



Weierstrass Institute for
Applied Analysis and Stochastics



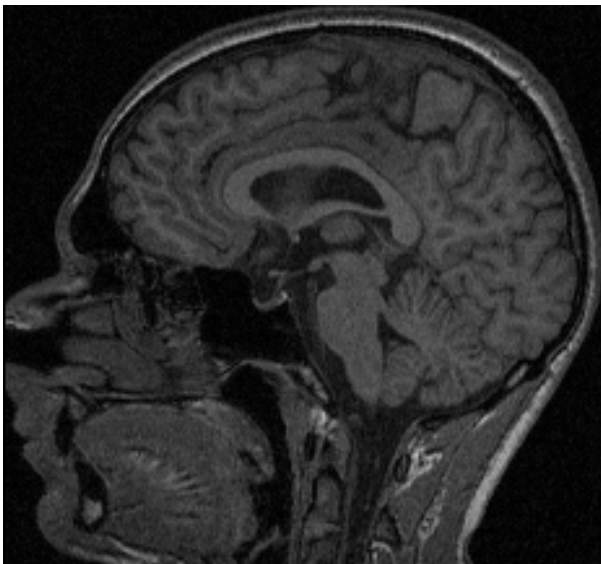
DFG-Forschungszentrum MATHEON
Mathematik für Schlüsseltechnologien

Statistical Parametric Maps for Functional MRI

Experiments in R: The Package fmri

Karsten Tabelow

UseR!2011



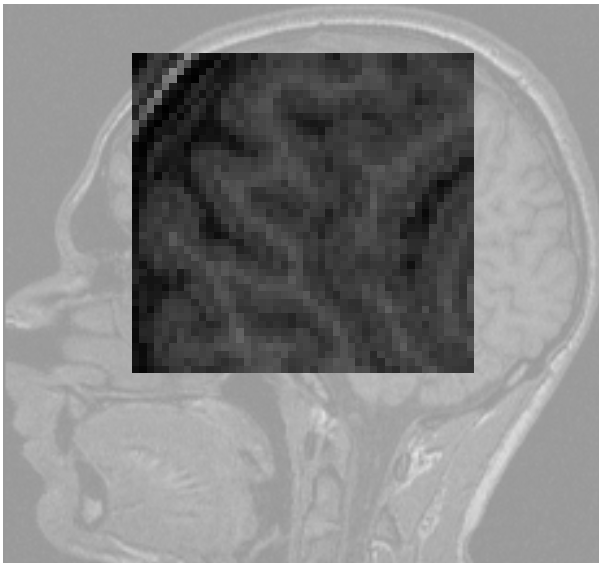


Image Denoising Methods

- Kernel estimators
- Wavelets, Curvelets, ...
- MCMC, SANN
- Diffusion methods
- Scale-space methods
- Scanner upgrade

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Structural Adaptive Smoothing

- Local parametric model for data
- Iterative estimation of parameters and homogeneity regions
- Dimension-free, structure preserving/enhancing

General setup / Local parametric model

- Design: $x_1, \dots, x_n \in \mathcal{X} \subseteq \mathbb{R}^p$
- Observations: $Y_1, \dots, Y_n \in \mathcal{Y} \subset \mathbb{R}^q$

$$Y_i \sim \mathbf{P}_{\theta(x_i)} \quad \theta : \mathbb{R}^p \rightarrow \Theta \quad (\text{i.i.d.})$$

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Structural assumption

- \exists Partitioning $\mathcal{X} = \bigcup_{m=1}^M \mathcal{X}_m$ such that

$$\theta(x) = \theta(x_i) \Leftrightarrow \exists m : x \in \mathcal{X}_m \wedge x_i \in \mathcal{X}_m$$

i.e. θ constant on each \mathcal{X}_m

- Some components of θ may be global parameters.

General idea

- Determine *structure* and *estimates* in an *iterative* procedure
- Weighting scheme $W(x_i) = (w_{i1}, \dots, w_{in})$
- $W(x_i)$ describes a local model

$$w_{ij}^{(k)} = K_{loc}(\|x_i - x_j\|^2/h_k^2) \cdot K_s(s_{ij}^{(k)}/\lambda)$$

s_{ij} measures the difference between estimates $\hat{\theta}(x_i)$ and $\hat{\theta}(x_j)$

- Four steps:
 - A) Initialize estimates $\hat{\theta}(x_i)$ and describe initial partitioning $W(x_i)$ ideally by observed values and $w_j(x_i) = w_{ij} = \delta_{ij}$
 - B) “Learn” about $\theta(x)$ from information on partitioning
 - C) “Learn” about partitioning from estimates $\hat{\theta}(x)$
 - D) Iterate while inspecting scale space from local to global

Results

- *Propagation* under homogeneity: For a homogeneous situation and $\mu > 4$ the last step estimate $\hat{\theta}_i^{k^*}$ fulfills $\mathbf{P}(\bar{N}_i^{(k^*)} \mathcal{K}(\hat{\theta}_i, \theta) > \mu \log(n)) \leq 2k^*/n$.
- *Propagation* under local homogeneity: Similar results for interior points of local homogeneous regions.
- *Stability* of estimates: The quality of the best intermediate result holds (up to a constant) for the final estimate.
- *Optimal adaptive rate* of estimation under smoothness conditions on f_θ .
- *Separation* property

Parameters

- λ can be selected by a *propagation condition*, independent of data
- k^* determines the maximum bandwidth
- $c_h = 1.25^{1/p}$ provides exponential growth of sum of location weights
- Kernels $K_{loc}(z) = (1 - z)_+$ and $K_s(z) = \min(1, 2(1 - z))_+$

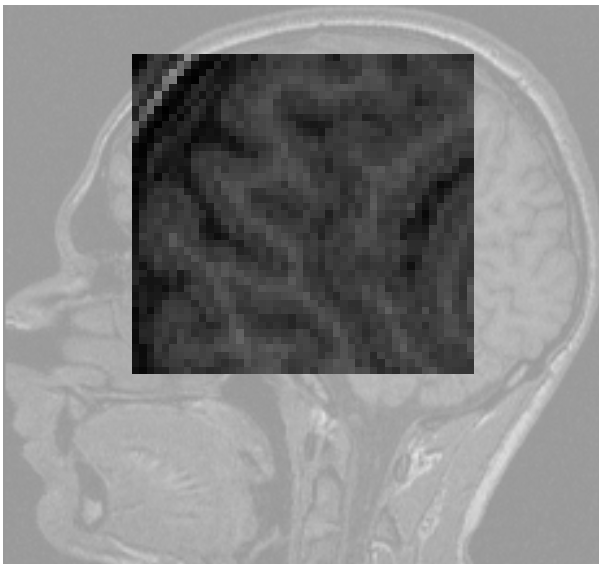
Propagation condition

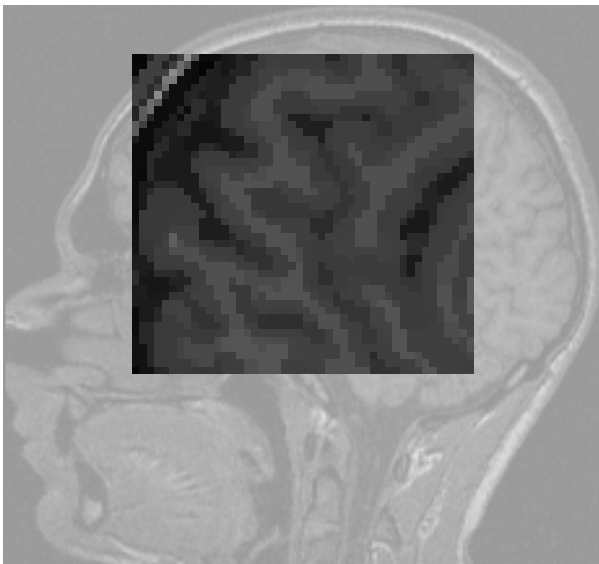
$\hat{\theta}^{(k)}(x)$ PS estimate of $\theta(x)$ from iteration k ,

$\tilde{\theta}^{(k)}(x)$ Kernel estimate of $\theta(x)$ using bandwidth $h^{(k)}$

Let $\theta(x) \equiv \theta$. Select parameters such that $\forall k$ and some $\alpha > 0$

$$\mathbf{E} |\hat{\theta}^{(k)}(\mathbf{x}) - \theta| \leq (1 + \alpha) \mathbf{E} |\tilde{\theta}^{(k)}(\mathbf{x}) - \theta|.$$





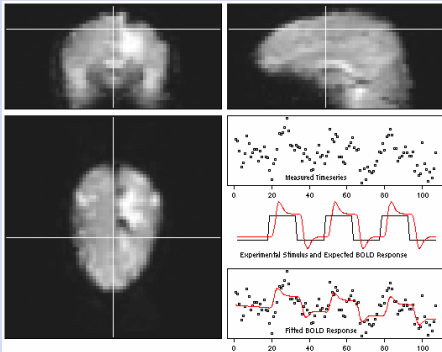
Human Eye vs. Structural adaptive smoothing

- Human eye is a very good denoiser
- Experience: Structural adaptive smoothing compares well with eyes.
- Limited use for visual inspection and for volumetric (integral) quantities

Applications

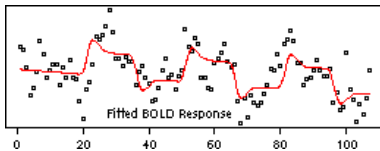
- Suitable for higher dimensional data
- Structure enhancement, structure identification.
- functional MRI - package **fmri**

fMRI data = 3D + T



How to smooth fMRI data?

- Blurring effect in Gaussian filter
- T2-volumes have limited structure.
- Local tests inefficient in 3D+T space
- Interesting structure is activation structure
- We have a model about HRF



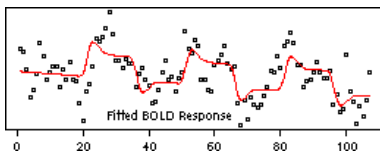
Data Preparation

Registration

Motion correction

Normalization

Smoothing



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Motion correction

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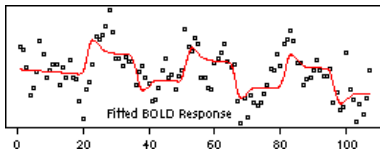
Smoothing

Linear Modeling

$$Y = X\beta + \epsilon$$

Prewhitening

$$\tilde{Y} = \tilde{X}\beta + \tilde{\epsilon}$$



Data Preparation

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Motion correction
Normalization

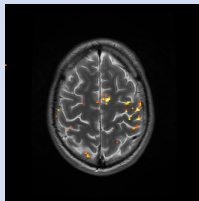
Linear Modeling

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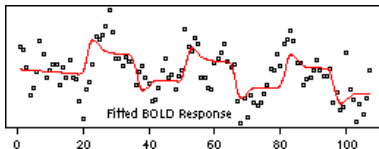
$$\tilde{Y} = \tilde{X}\beta + \tilde{\epsilon}$$

Smoothing



Thresholding

t-statistic ...
multiple test problem



Data Preparation

- Registration
- Motion correction
- Normalization

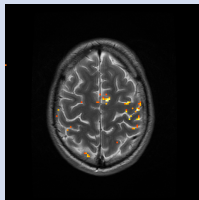
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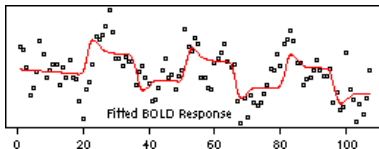
Smoothing



no filter

Thresholding

- t-statistic
- FDR



Data Preparation

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- Motion correction
- Normalization

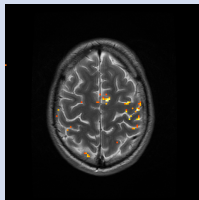
Linear Modeling

$$Y = X\beta + \epsilon$$

Prewhitening

$$\tilde{Y} = \tilde{X}\beta + \tilde{\epsilon}$$

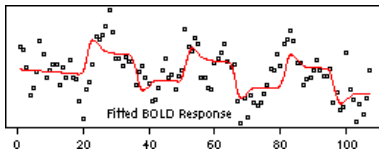
Smoothing



no filter

Thresholding

- t-statistic
- RFT



Data Preparation

Registration
Motion correction
Normalization

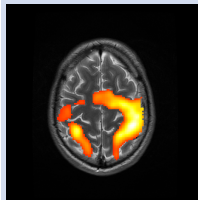
Linear Modeling

$$Y = X\beta + \epsilon$$

Prewhitening

$$\tilde{Y} = \tilde{X}\beta + \tilde{\epsilon}$$

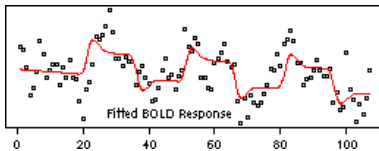
Smoothing



Gaussian filter

Thresholding

t-statistic
RFT



Data Preparation

Registration
Motion correction
Normalization

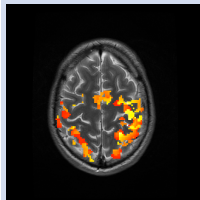
Linear Modeling

$$Y = X\beta + \epsilon$$

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$$\tilde{Y} = \tilde{X}\beta + \tilde{\epsilon}$$

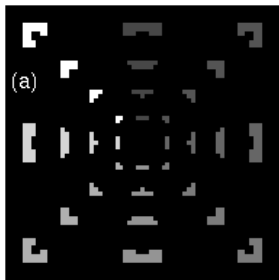
Smoothing



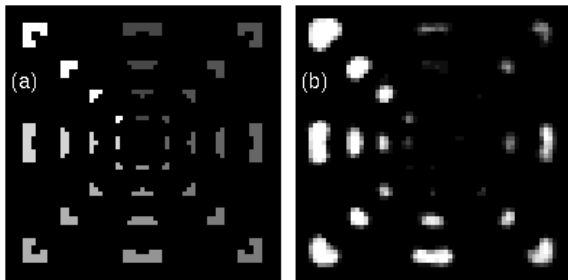
adaptive!

Thresholding

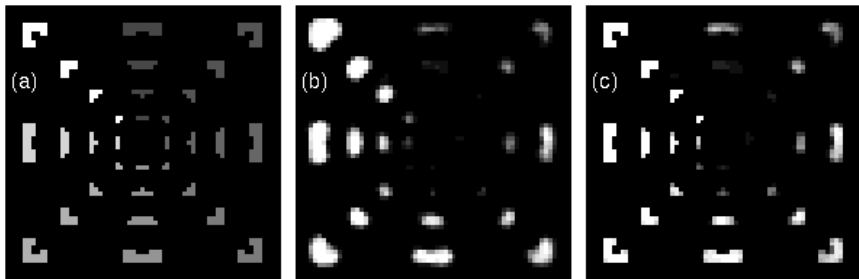
t-statistic
RFT



- (a) Signal location in Phantom data (3D) with different signal size
26 slices with 15 containing activations

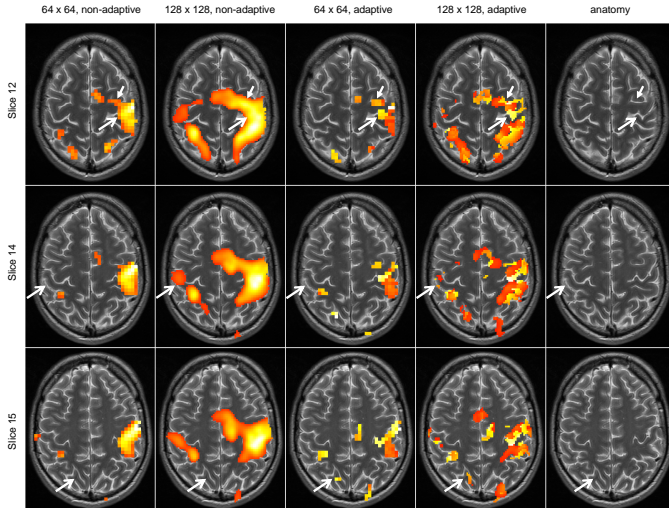


- (a) Signal location in Phantom data (3D) with different signal size
26 slices with 15 containing activations
- (b) Relative proportion of detection using non adaptive smoothing



- (a) Signal location in Phantom data (3D) with different signal size
26 slices with 15 containing activations
- (b) Relative proportion of detection using non adaptive smoothing
- (c) Relative proportion of detection using Structural Adaptive Smoothing

Finger tapping fMRI data at different resolutions



How to integrate smoothing and testing in on step?

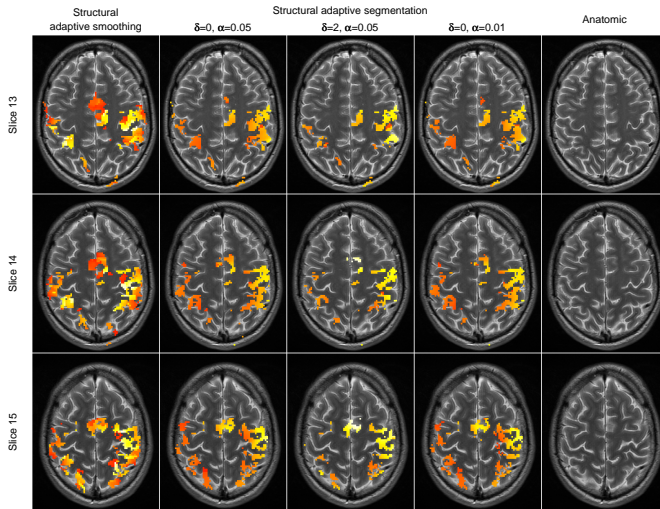
$$H : \max_{i \in V} \gamma_i \leq \delta \quad (\text{or } \max_{i \in V} |\gamma_i| \leq \delta).$$

$$T(\hat{\Gamma}^{\mathcal{H}}) = \max_{h \in \mathcal{H}} \max_{i \in V} \frac{(\hat{\gamma}_i^{(h)} - \delta)}{a_{n(h)}(\nu) \sqrt{\hat{\mathbf{D}} \hat{\gamma}_i^{(h)}} - \frac{b_{n(h)}(\nu)}{a_{n(h)}(\nu)}$$

Structural adaptation

- Distribution of $T(\hat{\Gamma}^{\mathcal{H}})$ under hypotheses by simulation, determine critical values
- Inspect scale space sequentially and segment as soon as critical value is exceeded
- Use adaptive smoothing as long as Hypothesis not rejected

Structural adaptive segmentation: Results



FMRI analysis

```
library(fmri);library(adimpro);

ds <- read.NIFTI("Imagination.nii")
anatomic <- extract.data(ds)[,,1]

scans <- 105
onsets <- c(16, 46, 76)
duration <- 15
tr <- 2
hrf <- fmri.stimulus(scans, onsets, duration, tr)
x <- fmri.design(hrf)
spm <- fmri.lm(ds, x)

spm.segment <- fmri.smooth(spm, hmax = 4, adaptation = "segment")

ds.ana <- read.NIFTI("MPRAGEco.nii")
for (slice in 2:30) {
  img <- plot(spm.segment, ds.ana, slice = slice)
  write.image(make.image(img, gammatype="ITU"), file=paste("result", slice, ".png", sep=""))
}
```

Package fmri

- Structural adaptive smoothing for fMRI (using RFT)
- Structural adaptive segmentation for fMRI
- Package **fmri**: I/O (cf. **oro.nifti**)
- Package **fmri**: Linear Modeling
- Package **fmri**: Smoothing
- Package **fmri**: Signal detection
- Package **fmri**: Publication ready images
- Package **fmri**: No group analysis
- Cf. packages **arf3DS4**, **AnalyzeFMRI**, **neuRosim**, **cudaBayesreg**

Joint Work with:

- Jörg Polzehl and Vladimir Spokoiny, WIAS and MATHEON
- Henning U. Voss, Weill Medical College, Cornell University

Cooperation:

- Citigroup Biomedical Imaging Center, Weill Medical College, Cornell University
- BNIC, Charité, Berlin

R-Community:

- CRAN Task View: Medical Image Analysis

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K. Tabelow, J. Polzehl, H.U. Voss, V. Spokoiny (2006).

Analyzing fMRI experiments with structural adaptive smoothing procedures.

Neuroimage, 33(1): 55–62.



K. Tabelow, V. Piëch, J. Polzehl, H.U. Voss (2009).

High-resolution fMRI: Overcoming the signal-to-noise problem.

Journal of Neuroscience Methods, 178(2): 357–365.



J. Polzehl, H.U. Voss, K. Tabelow (2010).

Structural adaptive segmentation for statistical parametric mapping.

Neuroimage, 52(2): 515–523.



K. Tabelow, J. Polzehl (2011).

Statistical parametric maps for functional MRI experiments in R: The package fmri.

Journal of Statistical Software, 44(11): 1–21

Special Volume of *Journal of Statistical Software* “MRI in R”.