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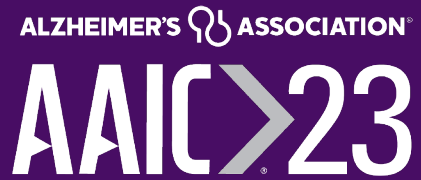
AAIC>23

ALZHEIMER'S ASSOCIATION INTERNATIONAL CONFERENCE®

JULY 16-20 > AMSTERDAM, NETHERLANDS, AND ONLINE

# ISTAART Neuroimaging PIA THE BASICS OF NEUROIMAGING SEMINAR SERIES





# BASICS OF NEUROIMAGING: FUNCTIONAL MRI

Luigi Lorenzini, PhD student  
Amsterdam UMC

Available on demand very soon!

## The Basics of Neuroimaging

Data Structure and Formats

**Moderator:**

Alexis Moscoso Rial, PhD

**Speaker:**

Ludovica Griffanti, PhD

Wednesday, April 5, 9 a.m. CT

## The Basics of Neuroimaging

Structural Magnetic Resonance Imaging (MRI)

**Moderator:**

Tavia Evans, PhD;  
Erasmus MC, Netherlands

**Panelists:**

David Cash, PhD;  
University College London, United Kingdom

Friday, April 14, 9 a.m. CT

## The Basics of Neuroimaging

Positron Emission Tomography (PET)

**Moderator:**

Lyduine Collij, Ph.D.

**Panelists:**

Tobey Betthausen, Ph.D.

Wednesday, April 19, 12 p.m. CT

## The Basics of Neuroimaging

Diffusion-Weighted Imaging (DWI)

**Moderator:**

Tom Veale, Ph.D.

**Panelists:**

Alexa Pichet Binette, Ph.D.

Friday, April 21, 9 a.m. CT

ADD MINE

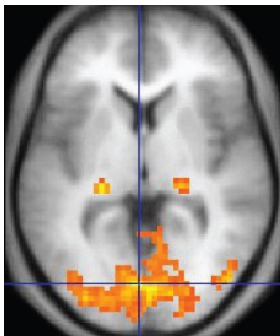
By the end of this session, you should be able to:

- Understand and discuss fMRI principles and measurements
- Outline the basic preprocessing steps needed fMRI data and typical issues
- Describe the relevance and current clinical application of fMRI



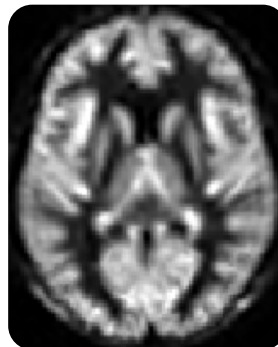
## Functional MRI: What are we measuring?

fMRI



BOLD signal

ASL



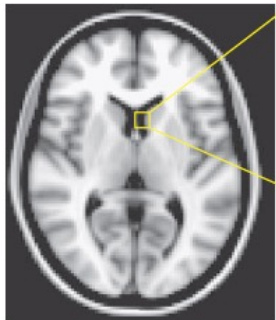
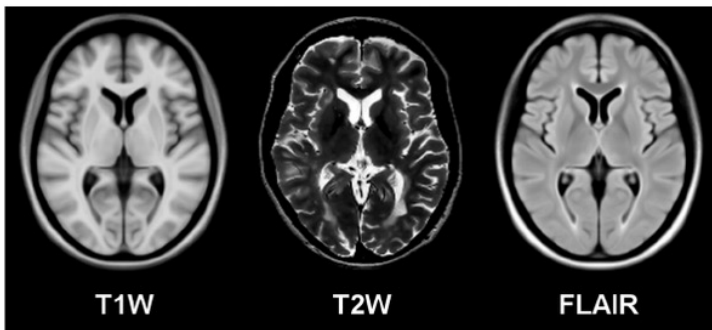
Cerebral blood flow (CBF)

EEG



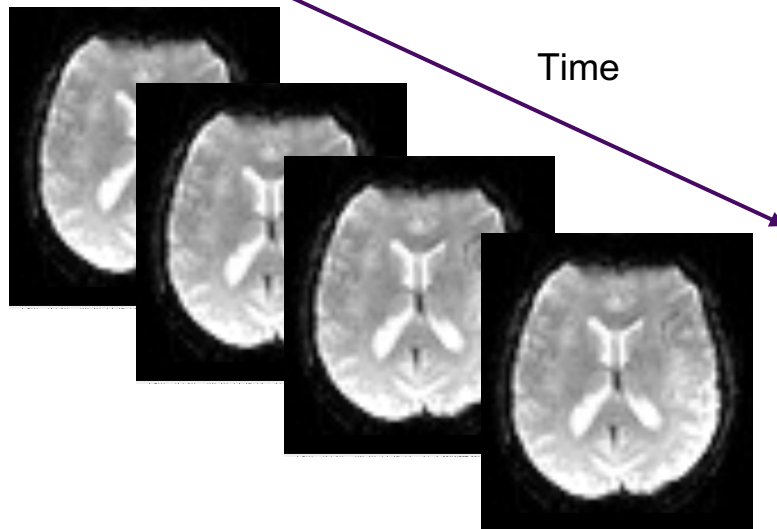
Electrical Signals

## Structural MRI

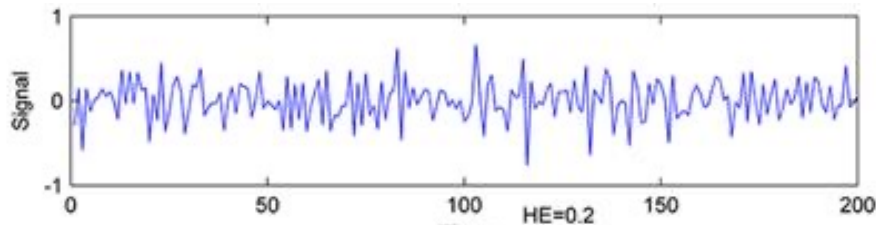


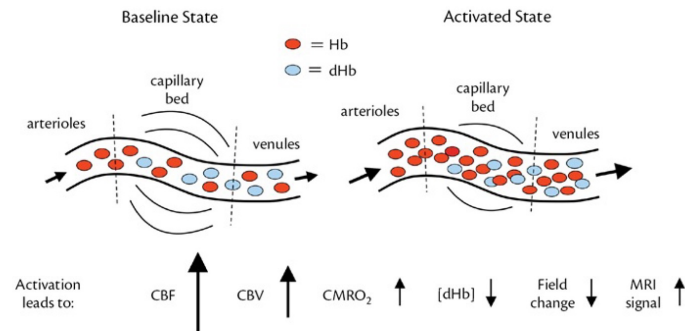
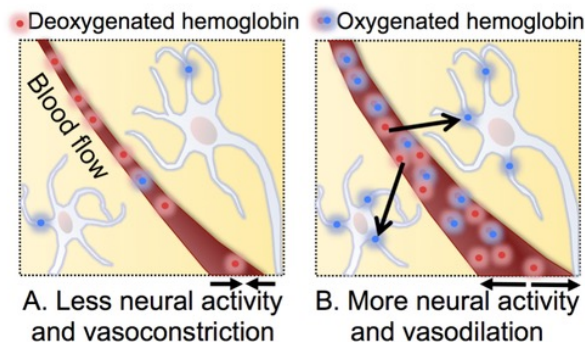
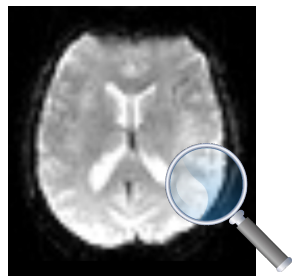
288	27	38	364	621
264	21	97	500	640
271	22	133	543	647
312	28	113	521	649
390	53	58	424	635

## Functional MRI



Frames = Volumes  
 TR = ~3 sec  
 T2\* contrast:  
 sensitive to BOLD



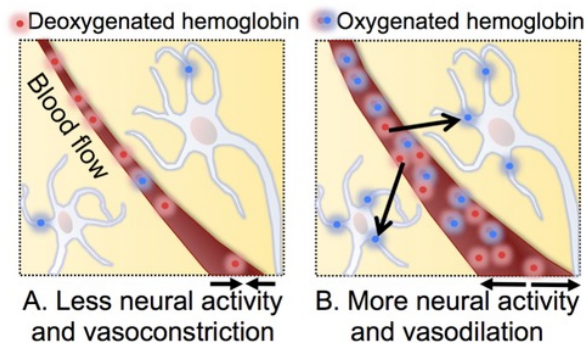
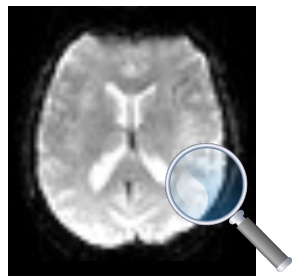
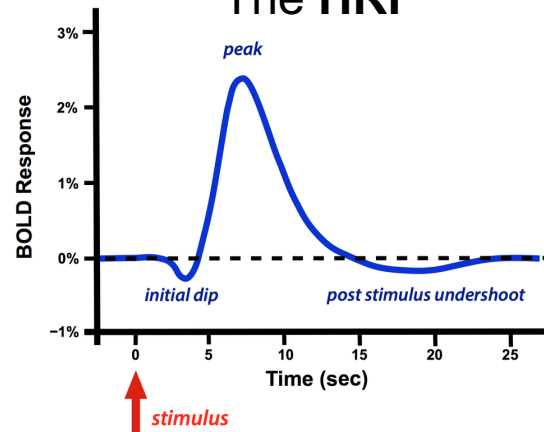
The **BOLD** SignalBlood oxygen level dependent (**BOLD**) signal

- Active neurons require oxygen! (action potentials are expensive)
- Blood levels (CBF) increase, capillaries dilate, to supply oxygen and glucose to activated neurons
- Unbalance between oxygenated and deoxygenated hemoglobin → BOLD
- The deoxygenated hemoglobin disturbs the local magnetic field
- Higher BOLD → More oxygenated hemoglobin → High activity



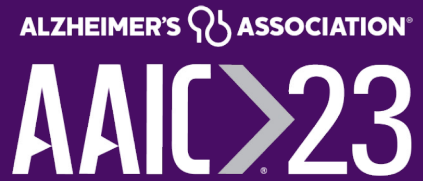
BOLD is an indirect measure of brain activation



The **BOLD** SignalThe **HRF**

## The hemodynamic response function (HRF)

- “Ideal” BOLD response function
- Peak around 6 seconds, take ~20 to baseline
- Low temporal resolution, difficult to know the exact time of neuronal changes
- However, good for image acquisition (no need to acquire 1 image every millisecond)



# Functional MRI: Basic Pre-Processing

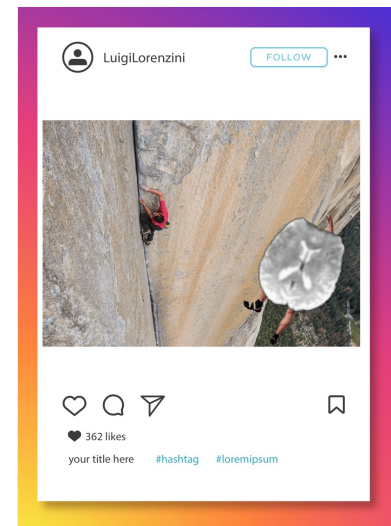
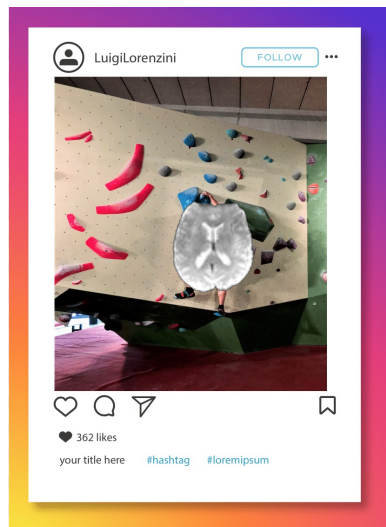
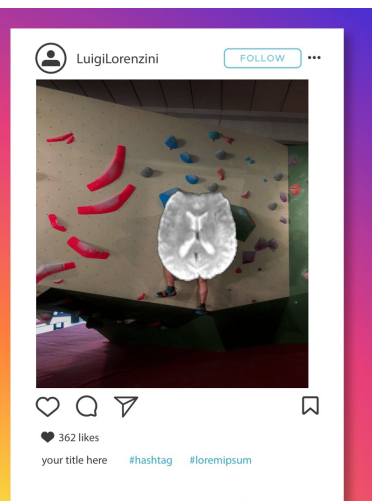
1. fMRI data are prone to a number of artifacts and sources of variability
2. Raw images are not usable for direct inspection or statistical analysis



Pre-processing



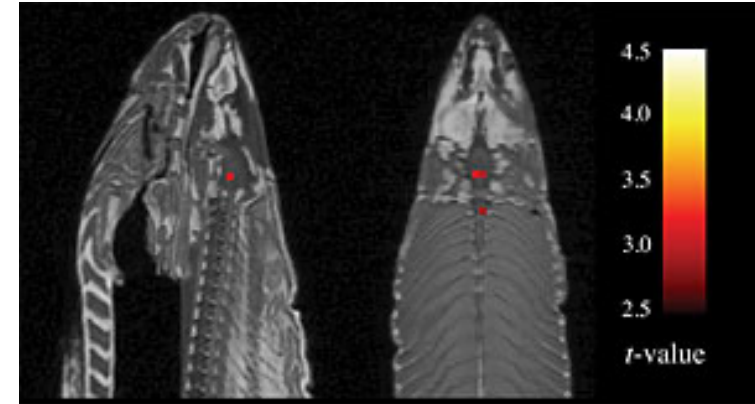
Extreme  
Pre-processing



1. fMRI data are prone to a number of artifacts and sources of variability
2. Raw images are not usable for direct inspection or statistical analysis

### What a dead salmon tells us about fMRI pre-processing and analysis

*“shown a series of photographs depicting human individuals in social situations. The salmon was asked to determine what emotion the individual in the photo must have been experiencing.”*



We want to clean “NOISE” and enhance our effect of interest

## What

A series of steps used to

1. Remove unwanted signal fluctuations and artefacts
  2. Clean desired effects
  3. Standardize data
- Before Statistical analysis

Distortion Correction

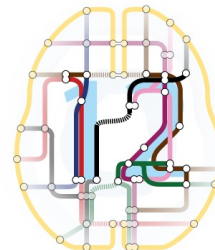
Motion Correction

Standard Space Mapping

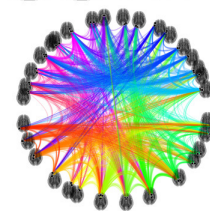
Spatial Smoothing

Temporal Filtering

## How

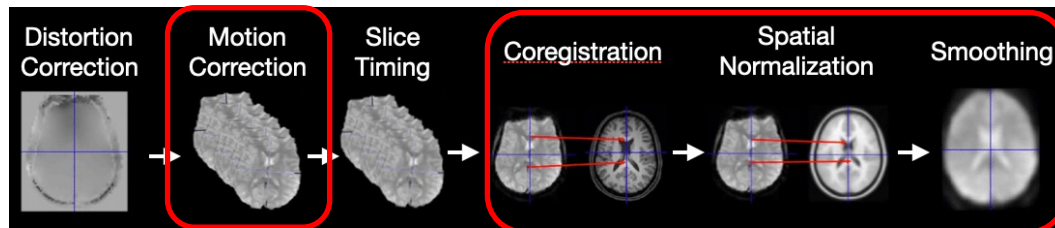


CONN



fMRIPrep

a robust preprocessing pipeline for task-based and resting-state fMRI data



### Variability in the analysis of a single neuroimaging dataset by many teams

Rotem Botvink-Nezer, Felix Holzmeister, ... Tom Schonberg [+ Show authors](#)

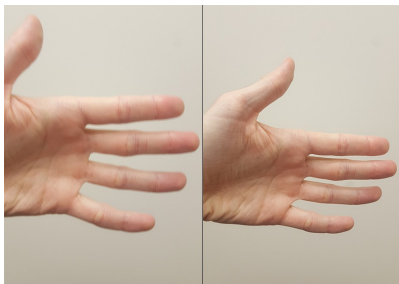
*Nature* 582, 84–88 (2020) | [Cite this article](#)

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# AAIC>23 MOTION CORRECTION

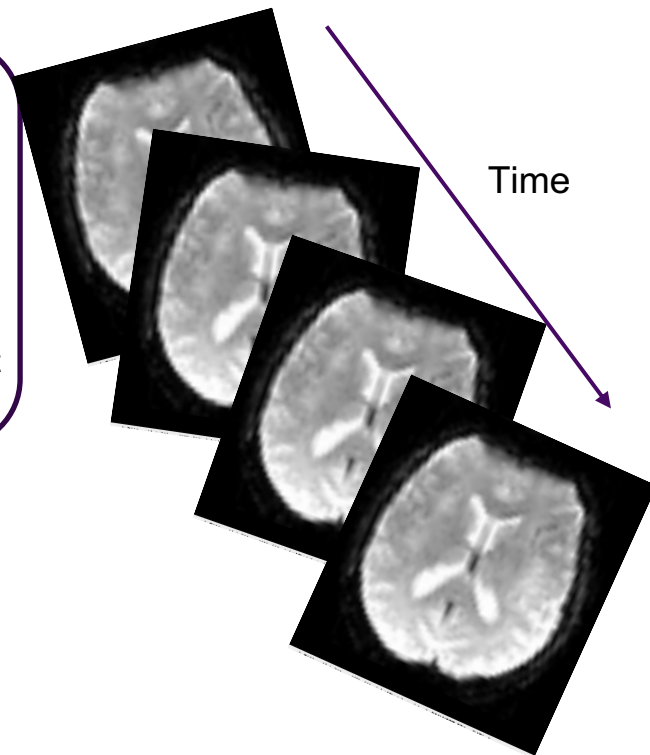
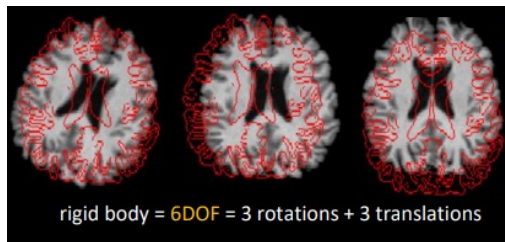
## Why

- As for real pictures, a moving target will look blurry
- We measure from moving voxels (different across the session)
- Introduce confounds (e.g. in response to a stimulus/ whole-brain correlations) that are stronger than physiological changes



## How

- Align all the volumes from a timeseries with a reference volume
- Usually, the reference volume is the first, middle or last of the TS
- Use rigid body transformation (6 DOF)
- Iterative process of finding the best alignment between 2 volumes (cost function)



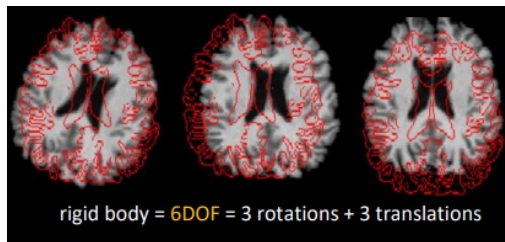
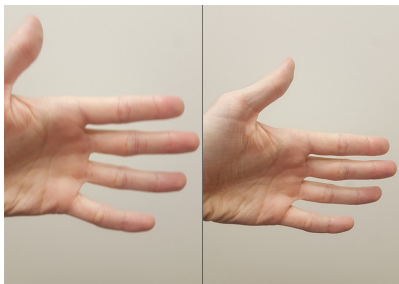
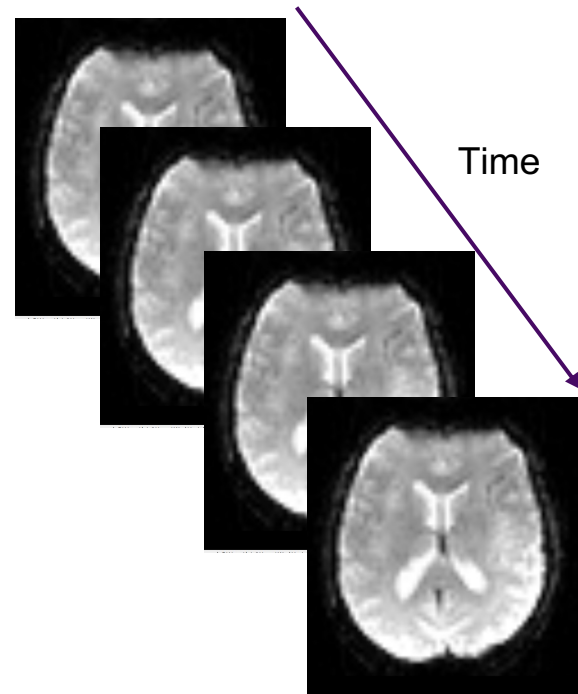
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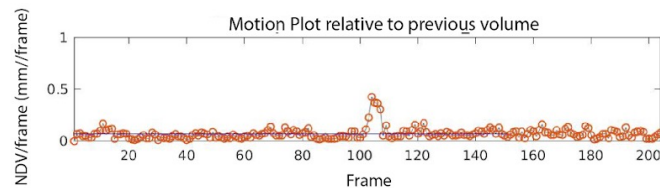
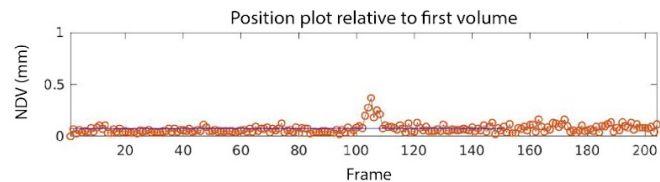
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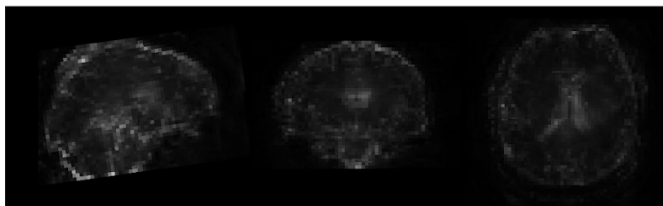


MOTION CORRECTION: MOTION  
PARAMETERS

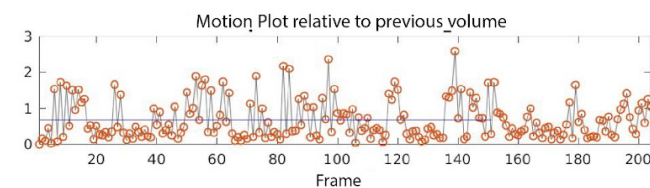
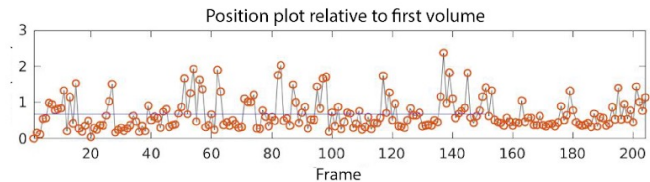
"Low-Motion rs-fMRI scan"



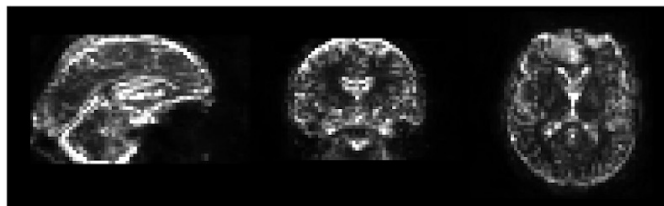
Standard Deviation Image



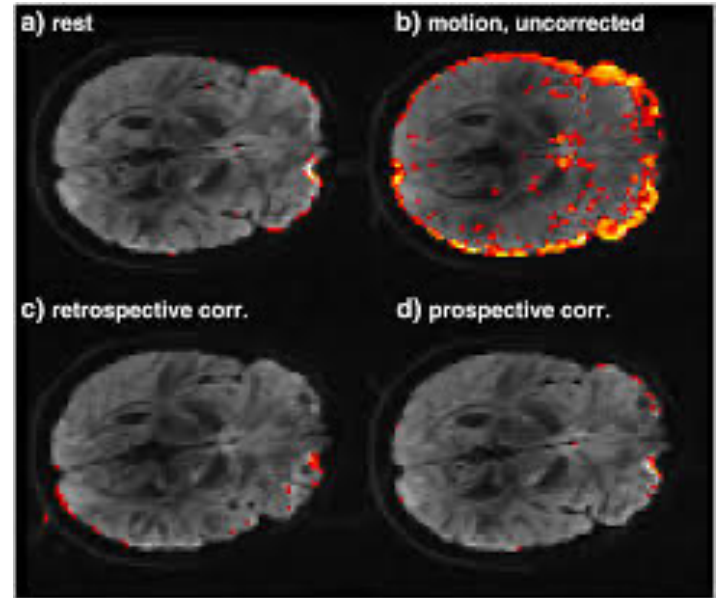
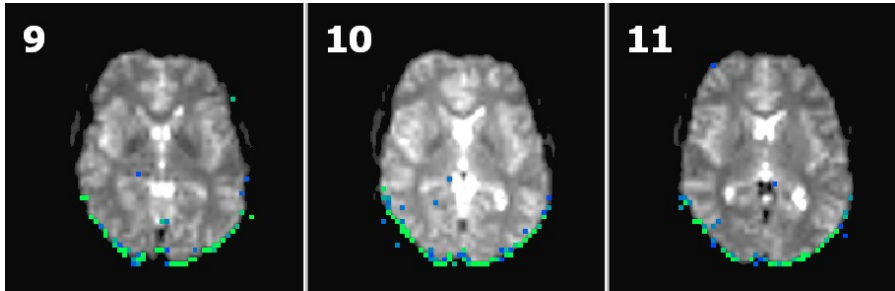
"High-Motion rs-fMRI scan"



Standard Deviation Image





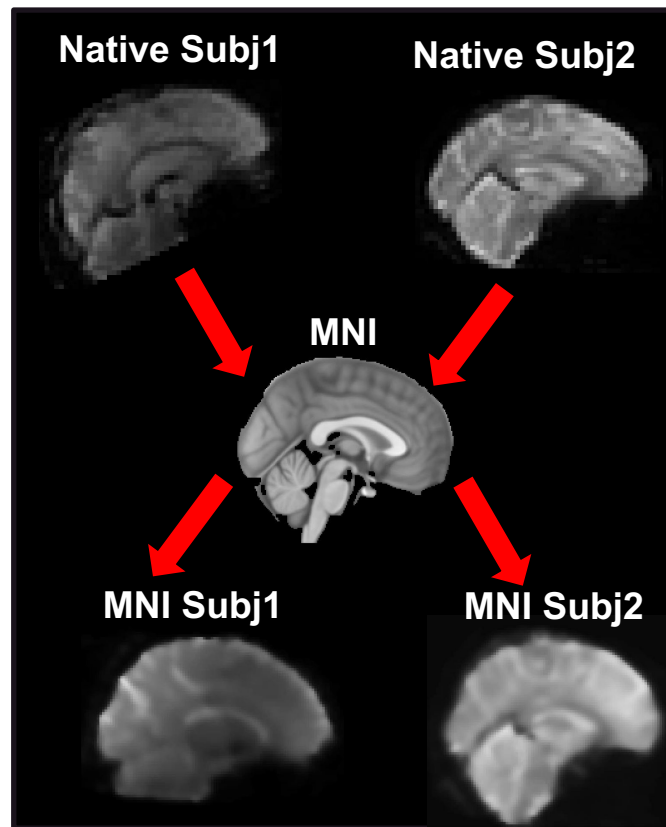
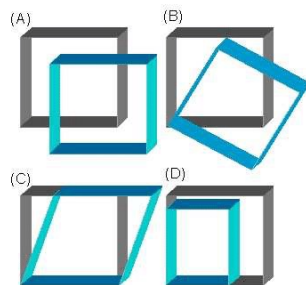
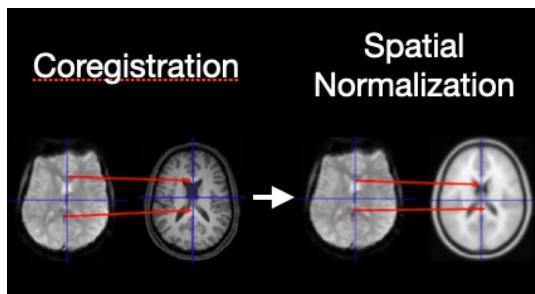


## Why

- Differences in brain size and shape exist between different individuals.
- For group analysis, we need voxels between different brains to correspond
- Registration or Normalization or Standard Space Mapping

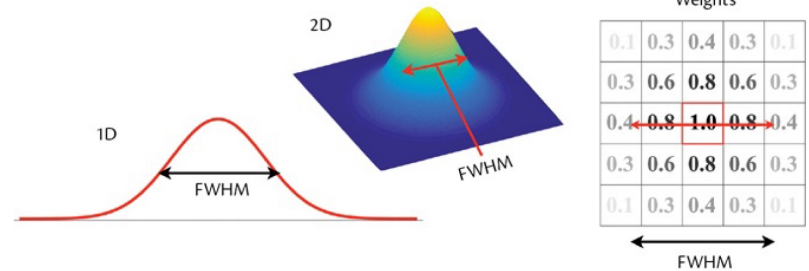
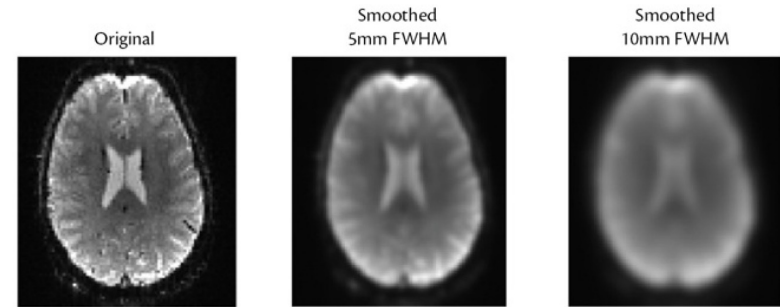
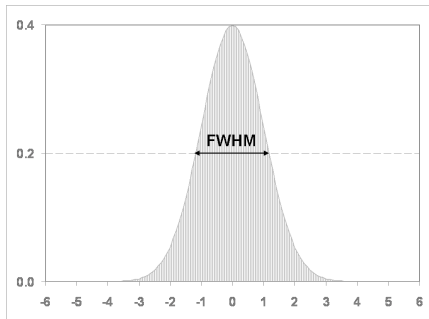
## How

- Affine Transformation, similar to rigid body but 12 DOF (allows zooms and shears)
- We register to the anatomical T1 (coregistration), previously registered to MNI space (normalization)
- Then we can apply the T1 -> MNI transformation to our fMRI



# AAIC>23 SPATIAL SMOOTHING

- Smooth functional data = replace the value at each voxel with a weighted average of that voxel's neighbors
- Lower resolution? Yes. But also, greater signal to noise!
- High frequencies of the signal are removed while enhancing low frequencies
- Gaussian kernel of specific width (FWHM) determines the amount of smoothing



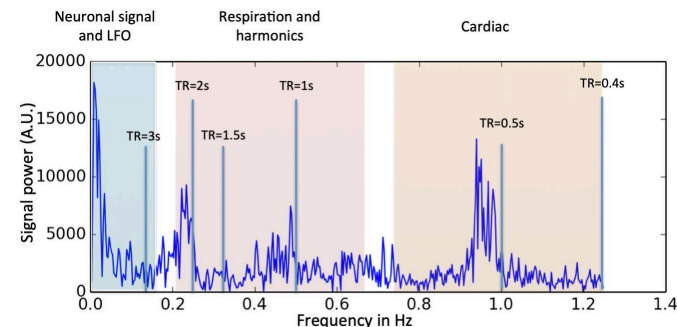
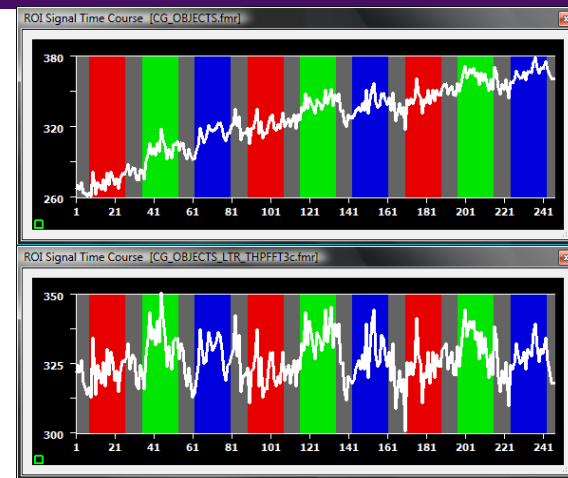
- Remove noise based on its frequency
- Low-frequency drifts: due to both physiological and physical (scanner-related) noise
- Linear and non-linear drifts
- Voxel's timecourses represented as frequency domain (e.g. Fourier transform), low drifts are set to 0

#### Examples of Scanner-related Noise:

- Hardware imperfections
- Heating of components

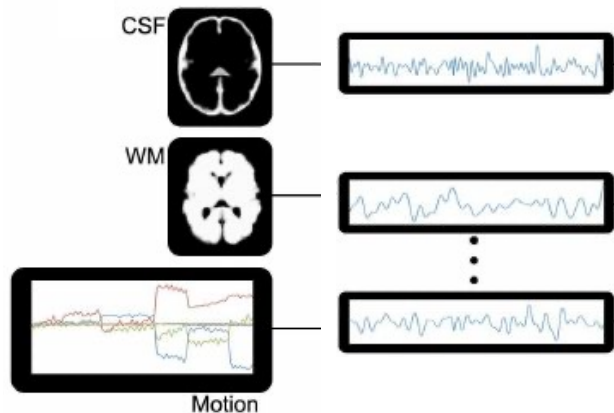
#### Examples of Physiological Noise:

- Cardiac pulsations
- Respiratory Cycle

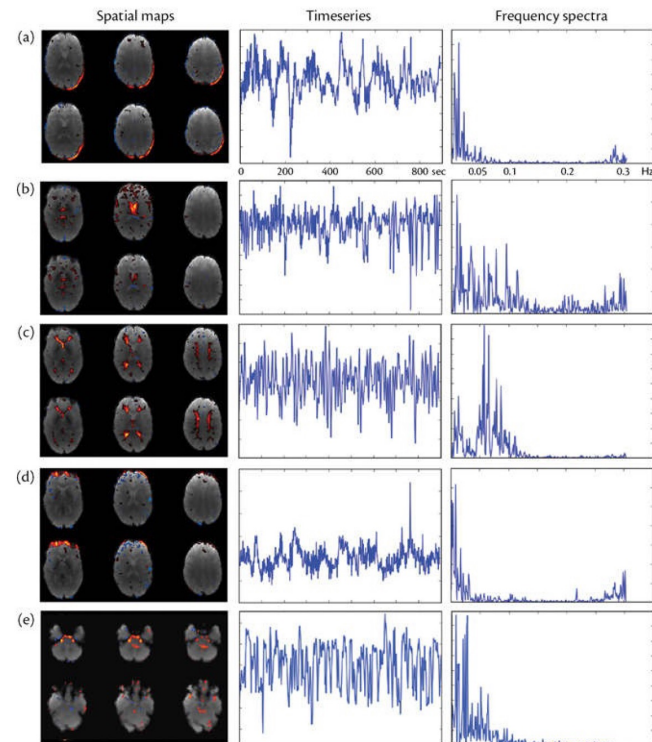


FILTERING OF PHYSIOLOGICAL NOISE:  
STATISTICAL APPROACHES

## GLM approach



## ICA approach

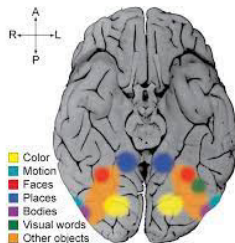
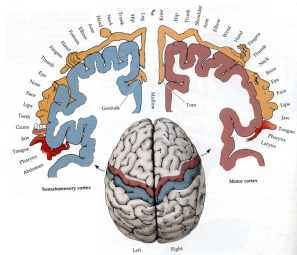




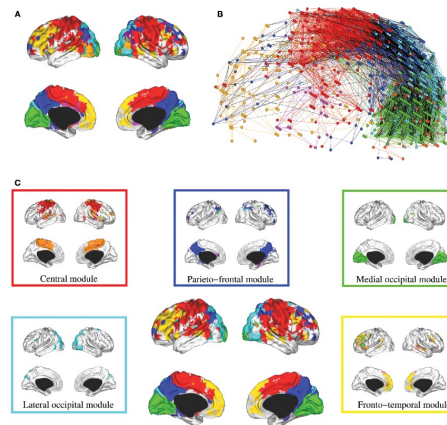
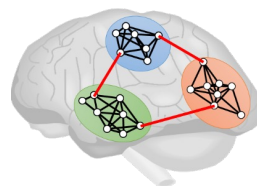
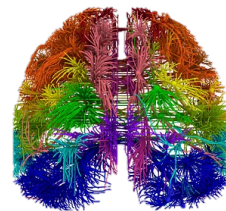
# Functional MRI: Derived data and applications



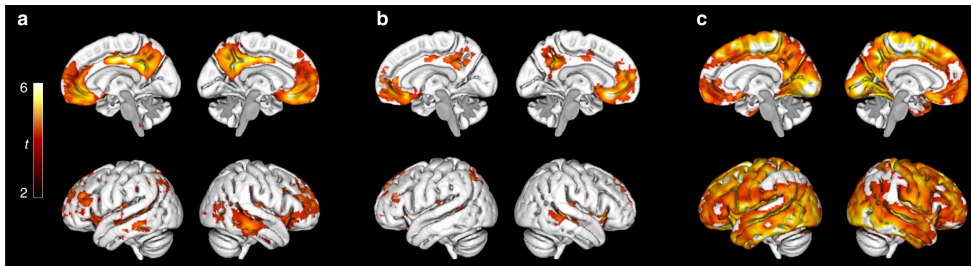
## Functional Localization



## Brain Networks

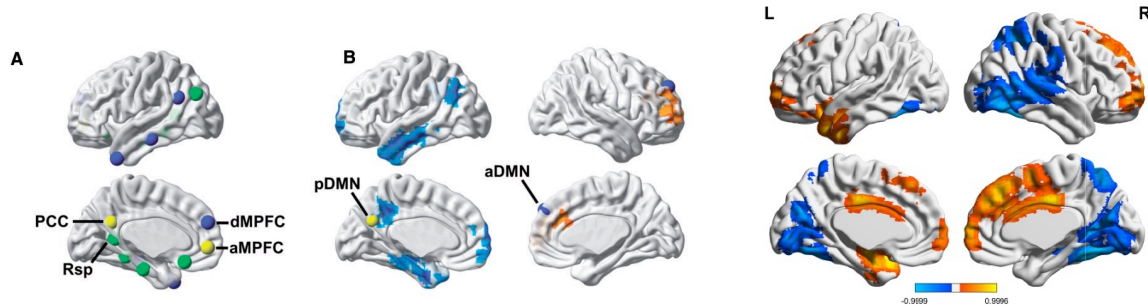
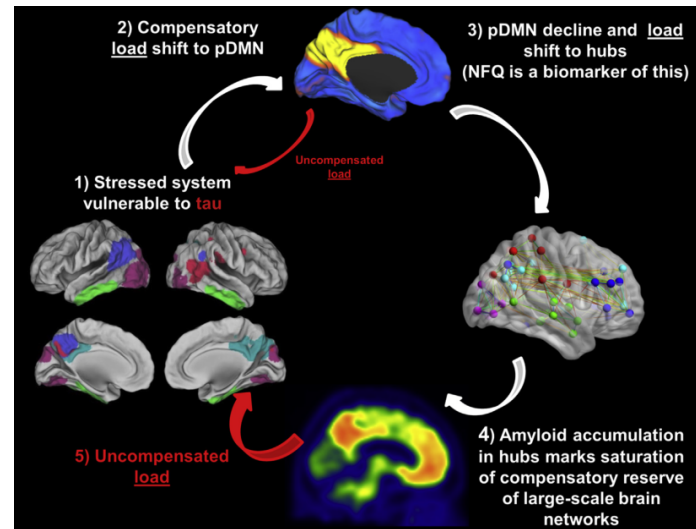


Meunier, D. et al (2019)



*Palmqvist, et al. (2017).*

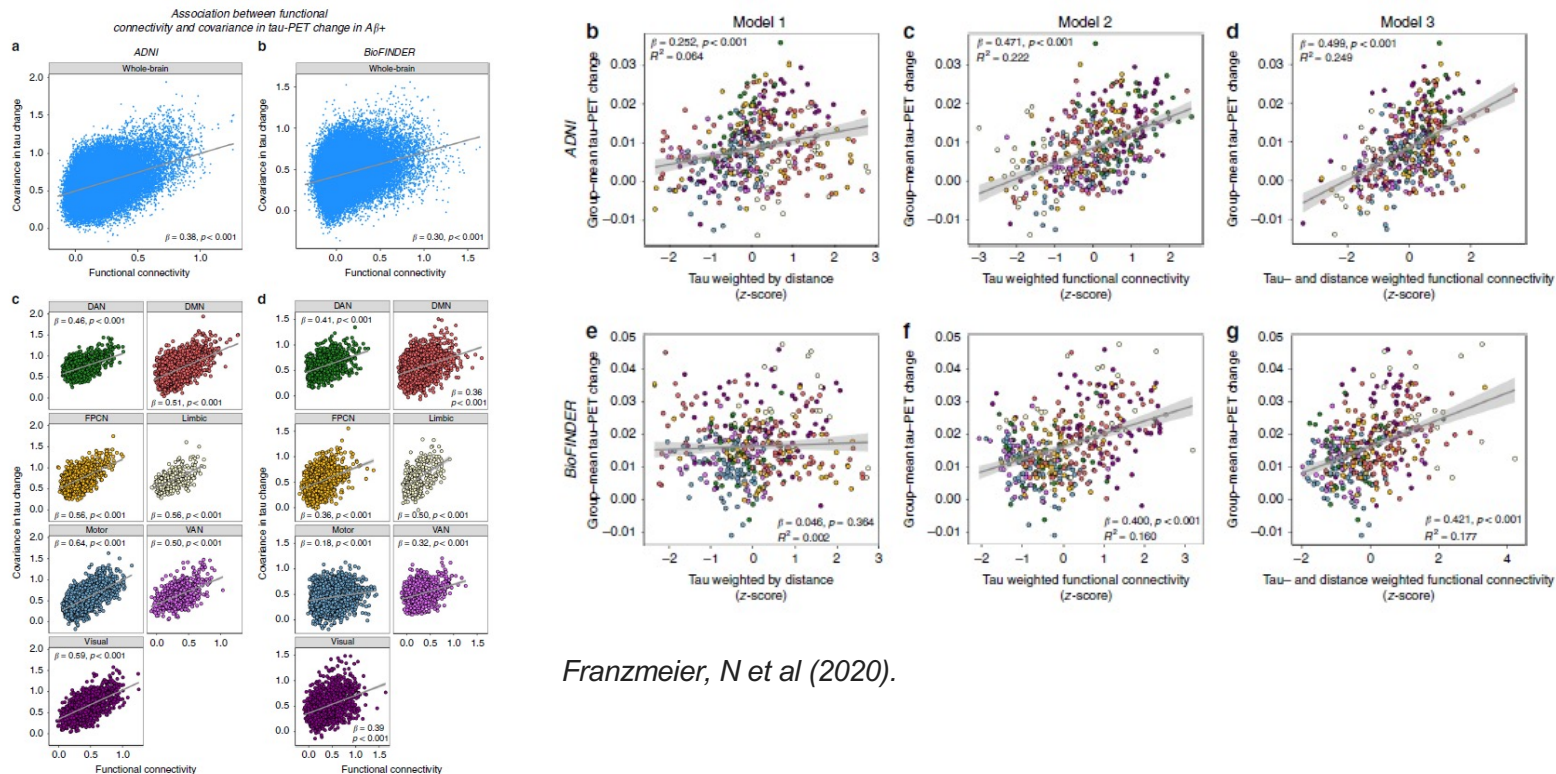
## Network failure cascade model



*Jones, D. et al. (2016).*

*Lorenzini et al. (2022)*



SPREADING OF PATHOLOGICAL PROTEINS  
ACROSS FUNCTIONAL CONNECTIONS

*Franzmeier, N et al (2020).*

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POP QUIZ!

fMRI motion correction works by:

- a) Aligning volumes between two fMRI time series
- b) Aligning volumes within on fMRI time series
- c) Registering the fMRI scan to a structural scan

Spatial smoothing:

- a) Decrease signal to noise ratio
- b) Has no effect on signal to noise ratio
- c) Increase signal to noise ratio

Which one of these statements is **false** about BOLD signal:

- a) It's a measure of neuronal electrical activity
- b) It's based on the unbalance between oxygenated and deoxygenated hemoglobin
- c) Measures brain hemodynamic response

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JULY 16-20 > AMSTERDAM, NETHERLANDS, AND ONLINE

**Thank You!**

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Amsterdam UMC,  
Department of Radiology and Nuclear Medicine  
[l.lorenzini@amsterdamumc.nl](mailto:l.lorenzini@amsterdamumc.nl)