

# BIOMEDICAL DATA FUSION

## using tensor-based blind source separation



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10-9-2018

# Contents Overview



## 1. Introduction

- Keytool: Blind Source Separation
- Biomedical Data fusion: Applications
- Tensor Decompositions



## 2. BIOTENSORS Project



## 3. Examples



## 4. Conclusions and Future Directions

# KEYTOOL : Blind source separation

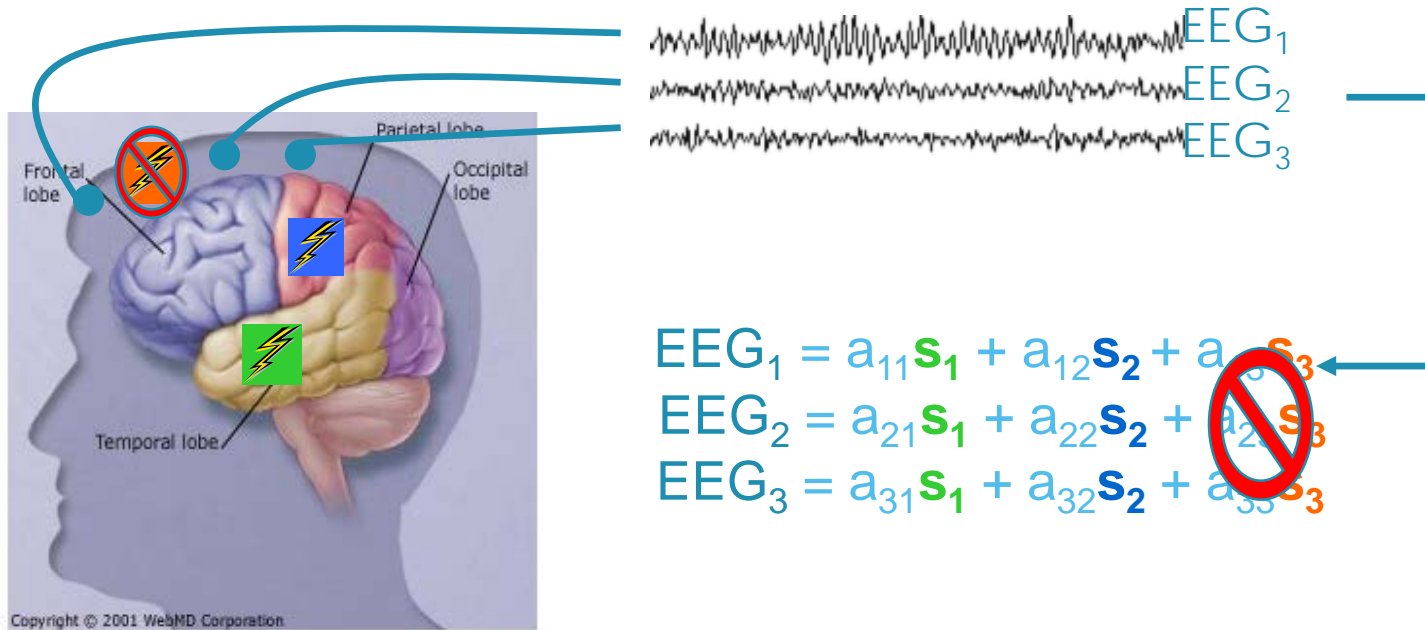
Signal analysis difficult because of artefacts → REMOVE

## Matrix based Blind Source Separation (BSS)

- **Non-unique** → Constraints are needed (orthogonal, independency)

## TENSOR based BSS: unique under mild conditions

ADD extra problem-specific constraints (nonnegative, sparse)



[illegible]


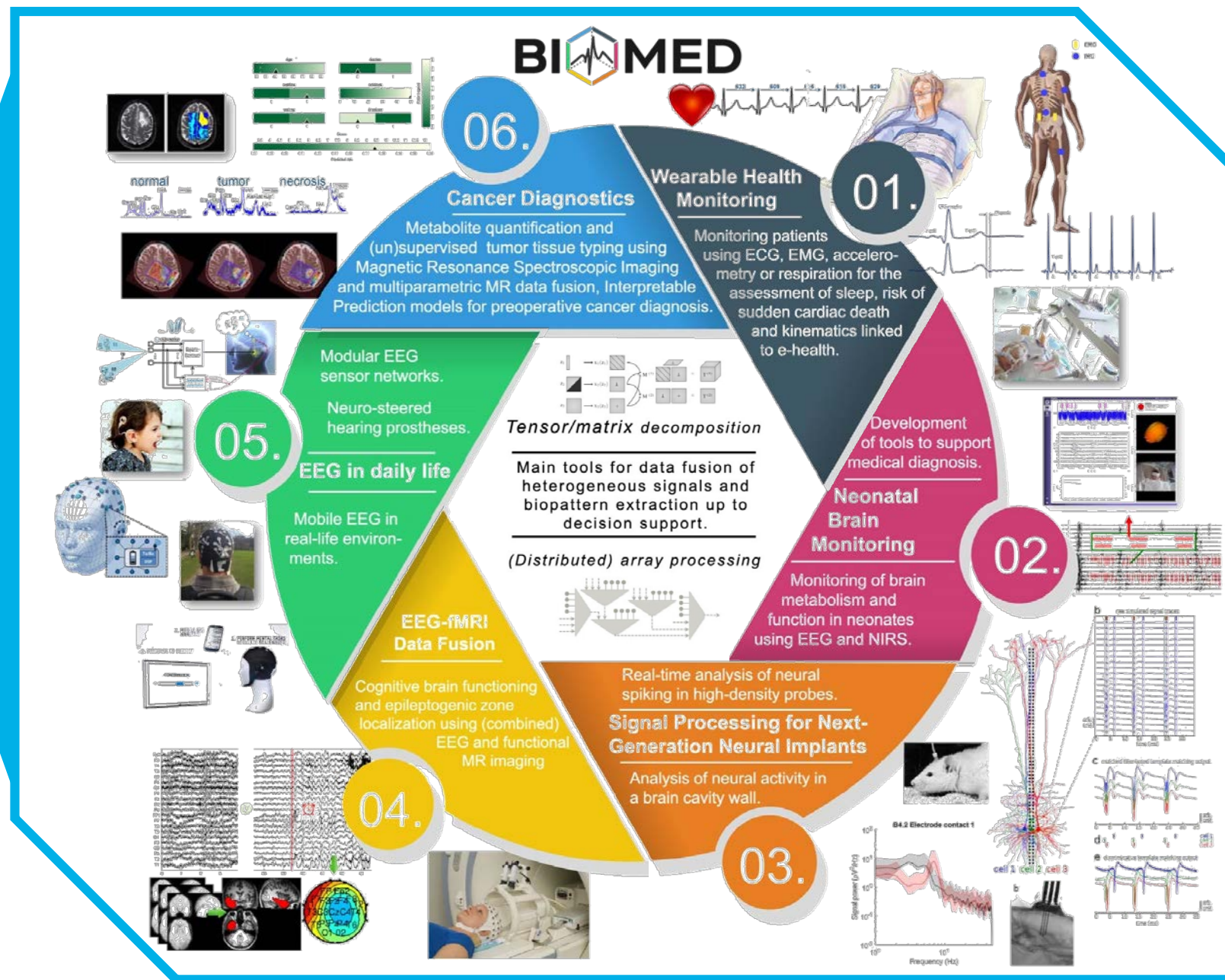
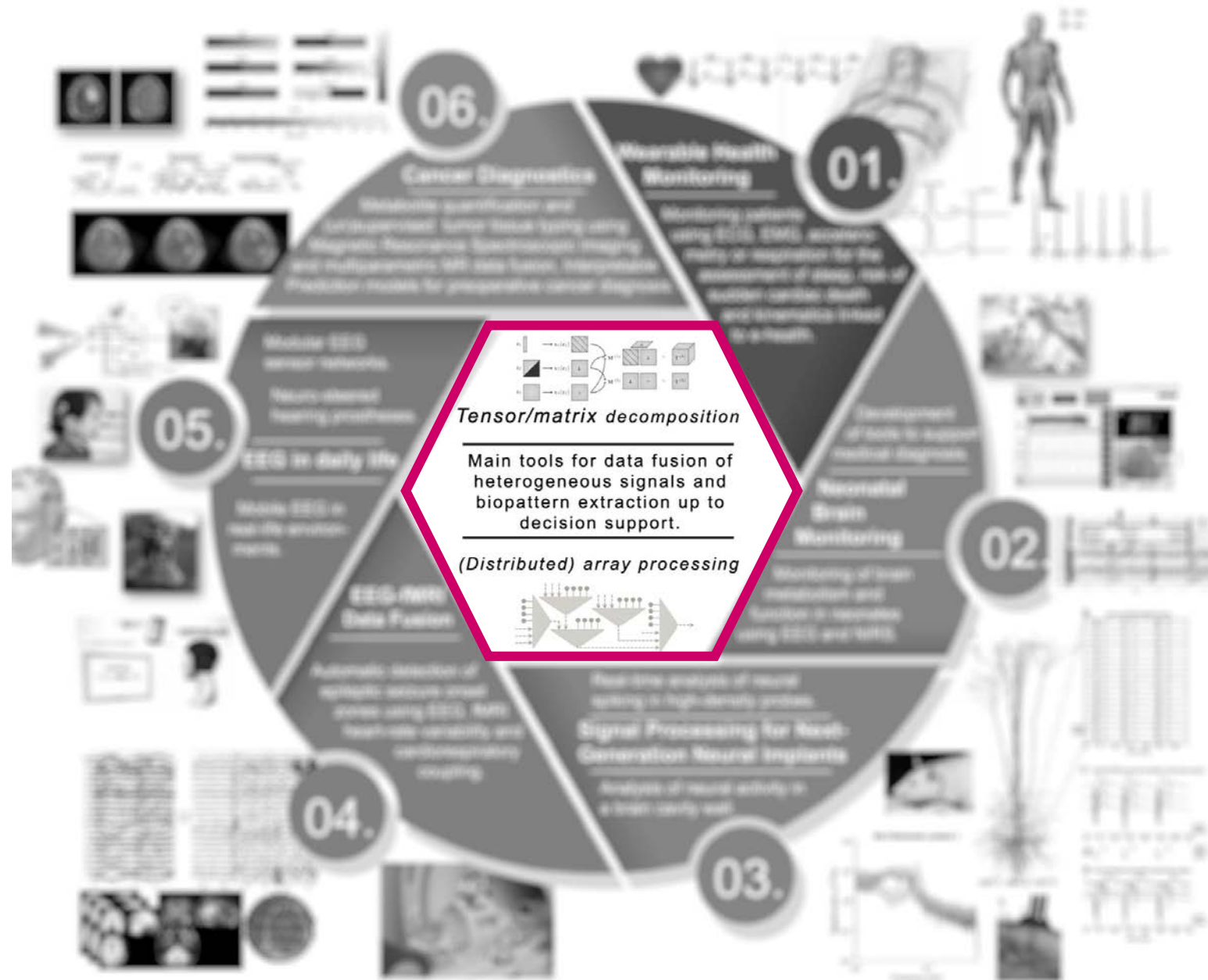


Diagram illustrating the relationship between EEG, A, and  $S^T$ :

$$\text{EEG} = A \quad S^T ?$$

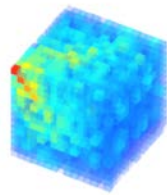
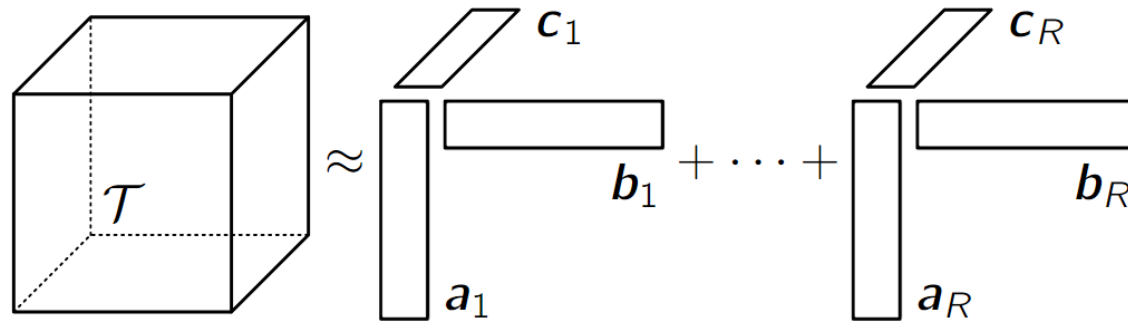






# Tensor Decompositions:

## Canonical Polyadic Decomposition - CPD

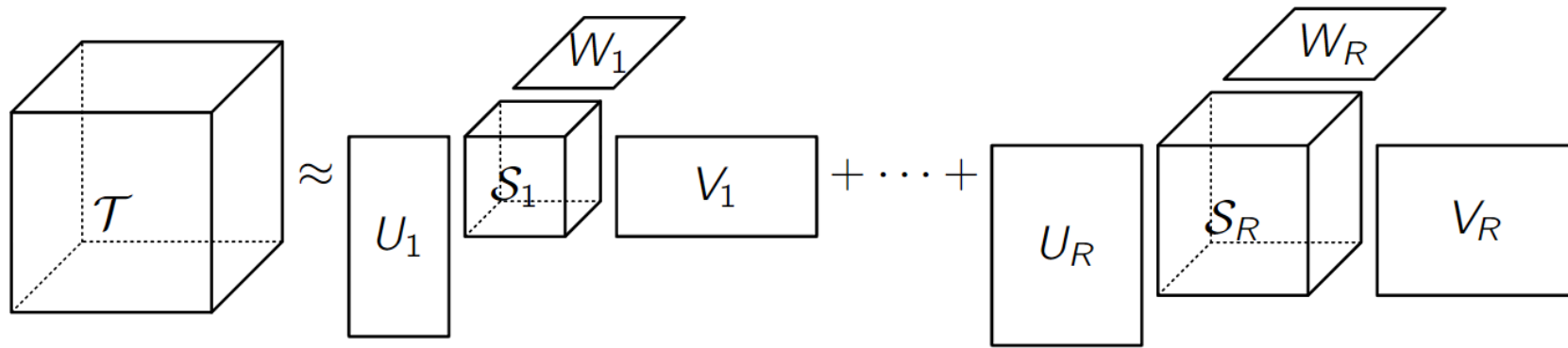


# Tensorlab

A MATLAB package for  
tensor computations.

[www.tensorlab.net](http://www.tensorlab.net)

# From CPD to Block Tensor Decomposition



De Lathauwer et al., SIMAX, 2008; Sorber et al., SIOPT, 2013

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**2. BIOTENSORS Project**



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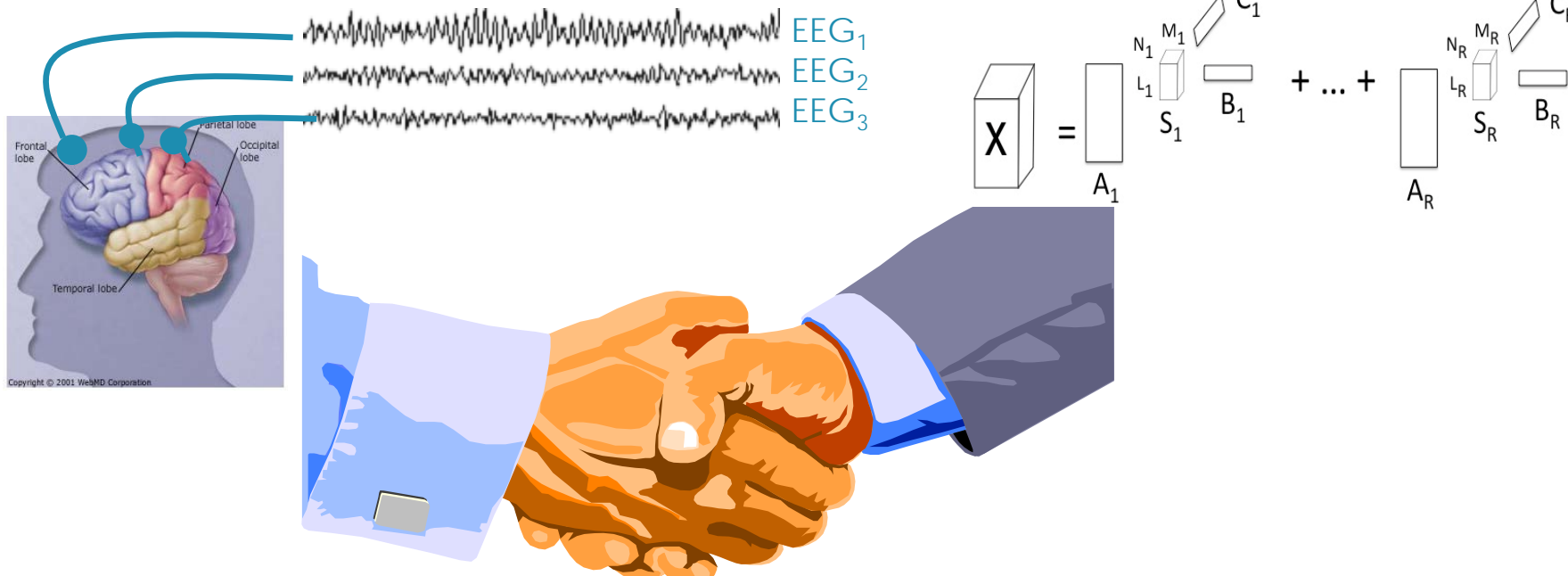


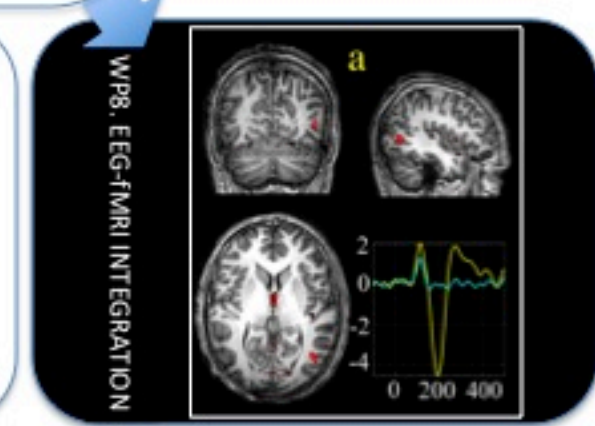
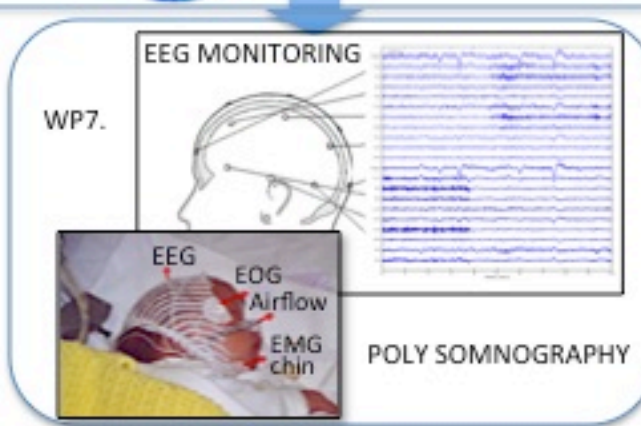
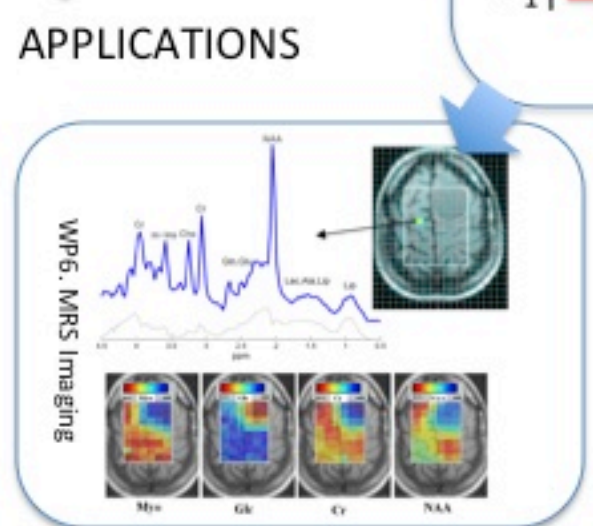
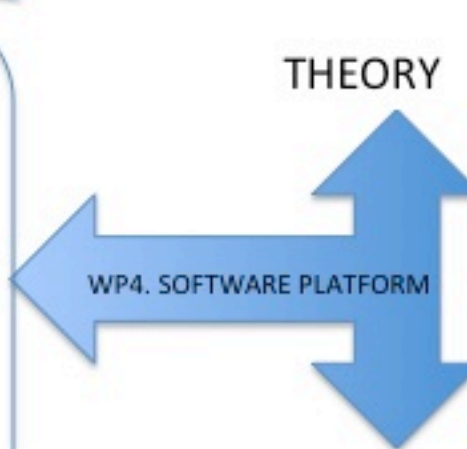
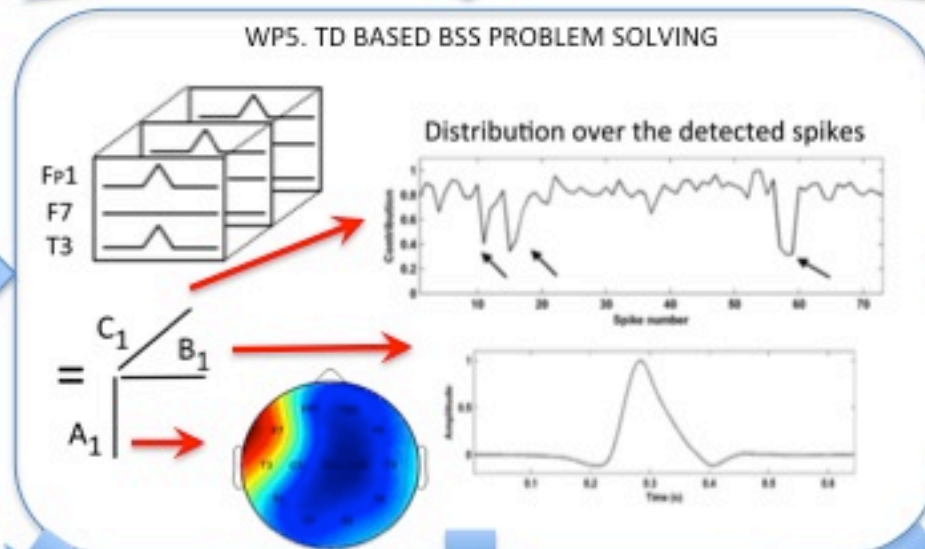
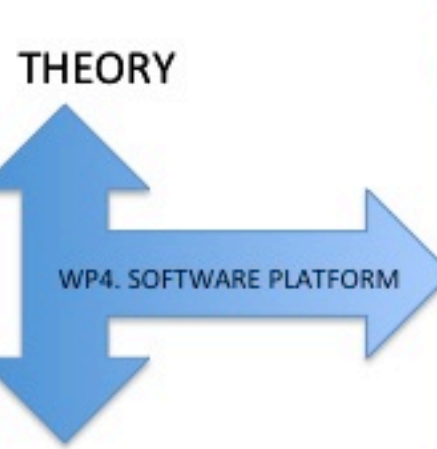
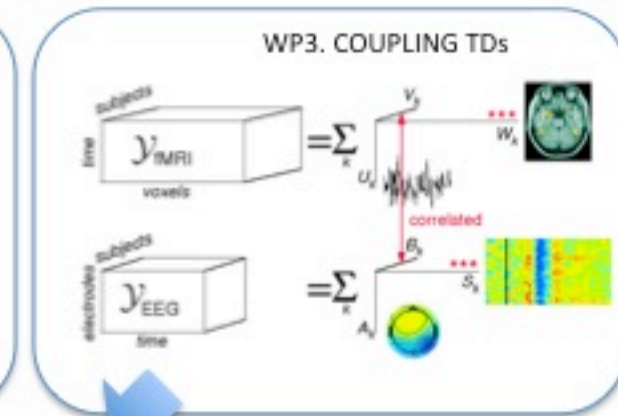
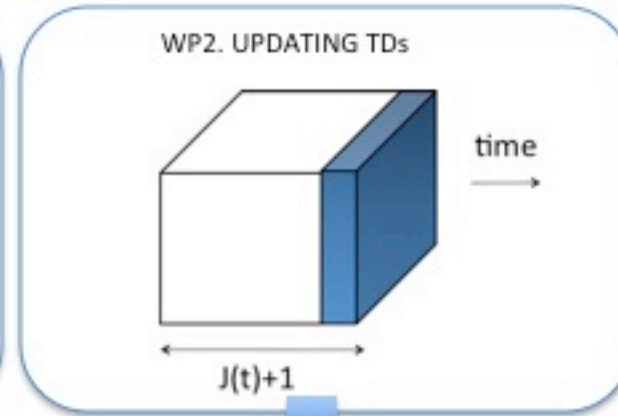
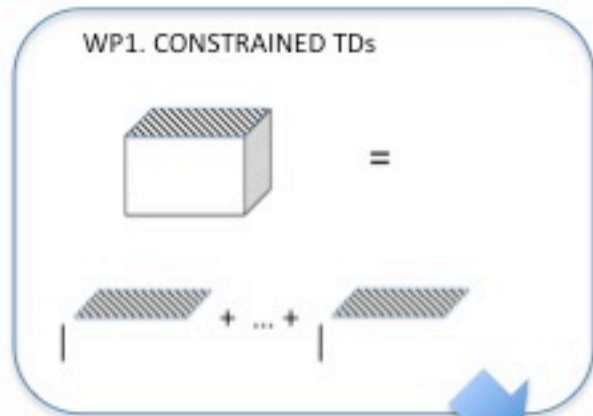
# Birth of BIOTENSORS

- > 1982: Advanced (multi)linear Algebra as CORE in SISTA (later: SCD, now: Stadius)
- > 1990: Birth Biomedical Data Processing Research in SISTA (HEADED by Sabine VH)
- > 1992: Birth MULTilinear Algebra Research in SISTA (HEADED by Lieven DL)

## ➤ **BIOTENSORS: Joining expertise of Lieven and Sabine**

- Idea: Christmas 2011
- ERC Adv.grant: Submitted (Feb & Nov 2012) → Accepted (July 2013) → Start (April 2014)





## Task 1.1 Basic algorithms

- *High-performance algorithms to solve Linear Systems with Kronecker Product-constrained solution (e.g. LS-CPD), providing broad framework for analysis of multilinear systems.*
- *Relaxed conditions (compared to Kruskal) for generic uniqueness proven for several tensor decompositions, including block terms, constraints and coupling. Extensions to missing fibers.*
- *New algorithms for the exact computation of the decomposition of a tensor into a sum of multilinear terms. Extended to CPD of large-scale tensors with missing fibers.*
- *Tensor optimization, improving computationally global minimization in line and plane search subproblems*
- *Exploiting sparsity, low-rank properties and incompleteness, allows decomposition of very large-scale tensors and even break the curse of dimensionality:*
  - *Randomized block sampling enabling decomposition of very large-scale tensors (e.g.  $10^{18}$  entries) by sampling very few entries (e.g.  $10^5$ ).*
  - *Numerical framework exploiting efficient tensor representation enable large computational gains.*
- *Efficient algorithm for weighted CPD using low-rank weights*

## Task 1.2 Constraints

- *New constraints embedded in tensor decompositions implemented: including nonnegativity, orthogonality, structure (Vandermonde, Kronecker, Khatri-Rao, exponential, Cauchy,...) and finite differences*

## Task 1.3 Source modeling

- *Exploiting Exponential polynomial model ( $\Sigma$  and/or  $\Pi$  of exponentials, sinusoids, polynomials), Rational model (Löwner matrices) and sparseness (compressive sampling)*
- *Segmentation is novel tensorization technique: useful for large-scale (instantaneous) blind source separation and (convolutive) blind system identification. It exploits property that sources/inputs and/or mixing vectors/system coefficients are modelled as low-rank matrices or tensors.*
- *New signal separation techniques presented in tensor framework: including exponential polynomials, rational functions and Kronecker-product structured sources. Corresponding structured tensor algorithms (CPD, Tensor Train, Hankel, Löwner...) optimized in speed and data storage (no need to expand to full tensor).*
- *Löwner-based BSS technique developed to enable deterministic separation of signals into rational functions with applications in fetal ECG extraction and water removal using MRSI.*
- *Computationally efficient algorithms developed for tensor-based convolutive signal separation.*
- *Methods comparing tensor factors without computing the decompositions have been developed*

**Contributors: Ignat Domanov, Otto Debals, Mikael Sorensen, Xiao-Feng Gong, Martijn Boussé, Paul Smyth, Frederik Van Eeghem, Marco Signoretto, Chuan Chen, Alwin Stegeman, Nico Vervliet, Michiel Vandecapelle**



# WP2: Updating tensor decompositions

## Task 2.1 Updating in one dimension

- *NLLS algorithm developed for CPD updating of Nth order tensor with(out) low-rank weighting of tensor entries.*  
→ *Outperforms state-of-the-art algorithm from Nion and Sidiropoulos (IEEE TSP 2009)*

## Task 2.2 Updating in several dimensions

- *NLLS Algorithm generalized to allow updating in any number of dimensions*

**Contributors: Geunseop Lee, Michel Vandecappelle, Nico Vervliet**

# WP3: Coupling tensor decompositions

## Task 3.1 Algorithms

- *Coupled CPD models extended to coupled multilinear rank- $(L_{r,n}, L_{r,n}, 1)$  terms with proven relaxed uniqueness conditions and allowing algebraic computation*
- *Coupled CPD modeling framework developed for solving Multidimensional Harmonic Retrieval problem and the Gaussian mixture parameter estimation.*
  - *Uniqueness conditions are most relaxed ones.*
  - *Very promising in sensor array processing enabling to exploit multiple spatial sampling structures (in contrast to ordinary CPD models)*
- *Algebraic Double Coupled-CPD algorithm presented based on a coupled rank-1 detection mapping for joint BSS, outperforming standard CPD based BSS methods*

## Task 3.2 Coupling constraints

- *Tensor lab introduces Domain specific language (DSL) to easily represent various couplings and facilitate creation of models with approx. equal factor matrices*

**Contributors: Lieven De Lathauwer, Laurent Sorber, Mikael Sorensen, Ignat Domanov, Frederik Van Eeghem, Nico Vervliet**

# WP4: Software platform for tensor-based BSS

*Powerful software tools allow to face current grand challenges in biomedical data fusion*

## Task 4.1 General purpose tensor toolbox

- *Algorithms in WP1-3 efficiently implemented in Tensorlab 3.0 and 4.0.*
- *Tensorlab allows to decompose structured tensors directly, avoids full tensor expansion*  
→ *very useful in tensorizations!*
- *Improved user friendliness by simplifying model construction and adding visualization routines, documentation and demos.*
- *Matlab-based GUI facilitates visual inspection of 3<sup>rd</sup> order tensor CPD*

## Task 4.2 Software platform for tensor-based biomedical source separation

- *Platform accessible to experts and non-experts (medical doctors)*
- *Integrate keytool Matlab implementations of WP6-8 as **easy-to-use toolboxes with GUI***

**Contributors: Laurent Sorber, Nico Vervliet, Otto De Bals, Martijn Boussé, Griet Goovaerts, Borbála Hunyadi, HN Bharath, Stijn Dupulthys, Rob Zink, Matthieu Vendeville, Vasile Sima**

# WP5: Tensor formulation of biomedical BSS problems (1)

*Translate biomedical problem into an “interpretable” tensor decomposition*

## Task 5.1 Artefact removal

- *Remove noise, irrelevant signals, ...*

## Task 5.2 Preprocessing

- *Low-pass filtering and downsampling effective measures to improve source extraction via CPD, e.g. in cognitive EEG*

## Task 5.3 Tensorization

- *Various ways of tensorization (Hankel, Löwner, decimation,...)*
- *Segmentation especially useful for both large-scale (instantaneous) blind source separation and large-scale (convolutive) blind system identification.*
- *Strong properties of tensorized data revealed, efficient Tensorlab implementations, promising applications in biomedical BSS problems, e.g.:*
  - *ECG: beat-by-beat, segmentation strategy, Löwner tensorization*
  - *EEG: Hilbert-Huang transformation, Hankel expansion, trial-by-trial, multiscale expansion*
  - *MRSI: symmetric  $XX^T$  expansion, Löwner tensorization, Hankel expansion*



# WP5: Tensor formulation of biomedical BSS problems (2)

## Task 5.4 Choice of decomposition type

- *CPD, BTD, MLSVD, convolutive mixture, ...? Trade-off between simplicity and model accuracy.*
- *Supervised decompositions, e.g. HODA, might outperform unsupervised ones (e.g. MLSVD) for machine learning, e.g. classification in the LS-CPD setting.*

## Task 5.5 Choice of decomposition parameters

- *Estimate rank and multilinear rank (Core Consistency Diagnostic, Pareto optimization, ...)*
- *Can be automated by thresholding ML singular values in MLSVD using energy constraints  
→ applied to multichannel ECG compression.*
- *Robustness improved by adding prior knowledge (e.g. templates).*

## Task 5.6 Definition of constraints

- *Introduce relevant mathematical constraints (orthogonality, NN, periodicity, ...)*

*General framework presented for making informed choices in case of epileptic EEG-fMRI*

**Contributors: Rob Zink, Borbála Hunyadi, Simon Van Eyndhoven, Otto De Bals, Martijn Boussé, Griet Goovaerts**

# WP6: Tensor based BSS in Magnetic Resonance Spectroscopic Imaging



## Task 6.1 Metabolite quantification and artefact removal

- *Exponential and Löwner modeling of MRSI signals/spectra*
- *All-at-once water component removal from 2D/3D MRSI → outperforms voxel-wise approach*

## Task 6.2 Brain tumour tissue typing

- *Extensions of NMF to NN CPD factorization*
- *Adjust multilinear rank to handle artefacts and incorporate prior knowledge*
- *Extension to multiparametric MRI (FLAIR, T1/T2, DWI, PWI)*

## EXTRA Results:

- *NN CPD algorithm to detect longitudinal pathological changes → multiple sclerosis lesions (**this talk!**)*
- *Supervised voxel classification using CNN and MLSVD-based regularization*
- *Supervised tissue segmentation using superpixel 2-stage random forest with MLSVD input*

**Contributors: Bharath HN, Diana Sima, Nicolas Sauwen, Claudio Stamile**

## Task 7.1 Epileptic seizure detection in multichannel EEG

- *Seizure localization & artefact removal (eye, ...) using wavelet expansion + CPD*  
→ *If nonstationary : use wavelet expansion+ BTD or Hankel expansion+ (L,L,1)-BTB*
- *Supervised Nonconvulsive seizure detector using signatures from CPD/BTD of HHT tensor*
- *Adult EEG, neonatal EEG → preterm EEG*

## Task 7.2 Neonatal Brain Monitoring

- *Supervised EEG background abnormality assessment using holistic tensor-based HODA approach*
- *Automated Sleep stage classification of preterm EEG using multiscale entropy + CPD (**this talk!**)*

**EXTRA:** *physical activity recognition from single arm-worn accelerometer using HODA approach (**this talk!**)*

**Contributors: Borbála Hunyadi, Ofelie De Wel, Stijn Dupulthys, Vladimir Matic, Yissel Aldana Rodriguez, Lieven Billiet**

# WP7: Tensor based BSS in Functional monitoring (2)

## EXTRA to Task 7.2: Cardiac Monitoring using multichannel ECG

- *T-wave alternans detection using beat-by-beat tensor CPD and PARAFAC2*
- *Irregular heartbeat detection using CPD and Kronecker Product Equations*
- *Atrial fibrillation detection using wavelet & beat-by-beat expansions and MLSVD*
- *Fetal heart extraction using Löwner tensor and segmentation approaches*
- *Prediction of in-hospital cardiac arrest using tensor CPD*

## EXTRA: Multiscale Analysis-by-synthesis approaches using MLSVD and multichannel ECG

- *ECG compression*
- *Detection & localization of myocardial infarction*
- *T-wave alternans Detection*

**Contributors: Griet Goovaerts, Carolina Varon, Sibasankar Padhy , Simon Geirnaert**



*Focus on 2 studies: Cognitive functioning and Seizure localization*

## Extra to Task 8.1: Cognitive Functioning using mobile EEG

- *Auditory P300 recognition in single-trial ERP without subject-specific calibration phase using structured CPD  
→ further extended to structured BTD approach (**this talk!**)*
- *Alpha and low-beta oscillatory wave extraction using wavelet transformed EEG + CPD*
- *Extraction of N100 and P300 across subjects using CPD*

## Task 8.2 New EEG-fMRI integration approaches based on (coupled) CPD/BTD

### Epileptic zone localisation

- *Coupled Matrix-Tensor Factorization (CMTF) approach outperforming (multichannel) joint ICA*

### Cognitive functioning (spatiotemporal brain path characterisation during visual detection task)

- *Multichannel extension of jointICA*
- *Structured CMTF with Toeplitz factor to model neural-hemodynamic coupling via flexible EEG-fMRI fusion*

**Contributors: Borbála Hunyadi, Wout Swinnen, Simon Van Eyndhoven, Rob Zink, Stijn Dupulthys**

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- WP6: Multiple Sclerosis Diagnostics
- WP7: Wearable Health Monitoring
- WP7: Neonatal Monitoring
- WP8: Mobile EEG



## 4. Conclusions and Future Directions

# Cancer & MS Diagnostics

UZ Leuven departments

Imaging & Pathology

Radiology

Neurosurgery

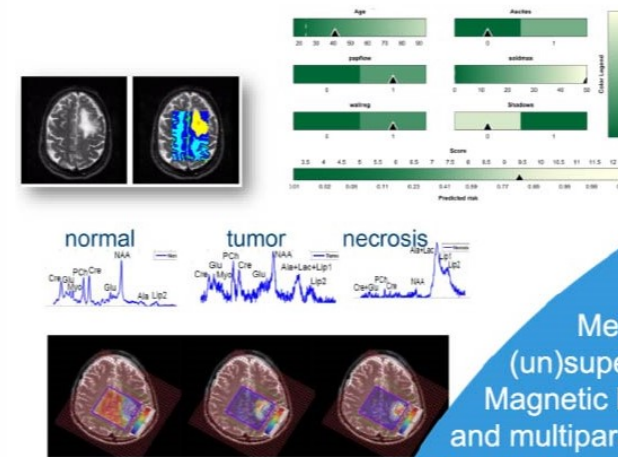
Development and Regeneration

Université Claude Bernard Lyon 1

Radiology

Neurology

EXAMPLE



06.

## Cancer Diagnostics

Metabolite quantification and (un)supervised tumor tissue typing using Magnetic Resonance Spectroscopic Imaging and multiparametric MR data fusion, Interpretable Prediction models for preoperative cancer diagnosis.



**CREATIS**

# TENSOR-BASED BSS FOR WHITE MATTER FIBER- BUNDLE ANALYSIS IN MULTIPLE SCLEROSIS

**Collaborator: joint PhD of Claudio Stamile**

(Université Claude Bernard Lyon 1 + KU Leuven)



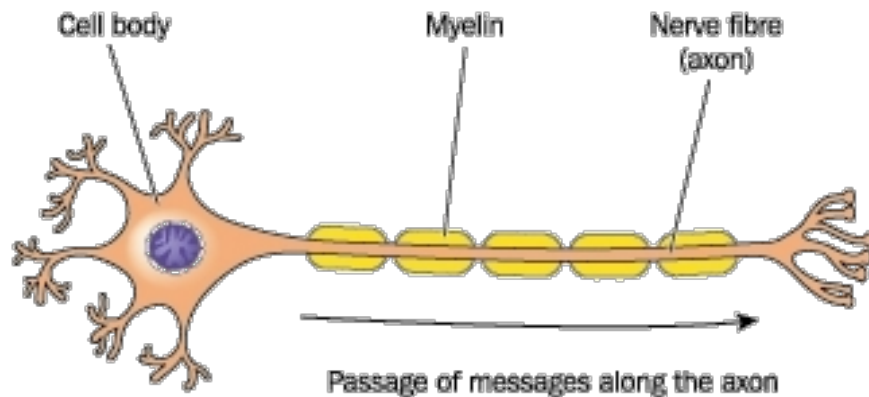
<http://tinyurl.com/stamile-phd>

# MULTIPLE SCLEROSIS

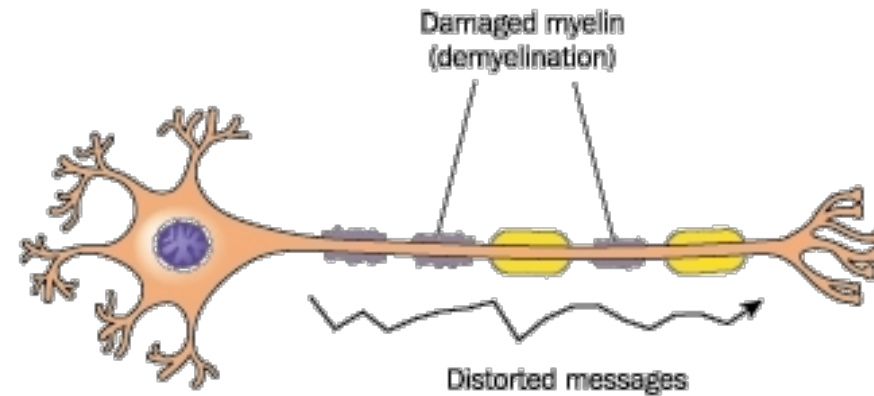
Multiple Sclerosis (MS) is a chronic disease of the central nervous system

- Demyelinating and Inflammatory Component
  - White Matter Lesions
- Neurodegenerative Component
  - Axonal loss

**Normal neuron**



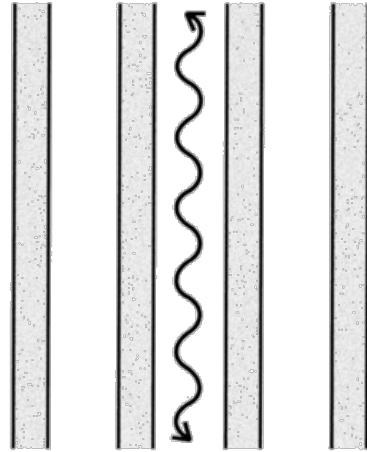
**Demyelination in MS**



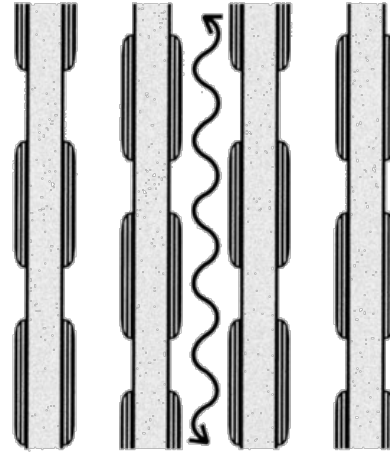


# DIFFUSION IMAGING

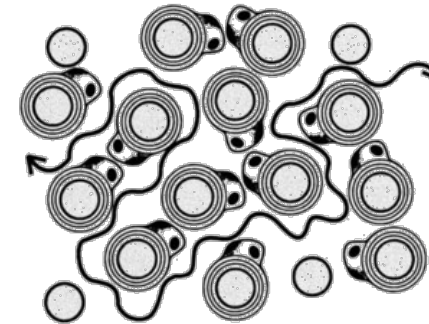
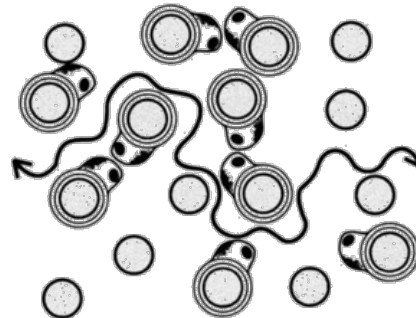
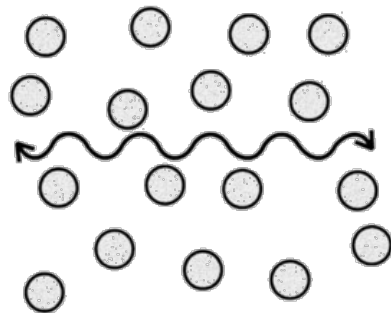
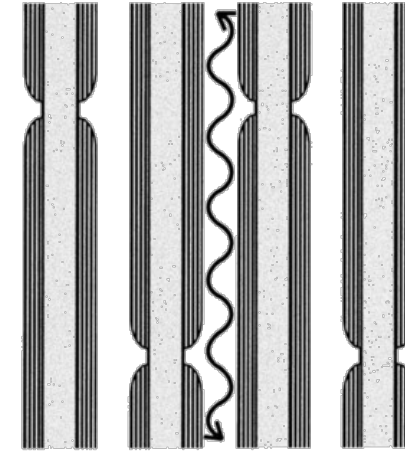
Unmyelinated  
tissue



Partly myelinated  
axons

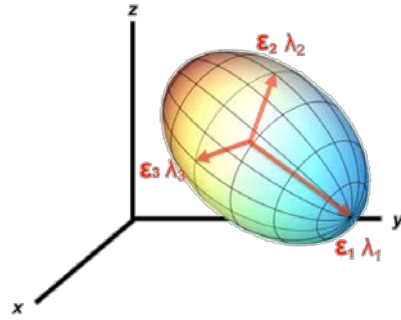


Myelinated  
tissue



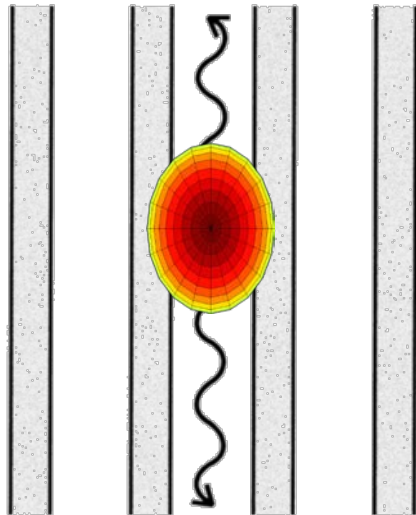
- Movement of the water inside the tissue
- Provide indirect measure of the tissue architecture

# DIFFUSION TENSOR IMAGING

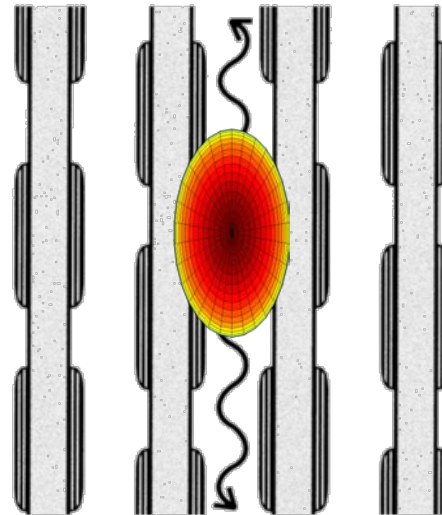


$$D = \begin{pmatrix} D_{xx} & D_{xy} & D_{xz} \\ D_{xy} & D_{yy} & D_{yz} \\ D_{xz} & D_{yz} & D_{zz} \end{pmatrix} = \begin{pmatrix} \lambda_1 & 0 & 0 \\ 0 & \lambda_2 & 0 \\ 0 & 0 & \lambda_3 \end{pmatrix} \begin{pmatrix} \vec{\epsilon}_1 \\ \vec{\epsilon}_2 \\ \vec{\epsilon}_3 \end{pmatrix}$$

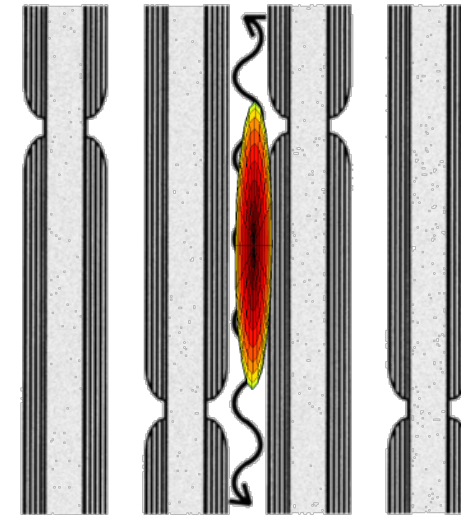
Unmyelinated  
tissue



Partly myelinated  
axons



Myelinated  
tissue

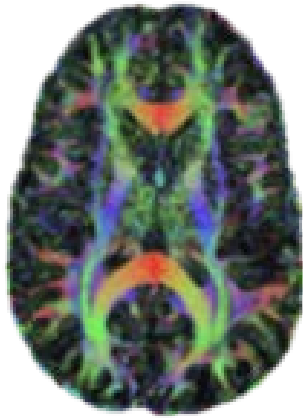


Shape of the  
ellipsoid  
changes  
according to  
the  
tissue  
structure

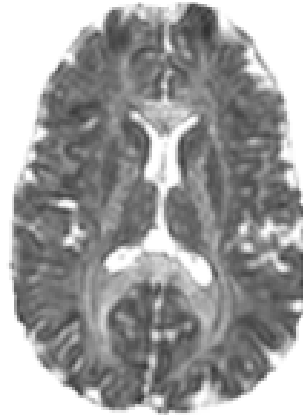


# DIFFUSION TENSOR IMAGING

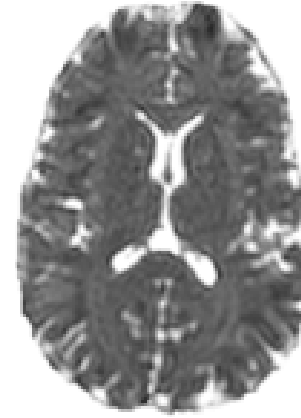
FA



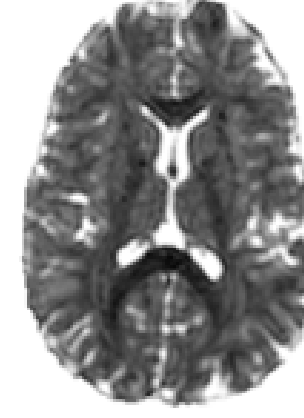
$\lambda_1$



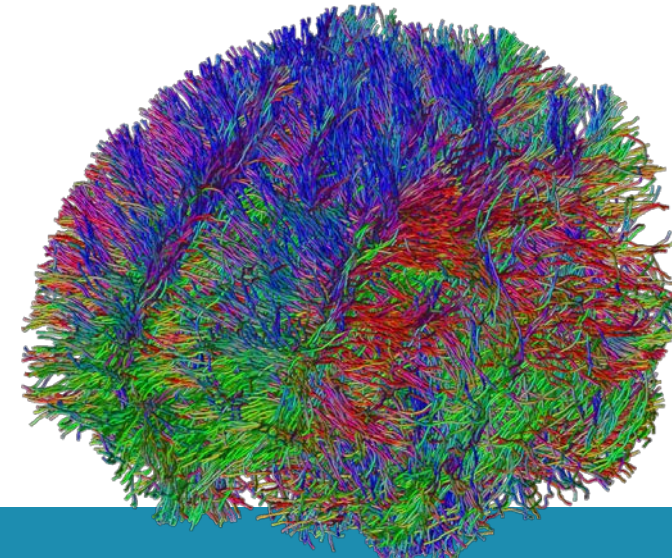
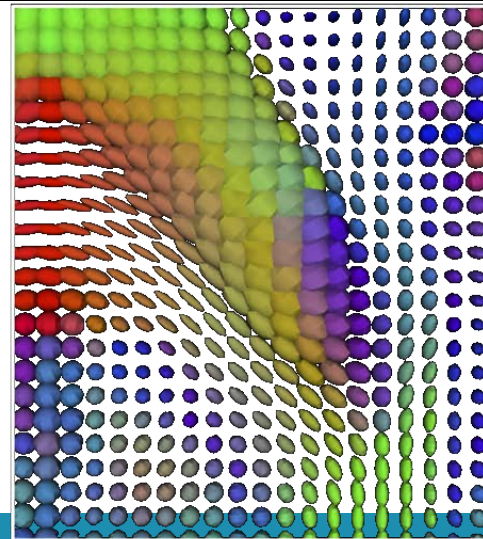
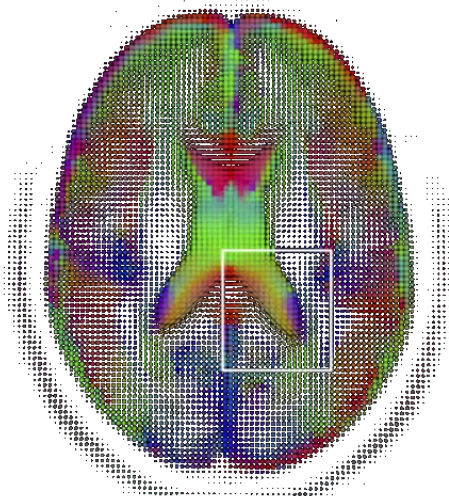
MD



$\lambda_r$



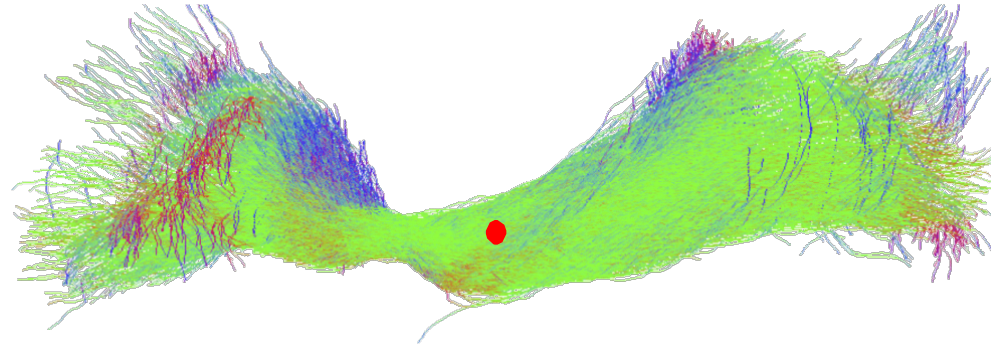
Quantitative information



Structural information

# MAIN OBJECTIVES

Demyelination and inflammation lead to alterations in fiber-bundles signal conduction

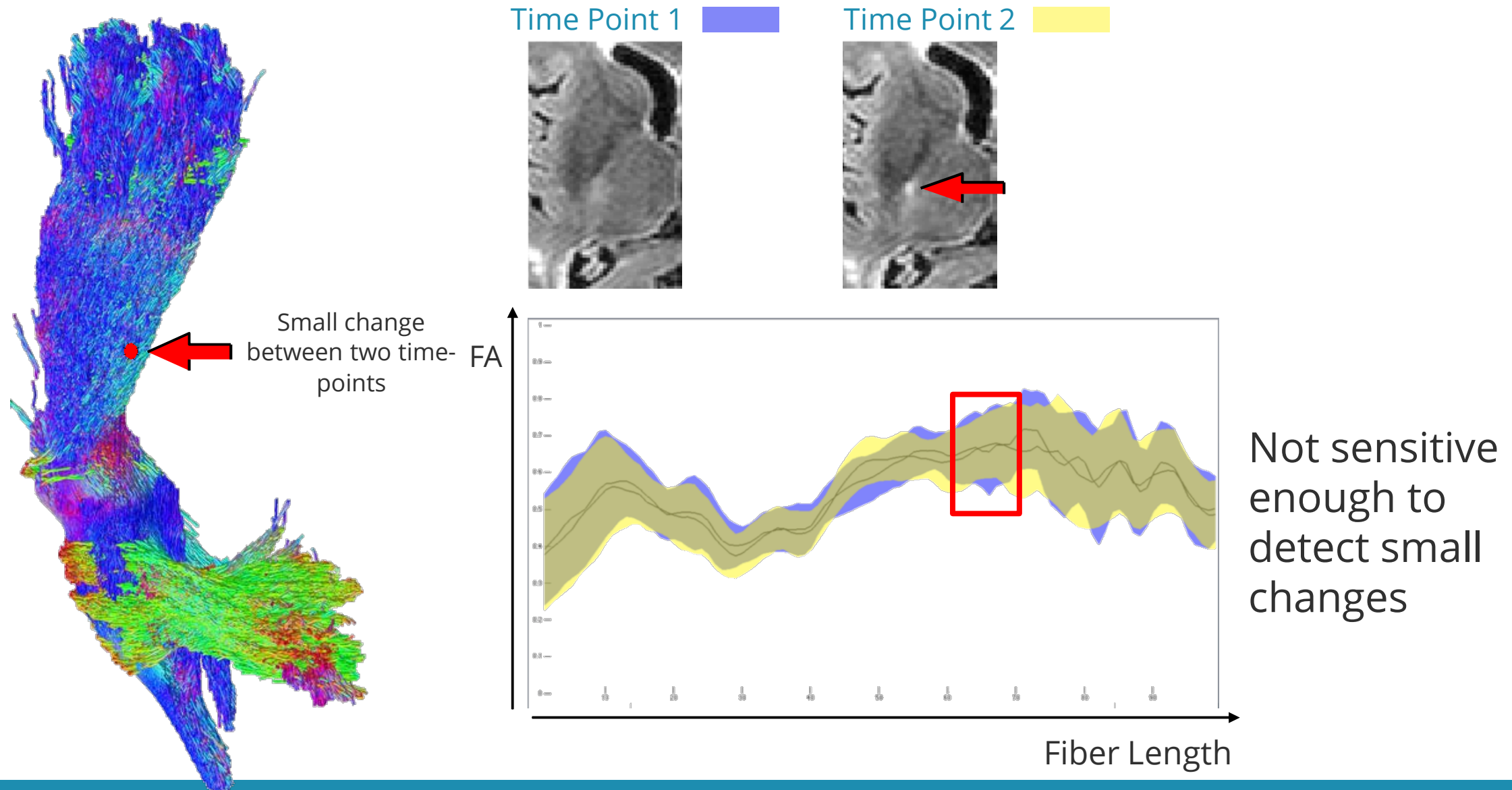


## Objectives :

1. Provide a better characterization of fiber-bundles profile:
  - Sensitive detection of WM lesions
  - Analyze afferent and efferent pathways
2. **Longitudinal fiber-bundle analysis → This talk!**

**Evidence-based decision making for personalized treatment of beginning MS patients to reduce disease severity at an early time-point.**

# "MEAN" FIBER PROFILE ANALYSIS

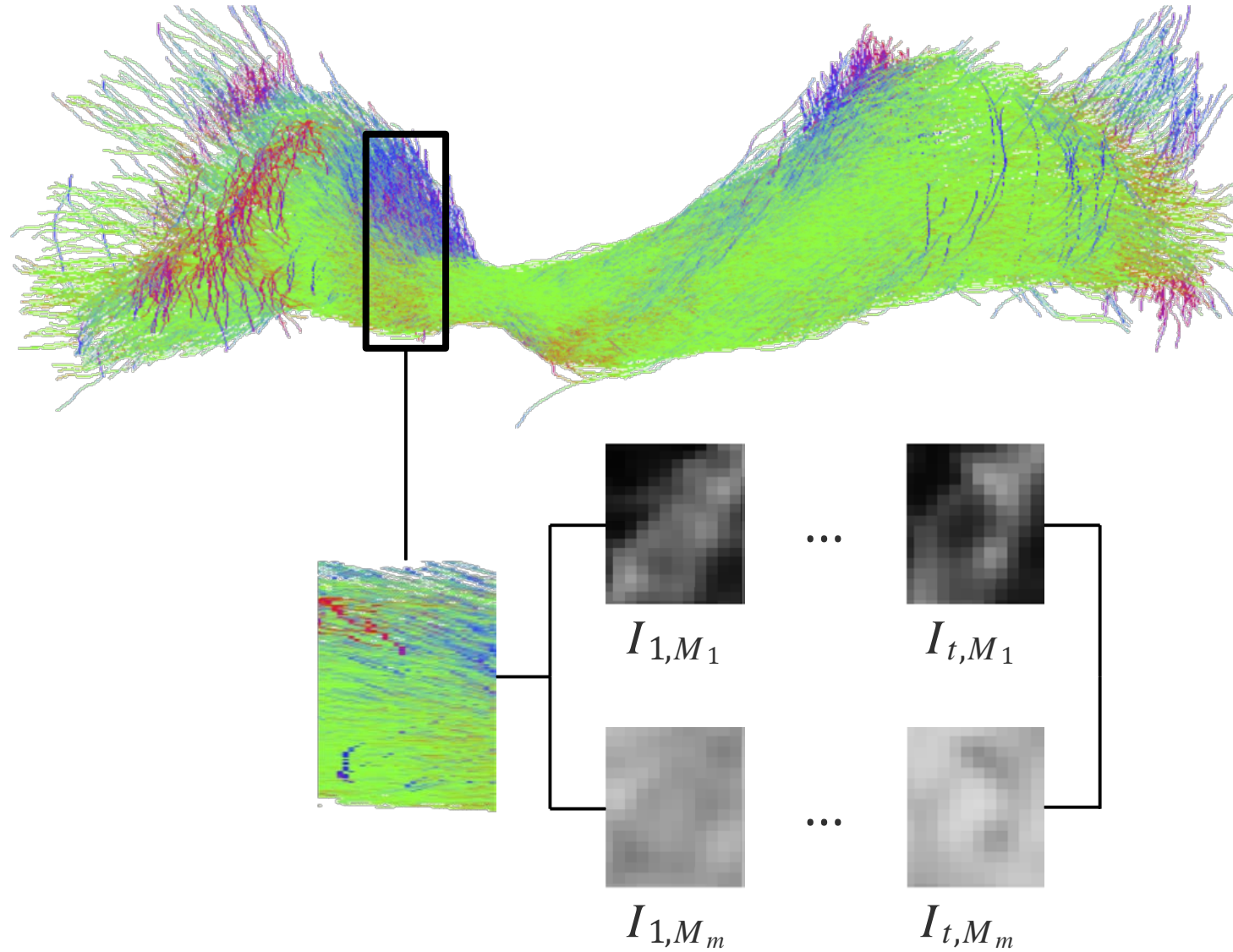


# LONGITUDINAL ANALYSIS

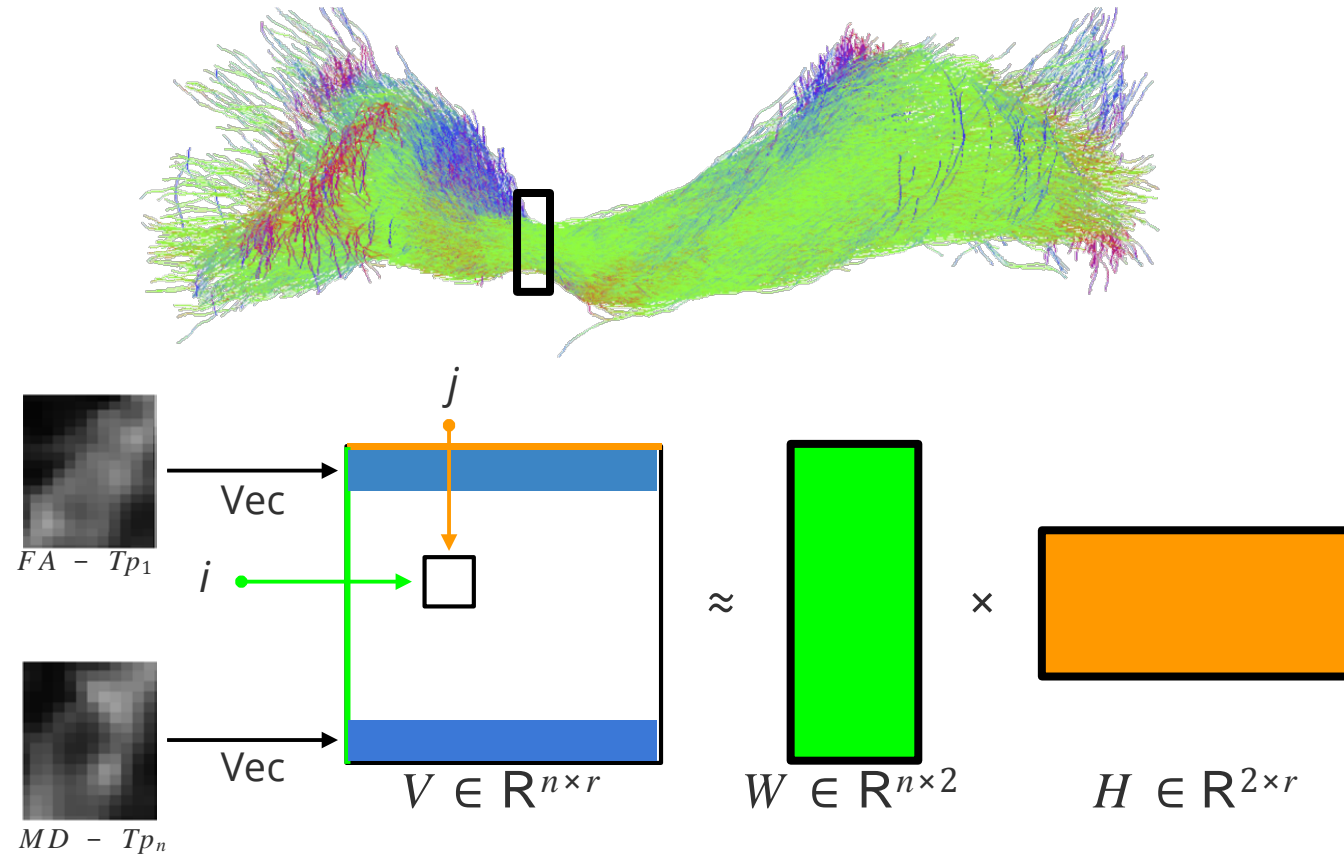
## NON-NEGATIVE MATRIX FACTORIZATION



# NON-NEGATIVE MATRIX FACTORIZATION



# NON-NEGATIVE MATRIX FACTORIZATION



$V_{i,j}$  = Value of the  $i$ -th modality/time-point for the  $j$ -th voxel of the cross-section

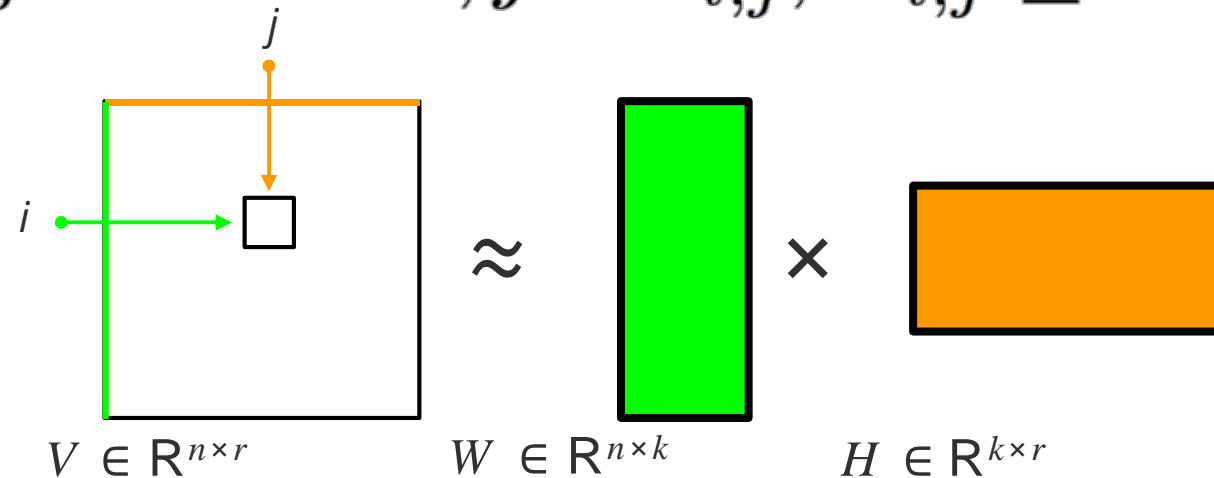
**W** Longitudinal modality contribution

**H** Segmentation of "normal" and "pathological" voxels

# NON-NEGATIVE MATRIX FACTORIZATION

$$\underset{\mathbf{W}, \mathbf{H}}{\text{minimize}} \quad f(\mathbf{W}, \mathbf{H}) = \frac{1}{2} \|\mathbf{V} - \mathbf{W} \times \mathbf{H}\|_F^2$$

$$\text{subject to} \quad \forall i, j : W_{i,j}, H_{i,j} \geq 0$$

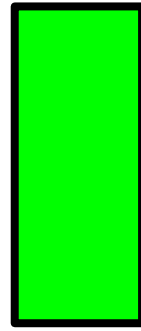


- Non Convex optimization problem → Apply sequentially = Recursive NMF
- Different algorithms proposed in literature:
  - Alternating Least Squares (ALS) , Multiplicative Update, Hierarchical ALS





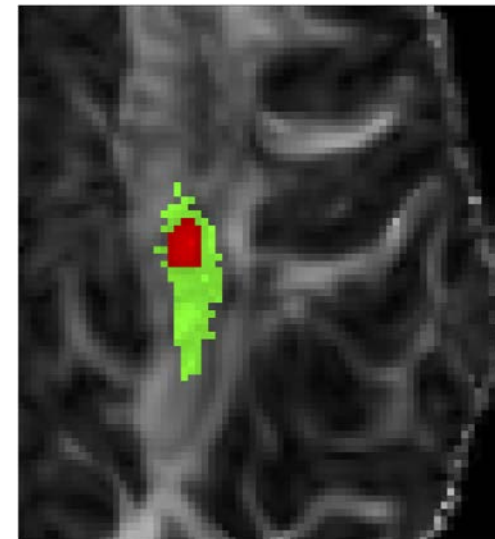
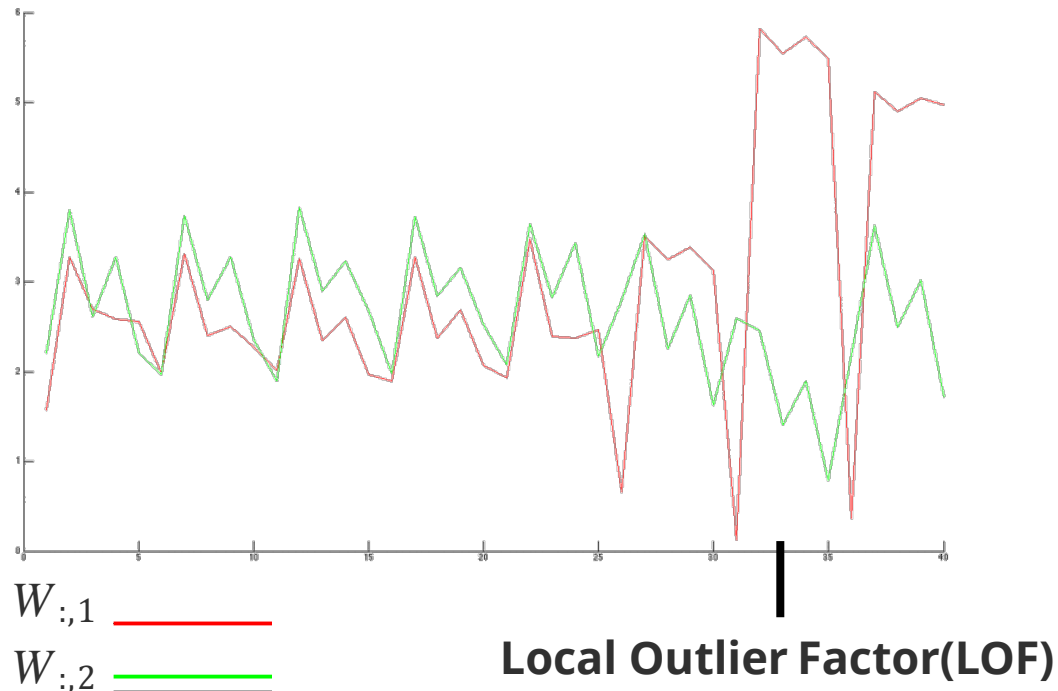
# NON-NEGATIVE MATRIX FACTORIZATION





$$W \in \mathbb{R}^{n \times 2}$$



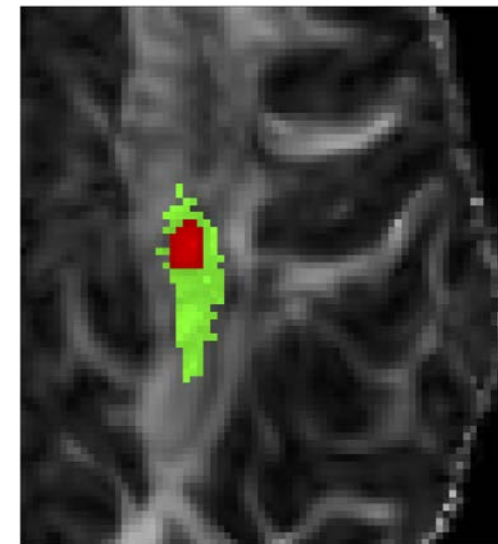
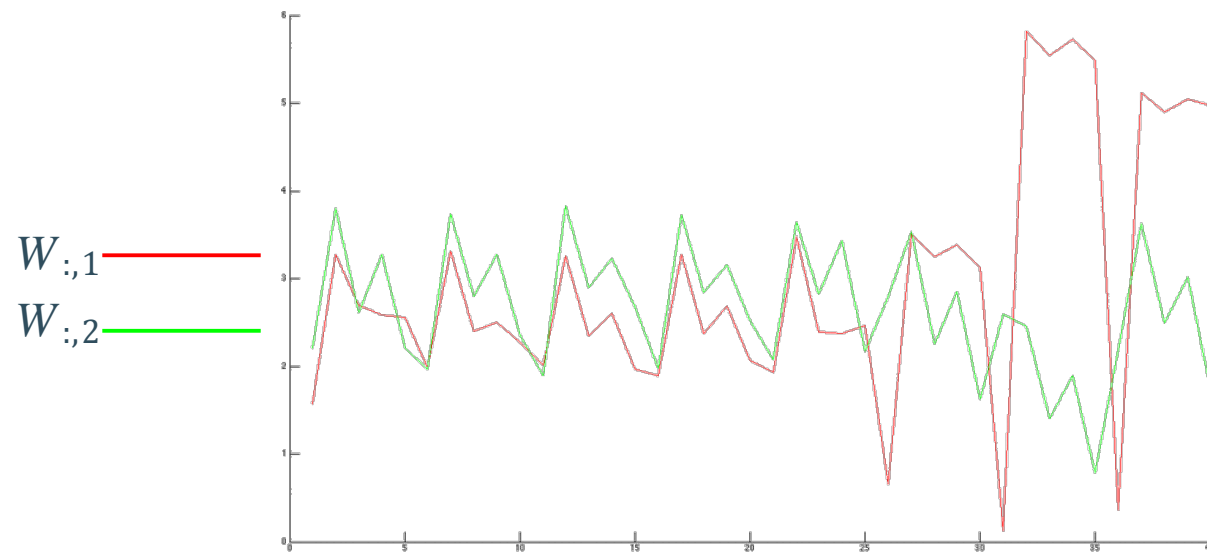
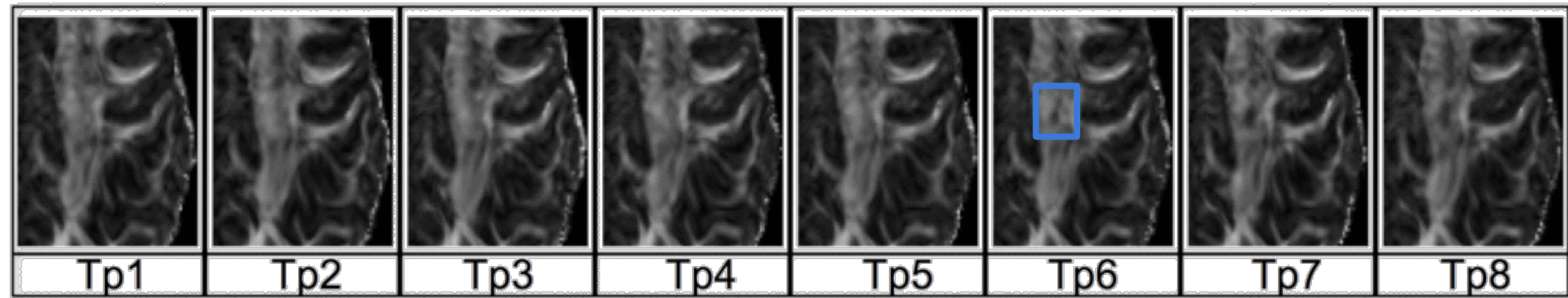
$$H \in \mathbb{R}^{2 \times r}$$



  $H_{1,:}$

  $H_{2,:}$

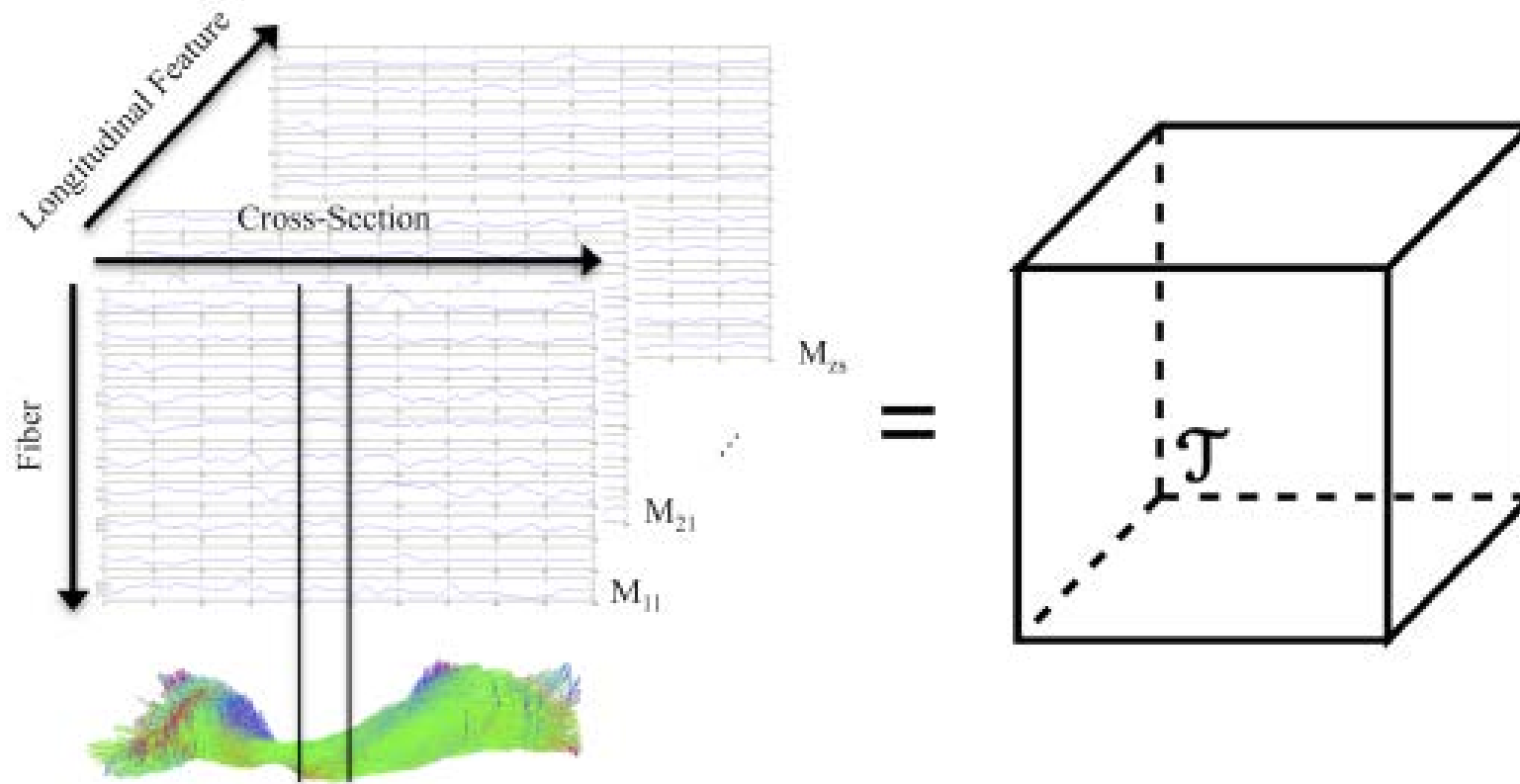
# APPLICATION ON MS PATIENTS



$H_{1,:}$  ■  
 $H_{2,:}$  ■

# LONGITUDINAL ANALYSIS

## NON-NEGATIVE TENSOR FACTORIZATION



# NON-NEGATIVE TENSOR FACTORIZATION

- $z$  : Number of features
- $s$  : Number of time-points
- $M_{ij}$  :  $i$ -th feature at  $j$ -th time-point

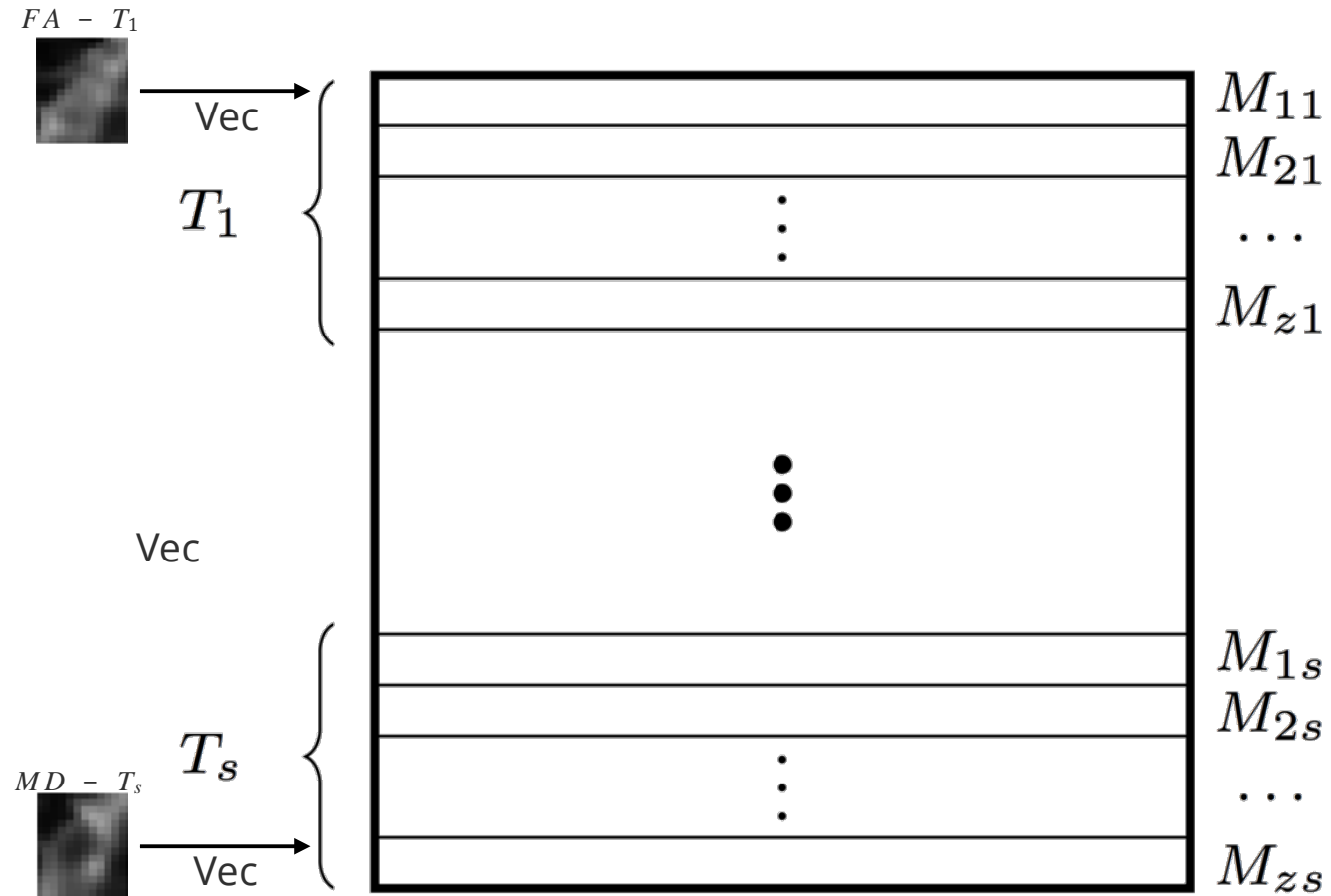
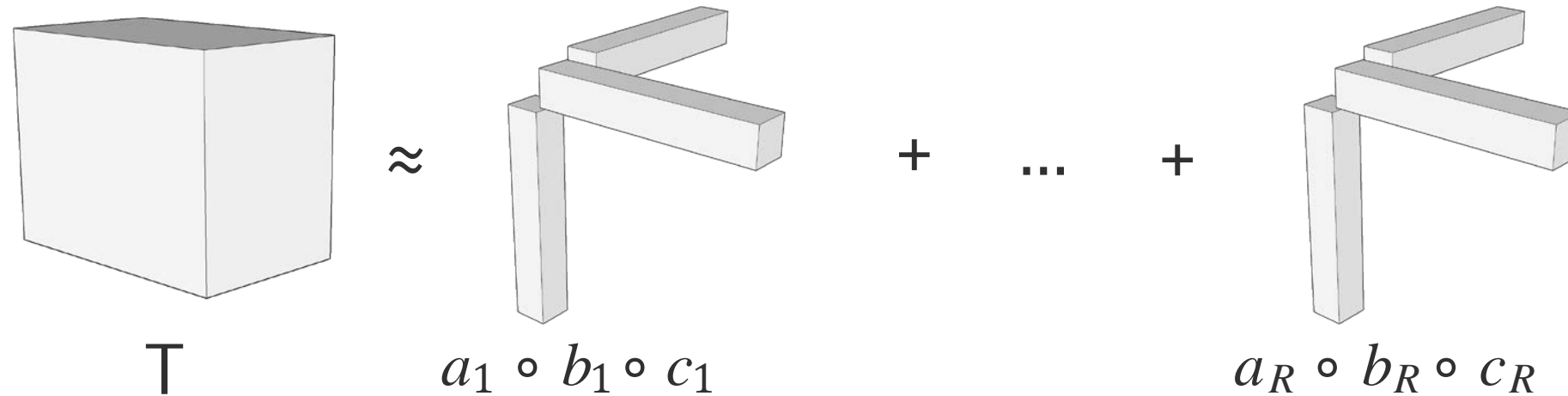


Illustration of the 3<sup>rd</sup> mode of the tensor  $T$

# NON-NEGATIVE TENSOR FACTORIZATION

Canonical Polyadic Decomposition (CPD) is a decomposition of a tensor  $\mathcal{T}$  into sum of rank-1 tensors


$$\mathcal{T} \approx a_1 \circ b_1 \circ c_1 + \dots + a_R \circ b_R \circ c_R$$

The rank ( $R$ ) of a CPD is the number of rank-1 tensors used to approximate  $a_i, b_i, c_i \geq 0$

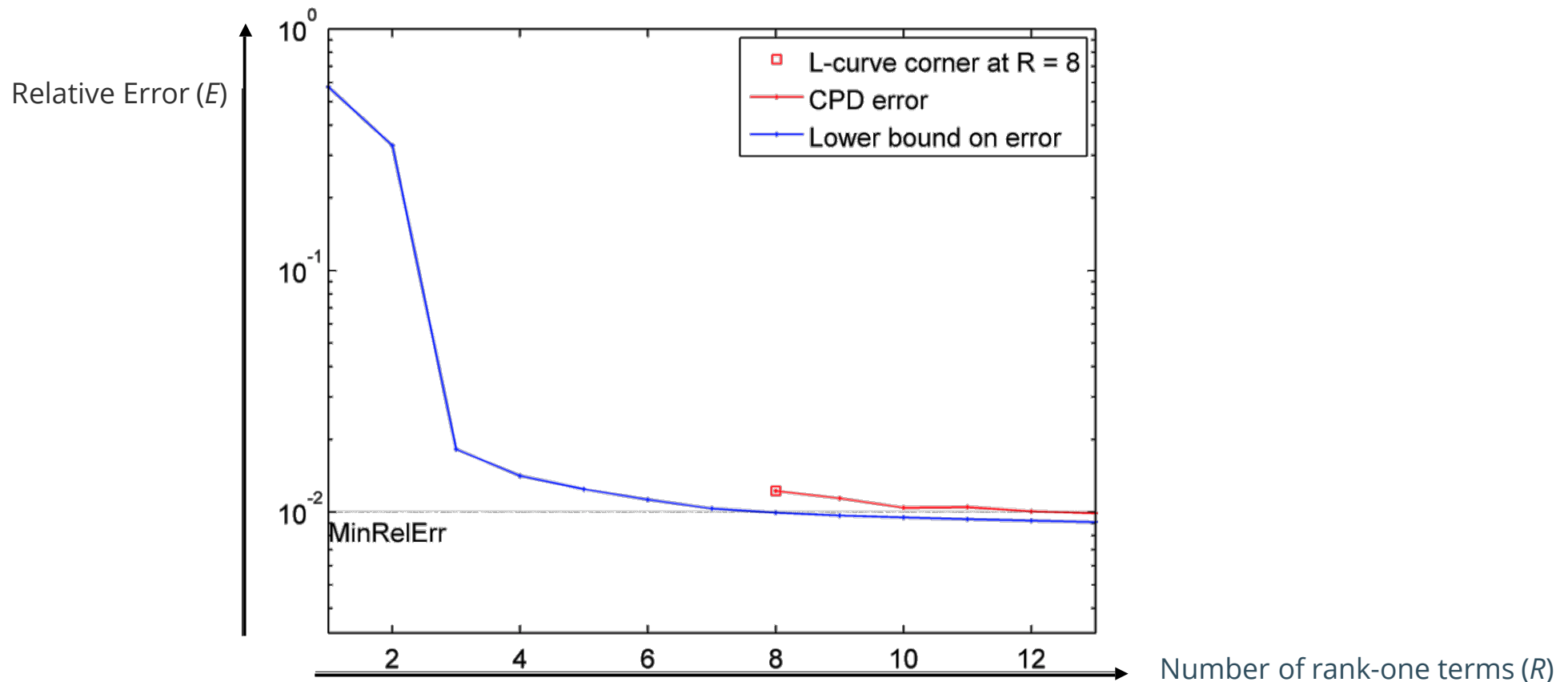
$$\min_{a_i \geq 0, b_i \geq 0, c_i \geq 0} \|\mathcal{T} - \sum_{i=1}^R a_i \circ b_i \circ c_i\|_F^2$$

- Non Convex optimization problem. A set of algorithms available in [TensorLab](#)

# NON-NEGATIVE TENSOR FACTORIZATION

**Rank estimation  $R$**  (using *rankest* from Tensorlab, based on L-curve)

1. The number of rank-one terms ( $R$ ) is increased
2. The relative error ( $E$ ) of the approximation is computed
3. If  $E < \text{MinRelErr}$  the rank is defined

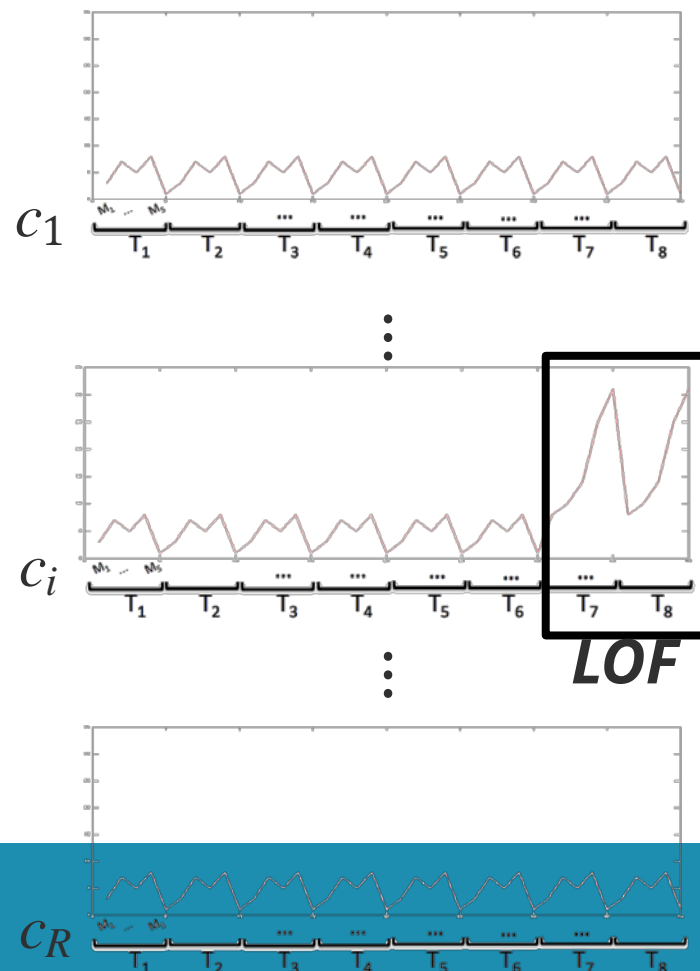




# NON-NEGATIVE TENSOR FACTORIZATION

## Identification of components

1. Identify the rank-one components containing information related to "pathological longitudinal changes"



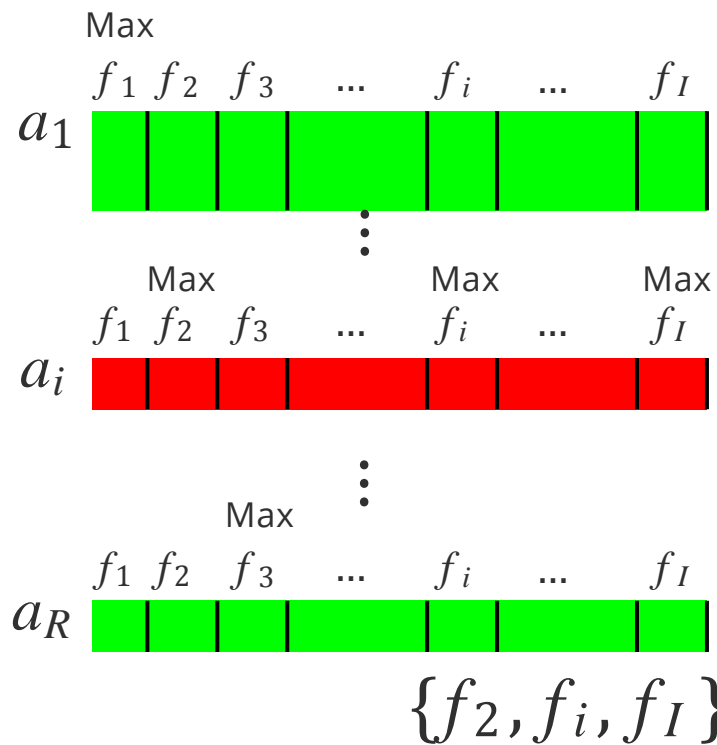
The algorithm collects the indices of all the rank-one terms containing outliers identified by LOF

$$P = \{c_i \mid 1 \leq i \leq R \mid c_i \text{ contains Outliers}\}$$

# NON-NEGATIVE TENSOR FACTORIZATION

## Pathological fibers identification:

1. Identifies the set  $P$  of fibers represented by rank-one terms identified as "outliers" by the previous step.



For each  $f_j \in F$ :

if  $a_{ij} = \max(A_{[i,:]}, i \in P)$

*is Pathological*

else

*is Normal*

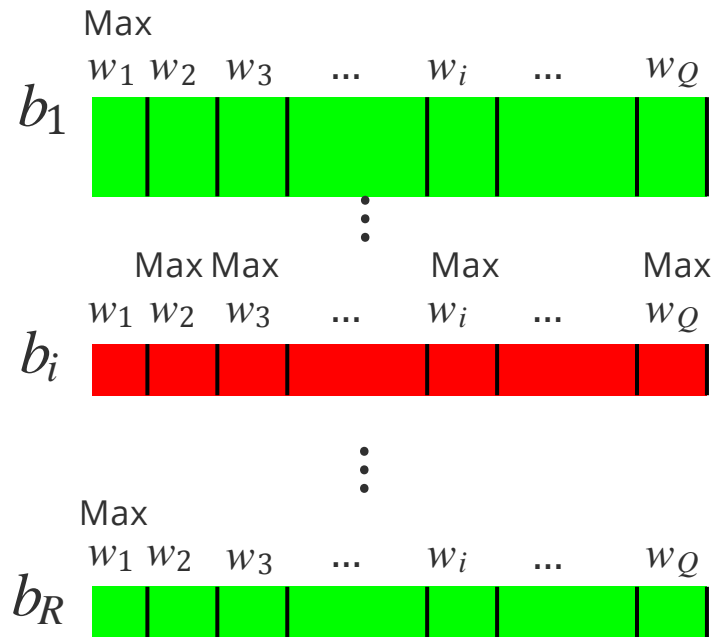
- They have maximal contribution in terms detected as outlier in the previous step



# NON-NEGATIVE TENSOR FACTORIZATION

## Pathological cross-sections identification:

1. Identifies the set  $H$  of cross-sections represented by rank-one terms identified as "outliers" by the previous step.



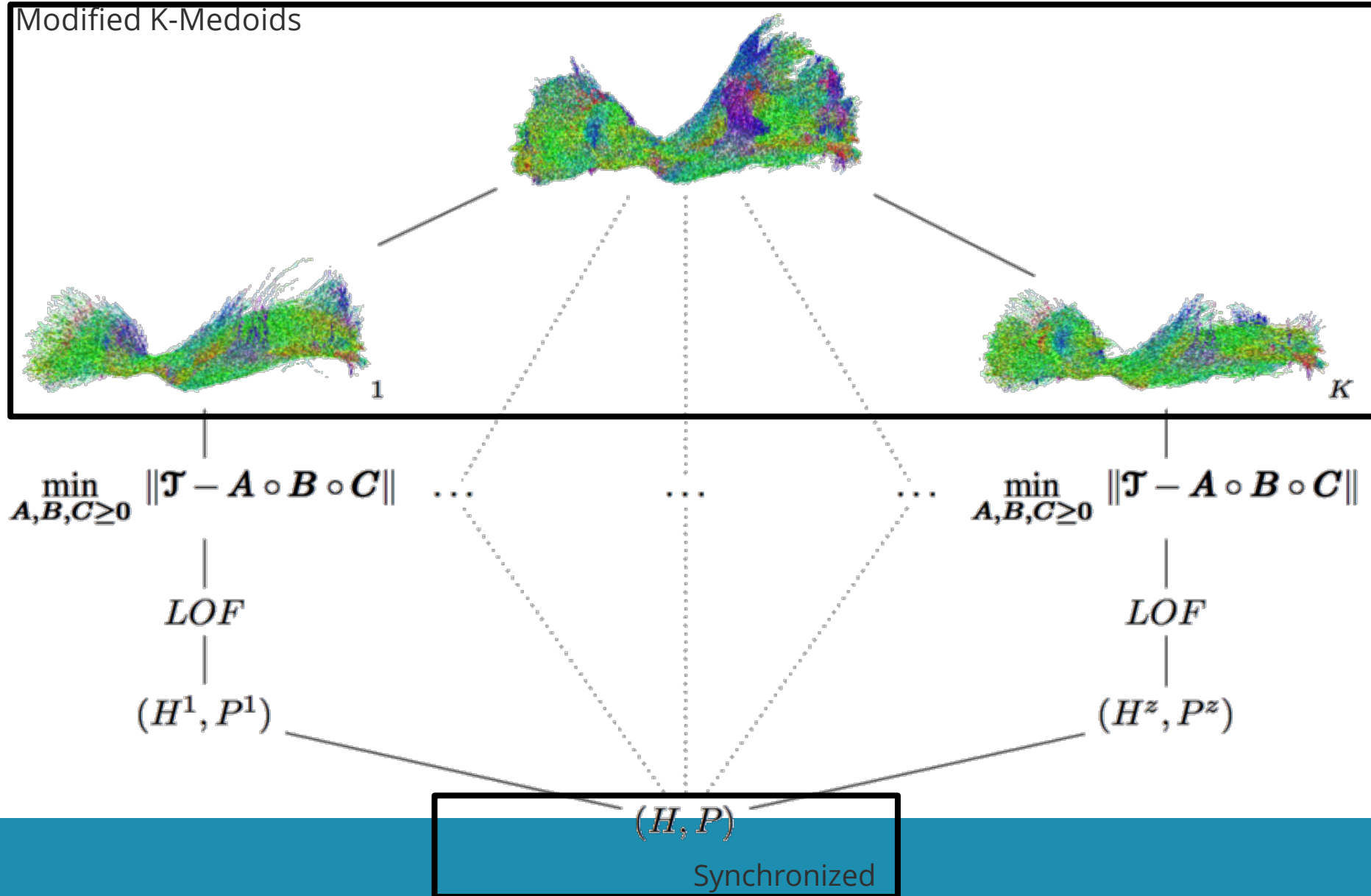
For each  $w_j \in F$ :  
if  $b_{ij} = \max(B_{[i,:]}, i \in P)$   
*is Pathological*  
else  
*is Normal*

$\{w_2, w_3, w_i, w_Q\}$  are detected as pathological cross-sections

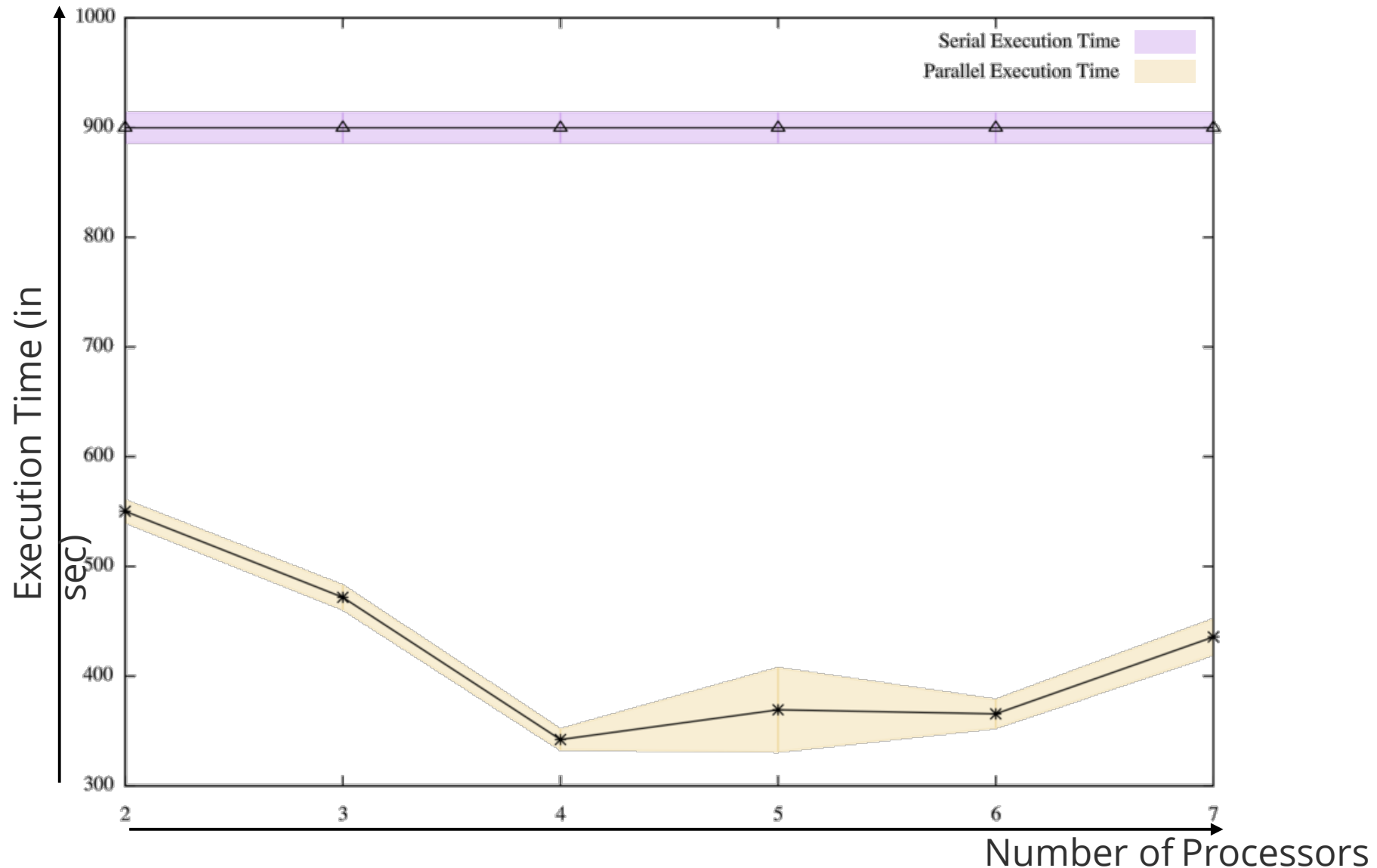
- They have maximal contribution in terms detected as outlier in previous step

# PARALLEL IMPLEMENTATION

Modified K-Medoids



# PARALLEL IMPLEMENTATION-RESULTS



- Min Speed-up factor: 1.63 ( $K=2$ )
- Max Speed-up factor: 2.60 ( $K=4$ )



# RESULTS ON SIMULATED DATA

- 100 longitudinal variations were simulated on control subject
- LVS simulation model was used
- Variations were generated along 10 different fiber-bundles
- Different features extracted from diffusion maps obtained from the data
- Extensive grid search was performed in order to find the best parameters for the algorithm



# RESULTS ON SIMULATED DATA

## Detection of pathological fibers

	Accuracy	Precision	Sensitivity
$\lambda_2, \lambda_3$	<b>0.97 (0.06)</b>	<b>0.79 (0.08)</b>	<b>0.40 (0.11)</b>
$\lambda_2, \lambda_3, \text{FA}$	0.77 (0.29)	0.73 (0.11)	0.20 (0.02)
$\lambda_2, \lambda_3, \text{FA, MD}$	0.79 (0.32)	0.76 (0.15)	0.23 (0.05)
KLA,AA	0.71 (0.23)	0.69 (0.13)	0.16 (0.01)
KLA,FA,AA	0.77 (0.20)	0.66 (0.15)	0.15 (0.04)

## Detection of pathological cross-sections

	Tensor Factorization			Non-Negative Matrix Factorization		
	Accuracy	Precision	Sensitivity	Accuracy	Precision	Sensitivity
$\lambda_2, \lambda_3$	0.63 (0.10)	0.98 (0.02)	<b>0.95 (0.15)</b>	0.65 (0.12)	0.60 (0.08)	0.92 (0.10)
$\lambda_2, \lambda_3, \text{FA}$	0.40 (0.12)	<b>1.00 (0.01)</b>	0.56 (0.12)	<b>0.74 (0.11)</b>	0.68 (0.04)	0.90 (0.09)
$\lambda_2, \lambda_3, \text{FA, MD}$	0.47 (0.13)	0.97 (0.03)	0.74 (0.23)	<b>0.74 (0.08)</b>	0.68 (0.12)	0.89 (0.11)
KLA,AA	0.30 (0.10)	0.96 (0.03)	0.64 (0.30)	0.59 (0.07)	0.55 (0.12)	0.91 (0.10)
KLA,FA,AA	0.32 (0.12)	0.96 (0.03)	0.63 (0.28)	0.56 (0.11)	0.53 (0.09)	0.85 (0.05)

## Detection of pathological time-points

	Tensor Factorization			Non-Negative Matrix Factorization		
	Accuracy	Precision	Sensitivity	Accuracy	Precision	Sensitivity
$\lambda_2, \lambda_3$	<b>0.84 (0.12)</b>	<b>0.93 (0.09)</b>	<b>0.96 (0.13)</b>	0.72 (0.30)	0.86 (0.18)	0.78 (0.31)
$\lambda_2, \lambda_3, \text{FA}$	0.60 (0.20)	0.76 (0.10)	0.54 (0.16)	0.57 (0.36)	0.75 (0.17)	0.53 (0.35)
$\lambda_2, \lambda_3, \text{FA, MD}$	0.73 (0.30)	0.87 (0.18)	0.81 (0.32)	0.67 (0.36)	0.84 (0.18)	0.78 (0.33)
KLA,AA	0.50 (0.30)	0.71 (0.21)	0.82 (0.30)	0.50 (0.36)	0.67 (0.19)	0.56 (0.37)
KLA,FA,AA	0.50 (0.32)	0.74 (0.19)	0.75 (0.20)	0.50 (0.36)	0.66 (0.19)	0.57 (0.34)

# APPLICATION ON MS PATIENTS

- **Subjects:**

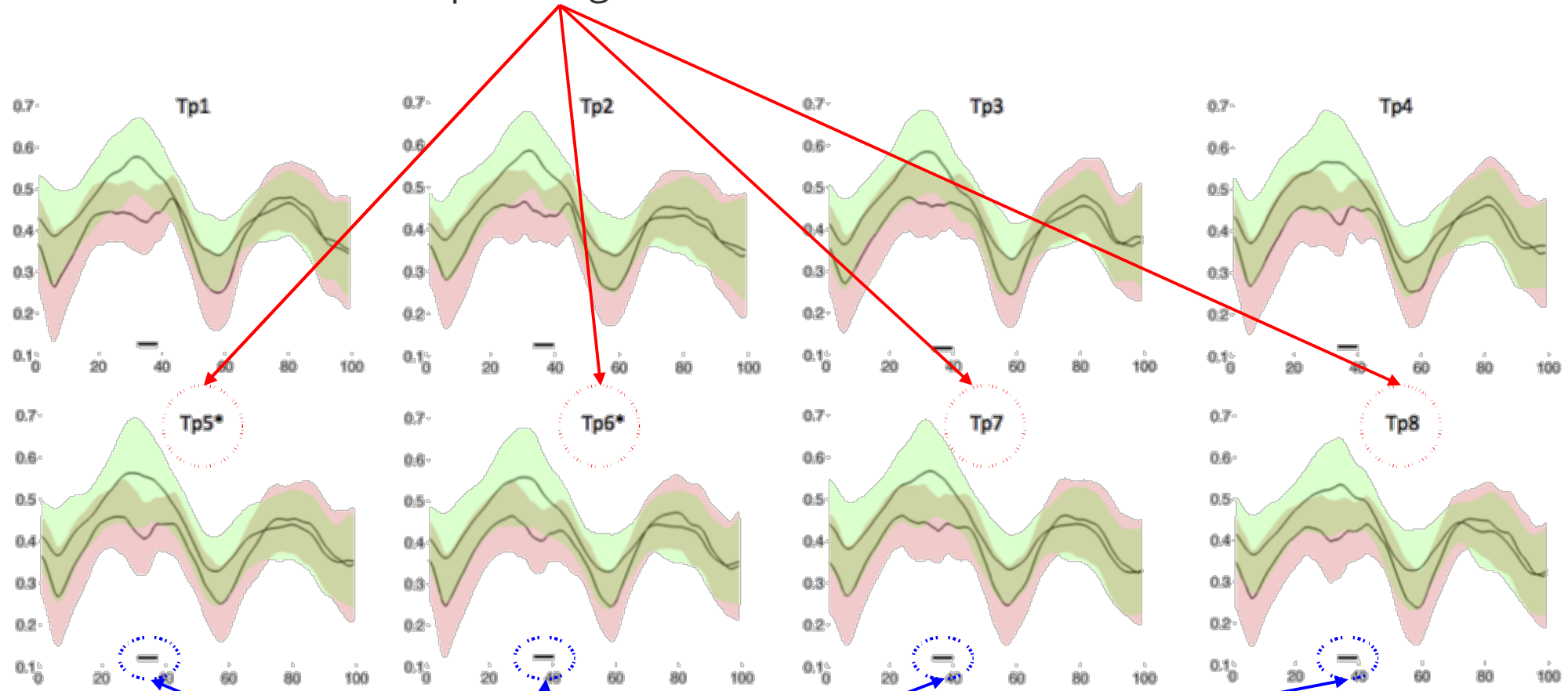
- 5 Relapsing Remitting (RR) Multiple Sclerosis Patients
- Lesions verified by expert neurologist
  - Along 10 fiber-bundles

- **Protocol:**

- EPI Acquisition: TE/TR = 60/8210 ms ; FOV = 224x224x120 $mm$
- 32 gradient directions  $b = 1000s.mm^{-2}$
- Voxel Size: 0.875x0.875x2.0 $mm^3$

# APPLICATION ON MS PATIENTS

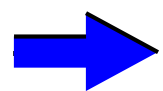
Time-points detected as  
"pathological"



Cross-sections detected as  
"pathological"

# CONCLUSION

- A new tensor-based framework for the analysis of longitudinal changes occurring along WM fiber-bundles.
- Multi time-points ( $> 2$ ), multi-features and multi dimensional:
  - Fibers, Cross-sections and Time-Points
- The method was tested on real simulated data with extensive parameters search
- Improved computational speed using parallel programming



**Tensor factorization is a potential tool to analyze multi-dimensional data in MRI based studies**

- **Limitation:** Low sensitivity in pathological fibers detection

# Wearable Health Monitoring

UZ Leuven departments:

Cardiology

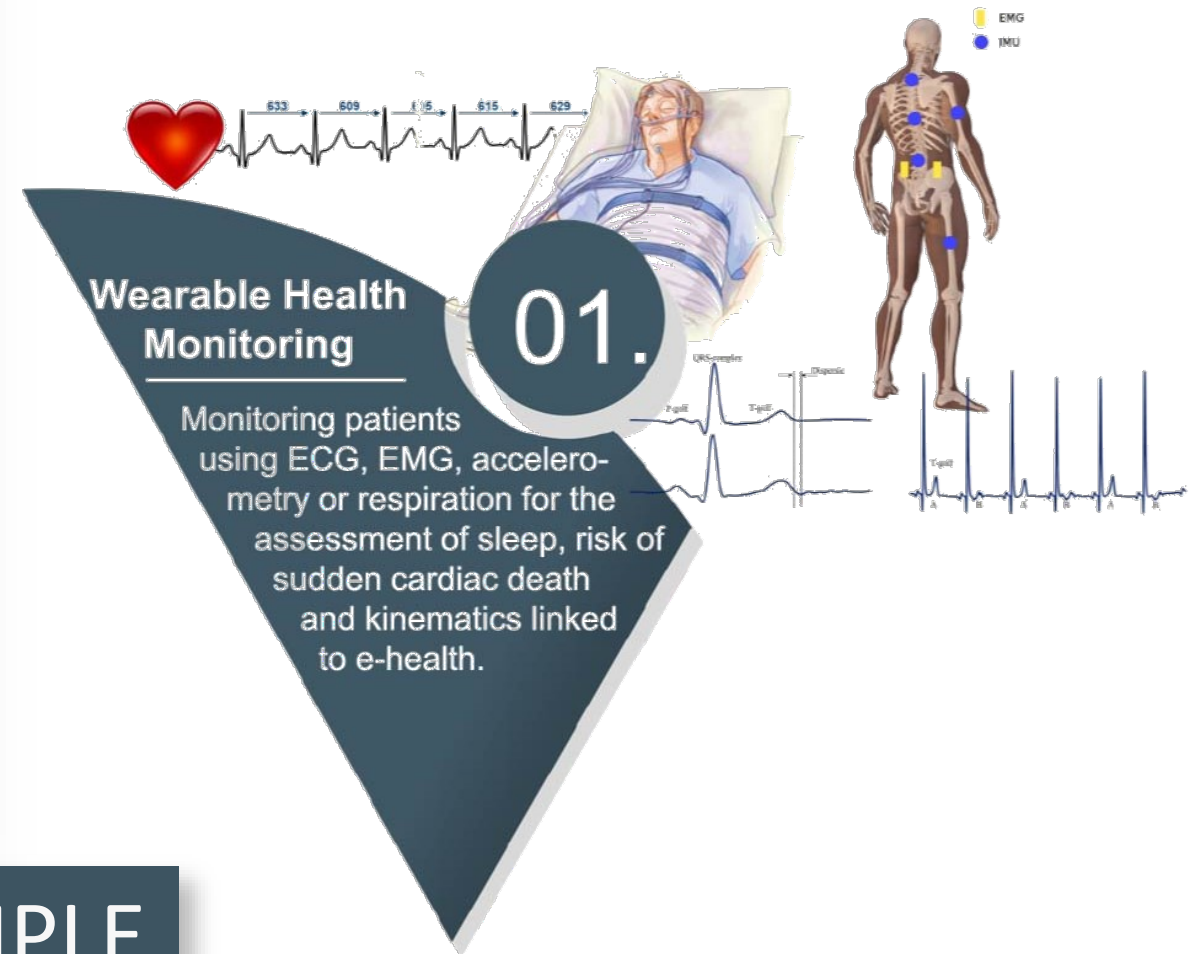
Neurology

Pneumology

**Rheumatology**

Pulderbos Revalidation Center

EXAMPLE





# Recognition of Physical Activities from Wearable Sensors: a Multiway approach

Collaborator: Lieven Billiet

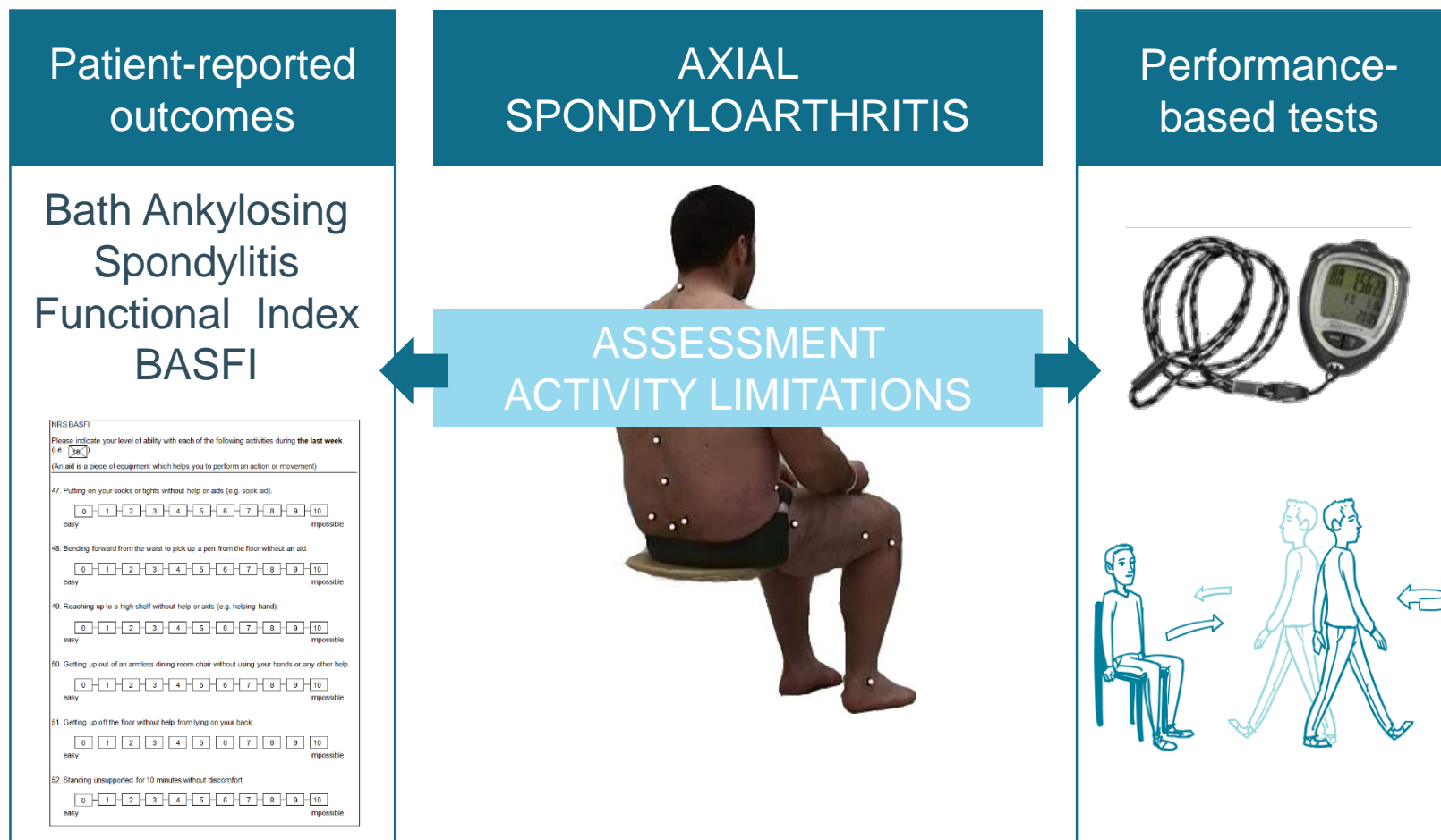
DISSERTATION (KU LEUVEN, OCT. 2018)

Clinical Decision Support: Interpretability and Applications in Patient Monitoring

Van Huffel, Sabine (Supervisor); Van Belle, Vanya (Cosupervisor)

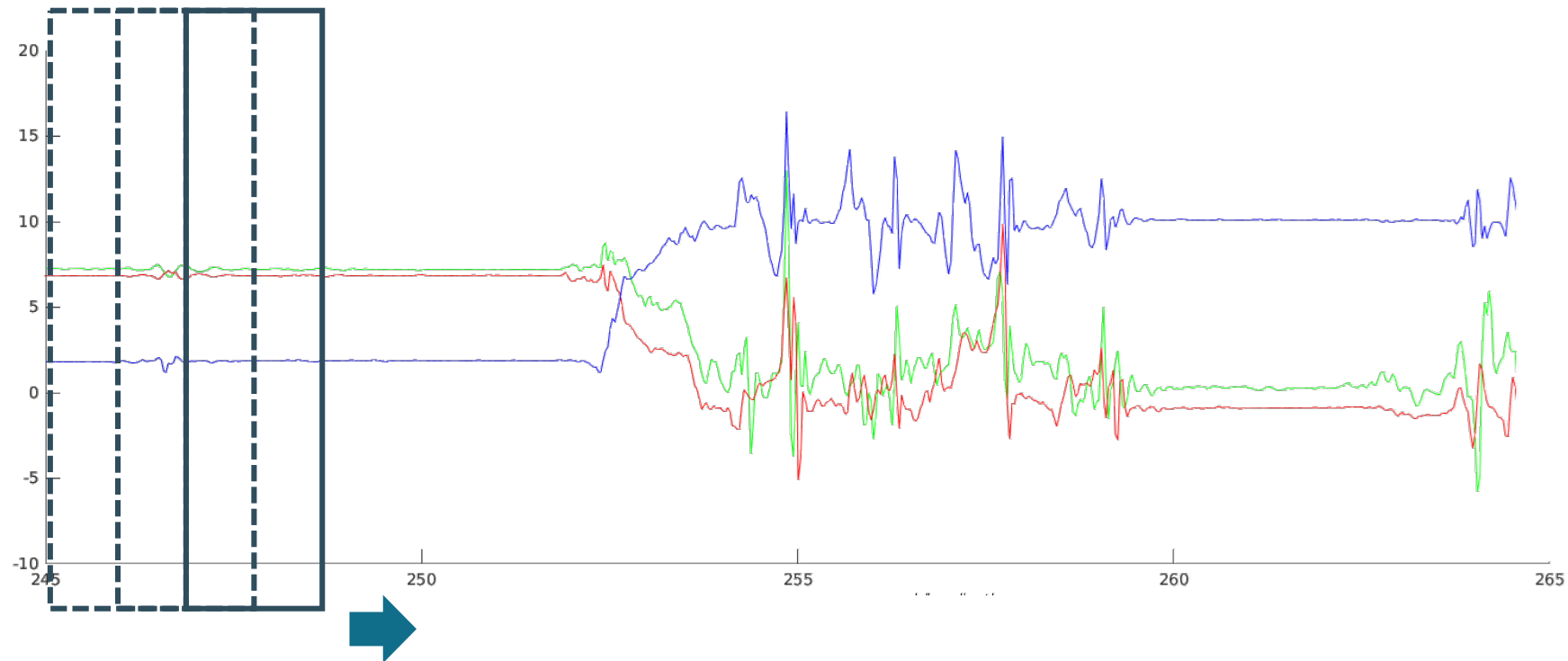


# Background





# Activity recognition: sliding windows



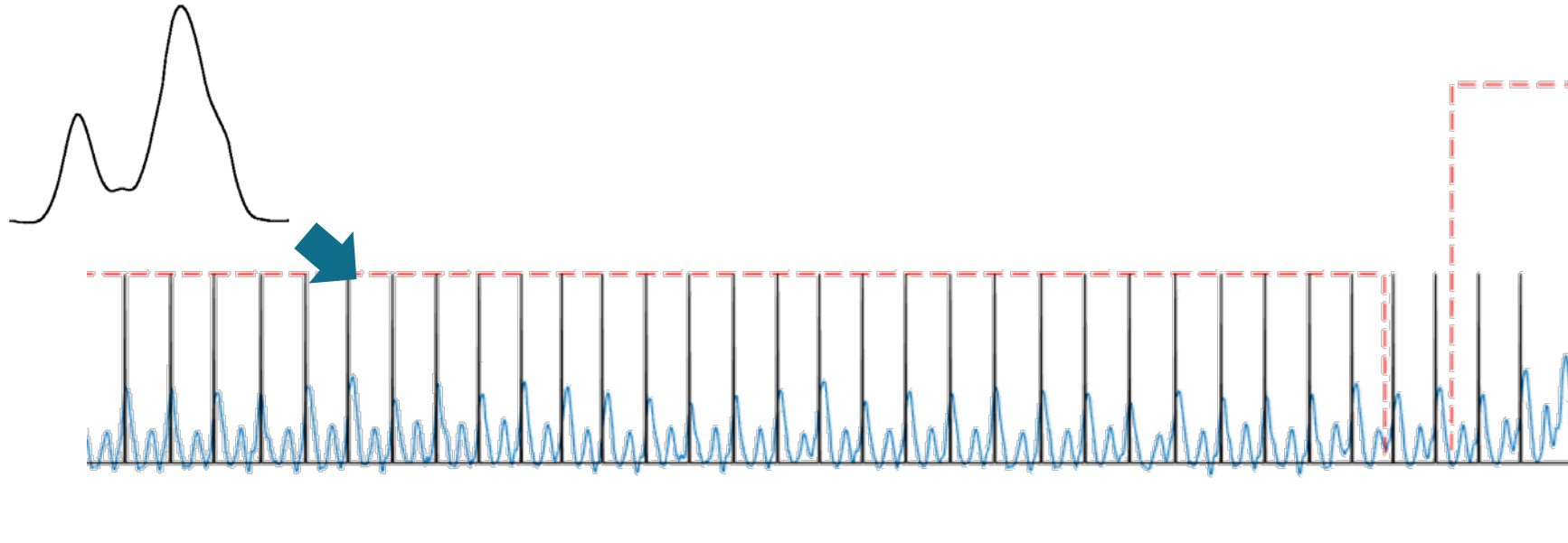
✓ Poses, repetitive activities

✓ Good accuracies

✗ Transitions

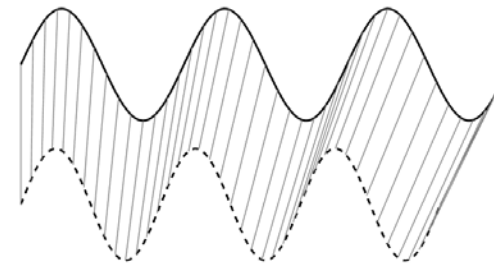
✗ Details

# Activity recognition: pattern matching



- Scale?
- Variability?

DTW  
➔





# Data

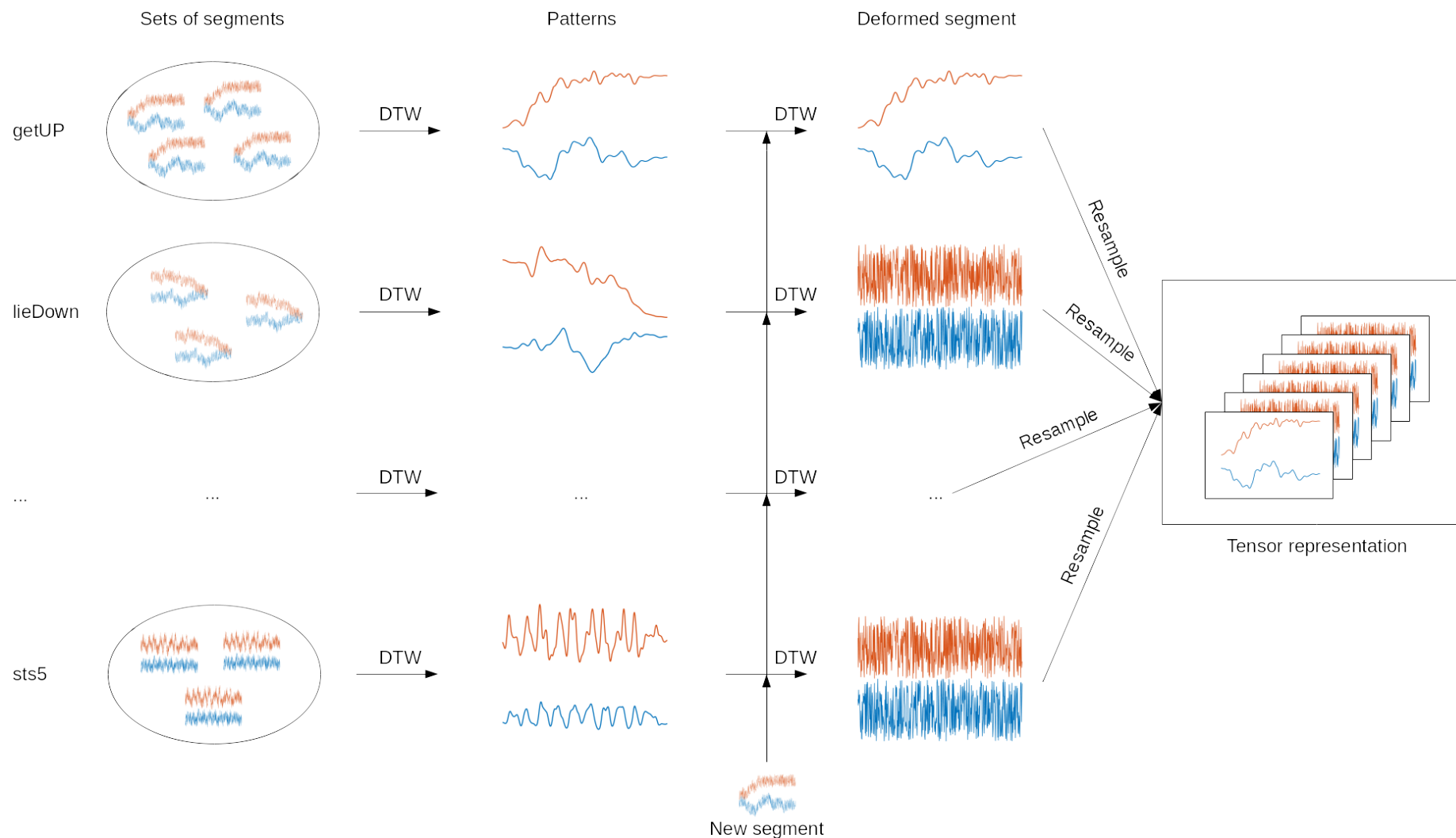


- 39 axSpA patients
- 28 equipped with SenseWear Pro (Bodymedia Inc.)  
10 equipped with Shimmer3 (Shimmer Inc)
- 6 BASFI activities

Get up	Maximal reach 5x
Lie down	Pick up a pen 5x
Maximal reach	Sit-to-stand 5x

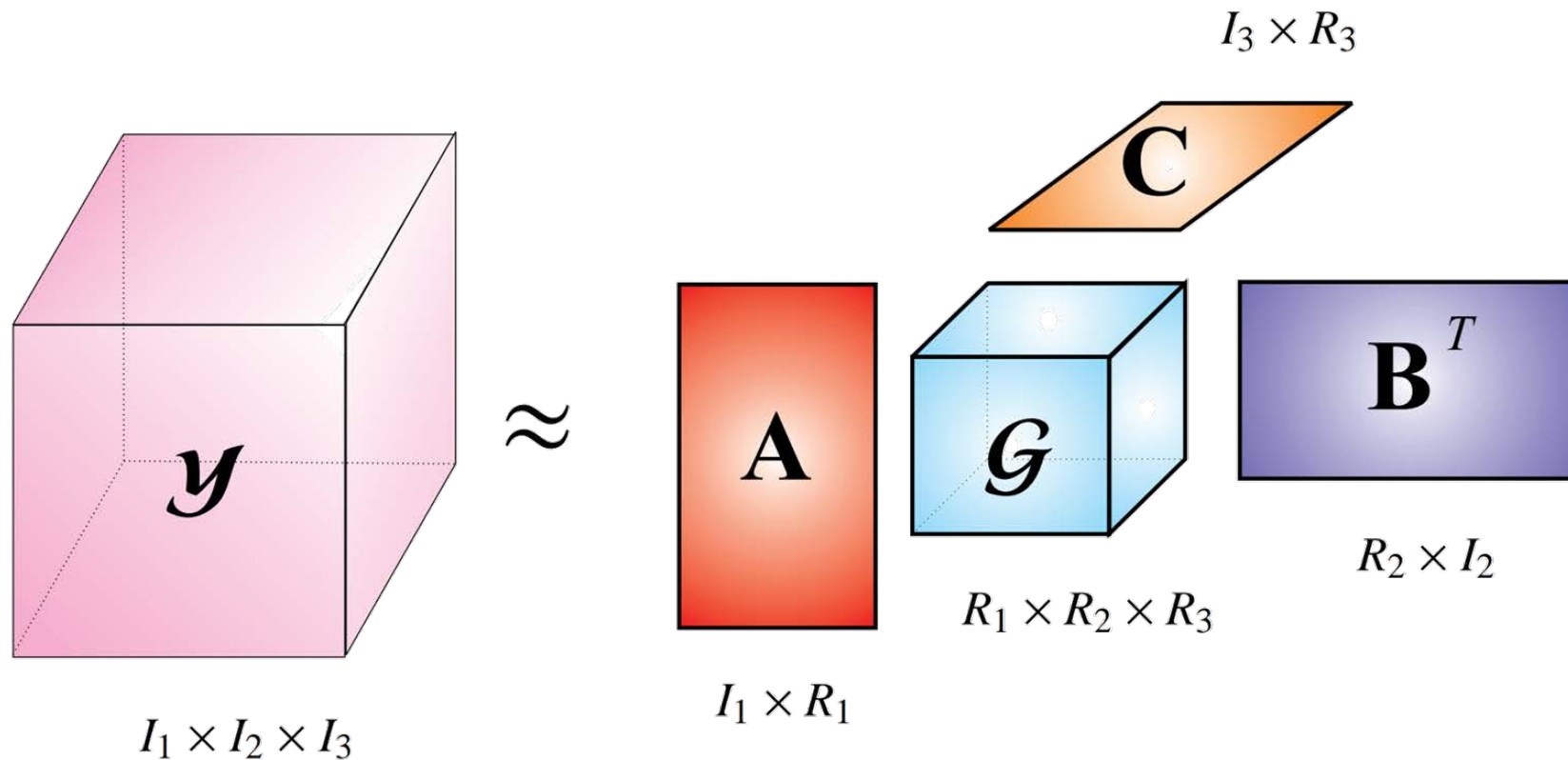
- 7-class problem (NULL class)

# Data representation



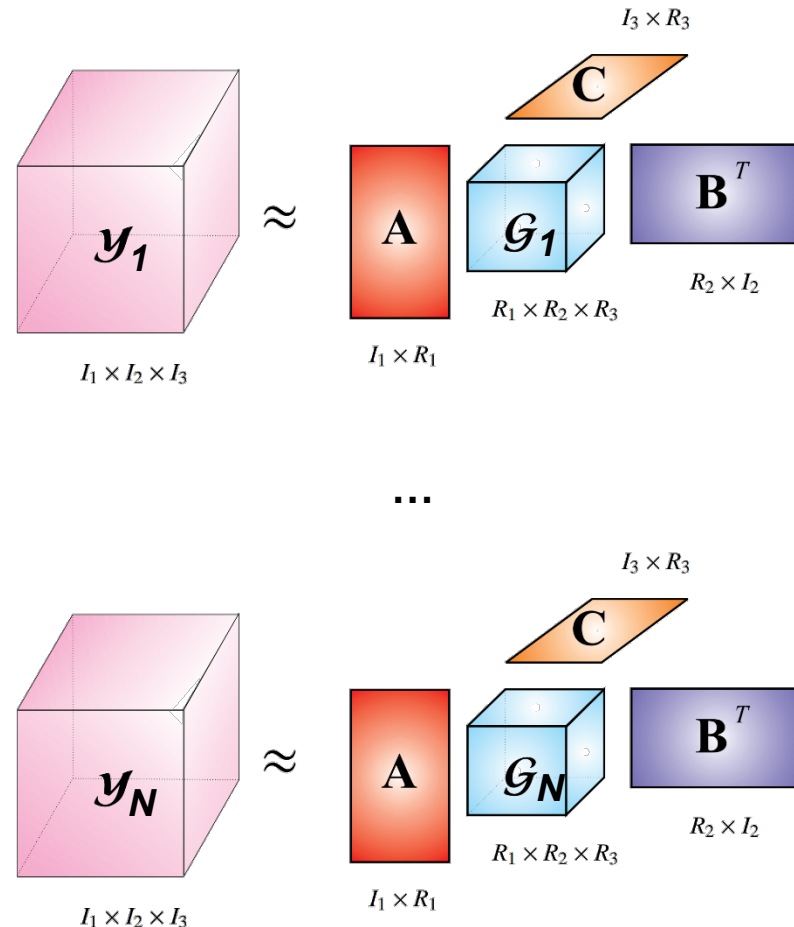
# HODA

- Tucker decomposition



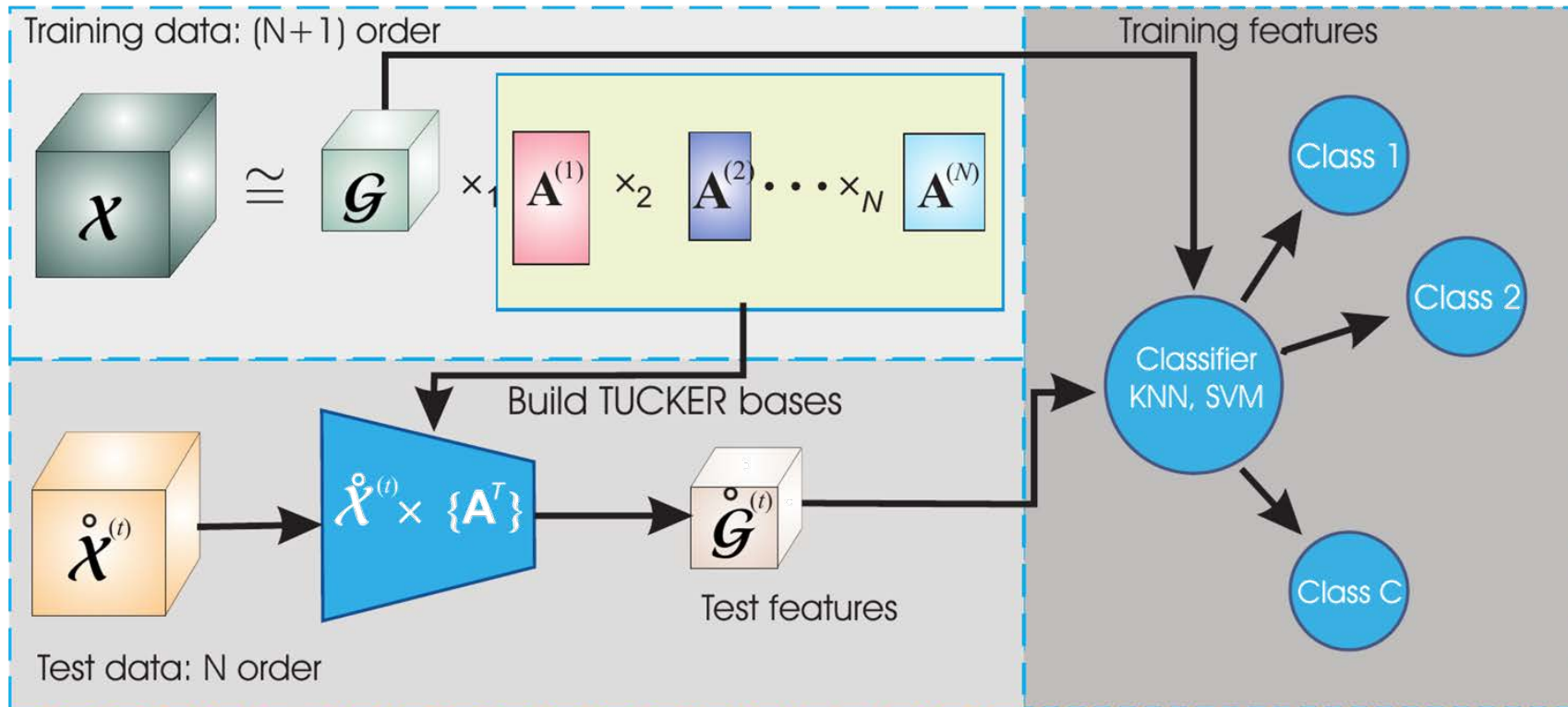
# HODA

- Joint Tucker decomposition for data compression



# HODA

- Joint Tucker decomposition with discriminant analysis







# Results

- Detection True Positive Rate:  $85.4 \pm 15.8\%$
- Number of false detections:  $2.90 \pm 3.32$
- Dice coefficient:  $89.3 \pm 12.0\%$
- | LoSo                    | HODA                                | SIMPLE (DTW distances) |
|-------------------------|-------------------------------------|------------------------|
| Accuracy <sub>1</sub> : | <b><math>86.7 \pm 14.2\%</math></b> | $82.9 \pm 20.4\%$      |
| Accuracy <sub>2</sub> : | <b><math>62.3 \pm 25.0\%</math></b> | $44.1 \pm 20.7\%$      |
- SenseWear 

LoSo	HODA
• Accuracy <sub>1</sub> :	<b><math>91.2 \pm 7.9\%</math></b>
• Accuracy <sub>2</sub> :	<b><math>74.5 \pm 17.0\%</math></b>
- Drawback: high computational load due to DTW, not suitable for real-time use

# Conclusions

- Patterns are a viable option for recognition of physical activity
  - Combined with statistical features
  - Using structure/HODA
    - quit segmentation/rejection?
- Future work
  - Impact of different sensors
  - Combining structure and statistical features

# Neonatal Brain Monitoring

UZ Leuven departments:

**Neonatology**

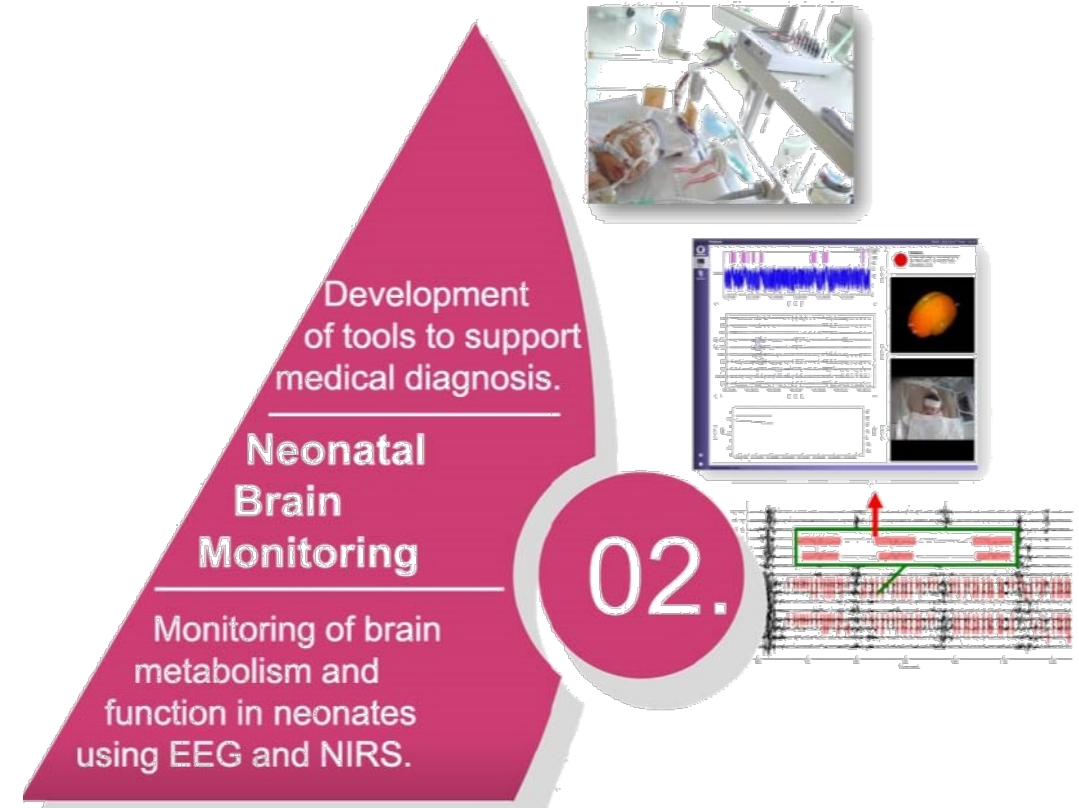
**Pediatric neurology**

Eindhoven University of Technology

Maxima Medical Centre

Oxford University:

Dept. of Engineering Science



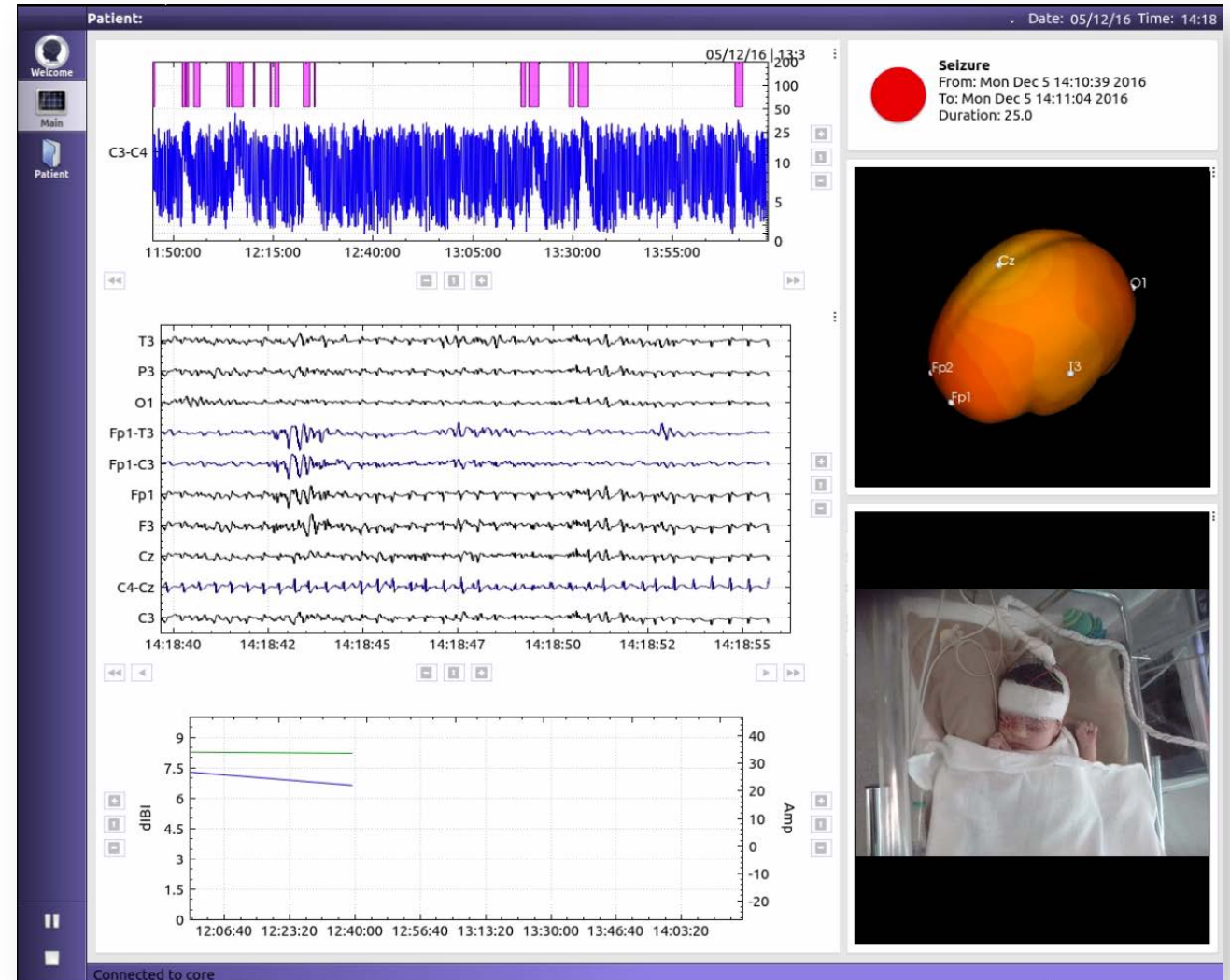
EXAMPLE

# NeoGuard : decision support

## Partners

KU Leuven-ESAT (Stadius & MICAS), UZ Leuven neonatology, EMC Rotterdam, ZNA Middelheim, Ghent University (TELIN)

- Brain injury estimate
  - Detection of neonatal epileptic seizures
  - Seizures localization
  - Inter-burst intervals
- Incorporated expertise
  - Knowledge of neurophysiologists are incorporated into algorithms
- Monitoring
  - Recovery after brain damage
  - **Brain Maturation in prematures**
  - **Sleep staging**
- Outcome prediction





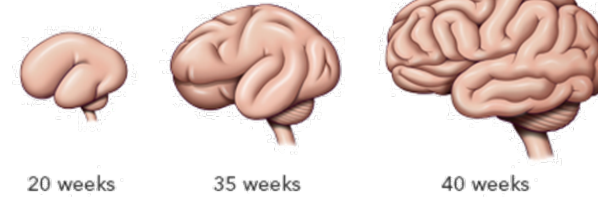
# Decomposition of a Multiscale Entropy Tensor for Sleep Stage Identification in Preterm Neonates

Collaborator: Ofelie De Wel

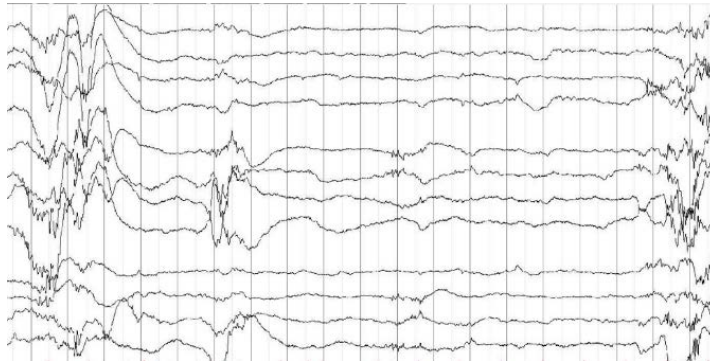
# Problem Statement

- Sleep → brain development
- Neonatal sleep
  - Quiet sleep
  - Nonquiet sleep (awake, active sleep & indeterminate sleep)

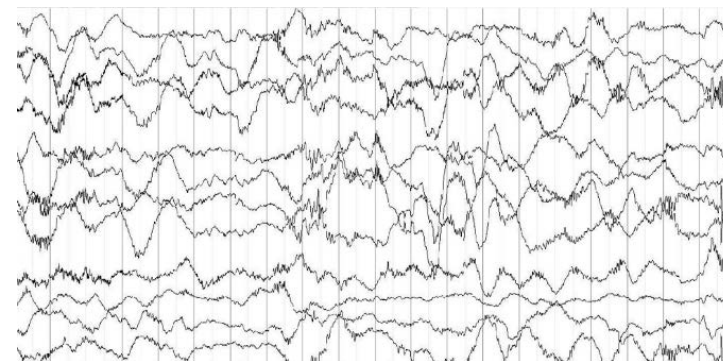
Infant brain growth



Quiet sleep

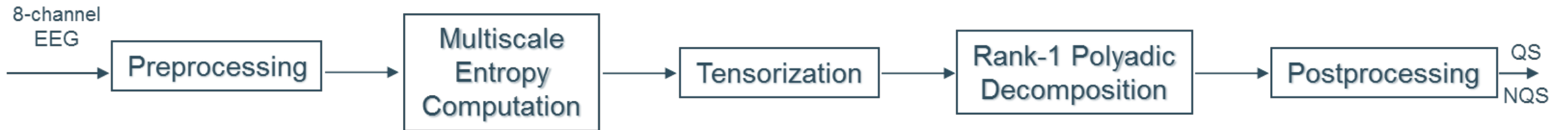


Nonquiet sleep



# Problem Statement

- Objective: discriminate quiet sleep from nonquiet sleep





# Database

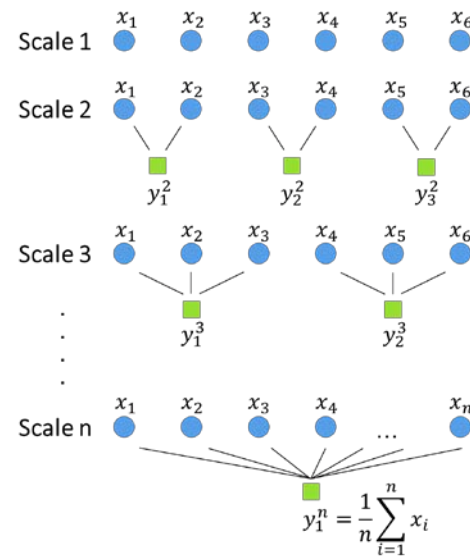
- 88 EEG recordings from 25 preterm neonates
- 8 monopolar EEG channels
- Postmenstrual age: 27 – 42 weeks
- Quiet sleep annotated by expert clinician

# Preprocessing

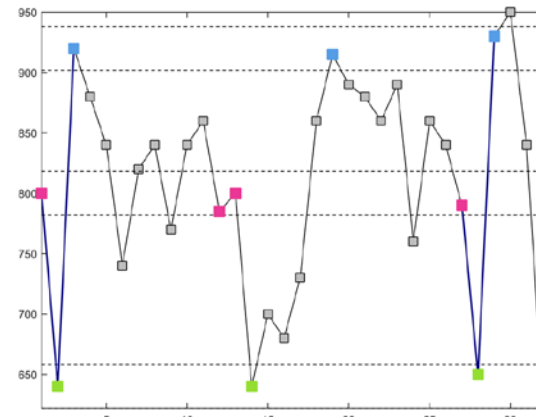
- 1-20 Hz bandpass filtering
- Resampling to 125 Hz
- Segmentation into 100s windows

# Multiscale Entropy Computation

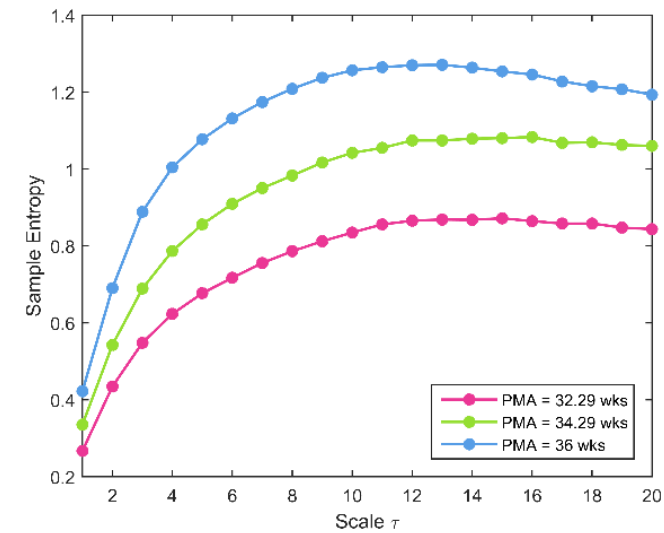
1) Coarse Graining



2) Sample Entropy Computation

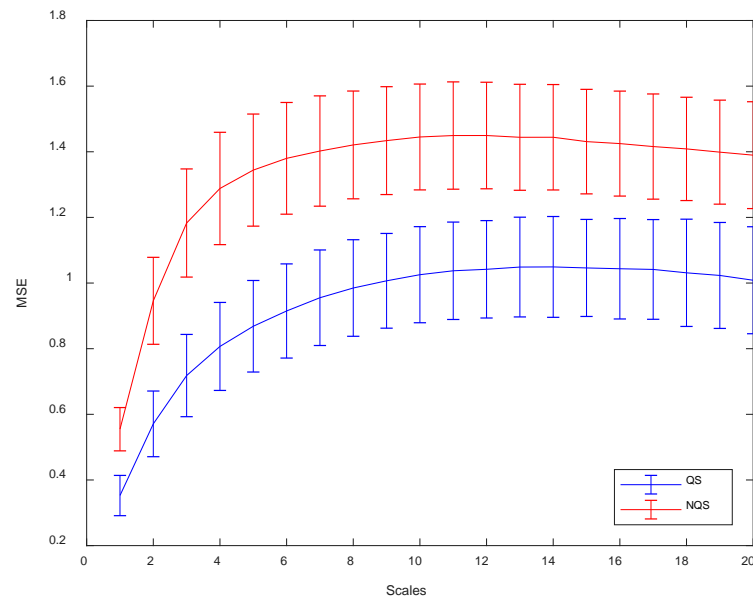


3) Multiscale Entropy Curve

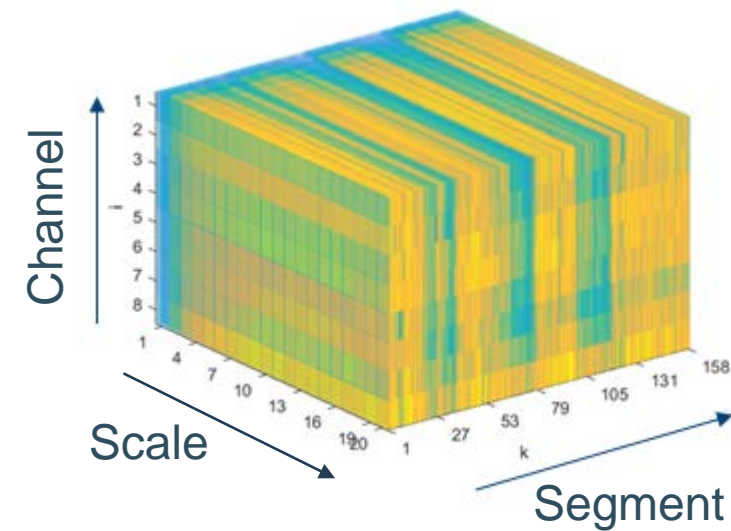


# Tensorization

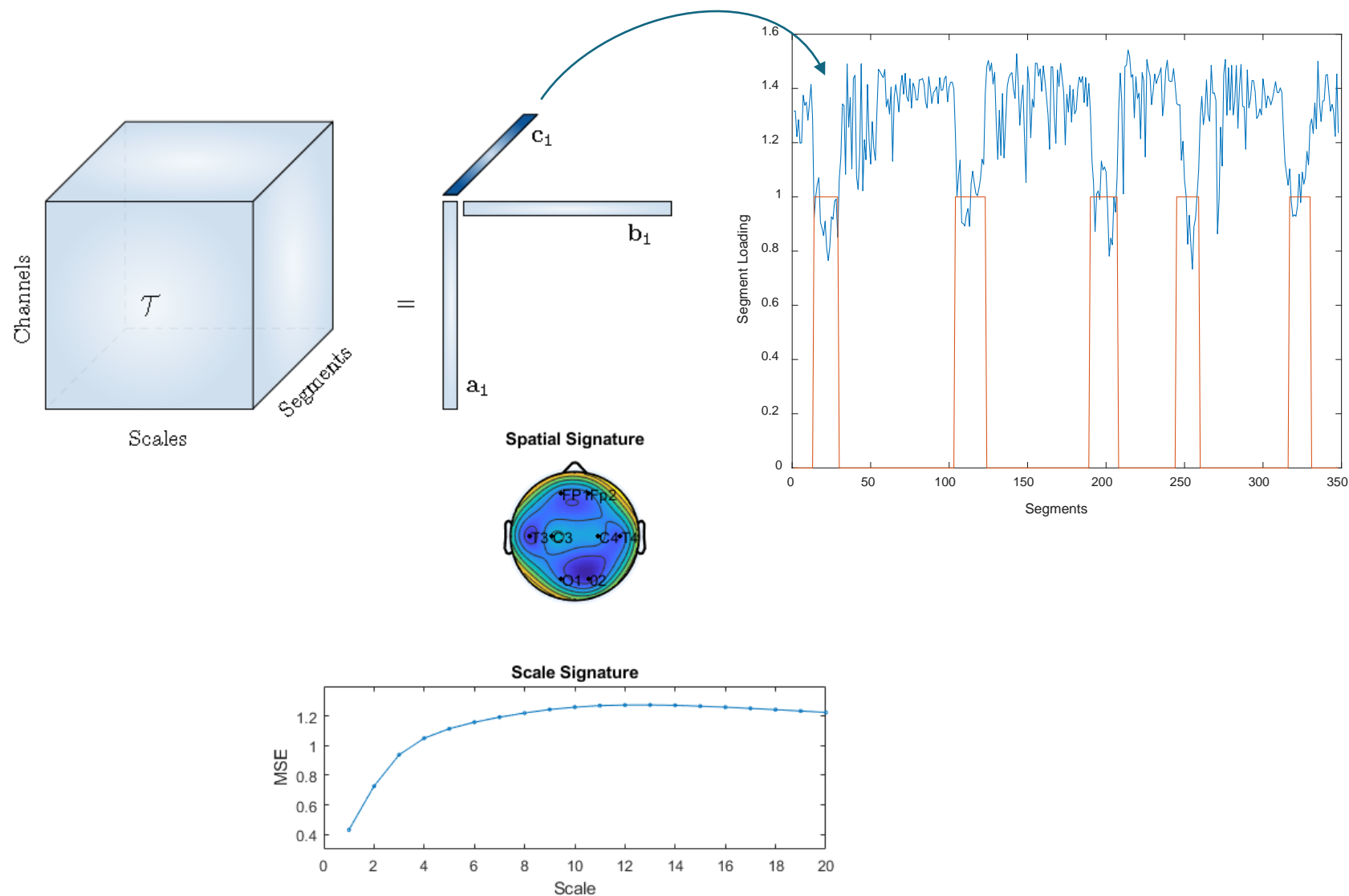
MSE curve is reduced during QS  
compared to NQS



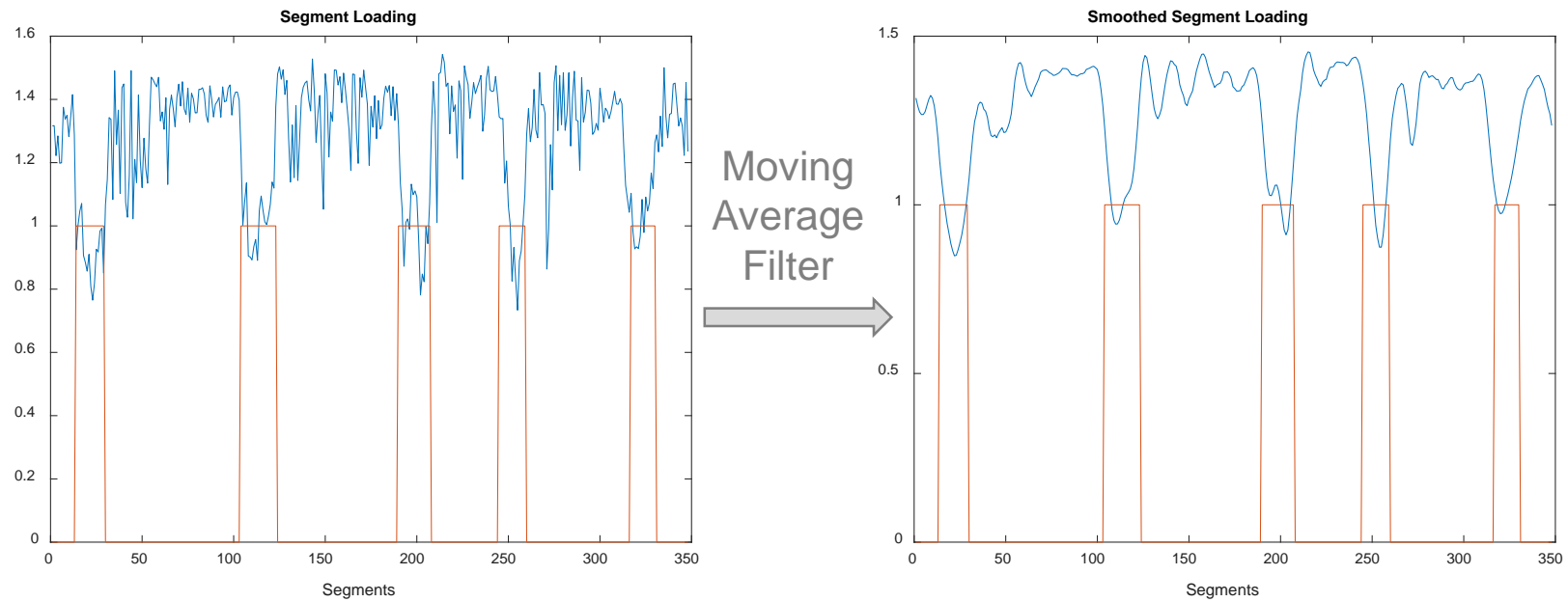
3rd order tensor:  
8 channels x 20 scales x segments



# Rank-1 Polyadic Decomposition



# Postprocessing



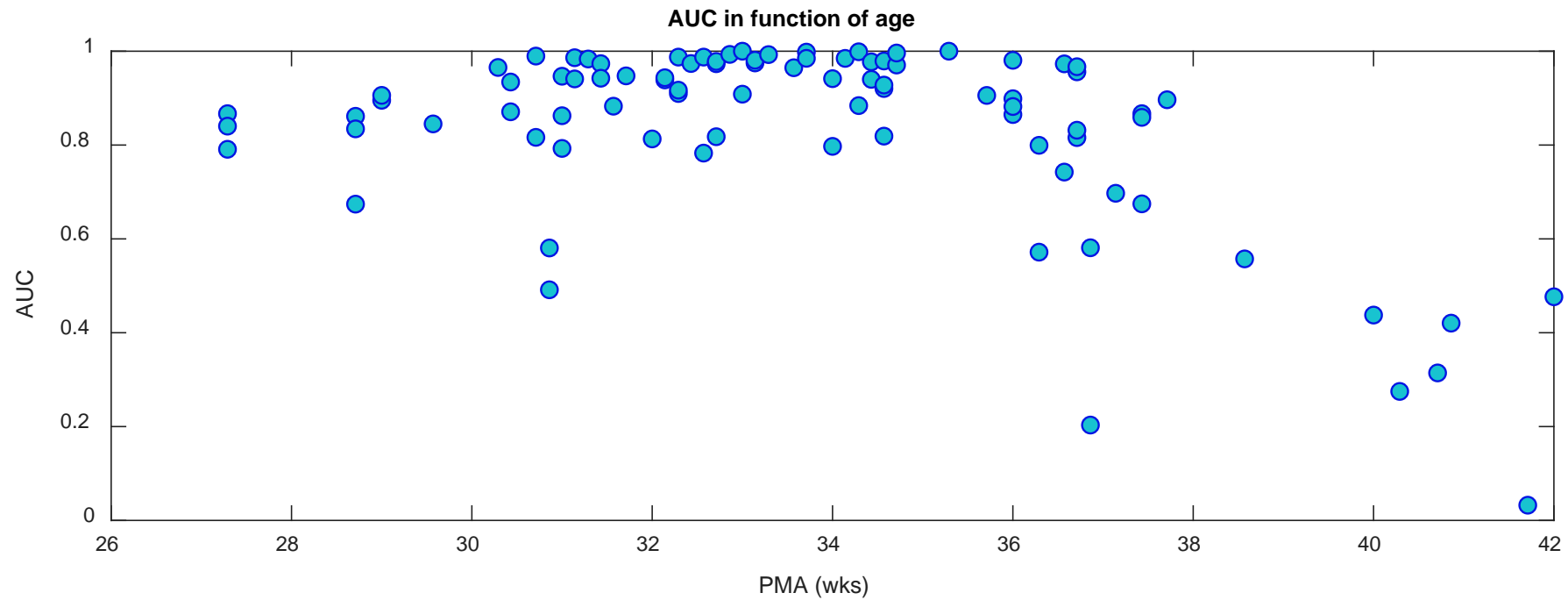


# Performance

- K-means clustering

Performance Measure	Average over all recordings
Silhouette Coefficient	0,81
Sensitivity	0,71
Specificity	0,82
Accuracy	0,79

# Performance

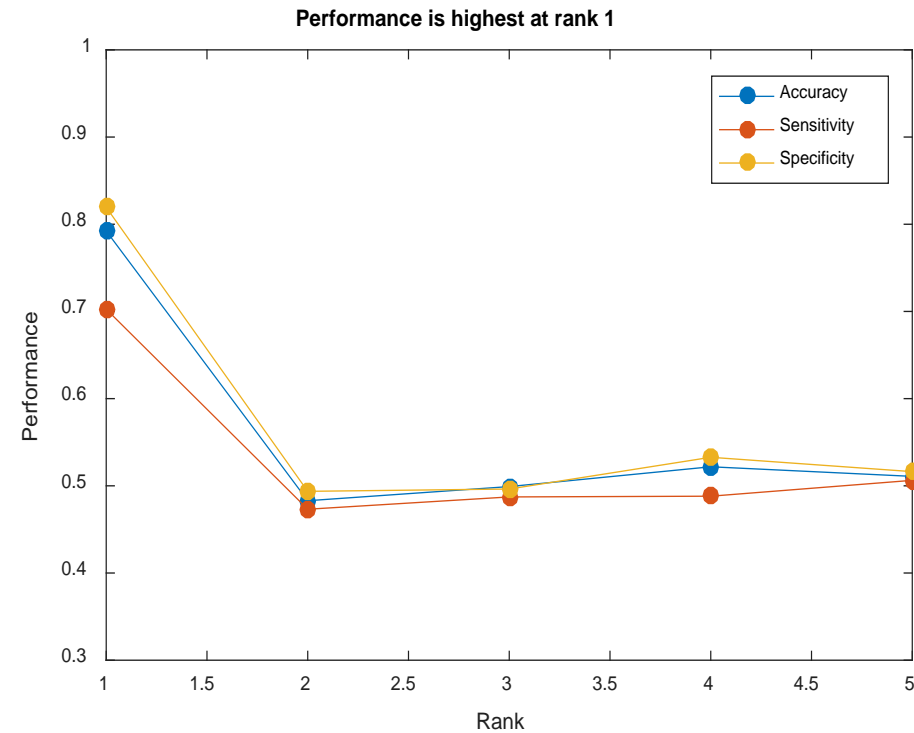
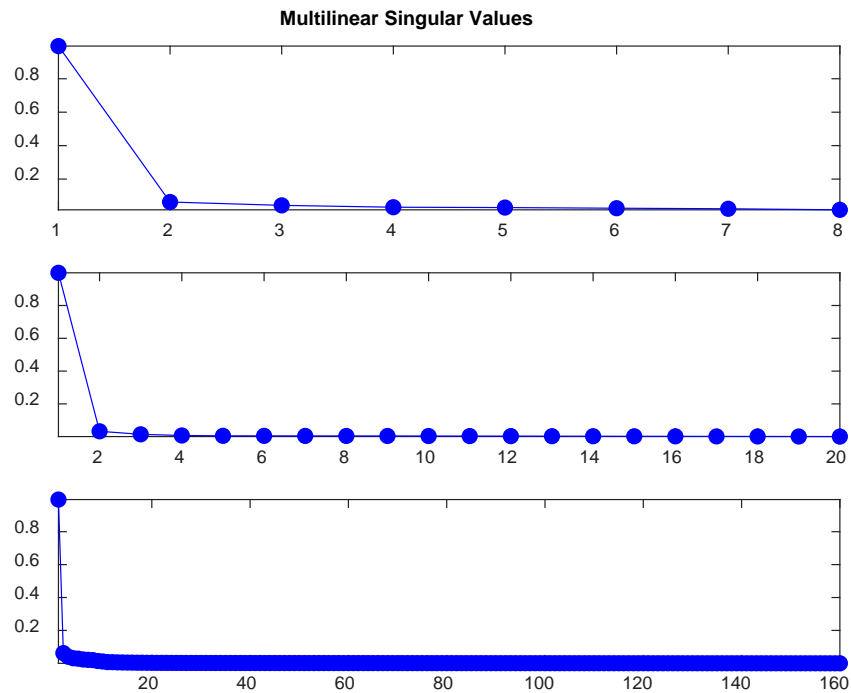


- Average AUC = 84%



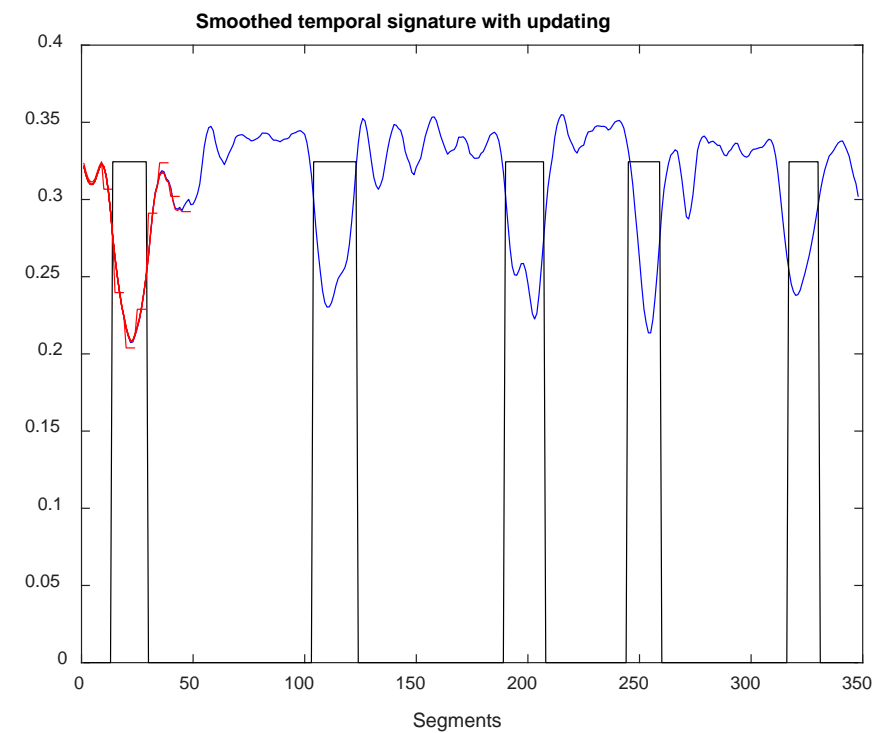
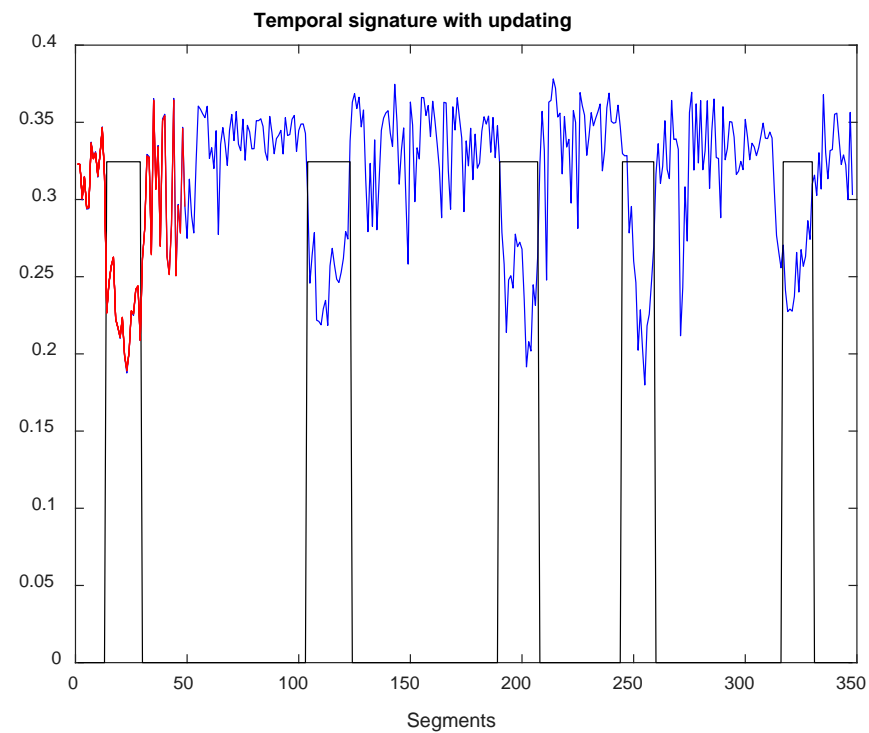
# Choice rank

- Mean Relerr = 0,11



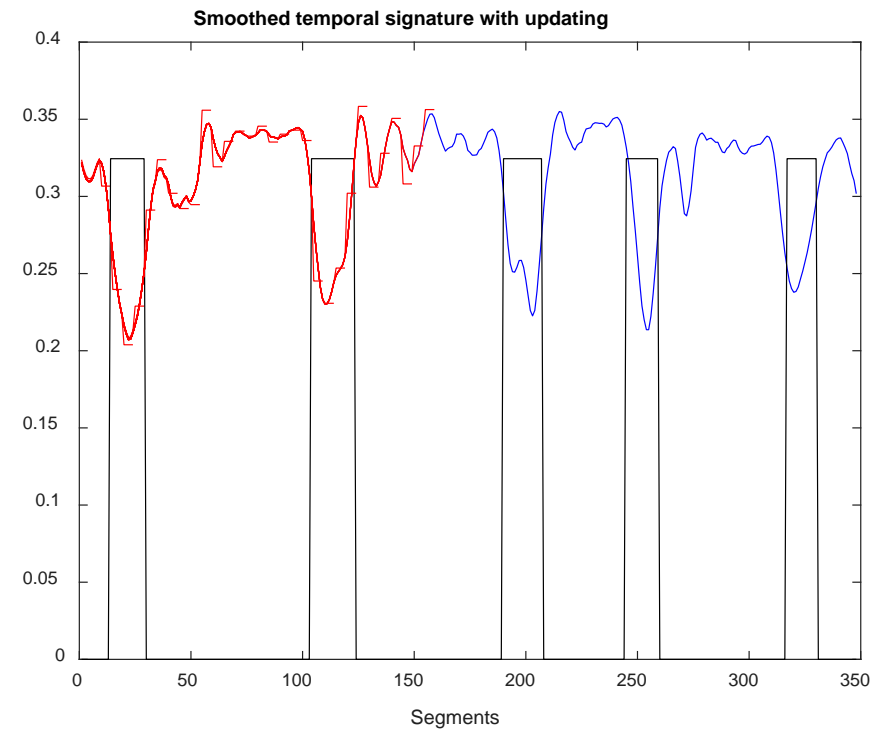
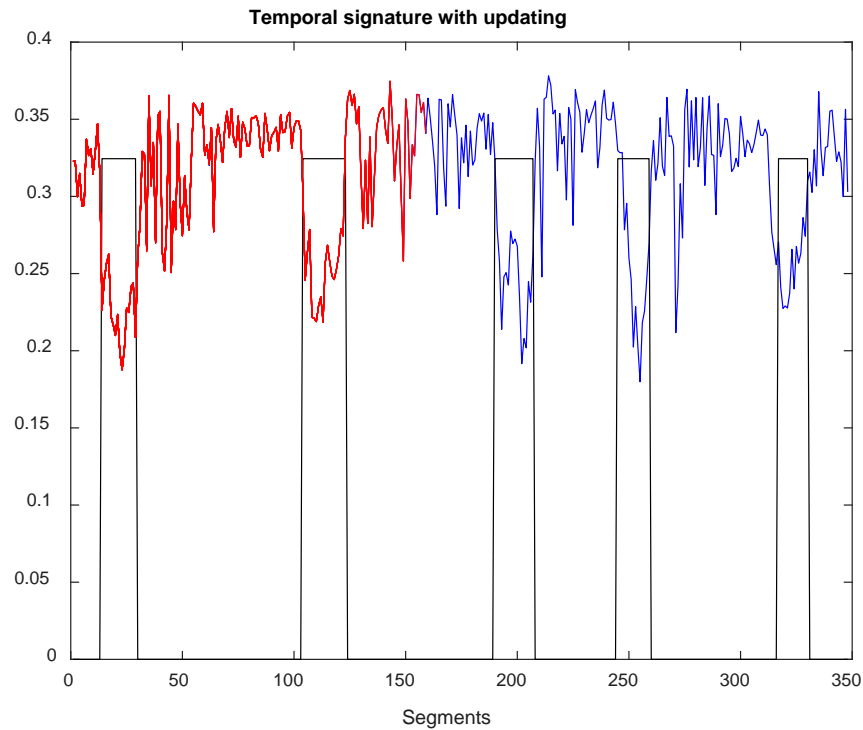
# Updating

- Real time sleep monitor → update CPD



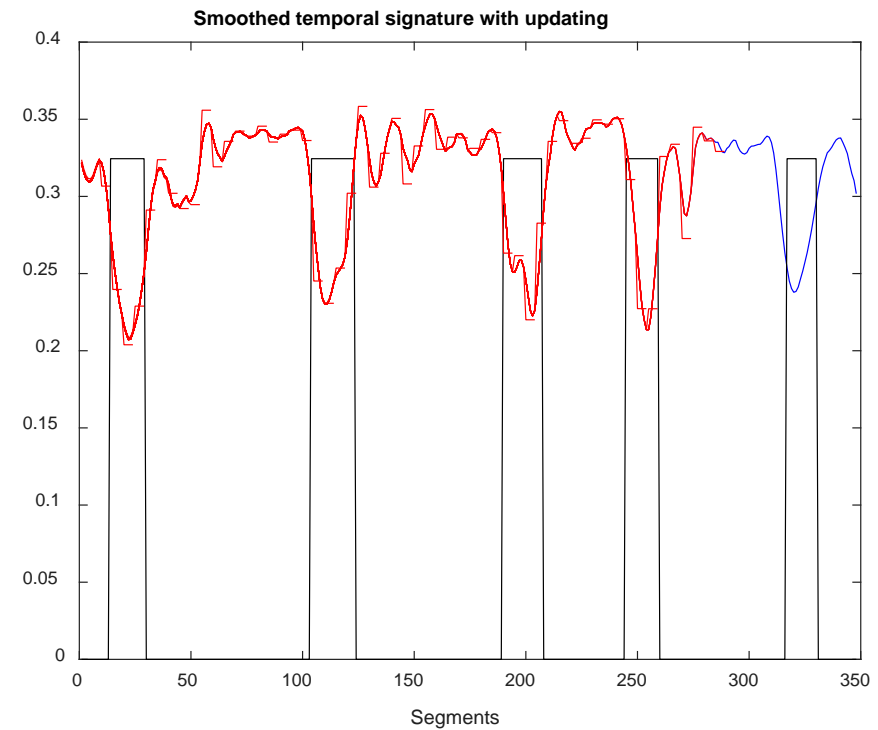
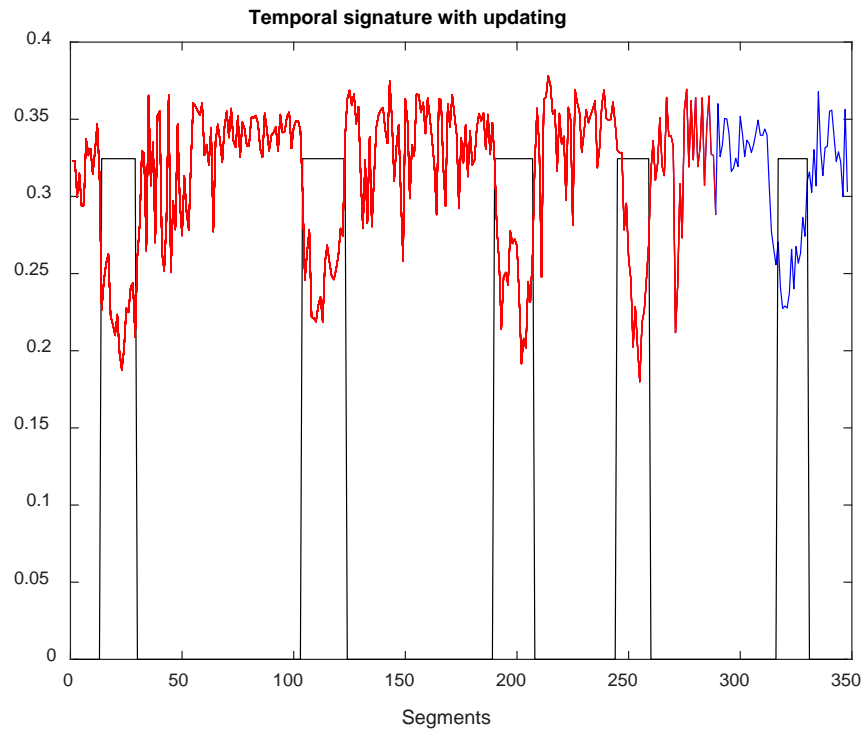
# Updating

- Real time sleep monitor → update CPD



# Updating

- Real time sleep monitor → update CPD



# Conclusions

- EEG complexity is different depending on sleep state
- Rank 1 CPD of multiscale entropy tensor can be used to discriminate QS from NQS in an unsupervised way.

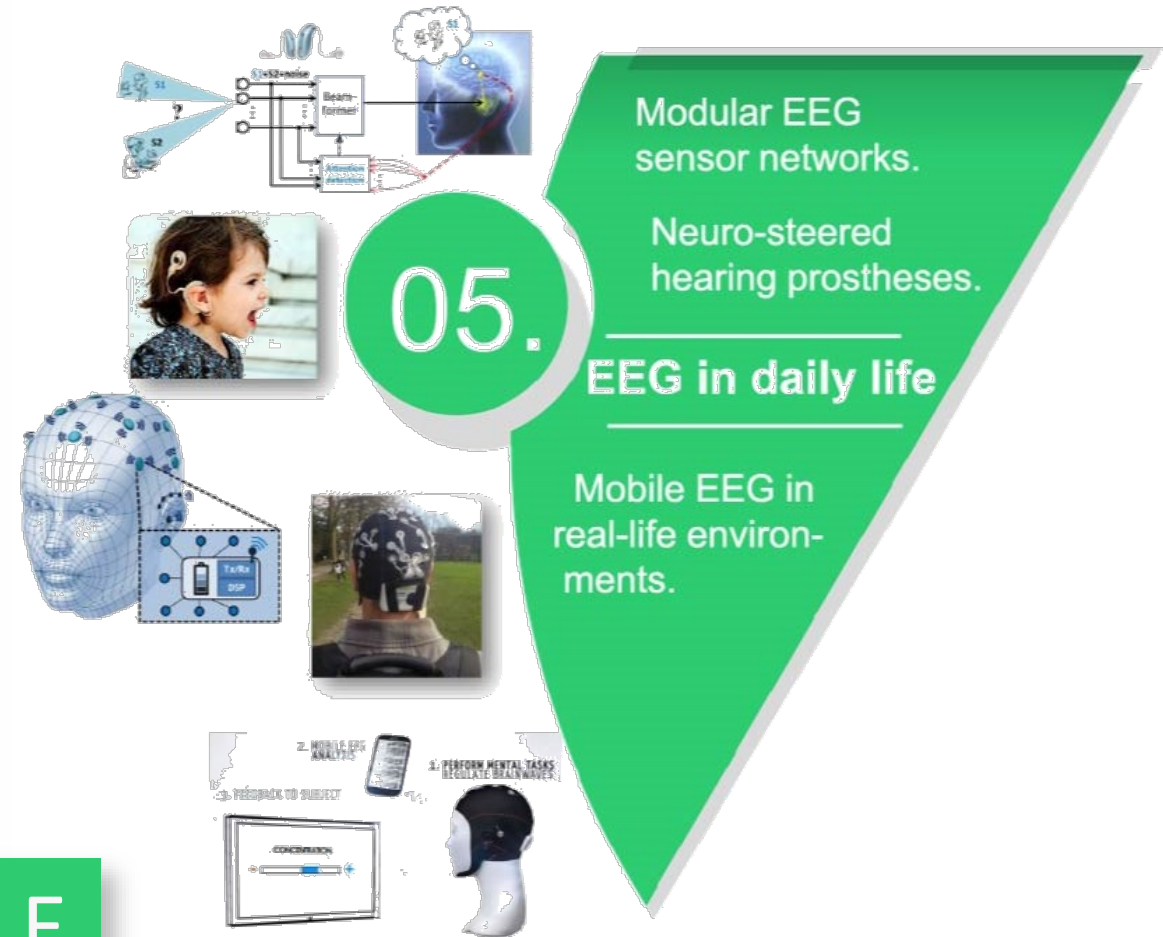
# EEG in daily life

**UZ Leuven partners:**

Tom Francart (experimental otorhinolaryngology)

Oxford University Dept. of Engineering Science:  
Maarten De Vos

EXAMPLE





# Use of Tensor Decompositions in auditory BCI (mobile EEG)

Rob Zink | KU Leuven



## DISSERTATION

Mobile EEG and Tensor Approaches for Auditory Attention Analysis in Real-life

Zink, Rob; Van Huffel, Sabine (Supervisor); De Vos, Maarten (Co-Supervisor)

December 2017





# Brain computer interfaces

Brain computer interfaces (**BCI**) **utilize brain activity patterns** to **control** a technical device by **thought alone**. Most BCI applications rely on electroencephalogram (**EEG**) signals in response to motor imagery or visual sensory input, but recently non-invasive auditory BCIs using **auditory attention** and imagery EEG responses have been developed.

Truly **mobile BCI systems** are emerging to allow BCIs that can be operated in **real-life environments** by means of **mobile EEG**.

# & Human performance

**Understand** the more complex EEG (i.e. *both signal and noise*) patterns in **real unconstrained environments**.

The **challenge** to **disentangle** observed **changes in EEG activity** due to increase in **cognitive demands** by the complex natural environment or due to **physical involvement**.



## Stimulus:



## Non-Attended Targets do not (fully)



**Attended Targets generate a P300 ERP**  
**Non-Attended Targets do not (fully)**



# Data Driven Classification

Structured decompositions of mobile EEG

## Divide Dataset into Train-Test set

Train classifier on Training Data → Obtain discriminative features/model

→ Apply classifier model on Test data

## Cons for (mobile) BCI applications:

1. **Always need Training Data** (costs time/effort subject)
- 2a. Classifier is **dependent on the Training data**.
- 2b. **High variability** between subjects and even within sessions (non-stationarity)

## Our aim:

**Can we model the P300 differences in a data-driven (structured) way, removing the training phase?**



# Data Driven Classification

Structured decompositions of mobile EEG

## Data (external):

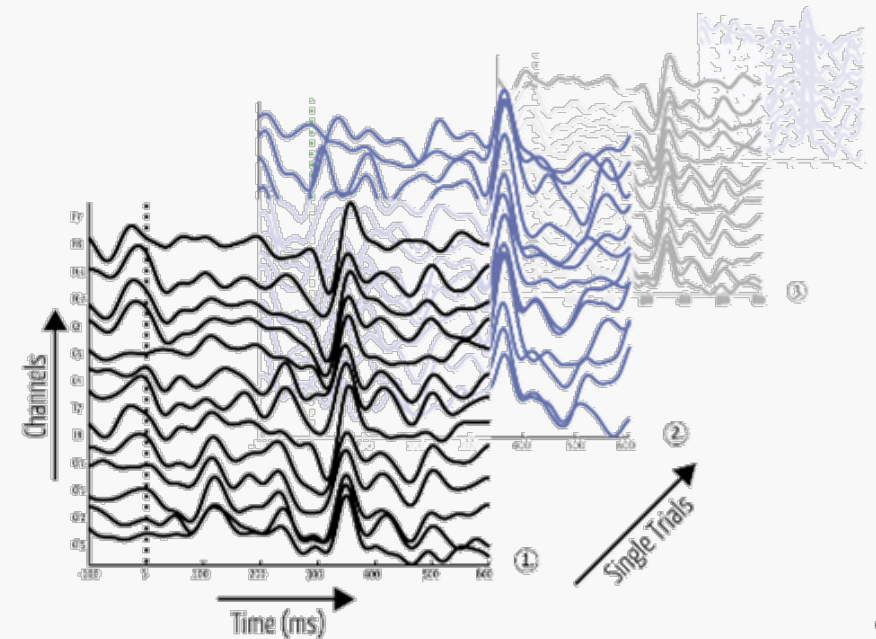
- 20 subjects (mean age 24.6 years)
- 14 channels (wet electrodes) – F3, Fz, C3, FPz, Tp10, Cz, O1, O2, F4, C4, TP9, Pz, P4, P3 (Afz, FCz).
- Sampling rate 128Hz
- Eye blinks removed using ICA
- Data epochs (trials) -200 to 800ms with respect to stimulus
- Baseline correction, Rereferencing, 1-20Hz BPfilter.
- **Sit and Walking outside**

## Data from:

De Vos, Maarten, Katharina Gandras, and Stefan Debener. "Towards a truly mobile auditory brain-computer interface: Exploring the P300 to take away." *International Journal of Psychophysiology* 91.1 (2014): 46-53.

Structure data as **channels x time x trials tensor**

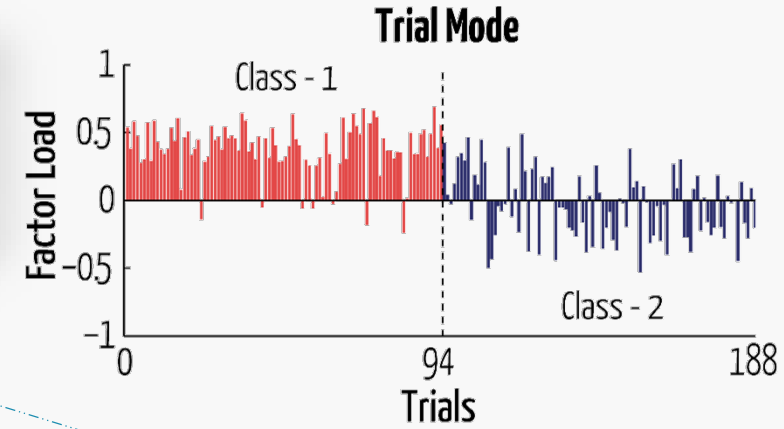
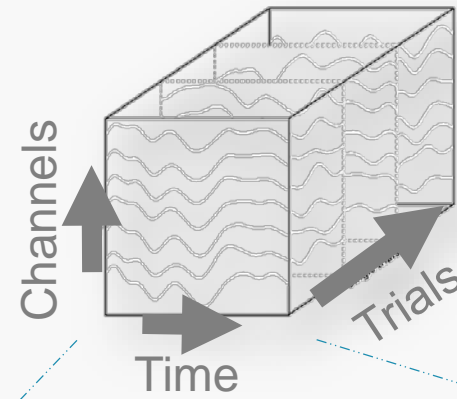
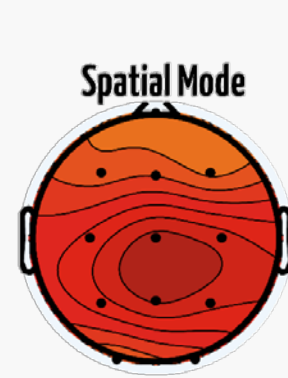
3D - Data tensor: Channels x Time x Trials



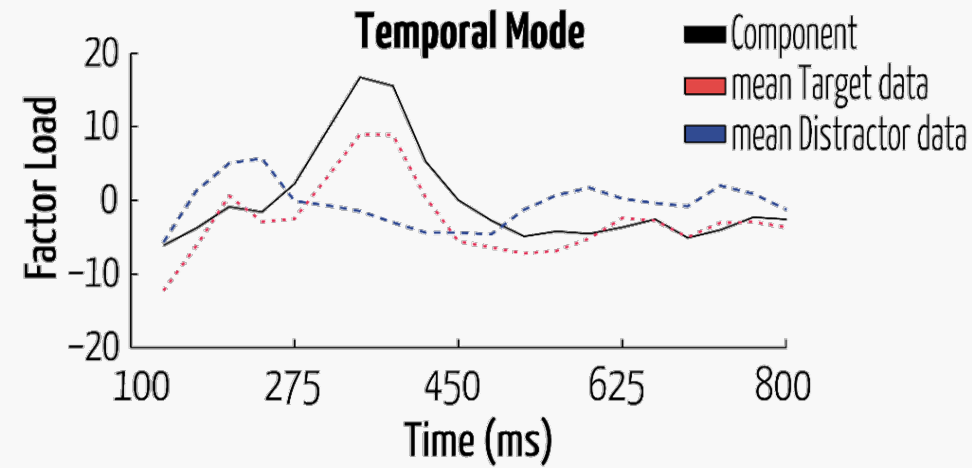


# Proof of concept of CPD on (auditory) ERP data

12 channels x 15 time points x 188 trials/epochs



**81% Accuracy**





# Data Driven Classification

## Parameters

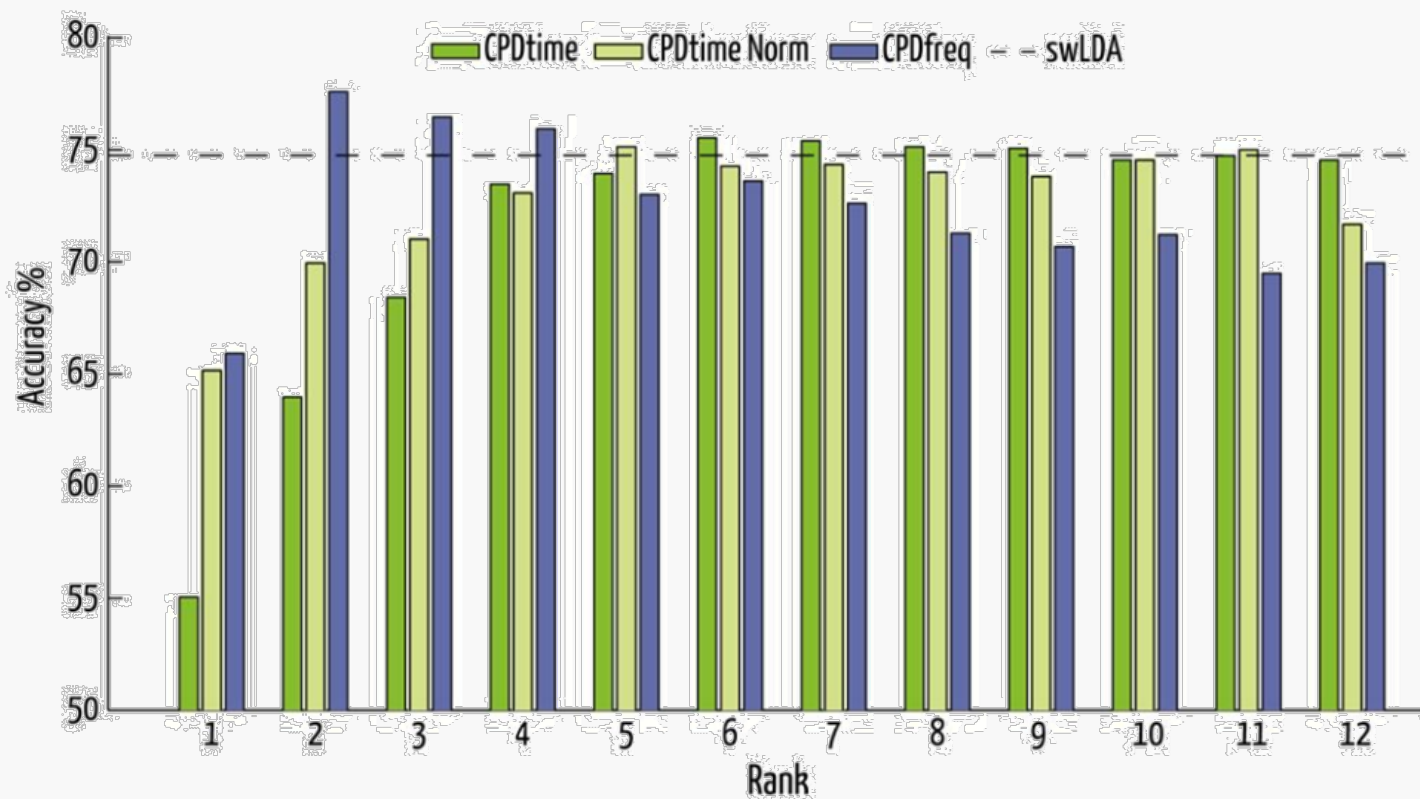
- Results based on 100 random **initializations**.
- The rank ( $R$ ) is expected to differ with the complexity of the dataset.  
(Semi)-automatic rank determination algorithms → tend to greatly overestimate the rank.
- First work we evaluated the results for **different ranks ranging from 1 to 12**
- **Channels x (norm)Time x trials**
- **Channels x Frequency x trials**





# Data Driven Classification

Proof-of-concept



## Published in:

Zink, R., Hunyadi, B., Van Huffel, S., & De Vos, M. (2015, April). **Exploring cpd based unsupervised classification for auditory BCI with mobile EEG**. In *7th International IEEE/EMBS Conference on Neural Engineering (NER)* (pp. 53-56). IEEE.

## Summary

- ✓ Proof-of-concept of CPD based models for ERPs → novel way to **analyze mobile auditory BCI data unsupervised**.
- ✓ Time and frequency decompositions provide signal components that represent approximately similar underlying source activation.
- ✓ Concept holds for small tensor with only few trials (ten trials). (not shown)
- × **Only possible offline scenarios**
- × inability to identify extracted clusters.

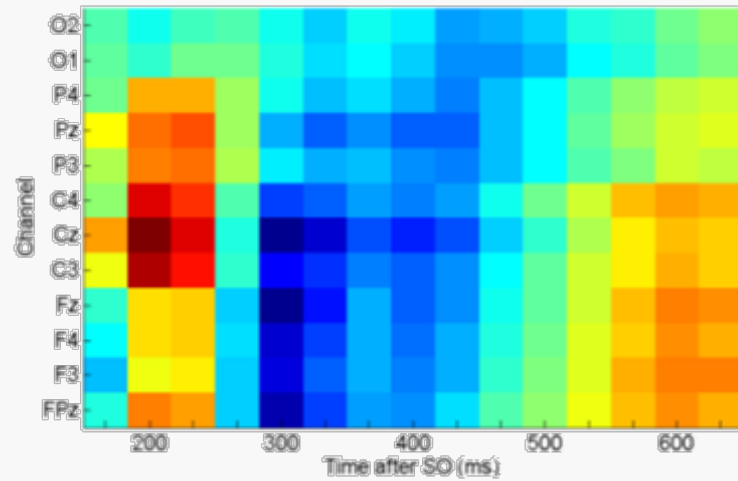


# Data Driven Classification

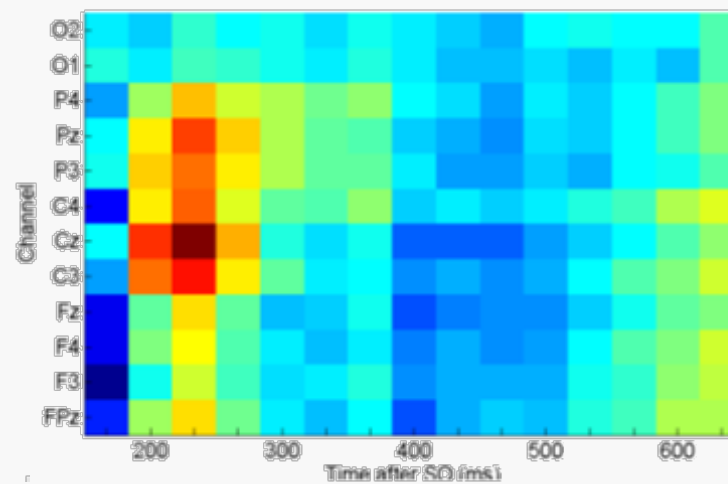
Adding general knowledge about stimuli classes

## Grand-Average ERP matrices

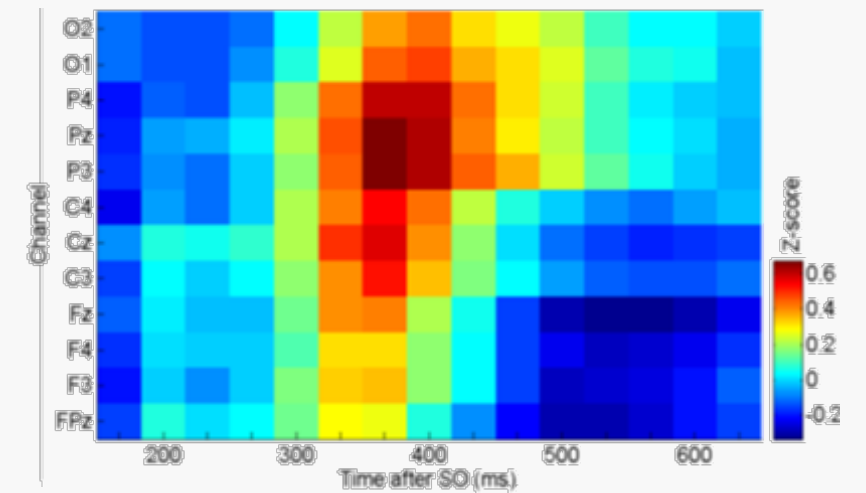
Baseline



Non-target

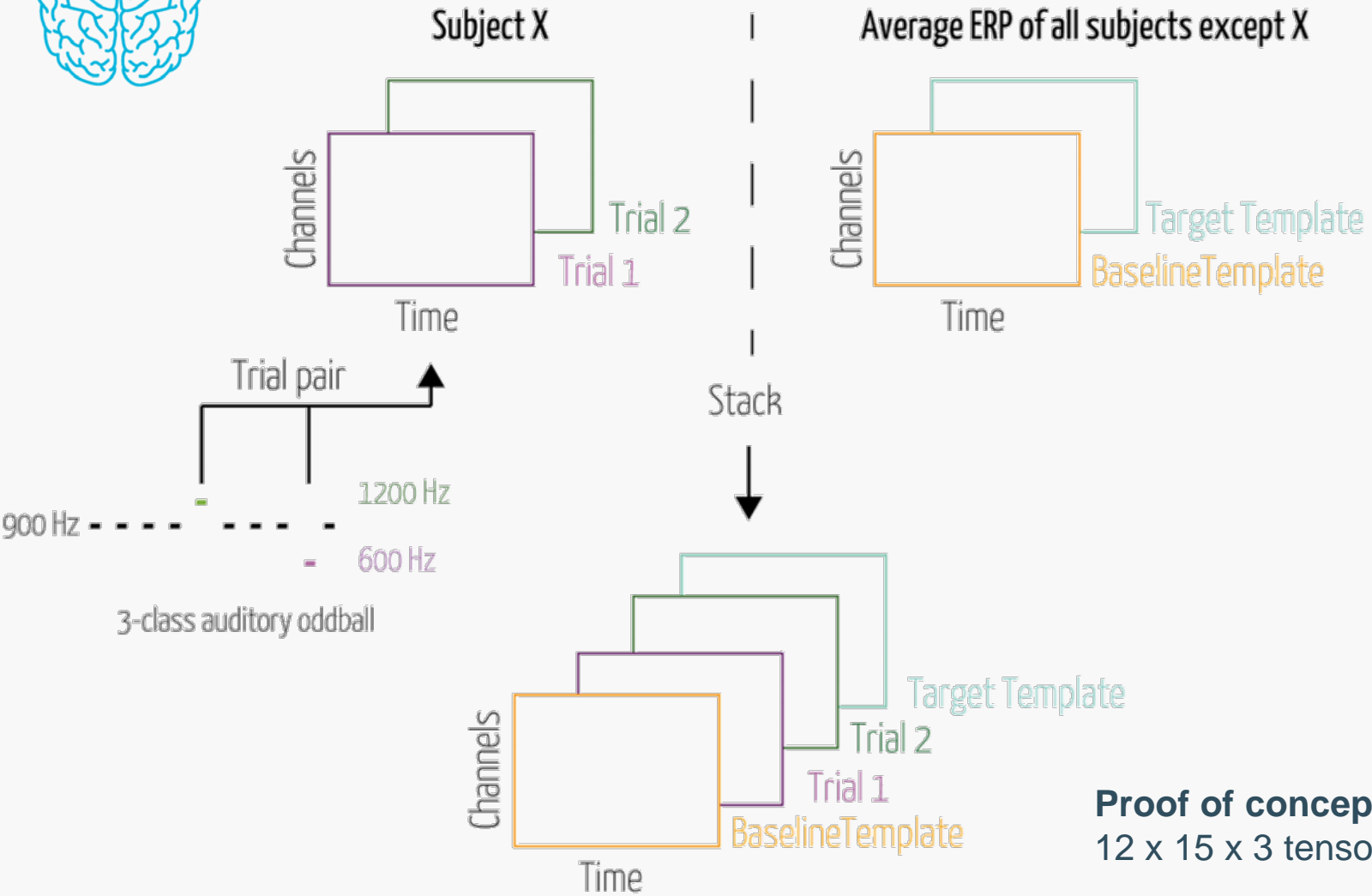


Target





**Online scenario:** Identify classes with non-specific templates

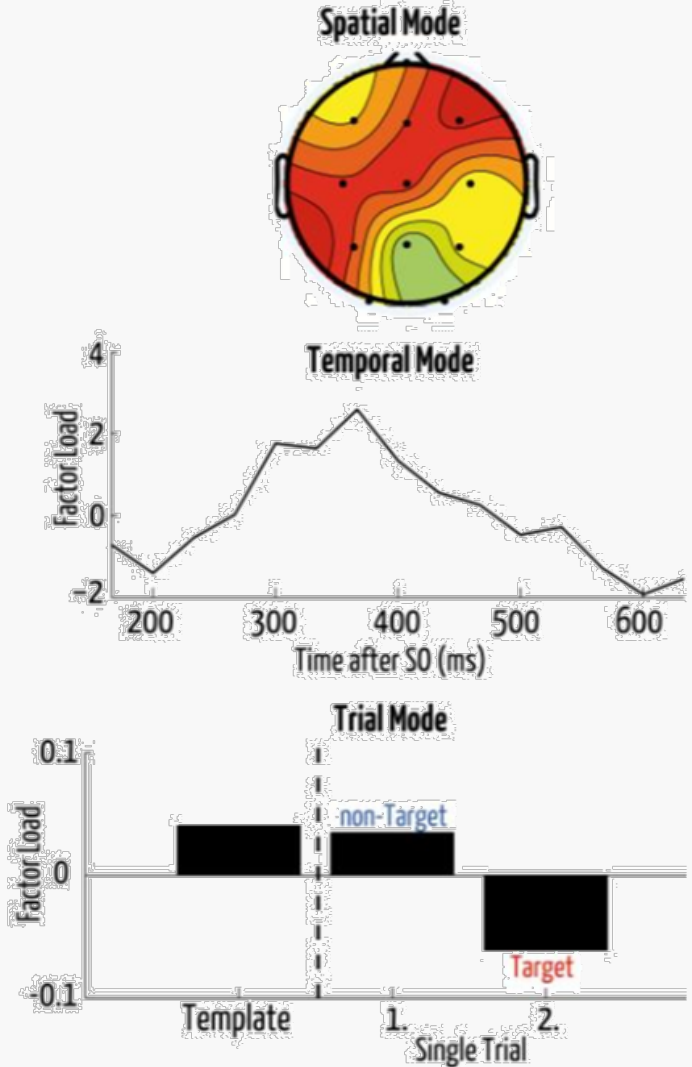


**Proof of concept**  
12 x 15 x 3 tensor

**Rank 1**

**Example Component**

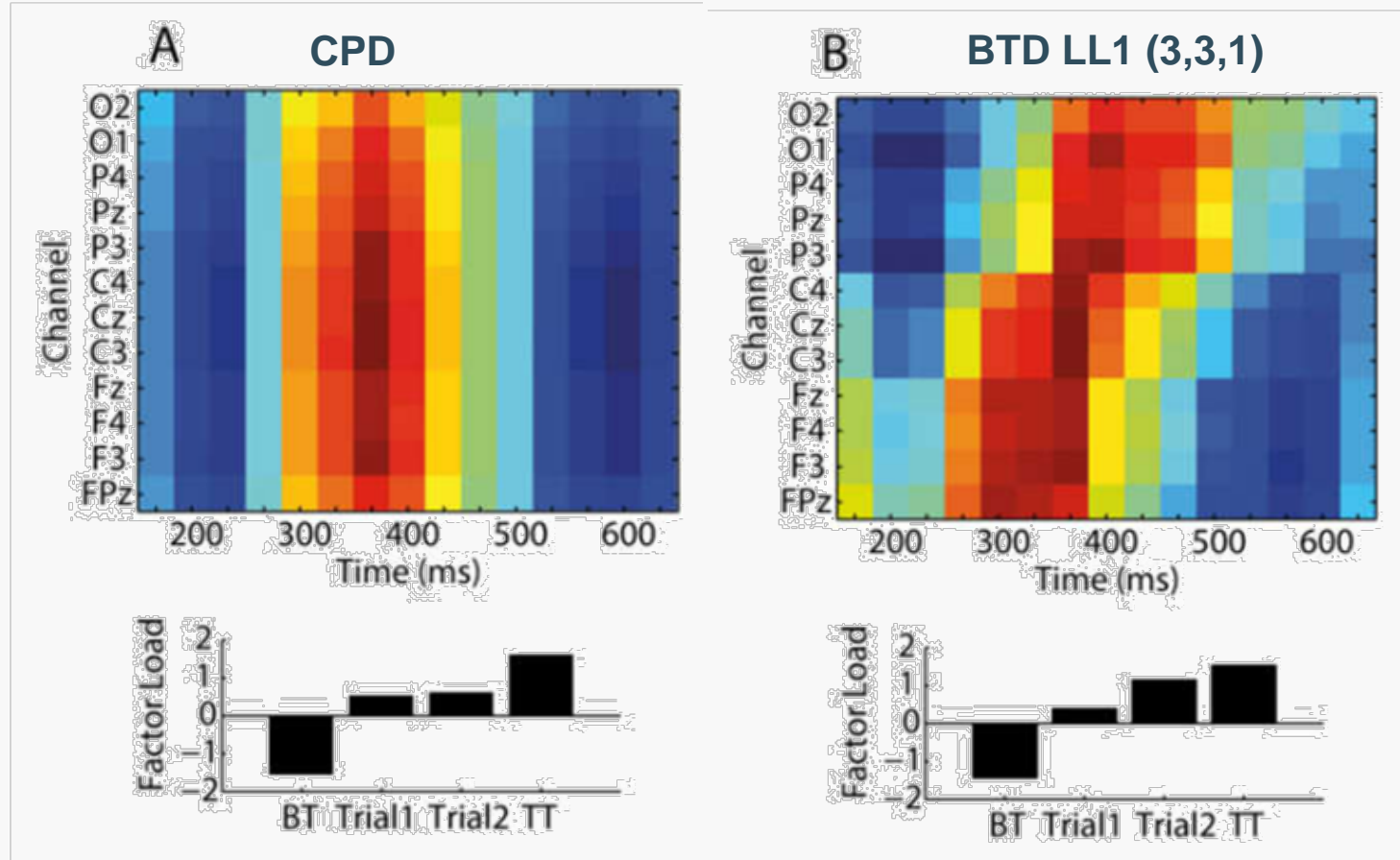
CPD: Channels x Time Norm x Trials





# Compare CPD and LL1-Block Tensor Decomposition on ERP data

## Example components



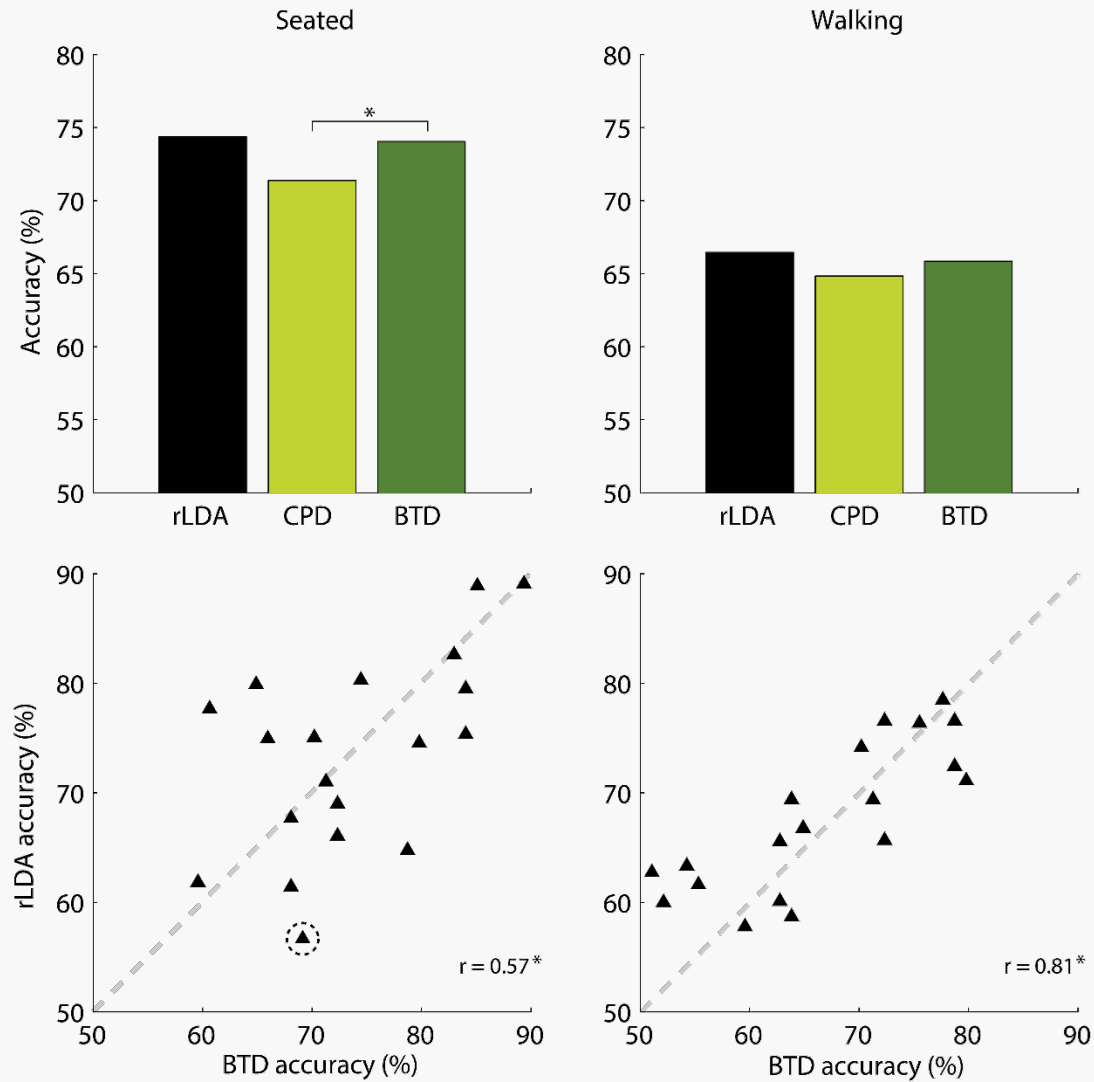
Parietal Occipital shift of activity.

Per binary command:  $12 \times 15 \times 4$  tensor:  
2 known-templates and 2 unknown-trials  
Values of 2 to 5 for  $L_r$  were explored.



# BTD to BCI

- Higher accuracy of BTD as compared to CPD
- Individual differences between rLDA and CPD/BTD

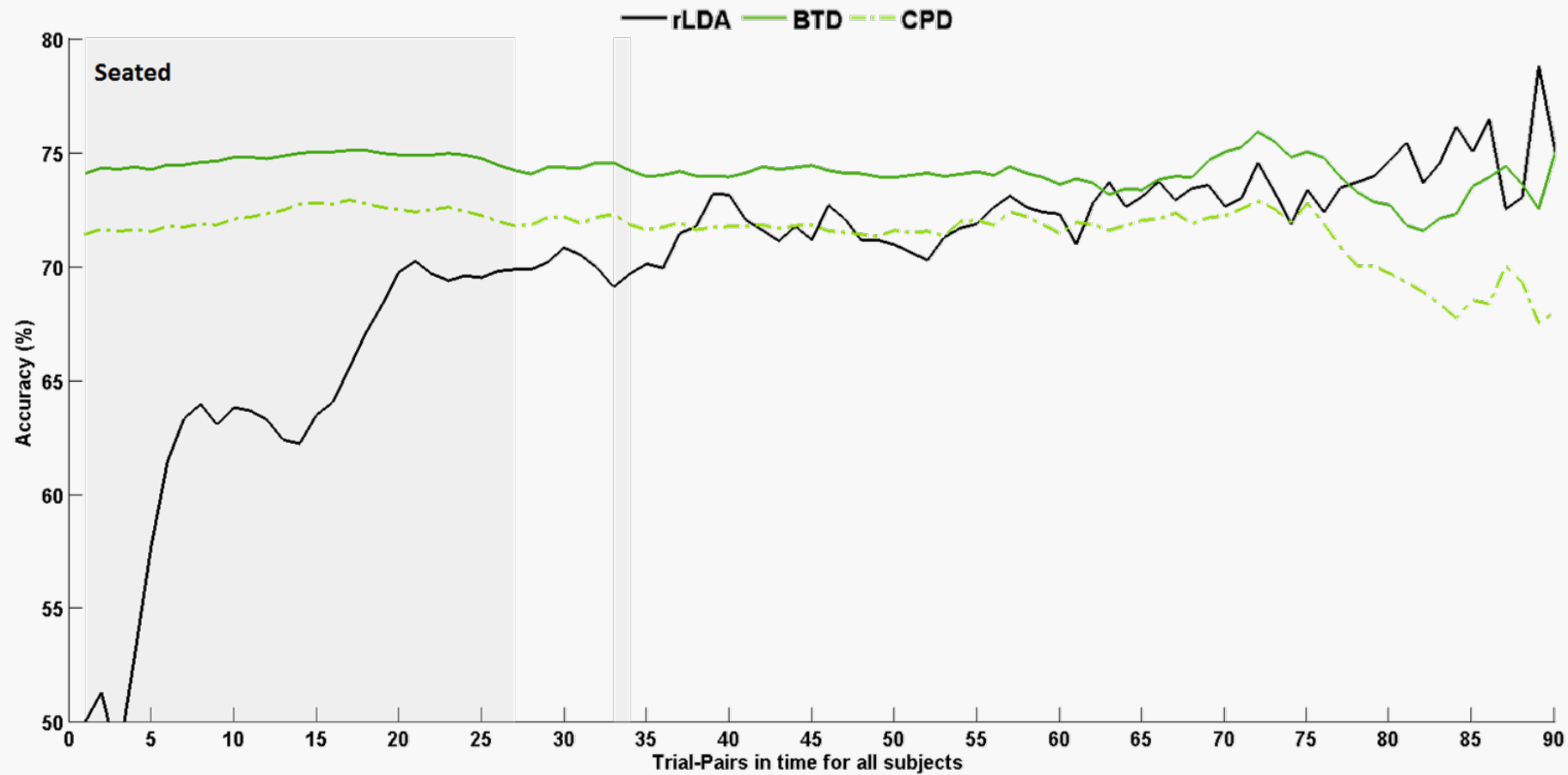


## Published in:

Zink, R., Hunyadi, B., Van Huffel, S., & De Vos, M. (2016). **Tensor-based classification of an auditory mobile BCI without a subject-specific calibration phase.** Journal of Neural Engineering, 13(2), 026005.



## Time-wise online analysis scenario Sit condition



Shaded area: significant differences to rLDA ( $p < 0.05$ )



## Summary

- ✓ The **BTD** method presented here has a **forthright link to the original P300 ERP** signal and achieves good classification results.
- ✓ Important step towards constructing intuitive classification methods by **exploiting data signatures from other subjects and structural paradigm information.**
- × **Dependence on the response patterns of the subject.** Between-subject differences remain. Walking condition lower accuracies.

## Future

- ◆ Understanding of the fluctuations in brain responses (and noise levels) on single trials.
- ◆ Assess the ‘maximum achievable’ classifications (Human Performance) – **100% accuracy is not possible on this type of data.**

### Published in:

Zink, R., Hunyadi, B., Van Huffel, S., & De Vos, M. (2016). **Tensor-based classification of an auditory mobile BCI without a subject-specific calibration phase.** Journal of Neural Engineering, 13(2), 026005.



# Contents Overview



1. Introduction



2. BIOTENSORS Project



3. Examples



**4. Conclusions and Future Directions**

# Conclusions

## MANY BLIND SOURCE SEPARATION PROBLEMS IN SMART PATIENT MONITORING OF LOW RANK

- *Solve via (constrained) matrix or tensor factorizations*
- *BIOTENSORS Project with TENSORlab: huge step forward*

### BIOTENSORS Achievement 1:

- *Development of **advanced algorithms for computation of tensor decompositions**, firmly based on numerical mathematics with proven uniqueness properties. This also includes more general studies, involving block terms, coupled data sets and various types of constraints, relevant for biomedical applications.*

### BIOTENSORS Achievement 2:

- *Introduction of **advanced approaches to BSS**, based on BTM instead of CPD, involving multimodal data, exploiting source structure as an alternative to statistical independence, and allowing exploitation of broad set of constraints.*

### TENSORlab Achievement:

- *Above algorithms efficiently implemented in Tensorlab 3.0 and 4.0. User friendliness improved significantly by simplifying model construction and adding visualization routines, documentation and demos.*

## ABOVE IMPROVEMENTS ALLOW TO FACE CHALLENGES IN BIOMEDICAL DATA FUSION

- *Successful examples: MS diagnostics, Neonatal Sleep Staging, Wearable Health monitoring, Mobile EEG*
- *Other examples: **See talks/posters at EURASIP Summer school Tensor-based Signal Processing***

# Future directions

- *Adaptive tensor decompositions, rank & structure estimation*
- *Multiscale multimodal approaches*
- *New emerging applications, e.g. in C(hr)onnectome analysis  
modelling dynamic brain connectivity networks*

→ *exploit full potential of existing Tensor(lab) **toolboxes***



# Thank you!

## Acknowledgments

- | University Hospitals Leuven Gasthuisberg
- | ZNA Middelheim, Queen Paola Children's hospital
- | EMC Rotterdam
- | KU Leuven, Dept. Electrical Engineering-ESAT, division STADIUS & MICAS
- | Ghent University, Dept. Telecommunication and Information Processing, TELIN-IPI
- | Eindhoven University of Technology



**ERC advanced grant 339804 BIOTENSORS** in collaboration with L. De Lathauwer and group

