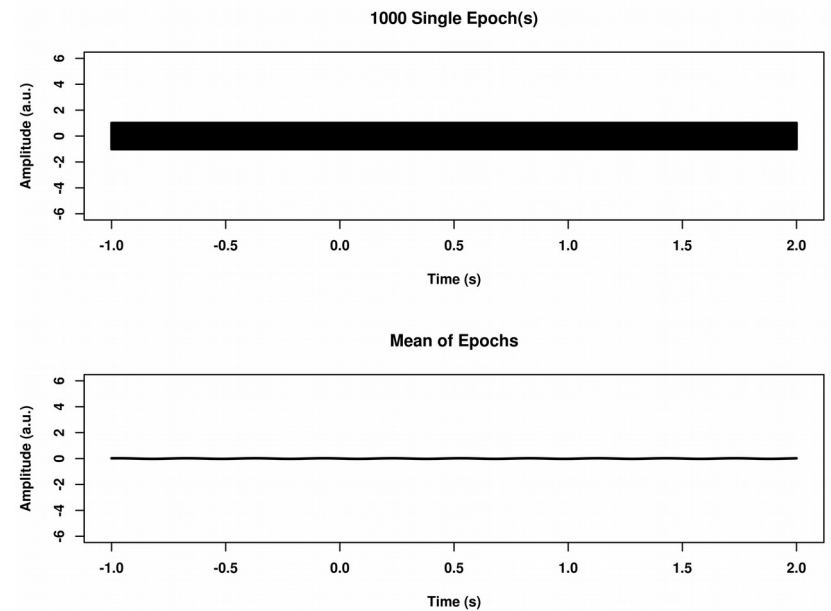
# IN-PHASE OR OUT-OF-PHASE WITH NOISE



IN PHASE

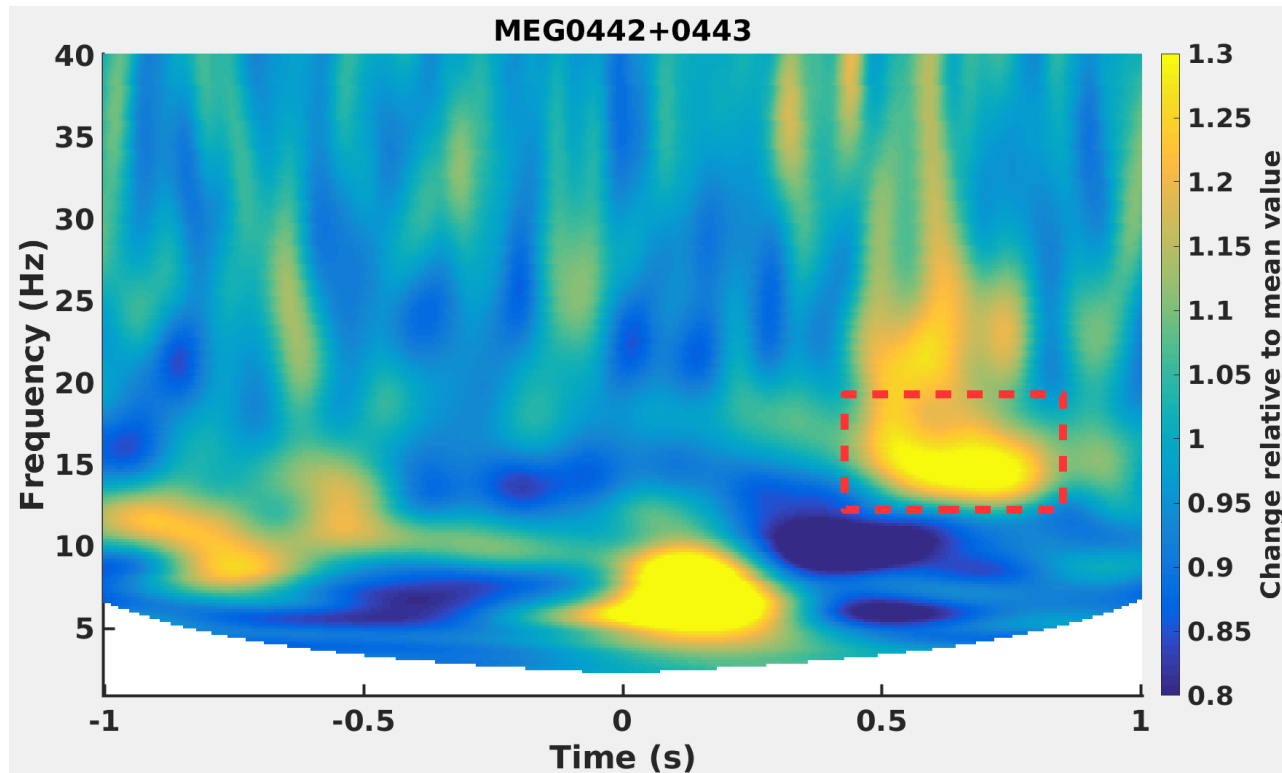
More realistic

Real Mean = 1.0

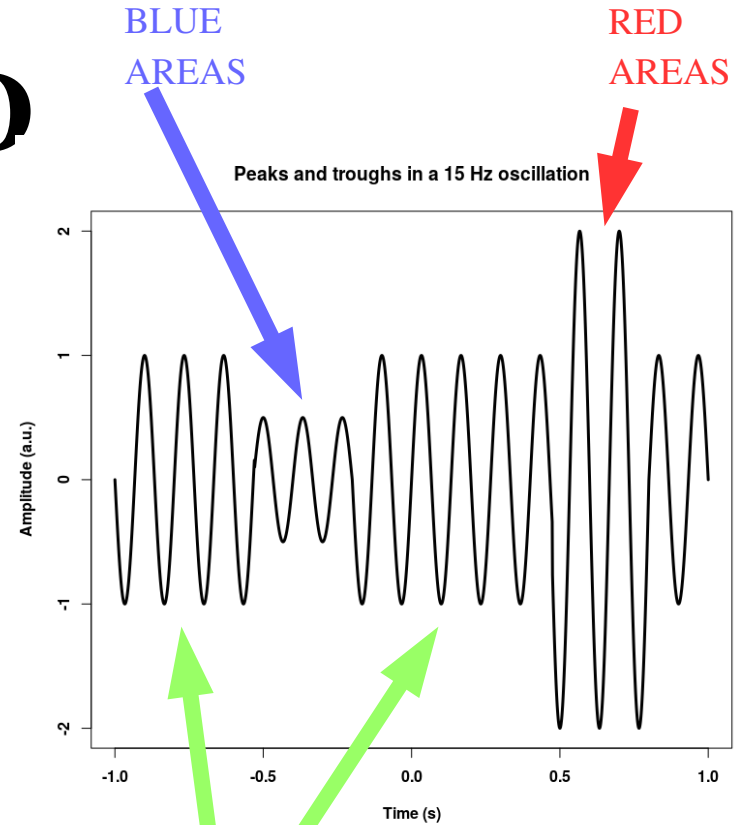
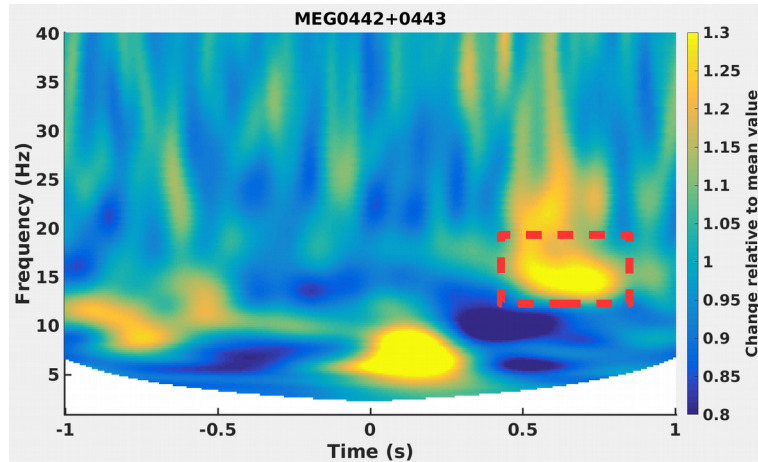


OUT OF PHASE

# BETA REBOUND – TIME-FREQUENCY REPRESENTATION (TFR)



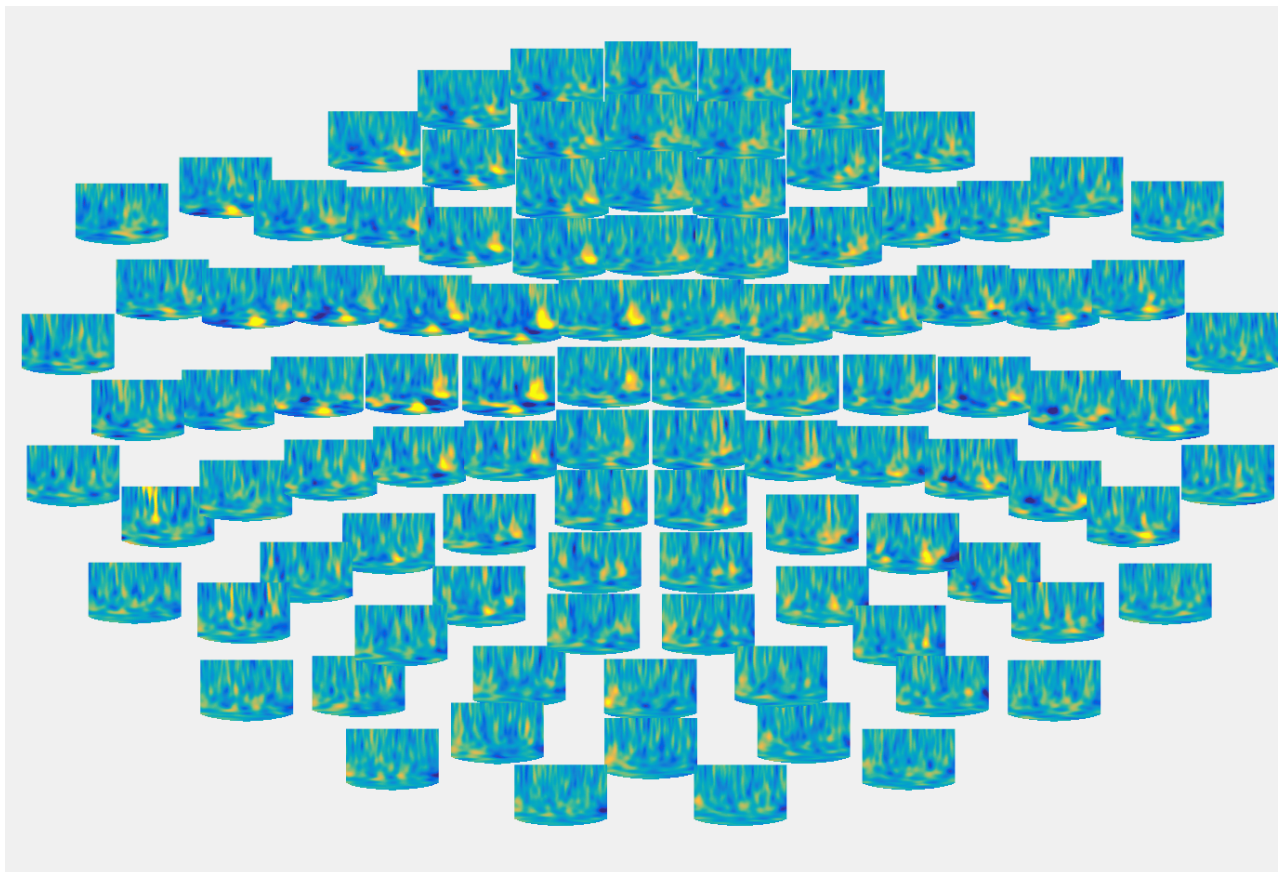
# BETA REBOUND



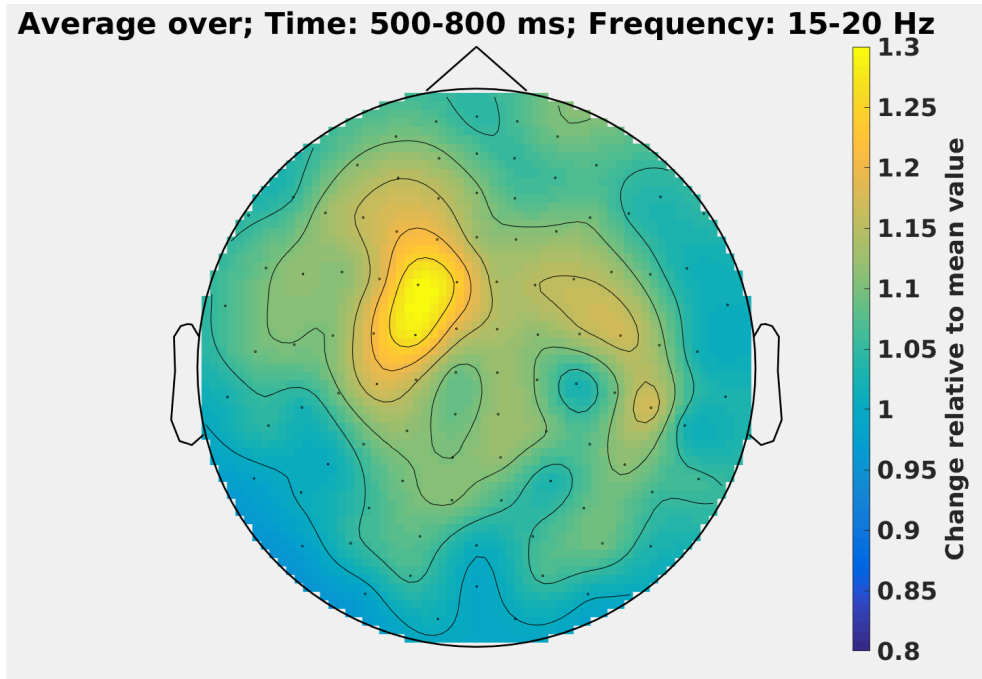
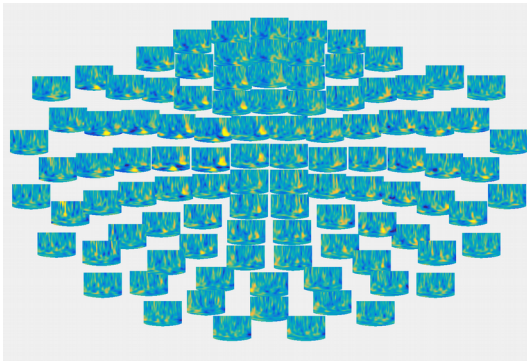
THESE CAN BE CREATED FOR ALL FREQUENCY BANDS



# OPERATIONS – TFR



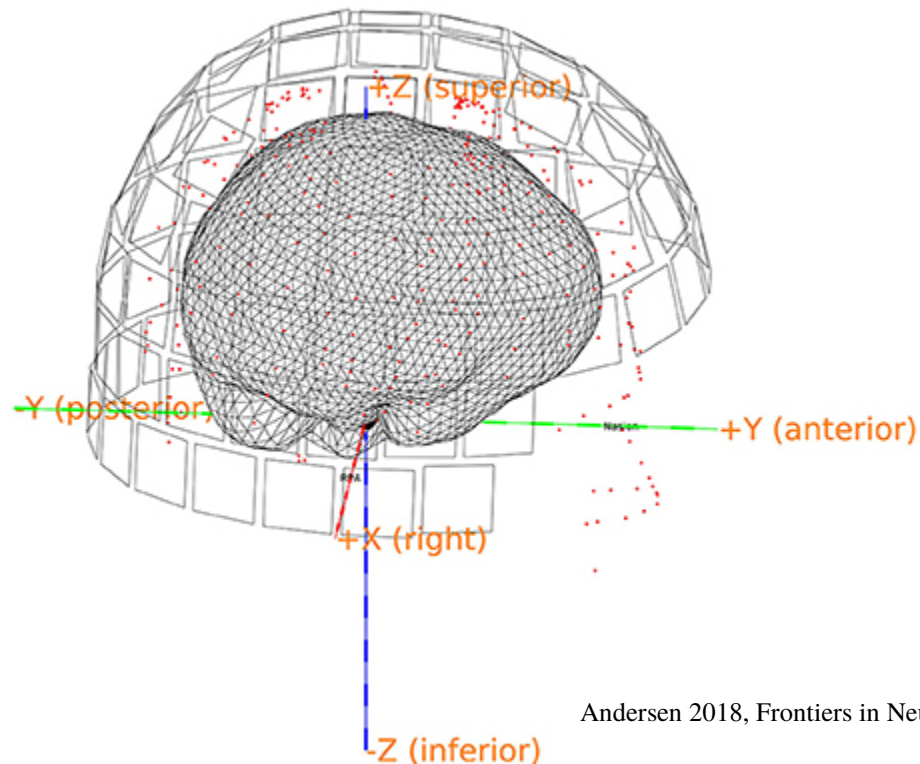
# OPERATIONS – TFR



# NEW QUESTION

Where does the beta rebound localize to?

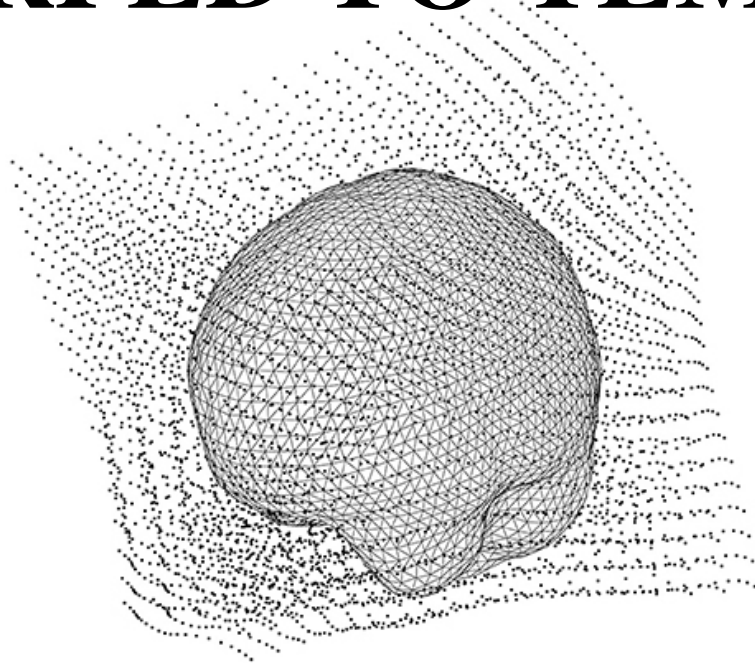
# OPERATIONS – CO-REGISTRATION



Andersen 2018, *Frontiers in Neuroscience*

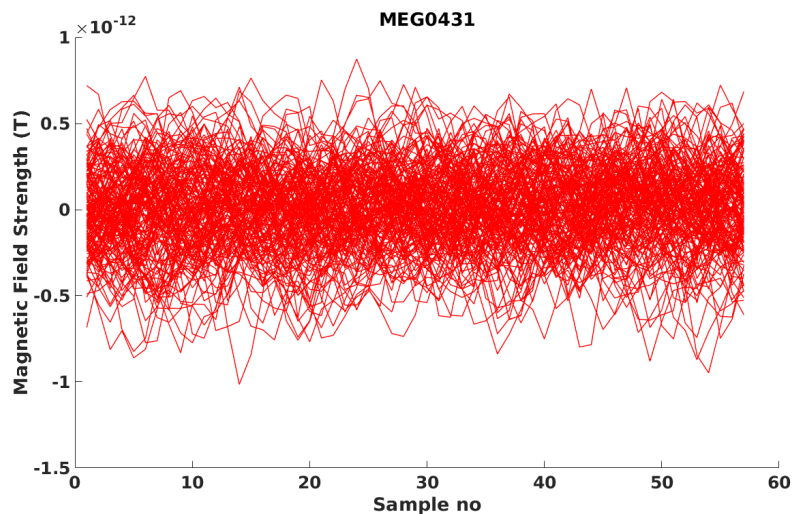
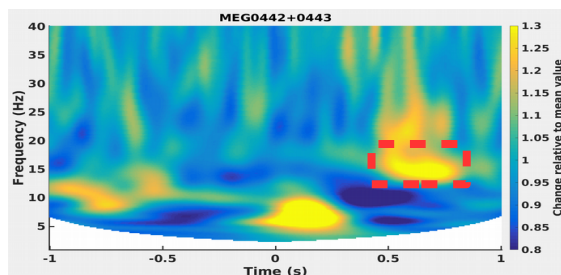
*How to analyse MEG?*

# OPERATIONS – LEADFIELD (FORWARD MODEL) WARPED TO TEMPLATE

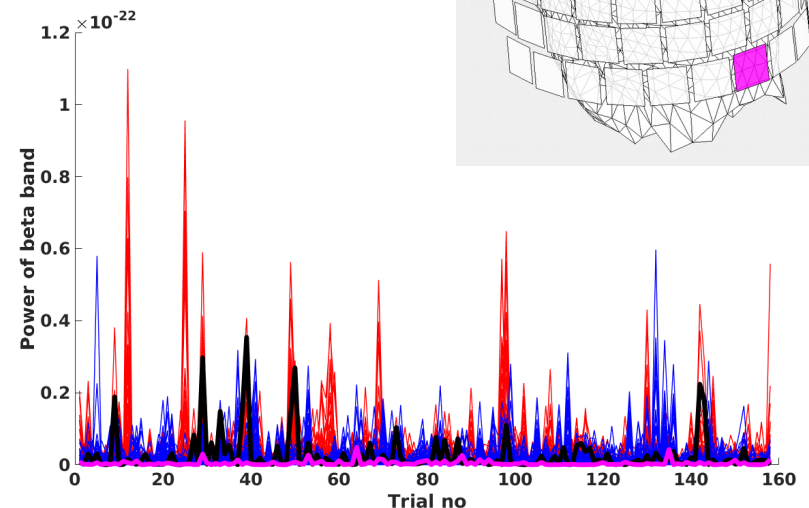
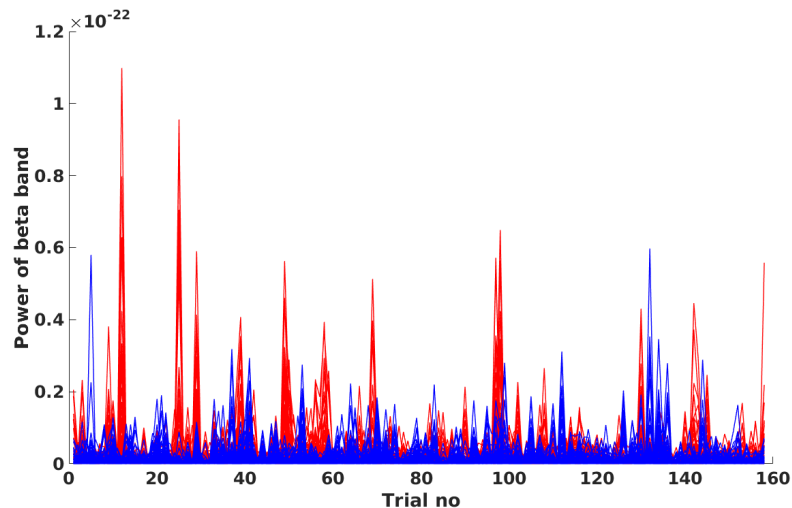
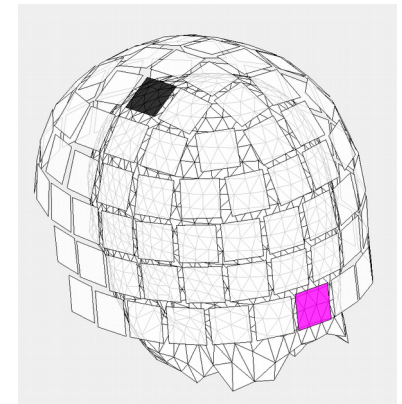
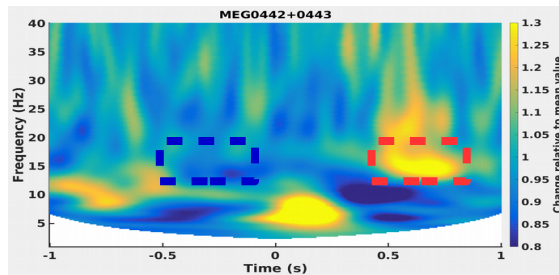


Andersen 2018, Frontiers in Neuroscience

# OPERATIONS – SIGNAL TO SOURCE RECONSTRUCT

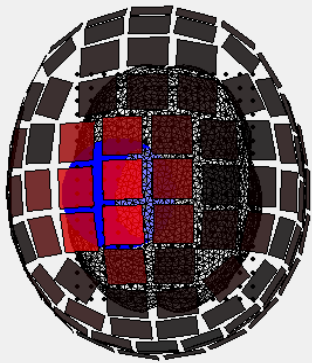


# OPERATIONS – SIGNAL TO SOURCE RECONSTRUCT

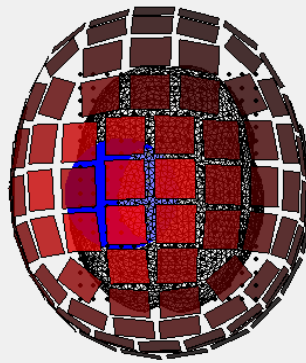


# OPERATIONS – LEADFIELD AND SPATIAL FILTER

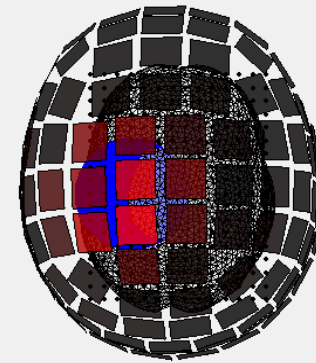
**Spatial filter**



**Leadfield**



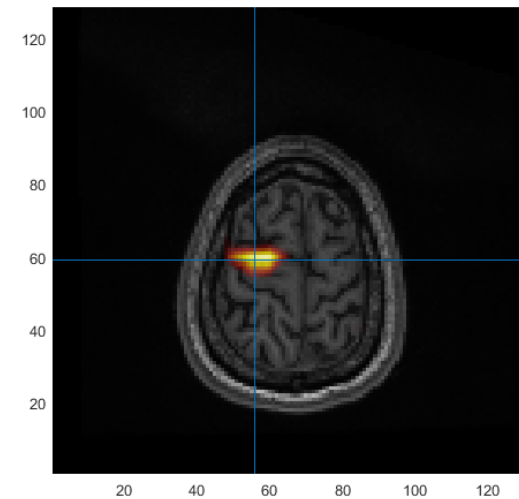
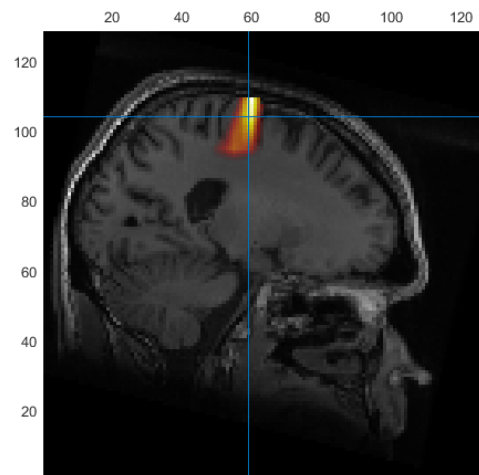
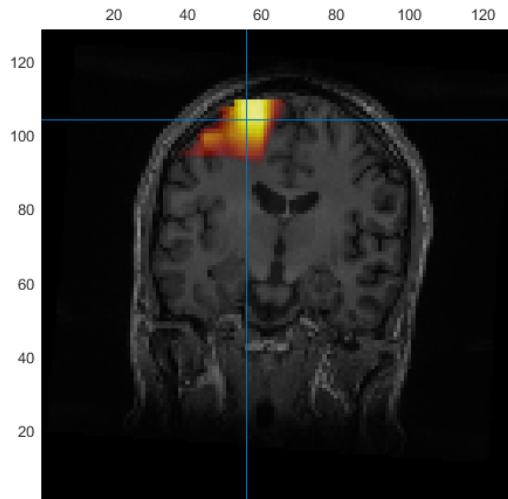
**Leadfield weighted by filter**



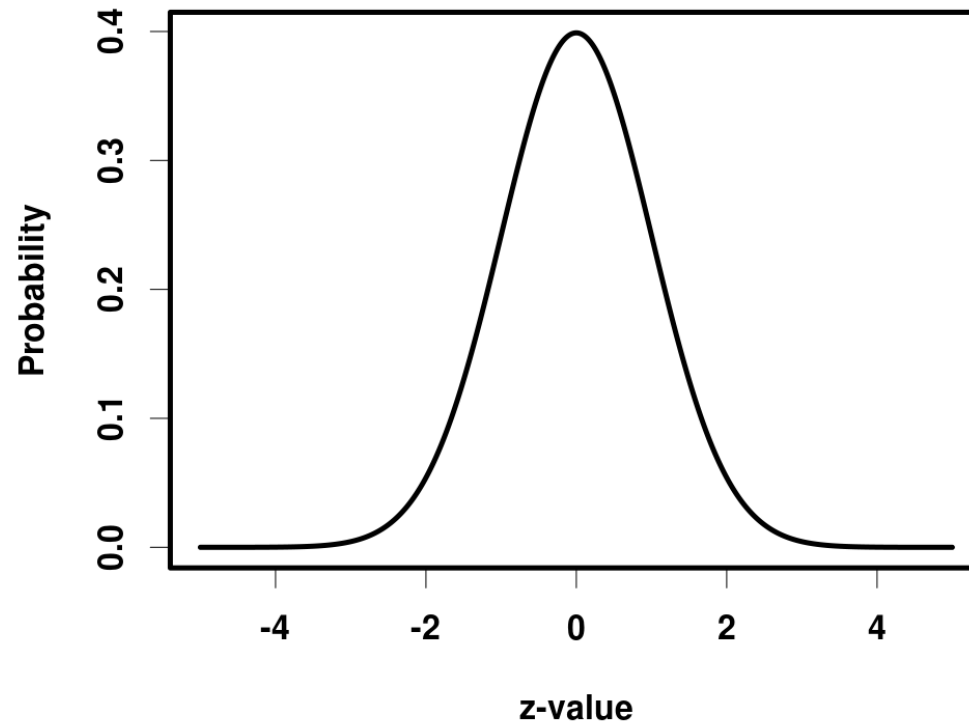


*How to analyse MEG?*

# OPERATIONS – BEAMFORMER



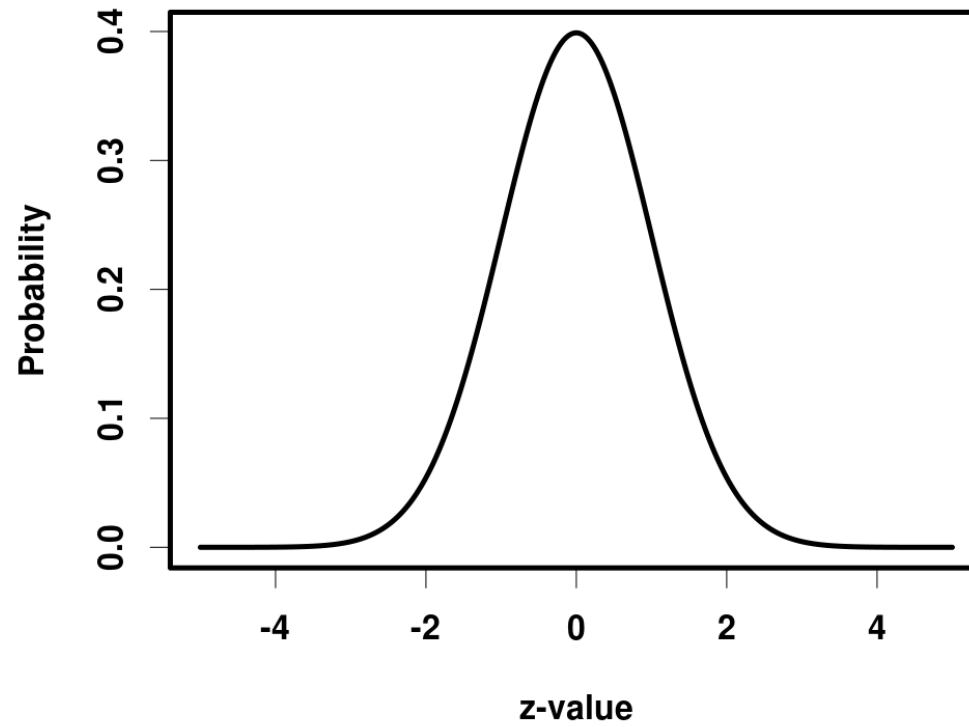
# Statistics



What is  
this?

# Statistics

Normal Distribution



# Statistics

$$Z = \frac{\bar{X} - P}{SE}$$

$$SE = \frac{\sigma}{\sqrt{n}}$$

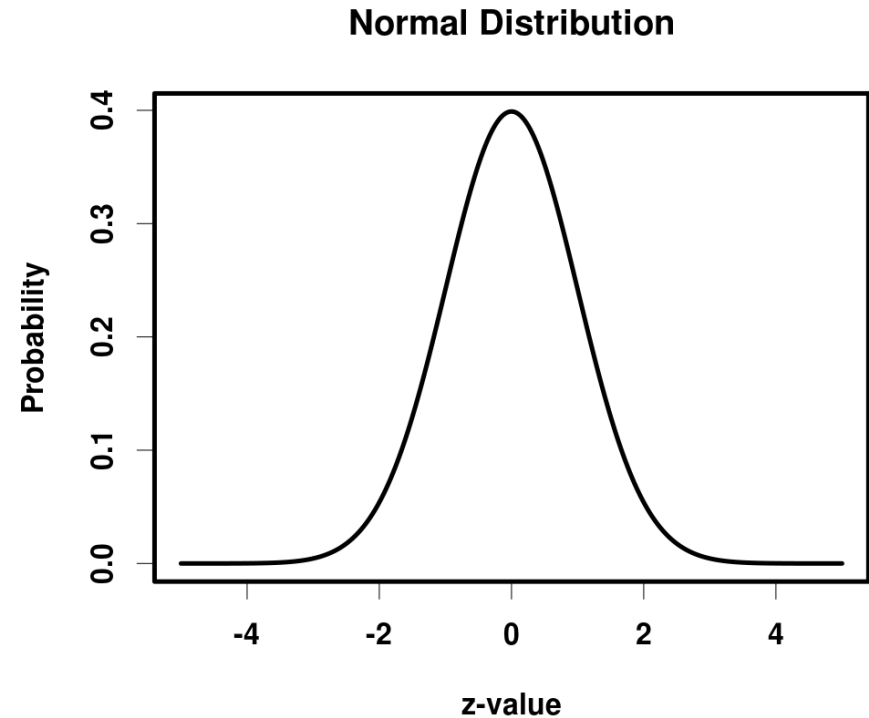
$\bar{X}$  : *sample mean*

$P$  : *population mean*

$SE$  : *standard error*

$\sigma$  : *standard deviation*

$n$  : *number of observations*



# Statistics

$$Z = \frac{\bar{X} - P}{SE}$$

$$SE = \frac{\sigma}{\sqrt{n}}$$

If we are testing the null hypothesis, then what is  $z$ ?

$\bar{X}$  : *sample mean*

$P$  : *population mean*

$SE$  : *standard error*

$\sigma$  : *standard deviation*

$n$  : *number of observations*

# Statistics

$$Z = \frac{\bar{X} - P}{SE}$$

$$SE = \frac{\sigma}{\sqrt{n}}$$

$\bar{X}$  : *sample mean*

$P$  : *population mean*

$SE$  : *standard error*

$\sigma$  : *standard deviation*

$n$  : *number of observations*

If we are testing the null hypothesis, then what is  $z$ ?

$$P = 0$$

$$z = \frac{\bar{X}}{SE}$$

# Statistics

What do we call the probability of observing a z-value or a z-value that is more extreme?

# Statistics

What do we call the probability of observing a z-value or a z-value that is more extreme?

*p*

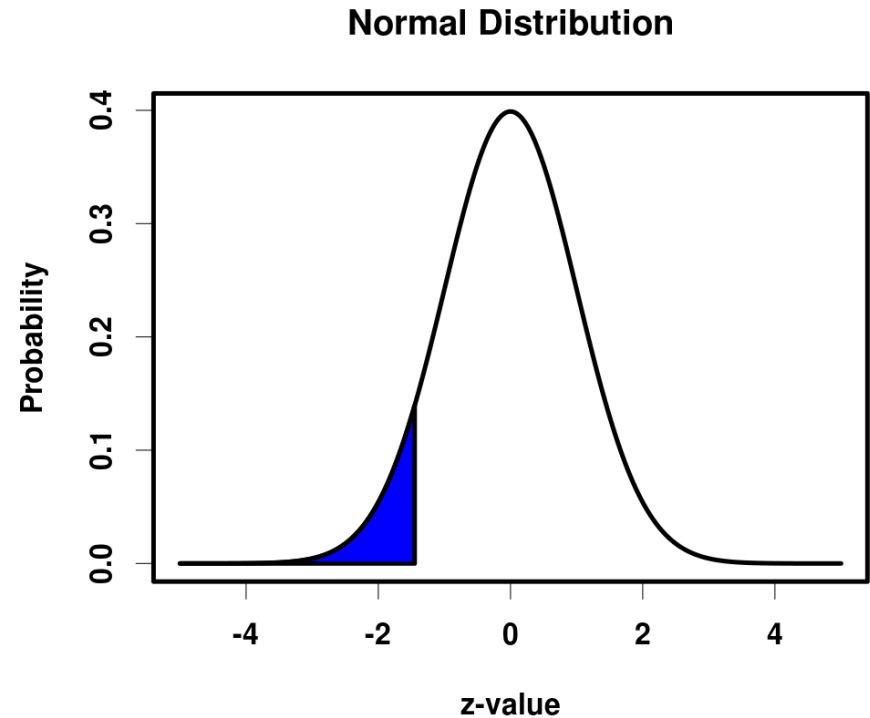


# Statistics

What do we call the probability of observing a z-value or a z-value that is more extreme?

$p$

If  $z = -1.45$ , what is  $p$  then?



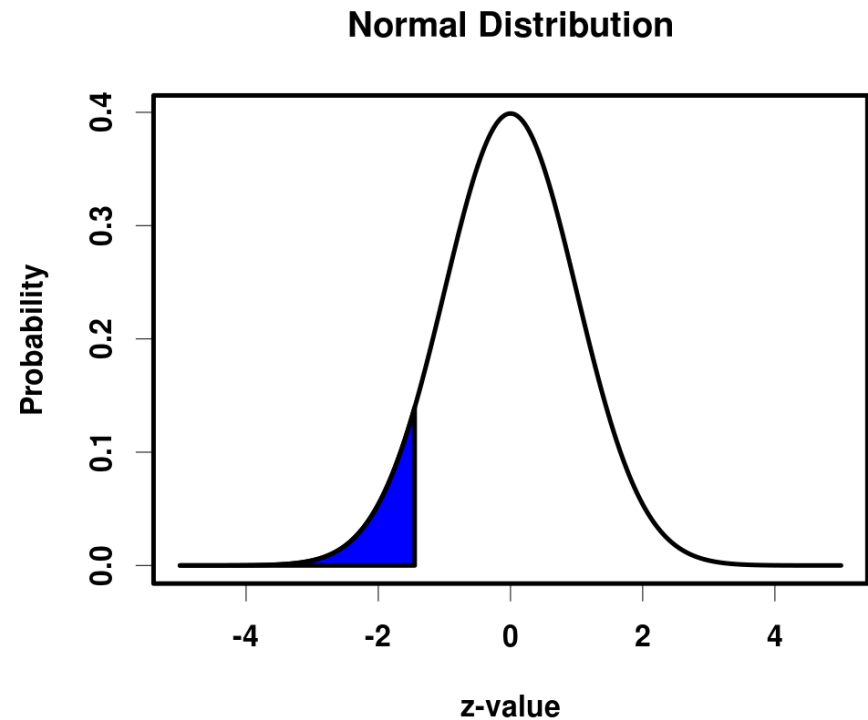
# Statistics

What do we call the probability of observing a z-value or a z-value that is more extreme?

$p$

If  $z = -1.45$ , what is  $p$  then?

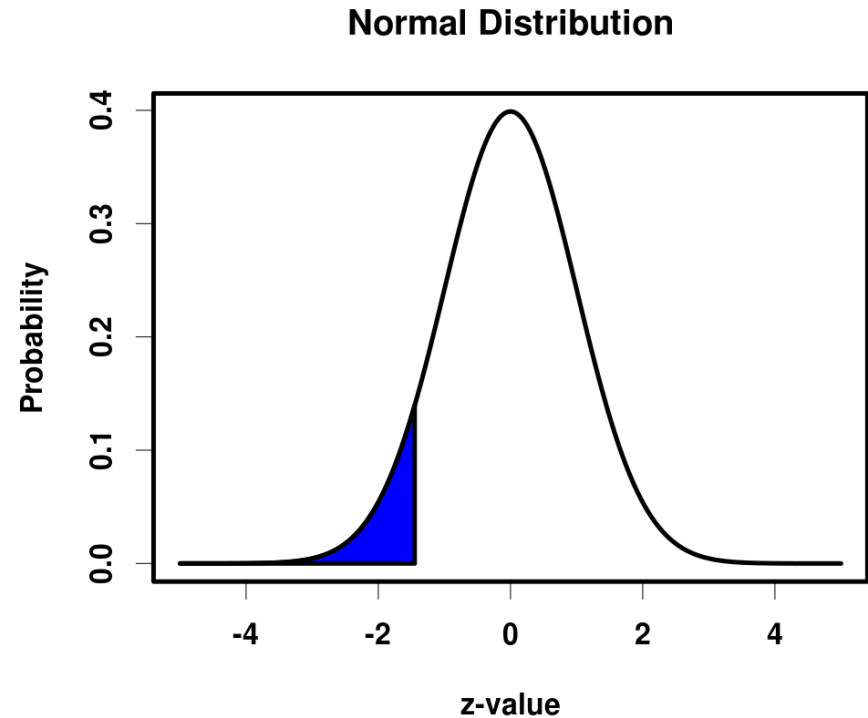
The area of the blue curve,  $p = 0.0735$



# Statistics

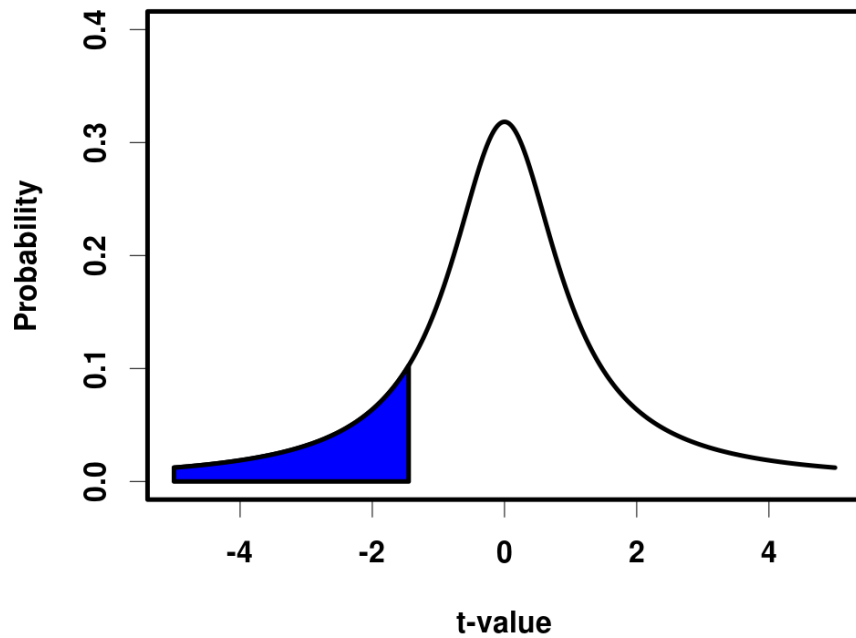
Crucial assumption when you are performing a z-test is that the population variance is known. When this is not known a *t*-test is more appropriate

We normally do not know the population variance



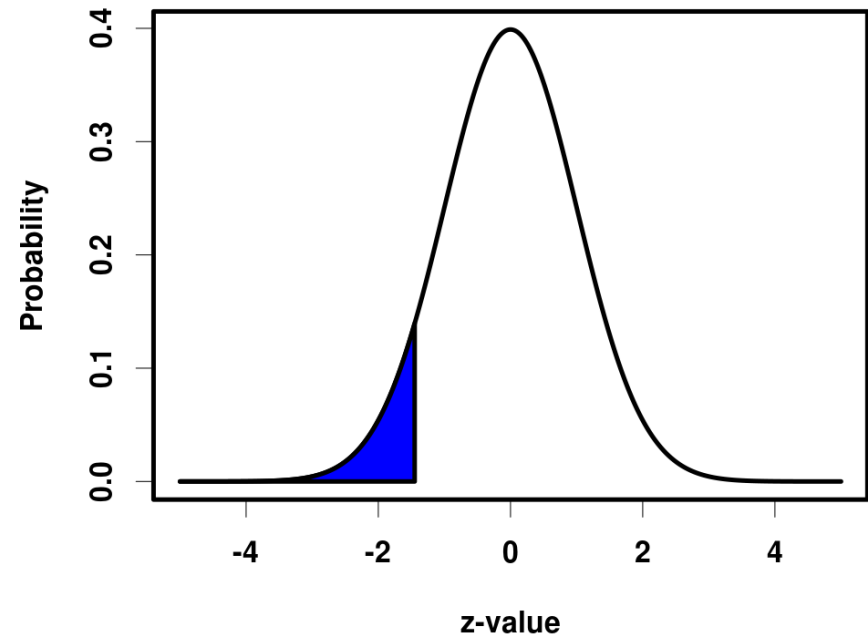
# Statistics – *t*-distribution

t-distribution, df=1



$$p = 0.192$$

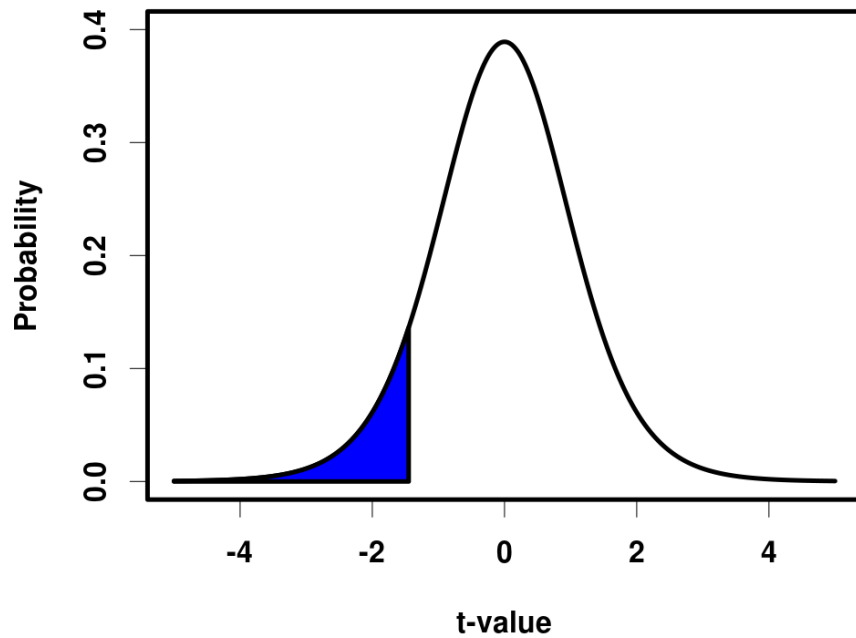
Normal Distribution



$$p = 0.0735$$

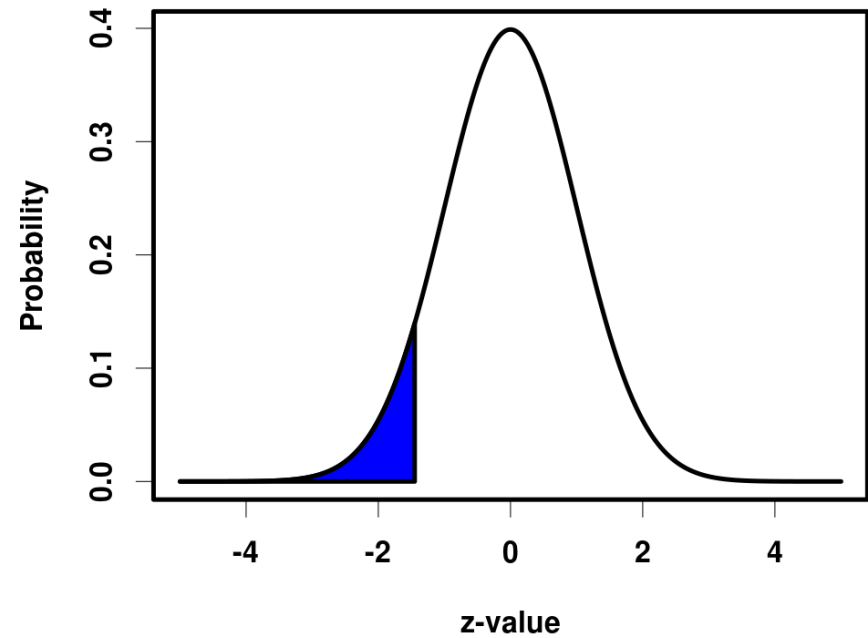
# Statistics – *t*-distribution

t-distribution, df=10



$$p = 0.0888$$

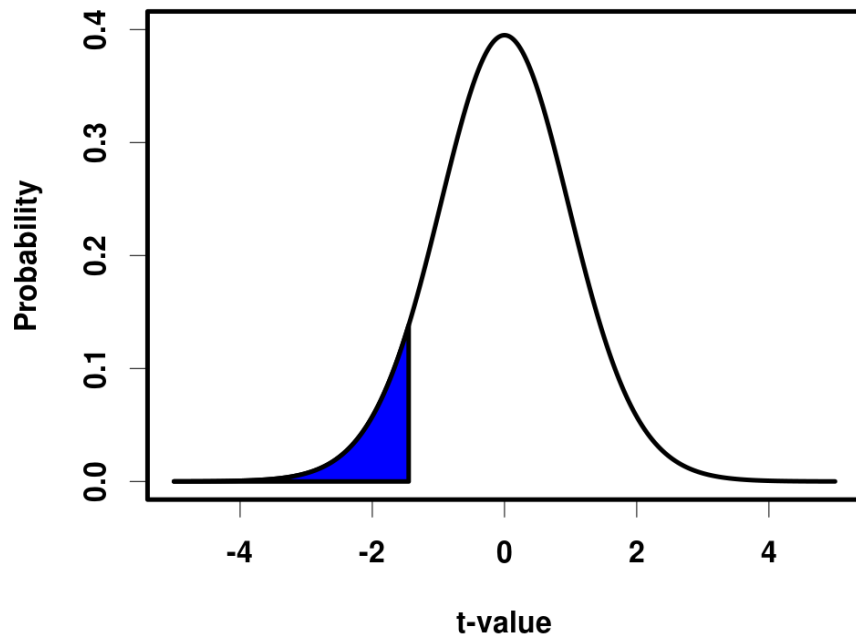
Normal Distribution



$$p = 0.0735$$

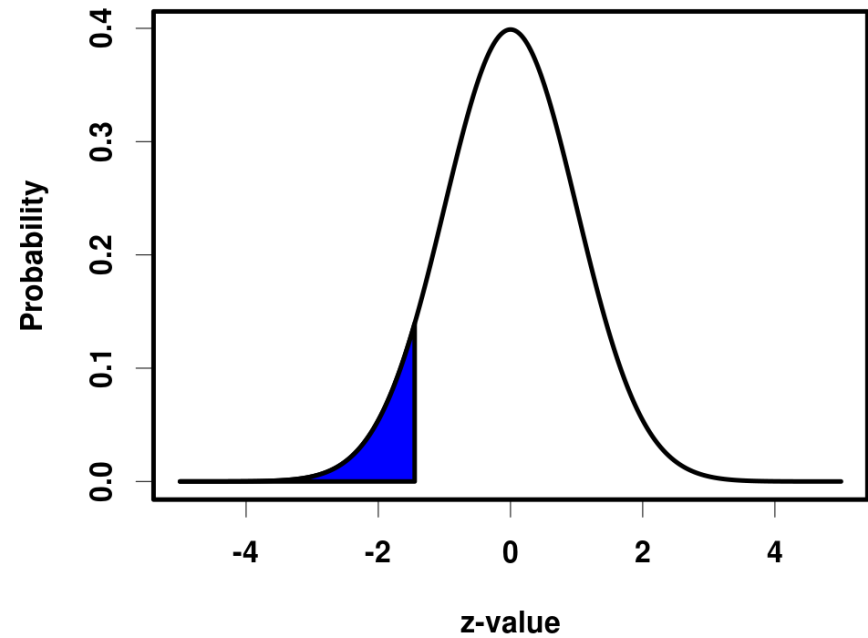
# Statistics – *t*-distribution

t-distribution, df=25



$$p = 0.0797$$

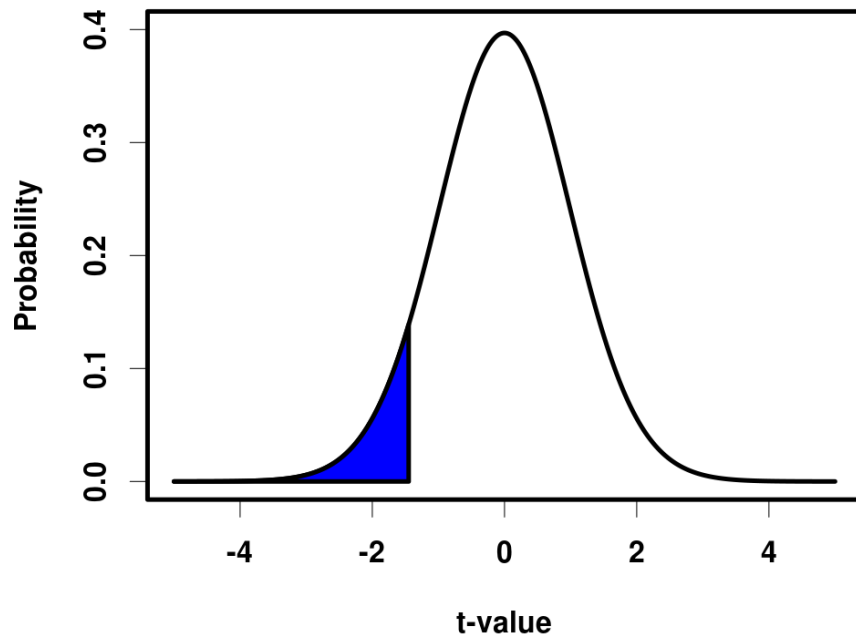
Normal Distribution



$$p = 0.0735$$

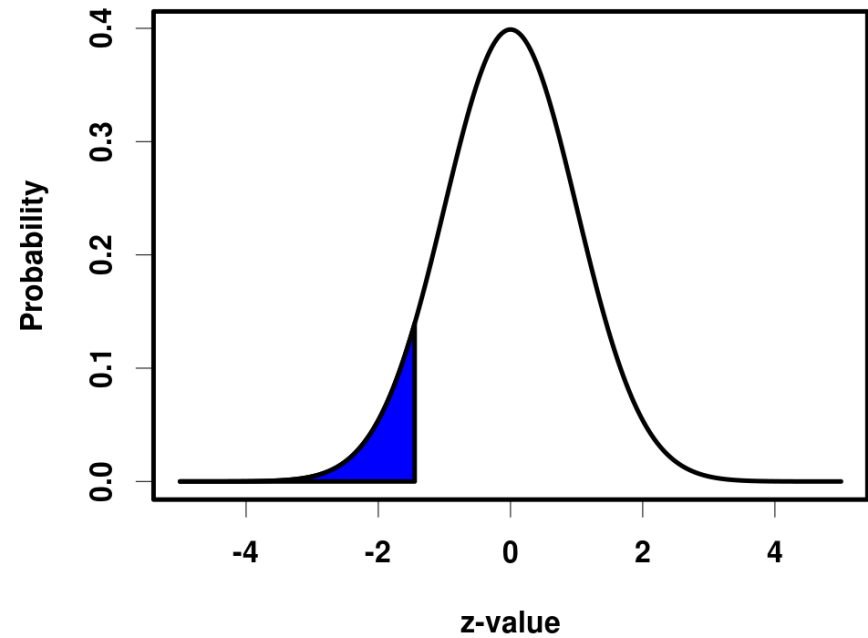
# Statistics – *t*-distribution

t-distribution, df=50



$$p = 0.0767$$

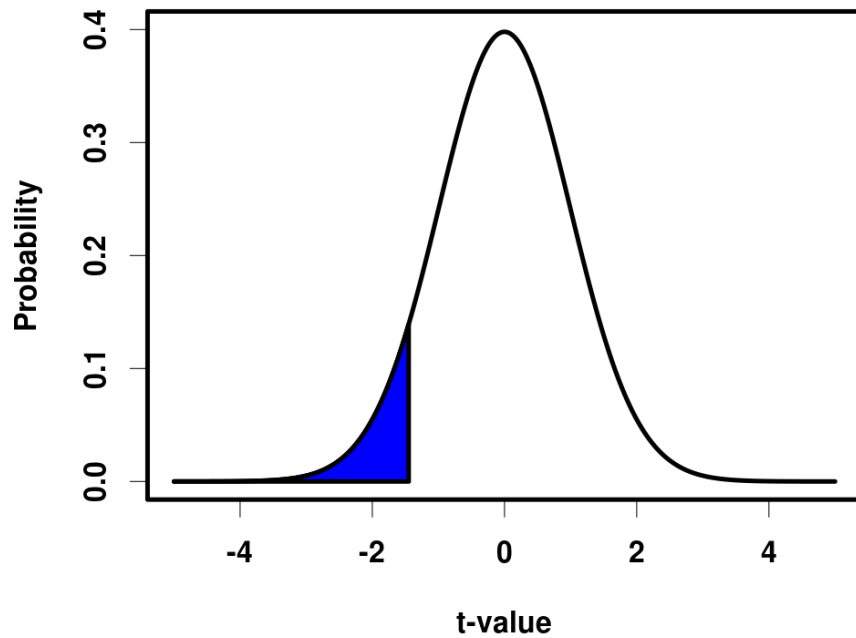
Normal Distribution



$$p = 0.0735$$

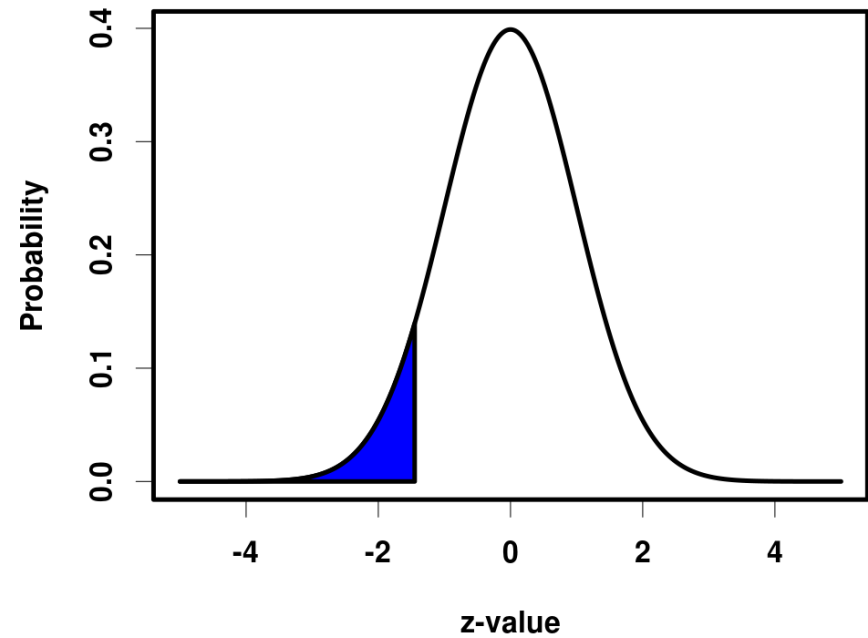
# Statistics – *t*-distribution

t-distribution, df=100



$$p = 0.0751$$

Normal Distribution

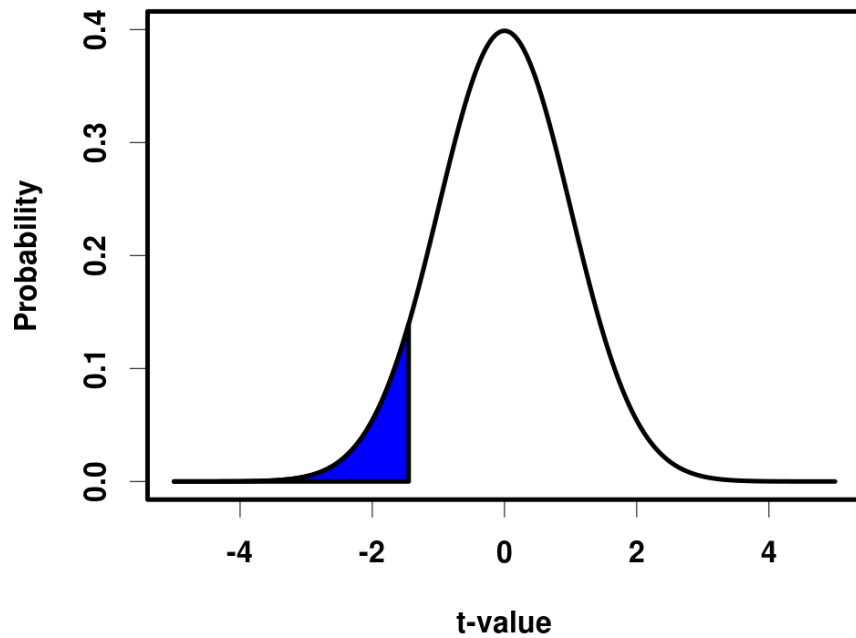


$$p = 0.0735$$



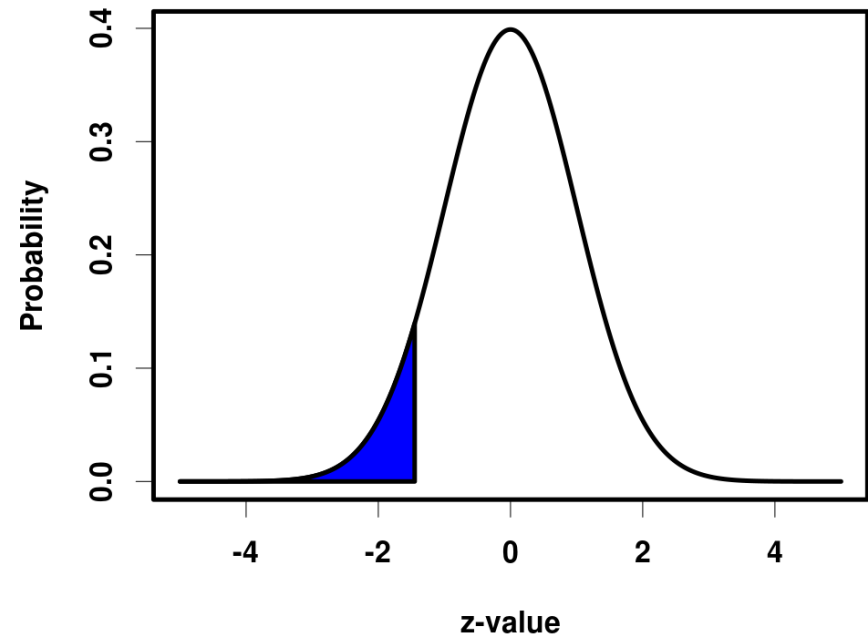
# Statistics – *t*-distribution

t-distribution, df=1000



$$p = 0.0737$$

Normal Distribution



$$p = 0.0735$$

# Two questions

- What does the  $t$ -distribution converge to when  $df$  goes to infinity?
- How are the  $t$ -value and the  $z$ -value related when doing null hypothesis testing?

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- What does the  $t$ -distribution converge to when  $df$  goes to infinity?
- How are the  $t$ -value and the  $z$ -value related when doing null hypothesis testing?

The  $z$ -distribution

$$t = z$$

# Two questions

- What does the  $t$ -distribution converge to when  $df$  goes to infinity?
- How are the  $t$ -value and the  $z$ -value related when doing null hypothesis testing?

The  $z$ -distribution

$$t = z = \frac{\bar{X}}{SE}$$

# Two questions

- What does the  $t$ -distribution converge to when  $df$  goes to infinity?
- How are the  $t$ -value and the  $z$ -value related when doing null hypothesis testing?

The  $z$ -distribution

$$t = z = \frac{\bar{X}}{SE}$$

Thus,  $t$ -values are independent of  $df$ . It is the  $p$ -value associated with the  $t$ -value that is dependent on  $df$

# $\alpha$

$\alpha$  signifies the cut-off point for  $p$ -values, which is used to decide whether to reject the null hypothesis and accept the alternative hypothesis, or whether to conclude that the null hypothesis could not be rejected

$\alpha$  also signifies the risk of making a Type-I error, i.e. the error of accepting a false positive, i.e. rejecting the null hypothesis when it is actually true, every time a test is made

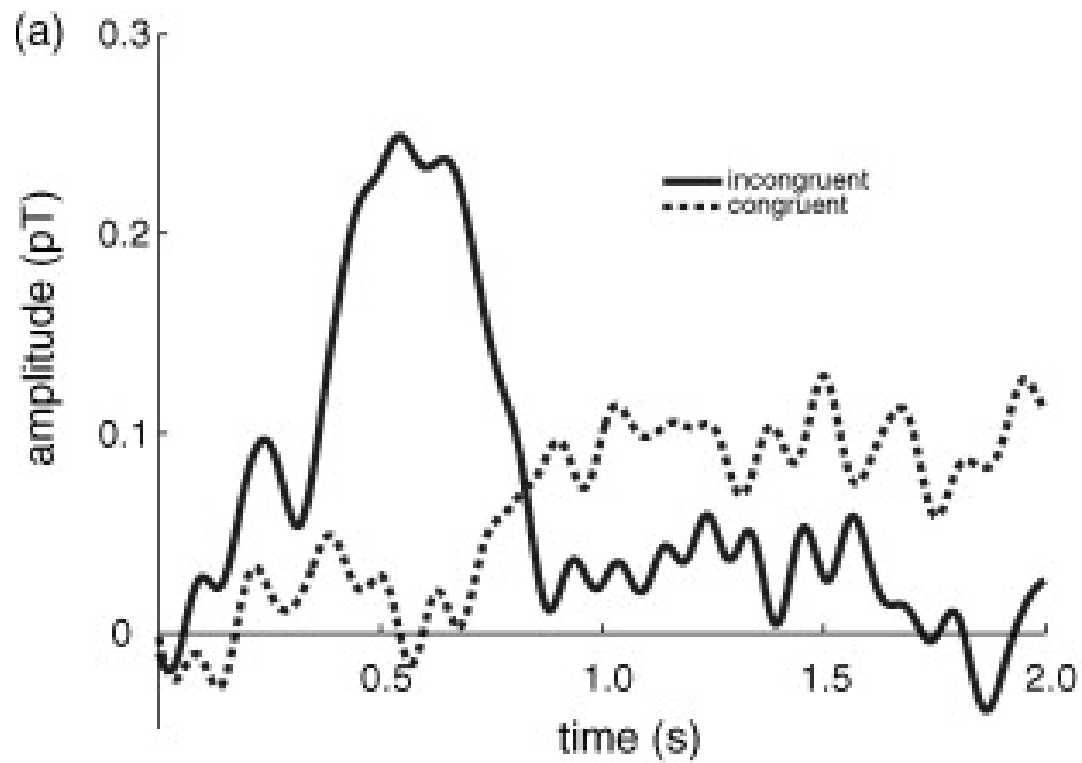
$\alpha$  is canonically set at 0.05 for reasons unbeknownst to man

# Statistics – $t$ -distribution

The  $t$ -test works fine for many things, for example testing the quality of beer batches (Guinness), which it is what is was originally developed for

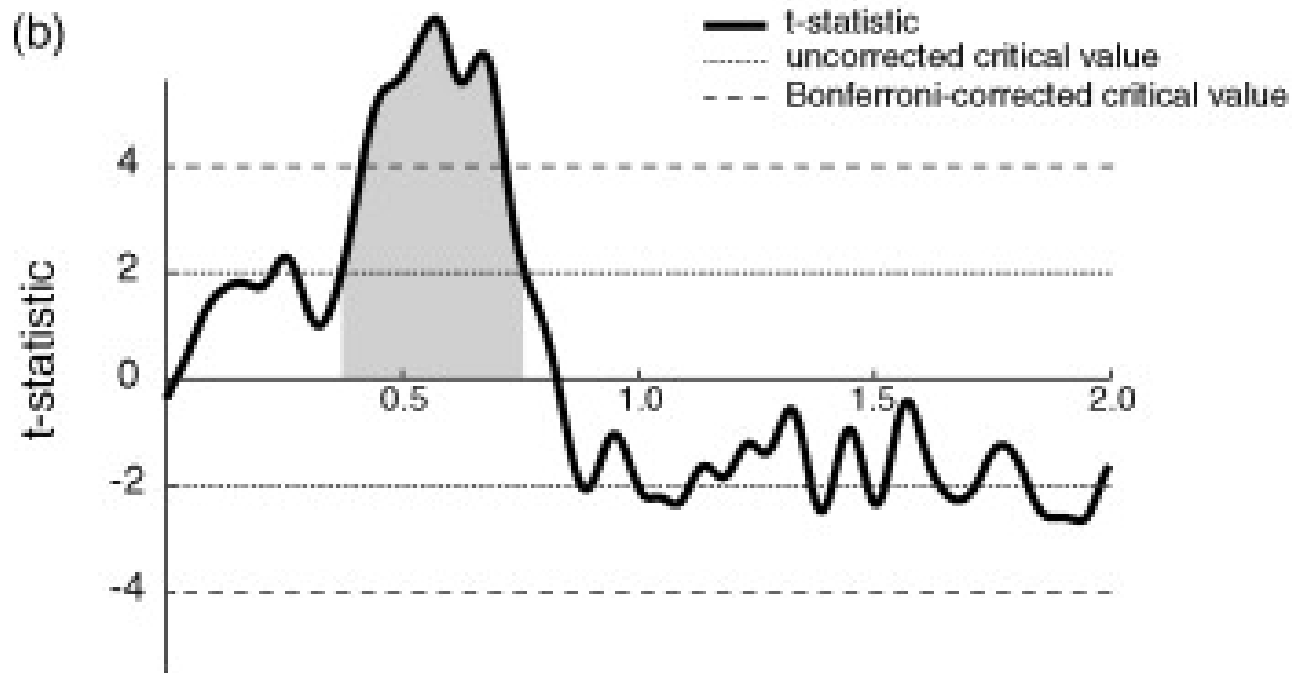


# Evoked example





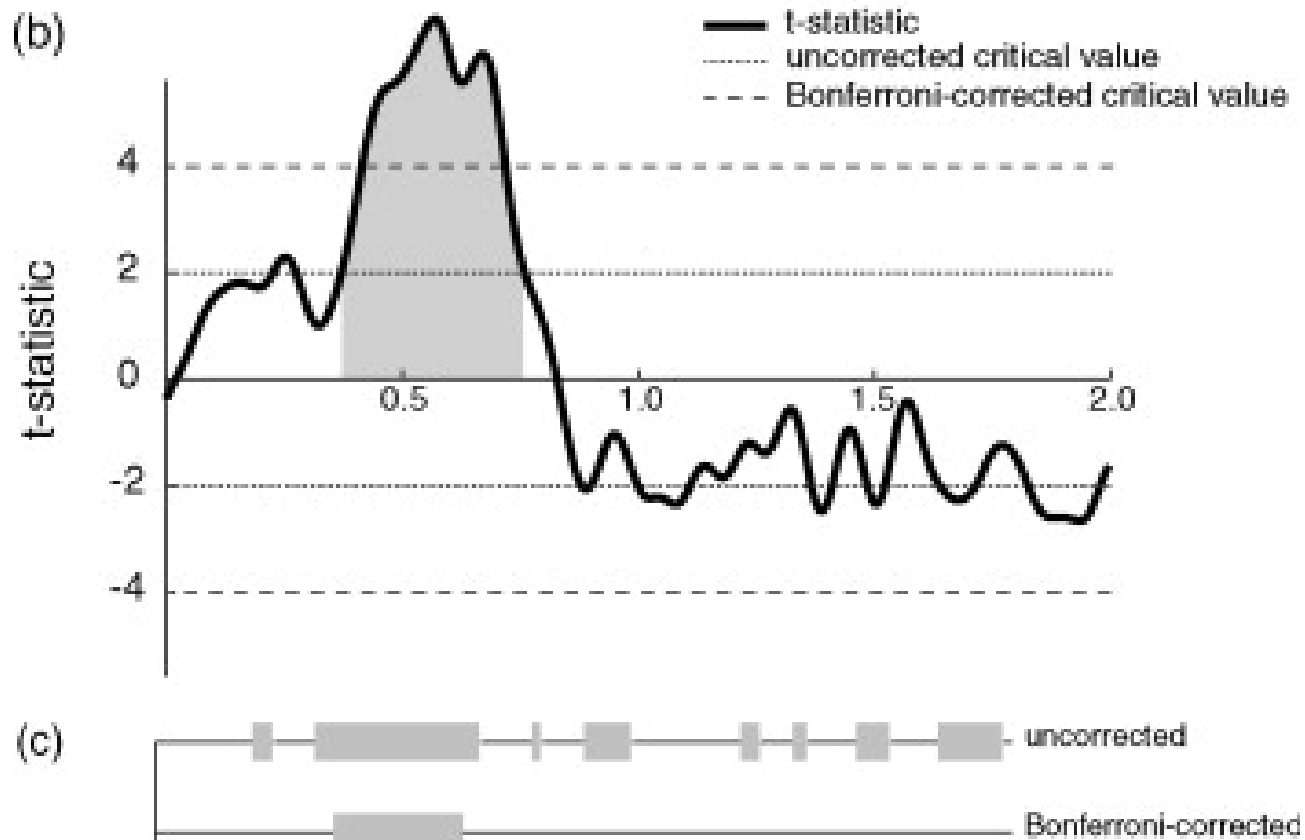
# Evoked example



$\alpha$  also signifies the risk of making a Type-I error, i.e. the error of accepting a false positive, i.e. rejecting the null hypothesis when it is actually true, every time a test is made

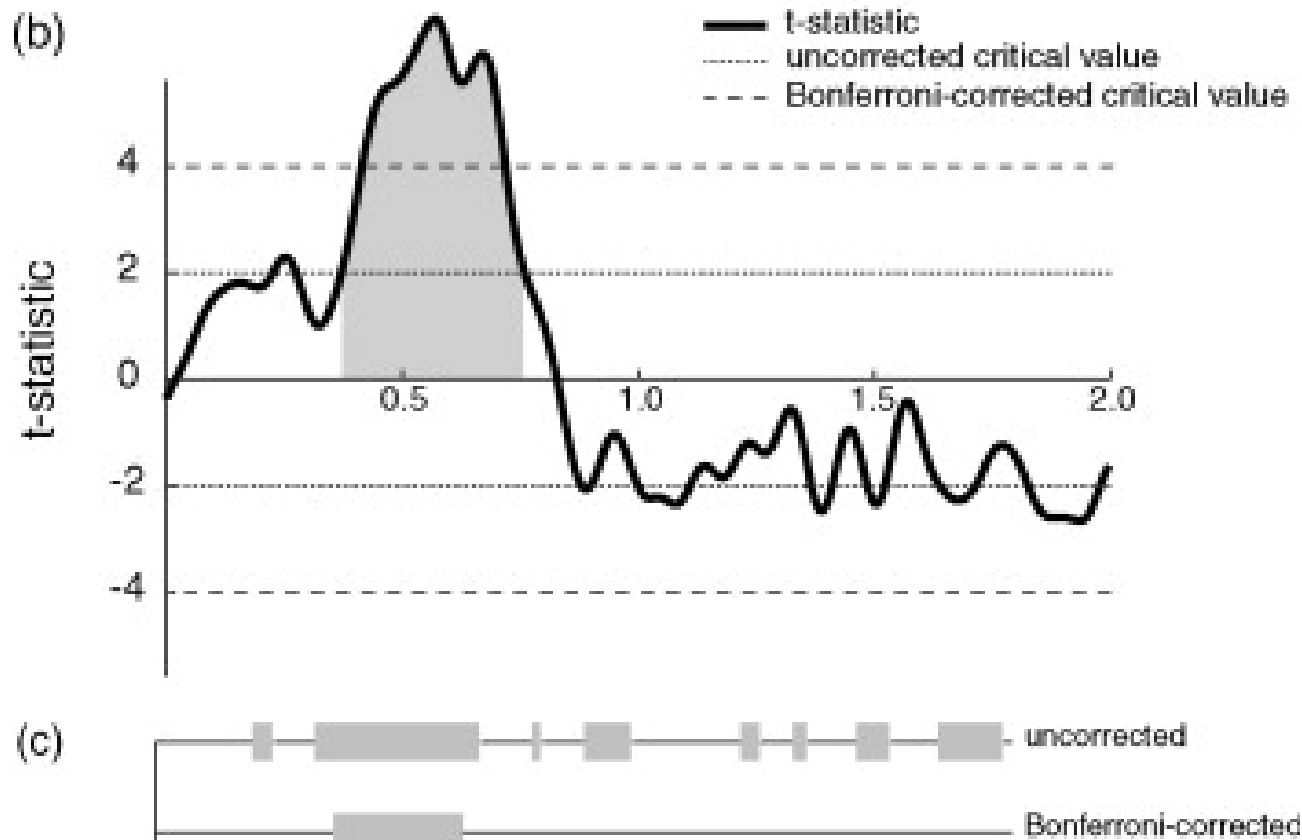
This leads to the Multiple Comparisons Problem

# Evoked example



$$\alpha_{BONF} = \frac{\alpha}{n \text{ tests}}$$

# Evoked example



$$\alpha_{BONF} = \frac{\alpha}{n \text{ tests}}$$

Goes well enough with one sensor

What about when we have 306  
MEG channels and 128  
electrodes each sampled 1000  
times per second?

What about when we have 306  
MEG channels and 128  
electrodes each sampled 1000  
times per second?

That amounts to 434,000 possible tests per second

Uncorrected testing will lead to way too many false  
positives

Bonferroni-corrected test will lead to no true positives

# Statistics – *t*-distribution

Can anyone name an assumption of the *t*-test that makes it less than optimal for MEEG data?

# Statistics – $t$ -distribution

Can anyone name an assumption of the  $t$ -test that makes it less feasible for MEEG data?

It requires that tested samples are independent from one another, which MEEG samples are evidently not. (MEEG data show strong spatio-temporal correlations)

# Permutation tests

They don't have strong assumptions about the underlying structure of the data



# Nonparametric statistical testing of EEG- and MEG-data<sup>☆,☆☆</sup>

Eric Maris<sup>a,b,\*</sup>, Robert Oostenveld<sup>b</sup>

<sup>a</sup> *NICI, Biological Psychology, Radboud University Nijmegen, Nijmegen, The Netherlands*

<sup>b</sup> *F.C. Donders Center for Cognitive Neuroimaging, Radboud University Nijmegen, Nijmegen, The Netherlands*

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## Abstract

In this paper, we show how ElectroEncephaloGraphic (EEG) and MagnetoEncephaloGraphic (MEG) data can be analyzed statistically using nonparametric techniques. Nonparametric statistical tests offer complete freedom to the user with respect to the test statistic by means of which the experimental conditions are compared. This freedom provides a straightforward way to solve the multiple comparisons problem (MCP) and it allows to incorporate biophysically motivated constraints in the test statistic, which may drastically increase the sensitivity of the statistical test. The paper is written for two audiences: (1) empirical neuroscientists looking for the most appropriate data analysis method, and (2) methodologists interested in the theoretical concepts behind nonparametric statistical tests. For the empirical neuroscientist, a large part of the paper is written in a tutorial-like fashion, enabling neuroscientists to construct their own statistical test, maximizing the sensitivity to the expected effect. And for the methodologist, it is explained why the nonparametric test is formally correct. This means that we formulate a null hypothesis (identical probability distribution in the different experimental conditions) and show that the nonparametric test controls the false alarm rate under this null hypothesis. © 2007 Elsevier B.V. All rights reserved.

*Keywords:* Nonparametric statistical testing; Hypothesis testing; EEG; MEG; Multiple comparisons problem

---

# Overview of logic

- 1) Create a design matrix indicating how data are organized and what conditions should be compared against one another
- 2) Do a  $t$ -tests on all samples of interest
- 3) Find the  $T$ -values for each cluster of  $t$ -values passing the analytic significance test
- 4) Permute the condition labels (between 1000-10000 times) using a random mechanism and find the  $T$ -values for each of the permutations.
- 5) Find the distribution of permutation-based  $T$ -values
- 6) Compare the  $T$ -value based on the design matrix with the permutation distribution and use the  $p$ -value together with a pre-set alpha to decide whether or not to reject the null hypothesis

# Design matrix

Unit variable	Independent variable
1	1
2	1
...	...
19	1
20	1
1	2
2	2
...	...
19	2
20	2

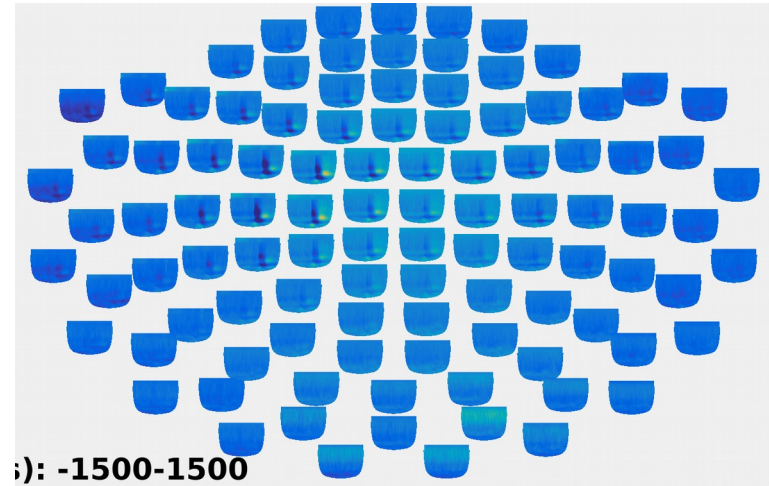
```
cfg = [];  
cfg.latency = [1.500 1.500];  
cfg.frequency = [1 100];  
cfg.parameter = 'powspctrm';  
cfg.method = 'analytic'; %% do a mass-univariate  
test  
cfg.alpha = 0.05; %% critical value around ± 2.09  
cfg.statistic = 'depsamplesT'; % use a dependent  
samples t-test  
cfg.design(1, :) = [1:n_subjects 1:n_subjects];  
cfg.design(2, :) = [ones(1, n_subjects) 2 * ones(1,  
n_subjects)];  
cfg.uvar = 1; % first row of cfg.design, containing  
the (u)nits (subjects)  
cfg.ivar = 2; % second row of cfg.design, the  
(i)ndependent events (3&15)  
stat = ft_freqstatistics(cfg, dataset_1, dataset_2);
```

Dependent variable is the frequency data

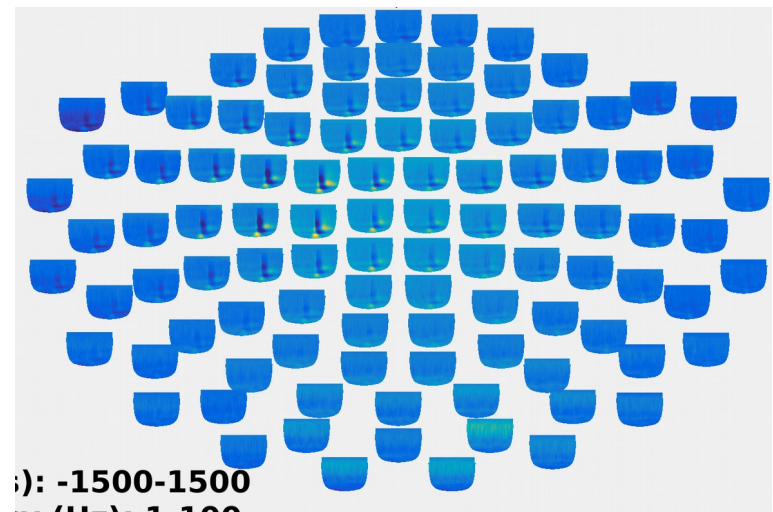
# Design matrix

Unit variable	Independent variable
1	1
2	1
...	...
19	1
20	1
1	2
2	2
...	...
19	2
20	2

Dependent variable is the frequency data



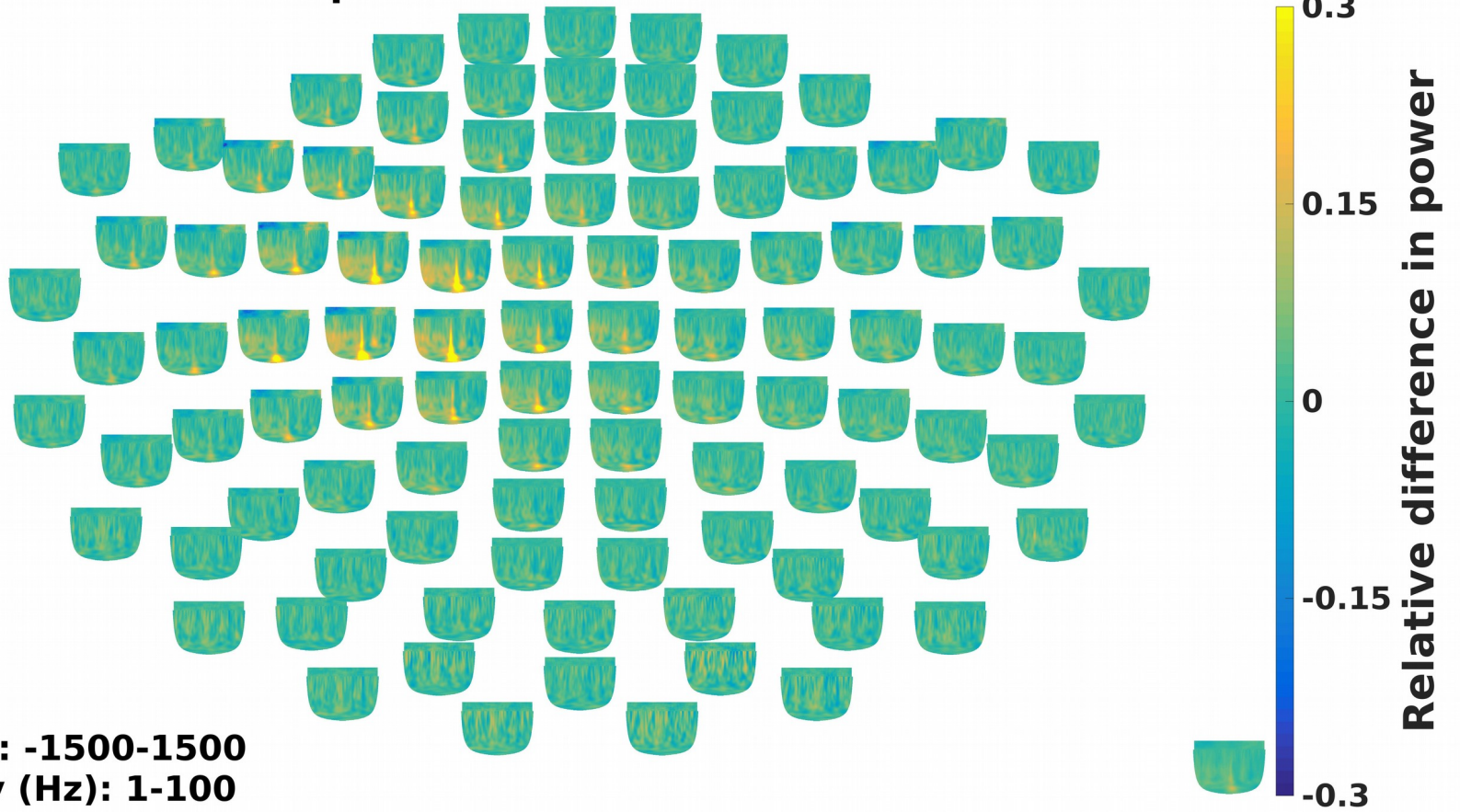
First Stimulation



Repeated Stimulation

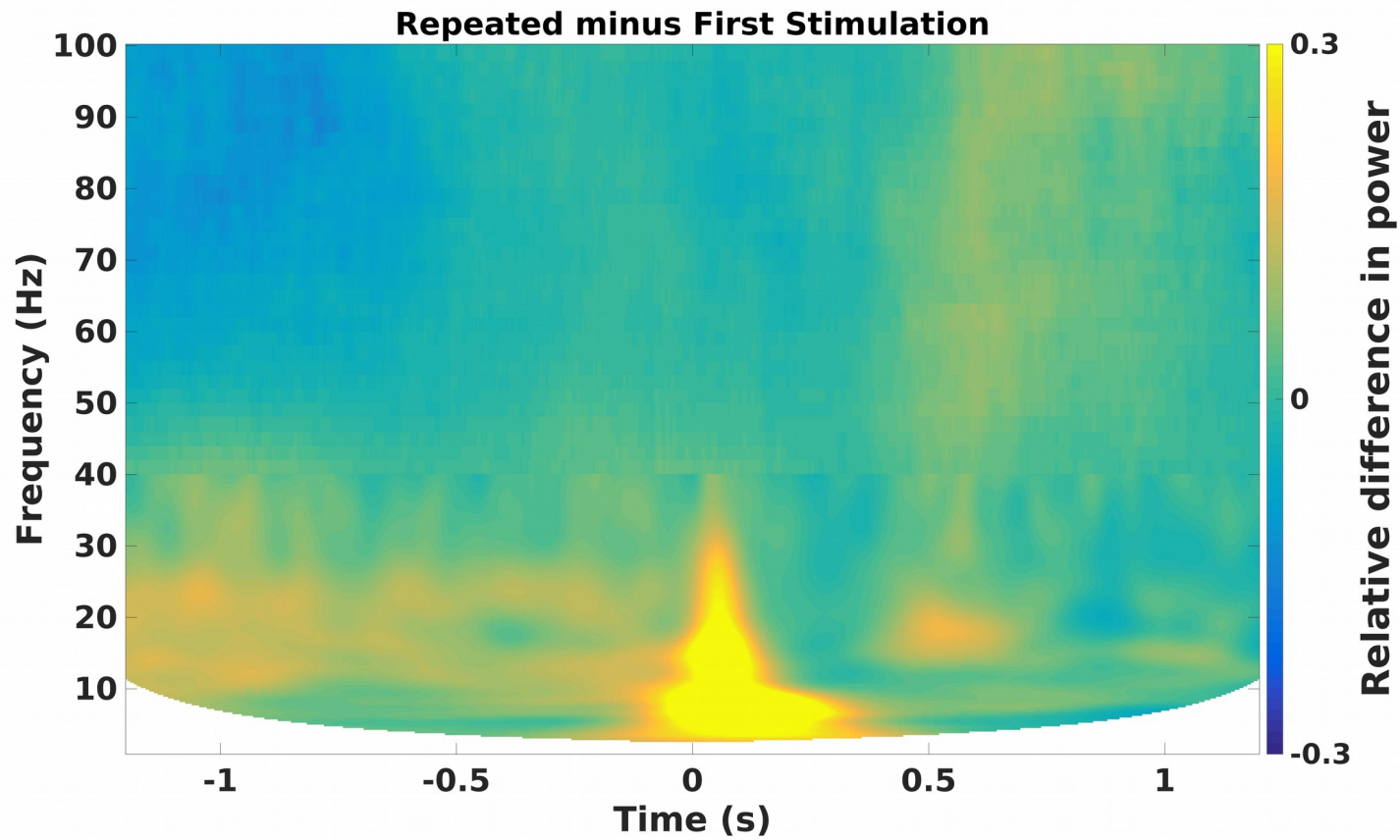
# Difference map

Repeated minus First Stimulation

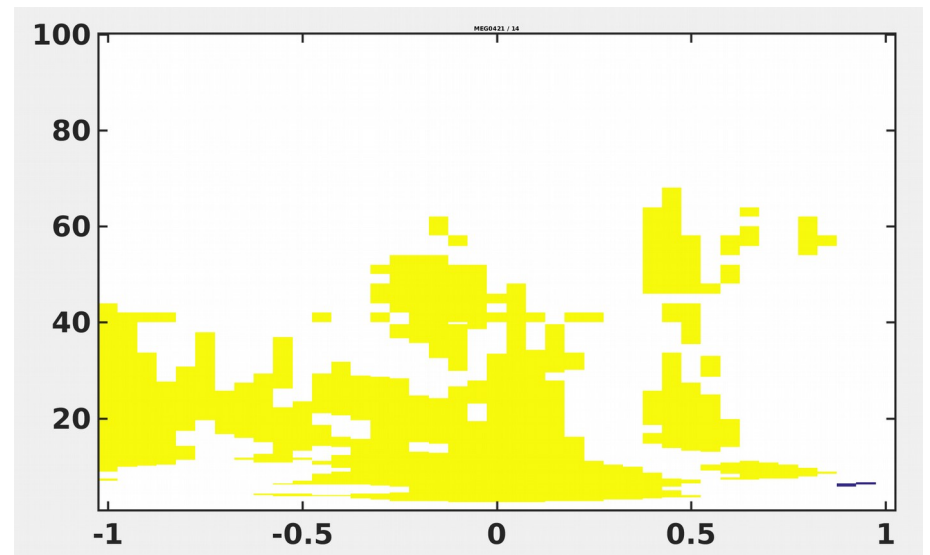
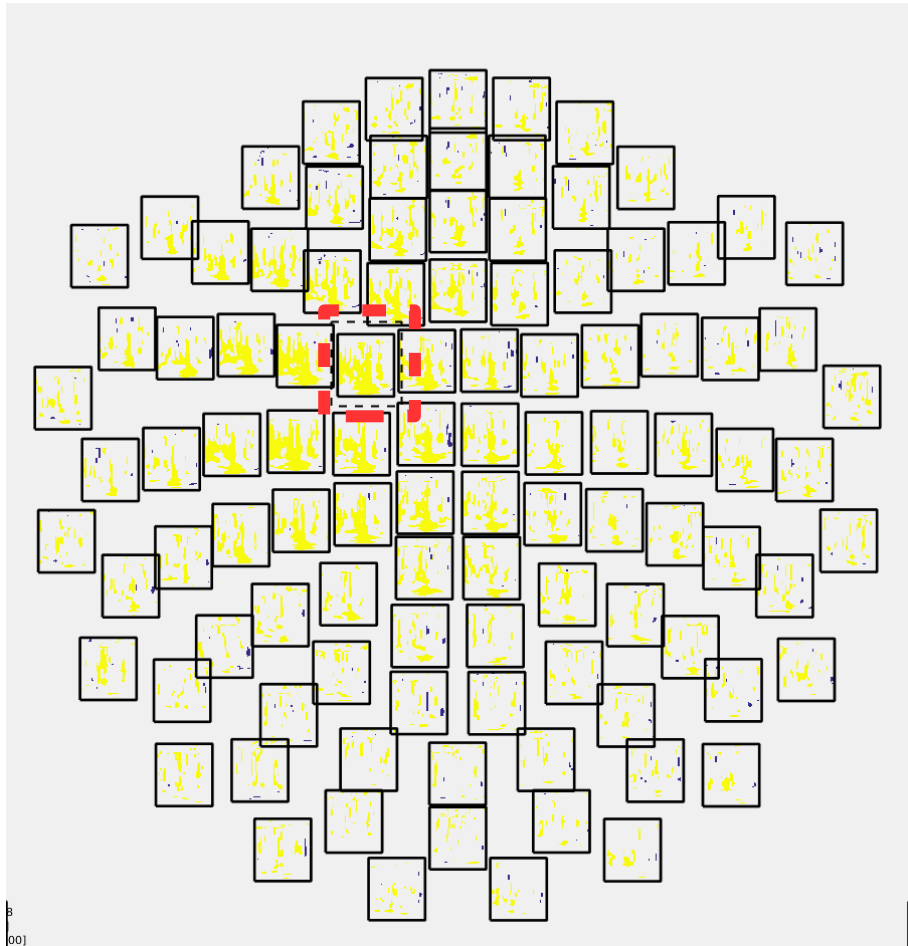


Time (ms): -1500-1500  
Frequency (Hz): 1-100

# Difference-map – single channel



# $t$ -value-map – masked



# Cluster analysis

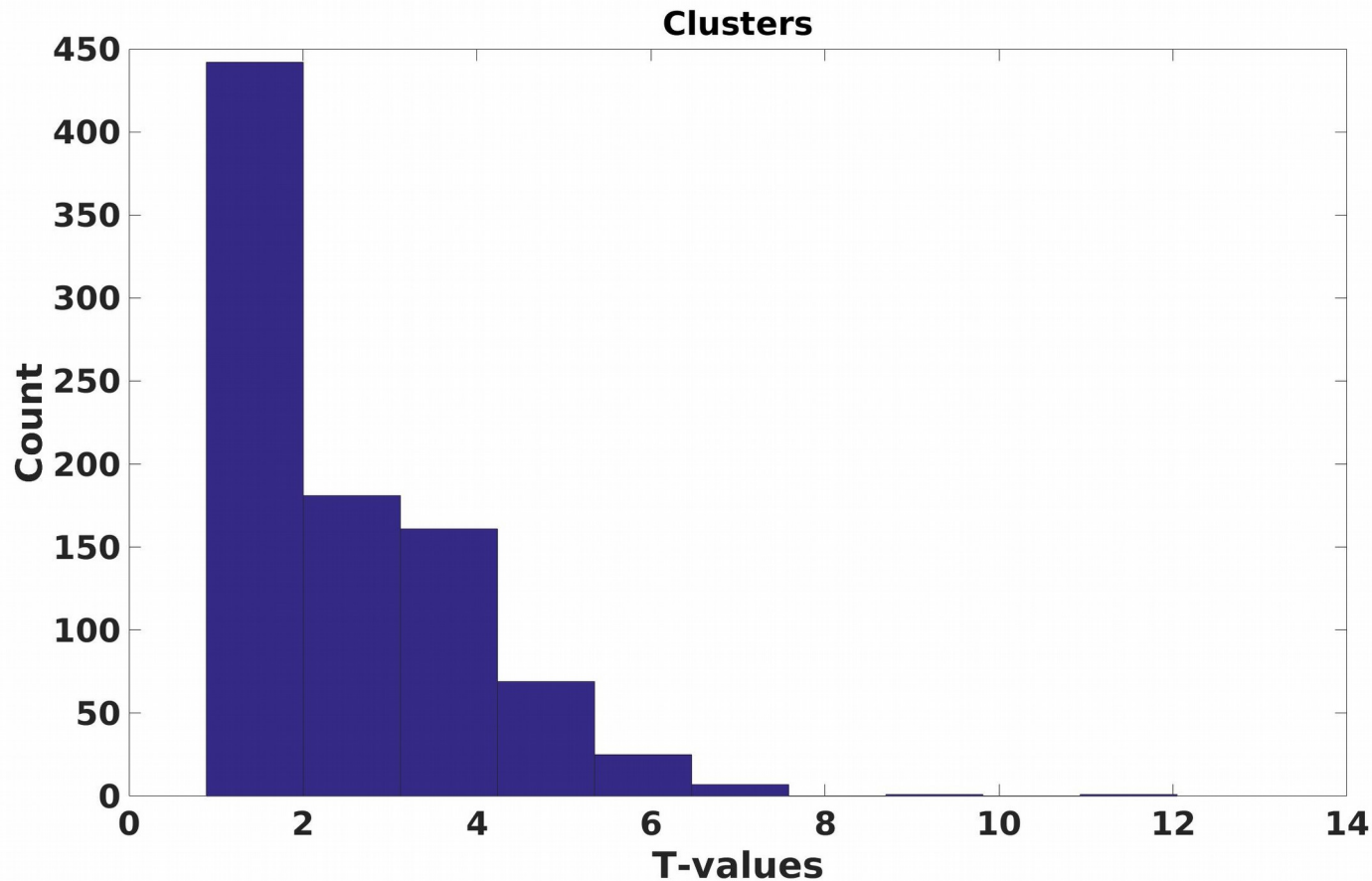
Unit variable	Independent variable
1	1
2	1
...	...
19	1
20	1
1	2
2	2
...	...
19	2
20	2

Dependent variable is the frequency data

```
cfg = [];  
cfg.latency = [1.500 1.500];  
cfg.frequency = [1 100];  
cfg.parameter = 'powspectrm';  
cfg.method = 'montecarlo'; %% do a permutation test  
cfg.alpha = 0.05; %% critical value around ± 2.09  
cfg.statistic = 'depsamplesT'; % use a dependent  
samples t-test  
cfg.design(1, :) = [1:n_subjects 1:n_subjects];  
cfg.design(2, :) = [ones(1, n_subjects) 2 * ones(1,  
n_subjects)];  
cfg.uvar = 1; % first row of cfg.design, containing  
the (u)nits (subjects)  
cfg.ivar = 2; % second row of cfg.design, the  
(i)ndependent events  
  
cfg.correctm = 'cluster'; % correct with cluster  
method  
cfg.clusteralpha = 0.025; % alpha for clusters  
cfg.clusterstatistic = 'maxsum'; % way to calculate  
T  
cfg.clustertail = 0;  
cfg.neighbours = neighbours; % a neighbour  
structure, see ft_prepare_neighbours  
cfg.numrandomization = 2000; % n randomizations  
  
stat = ft_freqstatistics(cfg, dataset_1, dataset_2);
```



# (Log)- $T$ -values



Now we have some  $T$ -values. But we do not have a distribution to compare it against, like the normal distribution or a  $t$ -distribution

# Cluster analysis – permutation 1

Unit variable	Independent variable
1	1
2	<u>2</u>
...	...
19	<u>2</u>
20	1
1	2
2	<u>1</u>
...	...
19	<u>1</u>
20	2

Dependent variable is the frequency data

```
cfg = [];  
cfg.latency = [1.500 1.500];  
cfg.frequency = [1 100];  
cfg.parameter = 'powspectrm';  
cfg.method = 'montecarlo'; %% do a permutation test  
cfg.alpha = 0.05; %% critical value around ± 2.09  
cfg.statistic = 'depsamplesT'; % use a dependent  
samples t-test  
cfg.design(1, :) = [1:n_subjects 1:n_subjects];  
cfg.design(2, :) = [ones(1, n_subjects) 2 * ones(1,  
n_subjects)];  
cfg.uvar = 1; % first row of cfg.design, containing  
the (u)nits (subjects)  
cfg.ivar = 2; % second row of cfg.design, the  
(i)ndependent events  
  
cfg.correctm = 'cluster'; % correct with cluster  
method  
cfg.clusteralpha = 0.025; % alpha for clusters  
cfg.clusterstatistic = 'maxsum'; % way to calculate  
T  
cfg.clustertail = 0;  
cfg.neighbours = neighbours; % a neighbour  
structure, see ft_prepare_neighbours  
cfg.numrandomization = 2000; % n randomizations  
  
stat = ft_freqstatistics(cfg, dataset_1, dataset_2);
```

# Cluster analysis – permutation 2

Unit variable	Independent variable
1	<u>2</u>
2	1
...	...
19	1
20	<u>2</u>
1	<u>1</u>
2	2
...	...
19	2
20	<u>1</u>

Dependent variable is the frequency data

```
cfg = [];  
cfg.latency = [1.500 1.500];  
cfg.frequency = [1 100];  
cfg.parameter = 'powspectrm';  
cfg.method = 'montecarlo'; %% do a permutation test  
cfg.alpha = 0.05; %% critical value around ± 2.09  
cfg.statistic = 'depsamplesT'; % use a dependent  
samples t-test  
cfg.design(1, :) = [1:n_subjects 1:n_subjects];  
cfg.design(2, :) = [ones(1, n_subjects) 2 * ones(1,  
n_subjects)];  
cfg.uvar = 1; % first row of cfg.design, containing  
the (u)nits (subjects)  
cfg.ivar = 2; % second row of cfg.design, the  
(i)ndependent events  
  
cfg.correctm = 'cluster'; % correct with cluster  
method  
cfg.clusteralpha = 0.025; % alpha for clusters  
cfg.clusterstatistic = 'maxsum'; % way to calculate  
T  
cfg.clustertail = 0;  
cfg.neighbours = neighbours; % a neighbour  
structure, see ft_prepare_neighbours  
cfg.numrandomization = 2000; % n randomizations  
  
stat = ft_freqstatistics(cfg, dataset_1, dataset_2);
```

# Cluster analysis – permutation ... 2000

Unit variable	Independent variable
1	1
2	<u>2</u>
...	...
19	1
20	1
1	2
2	<u>1</u>
...	...
19	2
20	2

```

cfg = [];
cfg.latency = [1.500 1.500];
cfg.frequency = [1 100];
cfg.parameter = 'powspectrm';
cfg.method = 'montecarlo'; %% do a permutation test
cfg.alpha = 0.05; %% critical value around ± 2.09
cfg.statistic = 'depsamplesT'; % use a dependent
samples t-test
cfg.design(1, :) = [1:n_subjects 1:n_subjects];
cfg.design(2, :) = [ones(1, n_subjects) 2 * ones(1,
n_subjects)];
cfg.uvar = 1; % first row of cfg.design, containing
the (u)nits (subjects)
cfg.ivar = 2; % second row of cfg.design, the
(i)ndependent events

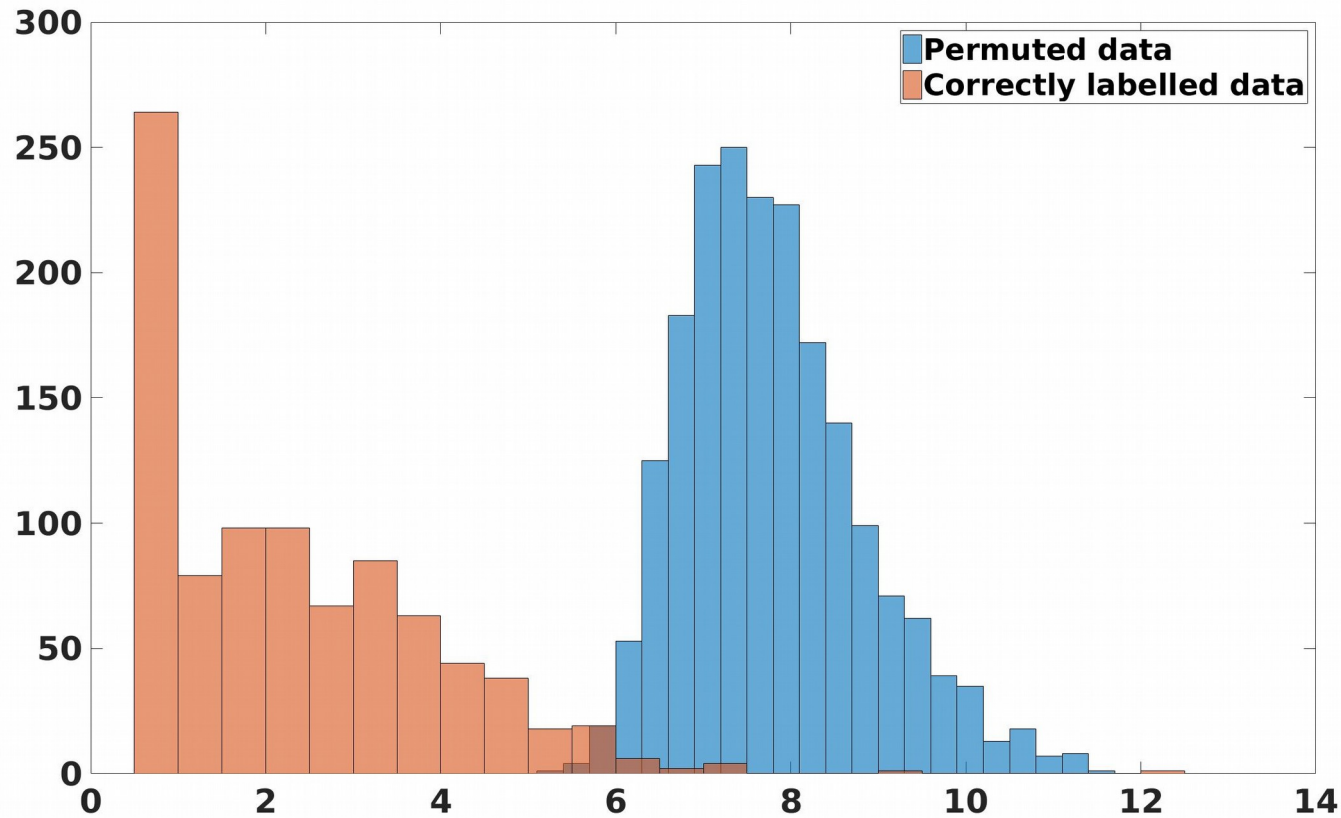
cfg.correctm = 'cluster'; % correct with cluster
method
cfg.clusteralpha = 0.025; % alpha for clusters
cfg.clusterstatistic = 'maxsum'; % way to calculate
T
cfg.clustertail = 0;
cfg.neighbours = neighbours; % a neighbour
structure, see ft_prepare_neighbours
cfg.numrandomization = 2000; % n randomizations

stat = ft_freqstatistics(cfg, dataset_1, dataset_2);

```

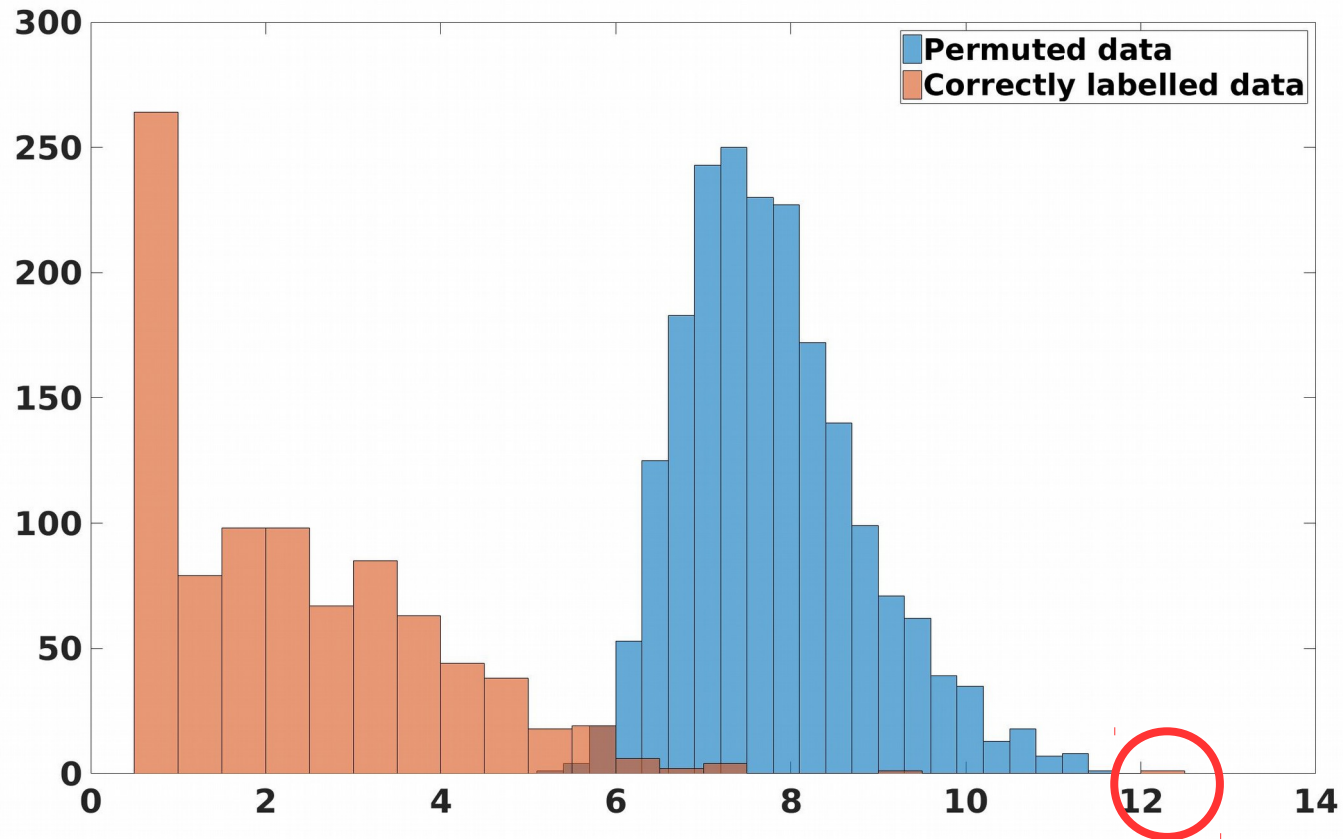
Dependent variable is the frequency data

# (Log)- $T$ -values



Let's run (many) permutations to sample the  $T$ -distribution

# (Log)- $T$ -values



$\log(T) = 12, p = 0.0005$

What is the null hypothesis when doing permutation based test?

# The null hypothesis

Unit variable	Independent variable
1	1
2	1
...	...
19	1
20	1
1	2
2	2
...	...
19	2
20	2

Null hypothesis: data is exchangeable → more informally: the way that you label your conditions do not matter

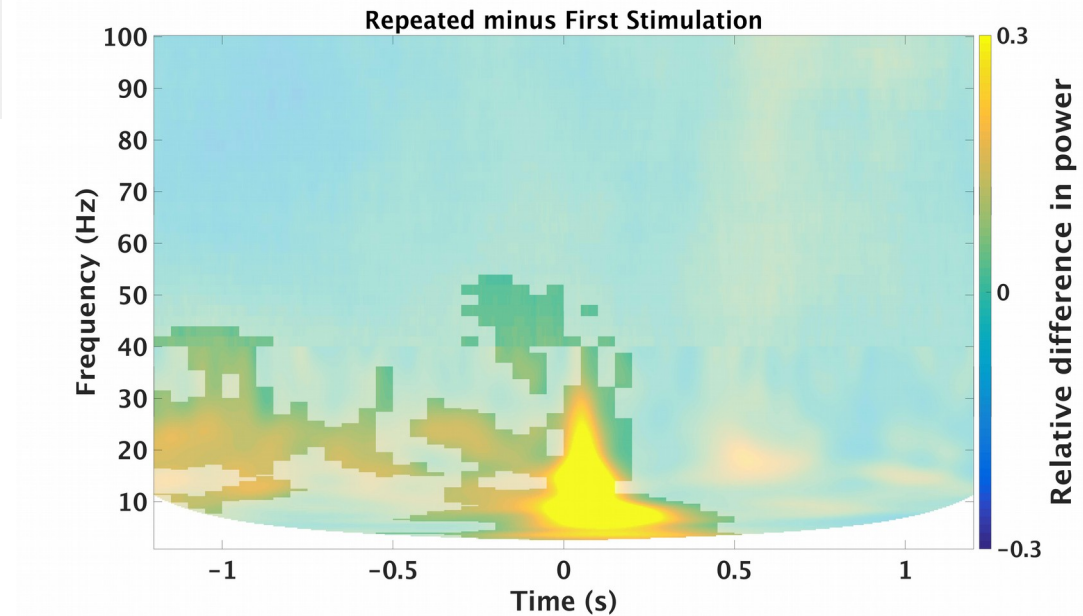
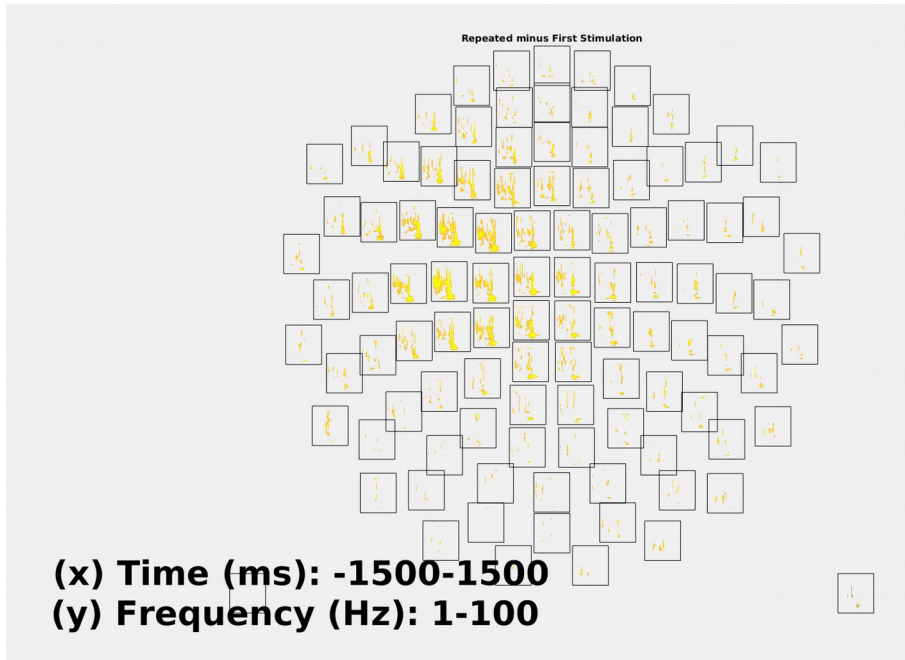
Alternative hypothesis informally: your way of labelling the conditions matter



This also means that it is nonsensical to speak of  
*statistically significant clusters*

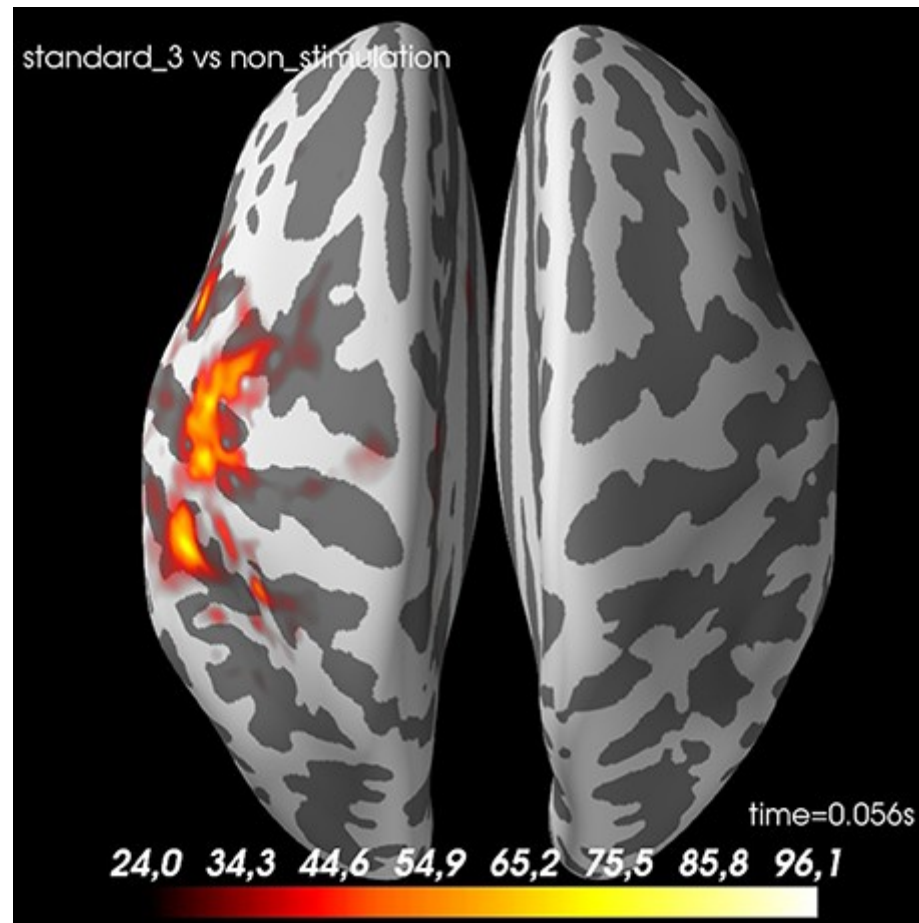
(It doesn't mean that clusters are not interesting to  
investigate further)

# Clusters



# Clusters

Clusters can also be used for source space



**Thank you!**