



Physiological Noise Correction for fMRI

- *an Introduction to the PhysIO Toolbox*

Lars Kasper

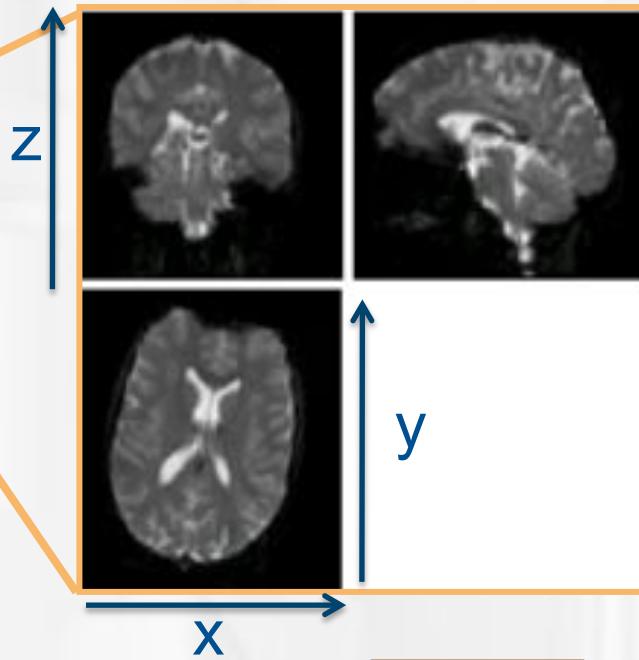
April 30th , 2014



The PhysIO Toolbox

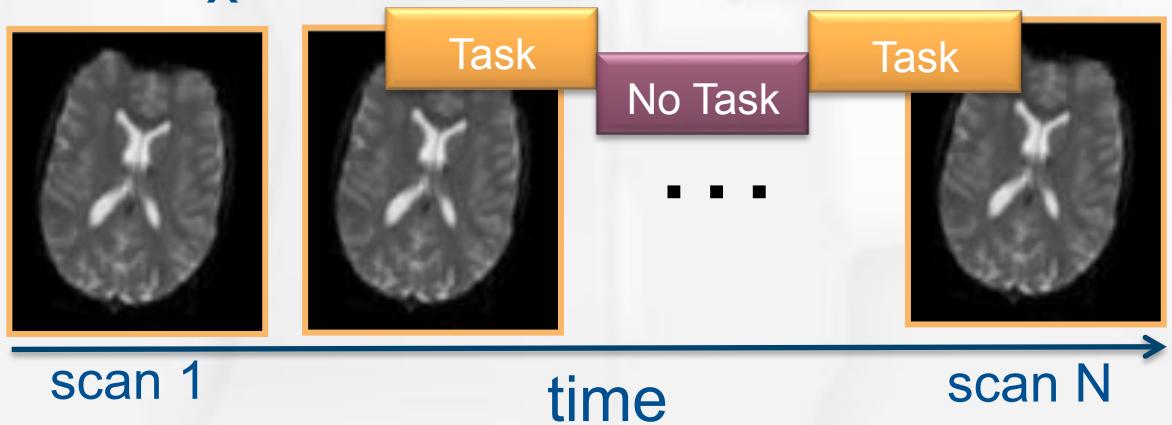
- Developed at the Translational Neuromodeling Unit (TNU) since 2008
 - Lead programmer: Lars Kasper (TNU)
 - Contributors: Jakob Heinze (TNU), Steffen Bollmann (KiSpi Zurich)
- Part of the TNU «TAPAS» software suite
- Used at the TNU, in Zurich and beyond by ~30 researchers
- Current version:
 - <http://www.translationalneuromodeling.org/tapas/>
- Documentation & Example Data:
 - <http://www.translationalneuromodeling.org/software/documentation/>
 - <http://www.translationalneuromodeling.org/software/tapas-data/>

fMRI = Acquiring Movies



- ...of three-dimensional Blood Oxygen-Level Dependent (BOLD) contrast images

- Run/Session:
Time Series of
Images



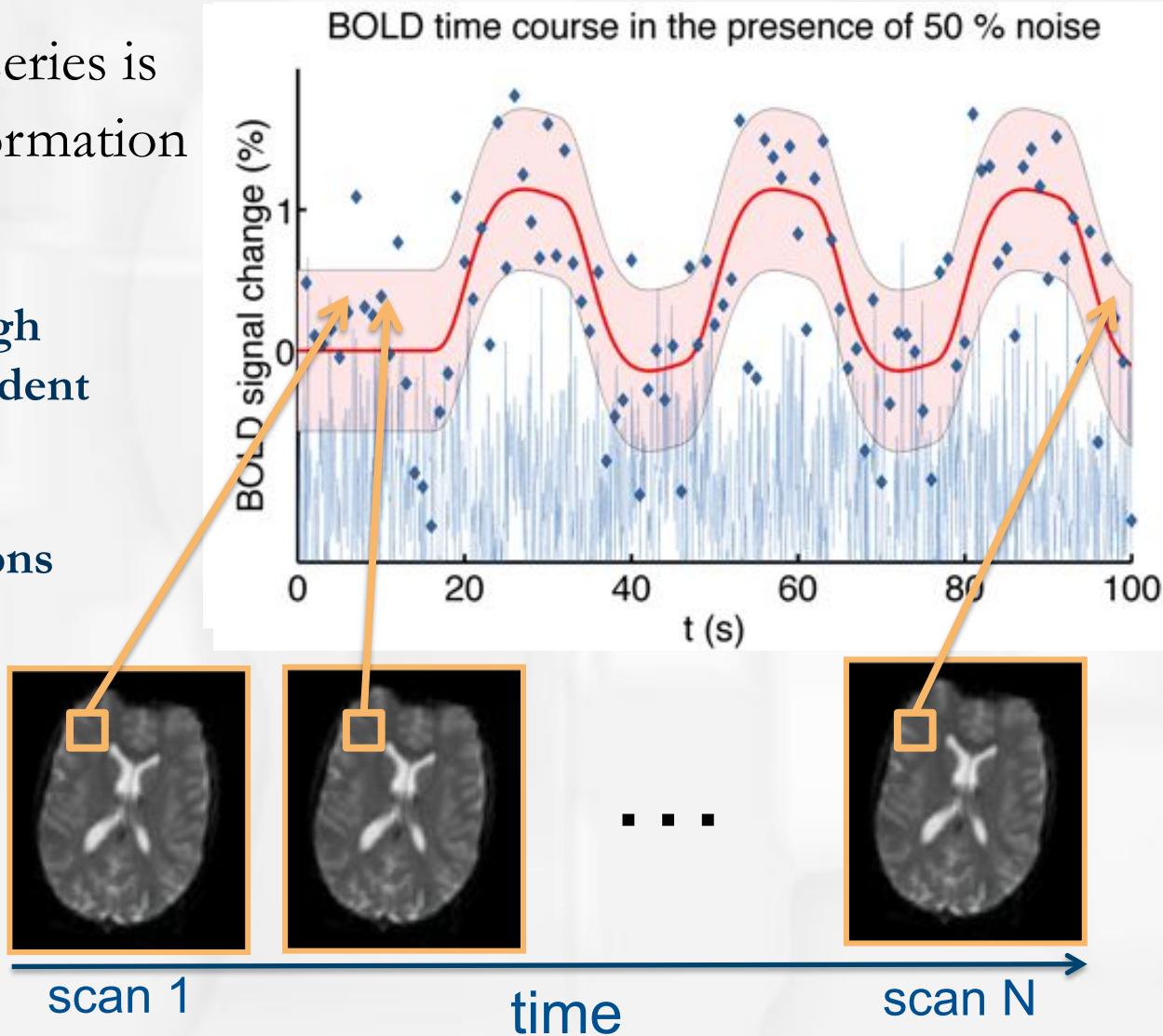
fMRI = Acquiring Movies

- The Localized Time-series is the Fundamental Information Unit of fMRI

Signal: Fluctuation through Blood oxygen level dependent (**BOLD**) contrast

Noise: All other fluctuations

- Run/Session:
Time Series of Images

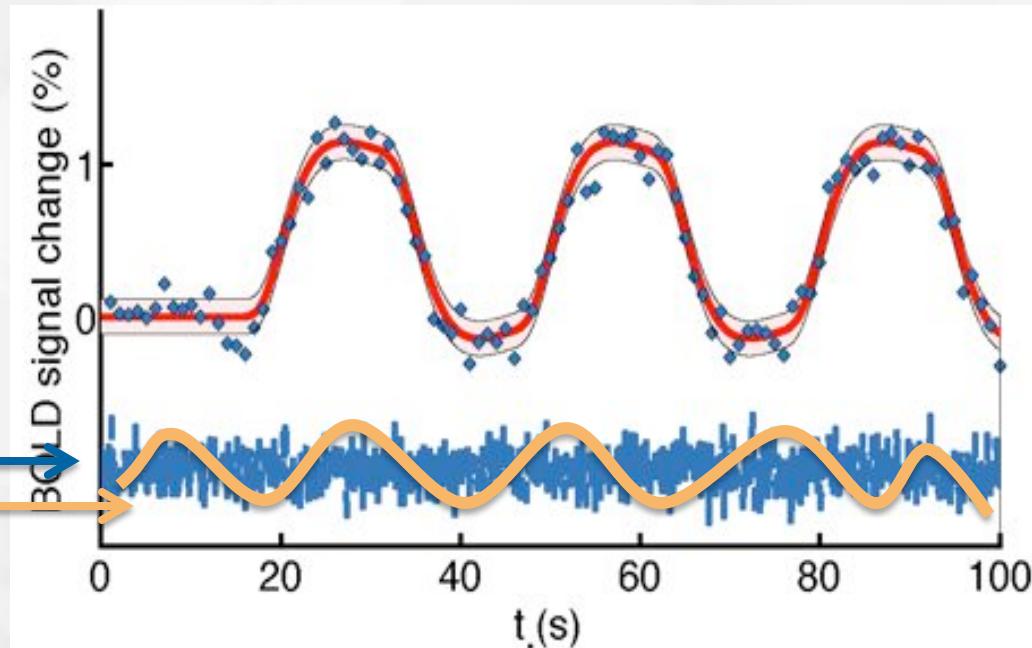


Noise Categories & Reduction

- Thermal Noise
 - temporally uncorrelated
 - reduced SNR → risk of false negatives
 - Remedy: Spatial Smoothing

Noise: All other fluctuations

- “Structured” Noise
 - temporally correlated
 - reduced SNR → risk of false negatives
 - correlated with task → risk of false positives
 - Remedy: Noise Modelling (e.g. GLM)



Inference = Signal-To-Noise

$$t = \frac{\beta}{\sqrt{\sigma_\varepsilon^2(X^T X)^{-1}}} = \frac{\beta \|x\|}{\hat{\sigma}_\varepsilon}$$

$$F = \frac{N - M}{M_1} \cdot \frac{(\sigma_S^2 + \sigma_N^2) - \sigma_N^2}{\sigma_N^2}$$

The Problem: Physiological Noise

- 
- Cardiac effects
 - Respiratory effects

The Problem: Physiological Noise

■ Cardiac effects

- Systole:
 - Blood pumped into brain, vessel volume increases: pulsatile vessels
 - CSF pushed down: pulsatile CSF
- Diastole:
 - Vessel volume decreases
 - CSF flows back into “void” brain volume

A Cardiac Cycle in the Brain

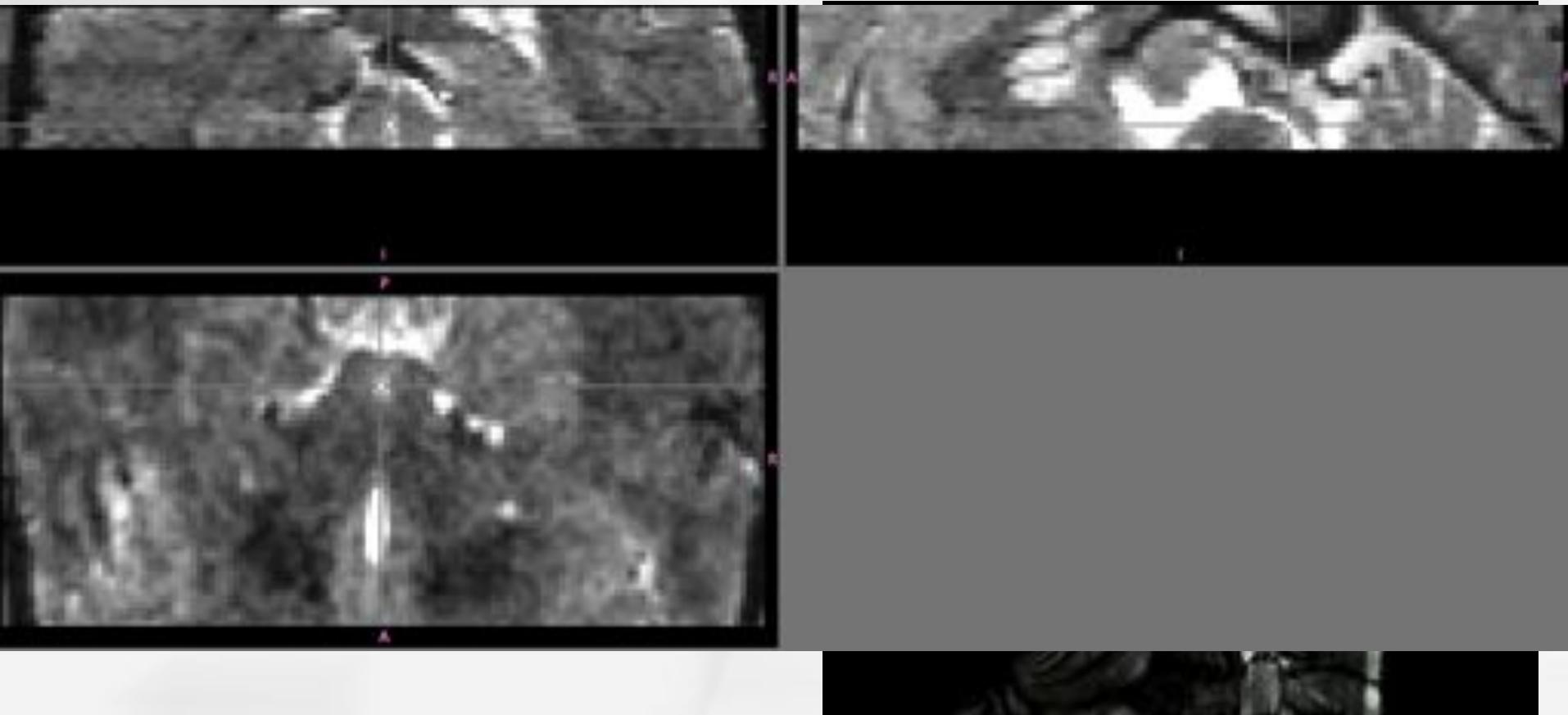


The Problem: Physiological Noise



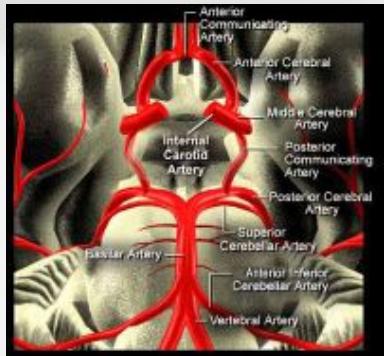
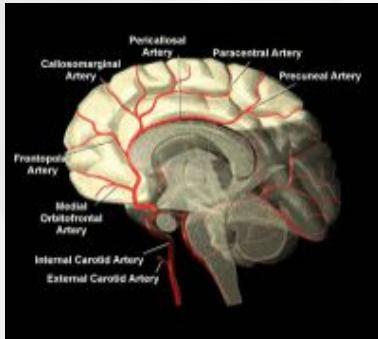
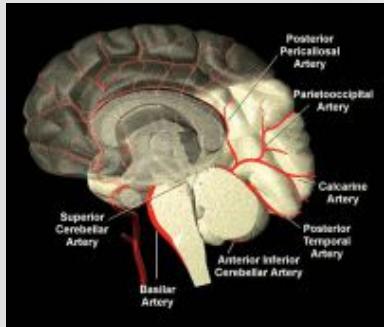
Triggered High-Resolution fMRI

A Cardiac Cycle in the Brain

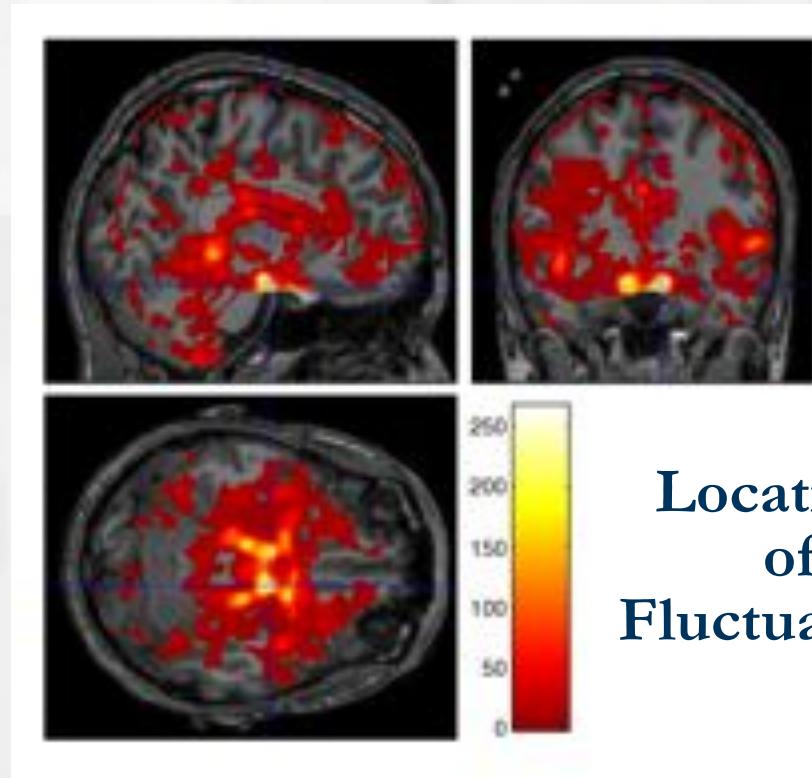


The Problem: Physiological Noise

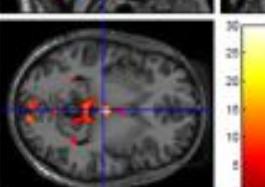
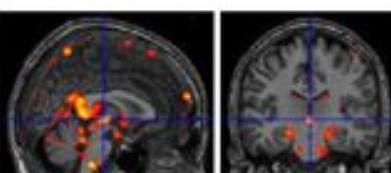
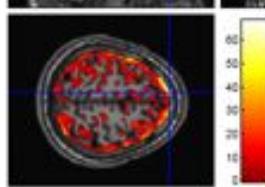
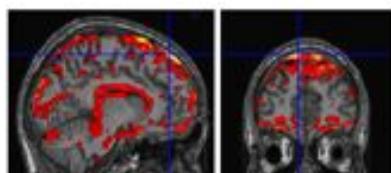
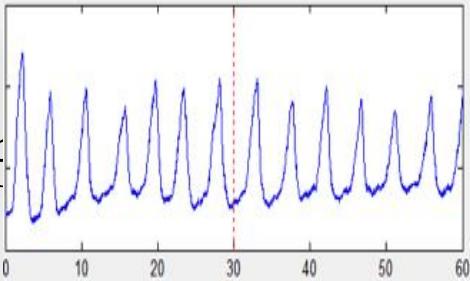
- Cardiac effects



Vessel
Anatomy



The Problem: Physiological Noise

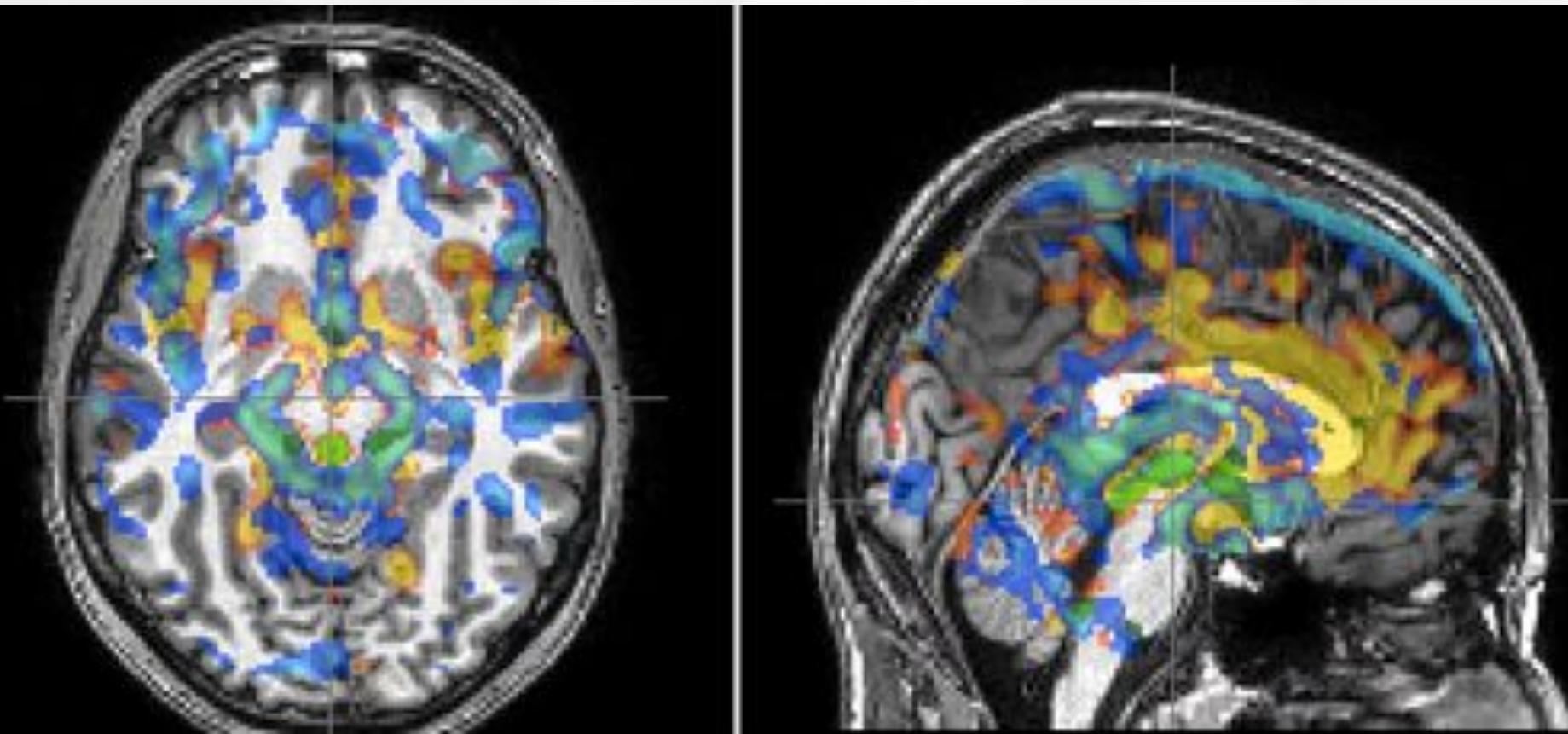


- Respiratory effects
 - Chest (&head) moves with respiratory cycle
 - Changes in lung volume change encoding magnetic field for MR
 - Geometric distortion/scaling
 - Respiratory-sinus arrhythmia
 - Heart beats faster during inhalation

Relevance for Neuromodulation



Translational
Neuromodeling
Unit

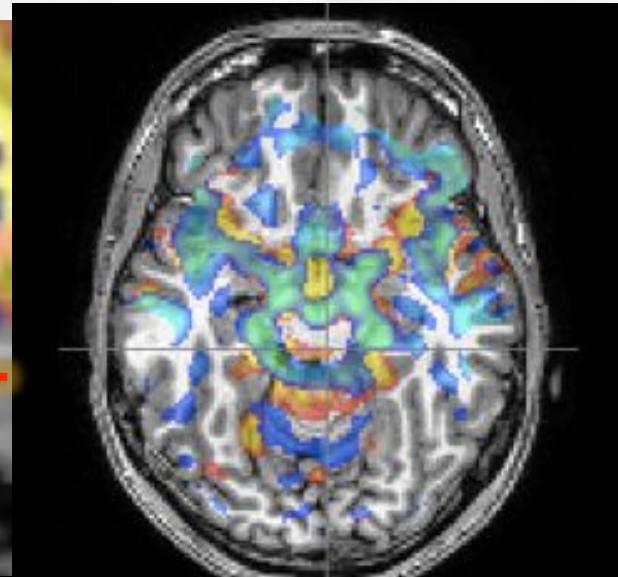
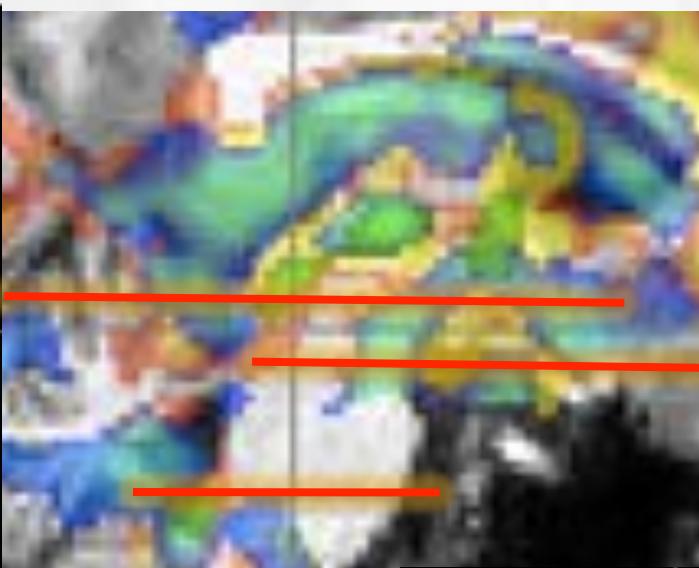
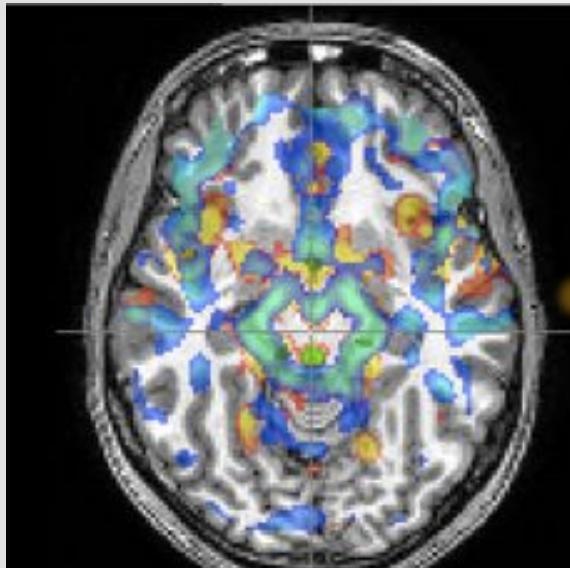


Relevance for Neuromodulation

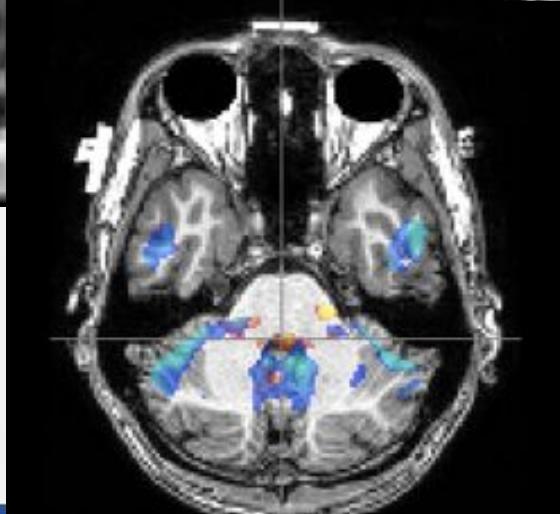


Translational
Neuromodeling
Unit

- VTA (DA)



- Raphe Nuclei (5-HT)



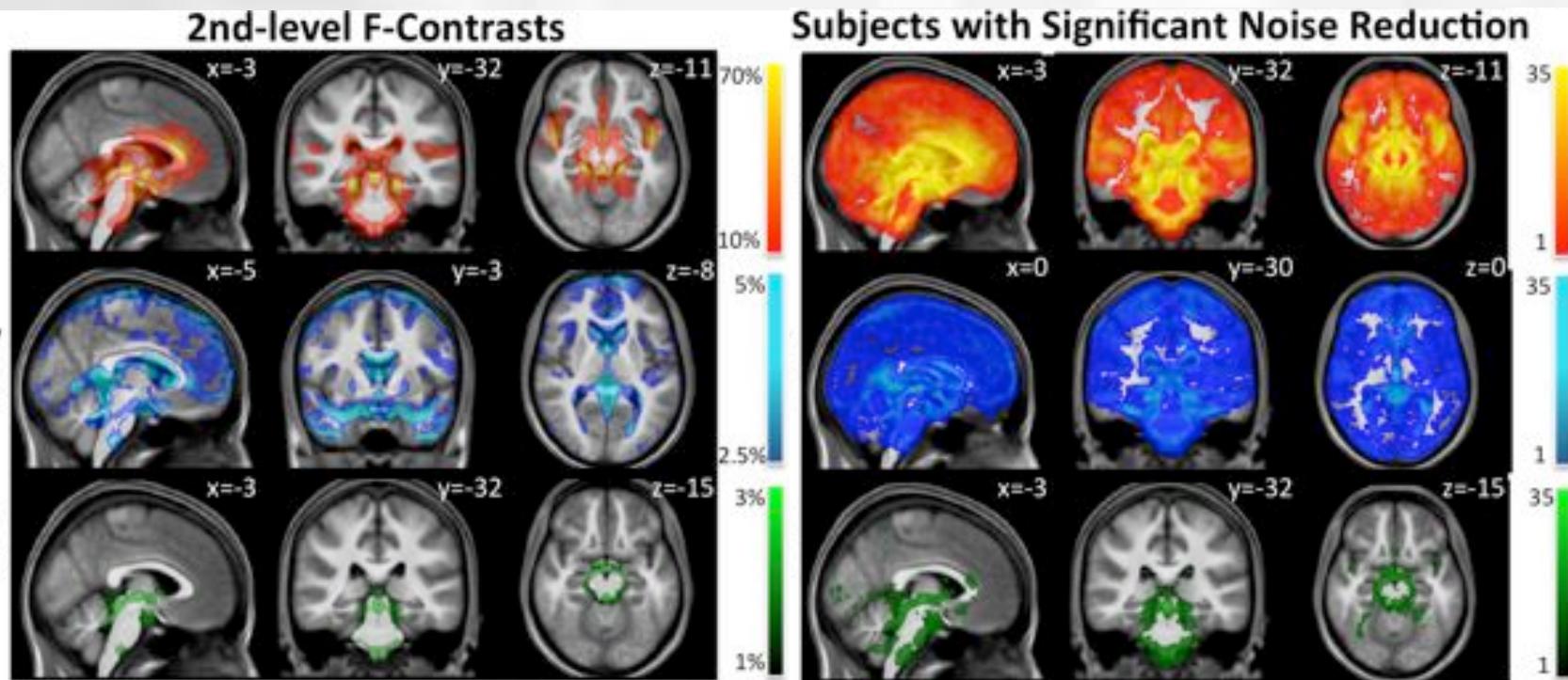
- Locus coeruleus (NA)

Group Level Impact PhysIO

- Andreea Diaconescu (TNU): Social Learning Experiment

2012-2014

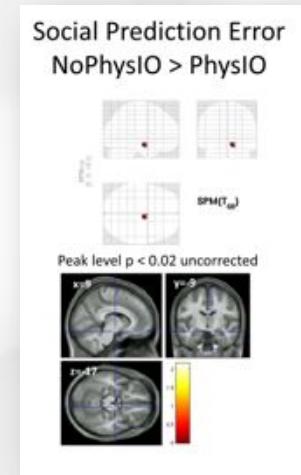
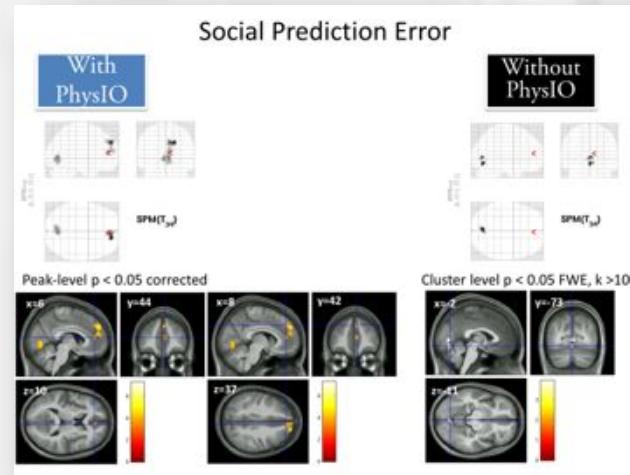
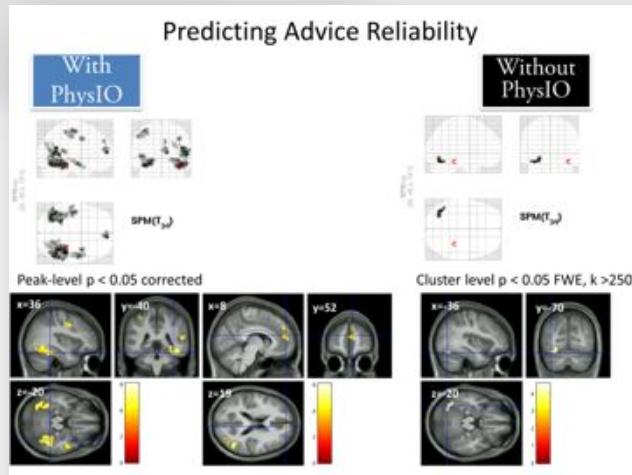
A



When? – PhysIO experience

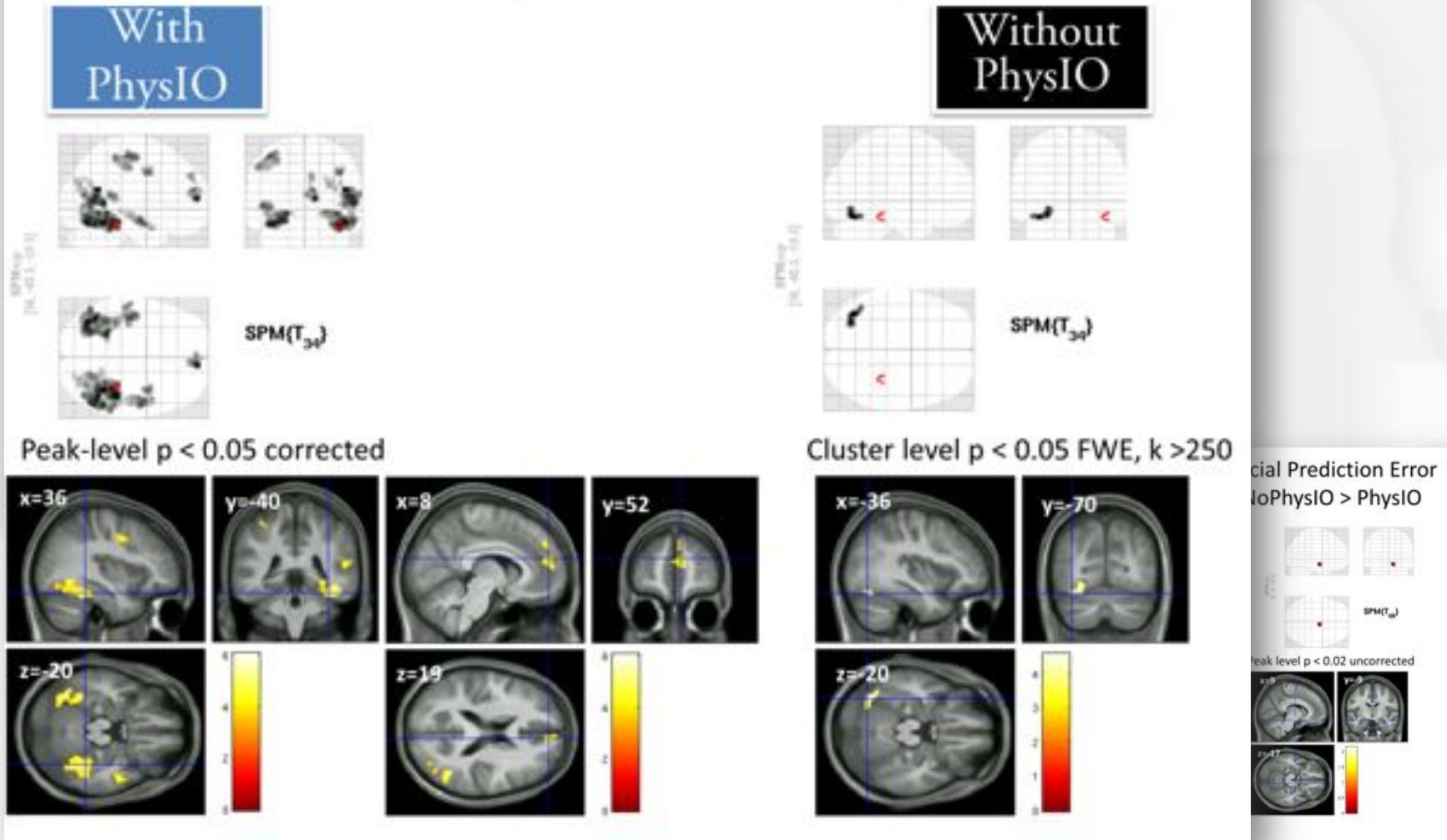
- Andreea Diaconescu (TNU): Social Learning

- Higher sensitivity for group effects (N=35)
 - Prediction of advice reliability: dmPFC, bilateral FFA
 - Prediction error: dmPFC
- Less false/ambiguous positives:
 - Brainstem (Substantia Nigra)



When? – PhysIO experience

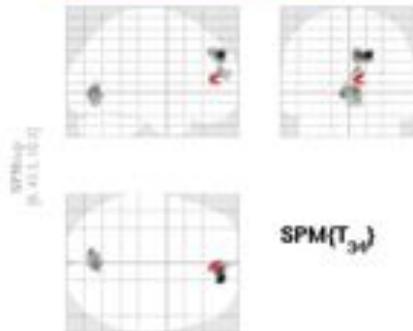
Predicting Advice Reliability



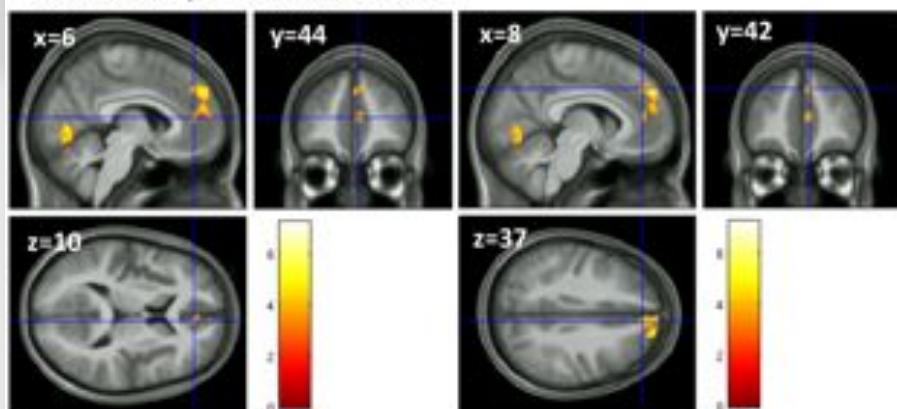
When? – PhysIO experience

Social Prediction Error

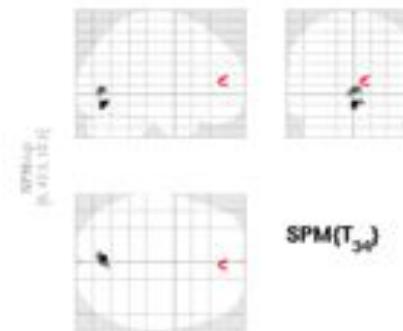
With
PhysIO



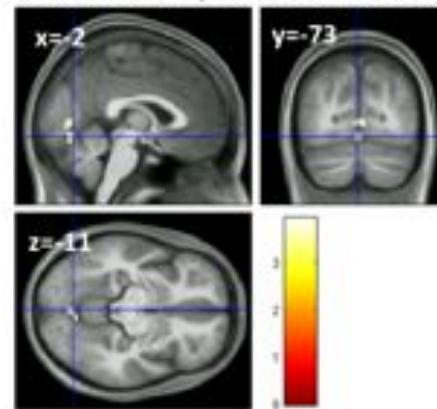
Peak-level $p < 0.05$ corrected



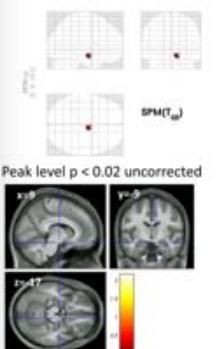
Without
PhysIO



Cluster level $p < 0.05$ FWE, $k > 100$



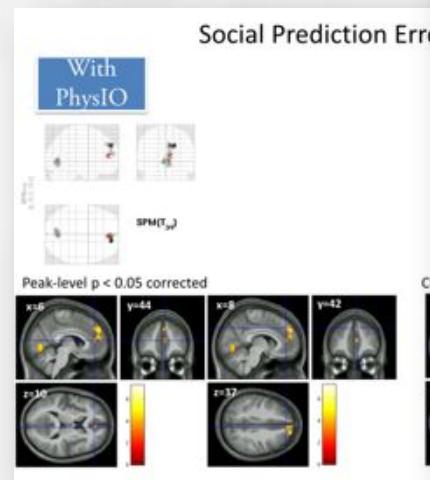
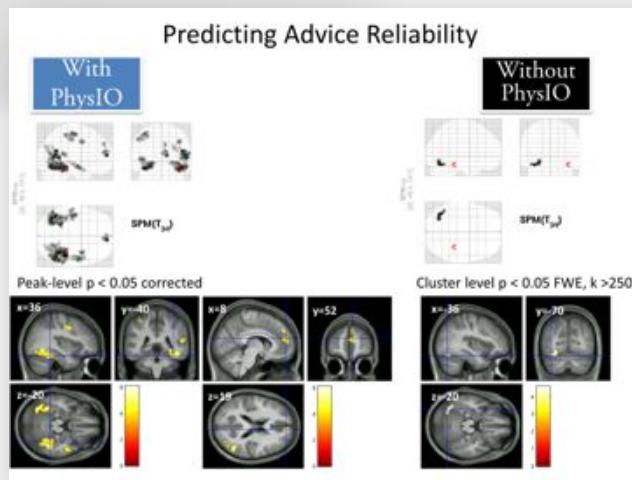
Social Prediction Error
NoPhysIO > PhysIO



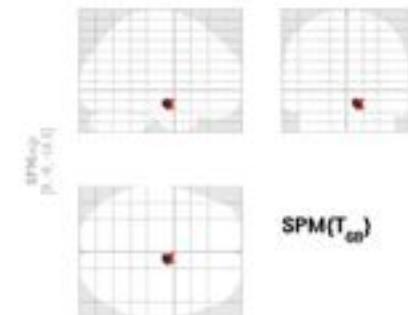
When? – PhysIO experience

- Andreea Diaconescu (TNU): Social Learning

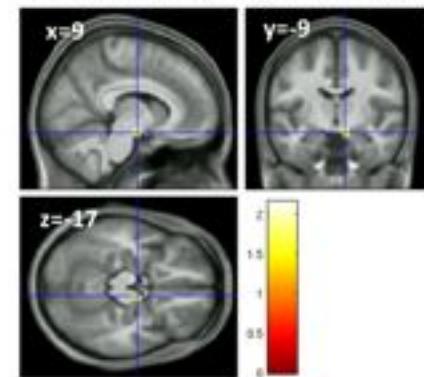
- Higher sensitivity for group effects (N=35)
 - Prediction of advice reliability: dmPFC, bilateral FI
 - Prediction error: dmPFC
- Less false/ambiguous positives:
 - Brainstem (Substantia Nigra)



Social Prediction Error
NoPhysIO > PhysIO



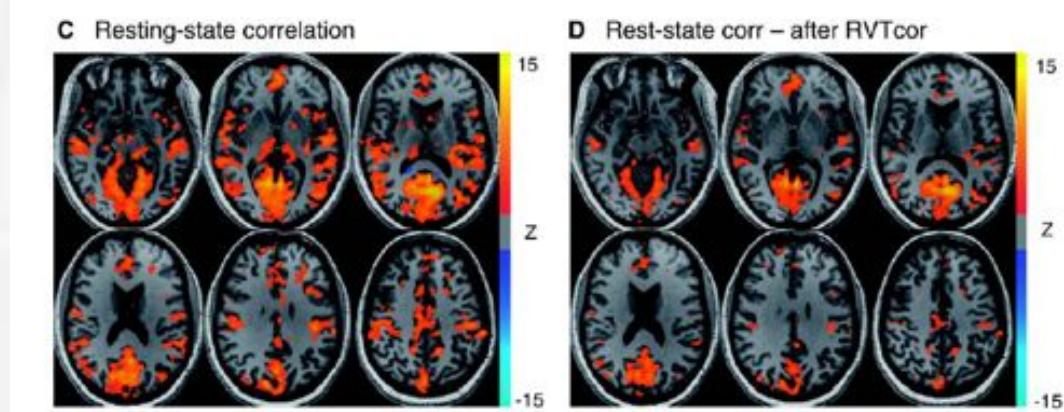
Peak level p < 0.02 uncorrected



When? – Literature Evidence

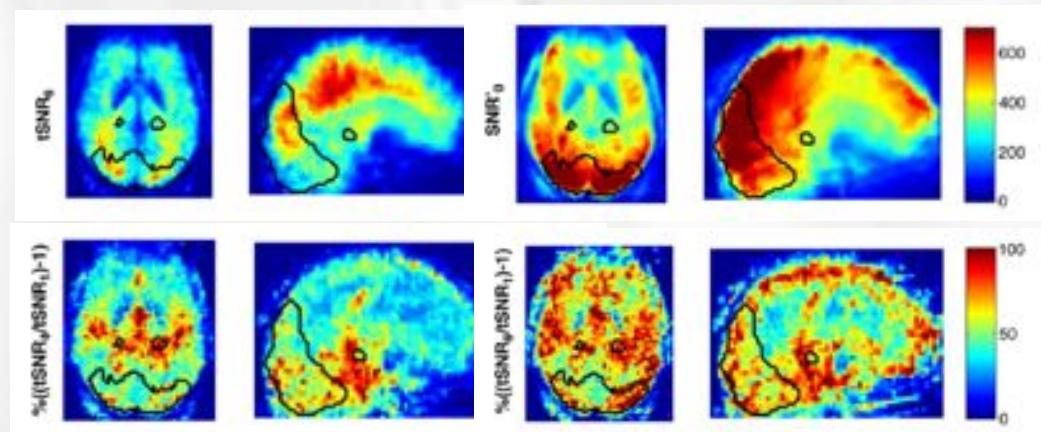
■ Resting-state:

- Birn, R. M. "The Role of Physiological Noise in Resting-state Functional Connectivity." *NeuroImage* 62, 2012
- Birn, R. M., et al. "Separating Respiratory-variation-related Fluctuations from Neuronal-activity-related Fluctuations in fMRI." *NeuroImage* 31, 2006

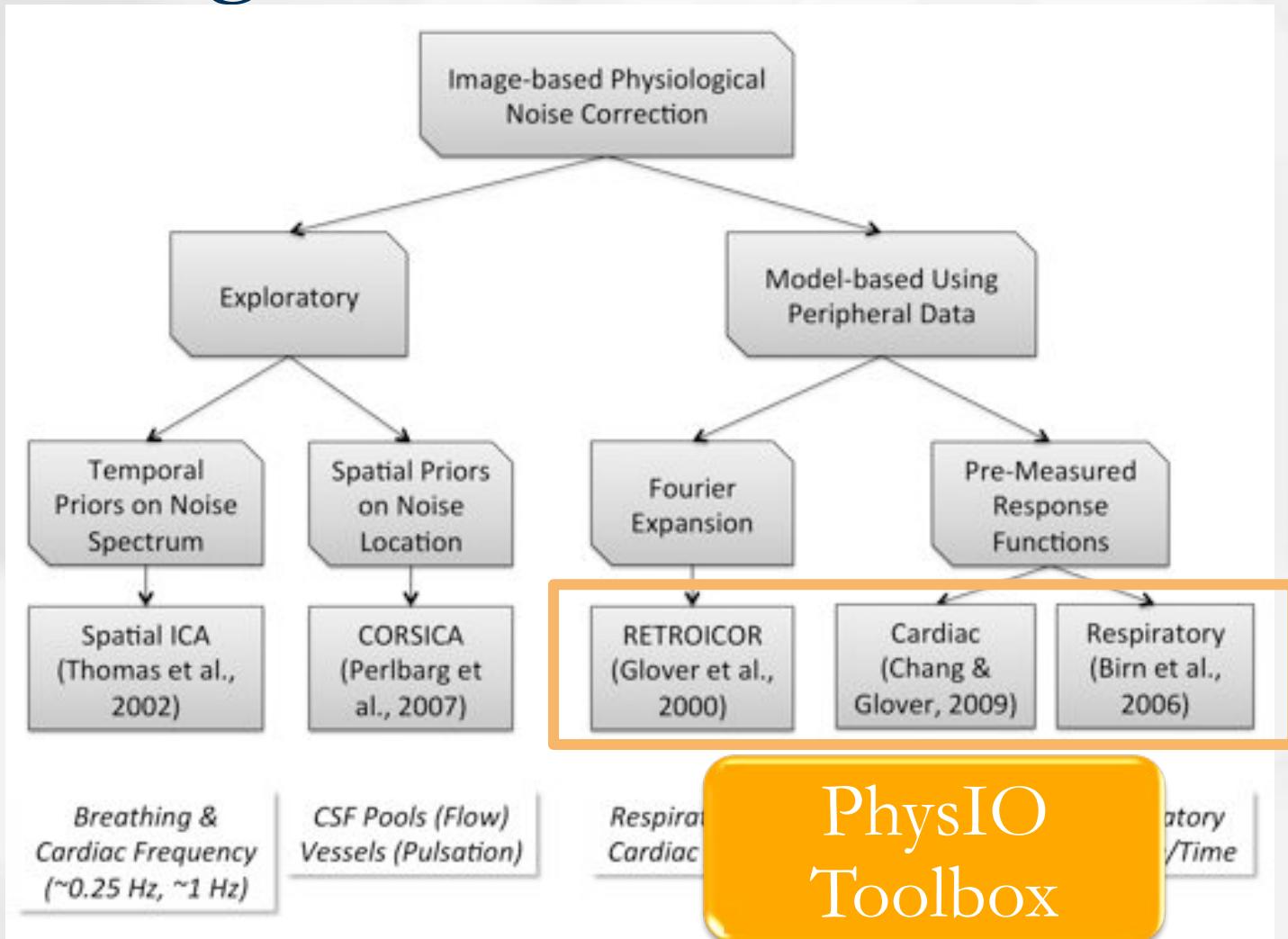


■ Task-based:

- Hutton, C., et al. "The Impact of Physiological Noise Correction on fMRI at 7 T." *NeuroImage* 57, 2011:



The Solution: Image-based Physiological Noise Correction



Noise Modelling

RETROspective
Image CORrection

Cardiac Response
Function

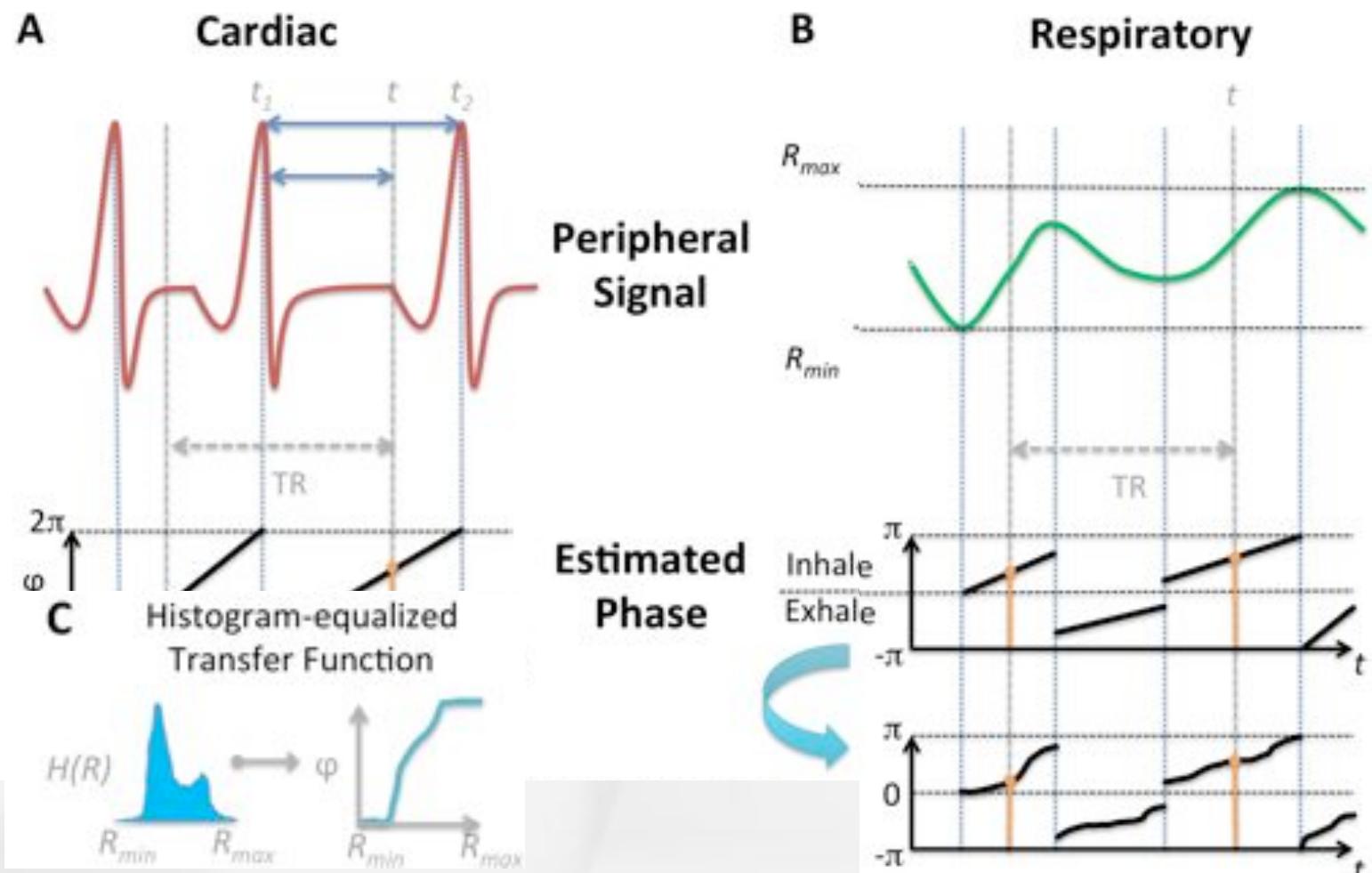
Respiratory
Response Function

- Cardiac/respiratory phase $\varphi_c \quad \varphi_r$
- Fourier expansion (cosine/sine)
- evaluated at 1 time point (slice) per volume = regressor
- Heart Rate
- convolved with CRF
- Resp. Volume per Time
- convolved with RRF

Noise Modelling

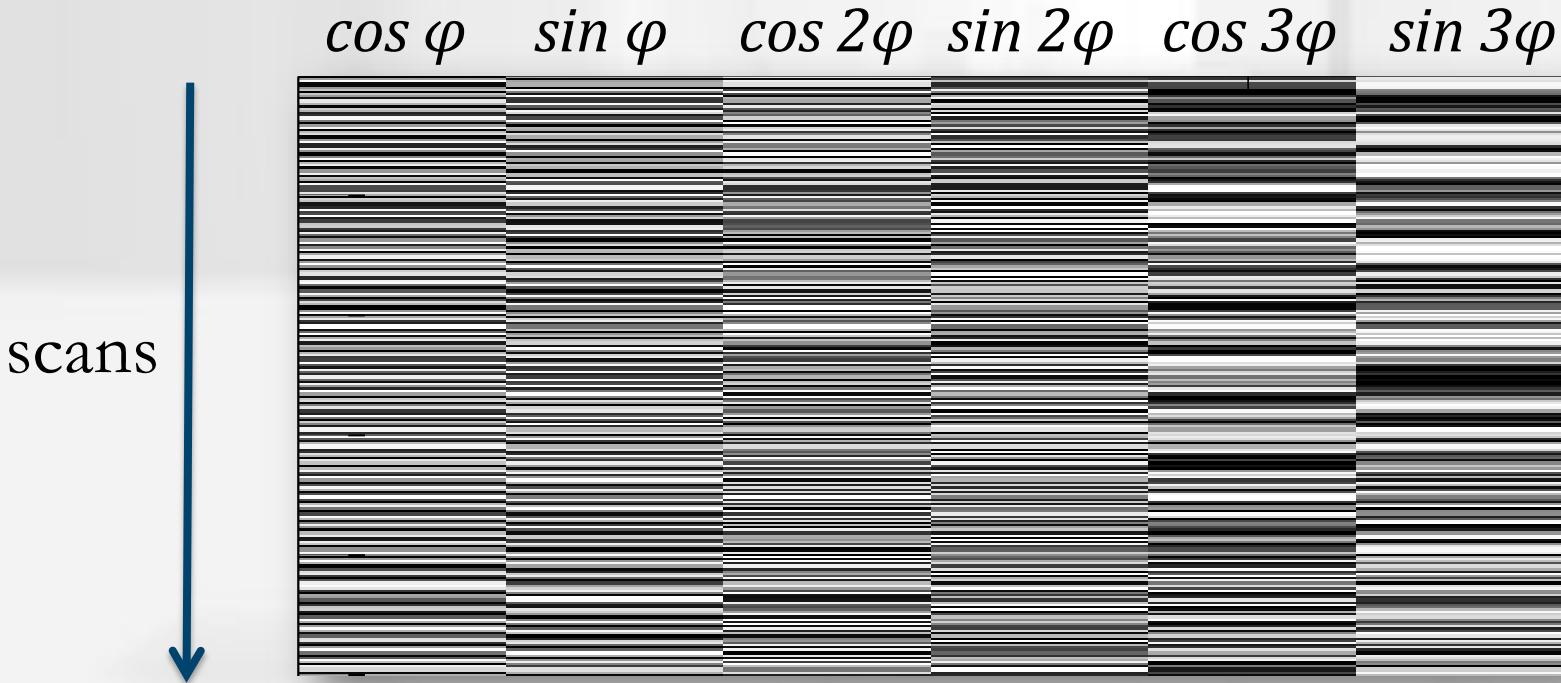
RETROspective
Image CORrection

■ Cardiac/Respiratory phase
Function φ_c Respiratory
Response Function φ_r



Noise Modelling: Fourier Phase Expansion

- cosine and sine to allow for constant phase shifts per voxel
- higher model orders to account for under-sampling of physiological frequencies with typical TR in fMRI



Noise Modelling

RETROspective
Image CORrection

Cardiac Response
Function

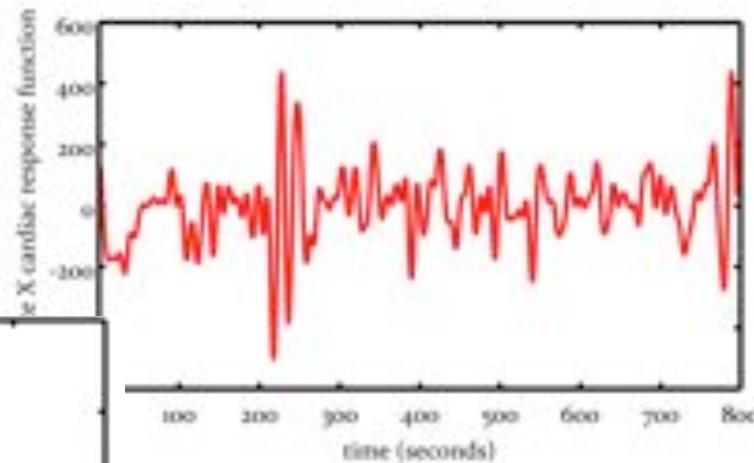
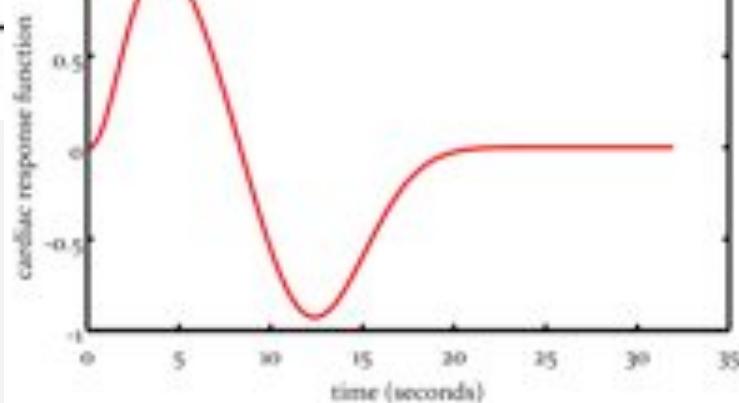
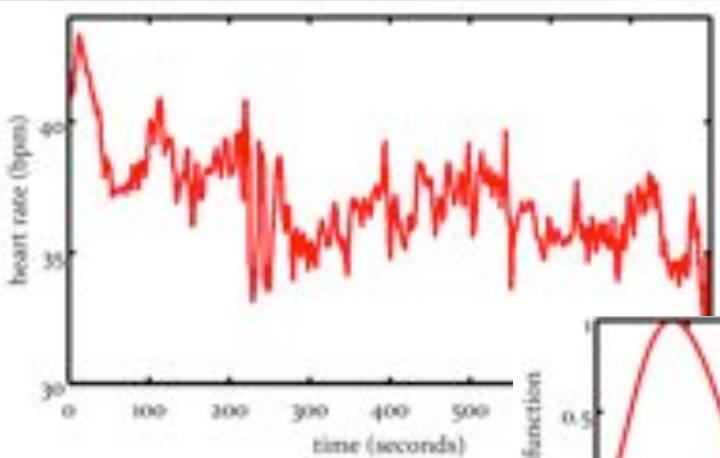
Respiratory
Response Function

- Heart Rate

- convolved with

CRF

- Heart Rate Variability
Response Regressor



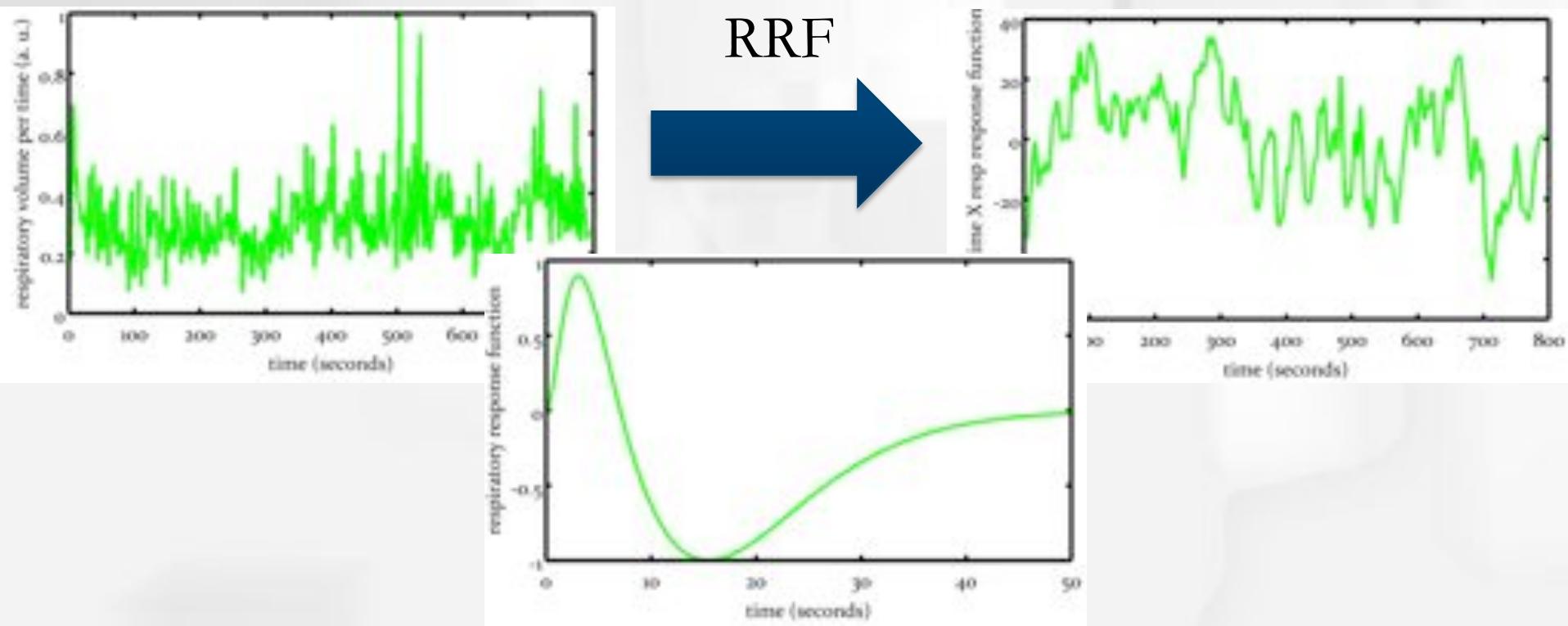
Noise Modelling

RETROspective Image CORrection

Cardiac Response Function

Respiratory Response Function

- Respiratory Volume per Time
- convolved with RRF



The Solution: Physiological Noise Correction

1.
Physiological
Monitoring

2.
Preprocessing
of
Physiological
Data

3. Model time
series
physiological
noise

4. Noise
Reduction and
Assessment

Peripheral
Devices

PhysIO Toolbox

SPM

ECG, PPU → Cardiac cycle

Breathing belt → Respiratory cycle

Confound
regressors

Workflow PhysIO Toolbox

Read logfiles

- Of peripheral physiological data
- Vendor-specific

Preprocess physiological data

- Filter noisy ECG & detect cardiac pulses
- Hand-pick missing pulses

Model time series physiological noise

- RETROICOR
- Respiratory Volume
- Heart Rate

Include confound regressors (GLM)

- Multiple_regressors file for SPM

Finally:

Check Influence of Physiological Noise (Correction) on Data

- SPM
- F-contrast on 1st and second level



Workflow PhysIO Toolbox

Read logfiles

Preprocess
physiological data

Model time series
physiological noise

Include confound
regressors (GLM)

Workflow PhysIO Toolbox

Read logfiles

Preprocess
physiological data

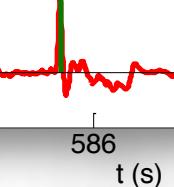
Model time series
physiological noise

Include confound
regressors (GLM)

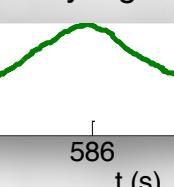
SCANPHYSLOG.log

```
## Universitaet Zuerich
## Wed 20-04-2014 12:14:39
## 2628 1214 775 39
## Dockable table -
# v1raw v2raw v1
-458 325 -494 2 0
-497 284 -527 32
-533 251 -560 2000
-571 219 -592 -104 0 -745 0 0 0 0000
-606 190 -623 -139 0 -745 0 0 0 0000
-636 159 -653 -173 0 -745 0 0 0 0000
-663 123 -680 -206 0 -724 0 0 0 0000
-688 82 -705 -239 0 -724 0 0 0 0000
-710 36 -726 -23000
-724 -9 -744 -32000
-733 -50 -758 -1000
-736 -85 -767 -380
-737 -116 -771 -1000
-736 -145 -770 -457 0 -693 582 0 0 0 0000
-731 -174 -765 -488 0 -693 0 0 0 0000
-725 -200 -757 -516 0 -693
```

ECG normalized

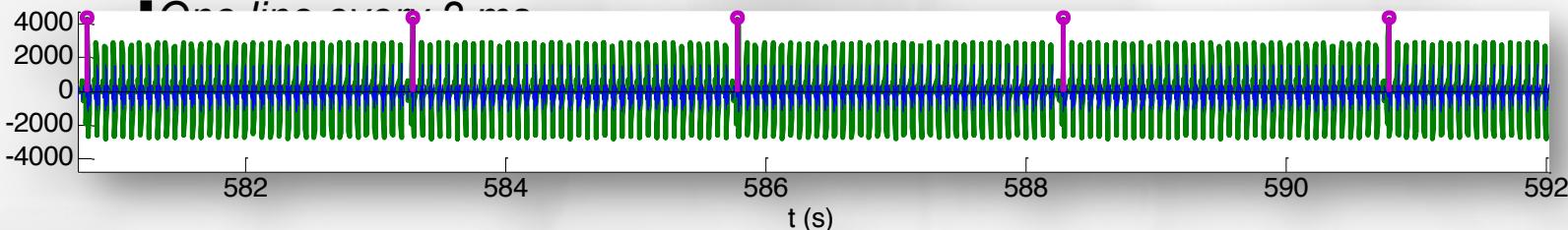


respiratory signal normalized



Gradient X,Y,Z

Open fMRI - 1.5T - 32ms



Workflow PhysIO Toolbox

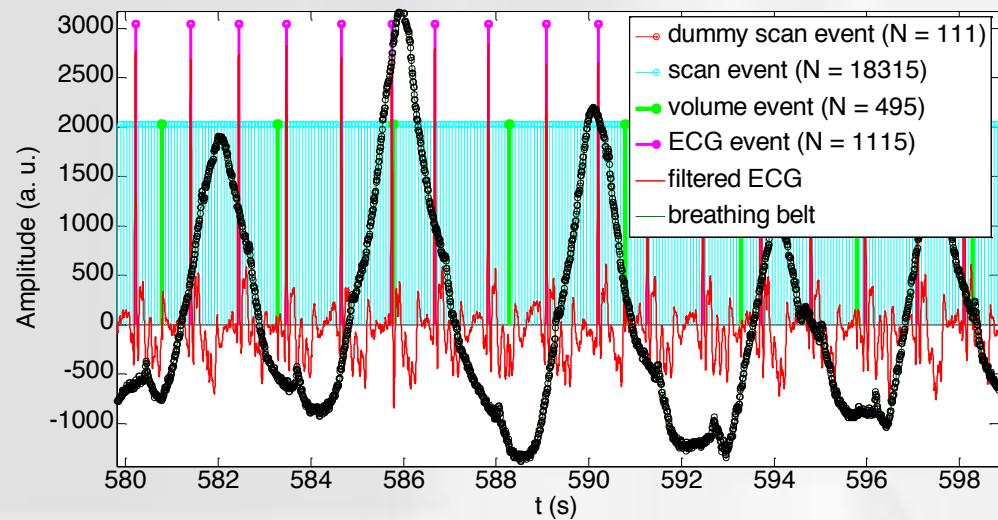
Read logfiles

Preprocess
physiological data

Model time series
physiological noise

Include confound
regressors (GLM)

Cutout region for RETROICOR



- Align scan timing to physiological time series

- Misdetected heartbeats



Workflow PhysIO Toolbox

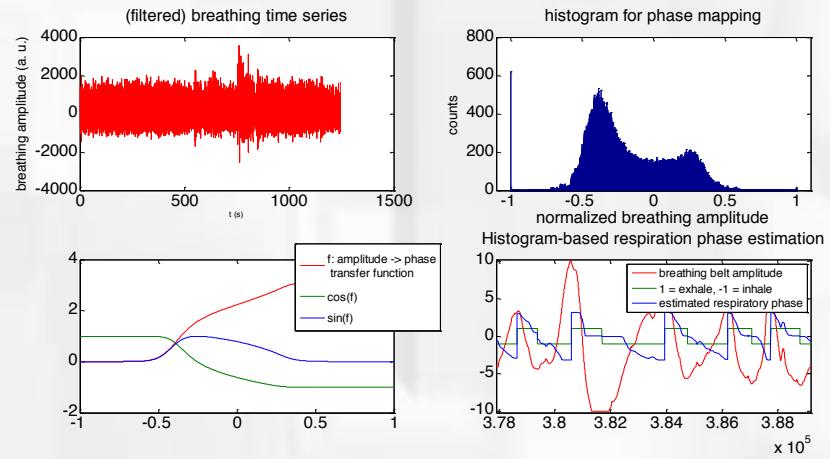
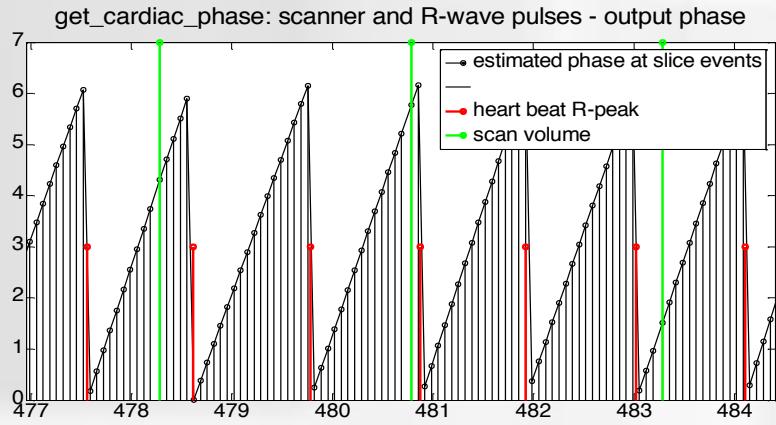
Read logfiles

Preprocess
physiological data

Model time series
physiological noise

Include confound
regressors (GLM)

■ Cardiac & respiratory phase estimation



■ Regressors via Fourier expansion of phases:

RETROspective Image CORrection (RETROICOR)

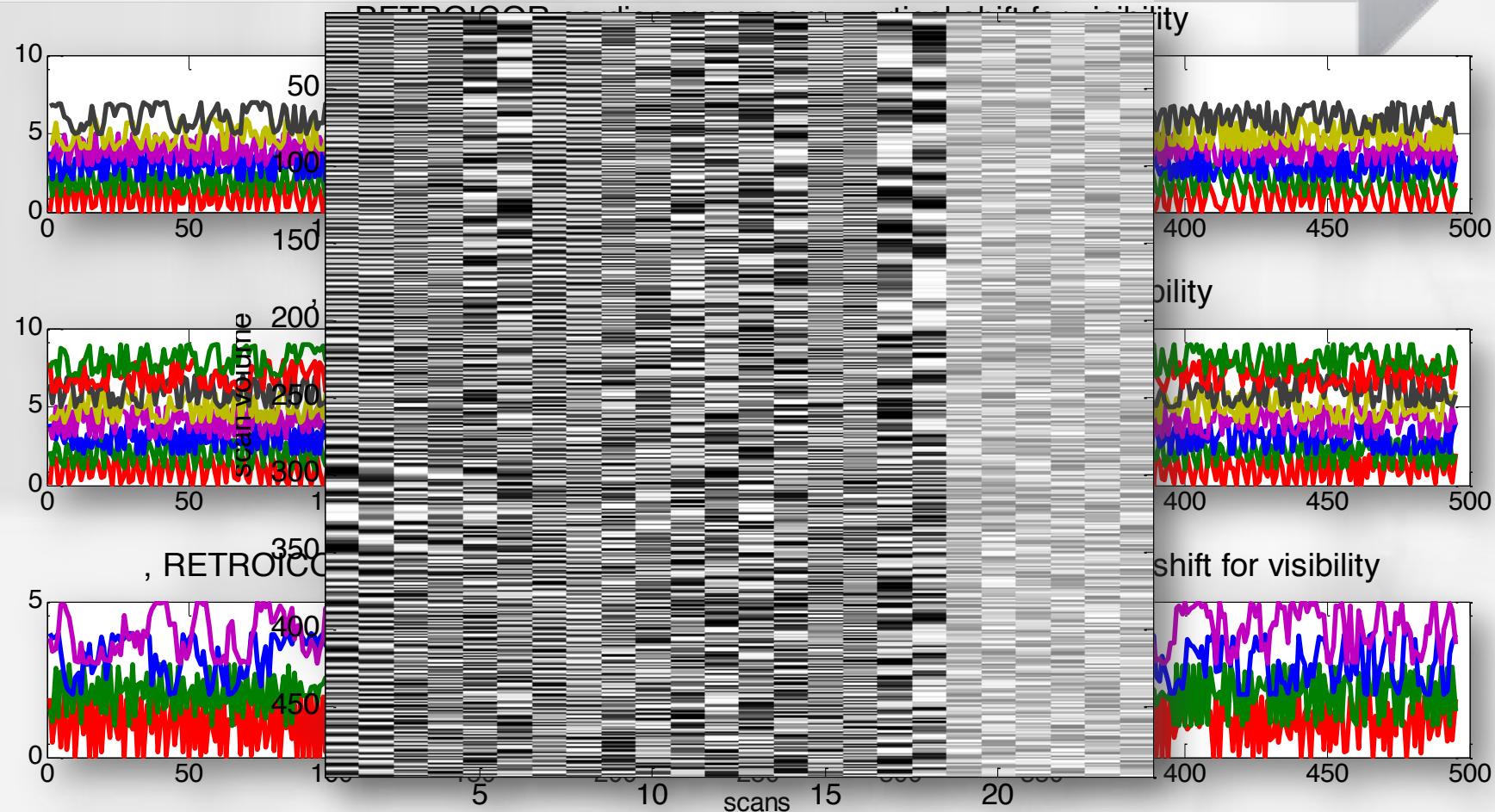
Workflow PhysIO Toolbox

Preprocess physiological regressors matrix for GLM

- specified regressors orthogonalized -

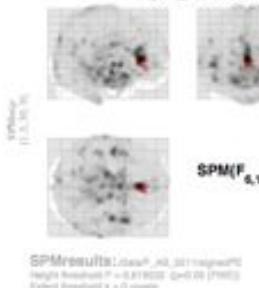
Model time series
ers matrix for GLM
physiological noise
orthogonalized

Include confound
regressors (GLM)

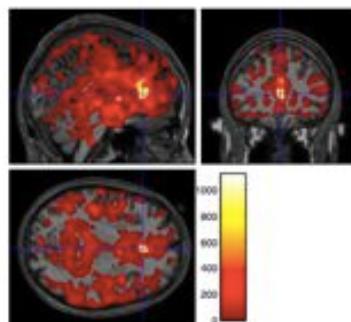


Model Check: SPM F-contrasts

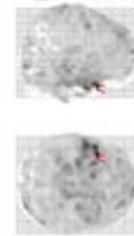
Cardiac regressors



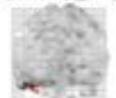
GPMresults.Loban^a .m4a (mp4video.m4a)



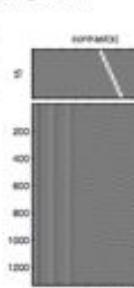
Respiratory regressors



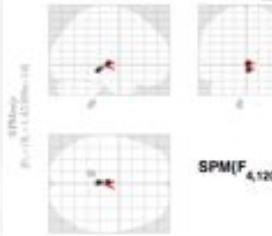
SPM16000C.mncP_AB_S21.mncP
Height threshold F = 8.7500000 (quadratic)
Elastic threshold k = 0.00000



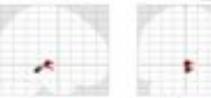
SPM|P



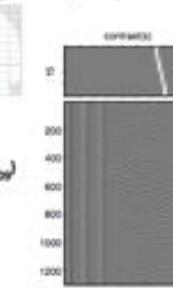
Cardiac x Respiratory



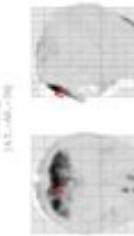
SDMresults: user_AC_2011new17
new model 1 & 2 models - 0.0000000000000000



SPM/F



Movement regressors



SPM(F)
6.120



contradict



Finally:

Check Influence of Physiological Noise (Correction) on Data

- SPM
 - F-contrast on 1st and second level

References

- **Birn**, Rasmus M., Jason B. Diamond, Monica A. Smith, and Peter A. Bandettini. 2006. “Separating Respiratory-variation-related Fluctuations from Neuronal-activity-related Fluctuations in fMRI.” *NeuroImage* 31 (4) (July 15): 1536–1548. doi:10.1016/j.neuroimage.2006.02.048.
- **Glover**, G H, T Q Li, and D Ress. 2000. “Image-based Method for Retrospective Correction of Physiological Motion Effects in fMRI: RETROICOR.” *Magnetic Resonance in Medicine: Official Journal of the Society of Magnetic Resonance in Medicine / Society of Magnetic Resonance in Medicine* 44 (1) (July): 162–7.
- **Harvey**, Ann K., Kyle T.S. Pattinson, Jonathan C.W. Brooks, Stephen D. Mayhew, Mark Jenkinson, and Richard G. Wise. 2008. “Brainstem Functional Magnetic Resonance Imaging: Disentangling Signal from Physiological Noise.” *Journal of Magnetic Resonance Imaging* 28 (6): 1337–1344. doi:10.1002/jmri.21623.
- **Hutton**, C., O. Josephs, J. Stadler, E. Featherstone, A. Reid, O. Speck, J. Bernarding, and N. Weiskopf. 2011. “The Impact of Physiological Noise Correction on fMRI at 7 T.” *NeuroImage* 57 (1) (July 1): 101–112. doi:10.1016/j.neuroimage.2011.04.018.
- **Josephs**, O., Howseman, A.M., Friston, K., Turner, R., 1997. “Physiological noise modelling for multi-slice EPI fMRI using SPM.” Proceedings of the 5th Annual Meeting of ISMRM, Vancouver, Canada, p. 1682
- **Kasper**, Lars, Sarah Marti, S. Johanna Vannesjö, Chloe Hutton, Ray Dolan, Nikolaus Weiskopf, Klaas Enno Stephan, and Klaas Paul Prüssmann. 2009. “Cardiac Artefact Correction for Human Brainstem fMRI at 7 Tesla.” In *Proc. Org. Hum. Brain Mapping* 15, 395. San Francisco.