

LIMO EEG: what are robust statistics? Application to a categorical design

D2.A1 & D2.A3. – 2.00 to 3.30

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SCIENTIFIC DATA

OPEN

SUBJECT CATEGORIES

» Electroencephalography

-EEG

» Brain imaging

A multi-subject, multi-modal human neuroimaging dataset

Daniel G. Wakeman^{1,2} & Richard N. Henson²

- *Scientific Data* **2**, Article number: 150001 (2015)
- doi:10.1038/sdata.2015.1

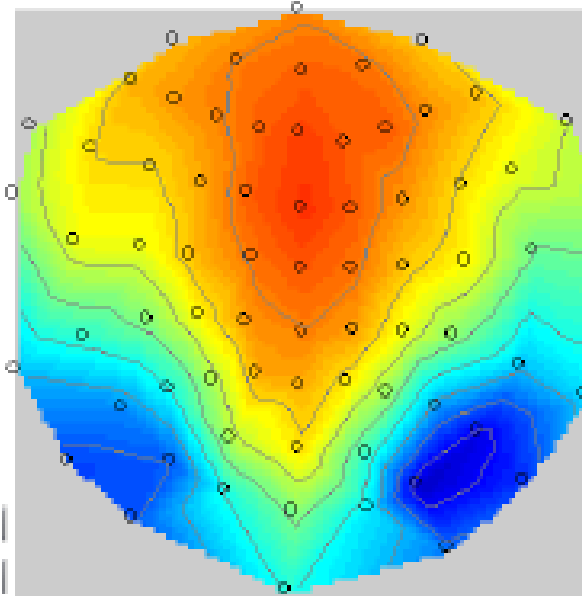
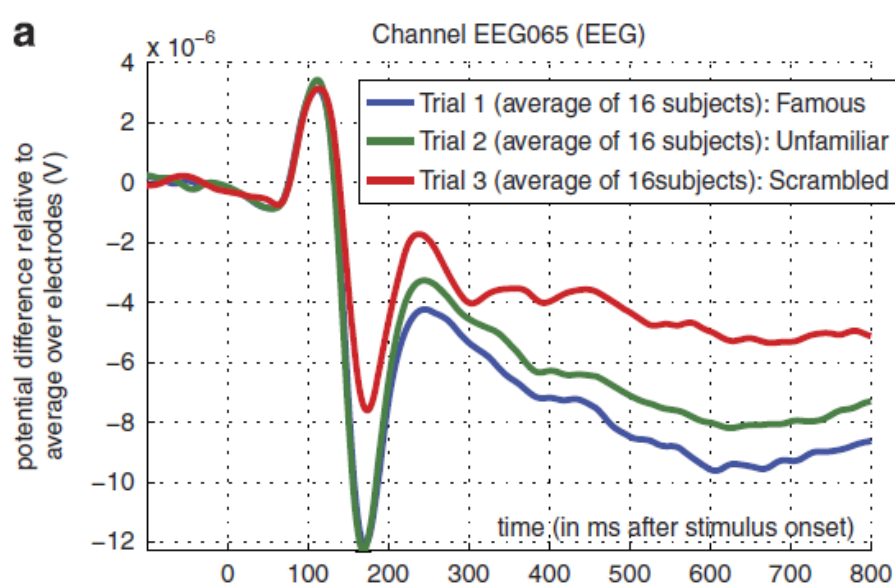
The Data

- 3 types of stimuli: Famous faces, Non-famous faces, Scrambled faces
 - 3 levels of repetition: 1st time, 2nd time (right after), 3rd time (delayed)
- Priming experiment with a possible interaction with the type of stimuli.

We need the conditions computed per subject (1st level) and then do the repeated measure ANOVA to test main effects and interactions.

What are we going to do?

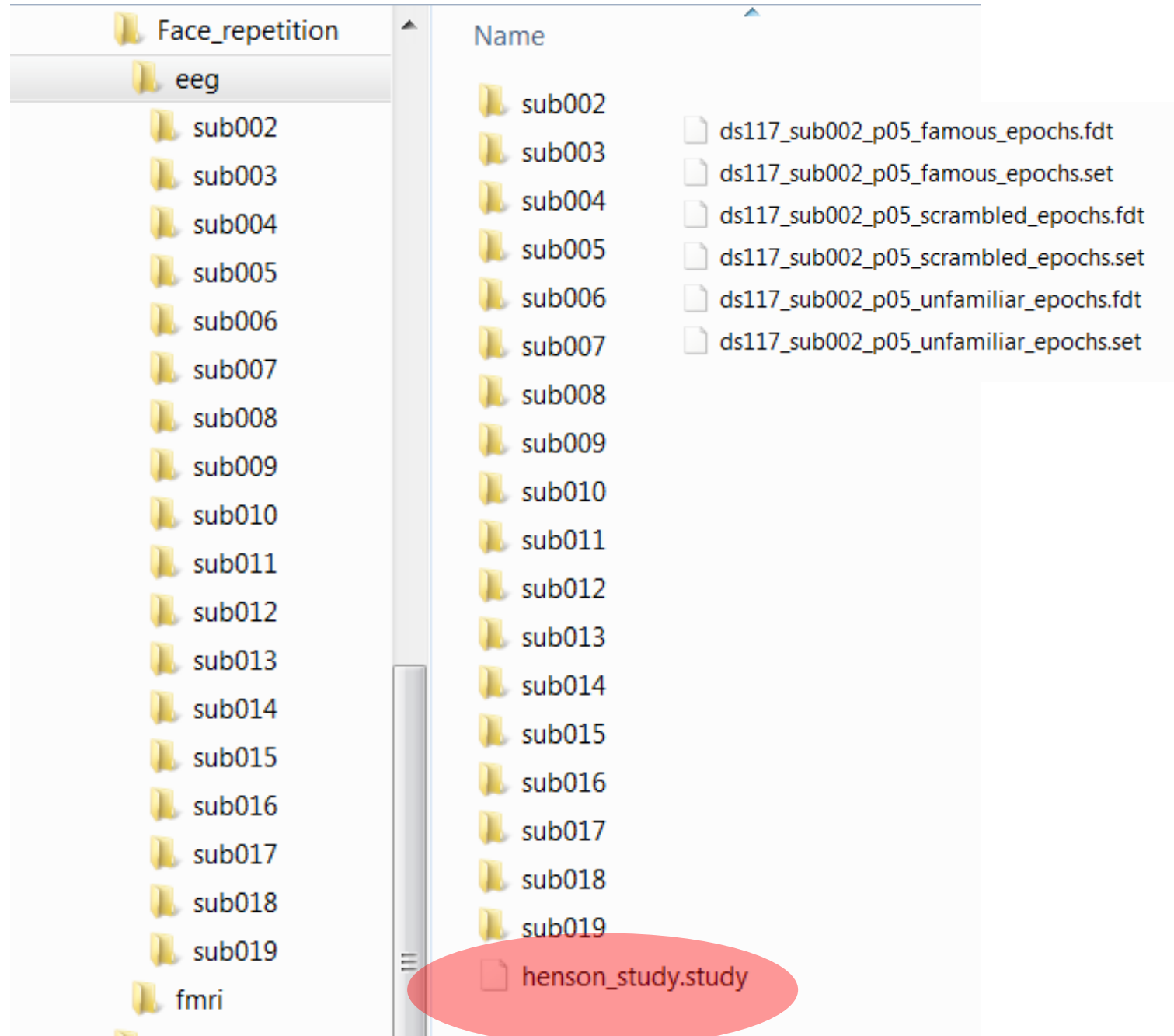
- 1 – Replicate Henson et al. – faces vs. scrambled



- 2 – learn about robust statistics
- 3 – see how to extend to a 3 (category) by 3 (repetition) design

Let's get started

- Open Matlab
- Move to the data
 - 18 subjects
- type 'eeglab'
- Load/Make the 'study'
 - file → load existing study



EEGLAB v14.x (dev)

File Edit Tools Plot Study Datasets Help

STUDY set: v

- Edit study info
- Select/Edit study design(s)
- Precompute channel measures
- Plot channel measures
- Linear Modeling EEG Data (LIMO/Channels) ▶
- Precompute component measures
- PCA clustering (original) ▶
- Edit/plot clusters
- Linear Modeling EEG Data (LIMO/Components) ▶

Study filename

Study task name

Nb of subjects

Nb of conditions

Nb of sessions

Nb of groups

1 per subject

Epoch consistency

yes

Channels per frame

60, 65, 66, 67, 68, 69, 70

Channel locations

yes

Clusters

21

Status

Pre-clustered

Total size (Mb)

228.1

```
warning: STUDY moved since last saved, trying to load...  
pop_loadset(): loading file E:\My_Documents\...  
Warning: STUDY moved since last saved, trying to load...  
pop_loadset(): loading file E:\My_Documents\...  
Warning: STUDY moved since last saved, trying to load...  
pop_loadset(): loading file E:\My_Documents\...  
Warning: STUDY moved since last saved, trying to load...  
pop_loadset(): loading file E:\My_Documents\...  
STUDY Warning: the trial information collection is incomplete.  
Done.  
Re-saving study file  
Done.
```

fx >>

Create a new STUDY set -- pop_study()

Edit STUDY set information - remember to save changes

STUDY set name:

wakeman_henson_study

STUDY set task name:

ScrambledVsNormalFace

STUDY set notes:

dataset filename

browse

subject

session

condition

group

Select by r.v.

1	E:\My_Documents\MATLAB\F..._rep001\pop_loadset\sub002\de	...	sub002	1	familiar	1	Comp.: 1 2 ...	Clear
2	E:\My_Documents\MATLAB\F..._rep002\pop_loadset\sub002\de	...	sub002	1	unfamiliar	1	Comp.: 1 2 ...	Clear
3	E:\My_Documents\MATLAB\F..._rep003\pop_loadset\sub002\de	...	sub002	1	scrambled	1	Comp.: 1 2 ...	Clear
4	E:\My_Documents\MATLAB\F..._rep004\pop_loadset\sub003\de	...	sub003	1	familiar	1	Comp.: 1 2 ...	Clear
5	E:\My_Documents\MATLAB\F..._rep005\pop_loadset\sub003\de	...	sub003	1	unfamiliar	1	Comp.: 1 2 ...	Clear
6	E:\My_Documents\MATLAB\F..._rep006\pop_loadset\sub003\de	...	sub003	1	scrambled	1	Comp.: 1 2 ...	Clear
7	E:\My_Documents\MATLAB\F..._rep007\pop_loadset\sub004\de	...	sub004	1	familiar	1	Comp.: 1 2 ...	Clear
8	E:\My_Documents\MATLAB\F..._rep008\pop_loadset\sub004\de	...	sub004	1	unfamiliar	1	Comp.: 1 2 ...	Clear
9	E:\My_Documents\MATLAB\F..._rep009\pop_loadset\sub004\de	...	sub004	1	scrambled	1	Comp.: 1 2 ...	Clear
10	E:\My_Documents\MATLAB\F..._rep010\pop_loadset\sub005\de	...	sub005	1	familiar	1	Comp.: 1 2 ...	Clear

Important note: Removed datasets will not be saved before being deleted from EEGLAB memory

<

Page 1

>

- ☐ Dataset info (condition, group, ...) differs from study info. [set] = Overwrite dataset info for each dataset on disk.
- ☐ Delete cluster information (to allow loading new datasets, set new components for clustering, etc.)

Help

Cancel

Ok

Date Modified

15/10/2016 23:23

15/10/2016 20:00

15/10/2016 17:56

15/10/2016 18:00

15/10/2016 18:05

15/10/2016 18:09

15/10/2016 18:14

15/10/2016 18:19

15/10/2016 18:23

15/10/2016 20:00

15/10/2016 19:58

15/10/2016 20:03

15/10/2016 20:07

15/10/2016 20:17

15/10/2016 20:21

15/10/2016 20:26

15/10/2016 20:30

15/10/2016 20:35

17/10/2016 21:23

MATLAB R2013a

EEGLAB v14.x (dev)

File Edit Tools Plot **Study** Datasets Help

STUDY set:

- Edit study info
- Select/Edit study design(s)
- Precompute channel measures**
- Plot channel measures
- Linear MOdeling EEG Data (LIMO/Channels) ▸
- Precompute component measures
- PCA clustering (original) ▸
- Edit/plot clusters
- Linear MOdeling EEG Data (LIMO/Components) ▸

Study filename: ...
 Study task: ...
 Nb of subjects: ...
 Nb of conditions: ...
 Nb of sessions: ...
 Nb of groups: 1 per subject
 Epoch consistency: yes
 Channels per frame: 60, 65, 66, 67, 68, 69, 70
 Channel locations: yes
 Clusters: 1
 Status: Pre-clustered
 Total size (Mb): 228

pop_loadset(): loading file E:\My_Documents\
 pop_loadset(): loading file E:\My_Documents\
 pop_loadset(): loading file E:\My_Documents\
 pop_loadset(): loading file E:\My_Documents\
 STUDY Warning: the trial information collected from datasets has changed; use STUDY menu to rec
 Done.

Select and compute component measures for later clustering -- pop_precomp()

Pre-compute channel measures for STUDY 'wakeman_henson_study'

- ☒ Spherical interpolation of missing channels (performed after optional ICA removal below)
- ☐ Remove ICA artifactual components pre-tagged in each dataset
- ☐ Remove artifactual ICA cluster or clusters (hold shift key) Parentcluster 1

List of measures to precompute

- ☒ ERPs Baseline ([min max] in ms)
- ☐ Power spectrum Spectopo parameters 'specmode', 'fft', 'logtrials', 'off' Test
- ☐ ERP-image ERP-image parameters 'nlines', 10, 'smoothing', 10 Test
- ☐ ERSPs Time/freq. parameters 'cycles', [3 0.8], 'nfreqs', 100, 'ntin' Test
- ☐ ITCs

☒ Overwrite files on disk

Help Cancel Ok

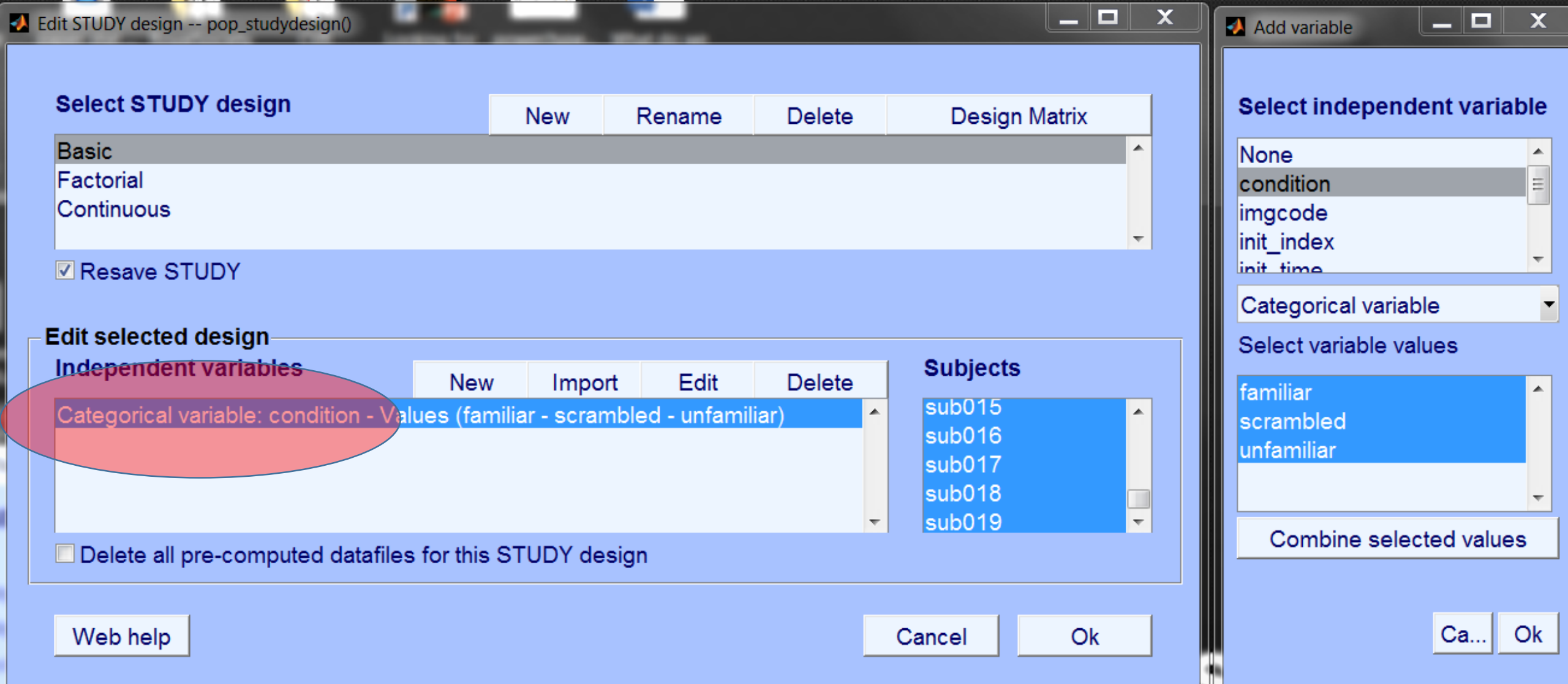
Value
<1x4 cell>
<1x54 struct>
10
16
8
[0.9300,0.9600,1]
10.6600,0.7600,11

Current Folder	Date Modified
ous_epochs.fdt	14/10/2016 23:45
ous_epochs.set	14/10/2016 23:45
mbled_epochs.fdt	14/10/2016 23:45
mbled_epochs.set	14/10/2016 23:45
amiliar_epochs.fdt	14/10/2016 23:45
amiliar_epochs.set	14/10/2016 23:45

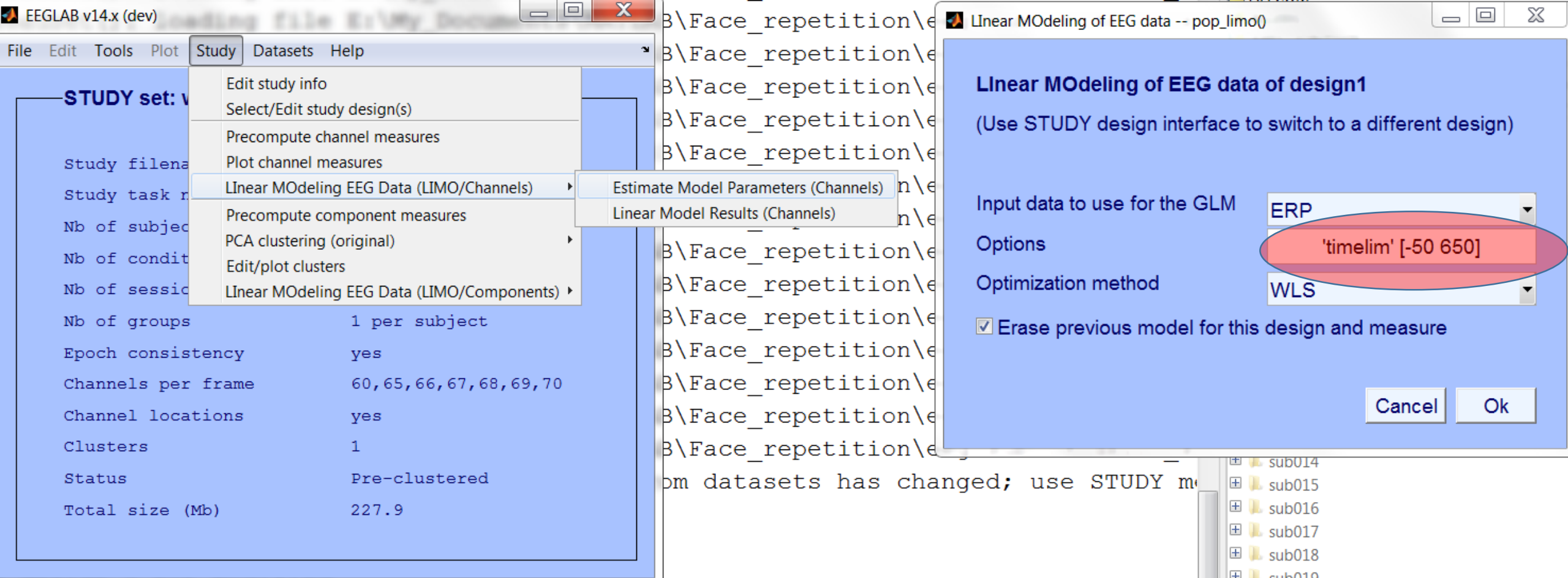
We now extract all the single trials

-- no interpolation of missing channels ; LIMO EEG handles missing data

-- data were processed with epochs -200/800



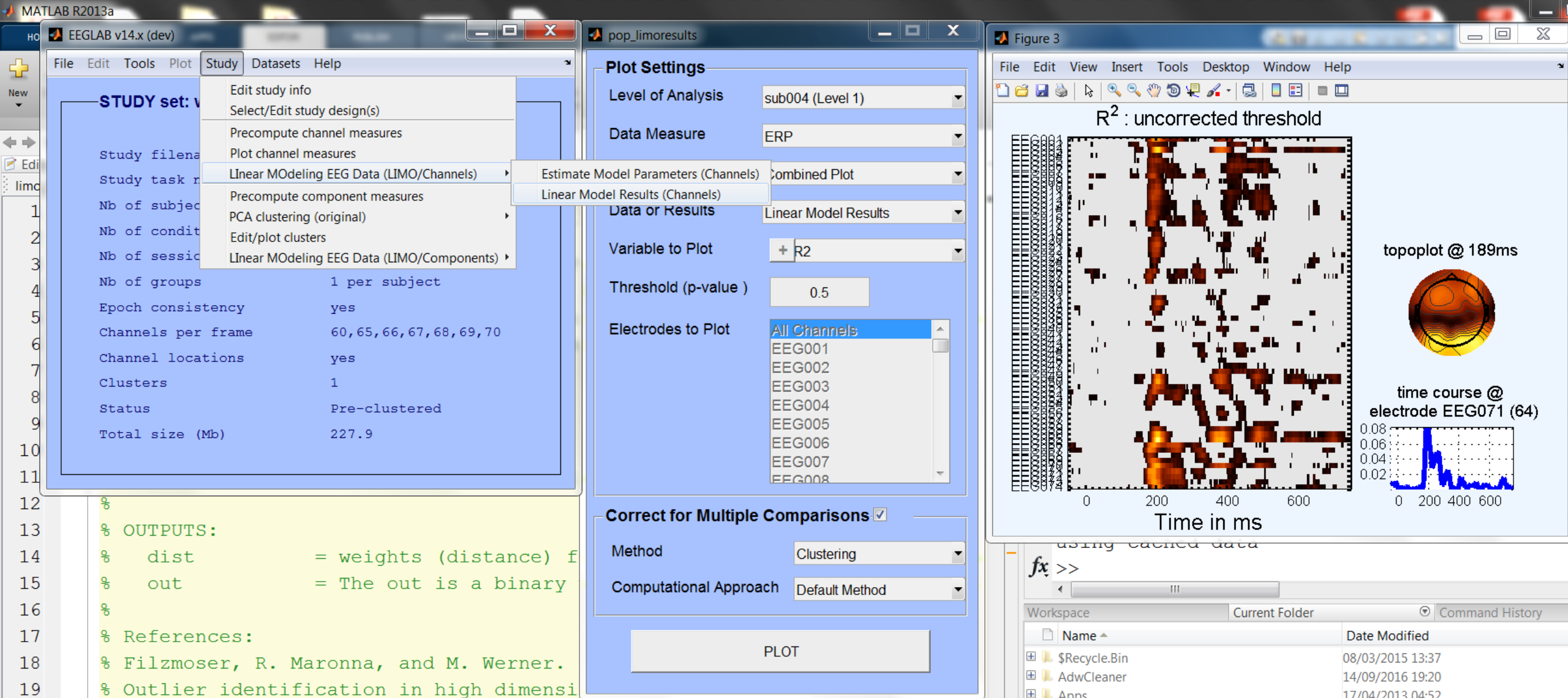
The new STUDY allows all sorts of designs
By default, you can model each and any condition / covariate
Here, we pick the categorical variable 'condition'



- 1 – we have created a study
- 2 – we have generated single trials for this study
- 3 – we made a design = statistical model
- 4 – now we have to do the stats = estimate the model parameters (b)
→ Restrict 'timelim' [-50 650] and use Weighted Least Square

Whilst it's computing, let's recap

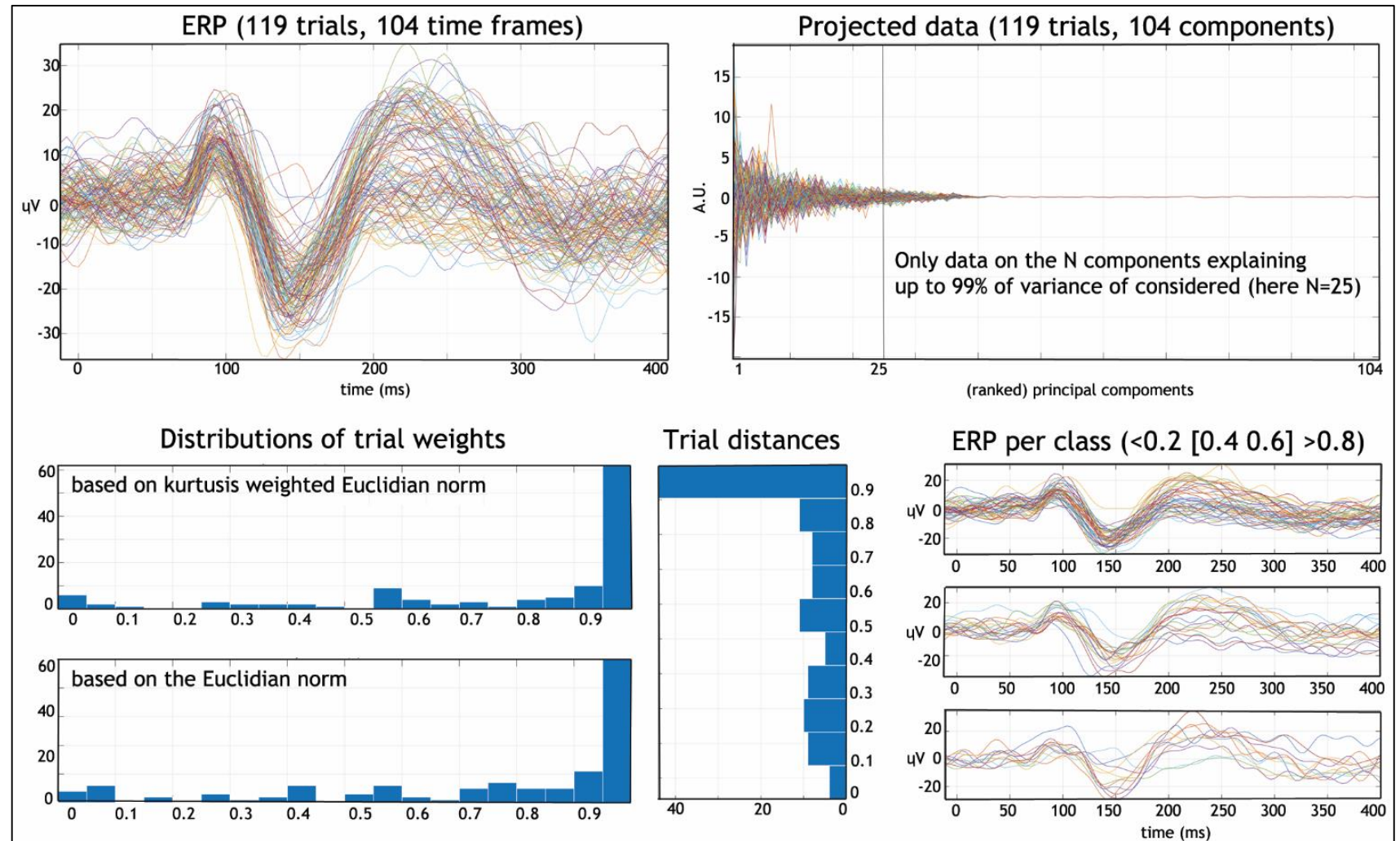
- For each subject, at each electrode we have a statistical model: $Y = XB + e$
- The Weighted Least Square solution is $B = \text{pinv}(WX) * WY$ for which the matrix W minimizes the influence of outliers in an otherwise (assumed) normally distributed data-set.
- Outliers are defined in a multivariate space, for instance a trial with a time course 'different' from others
- For each subject we have Y_r (data), LIMMO (model, weights, other info), $Betas$ (parameters), \hat{Y} ($LIMMO.design.X * Betas = \hat{Y}$), Res (residuals), R^2 (model fit), Condition_effect1 (=ANOVA F test across all conditions).



we can now easily look at each subject
the group level one-sample t-test is also always computed (not always meaningful)

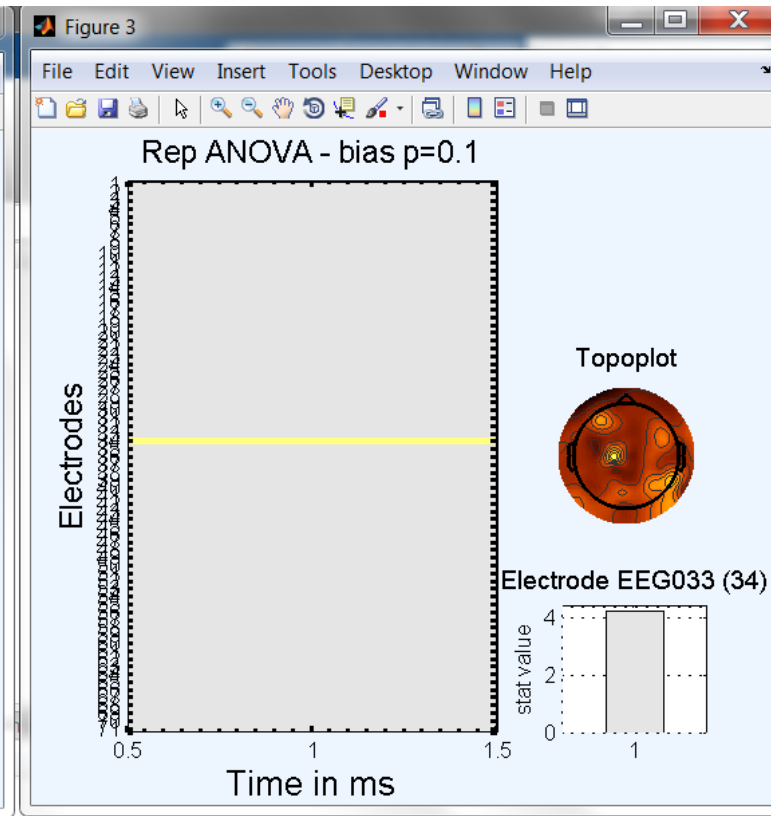
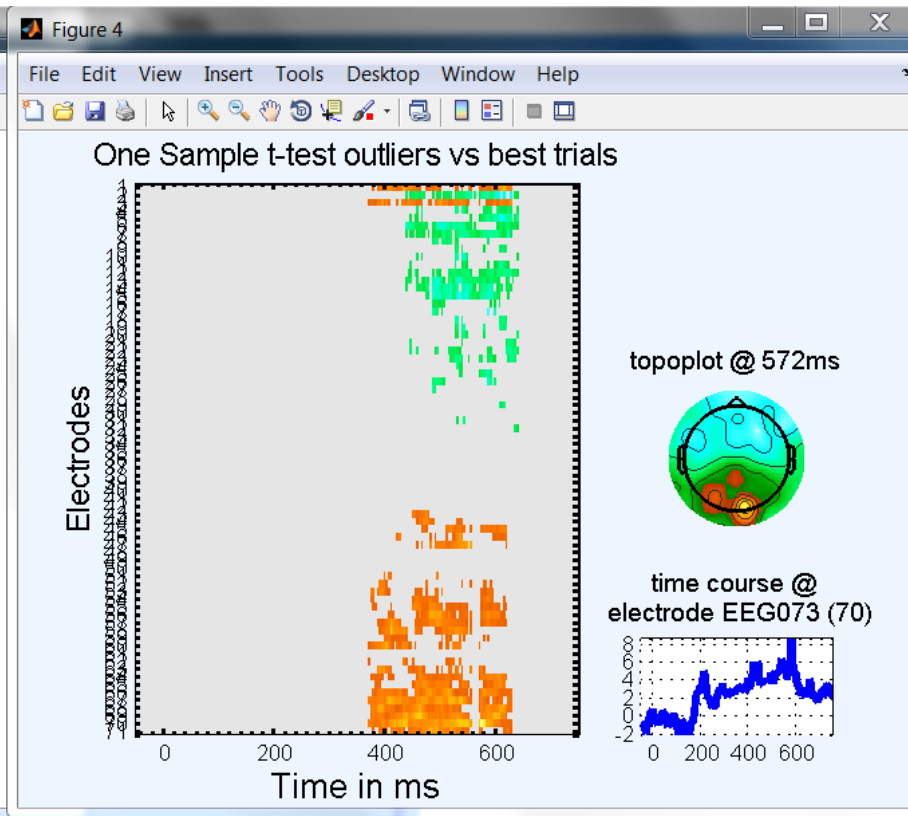
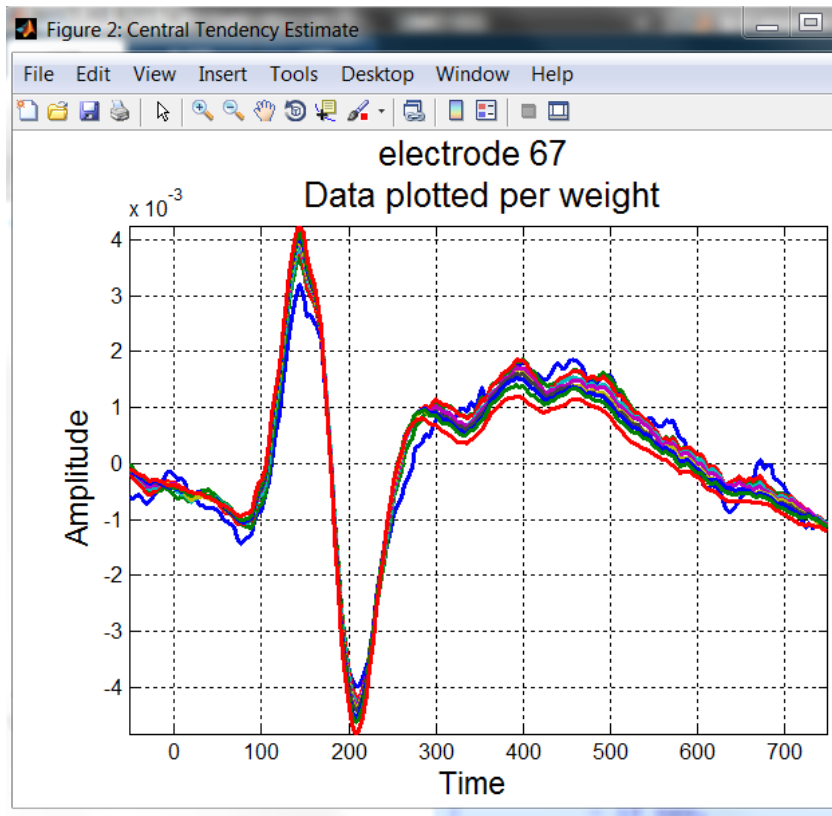
Weighted least squares in LIMO EEG

- Principal Component Projection method:
- PCA
- outlier detection on projected data points (Filzmoser et al., 2008)
- 1 weight per trial



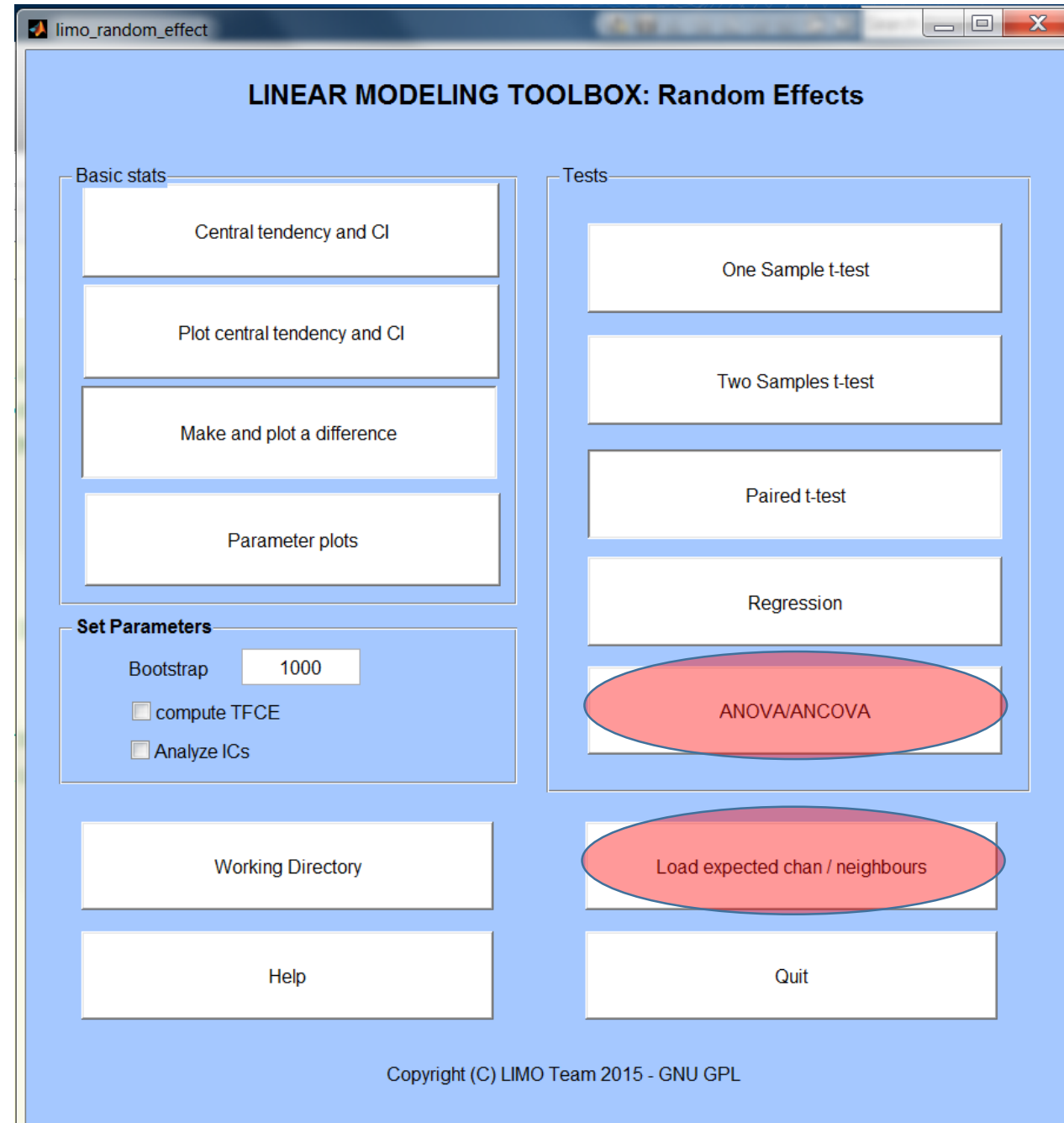
Let's check the weights

- In LIMO Tools, select 'Check weights'



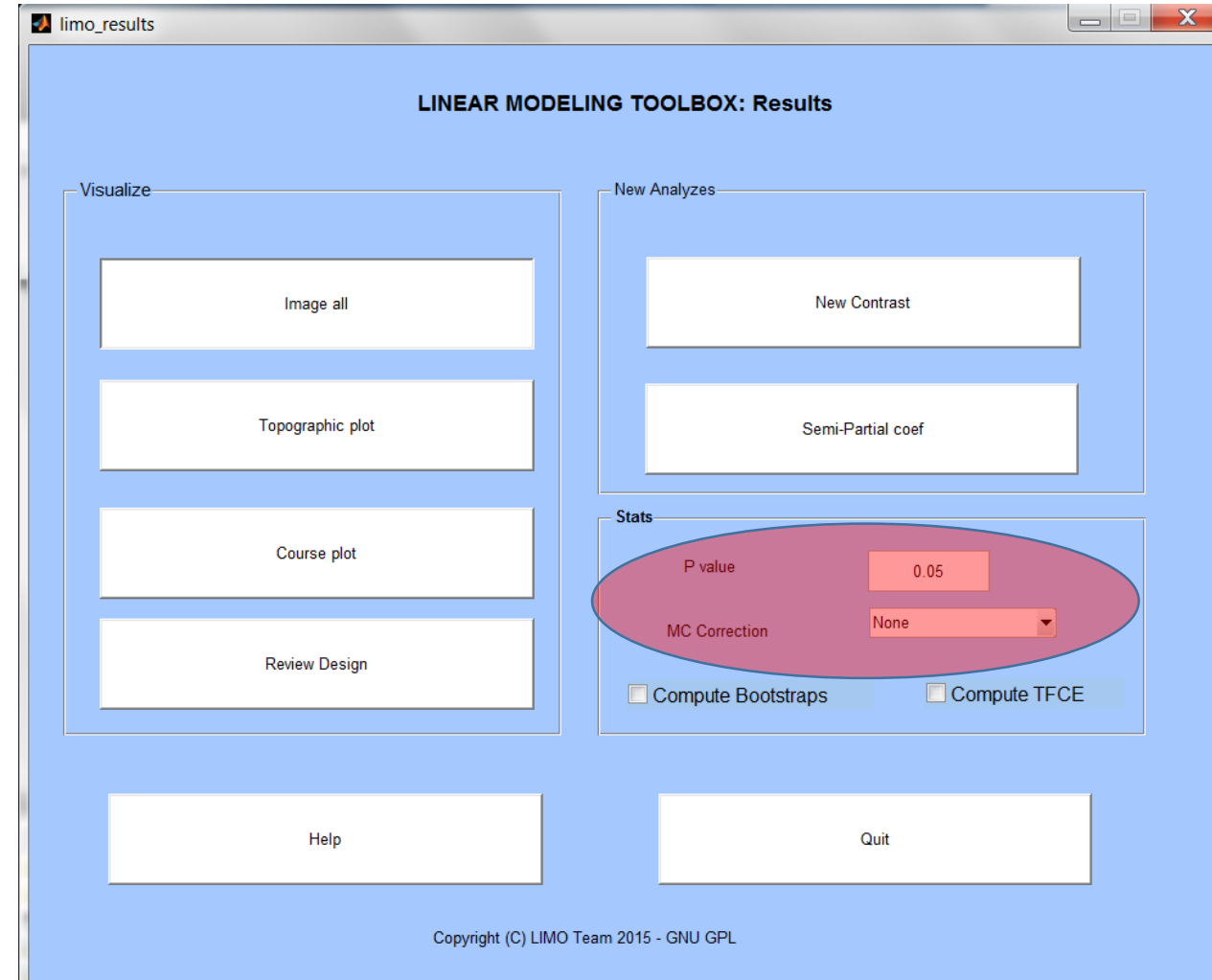
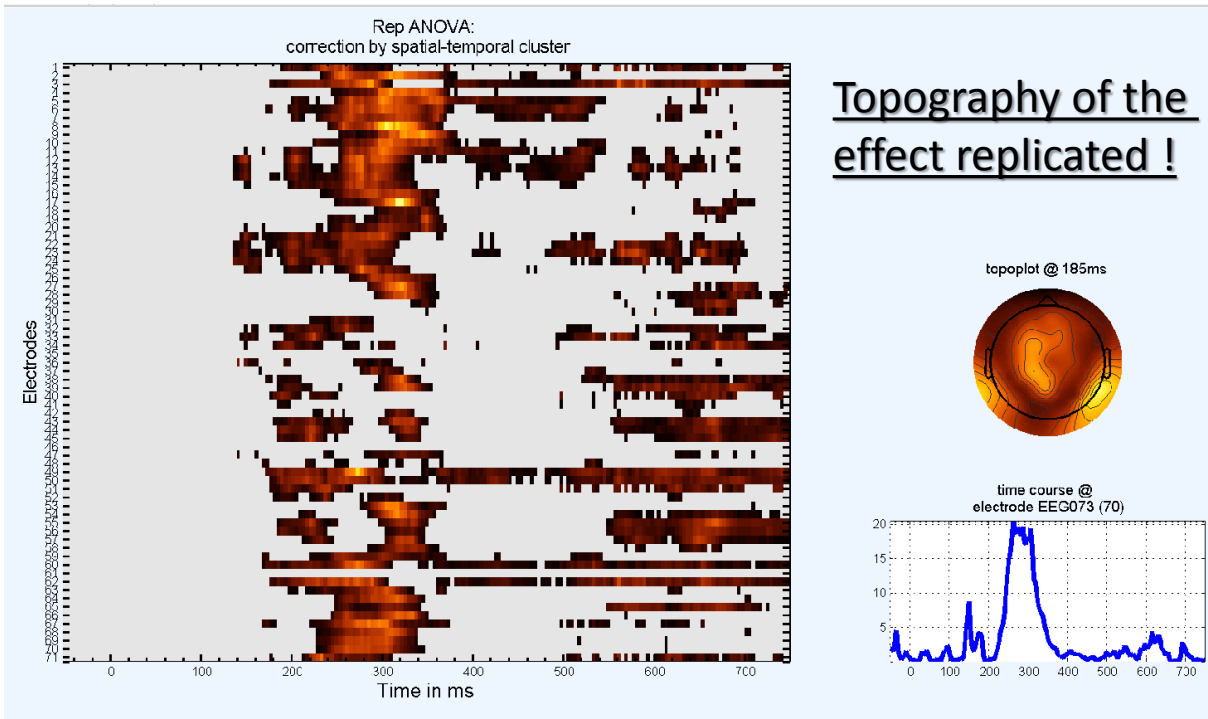
Group level analysis

- Just the same as with ERP but (i) we use betas (ii) LIMO EEG uses robust statistics (essentially 20% trimmed means – except *for now* the repeated measures ANOVA)
- Call the LIMO GUI, and select random effects
- Need a way to group subject with different channels (no interpolation) – the study created such file for you and it should be loaded by default – if not load:
→ *limo_gp_level_chanlocs.mat*
- Create a folder for the results and do a repeated measure ANOVA selecting the beta files [1 2 3]



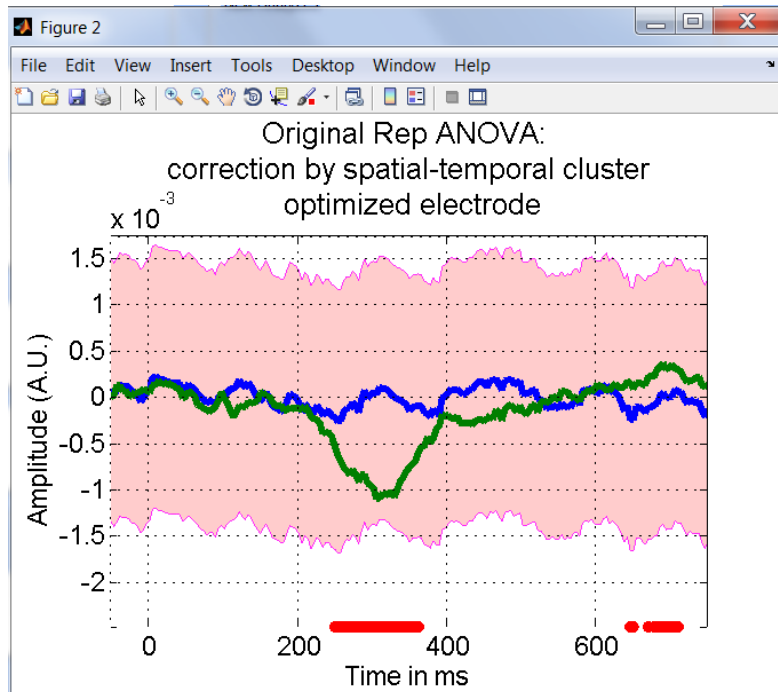
Group level analysis

- Call LIMO Results to look at the ANOVA results
- Note the choice between uncorrected and corrected p-values



Group level analysis

- If you check the 'course plot' all we have is the contrast used to get the F values
- Best check the 'real' data – plotting the betas and ERP → back to Random Effect



limo_random_effect

LINEAR MODELING TOOLBOX: Random Effects

Basic stats

Central tendency and CI

Plot central tendency and CI

Make and plot a difference

Parameter plots

Tests

One Sample t-test

Two Sample t-test

Paired t-test

Regression

ANOVA/ANCOVA

Set Parameters

Bootstrap 1000

☐ compute TFCE

☐ Analyze ICs

Working Directory

Help

Load expected chan / neighbours

Quit

what dat... type of analysis

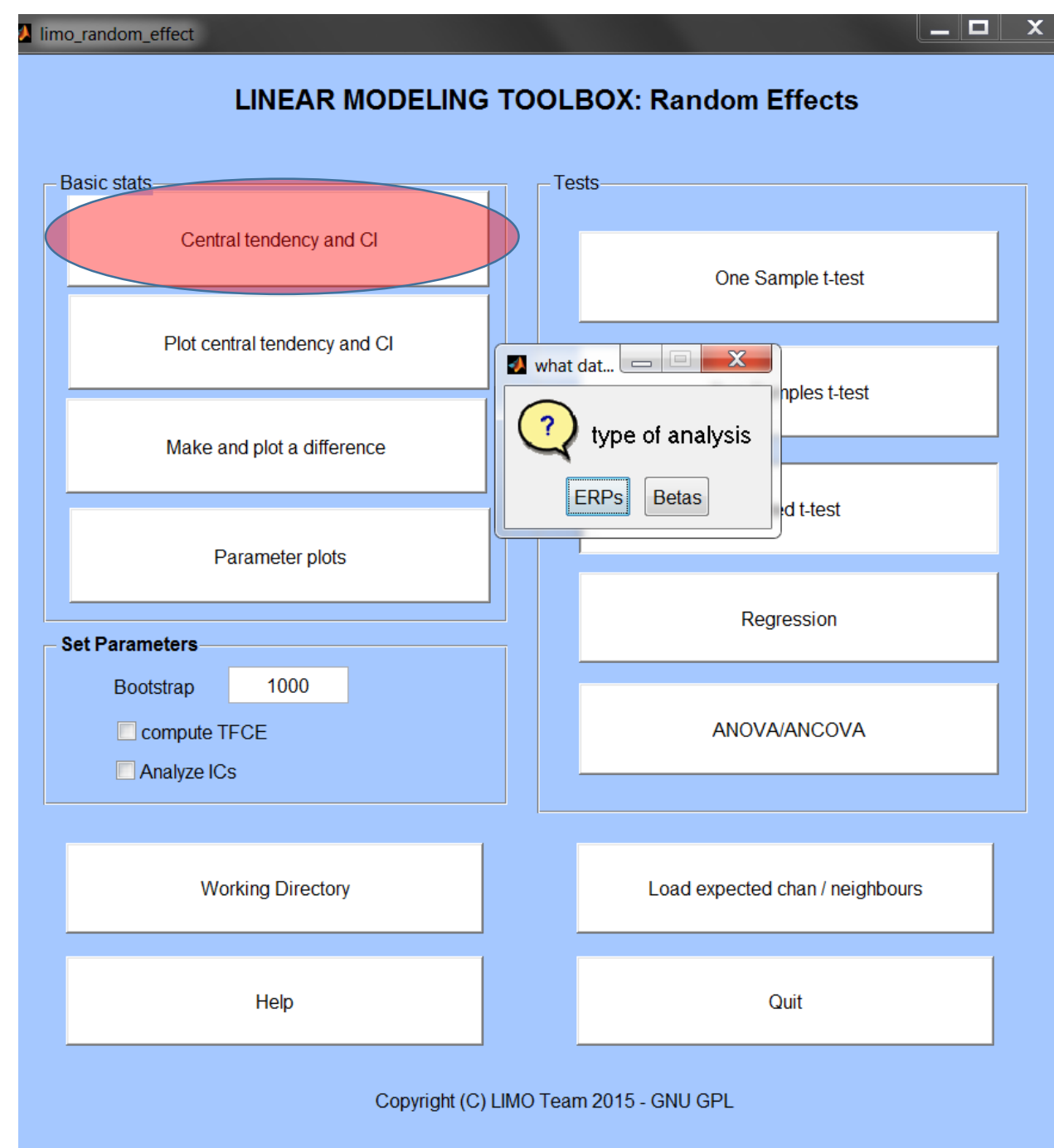
ERPs

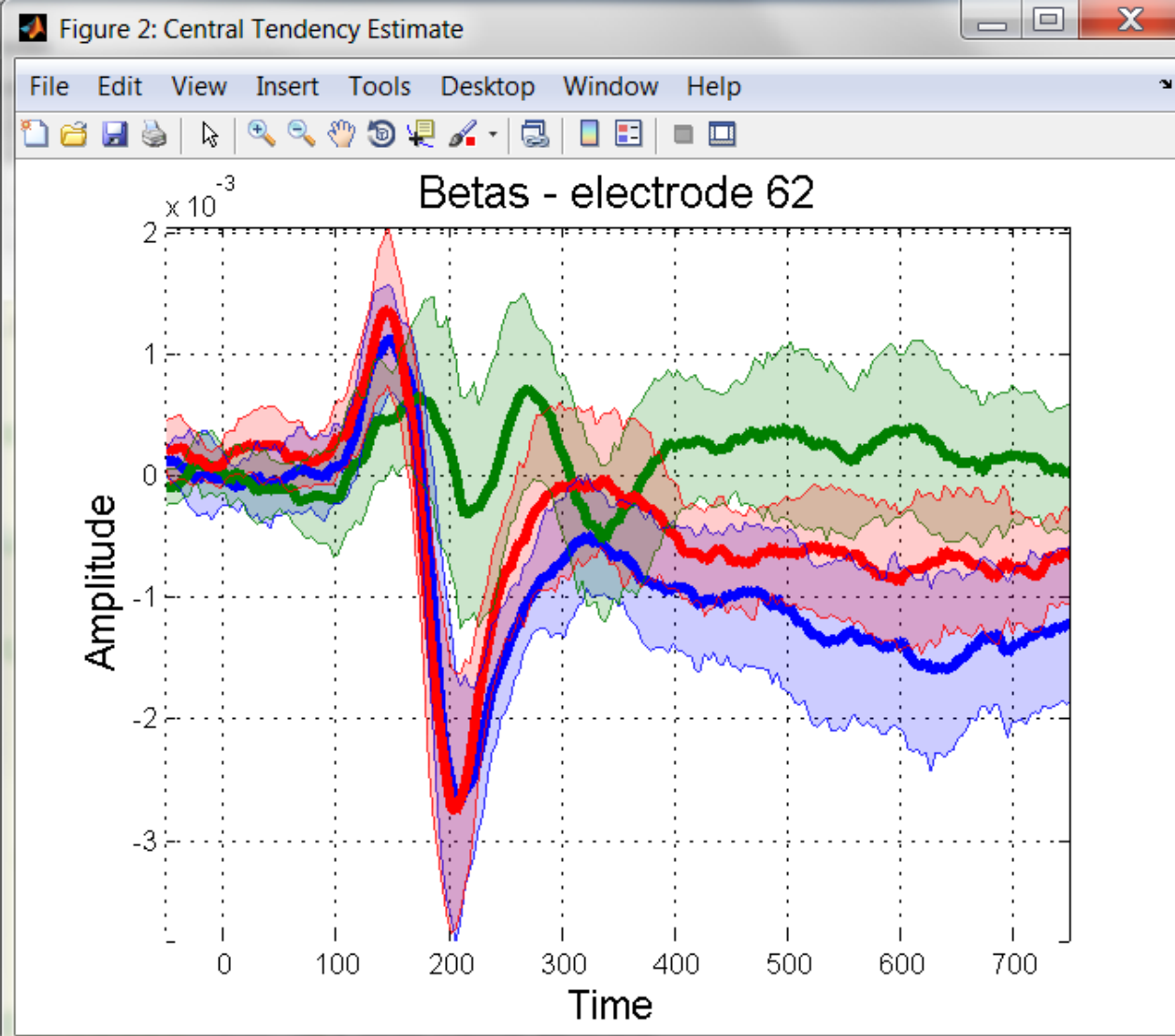
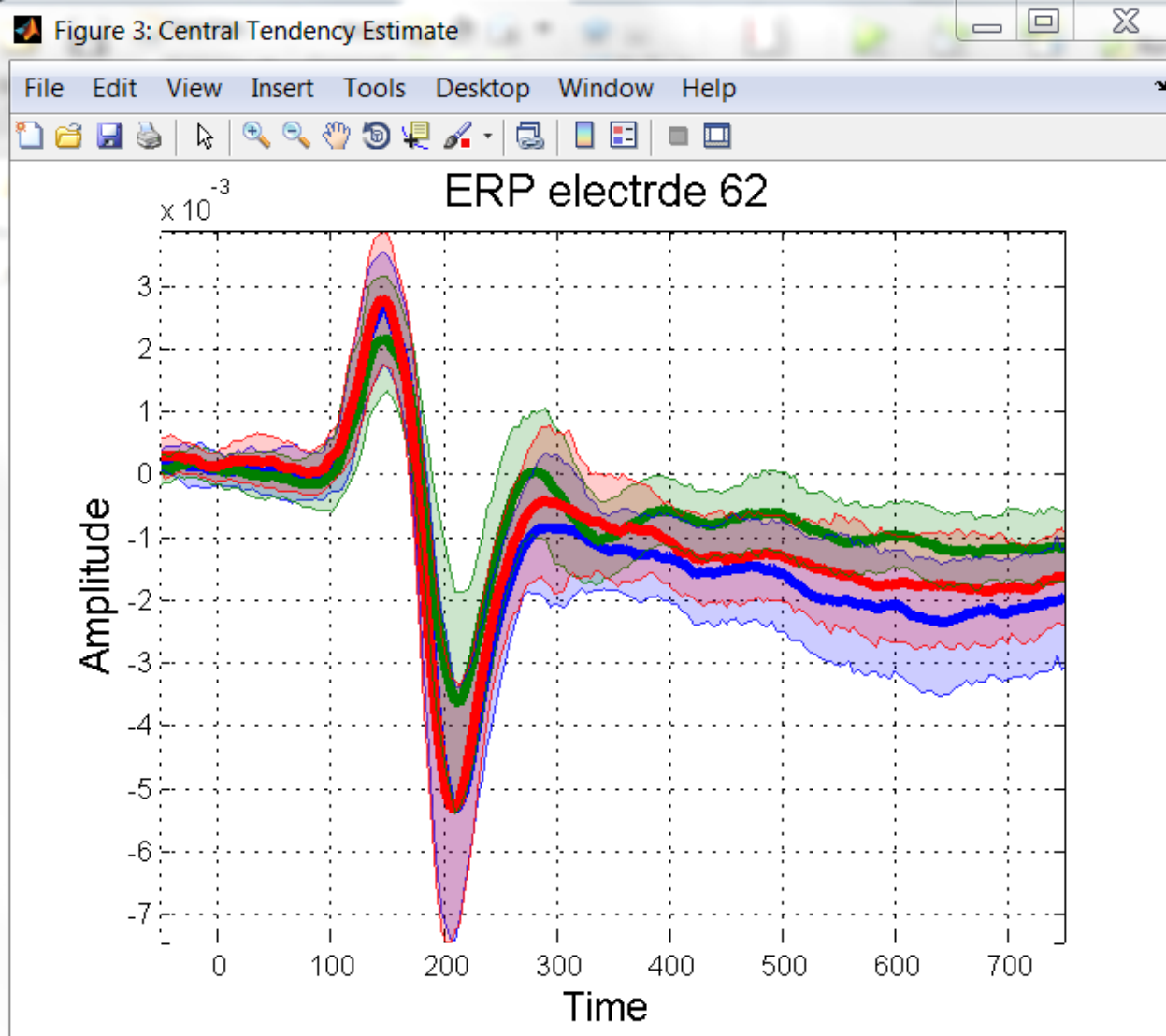
Betas

Copyright (C) LIMO Team 2015 - GNU GPL

Group level analysis

- For Betas – select the txt file listing betas of each subject (you can also load one by one if you like clicking)
 - For ERP – select the txt file listing LIMO files of each subject, use weights
- *Using Plot central tendency and CI, you should be able to:*
- 1 – see the mean beta 1 / 2 / 3 used in the ANOVA
 - 2 – see the corresponding ERP for Famous faces, scrambled faces and non famous faces

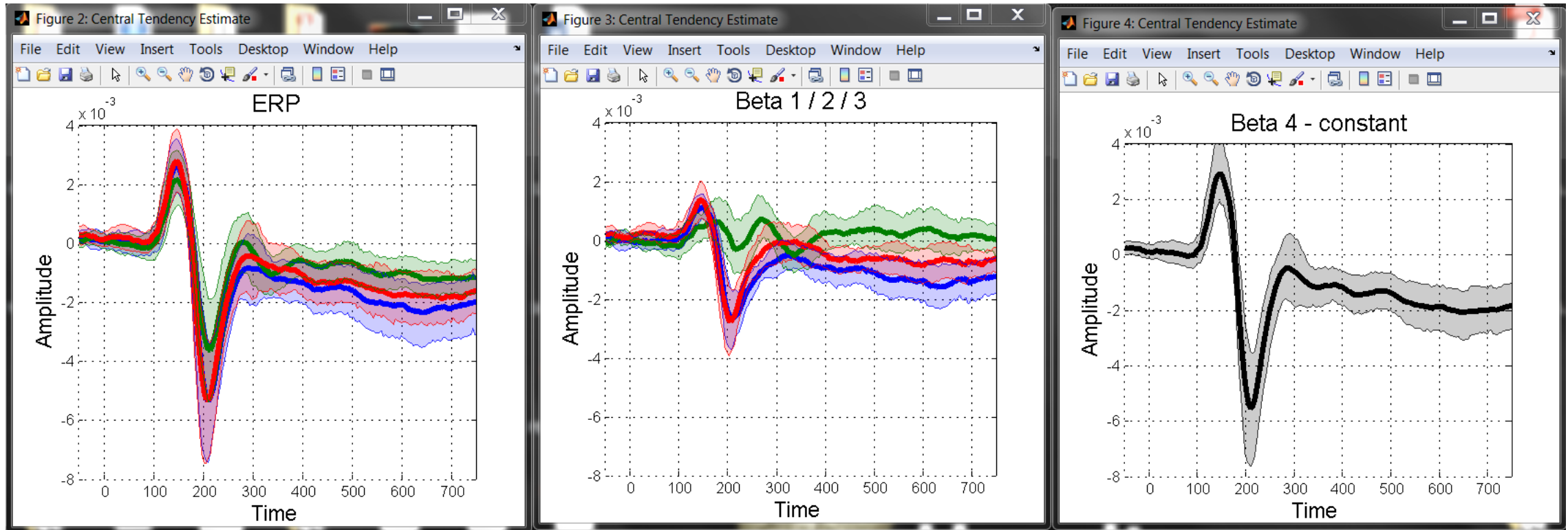




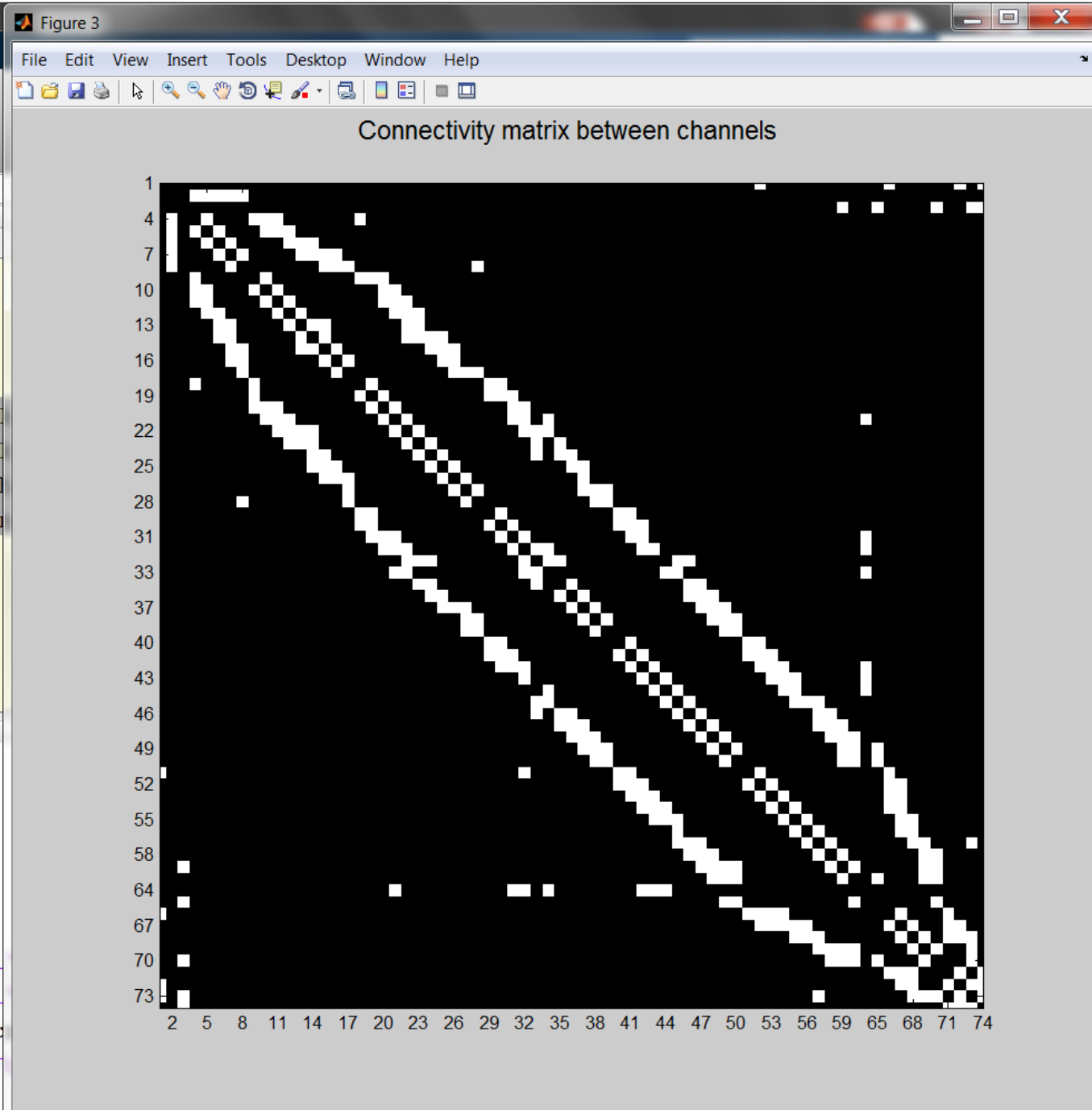
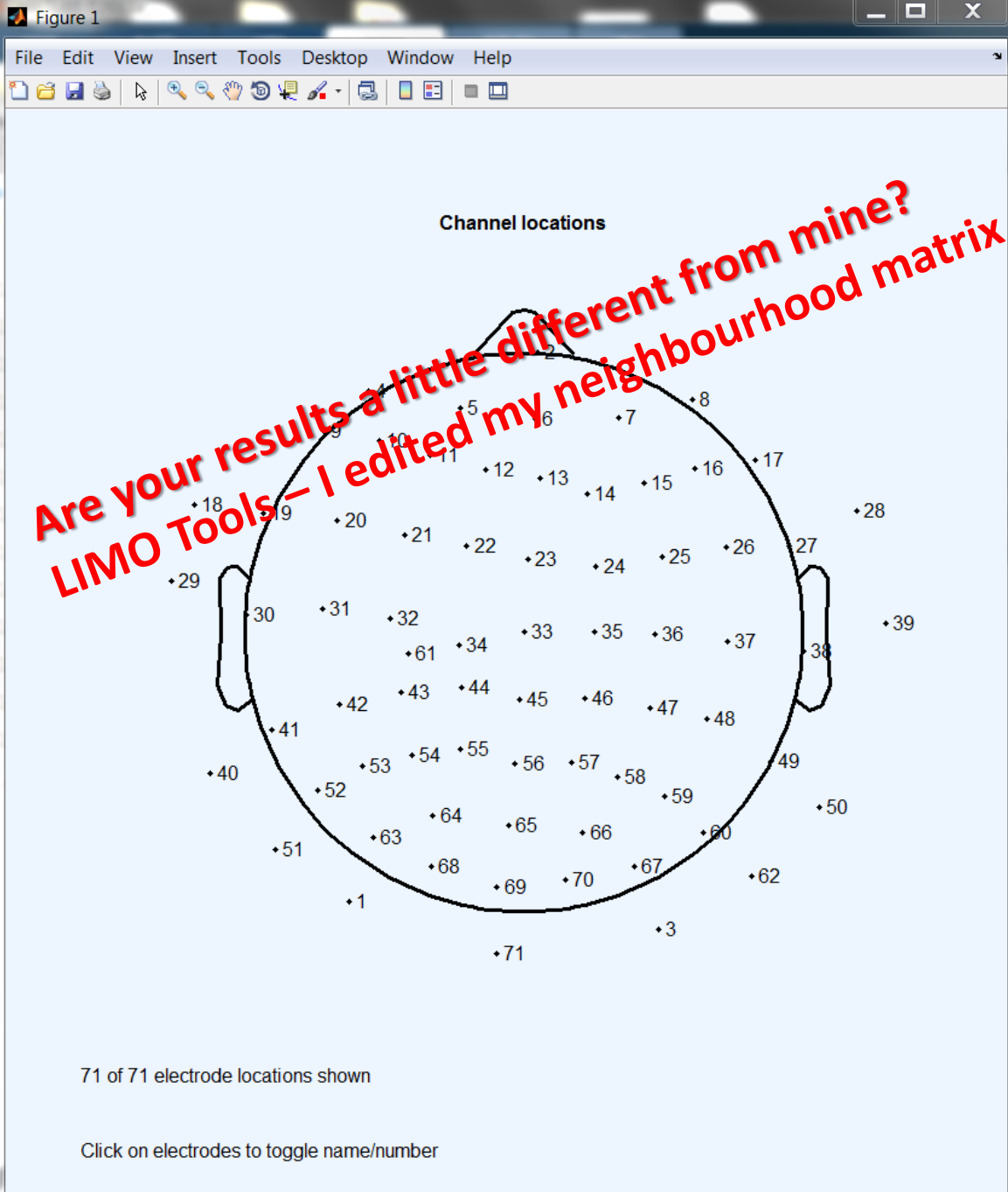
Our ERP looks a lot like the published results – replicated ! (but not so such betas ?)

Bonus: 95% CI are Bayesian = this is the probability of the estimator

Wait a minute ! The model was ...



$$Y \text{ (the data)} = FF * \text{Beta1} + SF * \text{Beta2} + NFF * \text{Beta 3} + \text{Beta 4} + e$$



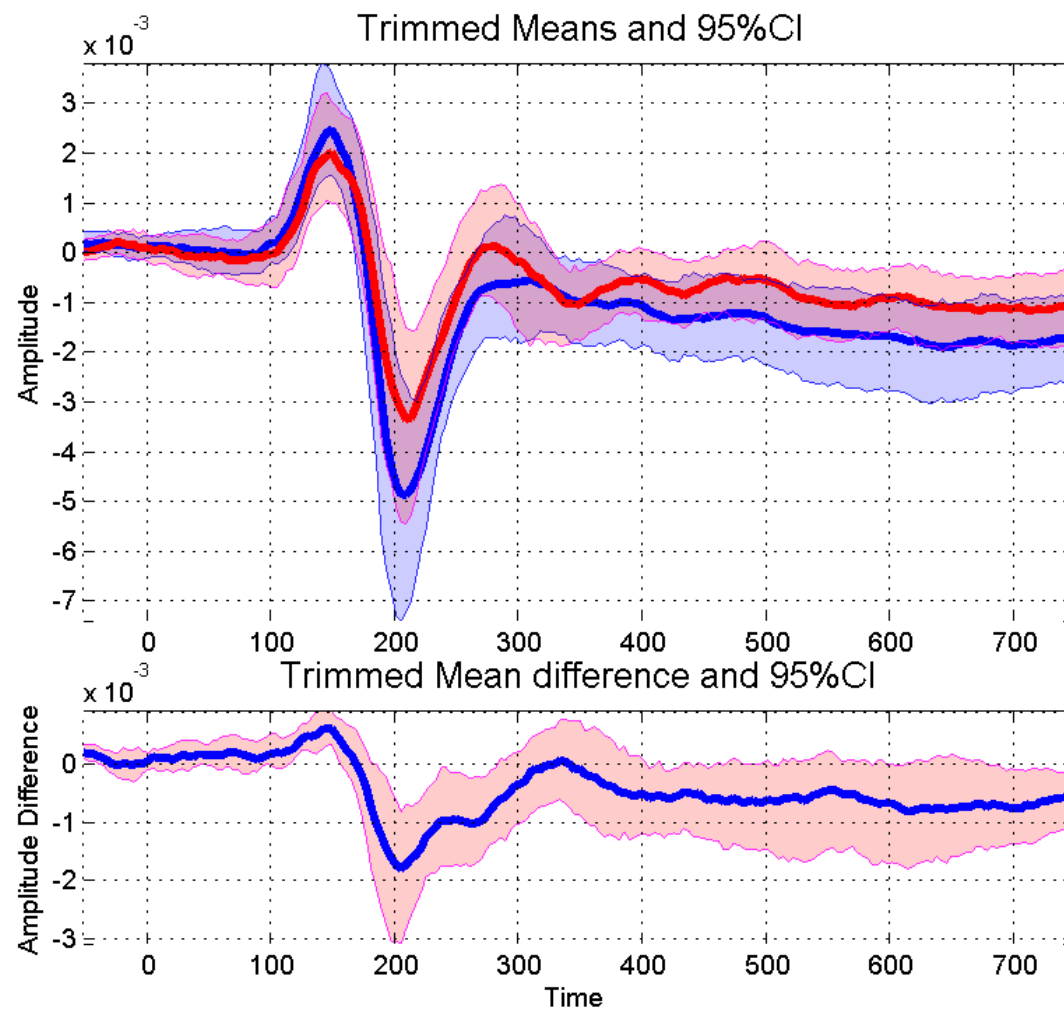
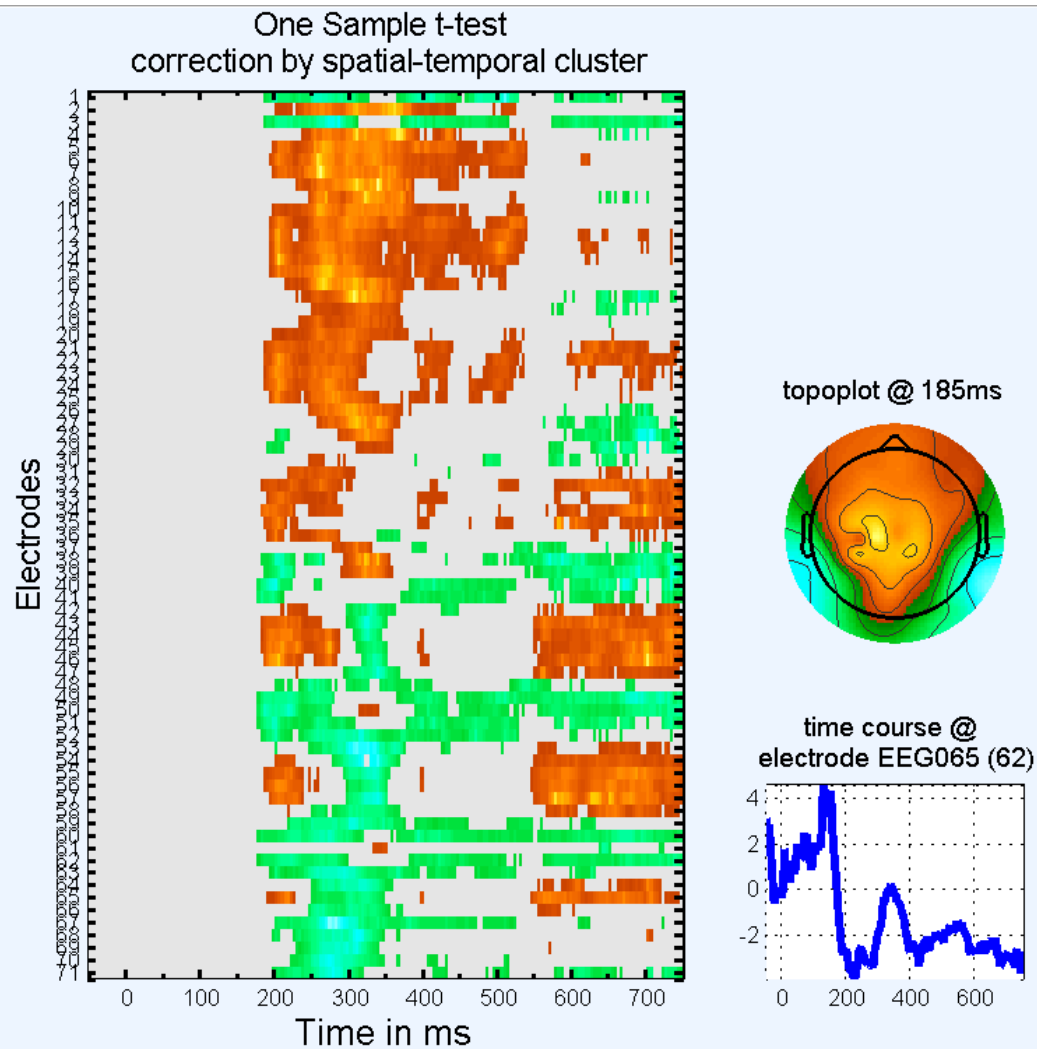
Let's try faces vs scrambled !

- Per subject compute a contrast famous + non-famous > scrambled
- $[1 \ -2 \ 1]$ tests $[\text{famous} - 2 * \text{scrambled} + \text{non-famous}] = 0$

LIMO BATCH

- At the group level, do a one-sample t-test
- At the group level, you can build an ERP pooling famous + non-famous

faces vs scrambled



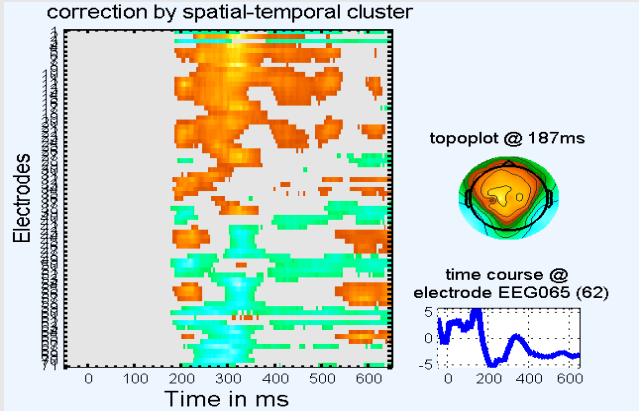
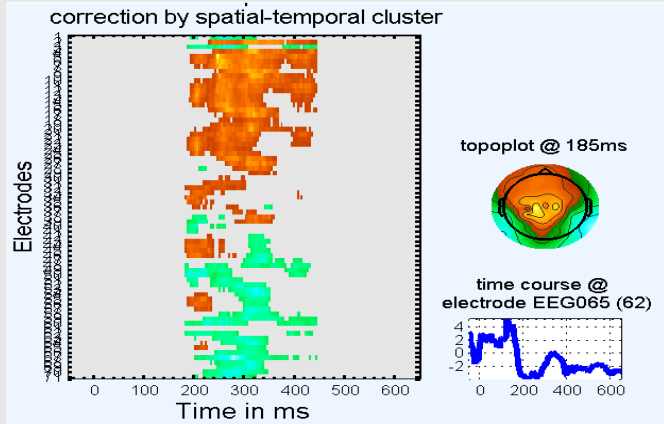
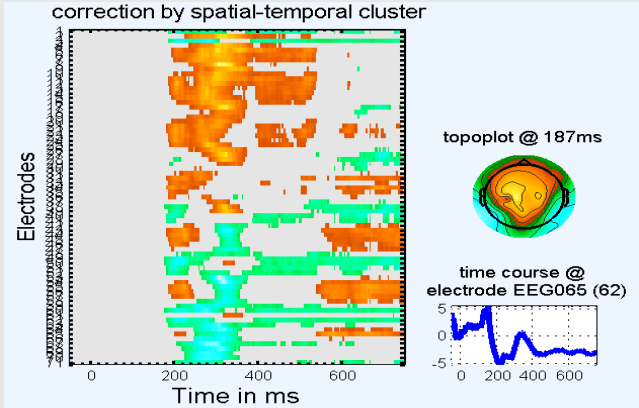
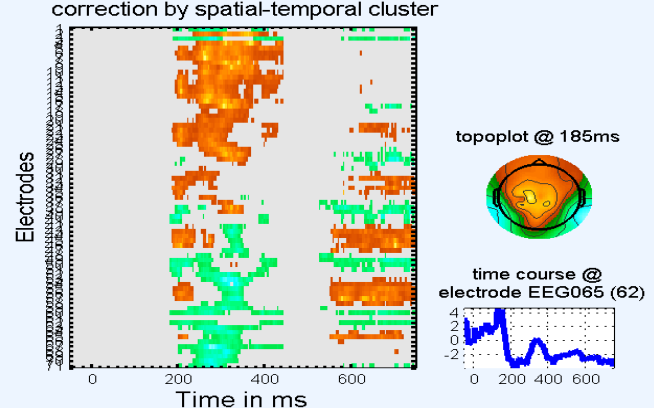
Why is LIMO Robust?

- Standard stats are all instantiations of a GLM using an Ordinary Least Square solution → implies looking at the mean
- the breakdown point of an estimator is the proportion of incorrect observations (e.g. arbitrarily large observations) an estimator can handle before giving an incorrect
- For data x_1 to x_n – the mean has a bkdp of 0 because we can make the mean large changing any x_i – the median has a bkdp of 50%

Why is LIMO Robust?

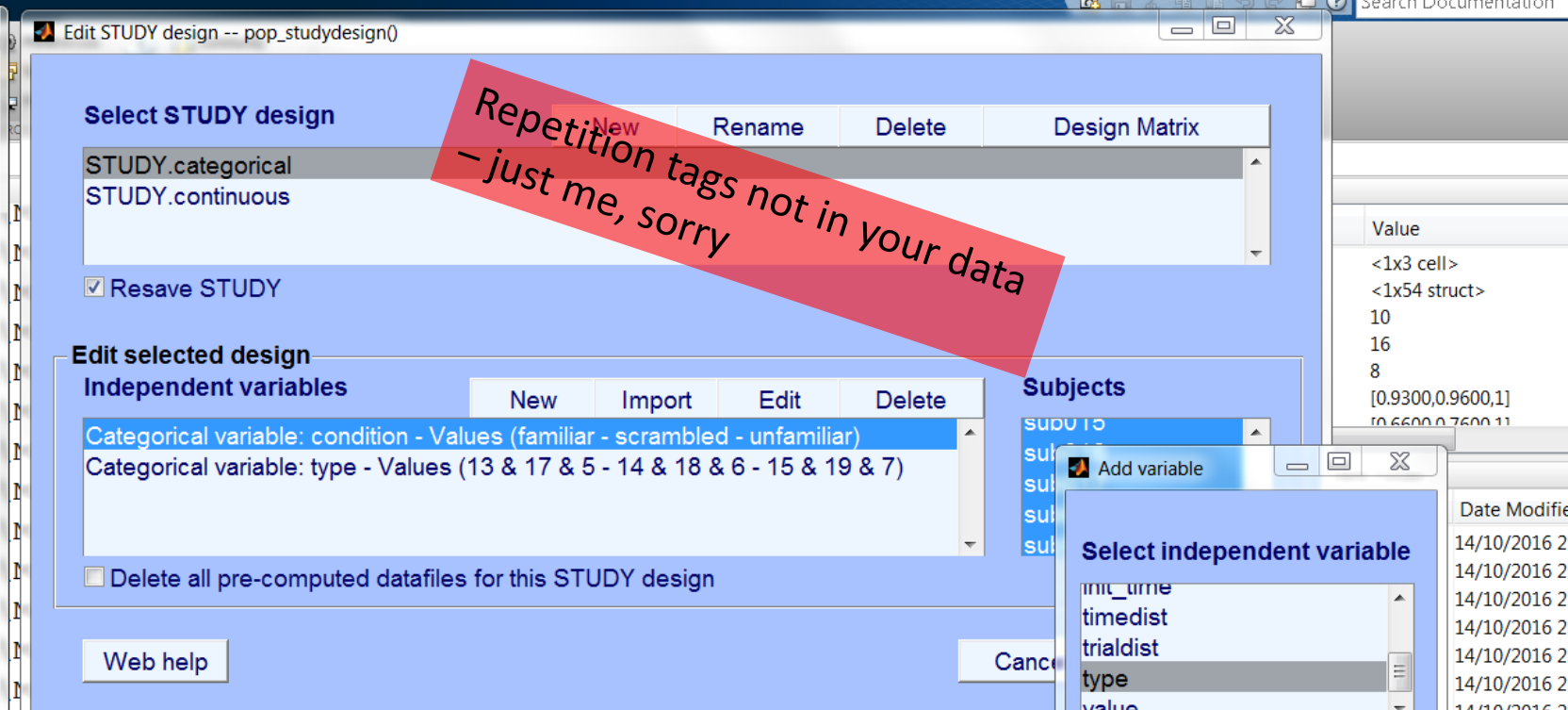
- Are you sure?
- [Micceri \(1989\). The Unicorn, The Normal Curve, and Other Improbable Creatures. Psych Bul. 105, 156-166](#)
- If the data are Gaussian, the median, the trimmed mean is the same as the mean !
So no reason not to use alternative techniques.
- 1st level, uses weighted least square (weights down bad trials – bkdp variable)
- 2nd level involves 20% trimmed mean (weights = 0 for bad subjects): t-tests, 1-way ANOVA, Repeated Measures ANOVA (soon)
- For regressions and N-way ANOVA/ANOVA we use an IRLS (all subjects have weights from 0 to 1 – bkdp variable)

Let's look again at faces vs scrambled

	T-test on mean	T-test on trimmed mean
OLS	<p>correction by spatial-temporal cluster</p>  <p>Electrodes</p> <p>Time in ms</p> <p>topoplot @ 187ms</p> <p>time course @ electrode EEG065 (62)</p>	<p>correction by spatial-temporal cluster</p>  <p>Electrodes</p> <p>Time in ms</p> <p>topoplot @ 185ms</p> <p>time course @ electrode EEG065 (62)</p>
WLS	<p>correction by spatial-temporal cluster</p>  <p>Electrodes</p> <p>Time in ms</p> <p>topoplot @ 187ms</p> <p>time course @ electrode EEG065 (62)</p>	<p>correction by spatial-temporal cluster</p>  <p>Electrodes</p> <p>Time in ms</p> <p>topoplot @ 185ms</p> <p>time course @ electrode EEG065 (62)</p>

3 x 3 repeated measures ANOVA

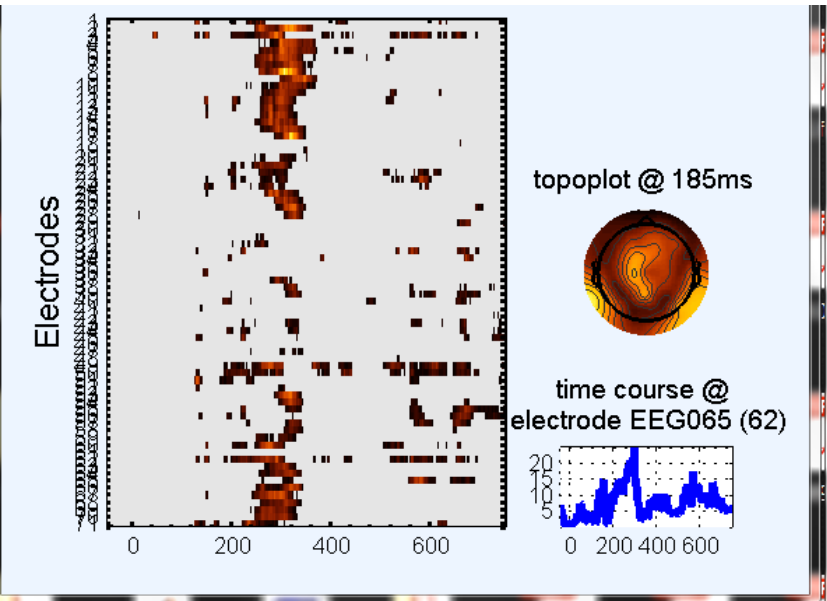
- Since we have 3 types of faces and 3 repetition levels – we can do a 3 by 3 ANOVA
- Question: How many 1st level regressors?
- Question: Why not modelling interaction at the subject level?



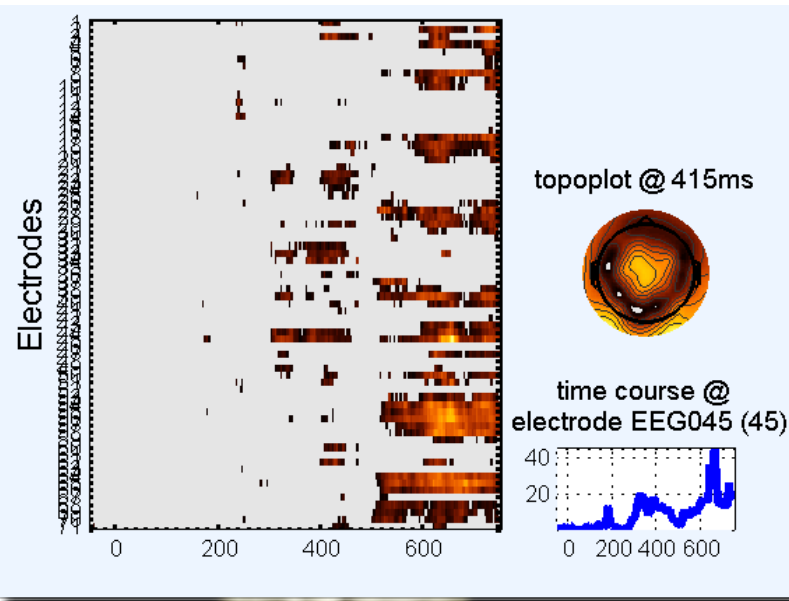
{ '17' '18' '19' } : Scrambled faces (1st presentation, 2nd presentation, 3rd presentation delayed)

3 x 3 repeated measures ANOVA

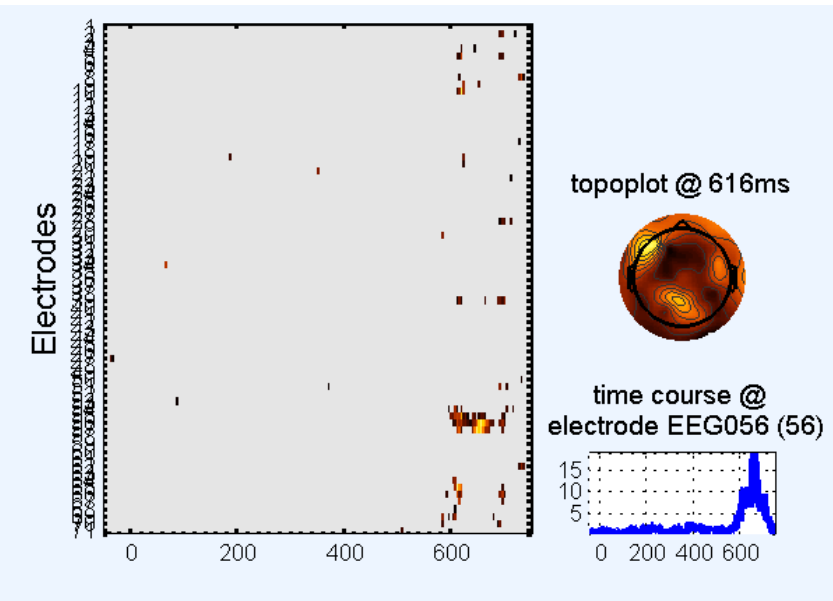
Category effect



Repetition effect



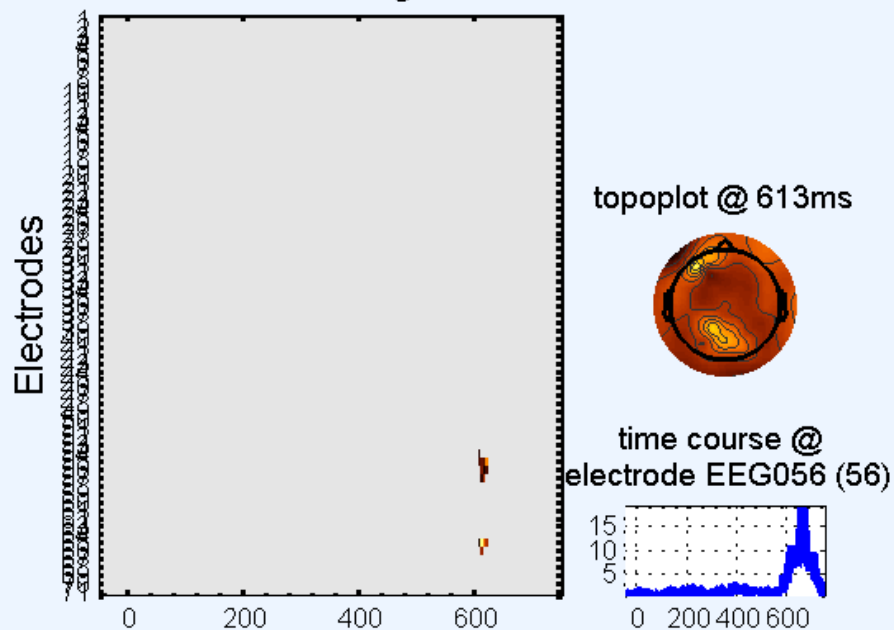
Interaction effect



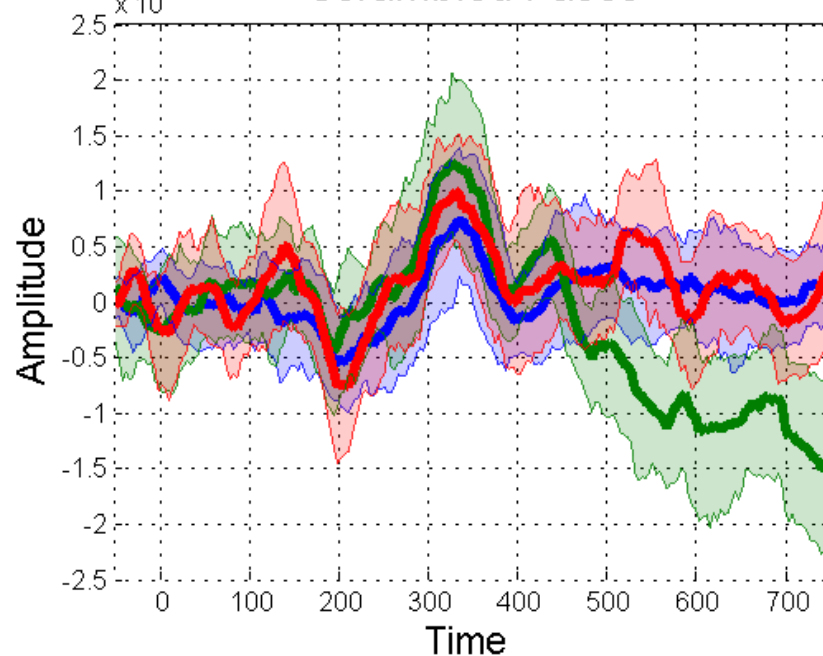
dataviz @ $p = 0.005$ uncorrected

Inference TFCE show category effect @ 200ms +, repetition effect @ 600ms +, and interaction at 600ms +

correction using TFCE



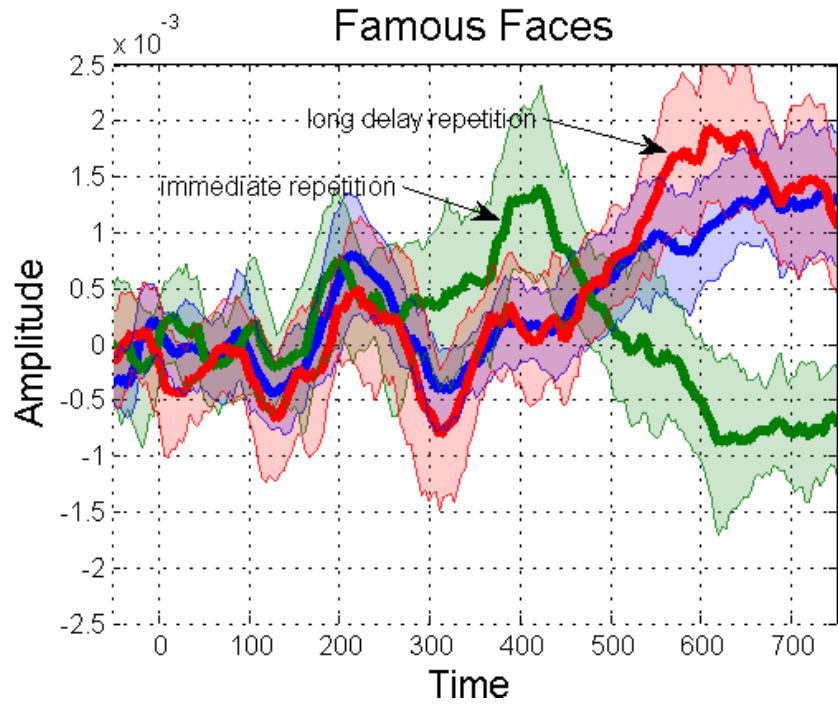
Scrambled Faces



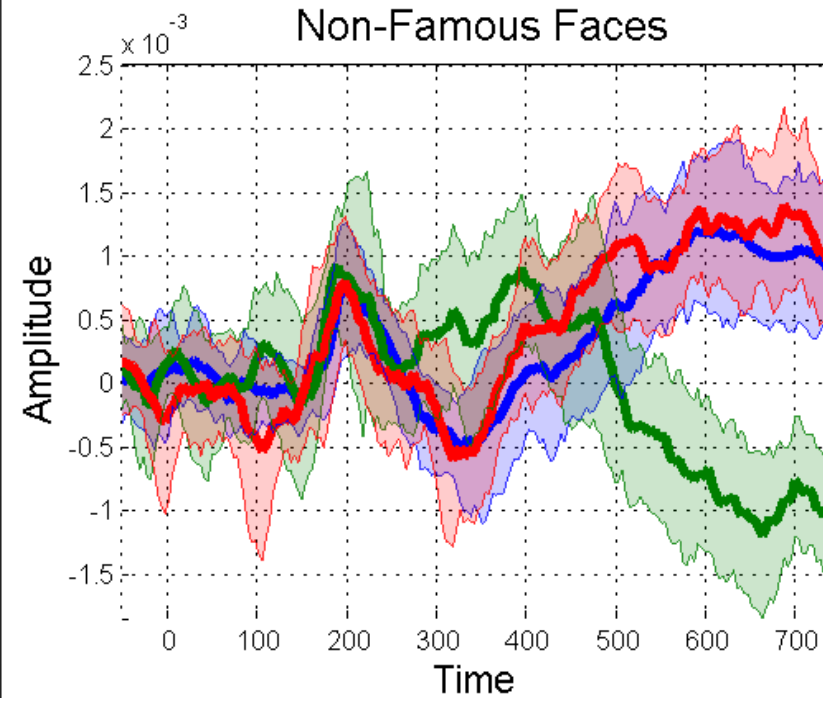
We have a main repetition effect driven by the direct repetition starting circa 380ms but sig only at 550ms

We have a small interaction effect with famous faces showing enhanced activity from circa 550ms

Famous Faces



Non-Famous Faces



The maths behind the GUI

- If you want to go for coffee now, it's fine
- 4 slides showing how standard and robust are different

One sample t-test

$$t = \frac{\text{Mean}}{\text{std}/\sqrt{n}}$$

$$p = 2 * \text{tcdf}(\text{abs}(t), \text{df})$$

$$\text{df} = n - 1$$

limo_ttest.m

$$t = \frac{\text{Trimmed Mean}}{\sqrt{\text{WinVar}/(1 - 2 * \text{trimming percentage})} * \sqrt{n}}$$

$$p = 2 * (1 - \text{tcdf}(\text{abs}(t), \text{df}))$$

$$\text{df} = n - 2 * \text{floor}((\text{trimming percentage}/100) * n) - 1$$

limo_trimci.m

Paired t-test

$$t = \frac{\text{Mean (difference)}}{\text{std (difference)}/\sqrt{n}} \quad p = 2 * \text{tcdf}(\text{abs}(t), \text{df}) \text{ with } \text{df} = n - 1$$

limo_ttest.m

$$t = \frac{\text{Difference of trimmed means}}{\sqrt{\frac{(WinVar1 * (n - 1)) + (WinVar2 * (n - 1)) - (2 * (n - 1) * WinCov)}{(n - 2) * n \text{ trim}}}}$$

$$p = 2 * (1 - \text{tcdf}(\text{abs}(t), \text{df})) \text{ with } \text{df} = ((n - 2) * n \text{ trim}) - 1$$

limo_yuend_ttest.m

Two-samples t-test

$$t = \frac{\text{mean}(gp1) - \text{mean}(gp2)}{\sqrt{\frac{\text{var}(gp1)}{n1} + \frac{\text{var}(gp2)}{n2}}}$$

$$p = 2 * \text{tcdf}(\text{abs}(t), df)$$

$$df = \frac{(s1 + s2)^2}{\frac{s1}{n1 - 1} + \frac{s2}{n2 - 1}}$$

limo_ttest.m

$$t = \frac{\text{Difference of trimmed means}}{\sqrt{\frac{(n1 - 1) * \text{WinVar1}}{n1 \text{ trim} * (n1 \text{ trim} - 1)} + \frac{(n2 - 1) * \text{WinVar2}}{n2 \text{ trim} * (n2 \text{ trim} - 1)}}$$

$$p = 2 * (1 - \text{tcdf}(\text{abs}(t), df))$$

$$df = \frac{(\text{Yuen } s1 + \text{Yuen } s2)^2}{\frac{\text{Yuen } s1}{n1 \text{ trim} - 1} + \frac{\text{Yuen } s2}{n2 \text{ trim} - 1}}$$

limo_yuen_ttest.m

IRLS

- **Limo_wls.m and limo_irls.m (for trials vs across subjects)**
- Start by OLS to obtain residuals
- Check outliers in standardized residuals (MAD)
- Compute weights (bisquare function)
- Recompute on weighted data
- Check residuals again until $E(e) = 0$
 - for eeg, iterate until $\max(\text{abs}(\text{oldRes} - \text{newRes})) < (0.0001)$

$$Wy = WX \beta + We, \quad E(e) = 0, \quad \text{Cov}(e) = \sigma^2 I$$