

# TRENTOOL workshop

# TRansfer ENtropy TOOLbox

Dr. Michael Lindner



# TRENTOOL

# TRENTOOL

- TRENTOOL
  - is a Matlab based toolbox
  - is designed to use transfer entropy (TE) on any kind of time series data.
  - is designed to make the non trivial handling of TE easy for all users.
  - offers solutions for optimizing parameters of TE calculation.
  - Infos and download: [www.trentool.de](http://www.trentool.de)

# The main functions of TRENTOOL

- TEprepare
- TEsurrogatestats
- TEconditionstatssingle
- TEgroup\_prepare
- TEgroup\_calculate
- TEgroup\_stats
- TEplot2D

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# The main functions of TRENTOOL

single subject  
analyses

- TEprepare
- TEsurrogatestats
- TEconditionstatssingle
- TEgroup\_prepare
- TEgroup\_calculate
- TEgroup\_stats
- TEplot2D

# The main functions of TRENTOOL

- TEprepare
- TEsurrogatestats
- TEconditionstatssingle

group  
analyses

- TEgroup\_prepare
- TEgroup\_calculate
- TEgroup\_stats

- TEplot2D

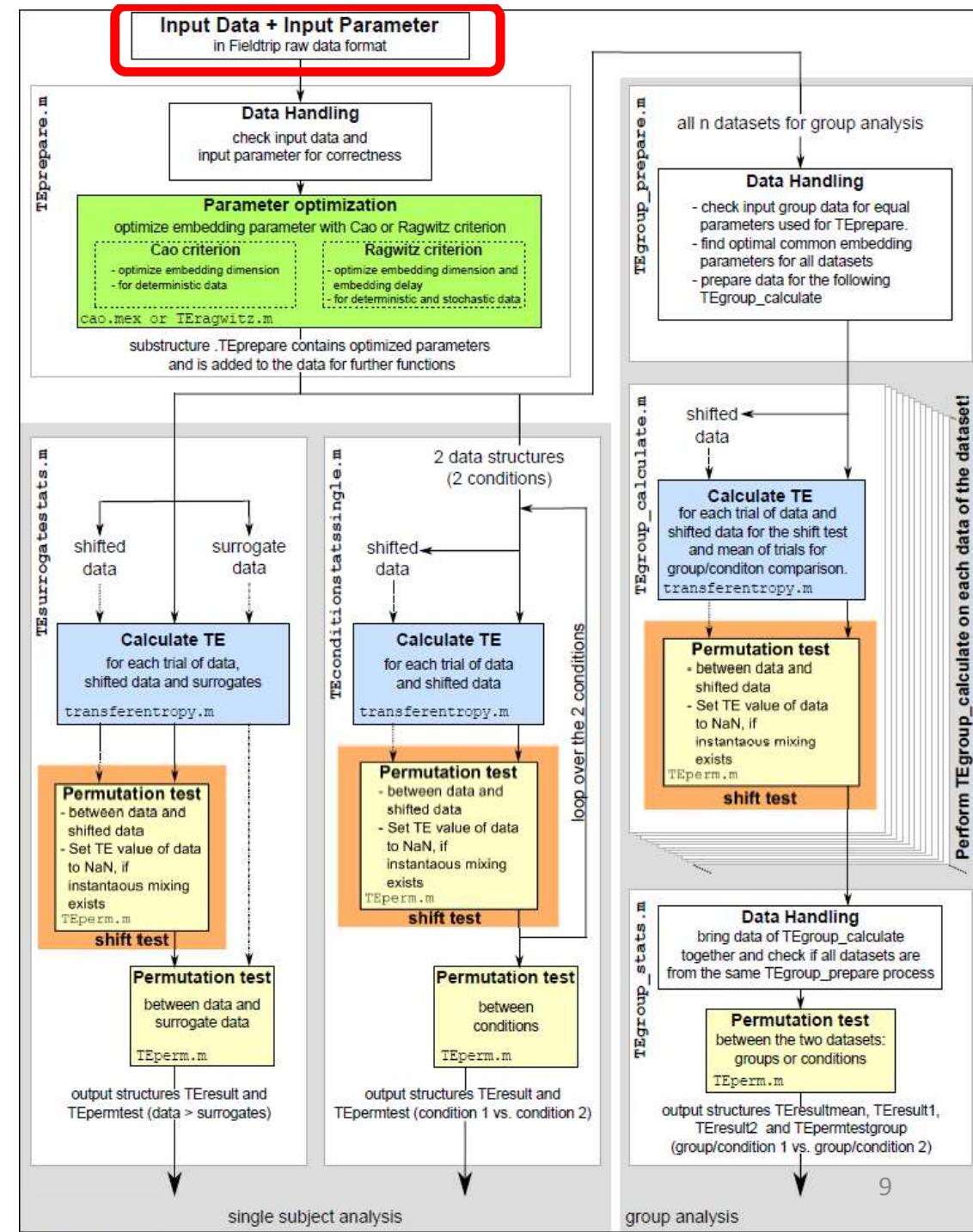
# The main functions of TRENTOOL

- TEprepare
- TEsurrogatestats
- TEconditionstatssingle
- TEgroup\_prepare
- TEgroup\_calculate
- TEgroup\_stats

Plotting results

- TEplot2D

# Overview TRENTOOL



# Input - Data

- In Fieldtrip raw data structure:

Matlab structure containing:

{ } = Cells

- {trials} ← (channel x time)
- {time} ← (time indices)
- {labels} ← (labels of the channels)
- fsample

# Input - Data

- In Fieldtrip raw data structure:

Matlab structure containing:

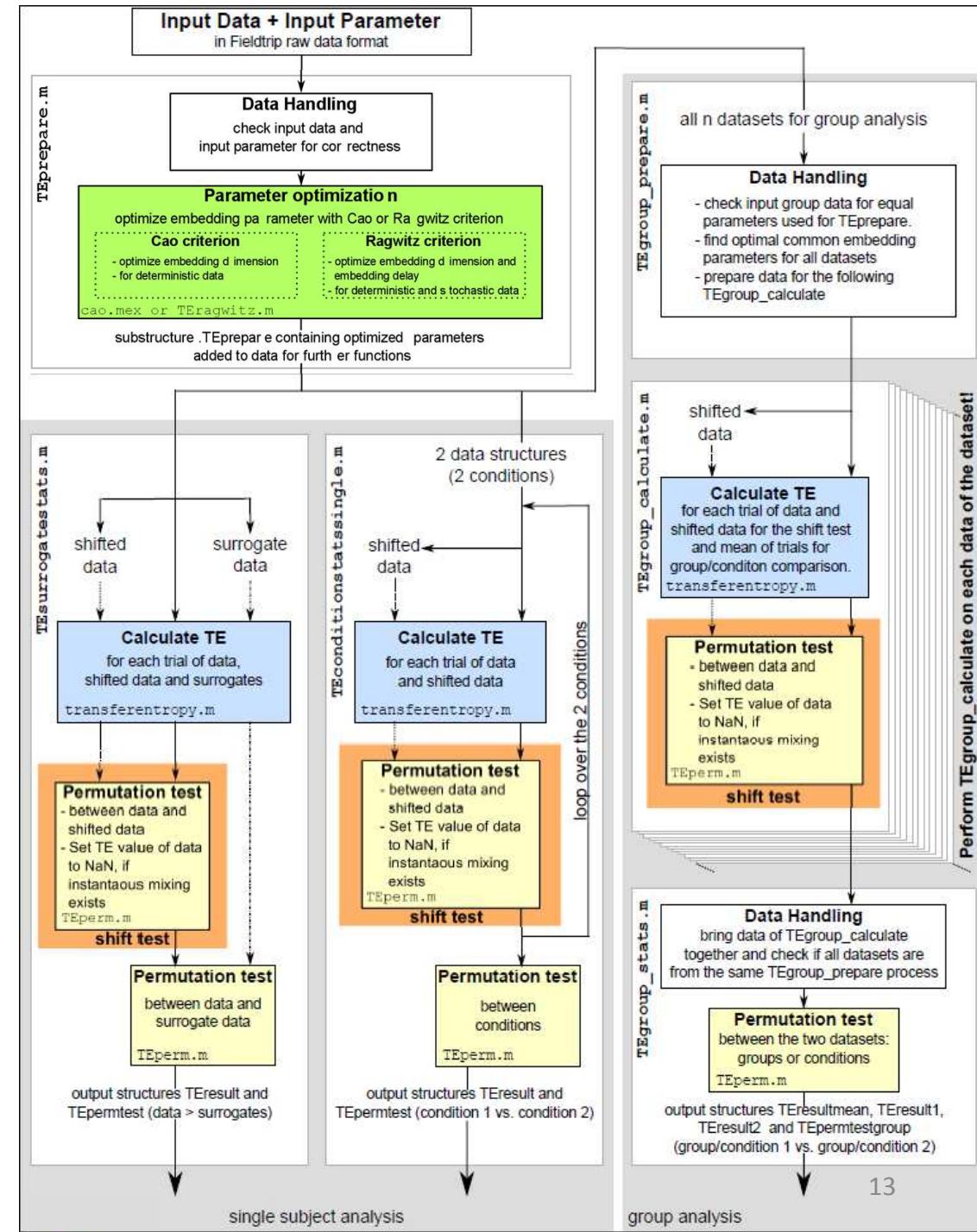
**help TEprepare** { } = Cells

- {time} (1xnumber of trials)
- {labels} (1xnumber of channels)
- fsample

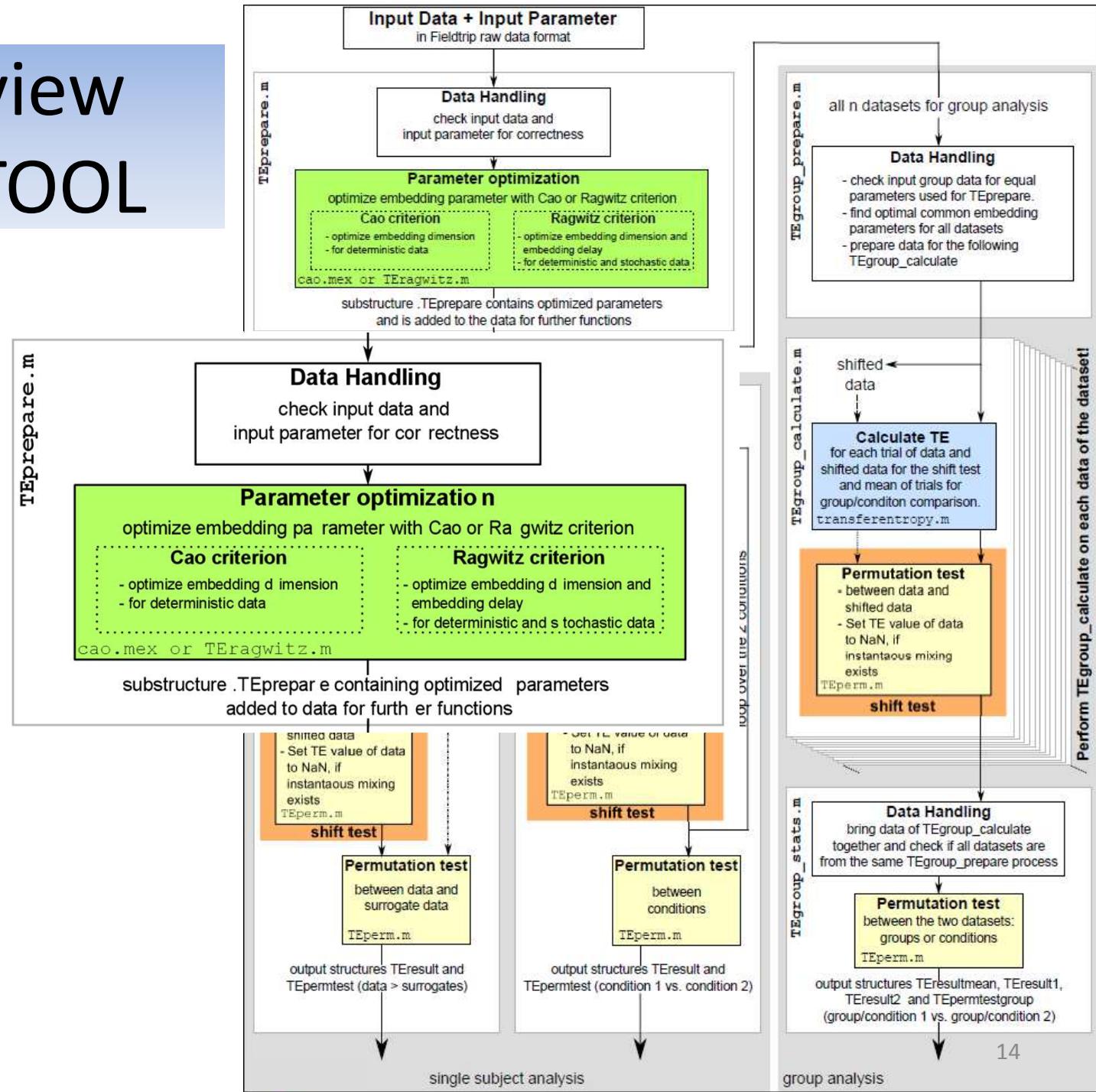
# Input - Parameter

- Parameters are given using a MATLAB structure (typically called „cfg“ - configuration)
- Type: `help <functionname>` to see which parameter is needed.

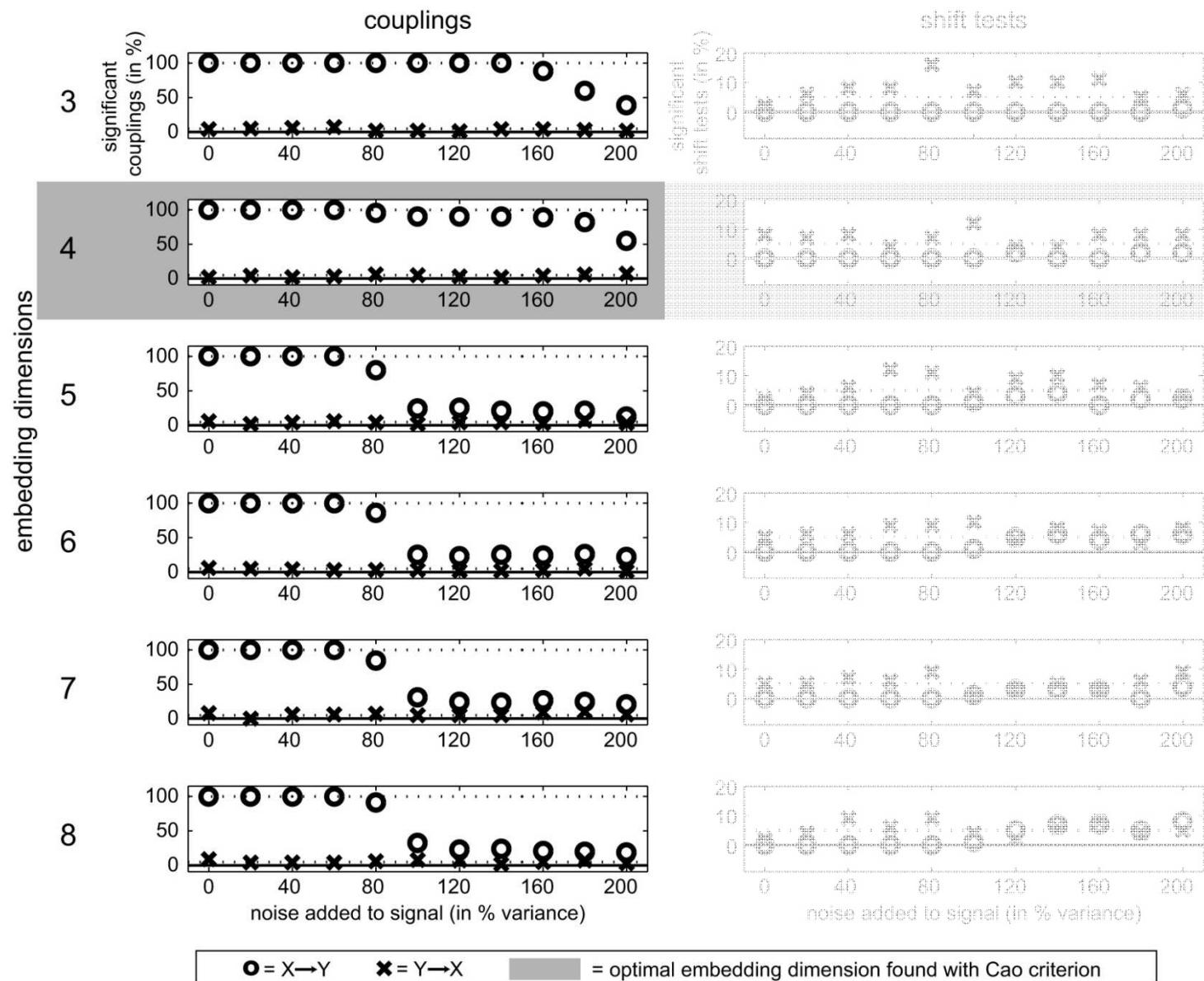
# Overview TRENTOOL



# Overview TRENTOOL



# Parameter optimization



# All parameters for TEprepare

**cfg.sgncmb** = list of channelpairs  
cell array (Nx(source, target))

or

**cfg.channel** = list of channels - testing will be done all-by-all

and

**cfg.Path2TSTOOL** = Path to the folder including the TSTOOL package

**cfg.toi** = the time range of interest (vector 1 x 2) in seconds  
e.g. (time\_from, time\_to) (units: seconds)

**cfg.predicttime\_u** = time ahead for the advance prediction (scalar, in ms)

**cfg.optimizemethod** = Method to optimize parameters: 'ragwitz' or 'cao'  
if you choose 'ragwitz':

**cfg.ragdim** = range of embedding dimensions to scan vector  
(1xnumdim)

**cfg.ragtaurange** = vector (1x2) of min and max embedding delays (in multiples of the autocorrelation decay time)

**cfg.ragtasteps** = number of equidistant steps in ragtaurange  
(min 5) (default = 10)

**cfg.flagNei** = 'Range' or 'Mass' type of neighbor search

**cfg.sizeNei** = Radius or mass for the neighbor search according to flagNeighborhood

**cfg.repPred** = repPred represents the number of points for which the prediction is performed (it has to be smaller than length(timeSeries)-(dimEmb-1)\*tauEmb-u)

if you choose 'cao' (recommended for fMRI Data):

**cfg.caodim** = range of embedding dimension to scan with stepwidth 1 (vector 1 x numdim)  
(default = [1,2,3,4,5,6,7,8,9,10])

**cfg.caokth\_neighbors** = number of neighbors for fixed mass search for cao (controls balance of bias/statistical errors) (default = 4)

**cfg.caotau** = embedding delay in units of ACT (x\*ACT)  
(default = 1.5)

**cfg.kth\_neighbors** = number of neighbors for fixed mass search  
(controls balance of bias/statistical errors)  
(default = 4)

**cfg.TheilerT** = number of temporal neighbors excluded to avoid serial correlations (Theiler correction) (default = ACT)

**cfg.trialselect** = ACT threshholding of trials - 'ACT' , 'range' or 'no'  
(default = 'ACT')

if you chose 'ACT' (or nothing):

**cfg.actthrvalue** = max threshold for ACT; min threshold

**cfg.minntrials** = minimum Nr of trials with ACT < actthrest used to calculate transfer entropy

if you chose 'range':

**cfg.trial\_from** = Inferior limit for the trials to be considered

**cfg.trial\_to** = Superior limit for the trials to be considered

**cfg.maxlag** = the range of lags for computing the auto correlation time: from -MAXLAG to MAXLAG (default = 1000)

**cfg.TEcalcype** = 'V' : self-prediction of the target signal and cross-prediction are both made from states in source and target that precede the target state to be predicted by cfg.predicttime\_u.

'VW' : the self-prediction time for the target is tau and cross-predictions are made from source states that precede the target state to be predicted by cfg.predicttime\_u.  
(to solve the problem of decreasing self-prediction accuracy for large prediction times)

(default = 'VW')

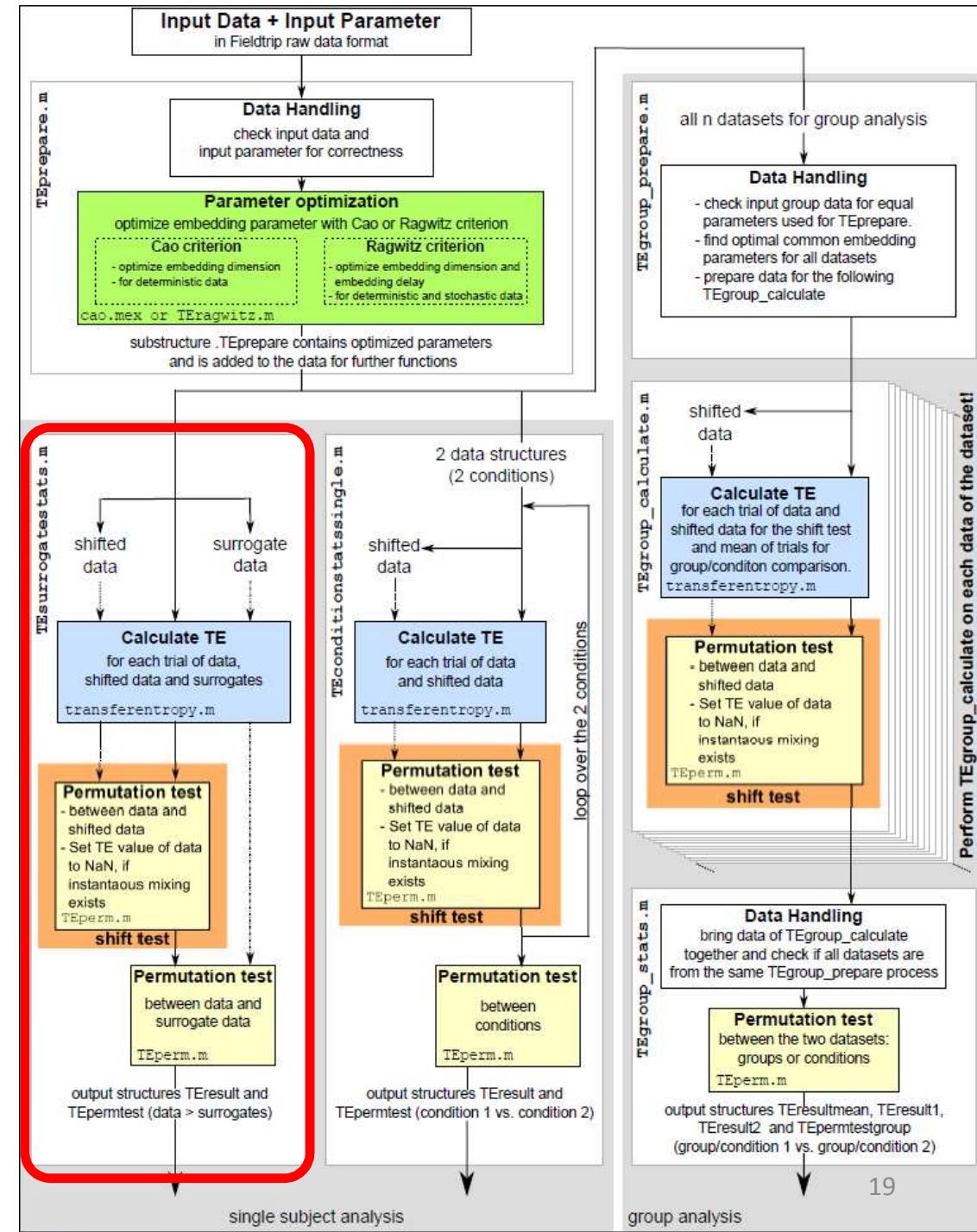
# Possible parameter set for TEprepare

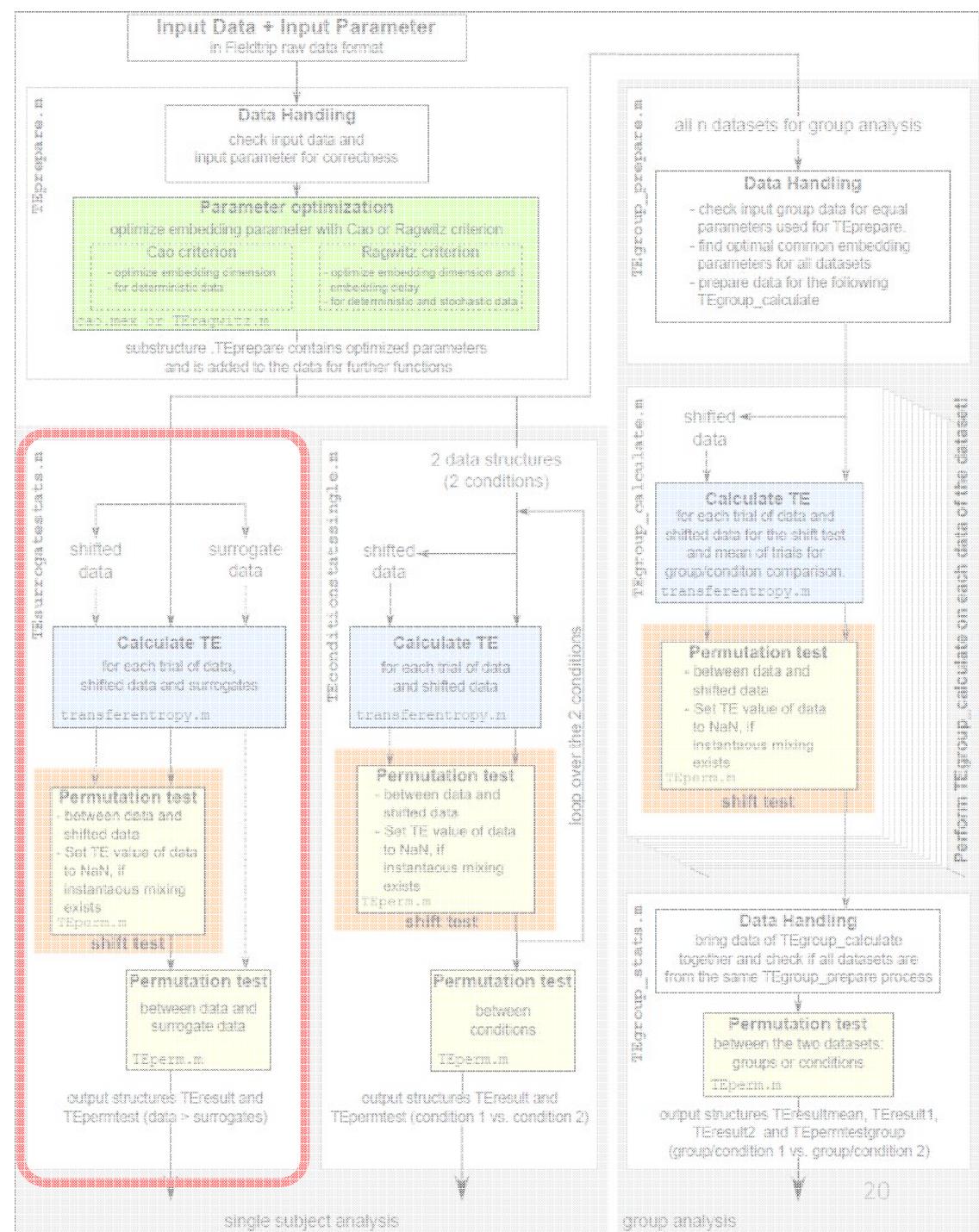
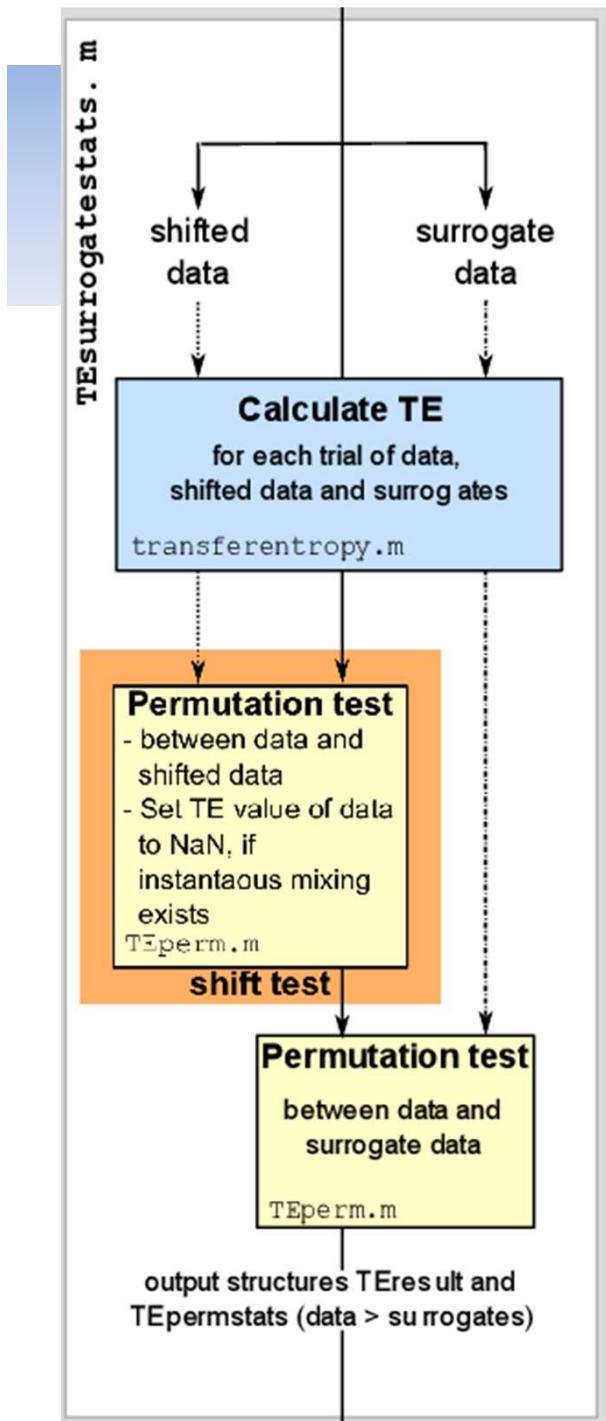
```
cfg.toi = [0 2.999];
cfg.sgncmb = {'X', 'Y';
              'Y', 'X'};
cfg.optimizemethod = 'cao';
cfg.caodim = 1:6;
cfg.trialselect = 'ACT';
cfg.actthrvalue = 120;
cfg.minnrtrials = 30;
cfg.Path2TSTOOL = '/data/common/OpenTSTOOL';
cfg.predicttime_u = 21;
```

# Run TEprepare

```
Data_prepared = TEprepare(cfg,Data)
```

# Overview TRENTOOL

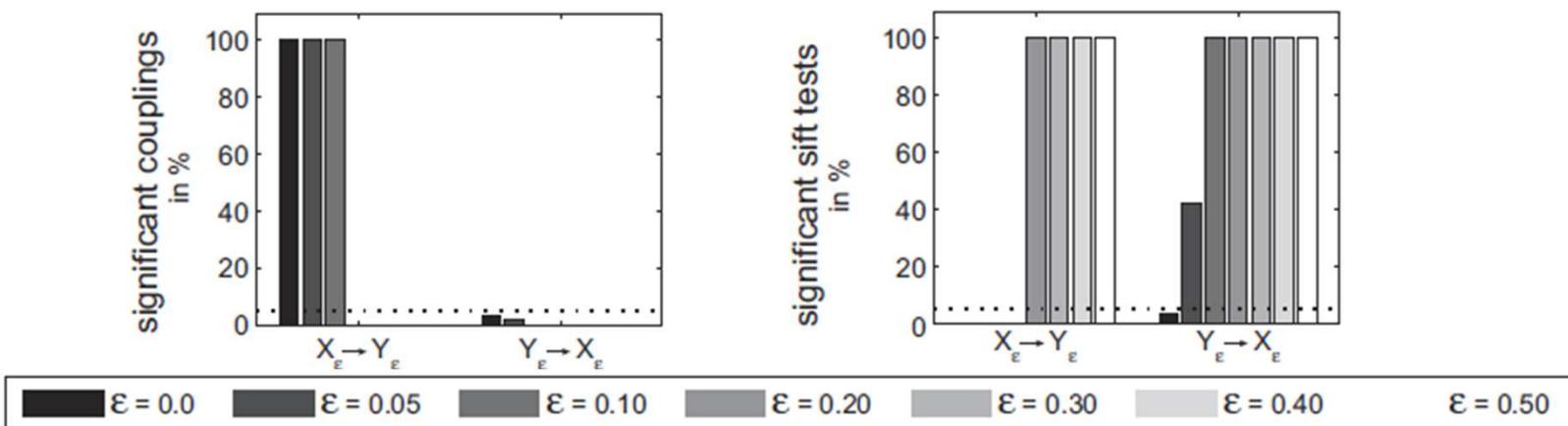




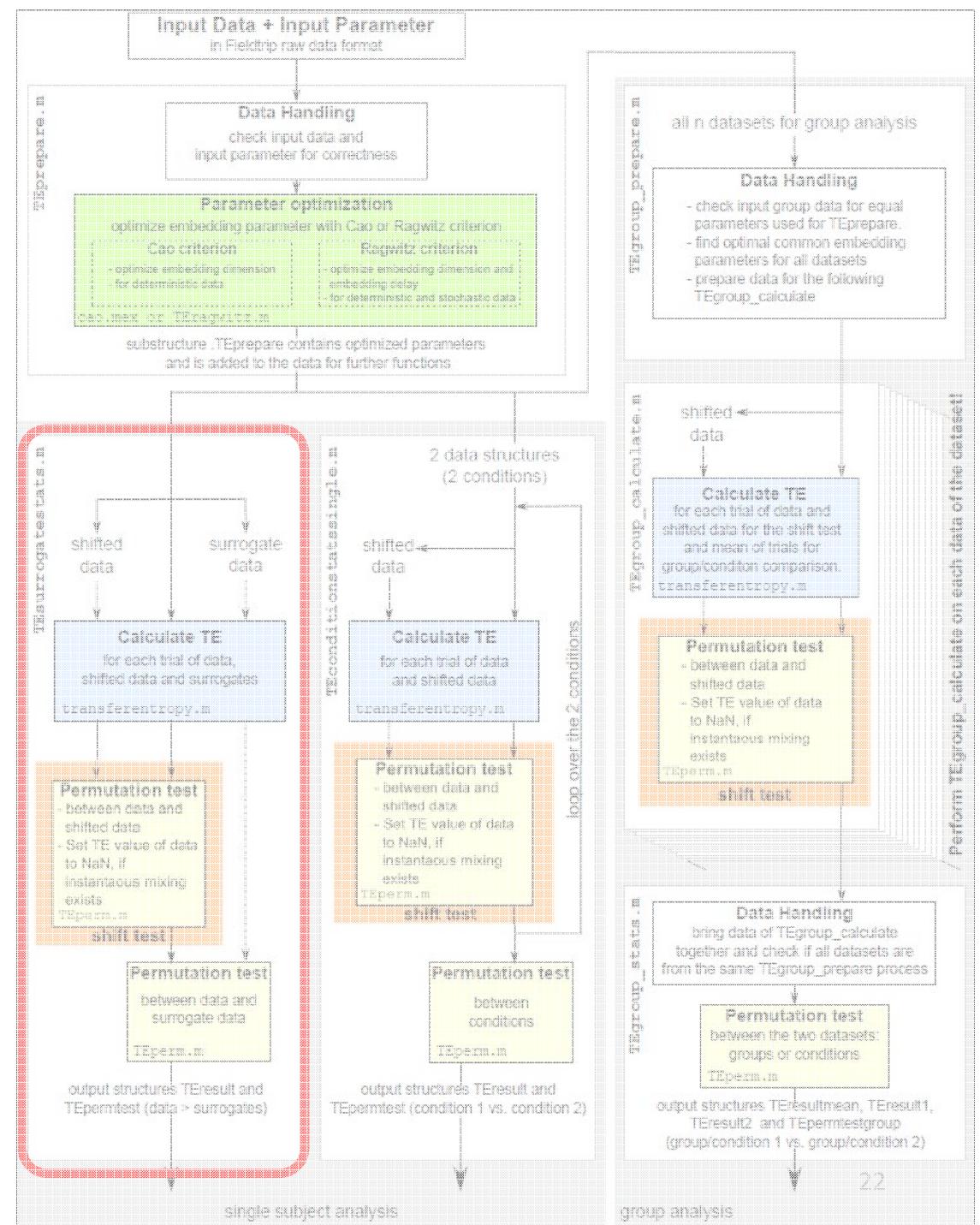
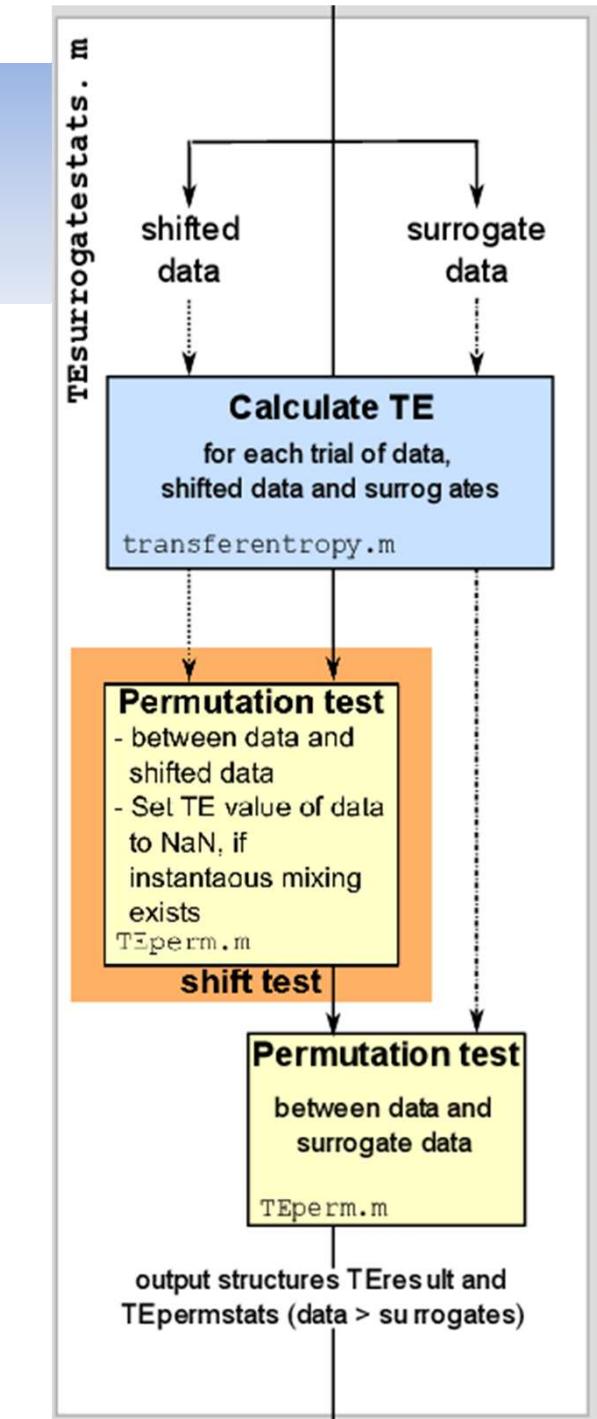
# Shift test

- detecting volume conduction in mixed signals (e.g. MEG)
- Permutation test between TE of the data and of the shifted data

$$\begin{array}{ccc} \begin{array}{c} X(t) \\ Y(t) \end{array} & \xrightarrow{\epsilon} & \begin{array}{c} X_\epsilon(t) \\ Y_\epsilon(t) \end{array} \\ (1-\epsilon) \quad \epsilon & & \epsilon \quad (1-\epsilon) \end{array}$$
$$X_\epsilon(t) = (1-\epsilon)X(t) + \epsilon Y(t) + \eta_{sx} \quad Y_\epsilon(t) = (1-\epsilon)Y(t) + \epsilon X(t) + \eta_{sy}$$
$$X(t) = AR(10) \\ Y(t) = AR(10) \\ \text{with } X \rightarrow Y$$



Wibral, Rahm, Rieder, Lindner, Vicente, Kaiser (2011)



# TEresult

- in file with the suffix ‘\_TE\_output’
- Matlab structure
- contains
  - .TEmat : Transfer entropy values
  - .Mlmat : Mutual information values
  - .act : autocorrelation decay time values
  - .trials : trial numbers used for TE calculation
  - .TEprepare
  - ...

# TEresult

- in file with the suffix ,\_TE\_output'
- Matlab structure
- contains
  - .TEmat : Transfer entropy values
  - .MIImat : Mutual information values
  - .act : autocorrelation decay time values
  - .trials : trial numbers used for TE calculation
  - .TEprepare
  - ...

# TEpermtest

- in file with suffix ,\_TEpermtest\_output'
- MATLAB structure
- contains
  - .TEpermvalues (number of channelpairs x 5) :
  - .TEprepare
  - (.TEgroupprepare)
  - ...

# TEpermtest

- in file with suffix ,\_TEpermtest\_output'
- MATLAB structure
- contains
  - .TEpermvalues (number of channelpairs x 5) :

- .TEprepare
- (.TEgroupprepare)
- ...

# TEpermtest

- in file with suffix ,\_TEpermtest\_output'
- MATLAB structure
- contains
  - .TEpermvalues (number of channelpairs x 5) :
    1. p-value
    2. significance (uncorrected)
    3. significance (corrected for multiple comparisons)
    4. statistic value (mean or t-value)
    5. Instantaneuos mixing exists or not
  - .TEprepare
  - (.TEgroupprepare)
  - ...



# All parameters for TEsurrogatestats

**cfg.optdimusage** = 'maxdim' to use maximum of optimal dimensions over all channels for all channels, or 'individm' to use the individual optimal dimension for each channel.  
In case of using ragwitz criterion also the optimal embedding delay tau per channelcombi is used.

**cfg.dim** = Value(s) for embedding dimension. In case of using cfg.optdimusage = 'maxdim' this has to be a scalar value. In case of cfg.optdimusage = 'individm' this has to be a vector of the size (channelcombi x 1). If not specified, the optimal dimension(s) found in TEprep will be used, which is the recommended option!

**cfg.tau** = embedding delay in units of act ( $x^*act$ ). If not specified (recommended option), the tau is used as followed:  
Depending optimizemethod in TEprep:  
'ragwitz' = optimal tau found via ragwitz criteron

'cao' = cfg.tau given by user in Teprep  
If not specified, the optimal embedding delay found in TEprep will be used, which is the recommended option!

**cfg.alpha** = significance level for statisical permutation test and correction for multiple comparison  
(default = 0.05)

**cfg.surrogatetype** = 'trialshuffling', 'trialreverse', 'blockresampling', 'blockreverse1', 'blockreverse2', or 'blockreverse3', surrogate data for trial(n) will be created as following:  
trialshuffling: trial(n+1)  
trialreverse: reverse of trial(n)  
blockresampling: cuts trial(n) at random point and resamples the trial  
blockreverse1: reverse after blockresampling  
blockreverse2: reverse first block after blockresampling  
blockreverse3: reverse second block after

blockresampling  
swapneighbors: pair odd trials with the higher neighbor and 3even with the lower neighbor

**cfg.shifttest** = perform shift test to identify instantaneous mixing between the signal pairs. Values: 'yes' or 'no'  
(default = 'yes')  
This shift test is important for EEG and MEG data, because linear mixing is always present in the data. In case of instantaneous mixing transfer entropy should not be calculated for the affected channelpairs with the corresponding parameter sets, because it could result in false positive results. Hence the TE values for these cases will be set to NaN and the corresponding p-values of the permutation test to 1.

**cfg.shifttesttype** = The shift test can be calculated for the direction TE value of original data > TE values of shifted data (value = 'TE>TEshift') or for the other direction (value = 'TEshift>TE'). In this case the alpha is set to 0.1 . (default = 'TE>TEshift')

**cfg.shifttype** = Shifting the data 'onesample' or the length of the 'predicttime' (default = 'predicttime')

**cfg.numpermutation** = nr of permutations in permutation test  
(default = 190100)

**cfg.permstatstype** = 'mean' to use the distribution of the mean differences and 'depsamplesT' or 'indepsamplesT' for distribution of the t-values. (default = 'indepsamplesT')

**cfg.tail** = 1 tail or 2 tailed test of significance (for the permutation tests) (default in TEsurrogatestats= 1)

**cfg.correctm** = correction method used for correction of the multiple comparison problem - False discovery rate 'FDR' or Bonferroni correction 'BONF' (default = 'FDR')

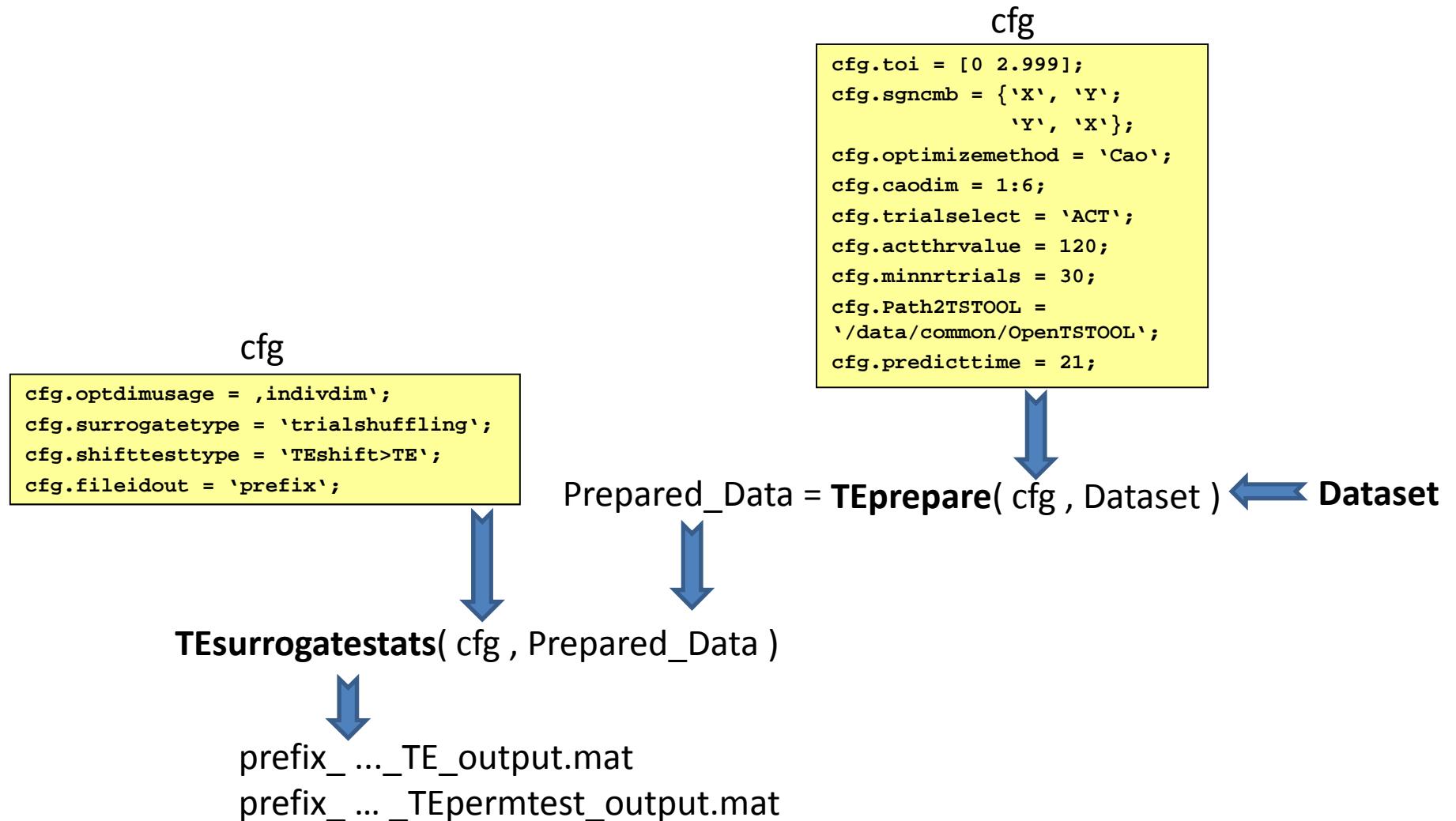
**cfg.fileidout** = string for the first part of the output filename.

# Example parameter set for TEsurrogatestats

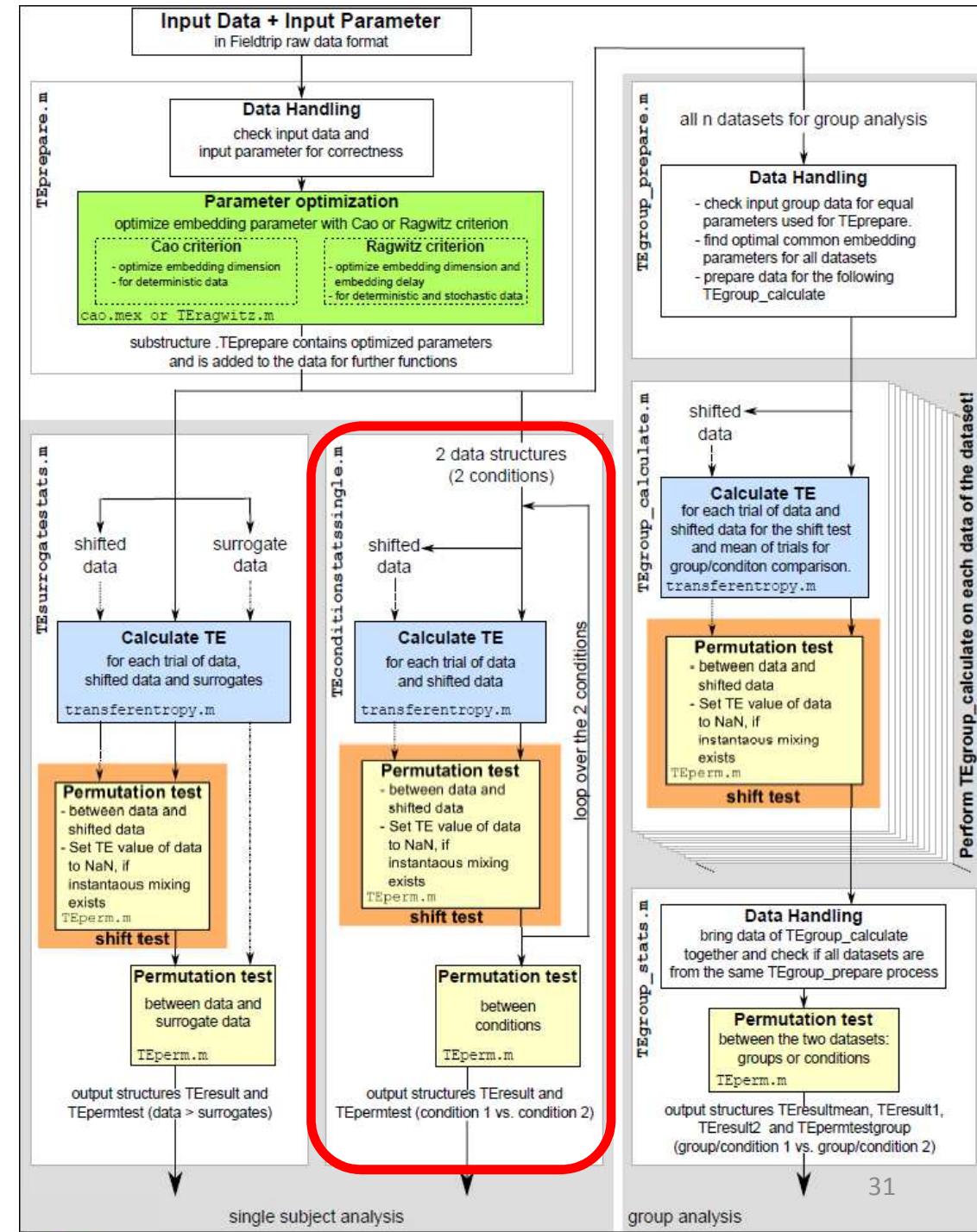
```
cfg.optdimusage = 'indivdim'  
cfg.surrogatetype = 'trialshuffling';  
cfg.shifttesttype = 'TEshift>TE';  
cfg.fileidout = 'your_choice';
```

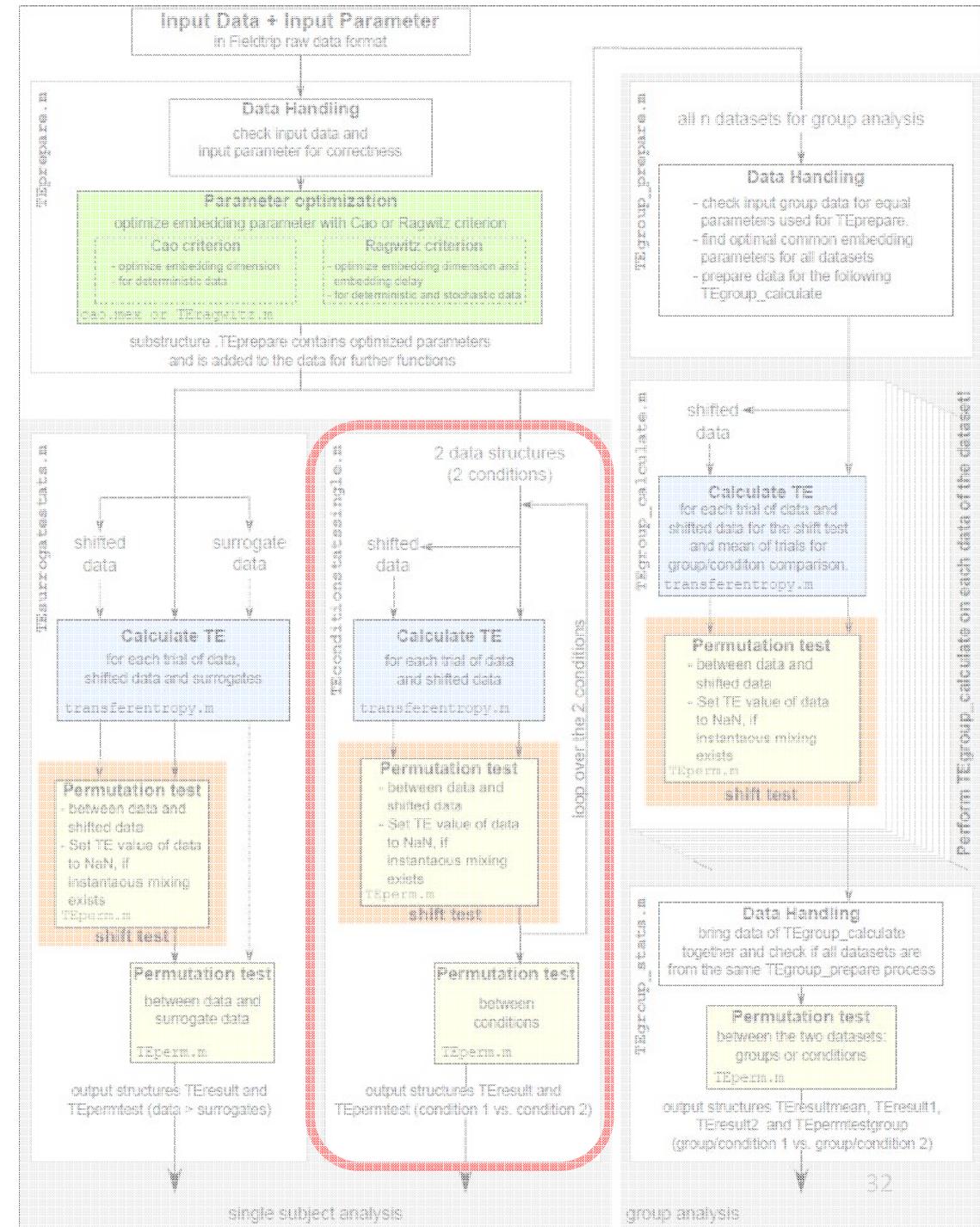
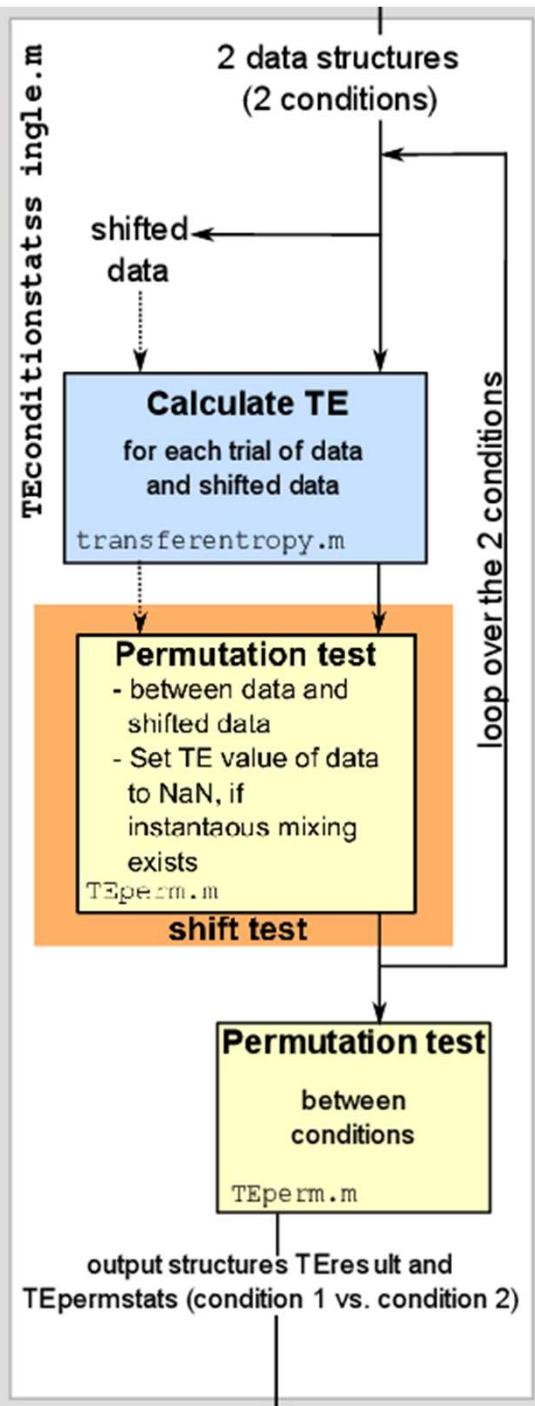
```
TESurrogatestats(cfg, Data_prepared)
```

# Example 1 - TEsurrogatestats



# Overview TRENTOOL

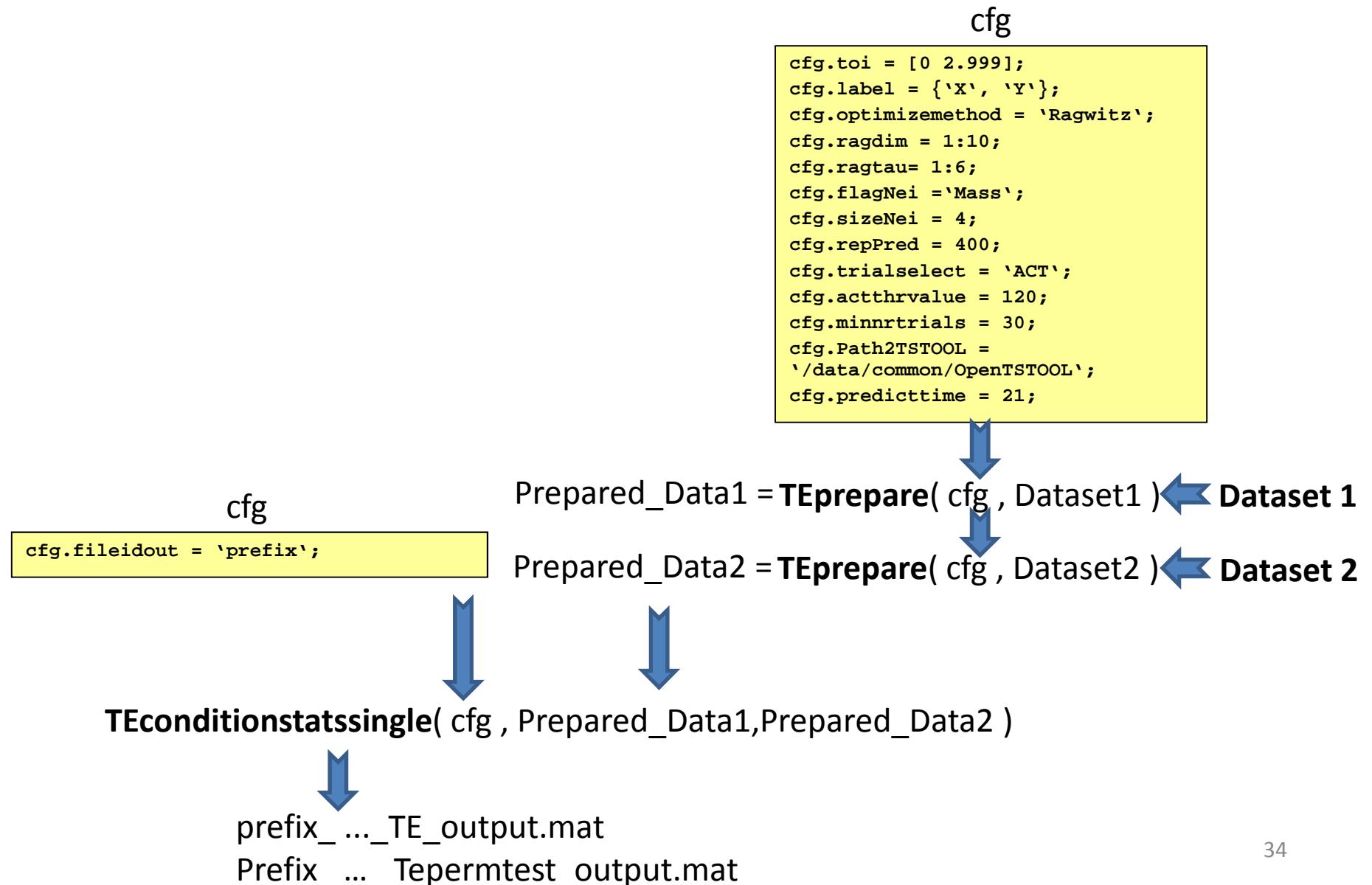




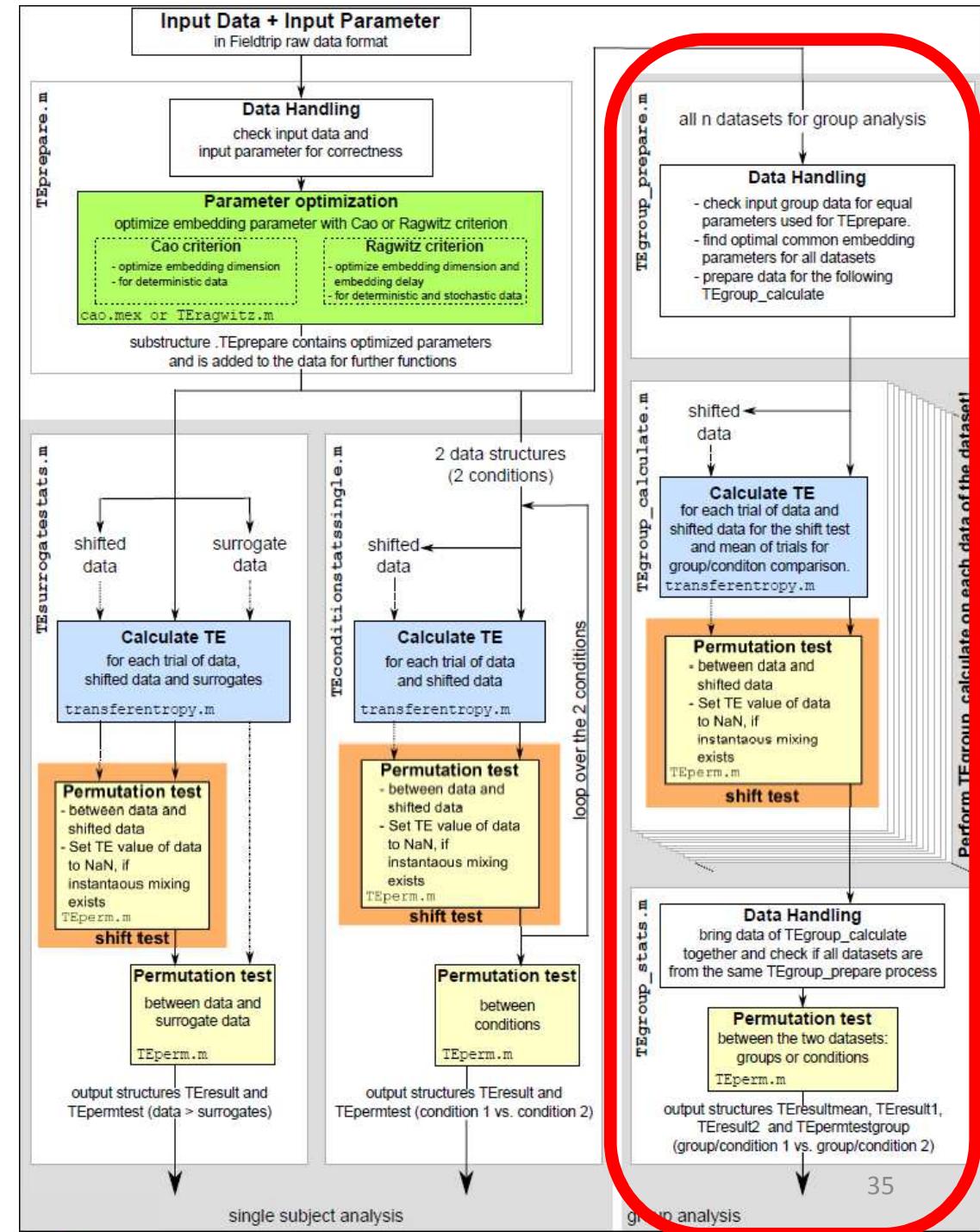
# All parameters for TEconditionstatssingle

<b>cfg.dim</b>	= Scalar value for embedding dimension. If not specified, the optimal dimension(s) found in TEprepare will be used, which is the recommended option!	<b>cfg.numpermutation</b> = nr of permutations in permutation test (default = 190100)
<b>cfg.tau</b>	= embedding delay in units of act ( $x^*act$ ). If not specified (recommended option), the tau is used as followed:  In case of optimizemethod in TEprepare: 'ragwitz' = optimal tau found via ragwitz critrion 'cao' = cfg.tau given by user in TEprepare	<b>cfg.permstatstype</b> = 'mean' to use the distribution of the mean differences and 'depsamplesT' or 'indepsamplesT' for distribution of the t-values. (default = 'mean')
<b>cfg.alpha</b>	= significance level for statisical shift test, permutation test and correction for multiple comparison (default = 0.05)	<b>cfg.tail</b> = 1 tail or 2 tailed test of significance (for the permutation tests) (default = 2)
<b>cfg.shifttest</b>	= perform shift test to identify instantaneous mixing between the signal pairs. Values: 'yes' or 'no' (default = 'yes')  This shift test is important for EEG and MEG data, because linear mixing is always present in the data. In case of instantaneous mixing transfer entropy should not be calculated for the affected channelpairs with the corresponding parameter sets, because it could result in false positive results. Hence the TE values for these cases will be set to NaN and the corresponding p-values of the permutation test to 1.	<b>cfg.correctm</b> = correction method used for correction of the multiple comparison problem - False discovery rate 'FDR' or Bonferroni correction 'BONF' (default = 'FDR') <b>cfg.fileidout</b> = string for the first part of the output filename.
<b>cfg.shifttesttype</b>	= The shift test can be calculated for the direction TE value of original data > TE values of shifted data (value = 'TE>TEshift') or for the other direction (value = 'TEshift>TE'). In this case the alpha is set to 0.1. (default = 'TE>TEshift')	
<b>cfg.shifttype</b>	= Shifting the data 'onesample' or the length of the 'predicttime' (default = 'predicttime')	

## Example 2 - TEconditionstatssingle



# Overview TRENTOOL



# Overview TRENTOOL

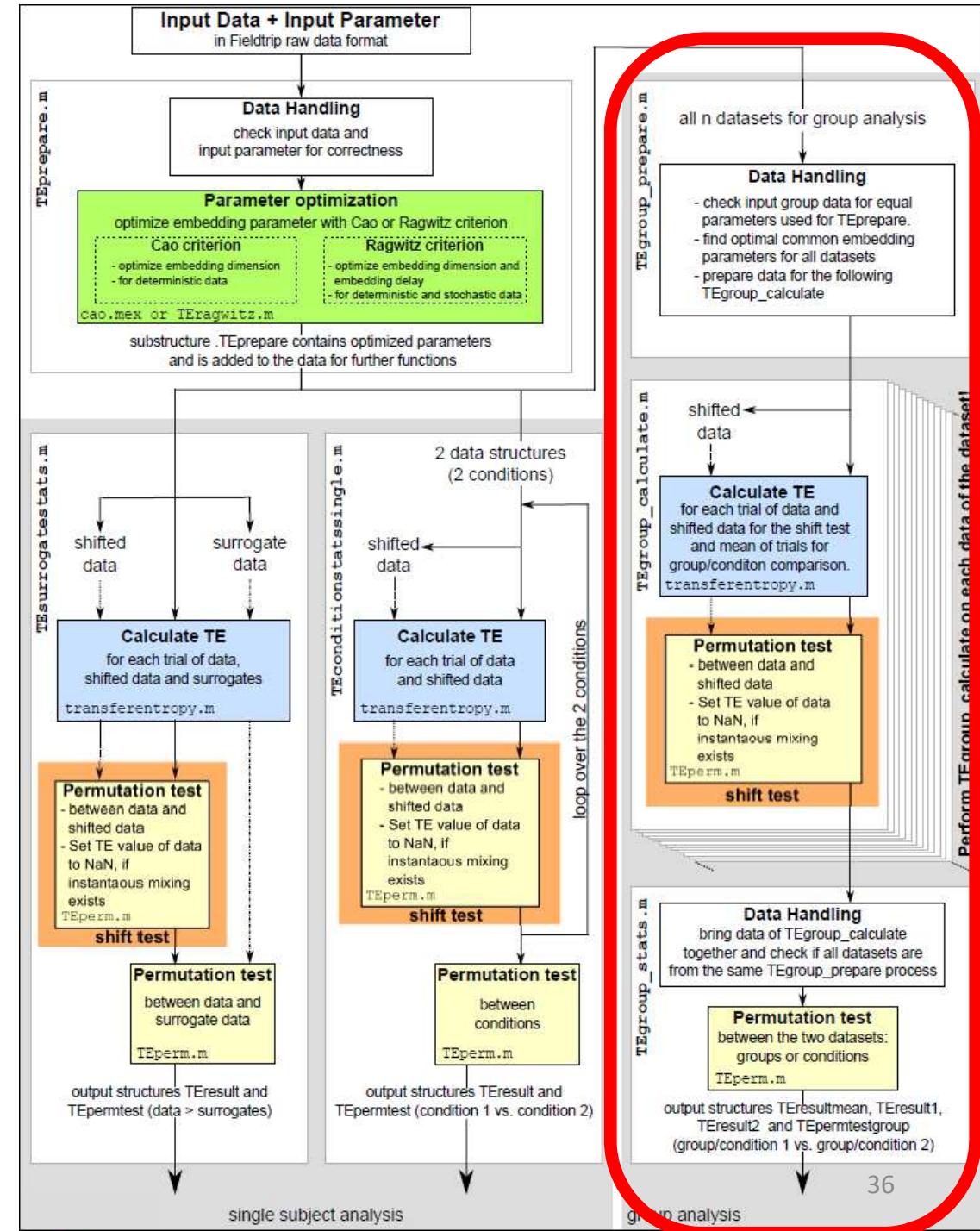
**Important:**

**Data\_prepared =  
TEprepare(cfg,Data)**

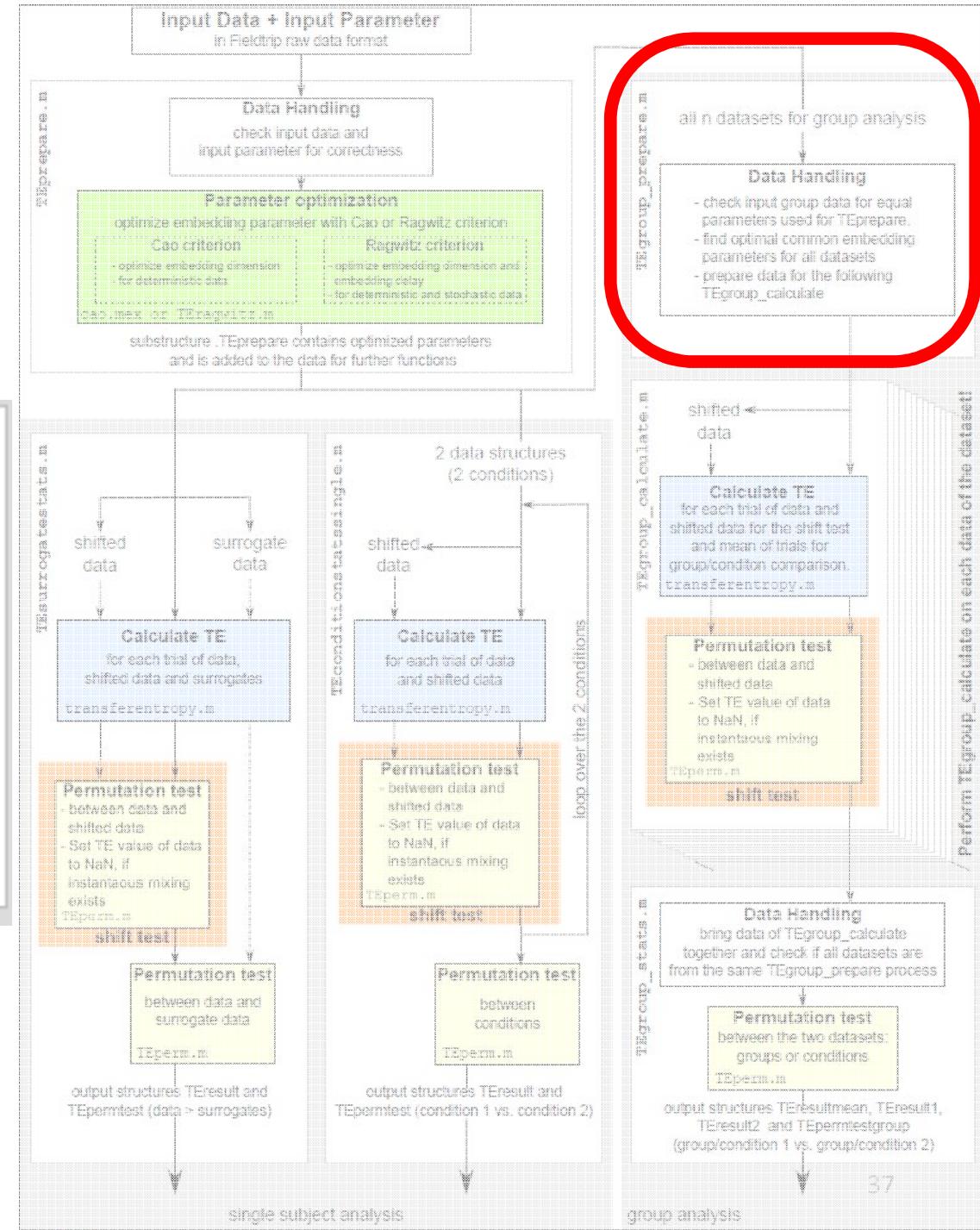
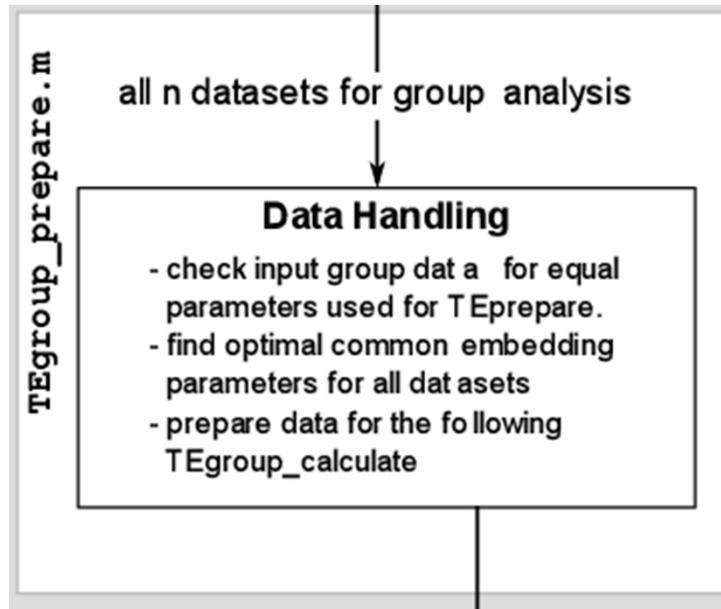
for EACH dataset with identical  
parameters!!!

&

SAVE EACH Prepared\_Data  
MANUALLY !!!



# Overview TRENTOOL



# All parameters for TEgroup\_prepare

**cfg.dim** = Number of embedding dimensions. If not specified, the maximum of the optimal dimensions found in Teprepare will be used, which is the recommended option!

**cfg.tau** = embedding delay in units of act ( $x^{\text{act}}$ ). If not specified (recommended option), the tau is used as followed:

In case of optimizemethod in TEprepare:

'ragwitz' = optimal tau found via ragwitz

critrion

'cao' = cfg.tau given by user in TEprepare

**cfg.shifttest** = perform shift test to identify instantaneous mixing between the signal pairs. Values: 'yes' or 'no'  
(default = 'yes')

This shift test is important for EEG and MEG data, because linear mixing is always present in the data.

In case of instantaneous mixing transfer entropy should not be calculated for the affected channelpairs with the corresponding parameter sets, because it could result in false positive results.

Hence the TE values for these cases will be set to NaN and the corresponding p-values of the permutation test to 1. Alpha is set to 0.05.

**cfg.shifttesttype** = The shift test can be calculated for the direction TE value of original data > TE values of shifted data (value = 'TE>TEshift') or for the other direction (value = 'TEshift>TE'). In this case the alpha is set to 0.1. (default = 'TE>TEshift')

**cfg.shifttype** = Shifting the data 'onesample' or the length of the 'predicttime' (default = 'predicttime')

# Run TEgroup\_prepare over all datasets

minimum parameter structure:

```
cfg = [ ];
```

```
TEgroup_prepare(cfg,FilesCell)
```

FilesCell is a MATLAB Cell containing all filenames of the already prepared Data sets for the group analyses

# TEgroup\_prepare

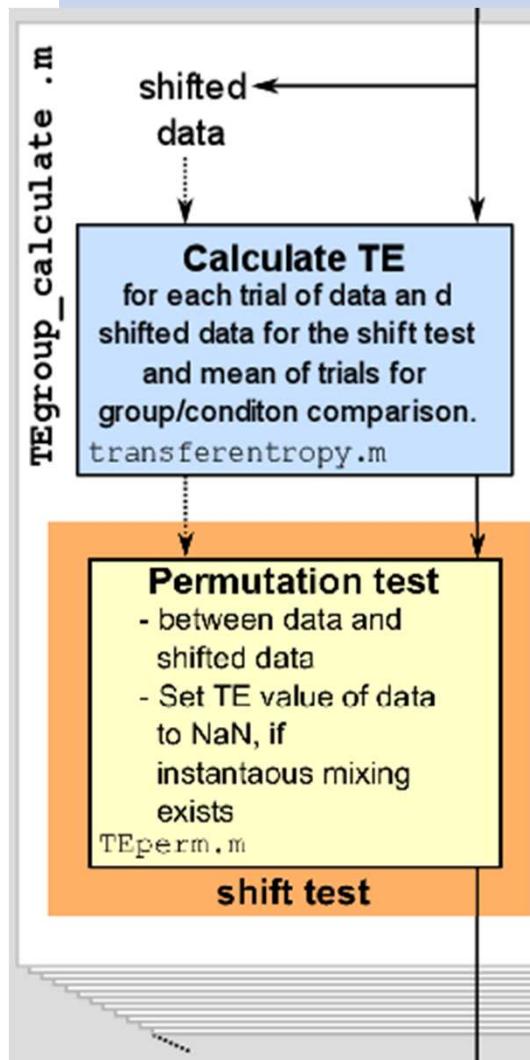
**Output :**

New version of each dataset:

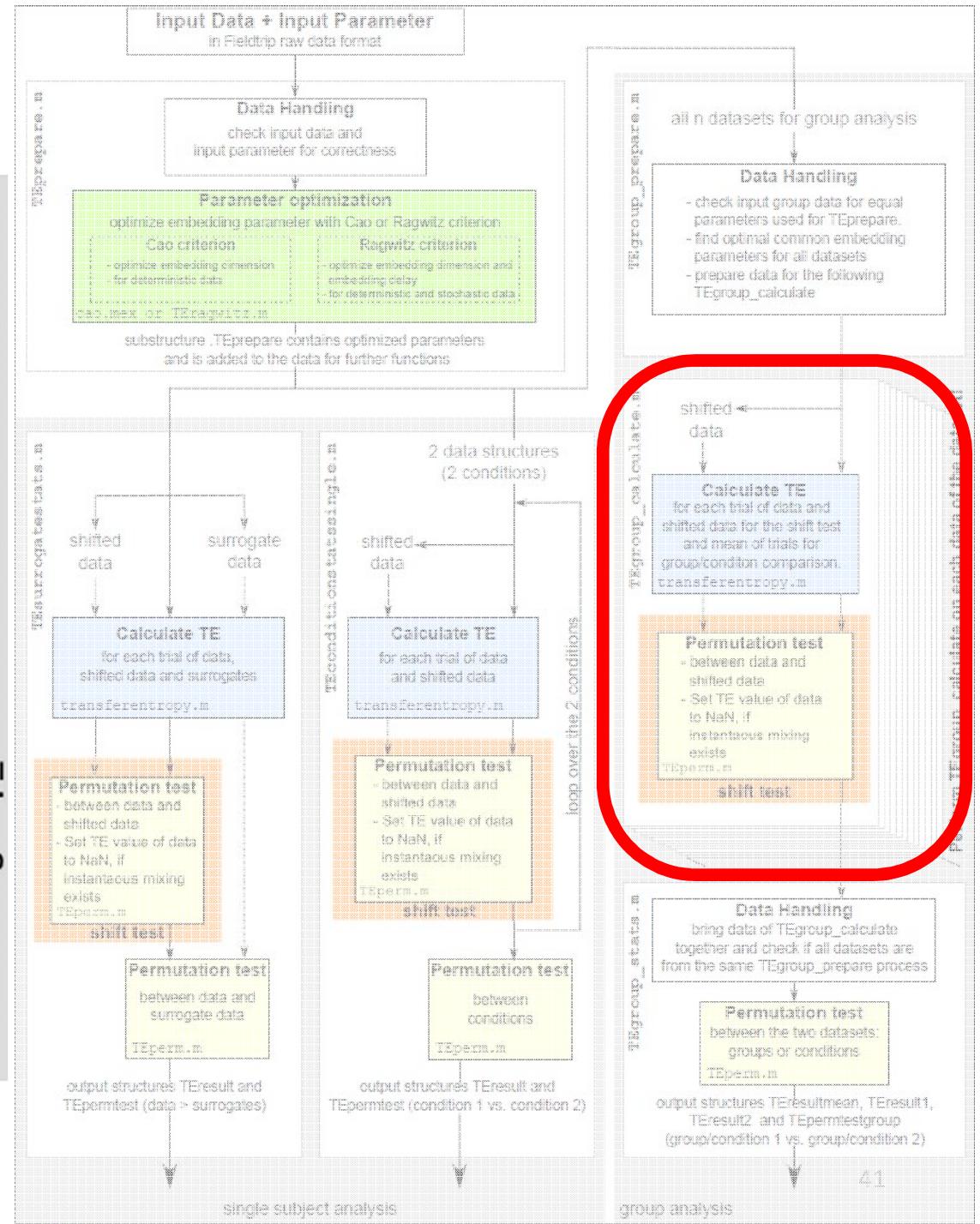
***filename\_for\_TEgroup\_calculate.mat***

containing an additional substructure .TEgroupprepare  
that is needed for the further functions.

# Overview



/ Perform TEgroup\_calculate on each data of the dataset!



## Run TEgroup\_calculate

for all datasets seperately  
(possibility to pseudo-parallelize):

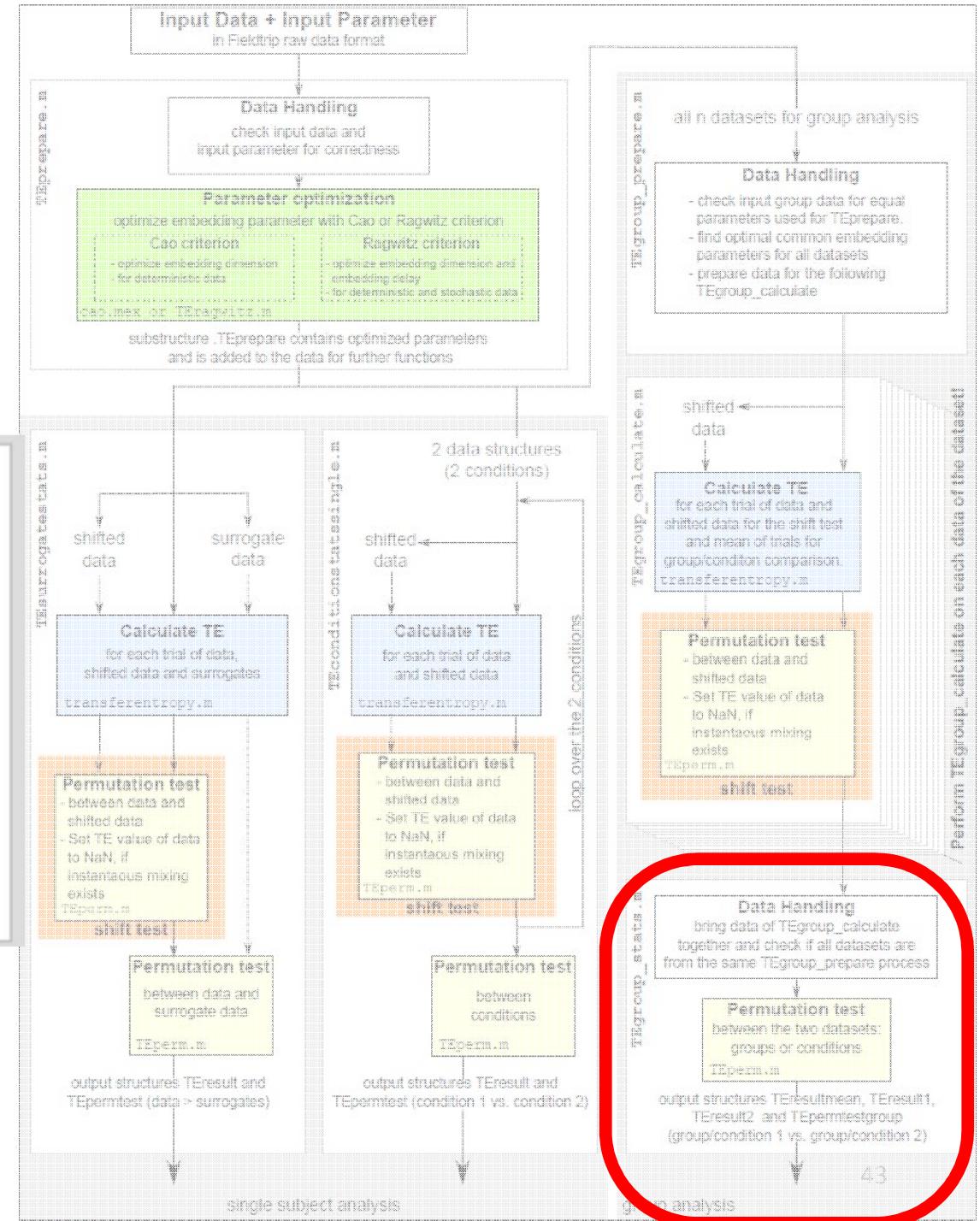
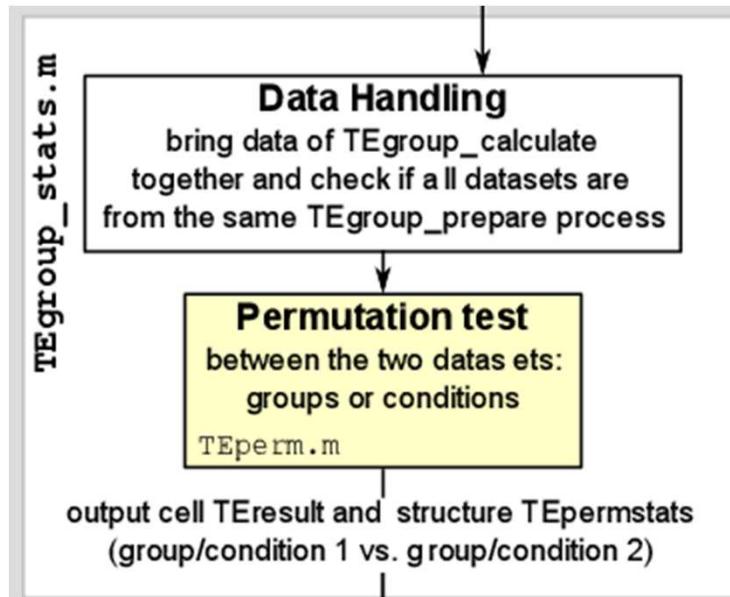
**TEgroup\_calculate(filename)**

Output:

*filename\_TE\_output.mat*

# Overview

# TRENTOOL



# All parameters for TEgroup\_stats

**cfg.design** = matrix containing a row with subject number and a row with independent variable representing the order of the data input.  
example:  
datasets: 1 2 3 4 5 1 2 3 4 5  
conditions: 1 1 1 1 1 2 2 2 2 2

**cfg.uval** = row in cfg.design which contains the dataset number (in the example: 1)

**cfg.ival** = row in cfg.design which contains the independent variable (in the example: 2)

**cfg.alpha** = significance level for statistical shift test, permutation test and correction for multiple comparison (default = 0.05)

**cfg.numpermutation** = nr of permutations in permutation test (default = 190100)

**cfg.permstatstype** = 'mean' to use the distribution of the mean differences and 'depsamplesT' or 'indepsamplesT' for distribution of the t-values. (default = 'mean')

**cfg.tail** = '1' tail or '2' tailed test of significance (for the permutation tests) (default = 2)

**cfg.correctm** = correction method used for correction of the multiple comparison problem - False discovery rate 'FDR' or Bonferroni correction 'BONF' (default = 'FDR')

**cfg.fileidout** = string for the first part of the output filename.

# Example parameter set for TEgroup\_stats

```
cfg.design =  
[1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,1,2,3,4,5,6,7,8,9,10,11,12,13,14,15;  
 1,1,1,1,1,1,1,1,1 ,1 ,1 ,1 ,1 ,1 ,1 ,1 ,2,2,2,2,2,2,2,2,2,2,2,2,2,2,2];  
cfg.uvar = 1;  
cfg.ivar = 2;  
cfg.permstatstype = 'depsamplesT';  
cfg.tail = 2;  
cfg.fileidout = 'your_choice';
```

Specify the order of files in the FilesCell  
One row codes the subjects (uvar)  
One row codes the conditions (ivar)

# Run TEgroup\_stats over all datasets

**TEgroup\_stats(cfg,FilesCell)**

Output:

*prefix\_ ... \_*

TEpermtestgroup\_output.mat

*prefix\_ ... \_ TE\_output.mat*

# Example 3 - Group analyses

Prepared\_Data1 = **TEprepare**( cfg , Dataset1 ) ← Dataset 1

Prepared\_Data2 = **TEprepare**( cfg , Dataset2 ) ← Dataset 2

...

...

...

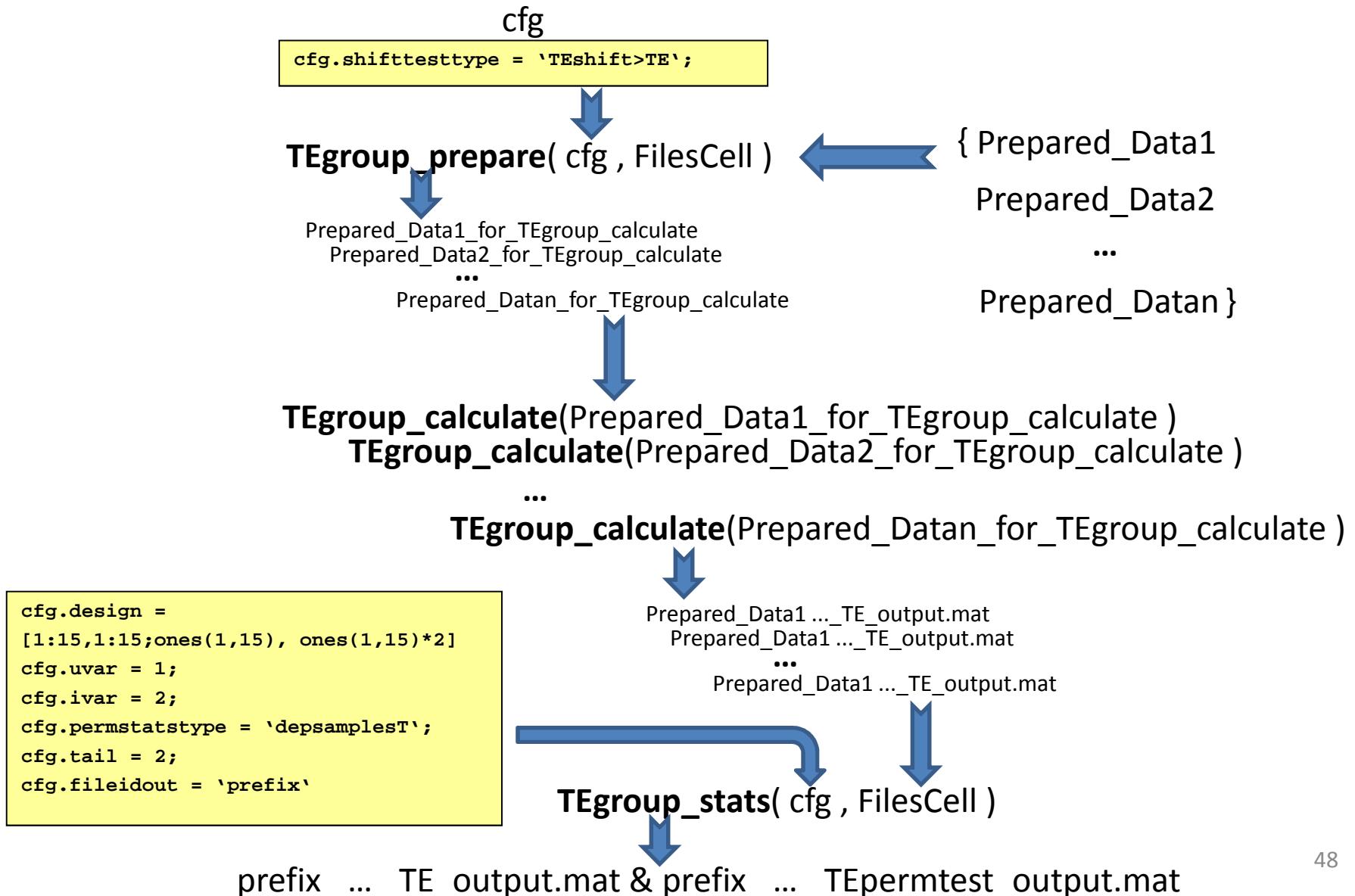
Prepared\_Datan = **TEprepare**( cfg , Datasetn ) ← Dataset n



cfg

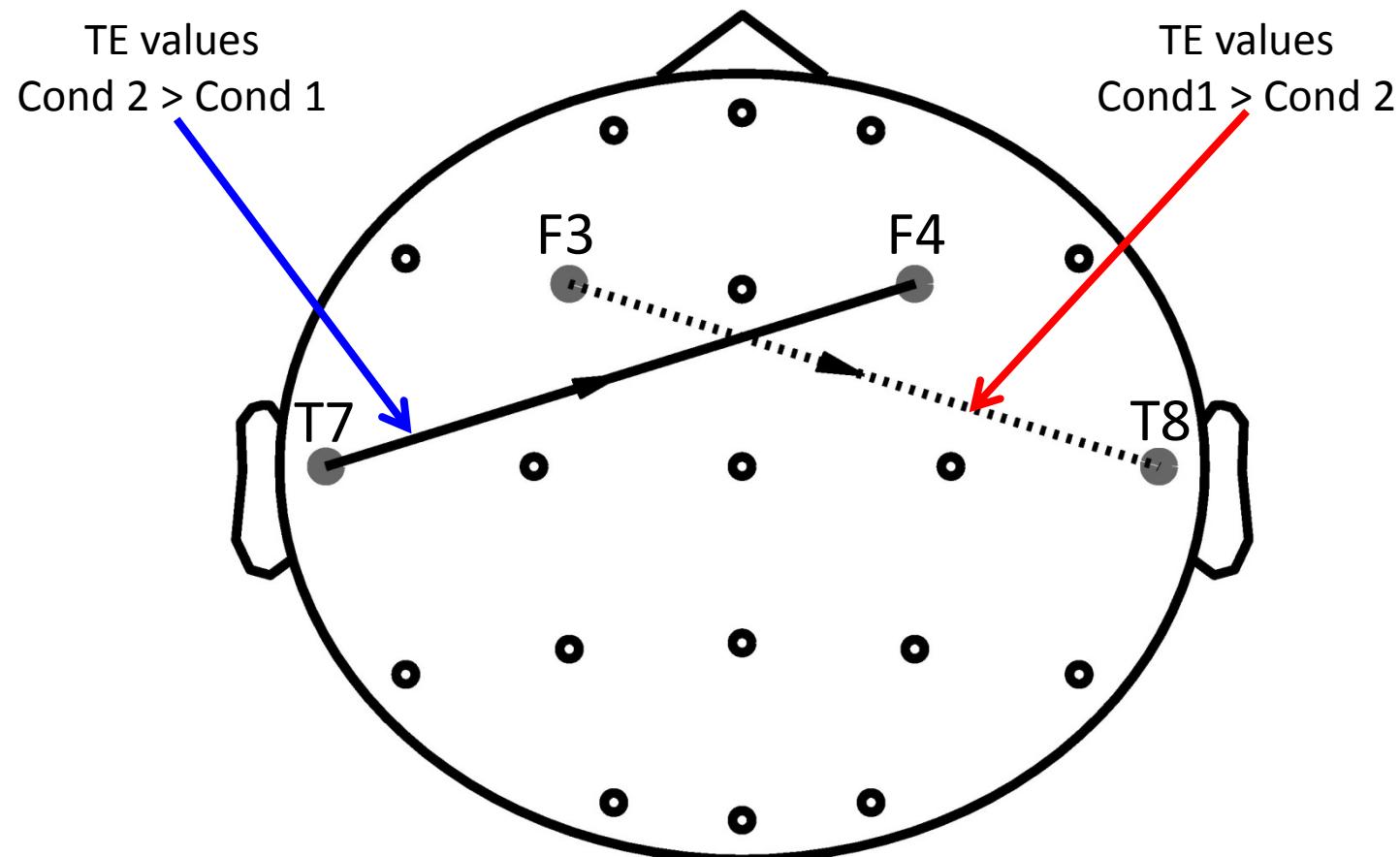
```
cfg.toi = [0 2.999];
cfg.labels = {'X', 'Y'};
cfg.optimizemethod = 'Cao';
cfg.caodim = 1:6;
cfg.caokth_neighbors = 3;
cfg.trialselect = 'ACT';
cfg.actthrvalue = 120;
cfg.minnrtrials = 30;
cfg.Path2TSTOOL =
'/data/common/OpenTSTOOL';
cfg.predicttime = 21;
```

# Example 3 - Group analyses



# TEplot2D

Example for the results of TEconditionstatssingle:



# Additional functions

- time reconstruction (detecting optimal prediction time  $u$ )

TEInteractionDelayReconstruction\_calculate

TEInteractionDelayReconstruction\_analyze

TEInteractionDelayReconstruction\_plotting

- Graph theoretical detection of cascade effects

## Some additional features we are working on:

- TRENTOOL for fMRI
  - Nifti to TRENTOOL data converter
  - Spatial embedding optimization
  - Usage for normal block or event related designs
- 3D plotting function
- ...

# Conclusion

- TRENTOOL
  - robustly detects the directed interactions.
  - is quite easy to use.
  - for neuroscientific applications: it works well on MEG data, single cell recording, intracranial recording, etc.
  - at the moment we are testing the usage of TRENTOOL on fMRI data

# Requirements

- MATLAB (<http://www.mathworks.de>)
  - statistic toolbox
  - (Parallel processing toolbox – only to make it much faster)
- Fieldtrip (<http://fieldtrip.fcdonders.nl>)
- OpenTSTOOL (<http://www.physik3.gwdg.de/tstool/>)
- Aplenty of calculating capacity

# Course program

- Day 1
  - TEprepare
    - Create cfg and script, run the function, inspect output
  - TEsurrogatestats
    - Create cfg and script , run the function, inspect output
  - Run prepared script for time reconstruction and let it run over night
- Day 2
  - Go through the results
  - Open questions

**Username = tuser**  
**password = 3Nt00l\_2**

### **Course folders:**

/data/home1/tuser/user01\_vnc28  
    .../user02\_vnc29  
    .../user03\_vnc30  
    .../user04\_vnc31  
    .../user05\_vnc32  
    .../user06\_vnc33  
    .../user07\_vnc34  
    .../user08\_vnc35  
    .../user09\_vnc36  
    .../user10\_vnc37  
    .../user11\_vnc38

### **Each user folder contains:**

- Toolboxes**
  - TRENTOOL2**
  - OpenTSTOOL**
  - Fieldtrip**
- Data**
  - data\_save**
- Prepared scripts (for time reconstruction)**

Example script:

```
addpath('~/user01_vnc28/toolboxes/TRENTOOL2')
addpath('~/user01_vnc28/toolboxes/fieldtrip')
ft_defaults

load('~/user01_vnc28/data/Lorenz_bidirectional_data.mat')

% TEprepare
cfg = [];
cfg.Path2TSTOOL =
'~/user01_vnc28/toolboxes/TRENTOOL2';
cfg.toi = [0.001 3];
cfg.channel = data.label;
cfg.predicttime_u = 46;
cfg.actthrvalue = 50;
cfg.minnrtrials = 12;
cfg.optimizemethod = 'ragwitz';
cfg.ragdim = 2:8;
cfg.ragtaurange = [0.5 1];
cfg.ragtausteps = 15;
cfg.repPred = 1000;
cfg.flagNei = 'Mass';
cfg.sizeNei = 4;

Prepared_Data = TEprepare(cfg,Data)
save('~/user01_vnc28/data/data_save/Prepared_Data.mat','Prepared_Data')

% TEsurrogatestats
cfg = [];
cfg.optdimusage = 'individ';
cfg.tail = 1;
cfg.numpermutation = 50000;
cfg.shifttesttype = 'TEshift>TE';
cfg.surrogatetype = 'trialshuffling';
cfg.fileidout =
'~/user01_vnc28/data/data_save/TEsur_output';

TEsurrogatestats(cfg,Prepared_Data)
```

Thank you for your attention and your upcoming  
directed interactions with the speaker!

Have fun with TRENTOOL!