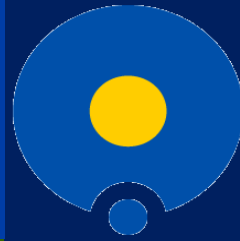




Young Universities
for the Future of Europe



Asymptotic spatiotemporal averaging of EEG power for schizophrenia diagnostics.



Włodzisław Duch

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Dept. of Informatics, Faculty of Physics, Astronomy & Informatics,
Nicolaus Copernicus University, Toruń, Poland

Search: Wlodzislaw Duch

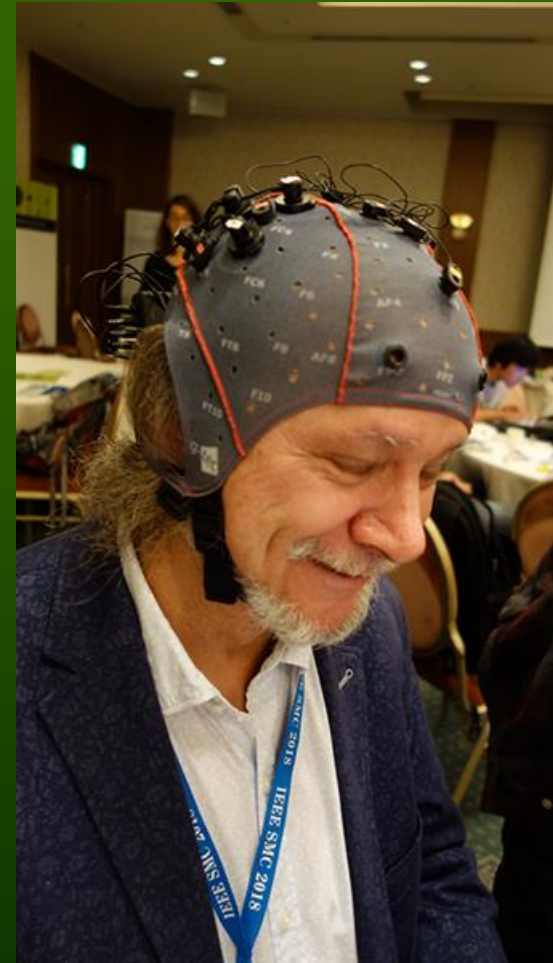
2023 Int. Conf. on Neural Information Processing, Changsha, China, 19-25.11.2023

On the threshold of a dream ...

Motivation: optimize brain processes!

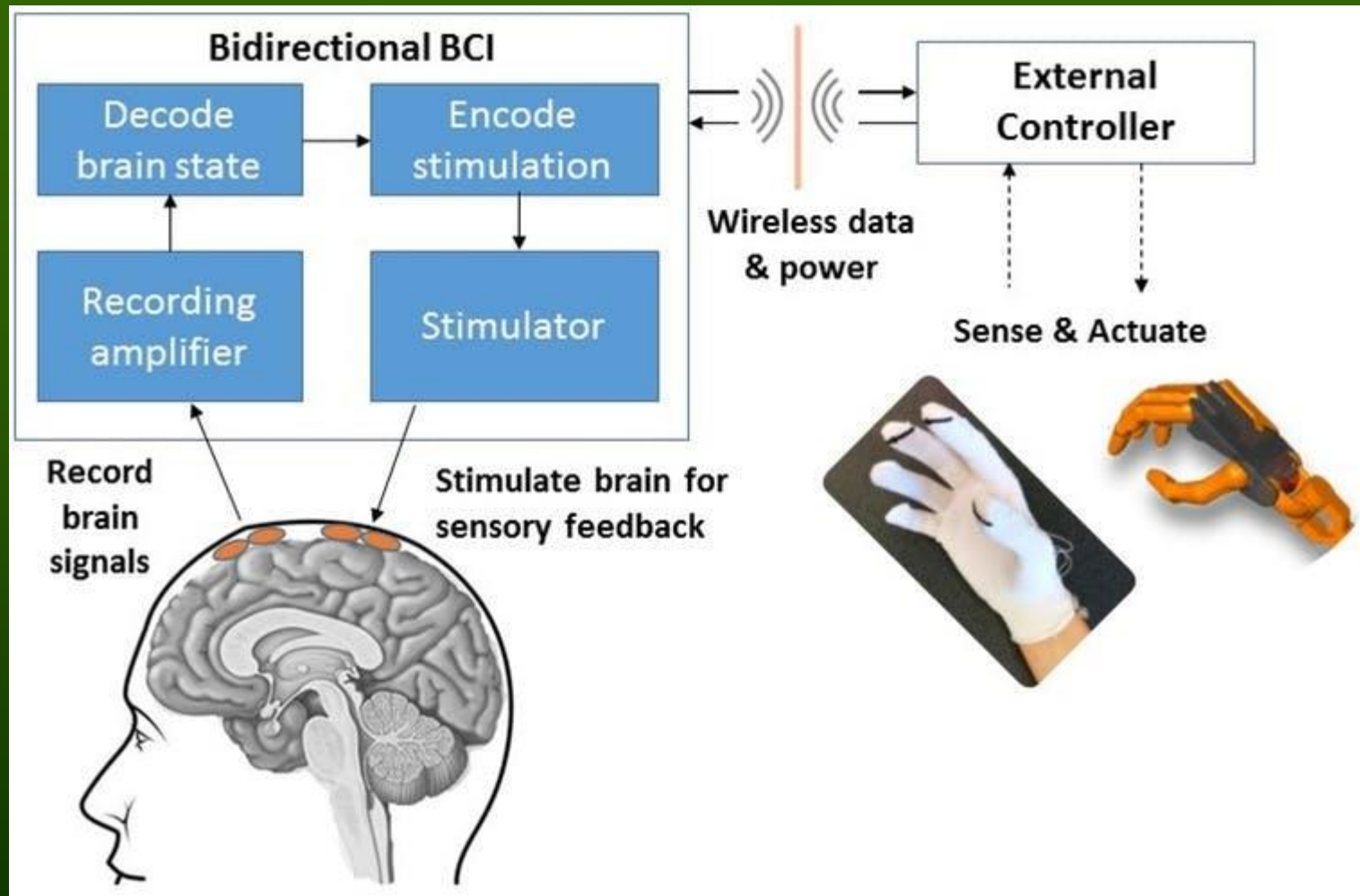
Repair damaged brains, increase efficiency of healthy brains! First we need to understand brain processes:

1. Find **fingerprints of specific activity** of brain structures using neurotechnologies.
2. Create **models of cognitive architectures** that help to understand information processing in the brain.
3. Create **new diagnostic and therapeutic procedures**.
4. Use **neurofeedback based on decoding and changes in connectivity** to stimulate the brain.
5. **Stimulate neuroplasticity** by monitoring brain activity and directly stimulating it (TMS, DCS, EM).



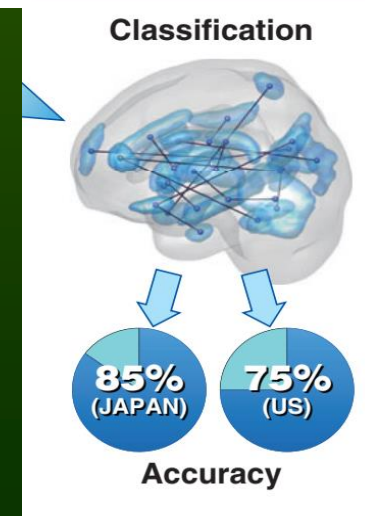
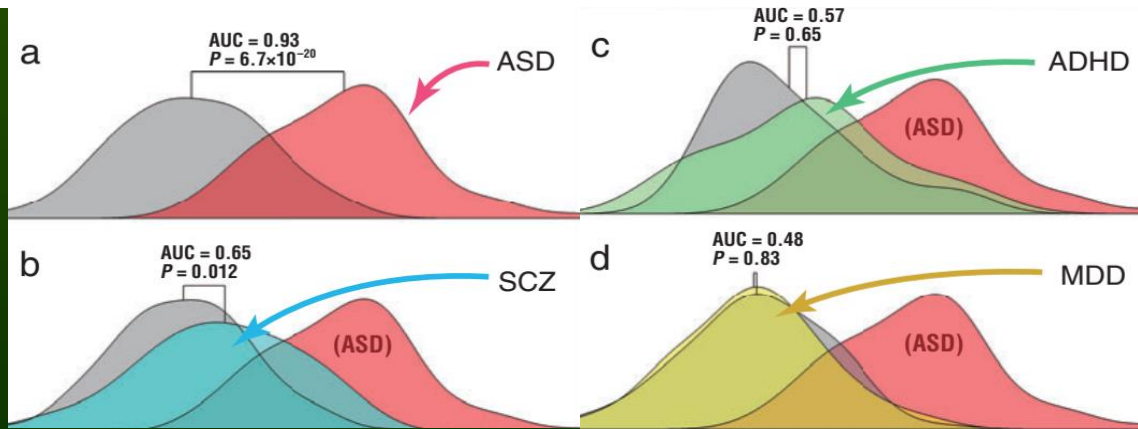
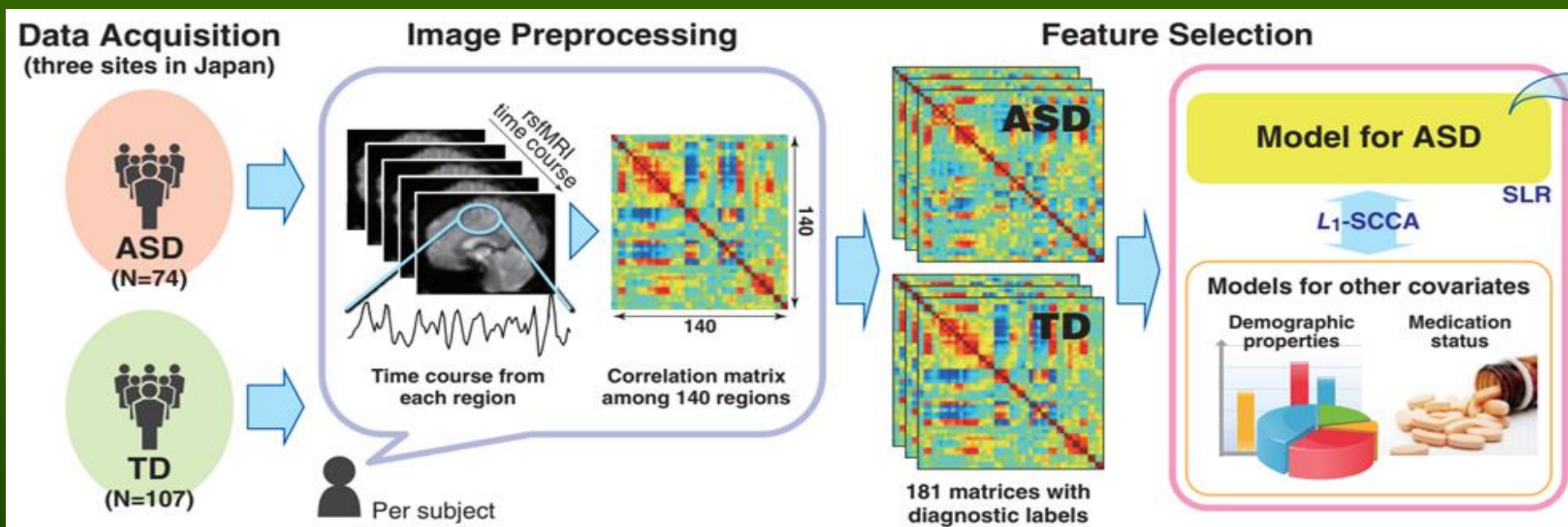
G-tec wireless NIRS/EEG on my head.

BCBI: Brain-Computer-Brain

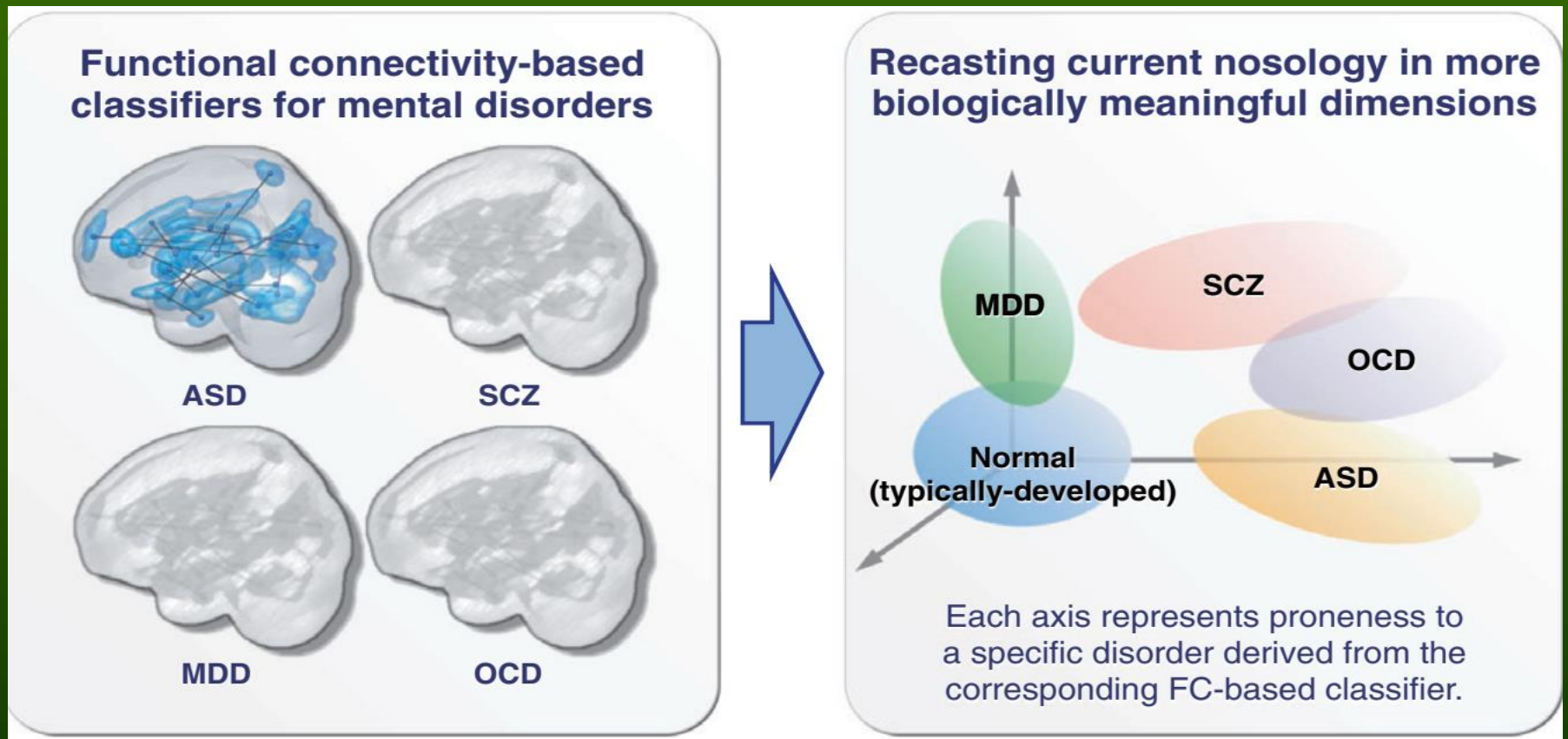


BCI + brain stimulation = BCBI – a closed loop through which the brain begins to restructure itself. The body can be replaced by signals in Virtual Reality.

Biomarkers from fMRI FCs



Biomarkers of mental disorders



fMRI biomarkers allow for objective diagnosis. MDD, deep depression, SCZ, schizophrenia, OCD, obsessive-compulsive disorder, ASD autism spectrum disorder. This should be most effective neurofeedback approach.

N. Yahata et al, *Psychiatry & Clinical Neurosciences* 2017; **71**: 215–237

Problems with EEG



- Sophisticated EEG analysis is rarely used in clinical practice.
- Reliable biomarkers for diagnosis of brain disorders are still unknown.

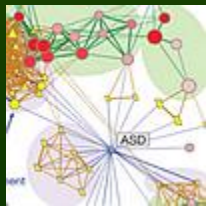
Why?

- Signals are non-stationary, even during short periods.
- Individual differences are quite large.
- Neurodynamics, especially in the resting state, depends on hundreds of confounds.
- Methods tested on a small datasets in real life do not generalize well.
100% accuracy on small datasets ~ little chance of being useful.

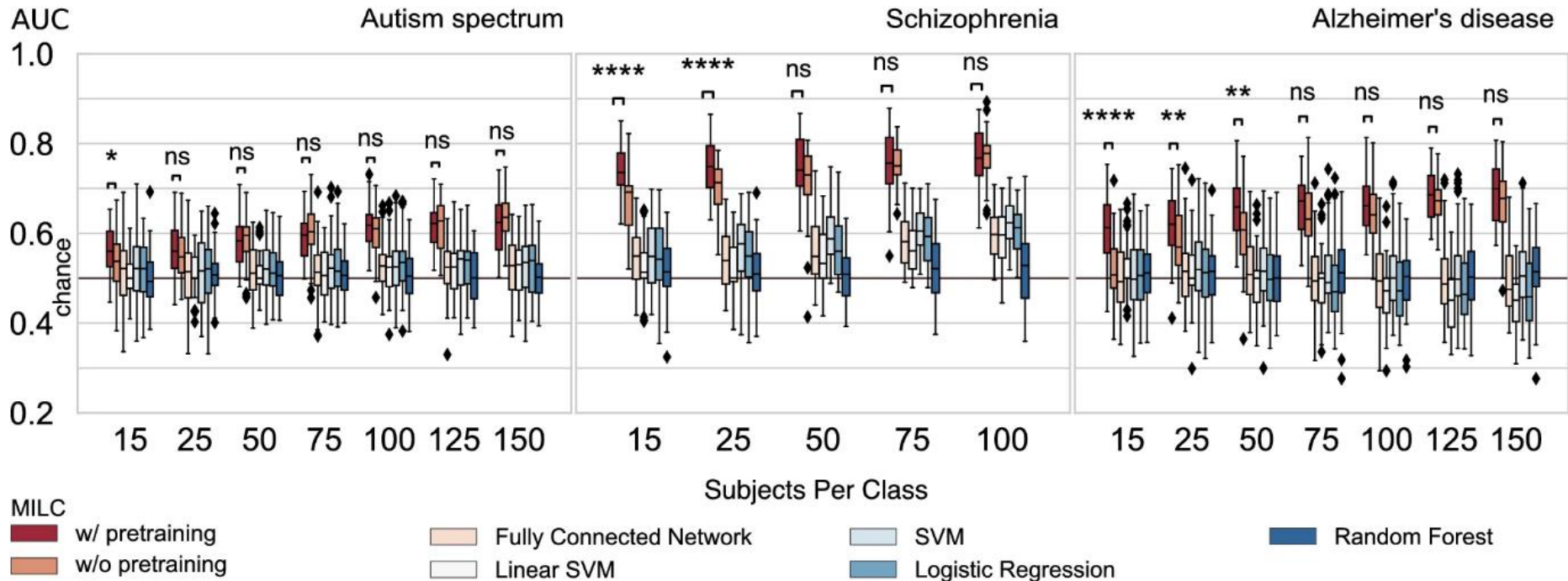
Why?

SFARI Human Gene Module contains 1140 genes related to ASD, and still new genes are added. Very large number of people had to be tested.

EEG diagnosis: without very large EEG databases of such heterogenic disorders as schizophrenia or MCI we shall not create good biomarkers.



MILC model



Rahman, ... & Plis, S. M. (2022). Interpreting models interpreting brain dynamics. *Scientific Reports*, 12(1), 12023.

Mutual information maximization between the whole sequence (~300 context embedding) and local windows (local embedding) from the same sequence.

Supervised pretraining scheme, which maximizes “Mutual Information Local to (whole) Context” (MILC). After pre-training on large fMRI data deep learning + MILC can learn directly from high-dim signal dynamics, even in small datasets.

Our goals



Understand:

- Limits of spatial averaging of power in the narrow frequency bands.
- Limits of temporal characterization of brain processes using a large number of microstates.
- Combination of spatial, temporal and frequency data in recurrence analysis.
- Show big influence of rare cases feature selection done on the whole data and just strictly on training partitions in CV on classification results.

Tests were made on a typical, small EEG dataset of 45 schizophrenic adolescents + 39 controls. Several papers can get 100% accuracy on such datasets. Perfect accuracy is reached when features are selected on the whole data, before training a classifier (which is a common practice).

Feature selection on the cross-validation training partition does not allow to identify rare cases that should be inspected separately.

Recent work

Where to look for information in EEG?

- Spatial distribution of power.
- Temporal dynamics of regions/subnetworks.
- Spectral (Frequency) Fingerprints.

Recent PhDs:

- Karolina Finc, Dynamics and plasticity of human brain functional network during working memory task performance.
- Kamil Bonna, Neural correlates of prediction errors during reward and punishment learning.
- Michał Komorowski, Locally specific human brain dynamics automatically modeled using spectral features of MEG/EEG signals.
- Ewa Ratajczak, Microstate neurodynamics in HRV biofeedback.

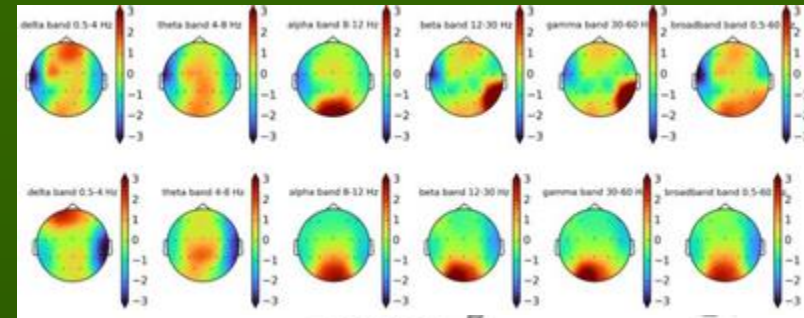
Furman Ł, Tołpa K, Minati L, Duch W. (2022) Short-Time Fourier Transform and Embedding Method for Recurrence Quantification Analysis of EEG Time Series. [European Physical Journal Special Topics](#), 1-15, (2022)

Brain fingerprinting

Find unique patterns of brain activity:

- brain regions of interest (ROI),
- active neural networks,
- mental states, tasks, processes.

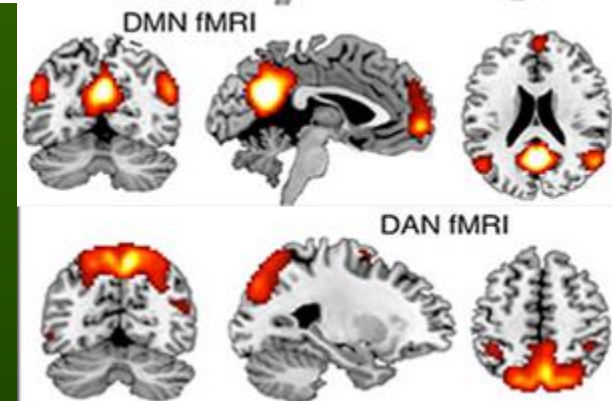
1



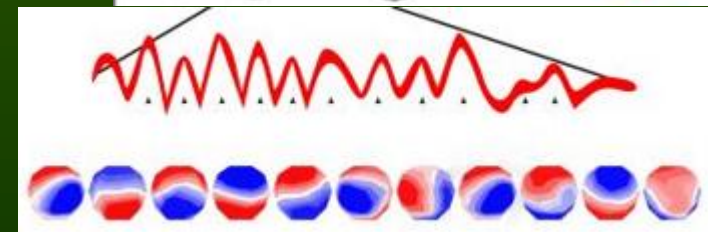
Several approaches:

1. Spatial distributions, power maps and spectral fingerprints (Keitel & Gross 2016)
2. Large scale networks seen in fMRI can be recreated from EEG (Yuan et al, 2015).
3. Temporal information, microstates and their transitions (Michel & Koenig 2018)
4. Recurrence plots and RQA, recurrence quantification analysis.
5. Connectivity, functional correlations and many more approaches...

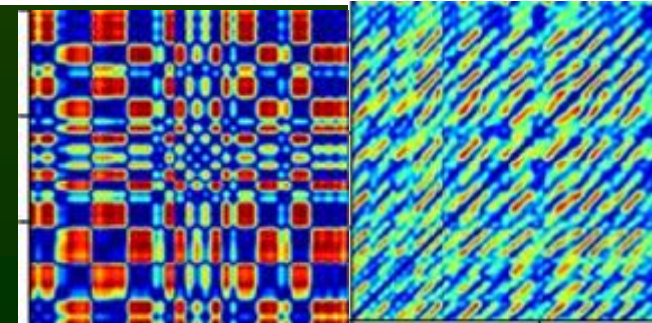
2

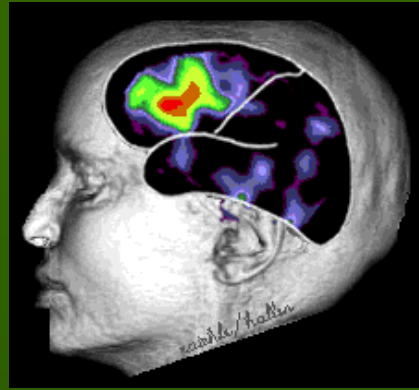


3

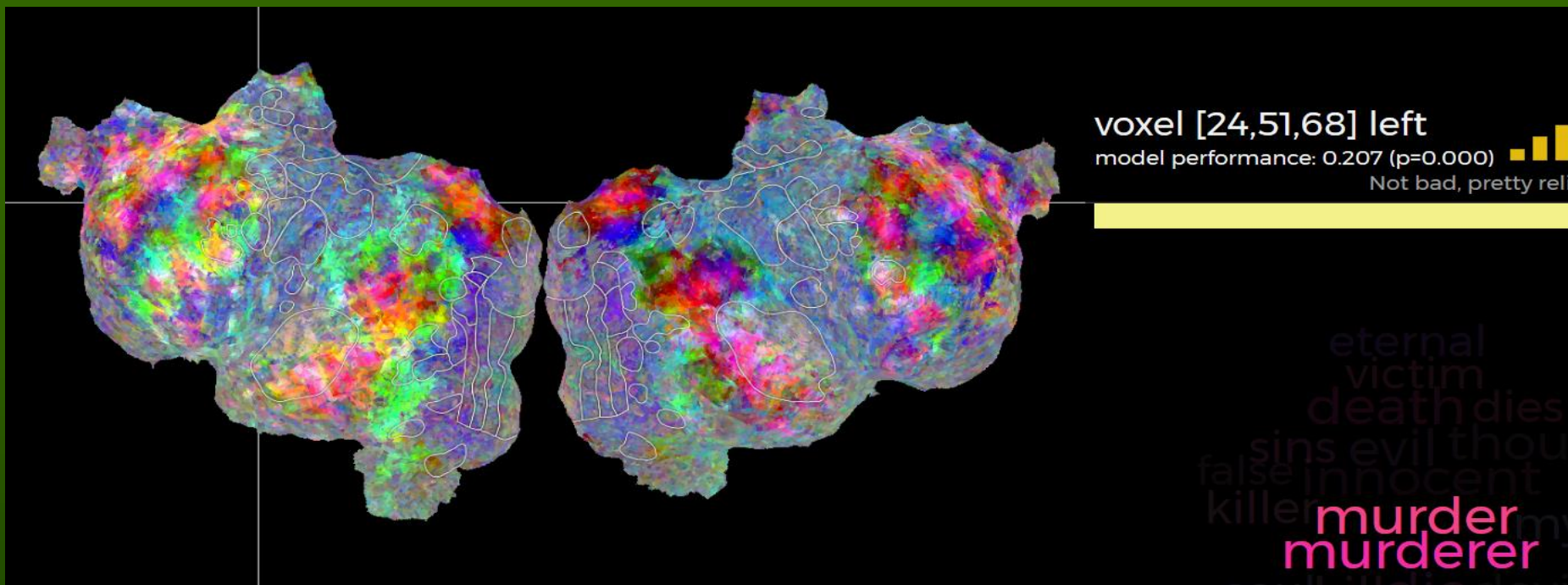


4





Brains: spatial aspects



Whole fMRI activity map for the word “murder” shown on the flattened cortex.

Each word activates a whole map of activity in the brain, depending on sensory features, motor actions and affective components associated with this word.

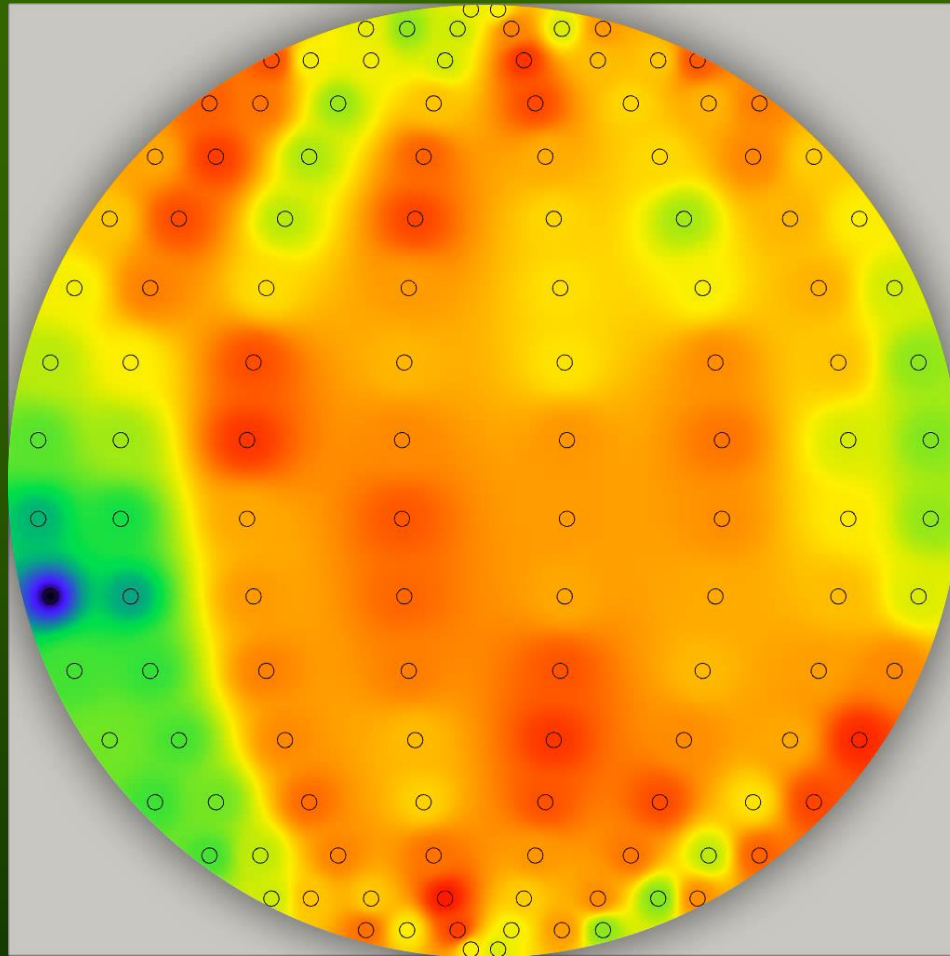
Why such activity patterns arise? Brain subnetworks connect active areas.

<http://gallantlab.org/huth2016/> and [short movie intro](#).

Can one do something like that with EEG or MEG?

Prompts invoke specific activation in LLMs.

Dynamics



Asymptotic EEG power in the resting state may show hypo/hyperactive brain regions, summarize overall brain activity. How well can it distinguish between brain disorders? Dynamics should be taken into account averaging activity of subnetworks.

Schizo reference results

Schizophrenia patients: 45 boys (10-14 y) diagnosed with schizophrenia and 39 healthy controls (Borisov et al. Human Physiology, 2005).

EEG: 16 electrodes, 125 Hz sampling rate, 60 sec. sequences.

[Phang et al.](#) A Multi-Domain Connectome Convolutional Neural Network for Identifying Schizophrenia from EEG Connectivity Patterns, IEEE JBHI 2020.

3 types of features:

Time-domain (VAR) autoregressive model coefficients,

Frequency-domain (PDC) partial directed coherence,

Network topology-based complex network (CN) measures.

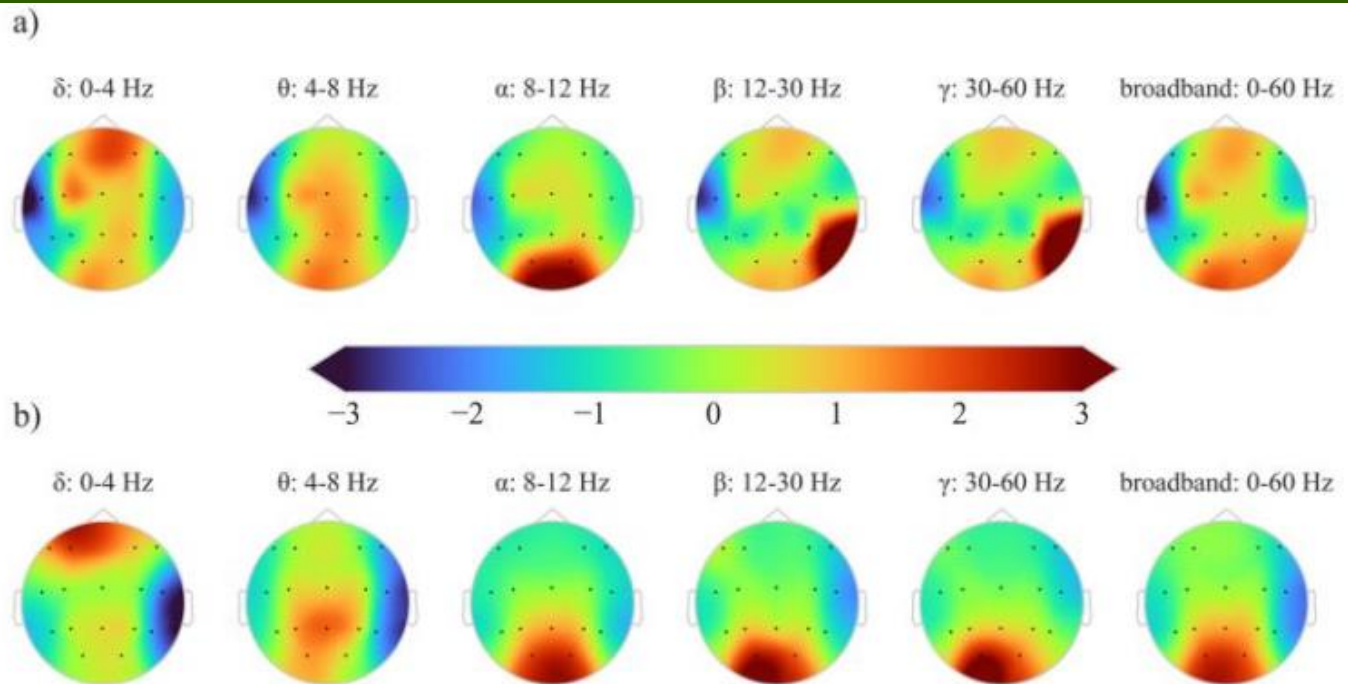
Total dim= $((2 \times 16 \times 16) + 34) \times 5 = 2730$

| | | | |
|------------|------|-----------------|-------------------------------|
| CNN CN | 170 | 81.0±4.4 | network topology graphs (TDA) |
| CNN VAR | 1280 | 81.0 | time-domain VAR |
| CNN PDC | 1280 | 89.2 | freq. domain PDC |
| SVM PDC | 1280 | 88.0 | freq. domain PDC |
| VAR+PDC+CN | 2730 | 91.7±4.7 | best CNN |

Average Power Plots (avPP)

Long-term temporal averaging of signals in each channel. Asymptotic values of average power distributions (avPP) shows activations of different brain regions for each frequency bands. We also have such maps in 1 Hz bands.

Schizo
S27w1



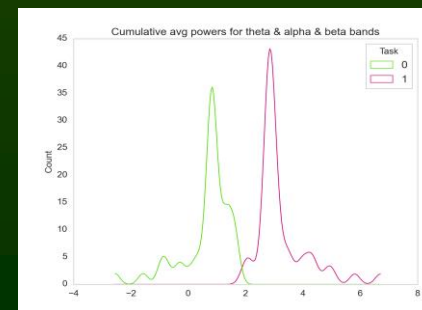
Control
719w1

Example of average resting state power in 60 sec segment for two subjects. Cumulative power is relatively stable. Individual differences are large.

L-SVM 5xCV avPP classification

| EEG bands | Selection on all data | | Selection on training | |
|-------------------------------------|-----------------------|-----------------|-----------------------|------------------|
| | N dim | Acc±Var % | N dim | Acc±Var % |
| Broadband | 10 | 72.5±20.6 | 10 | 68.0±22.8 |
| $\beta+\gamma$ | 3 | 73.8±19.6 | 3 | 72.7±20.5 |
| $\theta+\beta$ | 23 | 74.9±18.4 | 23 | 65.4±23.0 |
| $\delta+\theta$ | 19 | 68.9±21.0 | 19 | 65.6±23.6 |
| $\delta+\theta+\alpha$ | 19 | 90.5±8.6 | 19 | 76.2±19.2 |
| $\delta+\theta+\alpha+\beta$ | 21 | 95.2±4.6 | 21 | 78.5±17.8 |
| $\delta+\theta+\alpha+\beta+\gamma$ | 71 | 79.5±15.5 | 71 | 79.8±17.1 |

Number of features has been fixed for all folds, but in selection on the training partition features were different in each CV fold.



Schizophrenia avPP

Simple method, may be useful in clinical settings.

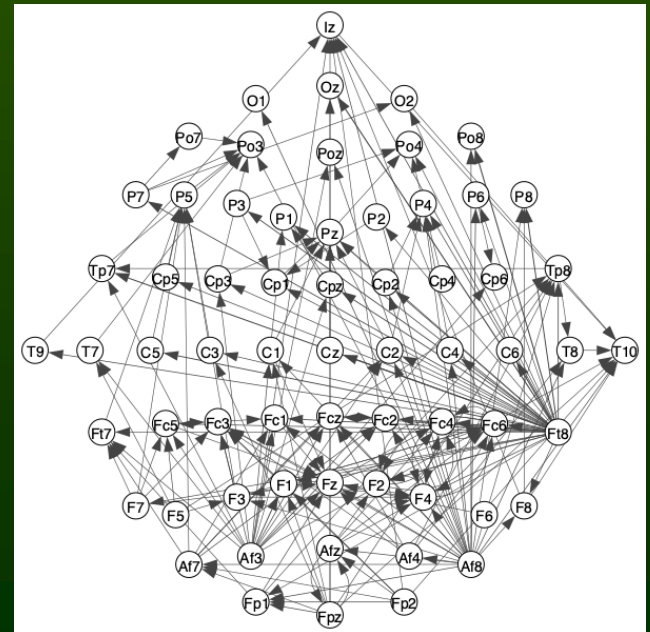
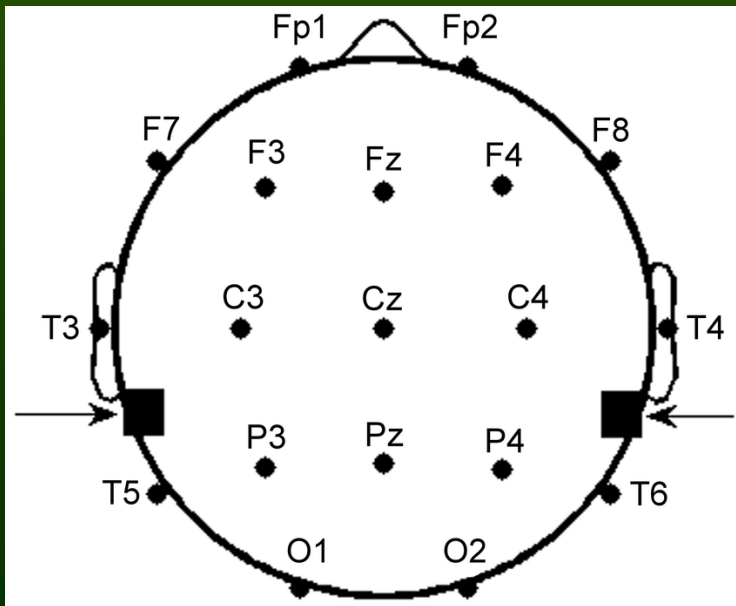
The power distribution maps are relatively stable for each individual, with clusters distinguishing schizophrenia patients vs. control subjects.

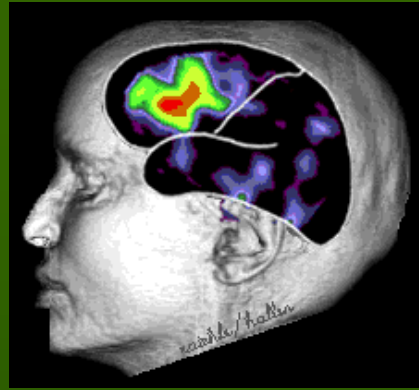
Electrode/frequency band selection is based on linear SVM weights.

The top 10 combinations :

T4 α , F8 α , P3 β , C3 β , O2 α , Pz α , F8 θ , P4 θ , T4 θ , T3 θ

Improve: add dynamics + coherence + topological data analysis.





Brains: temporal aspects

Understanding brains: microstates

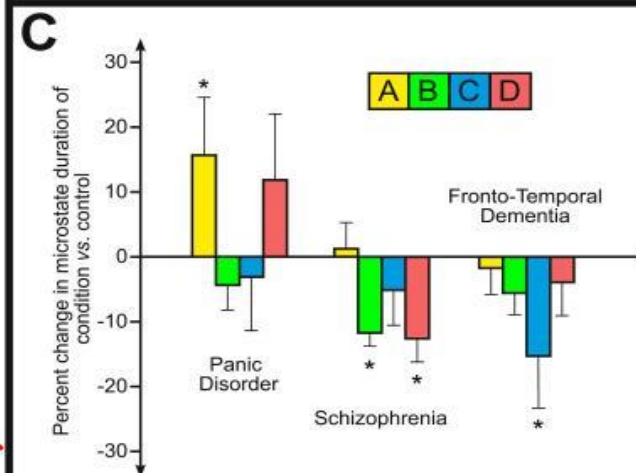
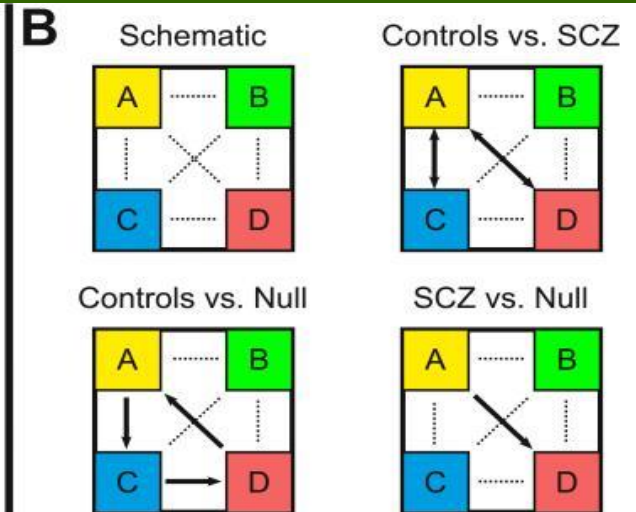
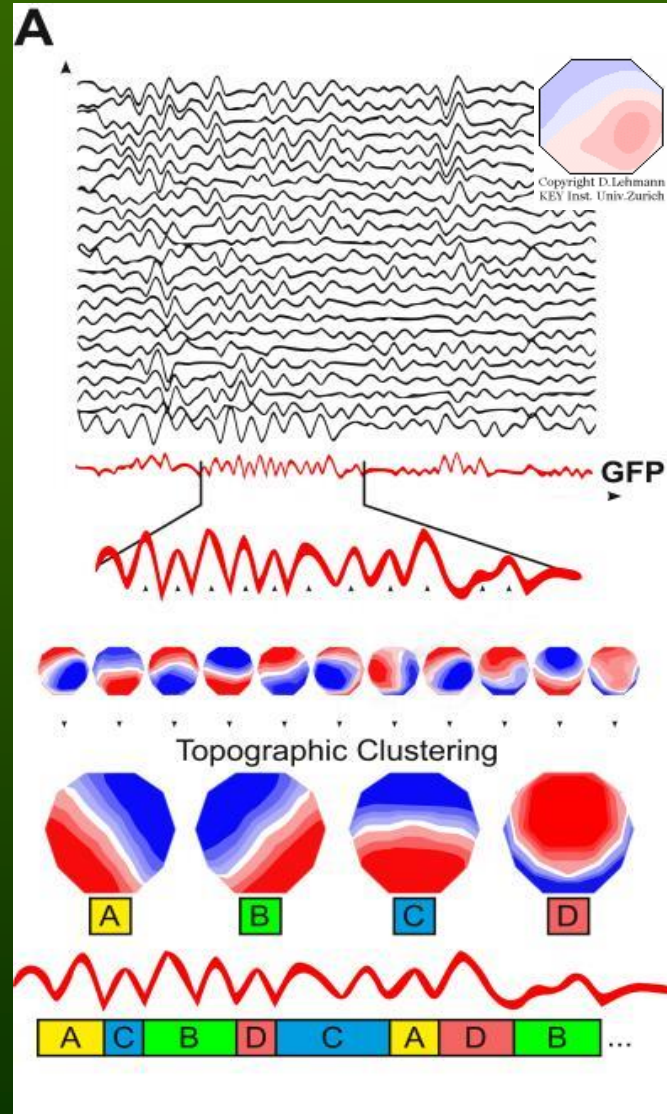
Global Field Power.
4-7 states, 60-150 ms.

Khanna et al. (2015)
Microstates in
Resting-State EEG.
*Neuroscience and
Biobehavioral Reviews.*

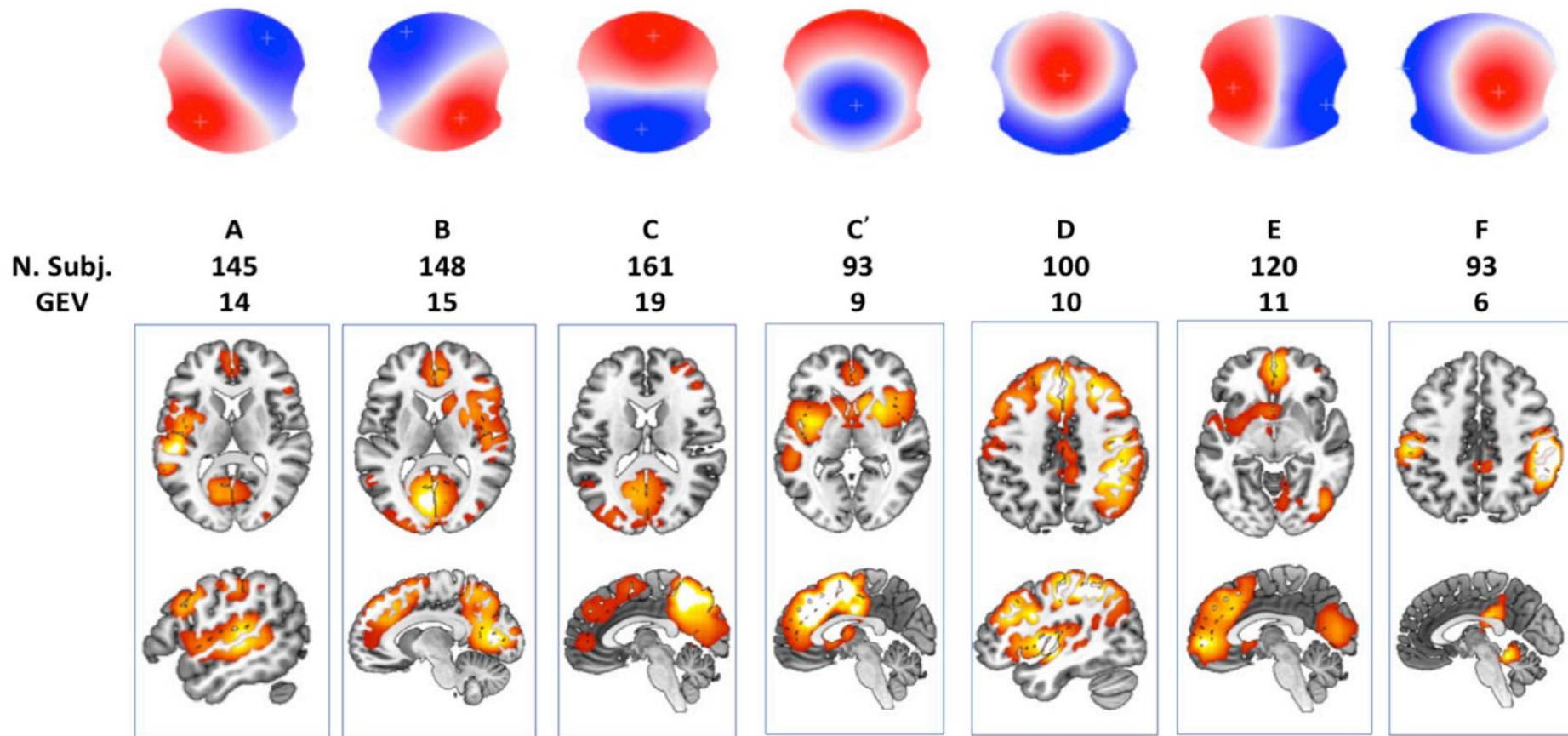
Symbolic dynamics:
statistics of A-D
symbol strings. Fuzzy
Symbolic Dynamics
(FSD) + visualizations.

Duch W, Doboşz K.
(2011). *Cognitive
Neurodynamics* 5, 145

Doboşz K, Duch W.
(2010). *Neural Networks,*
23(4), 487–496.



Microstates and their sources



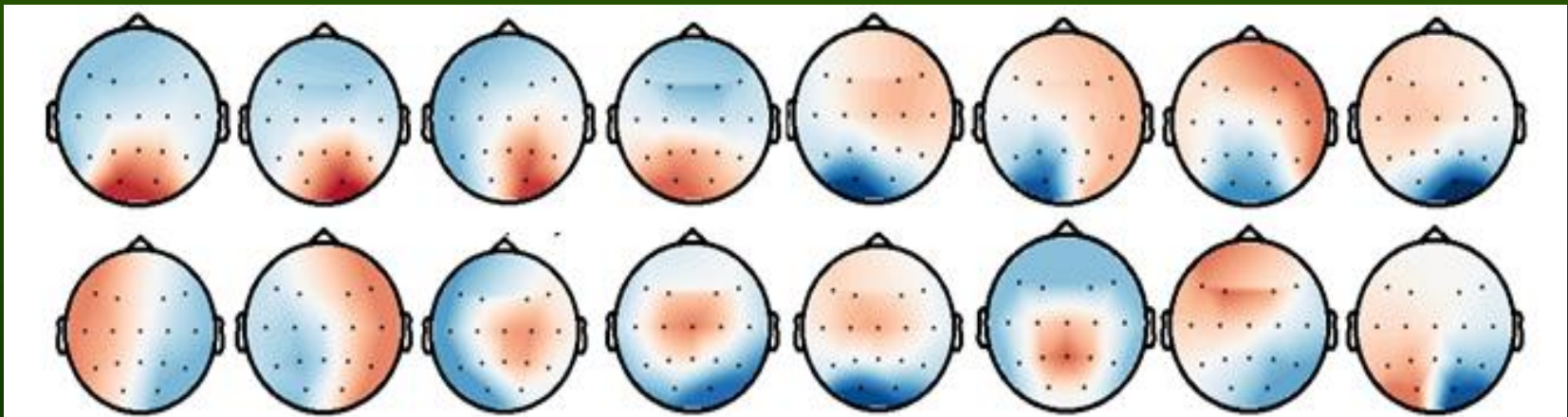
Michel, C. M., & Koenig, T. (2018). EEG microstates as a tool for studying the temporal dynamics of whole-brain neuronal networks: A review. *NeuroImage*, 180, 577–593.

Ewa Ratajczak, PhD thesis "Microstate neurodynamics in HRV biofeedback" (2022)

Microstates limits

T\We have created 4-20 microstates using two clusterization procedures. Features include occurrence (OCC), duration (DUR), coverage (Cov), global field potential (GFP), global explained variation (GEV), mean spatial correlation (MSC), transition probabilities between microstate classes (TP, 16x16).

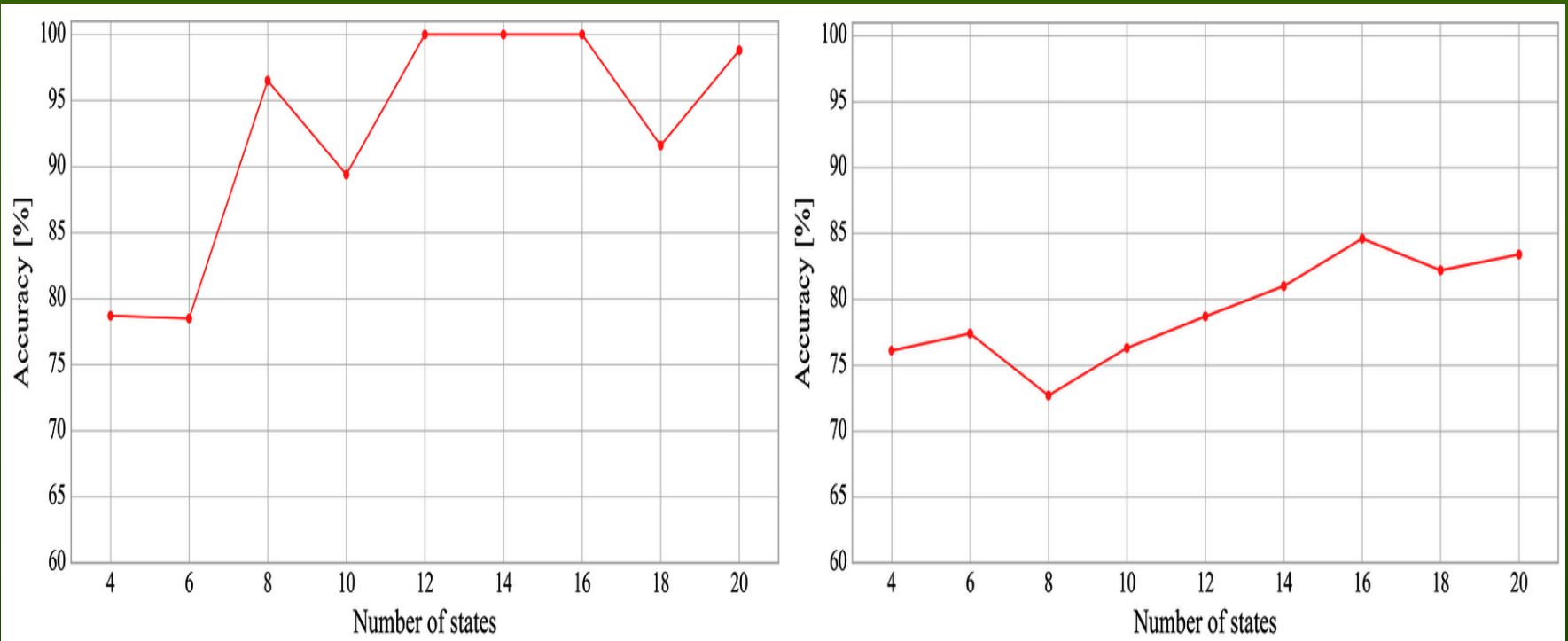
Below – 16 microstate example for schizophrenia. They hardly differ. Another choice: use avPP states as prototypes.



Microstates for classification

| Microstates | | Selection on all data | | Selection on training only | | |
|-------------|---------|-----------------------|-----------|----------------------------|------|-----------|
| N states | Type | N dim | Acc±Var % | Type | Ndim | Acc±Var % |
| 4 | TAAHC | 4 | 78.7±17.2 | TAAHC | 4 | 76.1±18.6 |
| 6 | TAAHC | 52 | 78.5±17.7 | TAAHC | 52 | 77.4±17.7 |
| 8 | TAAHC | 17 | 96.5±3.4 | TAAHC | 17 | 72.7±20.2 |
| 10 | TAAHC | 93 | 89.4±9.4 | TAAHC | 93 | 76.3±18.5 |
| 12 | K-means | 55 | 100 | K-means | 55 | 78.7±17.5 |
| 14 | K-means | 90 | 100 | K-means | 90 | 81.0±15.9 |
| 16 | TAAHC | 42 | 100 | K-means | 17 | 84.6±13.0 |
| 18 | TAAHC | 281 | 91.6±7.8 | TAAHC | 281 | 82.2±14.0 |
| 20 | TAAHC | 221 | 98.8±1.2 | TAAHC | 221 | 83.4±14.3 |

Microstates for classification

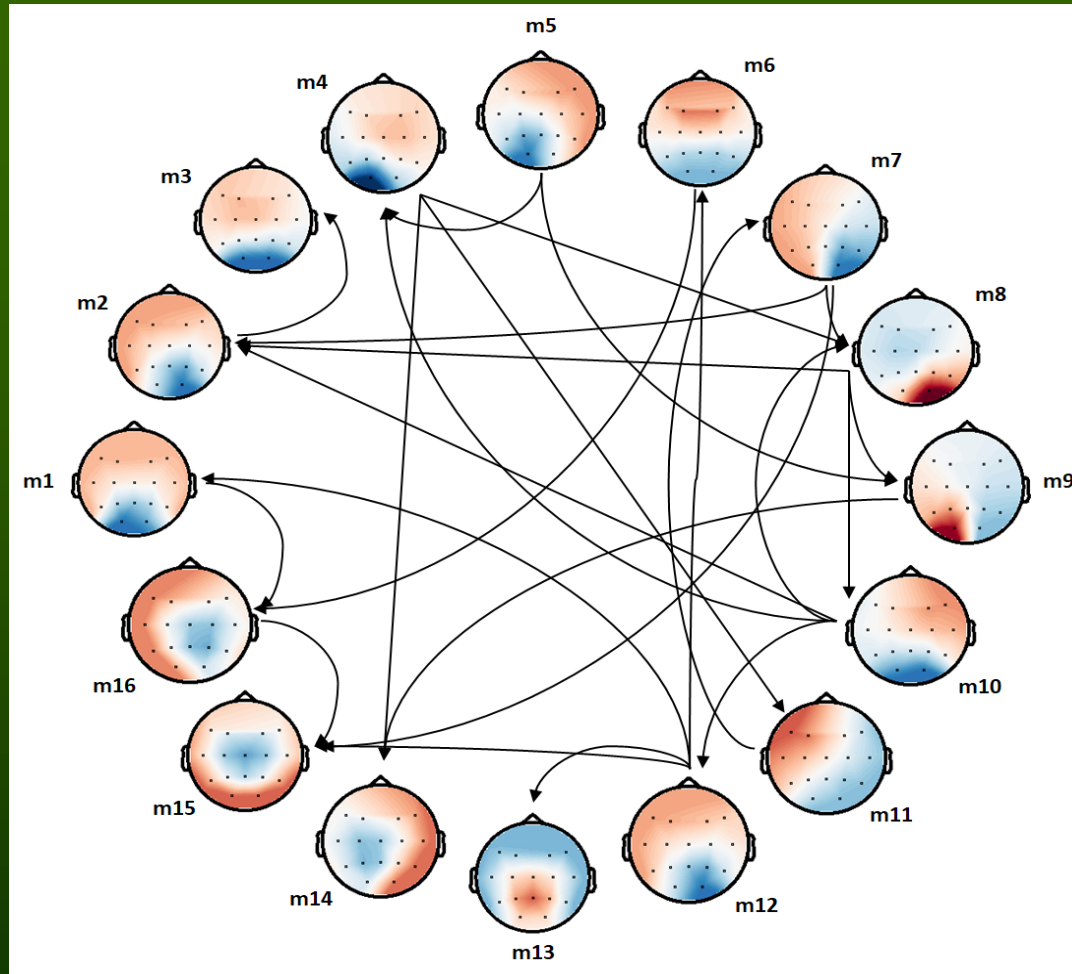


Left: features selected on the whole data, before training of LSVM.

Right: features selected on the training partition.

In both cases training of LSVM is in stratified crossvalidation.

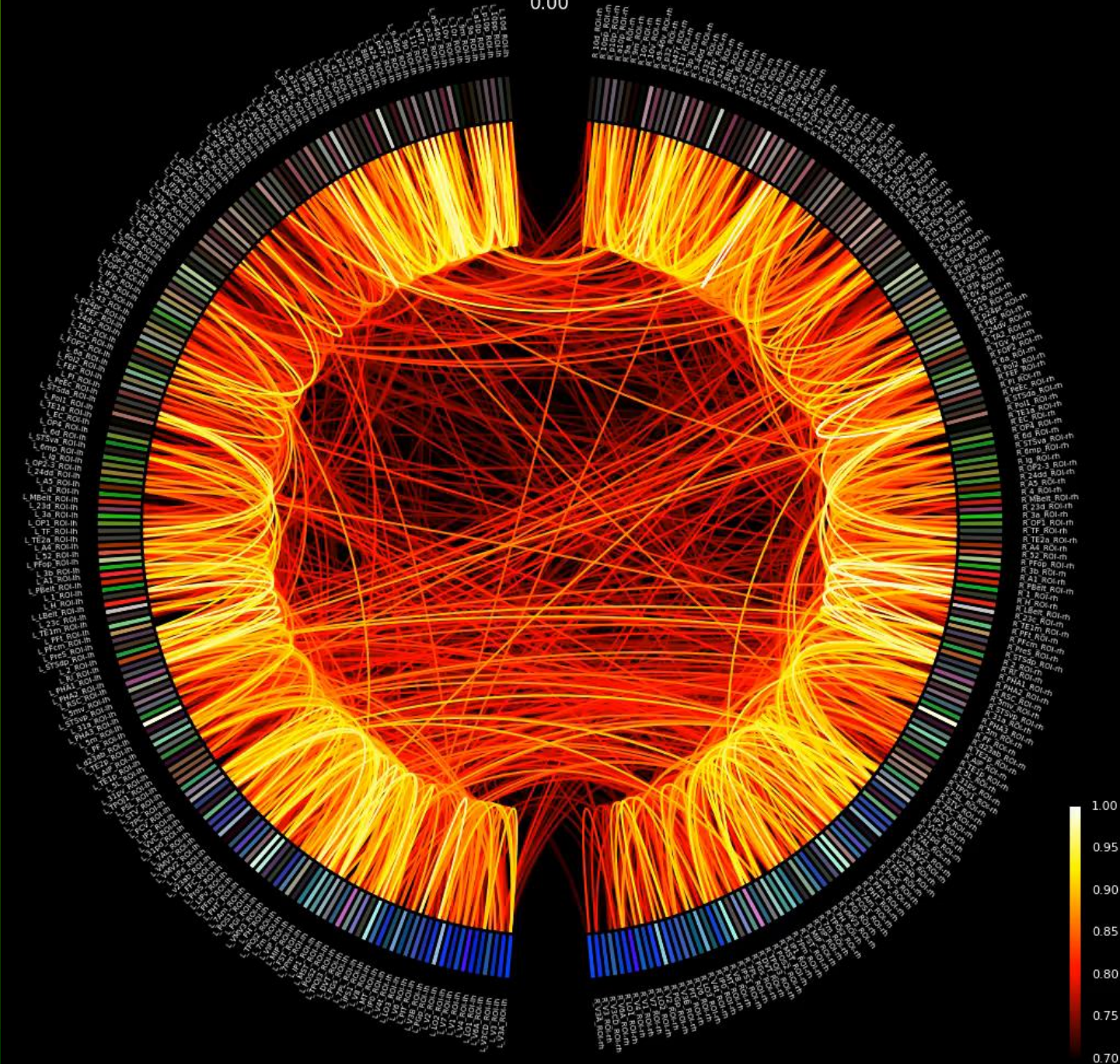
Microstate transitions

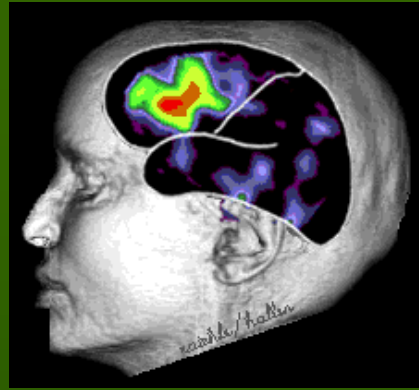


These transitions were selected most often.

PAC_Itest_inverse_circle_coh_8.0-12.0h_z_vmin0.7

0.00

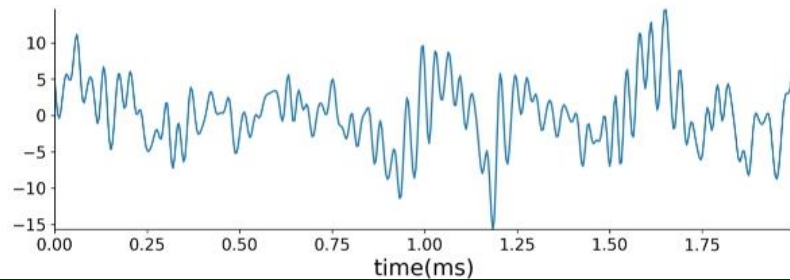
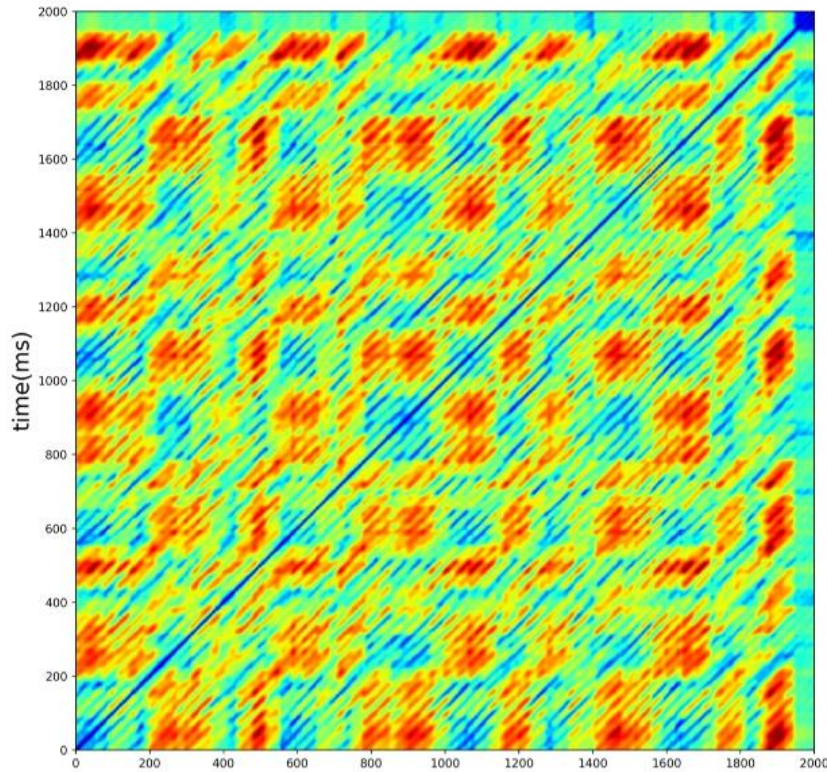
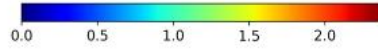




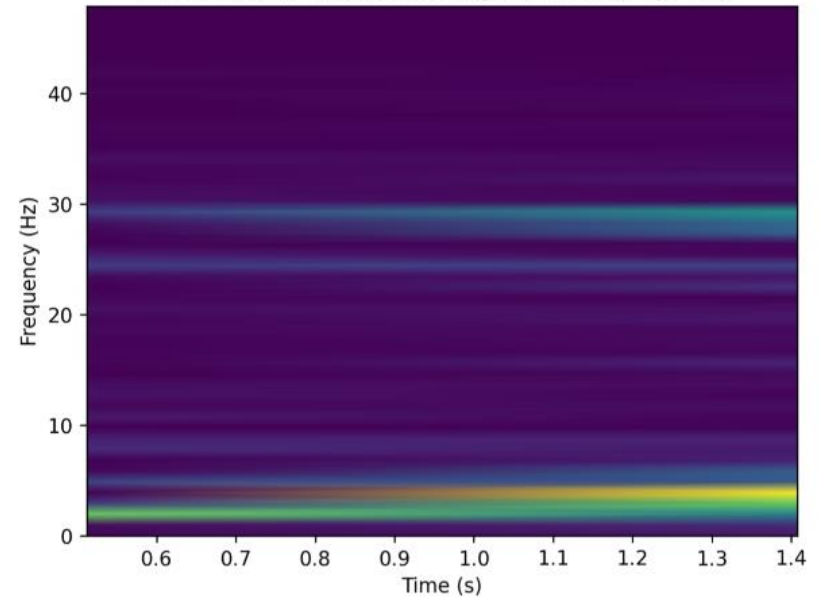
Brains: spatio-temporal aspects

Representations

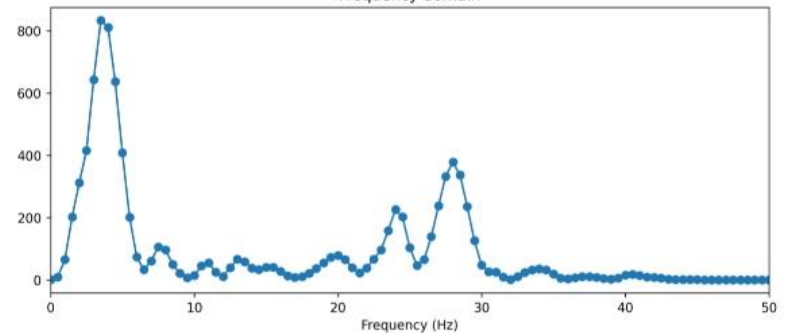
CP4, emb = 9 td = 2
sub: testSubject timestamp 26.0



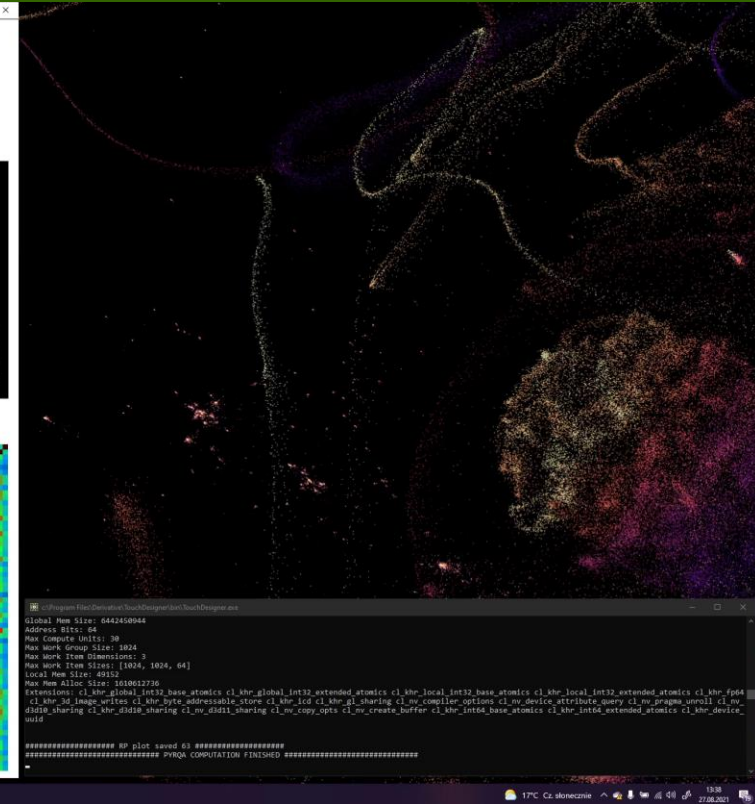
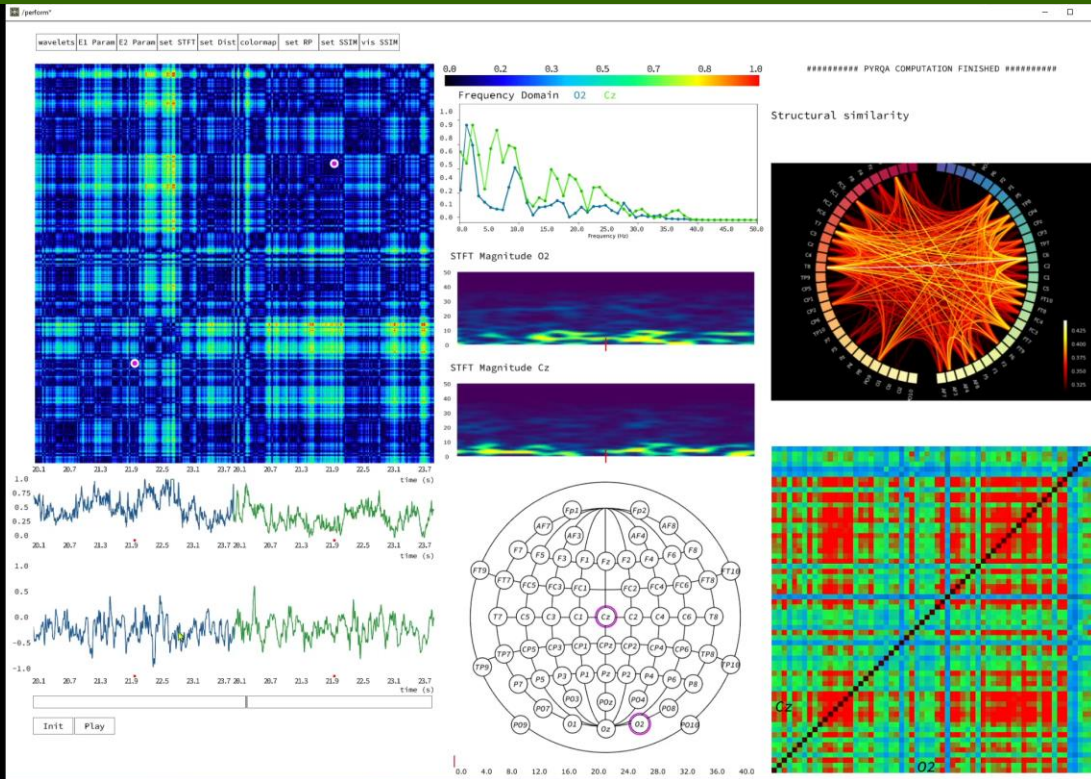
Electrode: CP4, sub: testSubject, timestamp: 26.0



Frequency domain

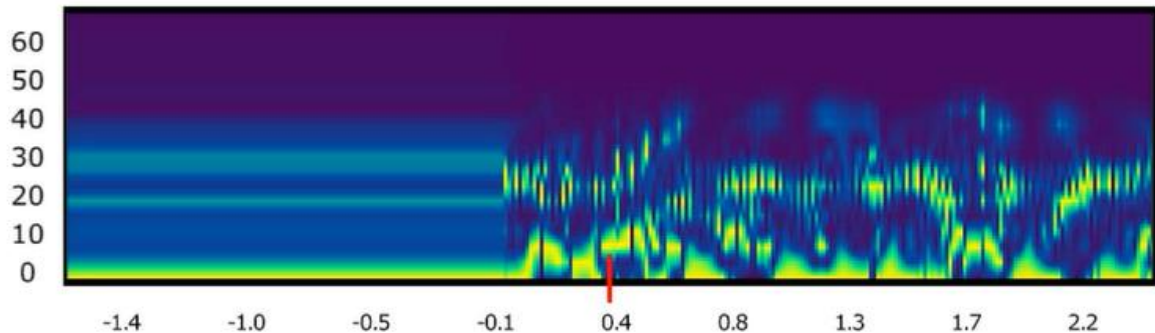


STFT EEG in real time

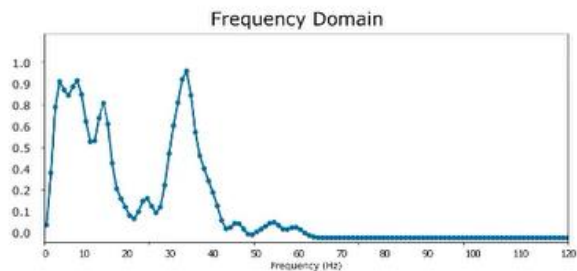
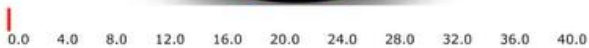
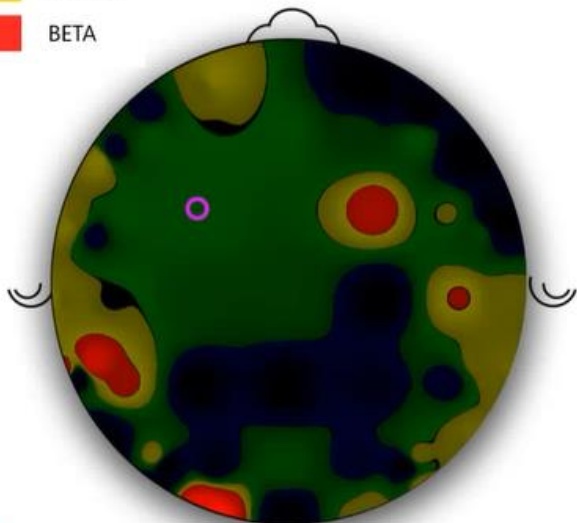


EEG data, 128 channels, recursion graphs, power spectrum for two electrodes, information flow and correlations between brain regions. BrainPulse (in development, Łukasz Furman).

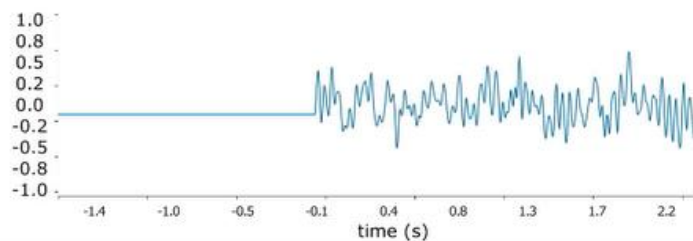
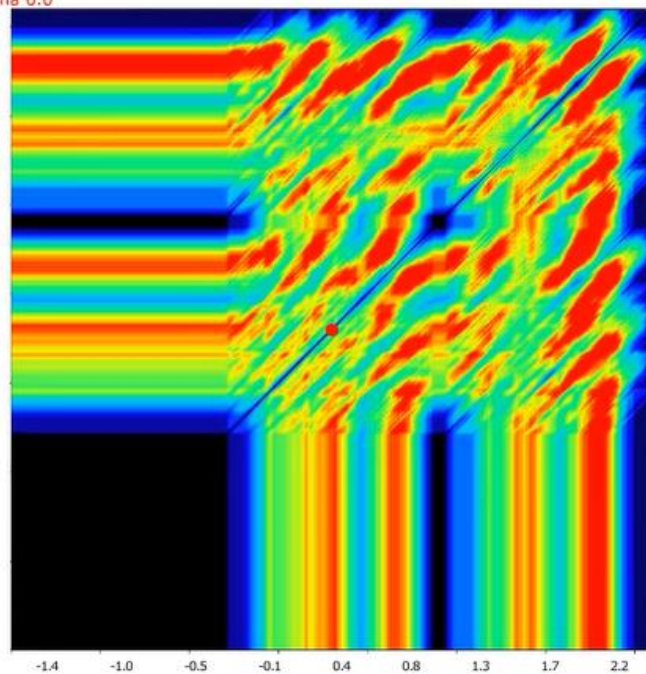
STFT Magnitude



- DELTA
- THETA
- ALPHA
- BETA



Electrode FC1 td=4 emb=28
norm=0.0

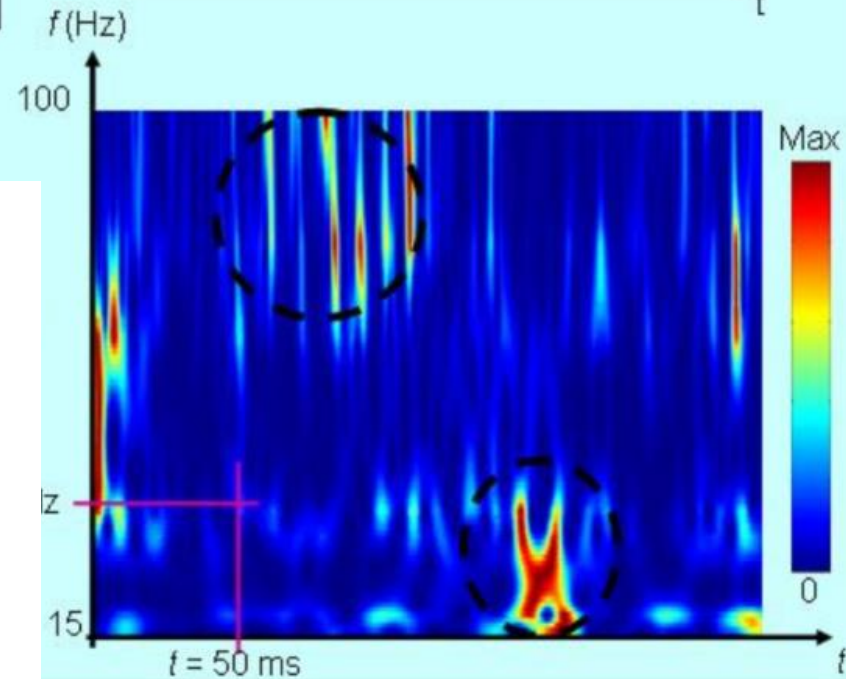
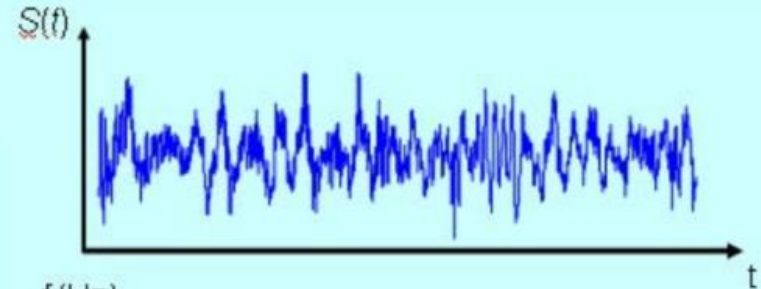


t/f rep and bumps

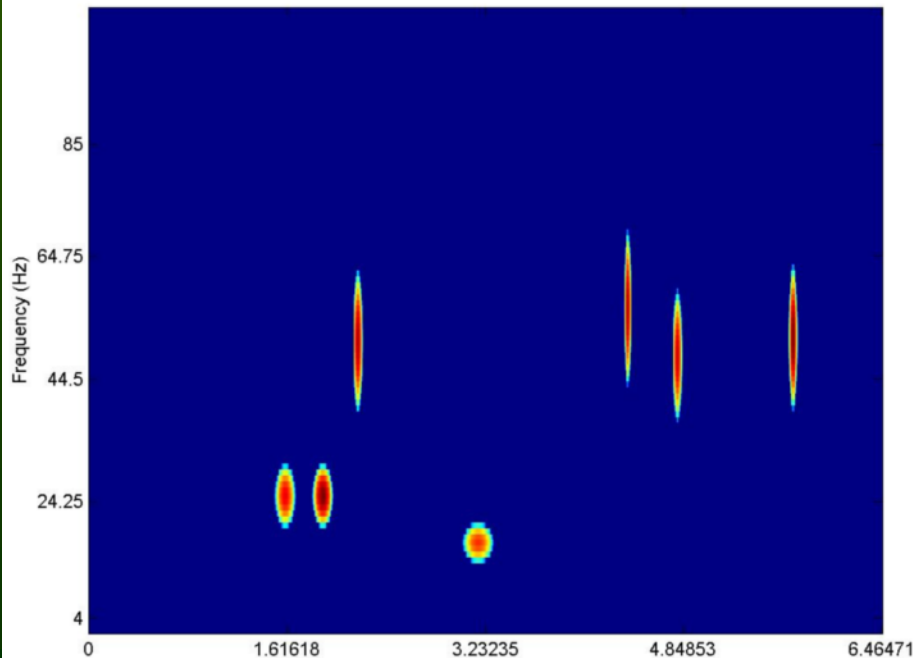
High frequency intermittent signals, and low beta strong activation, ECoG data, BCI Competition III . Msc thesis of M Szupke (2011), using EEGLab.

Wavelet transform

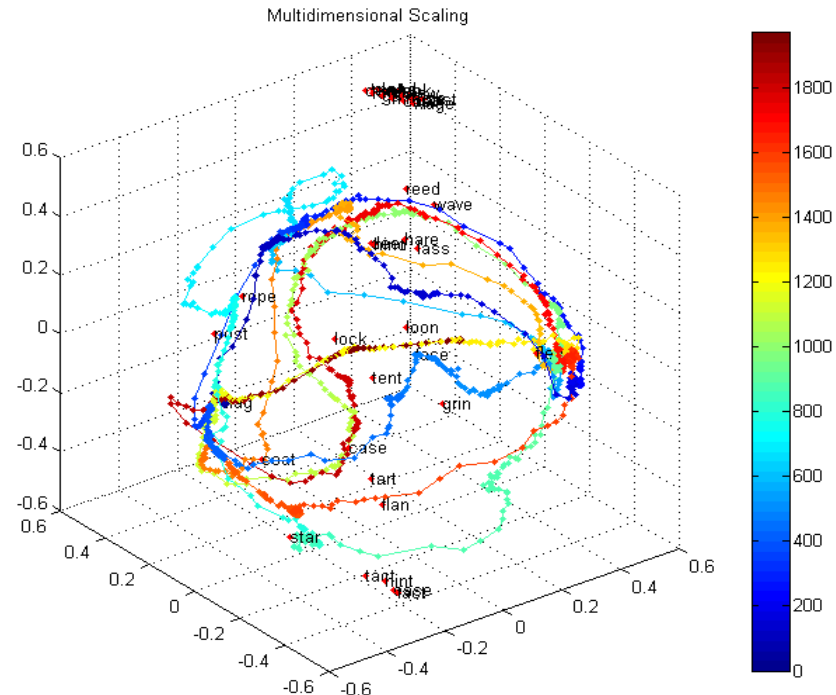
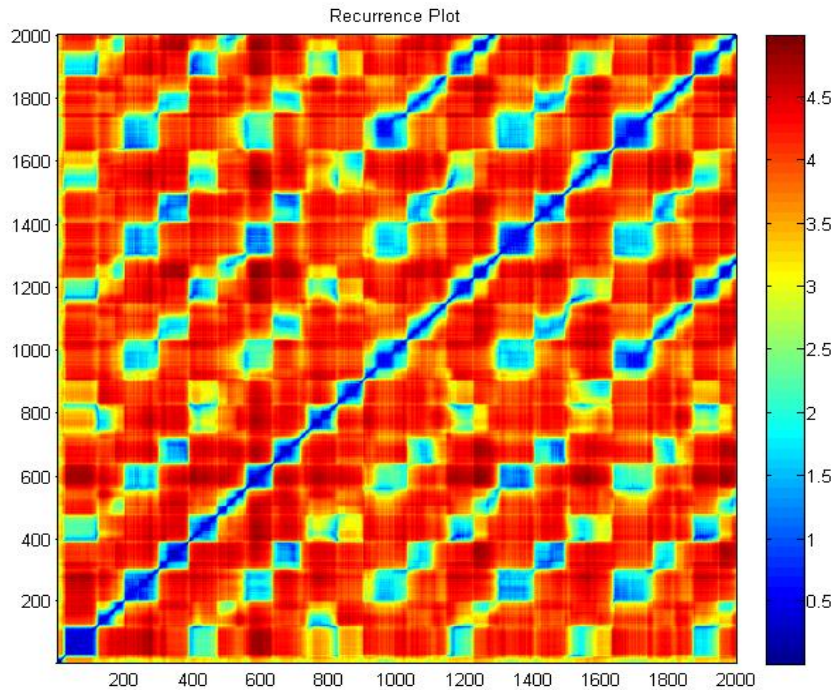
S signal, h wavelet
 W_f time-frequency map
 $W_f(s, \tau) = \int_{-\infty}^{+\infty} f(t) h_{s, \tau}^* dt$



Bump decomposition 1 / 1



Recurrence and trajectory visualization



Recurrence plots and MDS visualization of trajectories of the brain activity. Here evolution of 140-dim semantic layer activity during spontaneous associations in the 40-words microdomain is presented, starting with the word “flag”. Trajectories may be displayed using tSNE, UMAP, MDS or our FSD visualization. Identify metastable states, calculate trapping times, recurrence rates, entropy ...

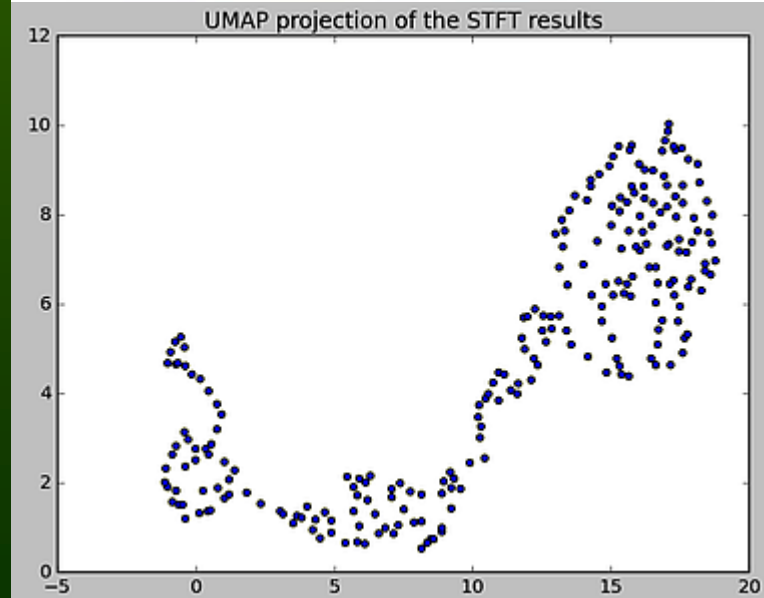
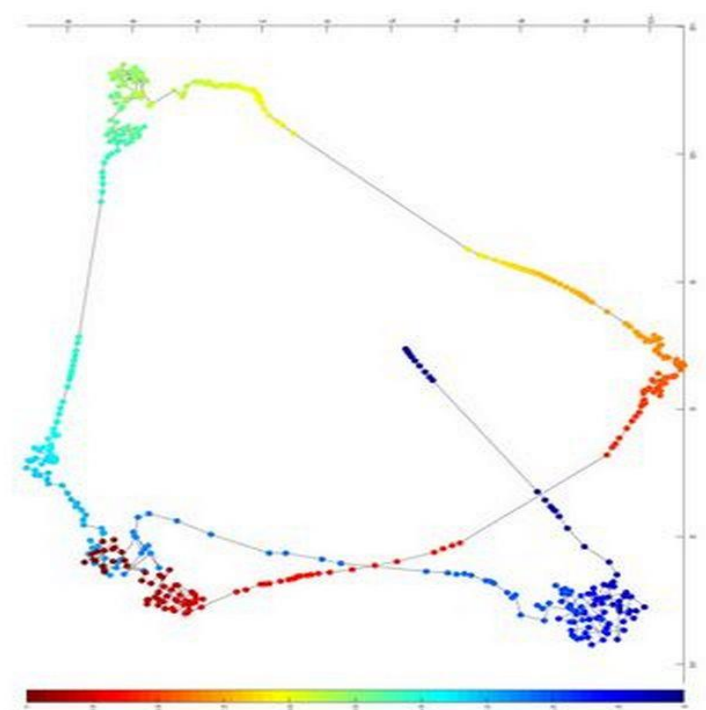
Trajectories

Can we characterize attractor states of the brain using EEG data?

tSNE for simulated attractor network,
color=time, each dot represents 140 ROIs.
Large and small attractor basins,
large clusters = long trapping time,
fast transitions between some states,
Recurrence near the end.

UMAP STFT visualization of real EEG data,
single channels/sources.

Some transitions and clusterization, but
several subnetworks with individual
trajectories (working memory patterns)
with separate trajectories (work in progress).

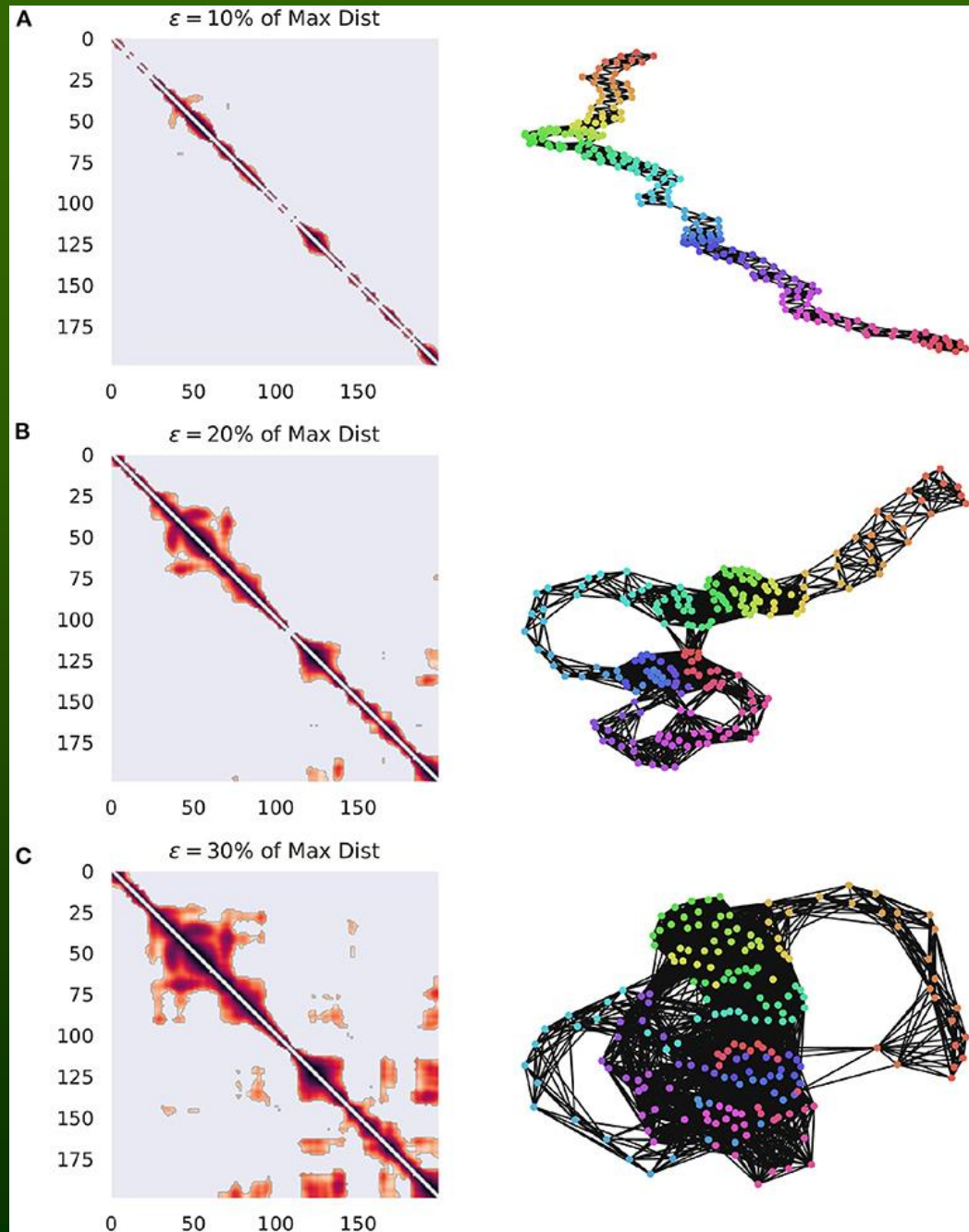


TDA

TDA quantifies complex network topology graphs.
Real brains, ECoG data: recurrence plots depend on the similarity threshold ε , cosine distance, Takens embedding of oscillatory data with dimension d and lag τ .

Caputi et al. (2021). Promises and pitfalls of **Topological Data Analysis** for brain connectivity analysis. *NeuroImage*, 238, 118245.

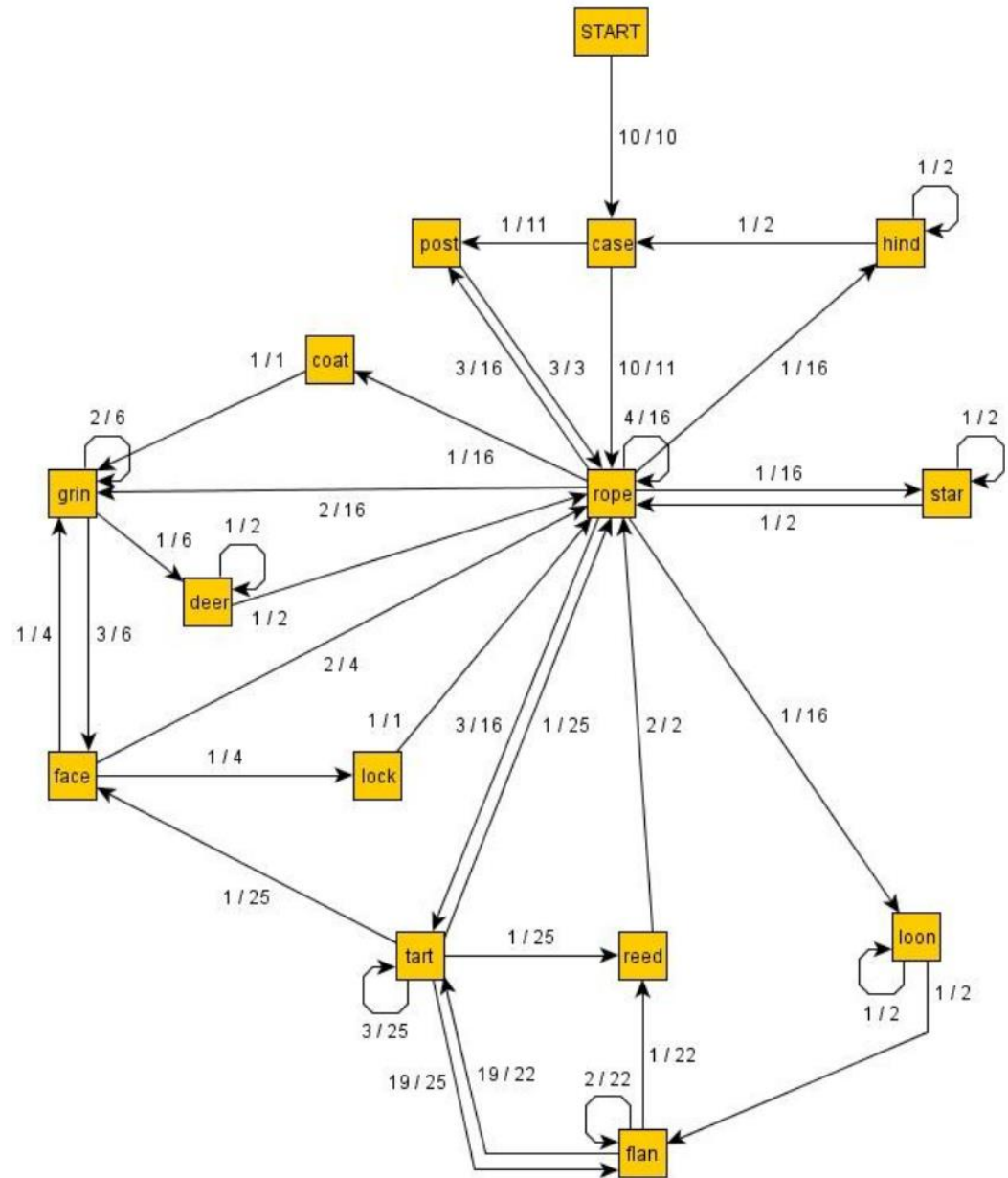
Varley, T. F., & Sporns, O. (2022). Network Analysis of Time Series: Novel Approaches to Network Neuroscience. [*Frontiers in Neuroscience*, 15.](#)



Multiple starts from the same word lead to different trajectories. Calculate transition probabilities between metastable states from frequency of transitions.

Why such transitions?

Linked state have patterns sharing few features, that recruit less active, but strongly connected neurons, and relax those currently active, making the previous state inaccessible for some time (refractory period).

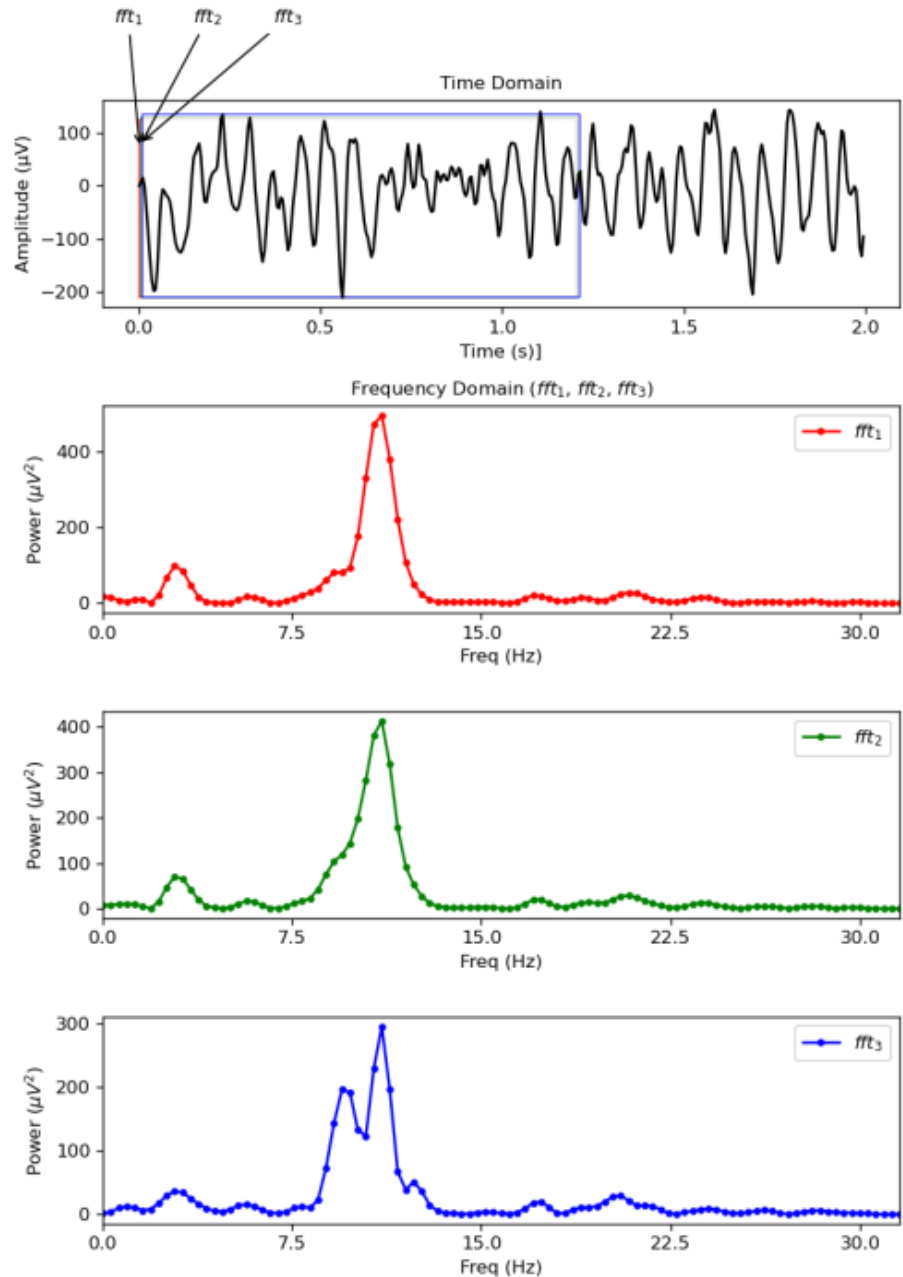


Labeling states

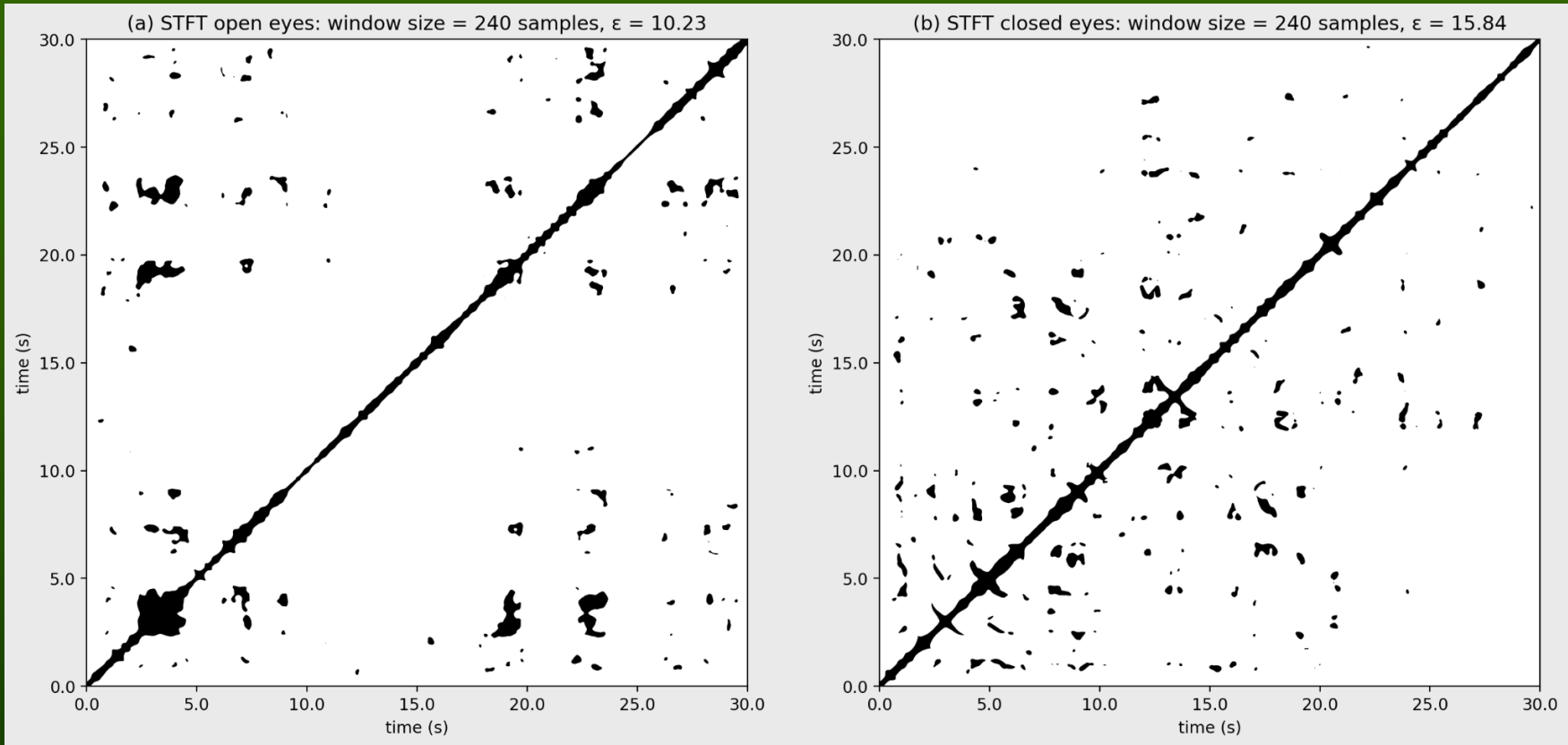
Example of STFT vectors, time windows ~ 1.5 -sec, showing a shift and split of the α peak frequency after 200 ms. O1 electrode (occipital area), eyes closed.

Compare spectra S_1, S_2 : many types of similarity measures may be used. Which channels have similar spectra? How long they are metastable (trapping time)? How frequent are transitions? Recurrence to the same state?

Ł. Furman, W. Duch, L. Minati, K. Tołpa, Short-Time Fourier Transform and Embedding Method for Recurrence Quantification Analysis of EEG Time Series. The European Physical Journal Special Topics (2022, p. 1-15).



RPs, O1 electrode



Example of recurrence plots, 30 s, electrode O1, subject S001.
Dark dots show distances inside small ϵ neighborhood.

RQA measures

RR, recurrence rate, density of recurrence points in a recurrence plot:

$$RR = \frac{1}{N^2} \sum_{i,j=1}^N R(i, j).$$

percentage of recurrence points which form diagonal lines in the recurrence plot of minimal length ℓ_{min} or predictability of the dynamical system.

$$DET = \frac{\sum_{\ell=\ell_{min}}^N \ell P(\ell)}{\sum_{\ell=1}^N \ell P(\ell)},$$

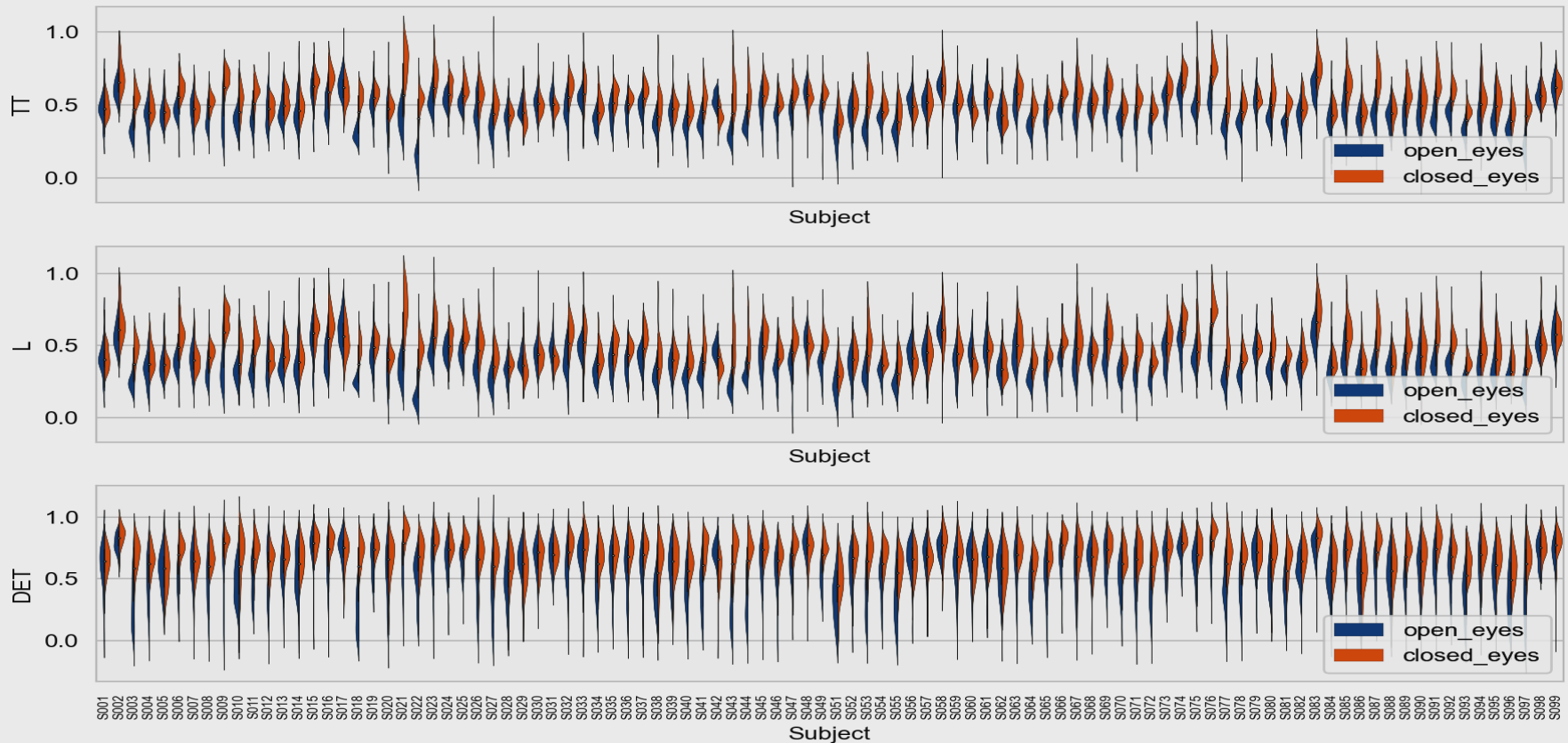
The averaged diagonal line length:

$$L = \frac{\sum_{\ell=\ell_{min}}^N \ell P(\ell)}{\sum_{\ell=\ell_{min}}^N P(\ell)}$$

Trapping time, measuring the average length of the vertical lines

$$TT = \frac{\sum_{v=v_{min}}^N v P(v)}{\sum_{v=v_{min}}^N P(v)}$$

RQA features for 64 electrodes



Distribution of trapping time, av. line length and determinism values for 64 electrodes shown for all 98 subjects. In some cases a single RQA feature allows for an easy separation of the two conditions. Variance is very different (focus? dreaming?), depending on the person. Linear SVM provides weights for (feature, electrode), facilitating selection of relevant EEG channels.

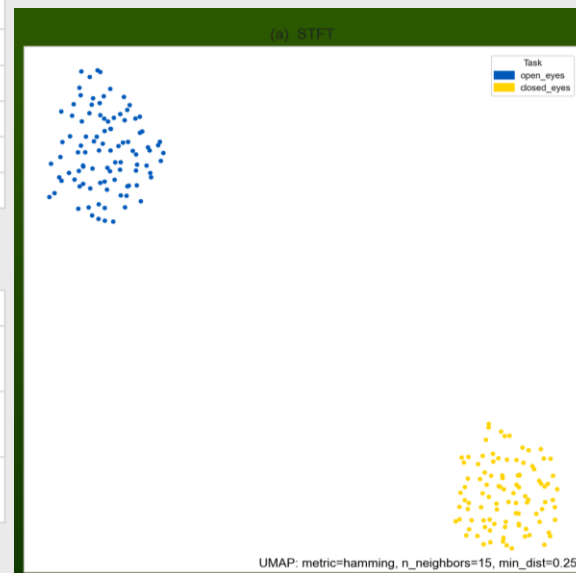
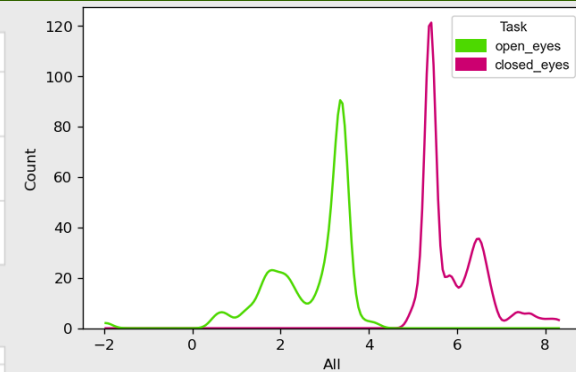
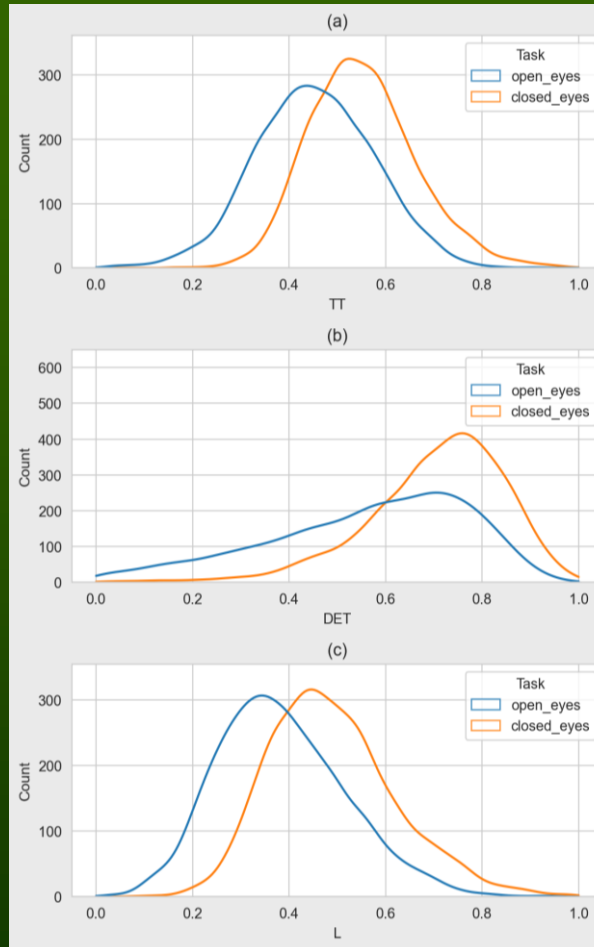
RQA features for 64 electrodes

Histograms of the RQA features for all 98 subjects:

TT (trapping time),
DET (determinism),
L (average diagonal line length).

Histograms of the projection of 320 FS feature values (5 RQA features x 64 electrodes), for all subjects, LSVM projection, for all data.

UMAP visualization of the 320-dimensional Z column vectors.



Labeling states

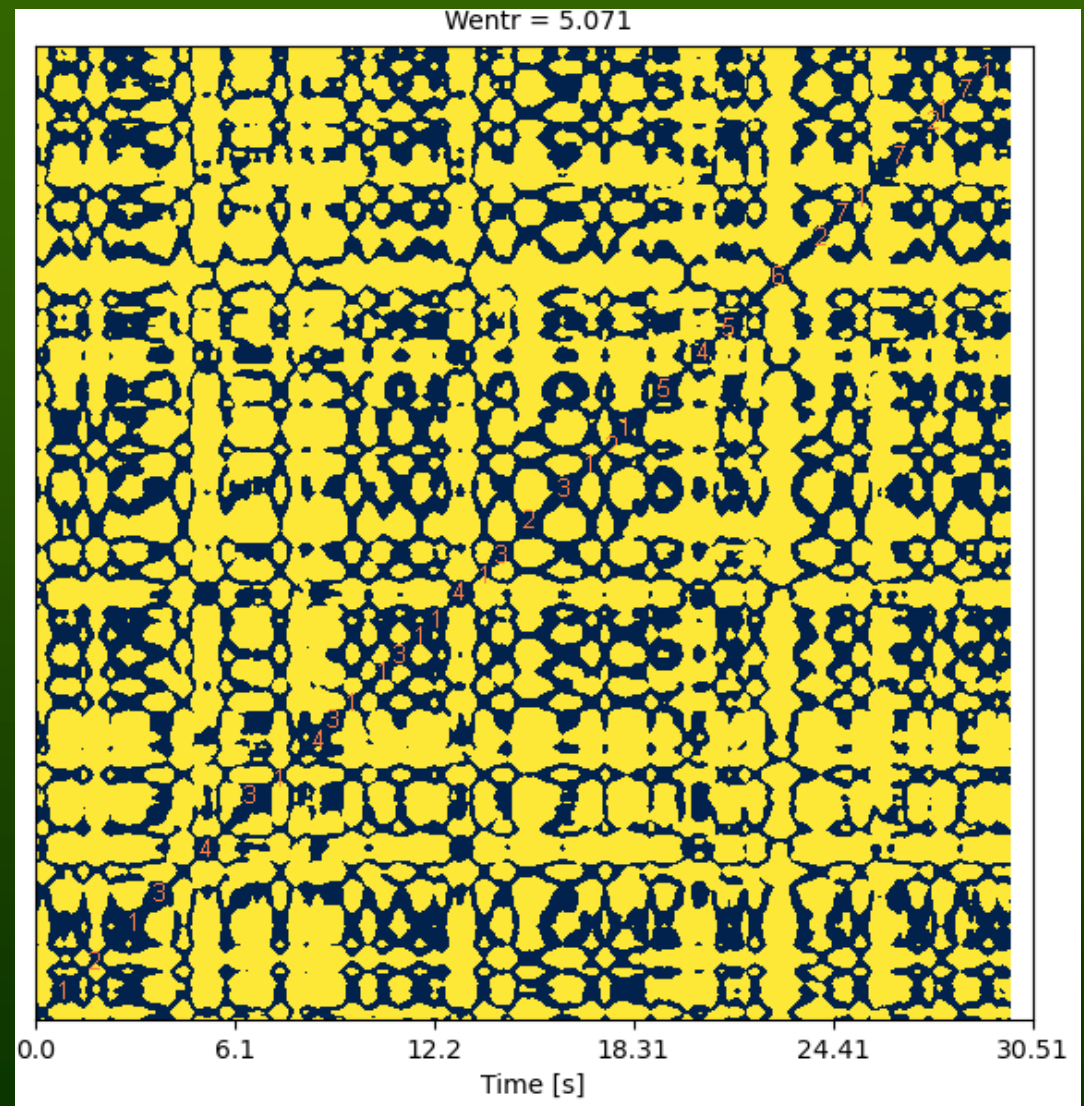
Automatic labeling of states and estimation of their recurrence may be important for biofeedback.

Metabolic costs of transitions between states may be important.

Ruminations? Pain states?
How external stimuli influence this dynamics?

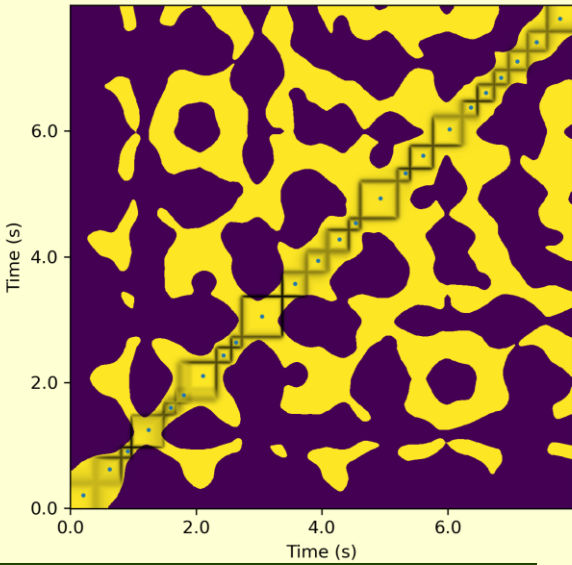
Needs automatic method for recognition of metastable, multivariate states.

More precise than microstates.

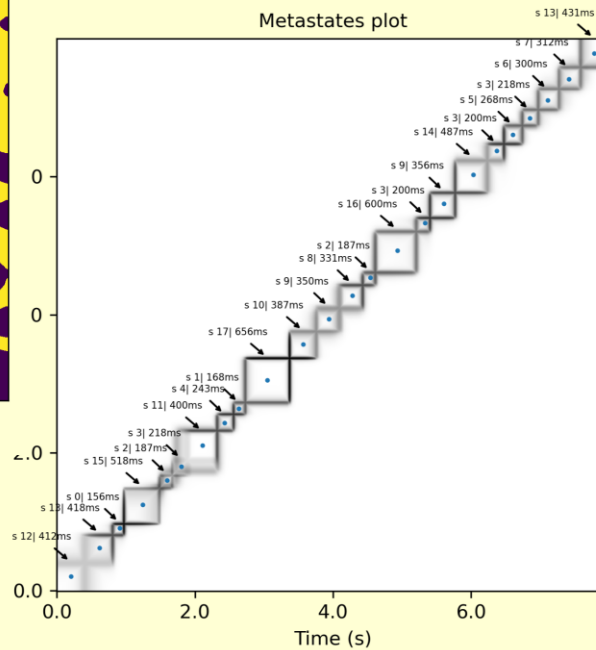


Segmentation of states

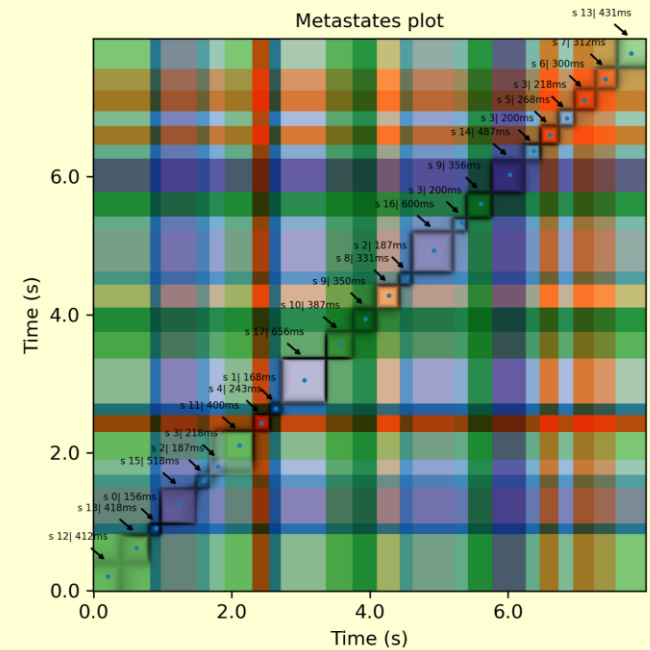
Metastates plot over recurrence plot



Metastates plot



Metastates plot



Labeling RPs

STFT rep, 1-50 Hz, represented by vectors X with 150 components.

Create RP matrices.

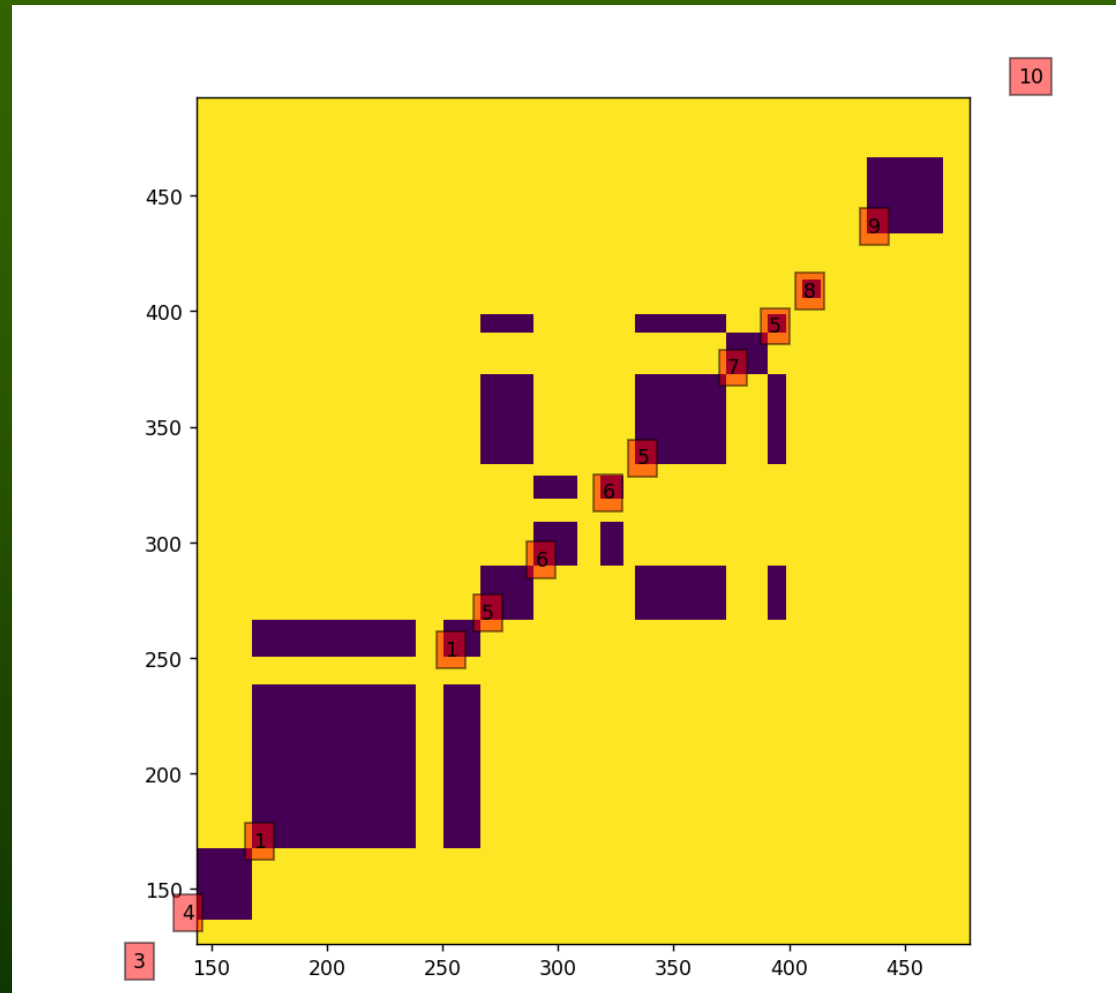
Smooth/erode data.

Identify stable regions along the diagonal.

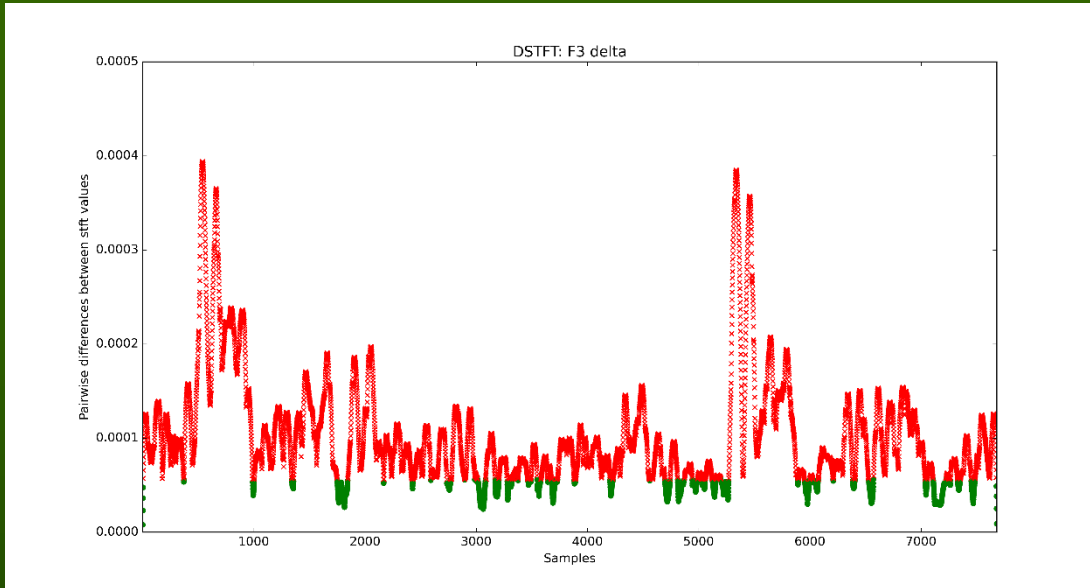
New state: no similar spectra in the past (row in upper part of the RP plot).

Problem: setting the threshold.

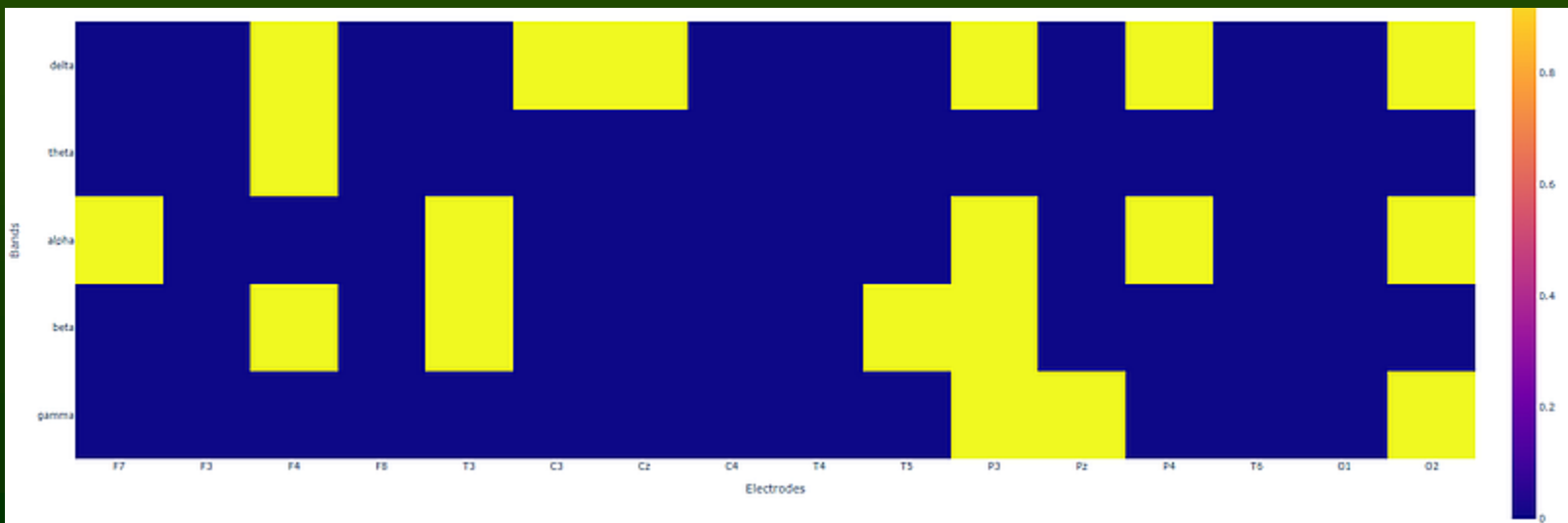
Compute power for individual states.



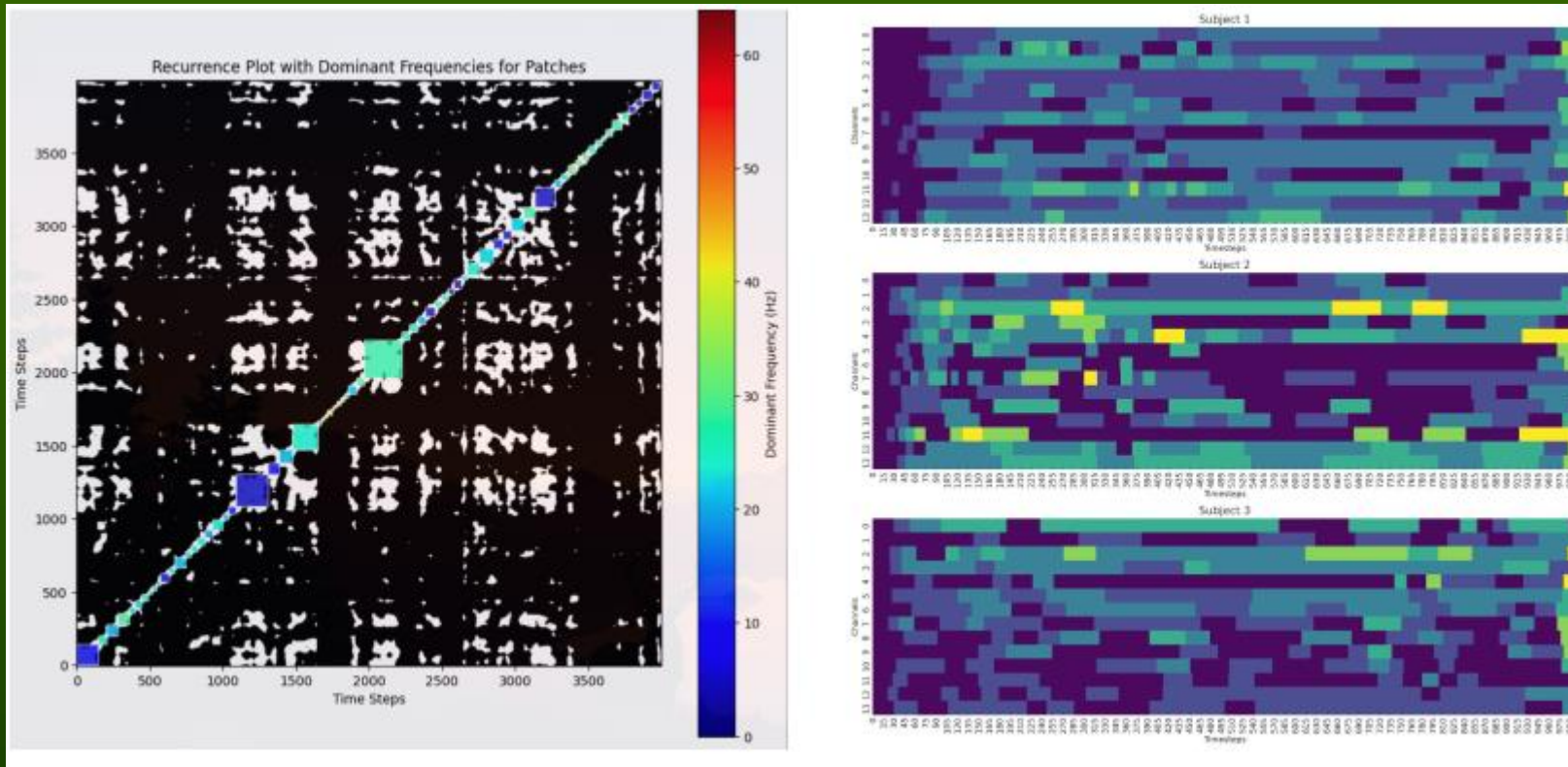
Labeling recurrent states



STFT rep, 1-50 Hz, represented by vectors X with 150 components.

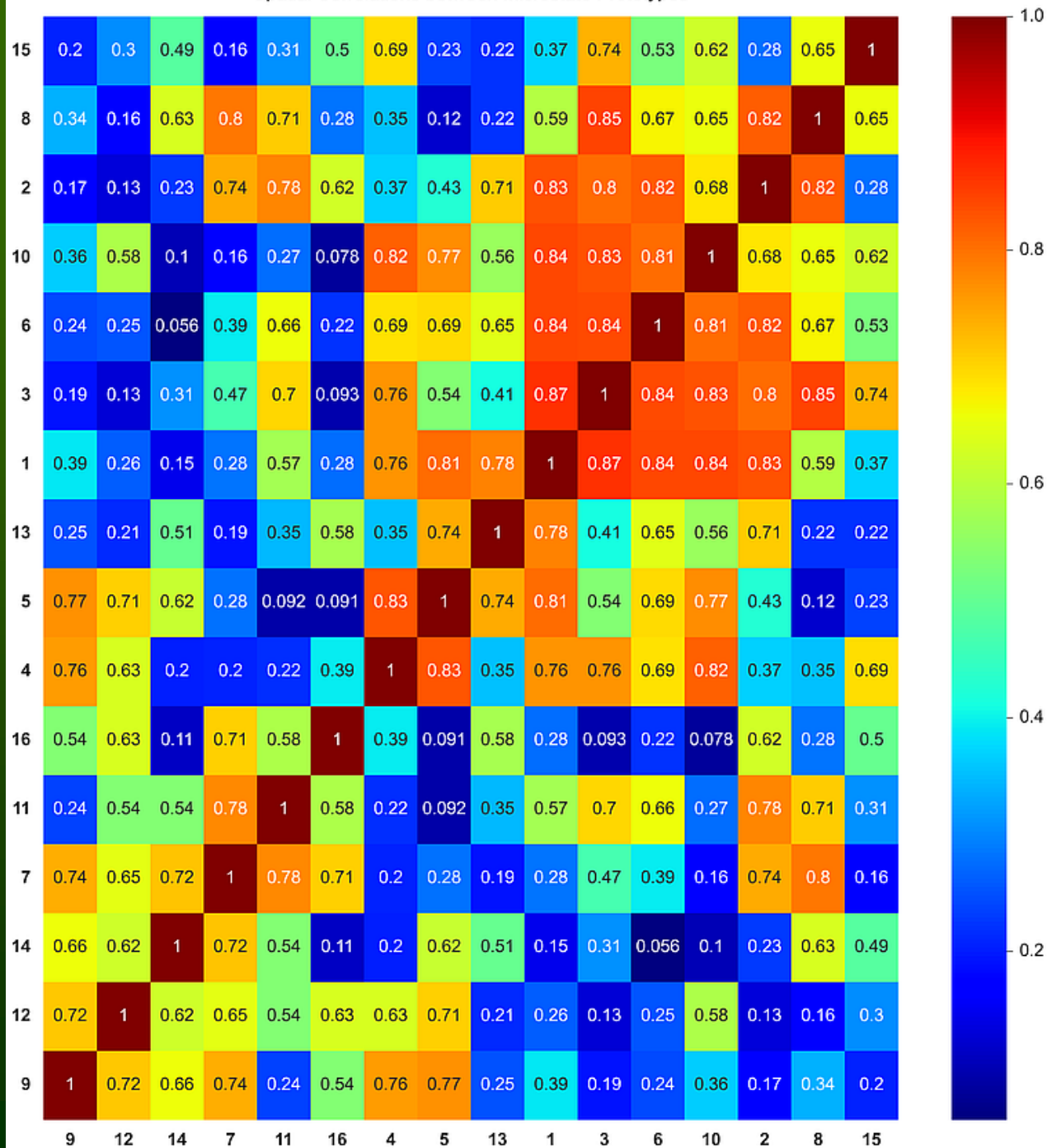


Labeled states



Example of RP with individual states, and patterns of states in discretized time steps for 3 subjects, 14 electrodes.

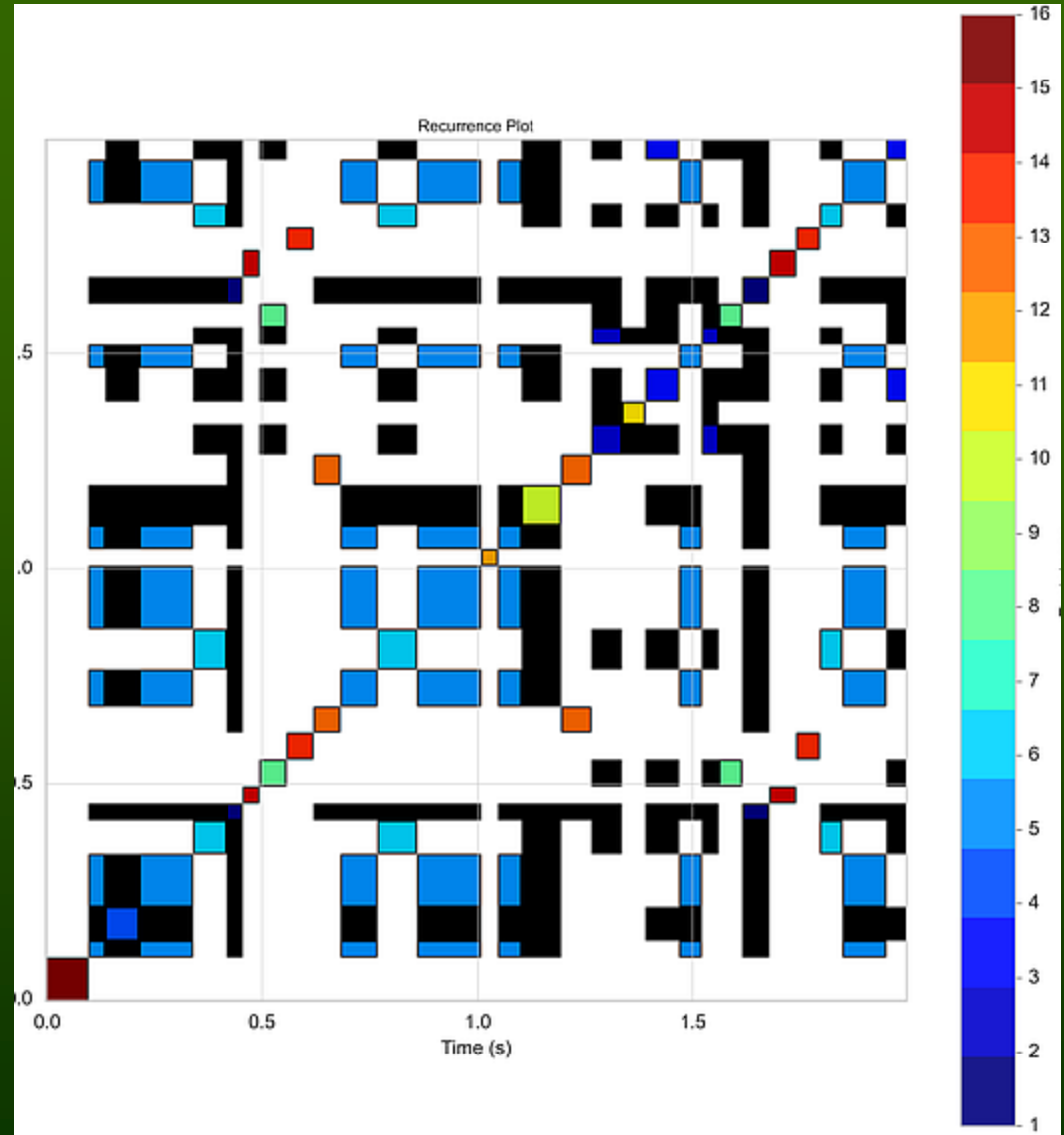
Spatial Correlations between Microstate Prototypes



Similarity of 16 microstate prototypes used to derive fuzzy versions of RQA measures.

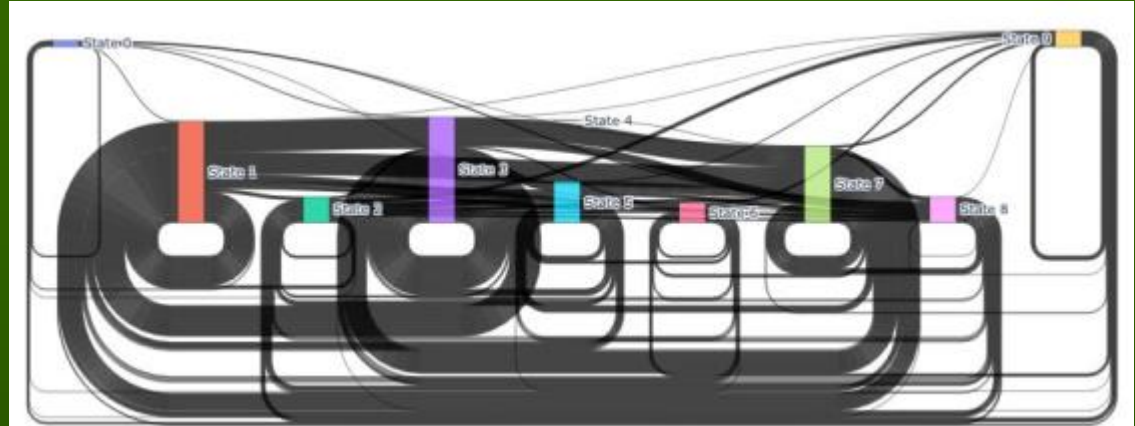
“Cleaned” recurrence plot of resting state dynamics based on 16 MS prototypes.

Good basis for identification of rare and recurrent states for biofeedback targets.

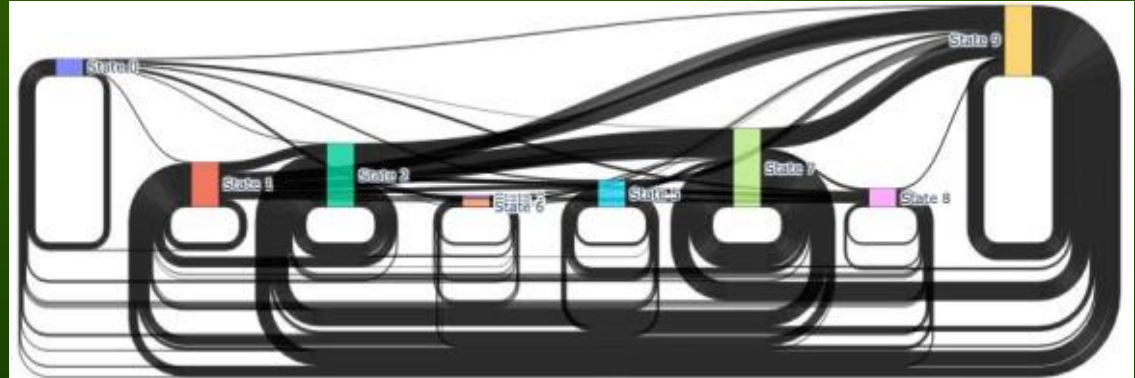


Sankey diagrams of dynamics

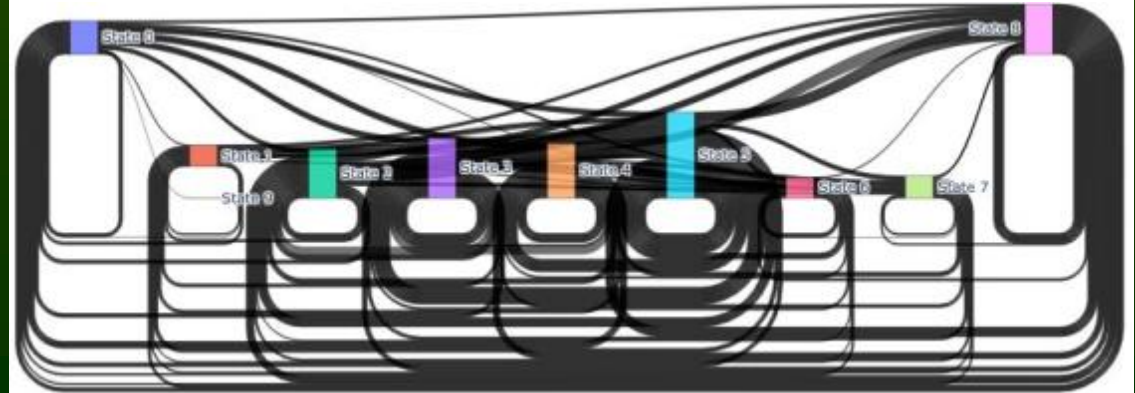
Control



ADHD



Schizophrenia



EEG diagnostics using symbolic dynamics

Transformer network with multi-head architecture attention mechanism,
applied to the state vectors.

Classification results (5xCV): recording from 16-19 electrodes only.

Schizophrenia dataset, 45/39 adolescent boys (10-14 years old).

Accuracy: 96.15%, Precision: 0.93-1.00, Recall: 0.92-1.00, F1-Score: 0.96-0.97

ADHD dataset, 61/60 control children 7-12 years old

Accuracy: 97.30%, Precision: 0.95-1.00, Recall: 0.95-1.00, F1-Score: 0.97

BCI Dataset, Motor imagery, hands vs feet, 106 subjects:

Accuracy: 99.72%, Precision: 0.99-1.00, Recall: 0.99-1.00, F1-Score: 1.0

Dobosz K, Duch W. (2010) Understanding Neurodynamical Systems via Fuzzy Symbolic Dynamics. Neural Networks Vol. 23 (2010) 487-496, 2010

VIRTUAL BR41N.IO HACKATHON

📅 April 17-18, 2021

during the

Spring School 2021*



*BR41N.IO and Spring School 2021 are part of g.tec's Teaching Plan 2021 with more than 140 hours of online courses and lectures.



1. PLACE WINNER

"NeuroBeat"

BCI application

Team members: Alicja Wicher, Joanna Maria Zalewska, Weronika Sójka, Ivo John Krystian Derezinski, Krzysztof Tołpa, Lukasz Furman, Sławomir Duda

IMPROVING HUMAN DAILY LIFE FUNCTIONING

NEUROHACKATOR 2021

21. - 23.
MAY 2021 //
ONLINE

SATURDAY

Project development
in groups



STARTS
10 a.m.

SUNDAY

Evaluation



ENDS
10 a.m.

FRIDAY

Organisers
presentation



workshops
with Judges

working 24h

REQUIREMENTS:

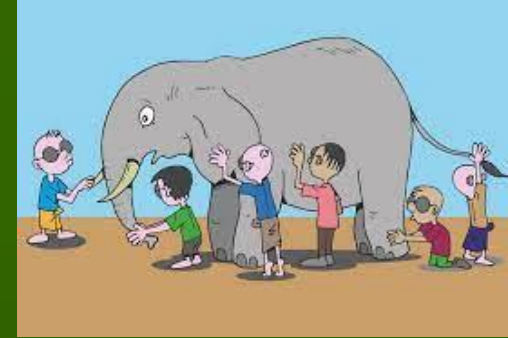
1. Create a team consisting of **3-5 people**.
2. Fill in the Registration Form (available on Facebook event).

DO YOU HAVE ANY QUESTIONS?

Write an e-mail:
NEUROTECTOR@GMAIL.COM

Neurotechnology Scientific Club
Center for Modern Interdisciplinary Technologies
at Nicolaus Copernicus University in Toruń
Wileńska 4 Street

Perspectives



- Optimization of brain processes is our biggest challenge! Medical diagnostics and closed loop systems for therapy of the brain disorders are the driving forces.
- Simple robust methods may be used in clinical applications. Asymptotic spatial distributions and temporal dynamics based on large number of microstates give surprisingly high diagnostic accuracy.
- Spatial, temporal and structural aspects of brain signals should be integrated.
- Promising way to such integration may be based on recurrence quantification analysis using spectral signal representation, RQA combined with microstates, and graphs of transitions between dynamical states.
- RQA may be based on comparison with different reference states, including microstates or avPP states.
- Feature selection done within crossvalidation partition may identify untypical cases that are not correctly recognized if specific features are not used.
- Models trained on small data will not give biomarkers of clinical value. Large databases are needed to handle idiosyncratic cases.

Towards Human-like Intelligence

IEEE Computational Intelligence Society Task Force (Mandziuk, Duch, M. Woźniak),
Towards Human-like Intelligence



IEEE Symposium on Computational Intelligence for Human-like Intelligence
(**IEEE SSCI CIHLI**), 12/2022 in Singapore, 12/2023 in Mexico City.

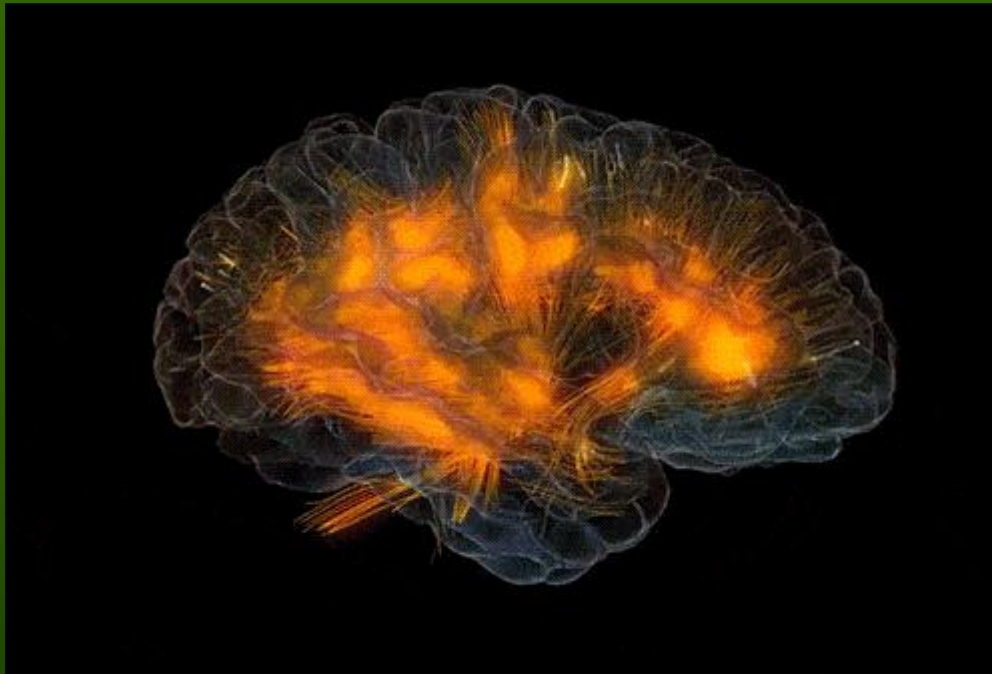
Distributed Artificial Brains (DAB) session (Duch, Mandziuk, Woźniak).

AGI: conference, 6/2023 in Stockholm

Journal of Artificial General Intelligence comments on Cognitive Architectures
and Autonomy: A Comparative Review (eds. Tan, Franklin, Duch).

BICA: Annual International Conf. on Biologically Inspired Cognitive Architectures,
13th Annual Meeting of the BICA Society, Guadalajara, Mexico 2023.

Thank you for synchronization of your neurons.



We have funds for internships and postdock support.

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