

fMRI Preprocessing & Noise Modeling

An SPM Tutorial

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- Download example code and data presented in this talk:
 - <http://www.tnu-zurich.org/team/lars-kasper/>
 - Section: Talk and Lecture Materials
- SPM12 (Statistical Parametric Mapping)
 - <http://www.fil.ion.ucl.ac.uk/spm/software/spm12/>
 - developed by the Functional Imaging Lab, UCL, London
- TAPAS PhysIO Toolbox (SPM or Matlab standalone)
 - <http://www.translationalneuromodeling.org/tapas/>
 - Documentation & Example Data (Philips/Siemens/GE):
 - <http://www.translationalneuromodeling.org/software/documentation/>
 - <http://www.translationalneuromodeling.org/software/tapas-data/>

- Setting up a Preprocessing Pipeline in SPM:
The Batch Editor
- Monitoring and Comparing Pipelines:
Quality Assurance
- Multi-subject Pipelines:
Automatisation
- Integrating Own Code and Toolboxes:
Physiological Noise Modeling and Evaluation

Setup Pipeline
DEMO 1

Quality Monitoring
DEMO 2

Multi-Subject
DEMO 3

Noise Modeling
DEMO 4

The SPM GUI

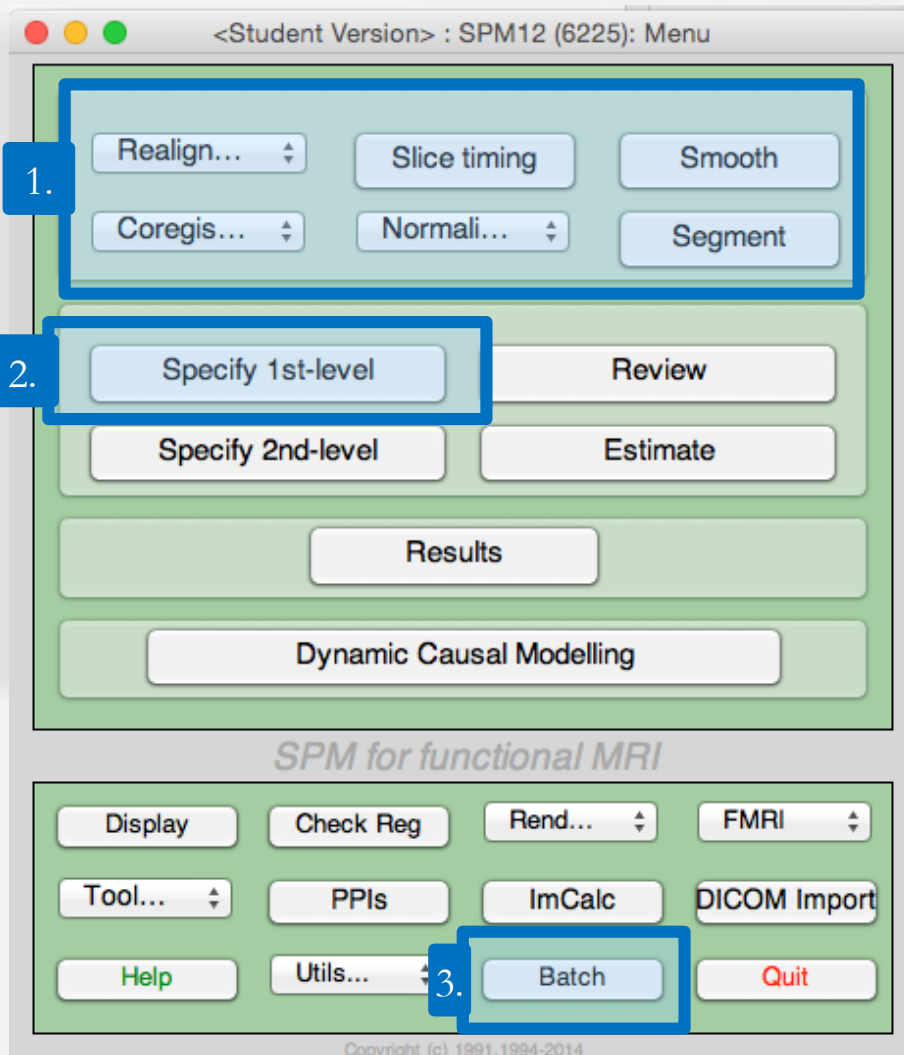


Setup Pipeline

Quality Monitoring

Multi-Subject

Noise Modeling



■ Preprocessing

- Realignment
- Slice-Timing Correction
- Co-Registration
- Unified Segmentation & Normalisation
- Smoothing...

■ Noise Modeling

- Physiological Confound Regressors

■ The Batch Editor

The Batch Editor in SPM



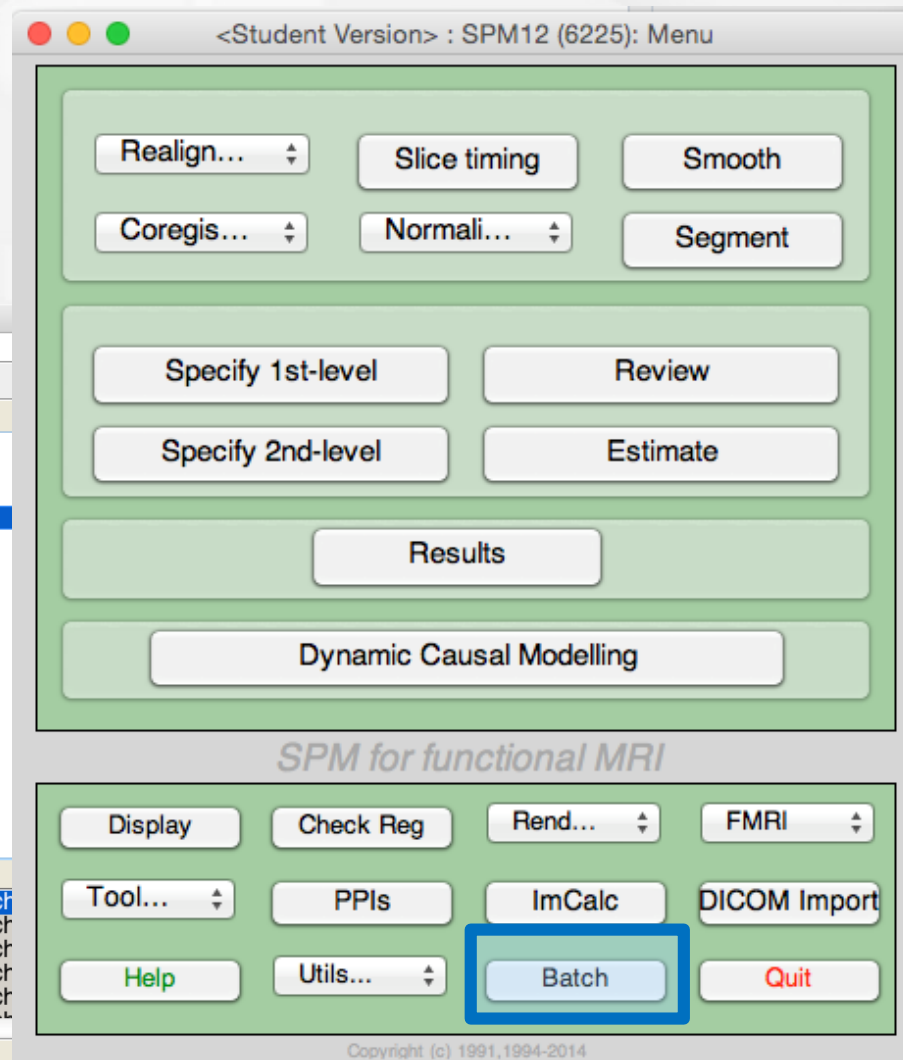
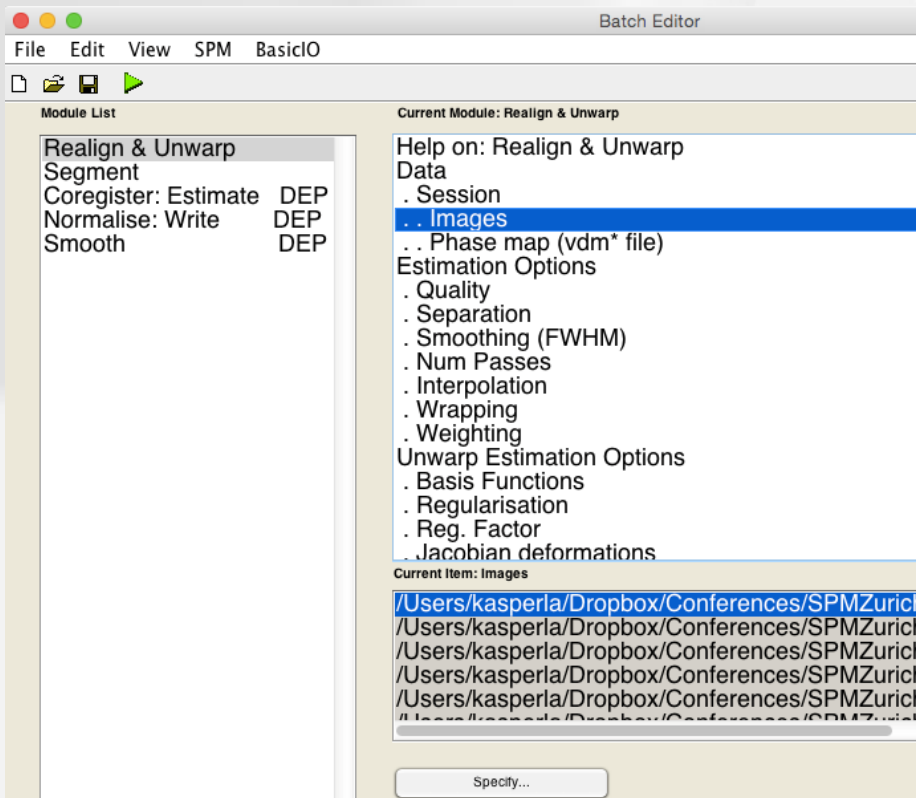
Setup Pipeline

Quality Monitoring

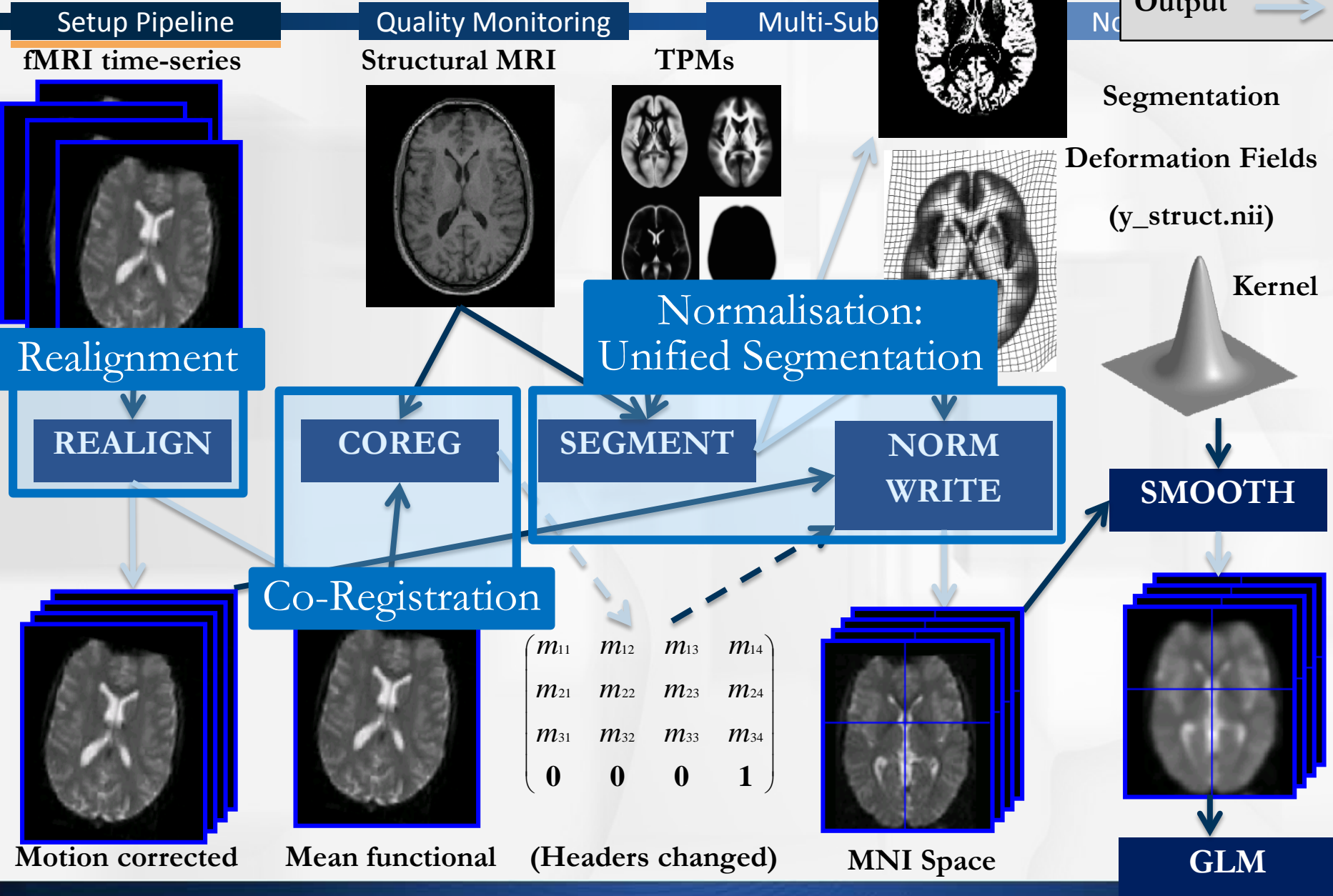
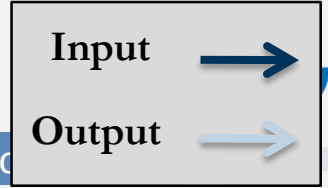
Multi-Subject

Noise Modeling

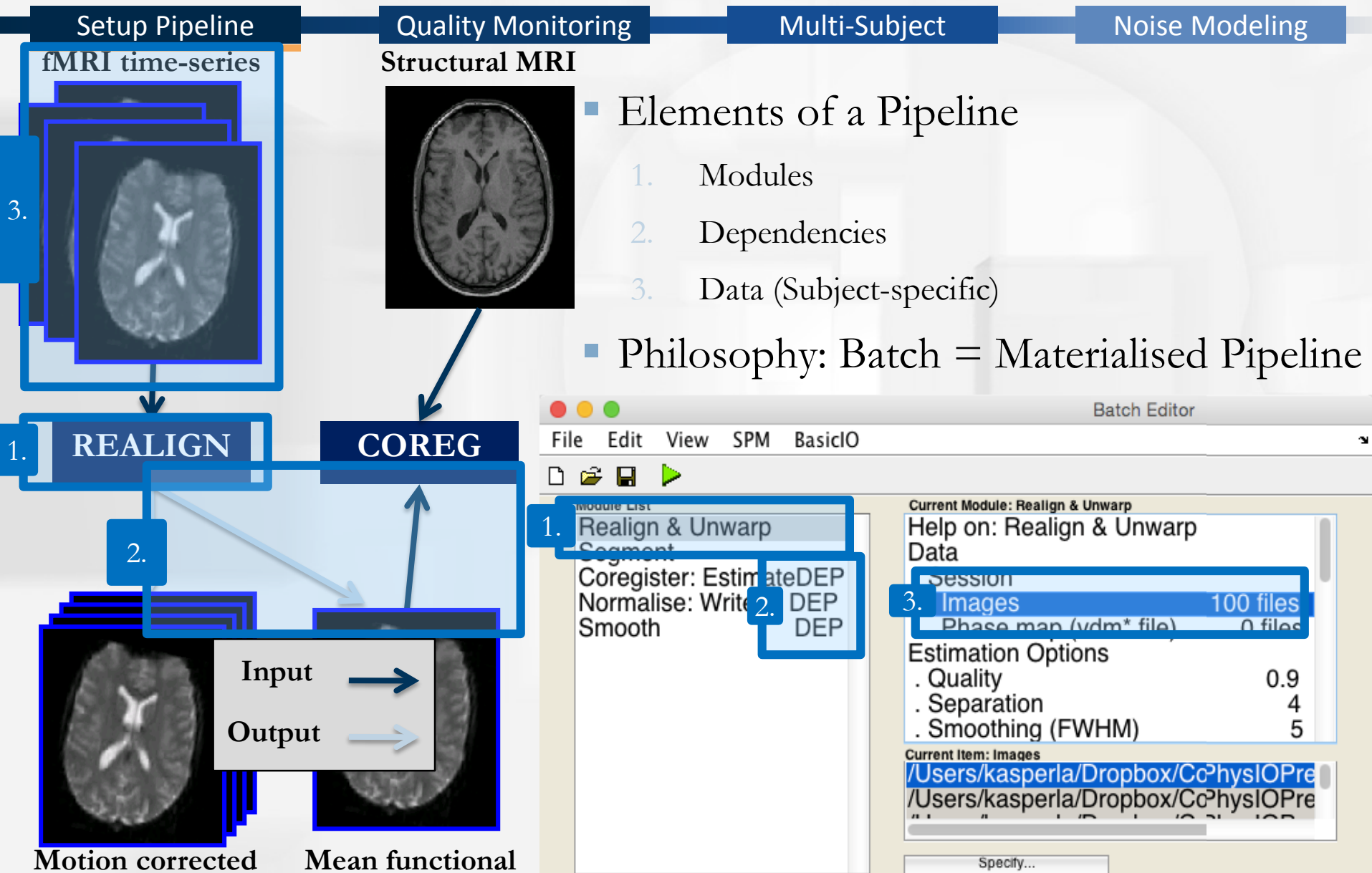
- Developer: V. Glauche
- Command line usage:
`spm_jobman`



SPM Preprocessing Pipeline



From Pipelines to Batches



- Setting up the data for all demos & checking it
 - Source: Social Learning (A. Diaconescu^[1], TNU Zurich, Philips 3T)
- After download:
 1. Unzip archive `examples_physio_short.zip`
 2. Open Matlab, run `code/init_reset_example.m`
 - subject01-folder created, Batches filled with right data (filenames and path) already
 3. Inspect raw functional *and* structural data carefully
 - Required for each subject
 - Preferred viewer: SPM CheckReg

[1] Diaconescu (2014), PLoS CB

Demo 0: CheckReg-Magic

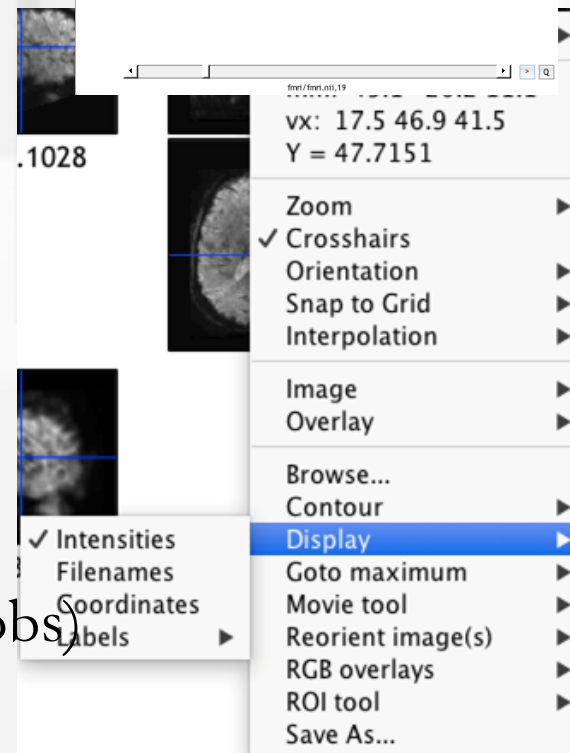
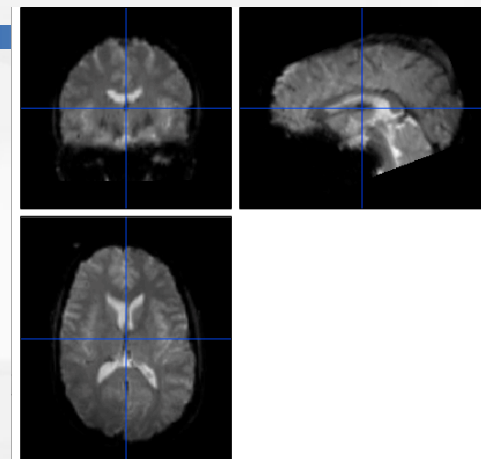


Setup Pipeline

Quality Monitoring

Multi-Subject

- Recommended Plotting: Check Reg
 - via Batch Editor: SPM => Util => Check Reg.
 - Matlab command line:
`spm_check_registration('img1.nii',
'img2.nii', ...)`
 - 4D NIFTI files:
 - show individual image tiles:
`spm_check_registration('fmri.nii,1',
'fmri.nii,2', ...)`
 - show movie (**NEW!** SPM12):
`spm_check_registration('fmri.nii')`
 - **Right click** reveals amazing features (edges, anatomic labels, header info, contrast, add blobs)



- Simple preprocessing pipeline for fMRI (G. Ridgway)
 - Based on `spm12/batches/preproc_fmri_simplified.m`
- Run via:
 1. Load `subject01/batches/demo01_simple_batch_preproc/batch_spm_preproc_fmri_simplified.m` in Batch Editor
 - Either via GUI or

```
spm_jobman('initcfg');  
spm_jobman('interactive',...  
'batch_spm_preproc_fmri_simplified.m' );
```
 2. Run Batch (Press Play)
- Exercise: Draw Pipeline Diagram
(Modules/Dependencies/Data)

[1] Diaconescu (2014), PLoS CB

Demo 1 – GUI Batch Editor



Setup Pipeline

Quality Monitoring

Multi-Subject

Noise Modeling

Batch Editor

File Edit View SPM BasicIO

Module List

- Realign & Unwarp
- Segment
- Coregister: Estimate DEP
- Normalise: Write DEP
- Smooth DEP

Current Module: Realign & Unwarp

Help on: Realign & Unwarp

Data

- . Session
- . . Images 100 files
- . . Phase map (vdm* file) 0 files

Estimation Options

- . Quality 0.9
- . Separation 4
- . Smoothing (FWHM) 5
- . Num Passes
- . Interpolation Register to first
- . Wrapping 2nd Degree B-spline
- . Weighting No wrap
- . . 0 files

Unwarp Estimation Options

- . Basis Functions 12x12x*
- . Regularisation 1
- . Reg. Factor Medium
- . Jacobian deformations No

Current Item: Images

/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproce
/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproce
/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproce
/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproce
/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproce
/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproce

Specify...

Demo 1 - Dependencies



Setup Pipeline

Quality Monitoring

Multi-Subject

Noise Modeling

Batch Editor

File Edit View SPM BasicIO

Module List

- Realign & Unwarp Segment
- Coregister: Estimate DEP
- Normalise: Write DEP
- Smooth DEP

Current Module: Coregister: Estimate

Help on: Coregister: Estimate

Reference Image DEP Segment: Bias Corrected (1)

Source Image DEP Realign & Unwarp: Unwarped Mean Image

Other Images ... Realign & Unwarp: Unwarped Images (Sess 1)

Estimation Options

- . Objective Function Normalised Mutual Information
- . Separation [4 2]
- . Tolerances 1x12 double
- . Histogram Smoothing [7 7]

Current Item: Source Image

Reference from

Realign & Unwarp: Unwarped Mean Image

Specify...

Dependency

Source Image

Demo 1 - Results



Setup Pipeline

Quality Monitoring

Multi-Subject

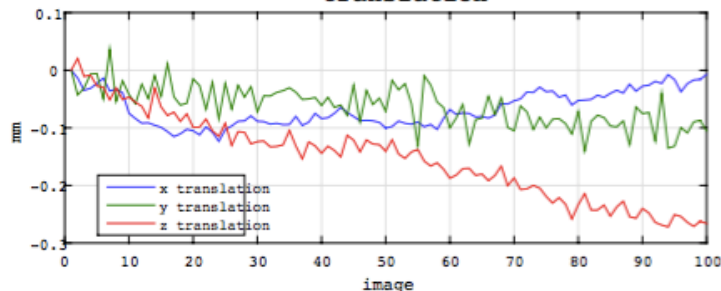
Noise Modeling

- Automatic status plots saved in `spm_<date>.ps`

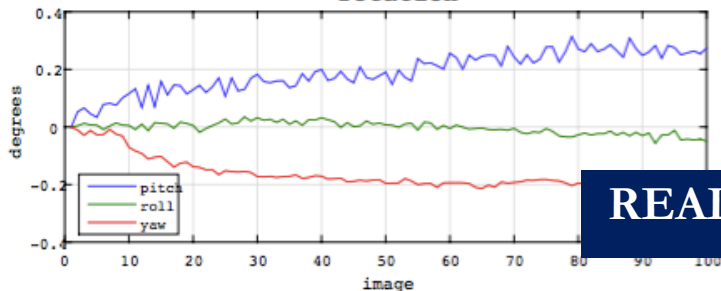
Image realignment

```
1 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
2 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
3 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
4 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
5 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
6 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
7 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
8 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
9 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
10 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
11 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
12 /Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreproces
..... etc
```

translation



rotation



REALIGN

Normalised Mutual Information Coregistration

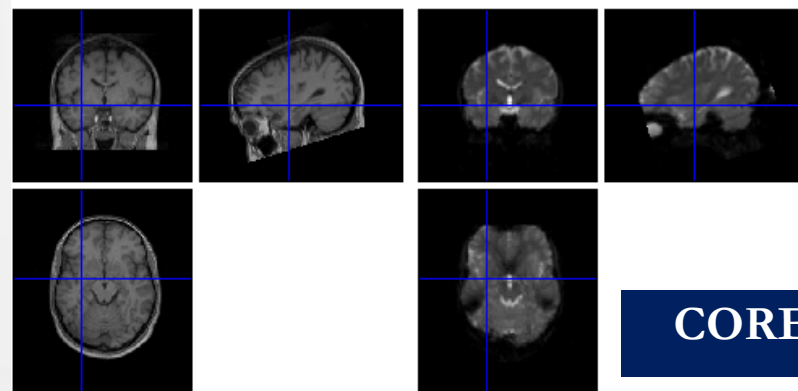
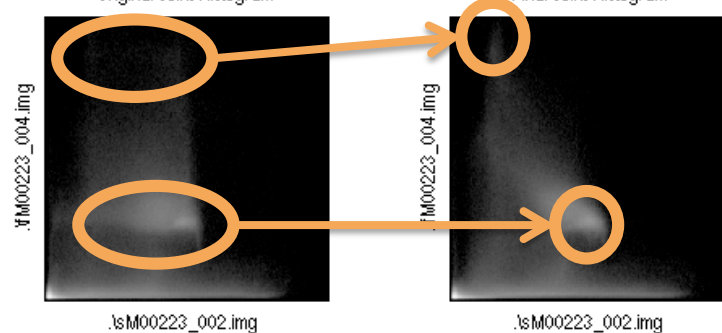
$$X1 = 3.000*X + 0.019*Y + 0.017*Z + 28.728$$

$$Y1 = -0.020*X + 3.000*Y + 0.043*Z + 32.827$$

$$Z1 = -0.006*X - 0.015*Y + 1.000*Z - 8.467$$

Original Joint Histogram

Final Joint Histogram



COREG

- Setting up a Preprocessing Pipeline in SPM:
The Batch Editor
- **Monitoring and Comparing Pipelines**
 - Performance Measures: Mean/SD/SNR/Diff Image
 - SPM Plotting Routines and Automatic Reporting
 - Spotting Failed Pipelines
 - Comparing Alternative Pipelines
- Multi-subject Pipelines
- Integrating Own Code and Toolboxes:
Physiological Noise Modeling and Evaluation

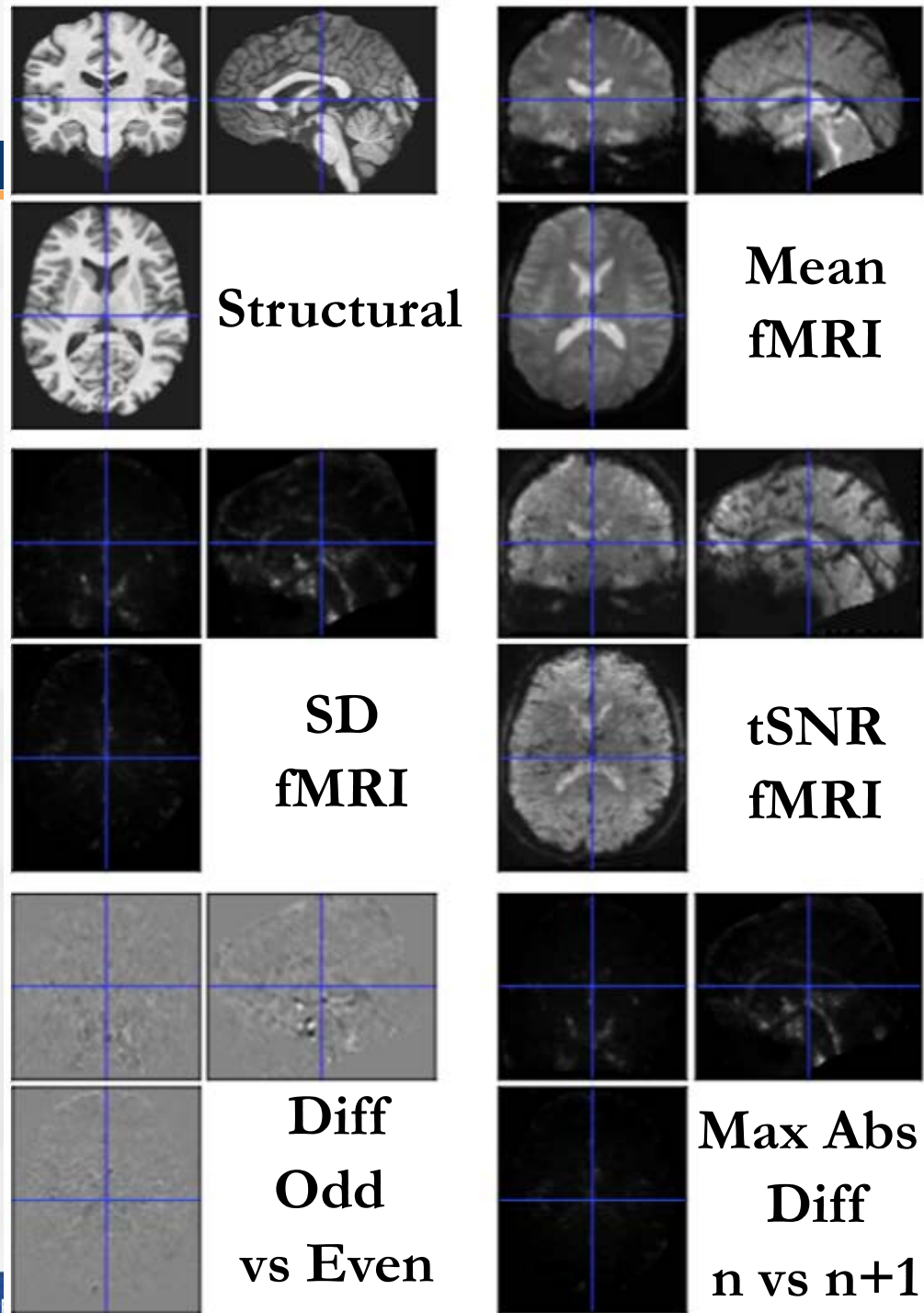
- Pipelines → Automatisatisation of Preprocessing
- When something goes wrong...how do we even notice?
 - Monitoring, but: cumbersome, when lots of data
 - Thus: Automate quality monitoring as well via pipelines
- Required: Suitable performance measures
 - Single image: visual inspection geometry/contrast/noise/SNR
 - structural image
 - Time series: Statistical Images (Mean, SD, tSNR, $\max(\text{abs}(\text{diff}))$)
 - functional images

Statistical Images

Setup Pipeline

Quality Monitoring

- Structural Image:
 - visual inspection geometry/contrast/noise/SNR
- Functional Time Series:
 - Mean \Rightarrow Artifact levels (localization)
 - SD \Rightarrow Fluctuation levels
 - tSNR = Mean/SD \Rightarrow sensitivity for BOLD signal changes
 - Diff = $\max(\text{abs}(\text{diff}))$ or odd – even \Rightarrow outlier detection, image noise



- Complete preprocessing & monitoring pipeline for fMRI
 - Based on `spm12/batches/preproc_fmri`
 - Includes `batch_report_quality.m` to visualize and save quality measures after each preprocessing step
- Run via
 1. Cleanup processed data of subject01: `init_reset_example(2)`
 2. Load `subject01/batches/demo02_simple_batch_preproc/batch_preproc_fmri_report_quality.m`
 3. Either via GUI or `spm_jobman('interactive', 'batch_preproc_fmri_report_quality.m')`
 - Alternative: Run batch directly from command line: `spm_jobman('run', 'batch_preproc_fmri_report_quality.m')`

Demo 2 - GUI



Setup Pipeline

Quality Monitoring

Multi-Subject

Noise Modeling

Batch Editor

File Edit View SPM BasicIO

Module List

| | |
|----------------------|-----|
| Named File Selector | |
| Named File Selector | |
| Realign & Unwarp | DEP |
| Segment | DEP |
| Get Pathnames | DEP |
| Image Calculator | DEP |
| Coregister: Estimate | DEP |
| Normalise: Write | DEP |
| Smooth | DEP |
| Normalise: Write | DEP |
| Make Directory | |
| Change Directory | DEP |
| Move/Delete Files | DEP |
| Run Batch Jobs | DEP |

Current Module: Run Batch Jobs

Help on: Run Batch Jobs

Job File(s) ...demo02_compare_batch_quality/batch_report_quality.m

Runs

- . Job Inputs
- . . Directory
- . . String
- . . NIfTI Images
- . . NIfTI Images
- . Job Inputs
- . . Directory
- . . String
- . . NIfTI Images
- . . NIfTI Images
- . Job Inputs
- . . Directory
- . . String
- . . NIfTI Images
- . . NIfTI Images
- . Job Inputs
- . . Directory

DEP Make Directory: Make Directory 'report_quality' 00_raw

DEP Move/Delete Files: Moved/Copied Files

DEP Named File Selector: Structural Image(1) - Files

DEP Make Directory: Make Directory 'report_quality' 01_realigned

DEP Realign & Unwarp: Unwarped Images (Sess 1)

DEP Named File Selector: Structural Image(1) - Files

DEP Make Directory: Make Directory 'report_quality' 02_coregistered

DEP Coregister: Estimate: Coregistered Images

DEP Named File Selector: Structural Image(1) - Files

DEP Make Directory: Make Directory 'report_quality'

Current Item: Job File(s)

/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreprocessing/exa

Specify...

Dependency

Demo 2 - Output



Setup Pipeline

Quality Monitoring

Multi-Subject

Noise Modeling

■ Output:

- `subject01/report_quality/report_quality.ps`
 - PostScript file with all output plots, generated from the following nifti image files
- `report_quality/00_raw/` => raw time series statistics
 - `mean.nii` => mean of time series (per pixel)
 - `sd.nii` => standard deviation (per pixel)
 - `snr.nii` => mean/sd (per pixel)
 - `diffOddEven.nii` => SumOddImages – SumEvenImages
 - `maxAbsDiff` => maximum delta image (vol n vs n+1)
- `subject01/01_realigned` => realigned time series stats
- ...
- `subject01/04_smoothed` => smoothed time series stats

- Example: Comparing temporal SNR throughout preprocessing
 - Temporal SNR per pixel in functional image time series after each preprocessing step
 - Command line code:

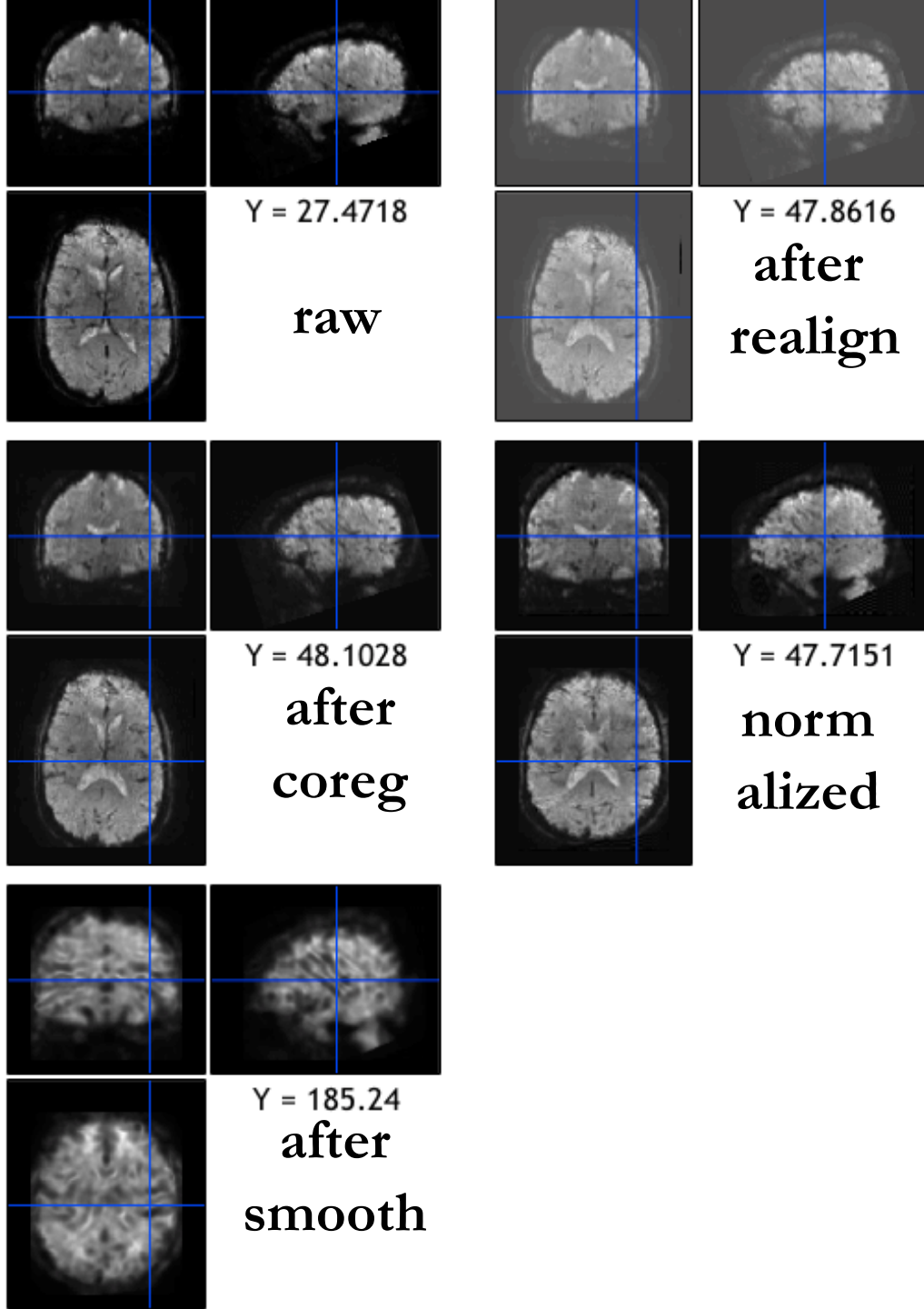
```
dirs = {'00_raw', '01_realigned',  
        '02_coregistered', '03_normalized',  
        '04_smoothed'}';  
files = 'snr.nii';  
fpFiles = strcat(dirs, '/', files);  
spm_check_registration(fpFiles{:})
```

Plotting Example

Setup Pipeline

Quality Monitoring

- Example: Temporal SNR per pixel in functional image time series after each preprocessing step
- Right click reveals amazing features
- Result: Increased SNR through realignment and smoothing



- Task: Spotting unusual quality report images
 - `run_demo02.../batch_run_artefact_subjects.m`
 - executes `code/create_artefact_subjects.m`
 - then: executes demo02-preprocessing batch for each subject
 - View for individual subjects: `report_quality.ps`
- What went wrong here?
 - subjectA1
 - subjectA2
 - subjectA3

Quiz...



Setup Pipeline

Quality Monitoring

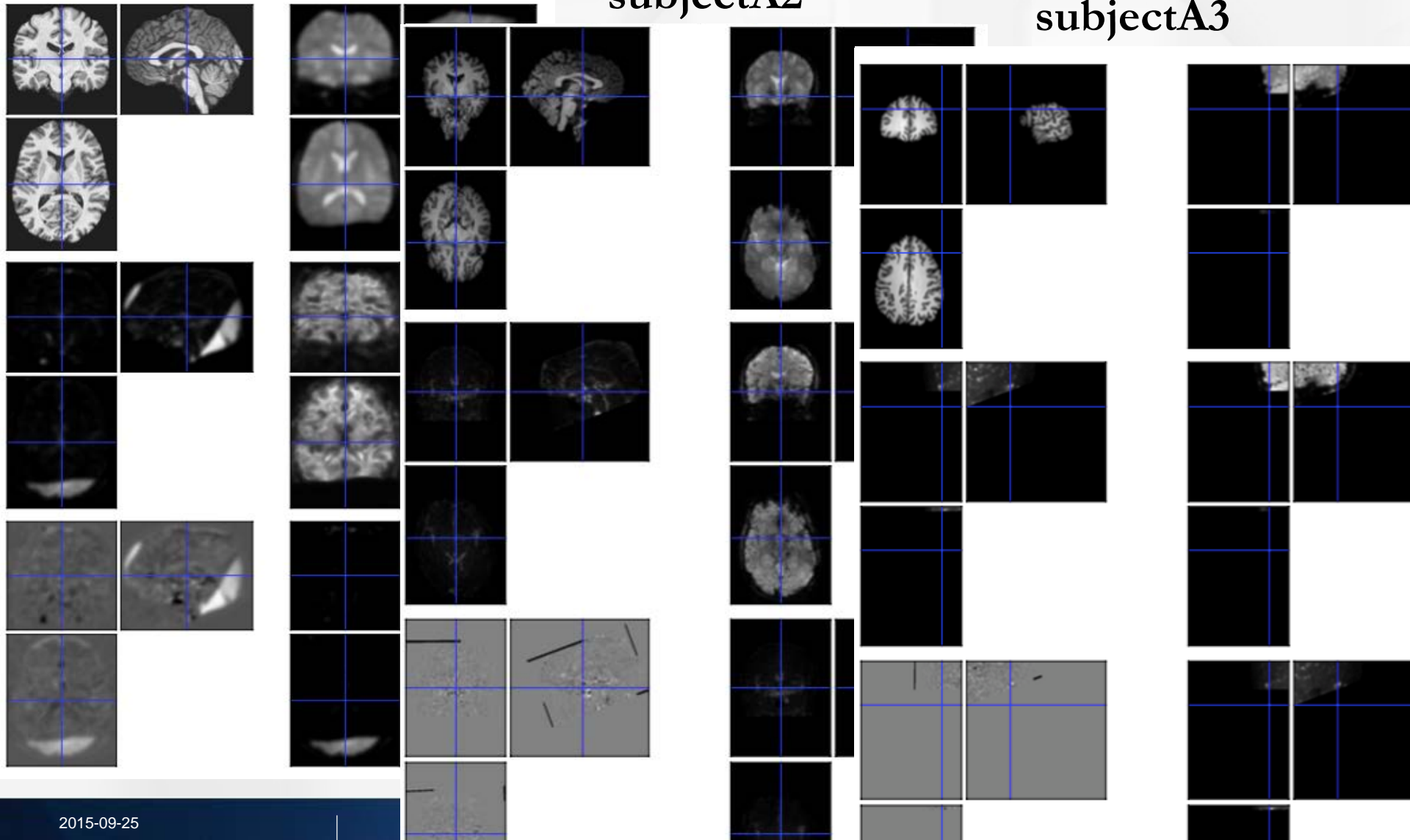
Multi-Subject

Noise Modeling

subjectA1

subjectA2

subjectA3



- Task: Spotting unusual quality report images
- What went wrong here?
 - subjectA1
 - realignment failed
 - one volume rotated (30 degrees around x-axis)
 - subjectA2
 - segmentation failed
 - structural mirrored compared to template
 - subjectA3
 - co-registration failed
 - functional image shifted by 10 cm (x, y, and z) compared to structural

- When running different preprocessing pipelines on the same subject:
 - Always start from raw nifti data (structural and functional)
 - Copy it to new processing folder, e.g.
 - `subject01/preproc_alternative/fmri/`
`/struct/`
 - Alter original batch file and save it under new name
 - Swapping preprocessing steps is not simple in the batch editor
 - Solution: save as .m-file and edit in Matlab
 - helper script (code folder):
`reorder_matlabbatch(fileBatch, indicesBefore, indicesAfter)`
 - Run different batch files via `run_job`

Comparing Pipelines



Setup Pipeline

Quality Monitoring

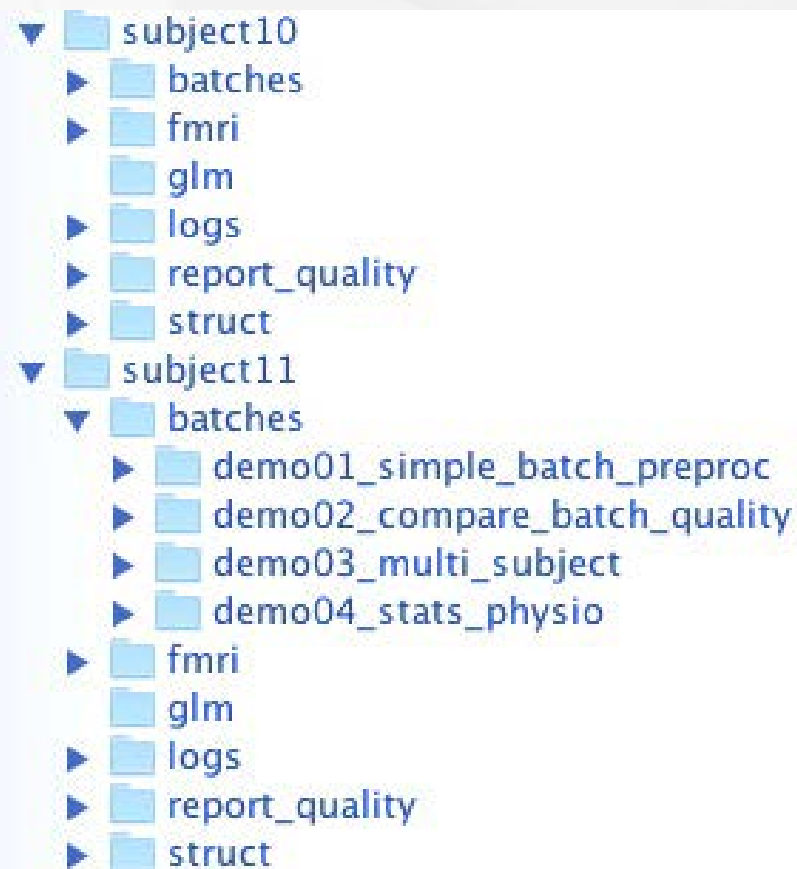
Multi-Subject

Noise Modeling

- Example: When to use Slice timing correction?
- Run:
 - `demo02_compare_batch_quality/batch_compare_preproc`
- this executes:
 - `batch_compare_template_stc_realign`
 - slice timing correction before realignment
 - `batch_compare_template_realign_stc`
 - realignment before slice timing
- Note: Proper Assessment only after statistical analysis possible

- Setting up a Preprocessing Pipeline in SPM:
The Batch Editor
- Monitoring and Comparing Pipelines
- **Multi-subject Pipelines**
 - Staying on Top: Organisation of Data
 - Looping Pipelines over Subjects' Data
 - Tips for Efficient Performance Monitoring
- Integrating Own Code and Toolboxes:
Physiological Noise Modeling and Evaluation

- One Study Folder
 - sub-folder for code, templates for subject-specific batches
 - modules and dependencies stay the same
- One Folder for each subject
 - therein: same sub-folder structure
 - reduces subject-specific data specification to changing the subject folder
 - batches/ => subject batch with data spec
 - fmri/ => functional data per session
 - struct/ => structural data



- glm/ => statistical analyses
- logs/ => behavioural response files, physiological recordings

Demo 3 – Example Loop Script



Setup Pipeline

Quality Monitoring

Multi-Subject

Noise Modeling

```
% List of open inputs
% Named File Selector: File Set - cfg_files
% Named File Selector: File Set - cfg_files
% Make Directory: Parent Directory - cfg_files

pathStudy      = '/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPrepr
dirSubjectArray = {
    'subject10'
    'subject11'
};
fileFunctional  = 'fmri/fmri.nii';
fileStructural  = 'struct/struct.nii';

nrun = 2; % enter the number of runs here
jobfile = {'/Users/kasperla/Dropbox/Conferences/SPMZurich15/PracticalsPhysIOPreprocessir
jobs = repmat(jobfile, 1, nrun);
inputs = cell(3, nrun);
for crun = 1:nrun
    pathSubject = fullfile(pathStudy, dirSubjectArray{crun});
    fullpathFunctional = fullfile(pathSubject, fileFunctional);
    fullpathStructural = fullfile(pathSubject, fileStructural);

    % functional file
    inputs{1, crun} = {fullpathFunctional}; % Named File Selector: File Set - cfg_files

    % structural file
    inputs{2, crun} = {fullpathStructural}; % Named File Selector: File Set - cfg_files

    % subject folder
    inputs{3, crun} = {pathSubject}; % Make Directory: Parent Directory - cfg_files
end
spm('defaults', 'FMRI');
spm_jobman('run', jobs, inputs{:});
```

- Performs Demo 2 (complete preproc + quality report) for multiple subjects (subject10 and subject11)
 - create subject folders: Run `code/create_multi_subject_data.m`
- Run via
 1. Load “job”:
`subject10/batches/demo03_multi_subject/batch_preproc_fmri_report_quality_job.m` - Inspect!
 2. Load & Run `batch_preproc_fmri_report_quality_gui.m`
 3. Alternative: Open in *Matlab* Editor, inspect construction of subject dependent file names, run in Matlab GUI:
`batch_preproc_fmri_report_quality_run.m`

- Setting up a Preprocessing Pipeline in SPM:
The Batch Editor
- Monitoring and Comparing Pipelines
- Multi-subject Pipelines
- **Integrating Own Code and Toolboxes:**
Physiological Noise Modeling and Evaluation
 - Executing Custom Matlab Code within the Pipeline
 - The TAPAS PhysIO Toolbox
 - Automatic (Noise) Modeling and Contrast Reporting

- Performs whole single-subject analysis except preprocessing
 - creates behavioral regressors (visual stimuli/button responses)
 - `multiple_conditions.mat`
 - creates nuisance regressors, including RETROICOR via the TAPAS PhysIO Toolbox
 - `multiple_regressors.mat`
 - Sets up 1st level analysis GLM & estimates it
 - Estimates F-contrasts of interests and plots them to .ps file
- Run via
 1. `spm_jobman('interactive',
subject01/batches/demo04_stats_physio/batch_physio_
glm_contrasts.m)`

Demo 4



Setup Pipeline

Quality Monitoring

Multi-Subject

Noise Modeling

Batch Editor

File Edit View SPM BasicIO

Module List

- Named File Selector
- Named File Selector
- Named File Selector
- Make Directory
- Named File Selector
- Named File Selector
- Call MATLAB function DEP
- TAPAS PhysIO Toolbox DEP**
- fMRI model specification DEP
- Model estimation DEP
- Contrast Manager DEP
- Results Report DEP

Current Module: TAPAS PhysIO Toolbox

Help on: TAPAS PhysIO Toolbox

save_dir ... Directory 'first_level_2BF'

log_files

- . vendor Philips
- . log_cardiac ...siological Logfile(1) - Files
- . log_respiration ...siological Logfile(1) - Files
- . log_scan_timing
- . sampling_interval
- . relative_start_acquisition
- . align_scan

sqpar (Sequence timing parameters)

- . Nslices 37
- . NslicesPerBeat
- . TR 2.5
- . Ndummies 5
- . Nscans 100
- . onset_slice 19
- . time_slice_to_slice
- . Nprep 6

thresh (Thresholding parameters for de-noising and timing)

Current Item: save_dir

Reference from

Make Directory: Make Directory 'first_level_2BF'

Specify...

Dependency

Pipeline Noise Modeling/Report



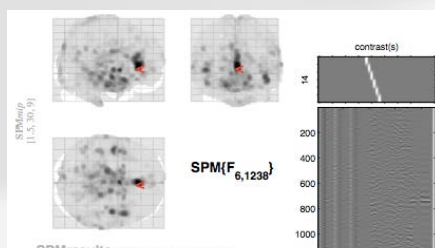
Setup Pipeline

Quality Monitoring

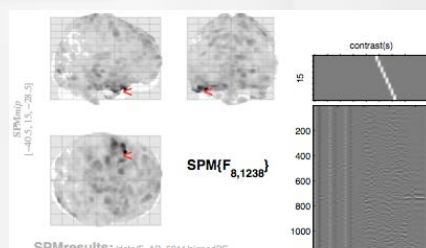
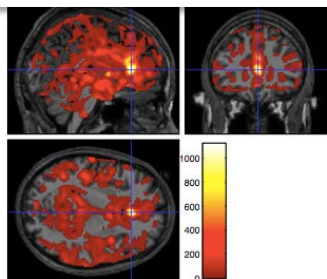
Multi-Subject

Noise Modeling

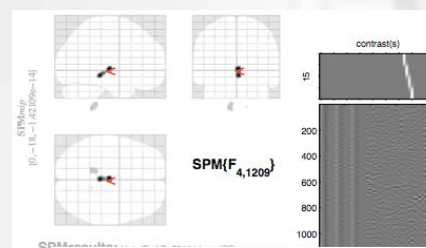
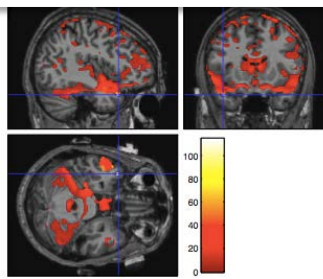
- Model physiological noise as nuisance regressors
- Check noise correction via explained variance (F-contrasts)
 - Automatic contrast creation and report (glass brain): Batch Editor
SPM => Stats => Contrast Manager/Results Report
 - With underlays: `tapas_physio_report_contrasts`



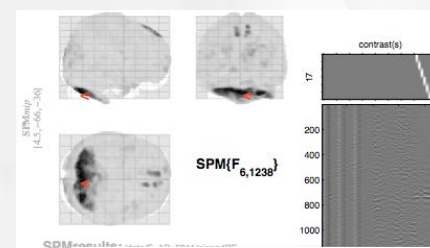
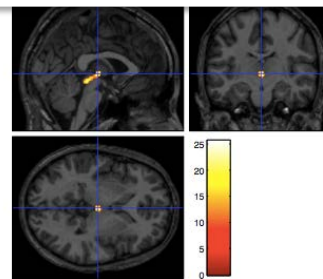
Cardiac



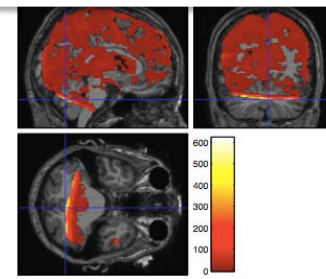
Respiratory



Card x Resp



Movement



- Setting up a Preprocessing Pipeline in SPM:

The Batch Editor

...best way for reproducible, documented preprocessing

- Monitoring and Comparing Pipelines

...within Batch Editor, CheckReg Tool, Stat Images per step

- Multi-subject Pipelines

...via template batch, same sub-folders & Matlab script (subj.-loop)

- Integrating Own Code & Physiological Noise Correction

...by calling Matlab functions within Batch Editor, e.g. PhysIO Toolbox