



#### Al for better brains



#### Włodzisław Duch

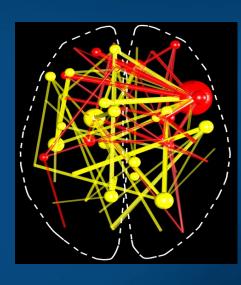
Neurocognitive Laboratory,
Center for Modern Interdisciplinary Technologies,
Dept. of Informatics, Faculty of Physics, Astronomy & Informatics,
Nicolaus Copernicus University

Google: Wlodek Duch

## On the threshold of a dream ... (50 y!)



Some ideas on how to optimize and repair human brains with AI.



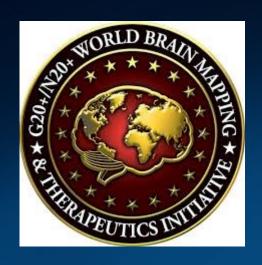
Duch W. (2012) Mind-Brain Relations, Geometric Perspective and Neurophenomenology, American Phil. Assoc. Newsletter 12(1), 1-7. Duch, W. (2019) Mind as a shadow of neurodynamics. Physics of Life Reviews



I've worked on many topics: computational intelligence algorithms, meta-learning in CI, neural networks, data understanding, NLP, similarity based methods, visualization, computational creativity, ASD, neuroinformatics, computational physics/quantum chemistry, philosophy of mind.

http://www.is.umk.pl/~duch/cv/WD-topics.html

# Global Brain Initiatives, or why is this so important?







The mission of IEEE Brain is to facilitate cross-disciplinary collaboration and coordination to advance research, standardization and development of technologies in neuroscience to help improve the human condition.

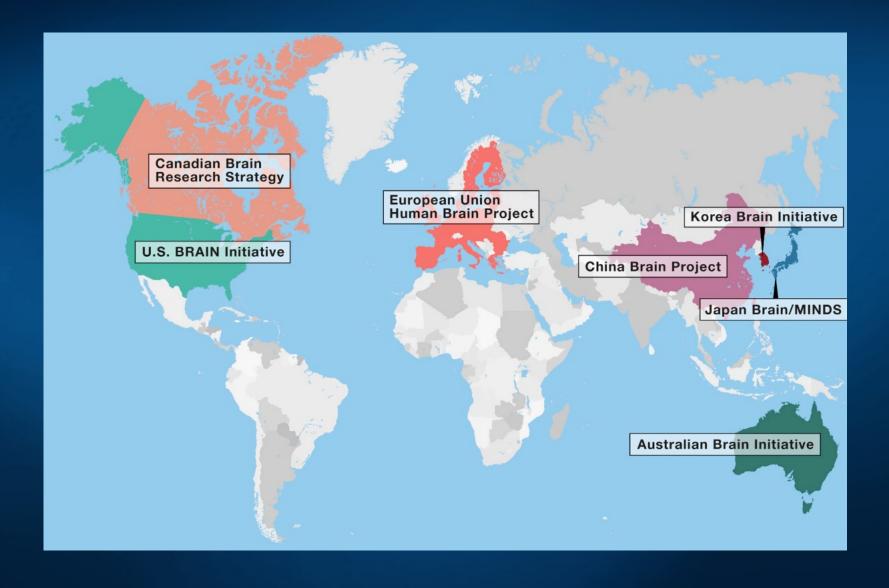
Al-neuroscience convergence! 20 IEEE Societies are involved, including:

**IEEE Computational Intelligence Society**; Computer Society; Consumer Electronics Society; Digital Senses Initiative; Robotics and Automation Society; Sensors Council; Signal Processing Society; Society on Social Implications of Technology; **Systems, Man, and Cybernetics Society**, International Neuroethics Society, and a few other societies.

Most these societies are also involved in artificial intelligence.

**Satya Nadella (CEO, Microsoft):** <u>examples of technology</u> that can be applied to empower more than one billion people with disabilities around the world.

### International Brain Initiatives



# Involvement in large EU Initiatives

# 2005, Beyond the Horizon **TG 5 group: Intelligent and Cognitive Systems**

Recommended work on mind-body co-evolution, materials and growth technologies, morphological computation, emerging behavior.

This was adopted in FP7, I wrote *votum separatum*, recommending work on artificial minds, NLP, bots and avatars, creativity, cognitive architectures.

2011, FET Work Programme, **Human Computer Confluence** panel. Merging Minds and Machines: Integrating AI with current Brain Research and future Neurotechnologies.

#### Two FET Flagship projects:

- 2010, The Mind and Brain Model Project.
- 2018 Future FET Flagship: Sapiens5.0, The science and technology for a 22<sup>nd</sup>-century humanity.

# Beyond the Horizon

**Anticipating Future and Emerging Information Society Technologies** 

#### Costs of brain diseases

Big ICT companies: Amazon, Apple, Google, Microsoft + Chinese giants Tencent, Baidu, and Alibaba, entering AI in health care (3 T\$ industry).

Brain research is most important: Al ⇔ Neuroscience ⇔ Neuropsychiatry.

Gustavsson et al. (2011). Cost of disorders of the brain in Europe 2010. *European Neuropsychopharmacology*, 21(10), 718–779.

179 million, or **1/3 of all European citizens**, had at least one brain disorder. 45% of the total annual health budget of Europe!

Total cost of brain disorders in EU estimated in 2010: 798 billion €/year.

China: >20% of population (~250 mln) suffering from some mental disorder.

European Brain Council (EBC) reports (2010; 2014).

Consensus Statement on European Brain Research (2015) includes a chapter

on Computational Neuroscience, data repositories and analytics.





#### Advance Neurotechnologies

Accelerate the development and application of new neurotechnologies.



Support multi-disciplinary teams and stimulate research to rapidly enhance current neuroscience technologies and catalyze innovative scientific breakthroughs.

Human Brain Project, EU Flagship, and Obama BRAIN Initiative (2013): Brain Research through Advancing Innovative Neurotechnologies.

"Develop new technologies to explore how the brain's cells and circuits interact at the speed of thought, ultimately uncovering the complex links between brain function and behavior. Explore how the brain records, processes, uses, stores, and retrieves vast quantities of information. Help bring safe and effective products to patients and consumers."

Since 2013 numerous exciting developments in neurotechnology and our understanding of the brain have been made by scientists across the globe.



This workshop was part of the Brain-Machines Interface Workshop and SMC2018 conference, organized by Mike Smiths (UC Berkeley).

Special meeting of **Global Current and Emerging Brain Initiative leaders was** attended by IEEE President, James Jefferies, President-elect Toshio Fukuda, and representatives from Australia to USA (NSF and NIH), **IEEE Brain Initiative**, International Neuroethics Society, industry, and other stakeholders.

#### Neuroscience => Al

Hassabis, D., Kumaran, D., Summerfield, C., Botvinick, M. (2017).

**Neuroscience-Inspired Artificial Intelligence**. *Neuron*, *95*(2), 245–258.

Affiliations: Google DeepMind, Gatsby, ICN, UCL, Oxford.

Bengio, Y. (2017). The Consciousness Prior. ArXiv:1709.08568.

Amoset al. (2018). Learning Awareness Models. ICRL, ArXiv:1804.06318.

#### Al Systems inspired by Neural Models of Behavior:

- (A) Visual attention, foveal locations for multiresolution "retinal" representation, prediction of next location to attend to.
- (B) Complementary learning systems and episodic control: fast learning hippocampal system and parametric slow-learning neocortical system.
- (C) Models of working memory and the Neural Turing Machine.
- (D) Neurobiological models of synaptic consolidation.

**SANO**, new Centre for Individualized Computational Medicine in Kraków (EU Team project, with Sheffield Uni, Fraunhofer Society, Research Centre Juelich.

#### BICA, Brain-Inspired Cognitive Architecture

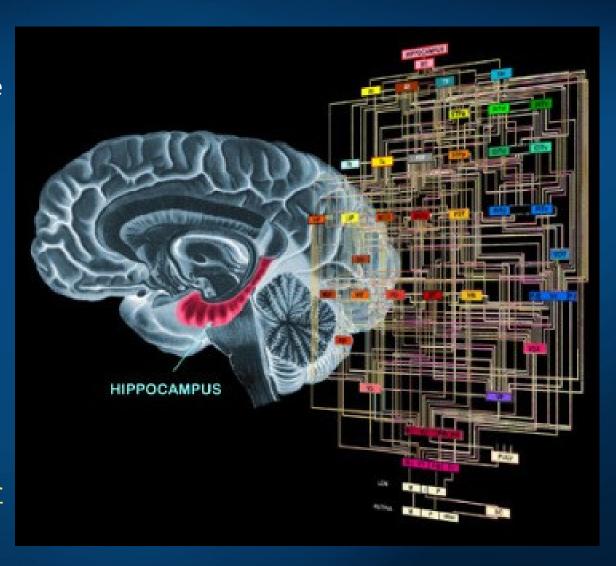
Understanding the brain from engineering perspective means to build a model of the brain showing similar functions.

Cognitive informatics, Neurocognitive Informatics.

BICA = Brain Inspired Cognitive Architecture.

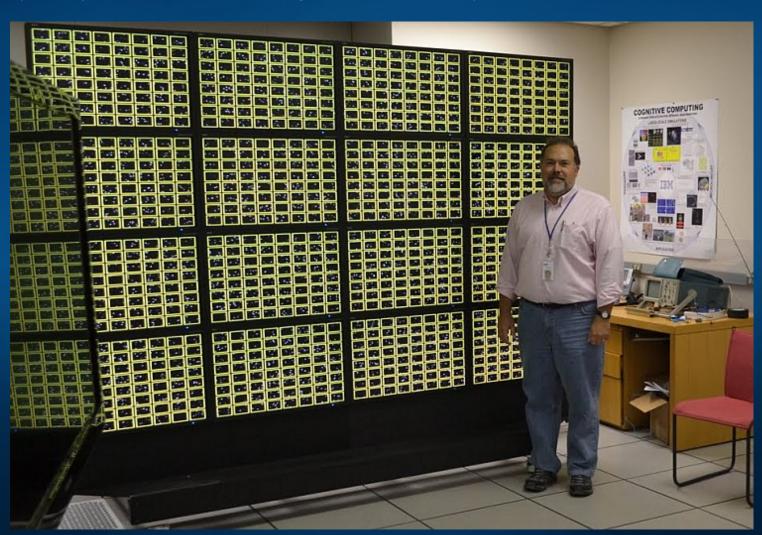
Review: Duch, Oentaryo,
Pasquier,
Cognitive architectures: wher
e do we go from here

? 2008



# Neuromorphic wall

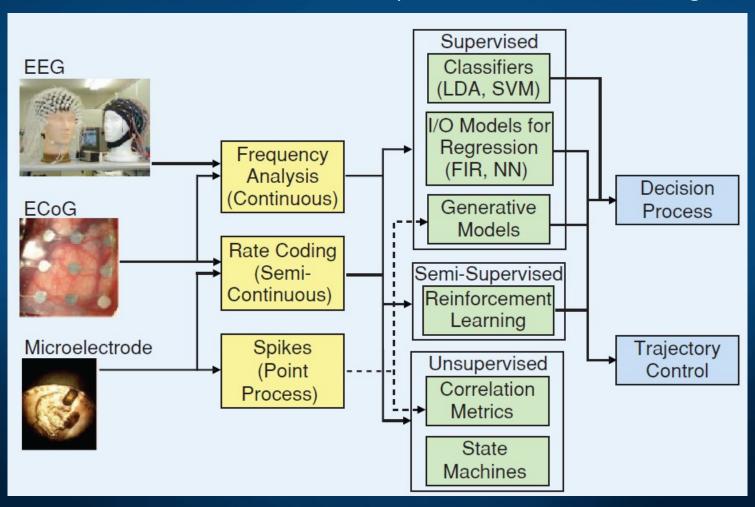
1024 TN neuromorphic chips, or 1B neurons and 256B synapses! Complexity ~ horse brain, 1/4 gorilla, 1/6 chimpanzee.



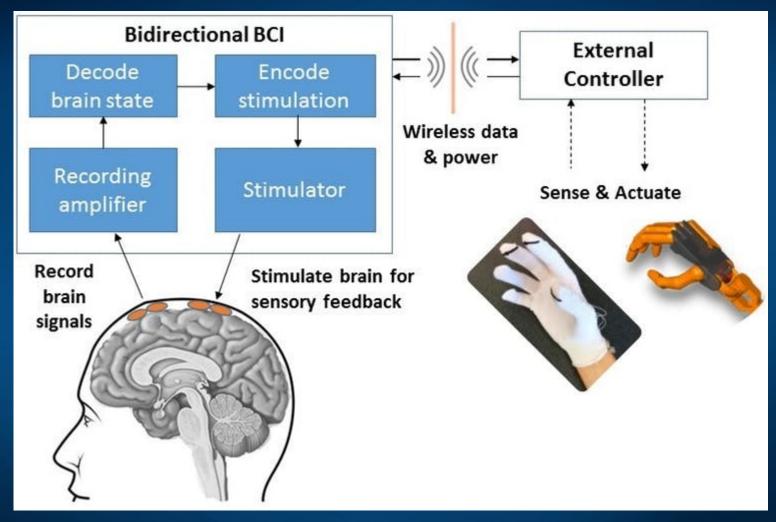
# Optimizing brains Neurocognitive technologies

#### BCI: wire your brain ...

Non-invasive, partially invasive and invasive signals carry progressively more information, but are also harder to implement. EEG is still the king!

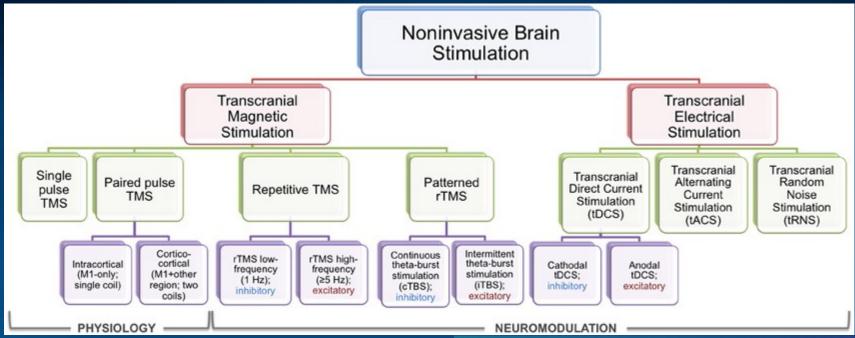


## Brain-Computer-Brain Interfaces



Neurofeedback + neuromodulation. Closed loop system with brain reading and stimulation for self-regulation. Sensory signals may com from Virtual Reality.

#### **Brain stimulation**



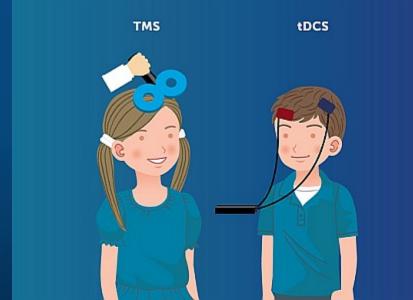
ECT – Electroconvulsive Therapy

VNS – Vagus Nerve Stimulation

Ultrasound, laser ... stimulation.

Complex techniques, but portable phones are also complex.

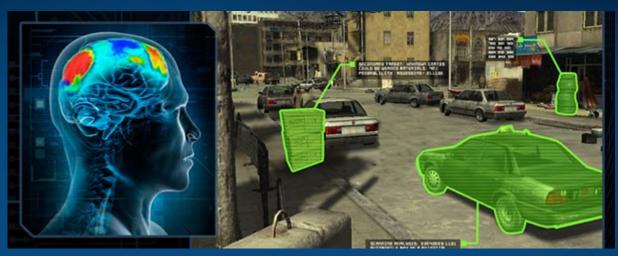
Attention? Just activate your cortex, no effort is needed!



### Trenowanie mózgu

Engagement Skills
Trainer (EST) to
procedury treningu
amerykańskich żołnierzy.

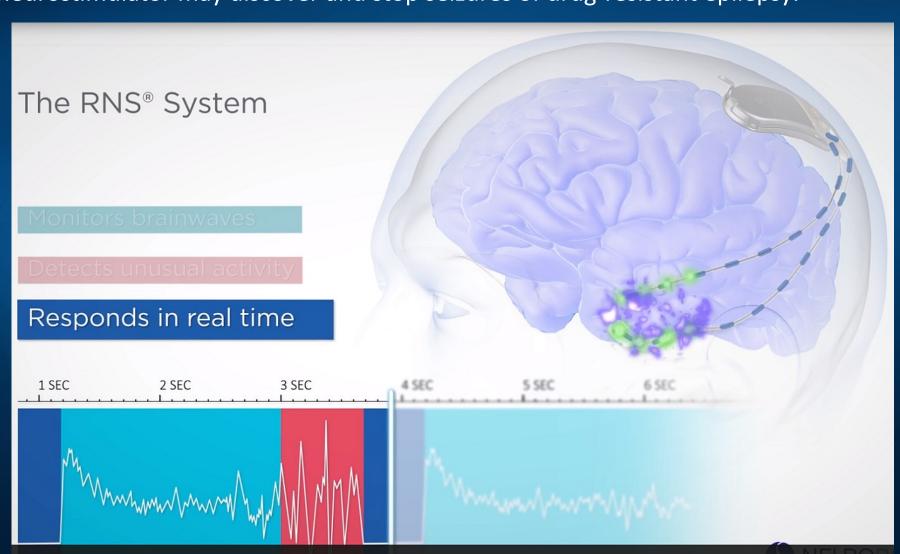
Intific Neuro-EST to technologia wykorzystująca analizę EEG i wielokanałowy stymulator przezczaszkowy (MtCS) do transferu umiejętności pomiędzy mistrzem i uczniem.





# **Epilepsy**

About 1% of people and some mammals suffers from epilepsy. Detector and neurostimulator may discover and stop seizures of drug-resistant epilepsy.



## Flow for depression

Complete at-home depression treatment: combining tDCS with behavioral therapy. 24% could overcome depression, 41% claimed 50% improvement.

#### https://flowneuroscience.com/

- Medication free. Home treatment.
- Reduces depression with a brain stimulation headset and free app for behavioural therapy.
- 18 sessions, each 30 min, 6 weeks.
- After 6 weeks, the activity in your frontal lobe is rebalanced and your depressive symptoms will have decreased.
- Approved for medical use in the EU and UK.

Early neurocognitive technology, but more precise analysis of individual brain activity with tDCS is needed for best adaptation.



## HD EEG/DCS?

EEG electrodes + DCS.

Reading brain states

- => transforming to common space
- => duplicating in other brains

#### Applications:

depression, neuro-plasticity, new neurofeedback, pain, psychosomatic disorders!

Multielectrode DCS stimulation with 256 electrodes induces changes in the brain increasing neuroplasticity.

Ex: Phillips Neuro EEG S400.



#### MemorEM

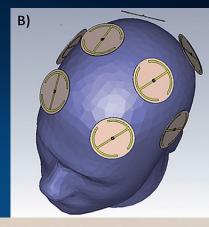
Transcranial Electromagnetic Treatment (TEMT)
MemorEM head device being worn by a subject.
Position of the eight electromagnetic emitters embedded between the device's two-layered head cap.
8 emitters 915 MHz, pulses 4.6 ms, 1.6 W/kg, provide global TEMT to the cortex and deeper structures.

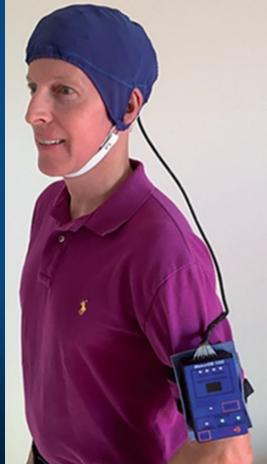
In AD transgenic mice TEMT prevents and reverses both cognitive impairment and brain amyloid- $\beta$  (A $\beta$ ) deposition. TEMT improves cognitive performance in normal mice. 3 disease-modifying and inter-related mechanisms of TEMT action:

1) anti-Aβ aggregation, both intraneuronally and extracellularly; 2) mitochondrial enhancement; and 3) increased neuronal activity.

8 mild/moderate AD patients were treated with TEMT, increased functional connectivity within CC area.

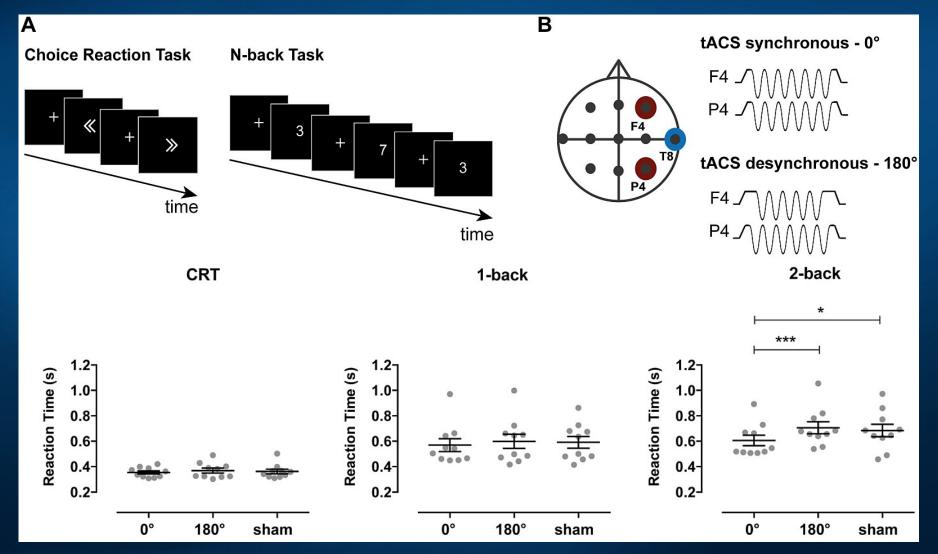
Arendash GW et. al. J. of Alzheimer's Dis 71 (2019) 57





## Synchronize PFC/PC to improve WM

Violante, I.R. et al. Externally induced frontoparietal synchronization modulates network dynamics and enhances working memory performance. ELife, 6 (2017).

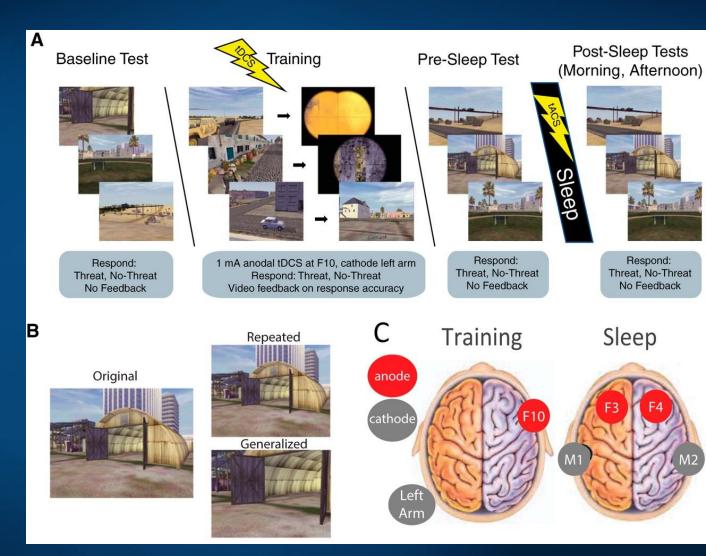


# **BCBI** and memory

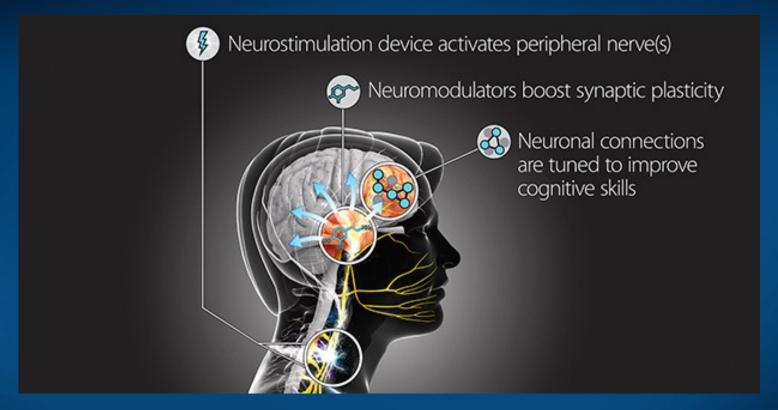
N. Ketz et al.,
Closed-Loop SlowWave tACS Improves
Sleep-Dependent
Long-Term Memory
Generalization
by Modulating
Endogenous
Oscillations.

J. Neuroscience 8 (33) 2018 Enhances the consolidation of

consolidation of recent experiences into long-term memory.



### Targeted Neuroplasticity Training

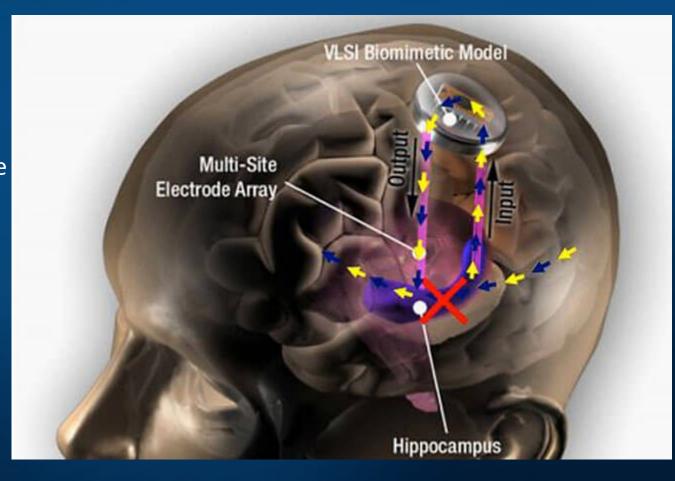


<u>DARPA (2017):</u> Enhance learning of a wide range of cognitive skills, with a goal of reducing the cost and duration of the Defense Department's extensive training regimen, while improving outcomes. TNT could accelerate learning and reduce the time needed to train foreign language specialists, intelligence analysts, cryptographers, and others.

#### Memory implants

Ted Berger (USC, <u>Kernel</u>): hippocampal neural prosthetics facilitate human memory encoding and recall using the patient's own hippocampal spatiotemporal neural codes. Tests on rats, monkeys and on people gave memory improvements on about 35% (<u>J. Neural Engineering 15, 2018</u>).

DARPA: Restoring **Active Memory** (RAM), new closedloop, non-invasive systems that leverage the role of neural "replay" in the formation and recall of memory to help individuals better remember specific episodic events and learned skills.

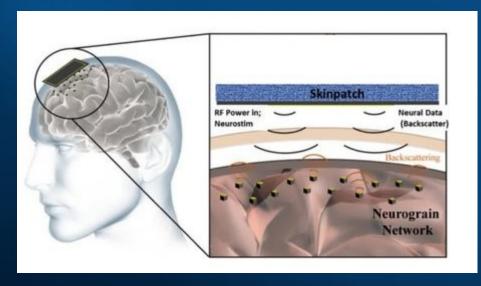


### Million nanowires in your brain?

DARPA (2016): **Neural Engineering System Design (NESD)**Interface that reads impulses of 10<sup>6</sup> neurons, injecting currents to 10<sup>5</sup> neurons, and reading/activating 10<sup>3</sup> neurons.

DARPA <u>Electrical Prescriptions</u> (<u>ElectRx</u>) project enables "artificial modulation of peripheral nerves to restore healthy patterns of signaling in these neural circuits. ElectRx devices and therapeutic systems under development are entering into clinical studies."

Neural lace i neural dust project for cortex stimulation.

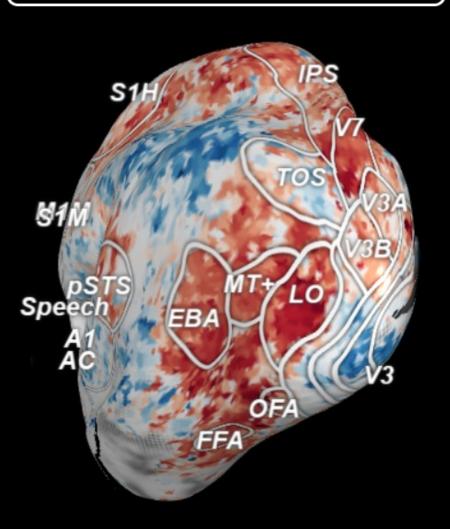


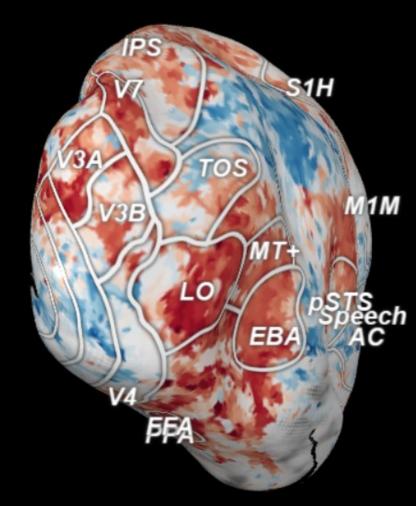


# Decoding mental states



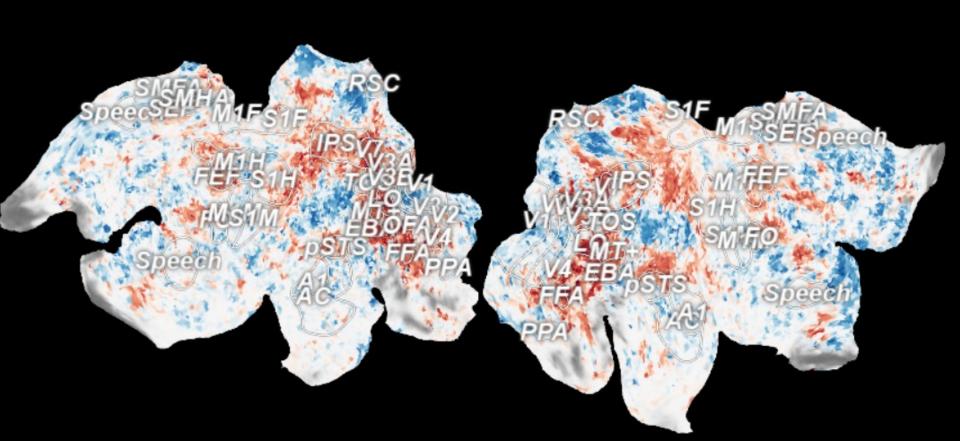
#### Category zebra: Passive Viewing





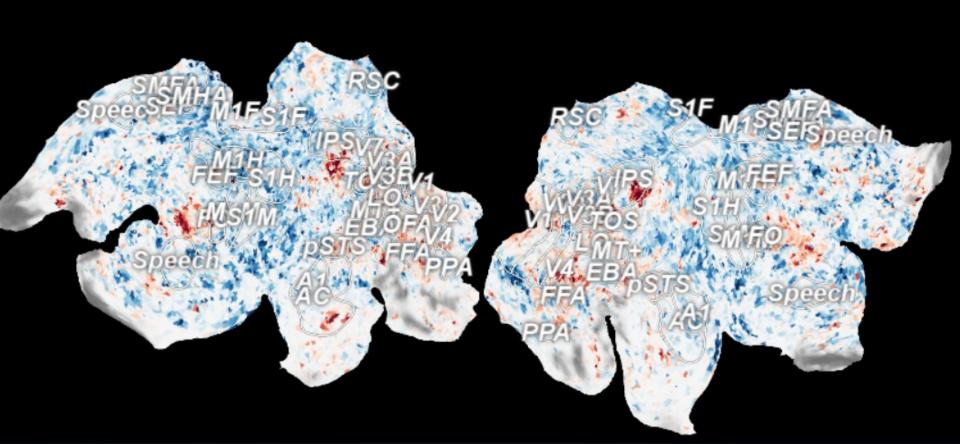


Category zebra: Passive Viewing



Category traffic light: Passive Viewing

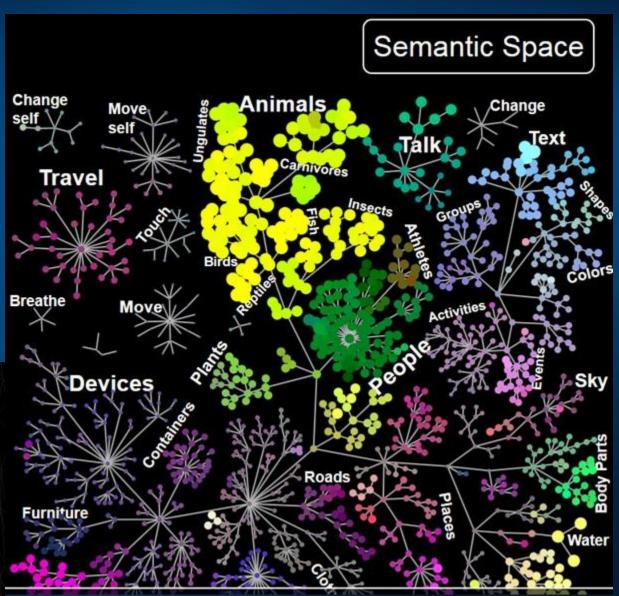




### Semantic neuronal space

Words in the semantic space are grouped by their similarity.
Words activate specific ROIs, similar words create similar maps of brain activity.
Video or audio stimuli, fMRI 60.000 voxel).
Gallant lab, Berkeley.





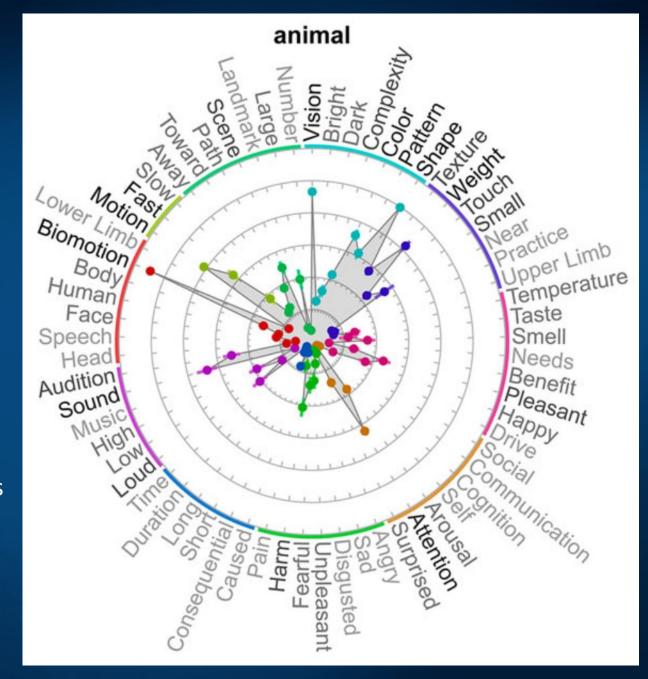
65 attributes related to neural processes;

Colors on circle: general domains.

J.R. Binder et al
Toward a Brain-Based
Componential Semantic
Representation, 2016

More than just visual objects!

Decompose brain signals for a given concept into components coding these attributes.



#### Brains Minds

**Cognitive neuroscience**: map  $S(M) \Leftrightarrow S(B)$ , as in BCI.

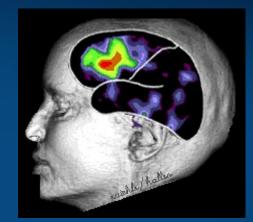
How do we describe the state of mind?

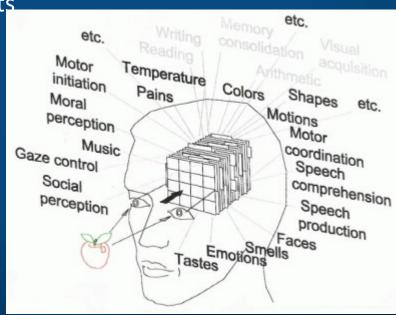
Verbal description is not sufficient unless words are represented in a space with dimensions that measure different aspects of experience.

Stream of mental states, movement of thoughts

trajectories in psychological spaces.

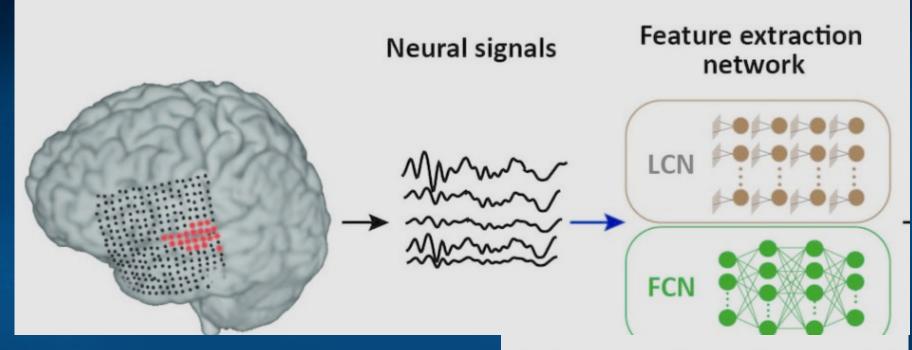
**Two problems**: discretization of continuous processes for symbolic models, and lack of good phenomenology – we are not able to describe our mental states. Neurodynamics: bioelectrical activity of the brain, neural activity measured using EEG, MEG, NIRS-OT, PET, fMRI ...



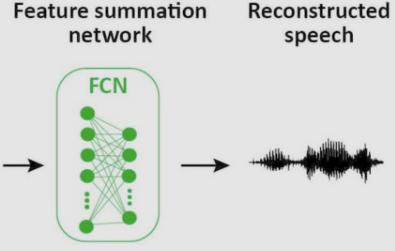


Duch W (1996) Computational physics of the mind. CPC 97: 136-153

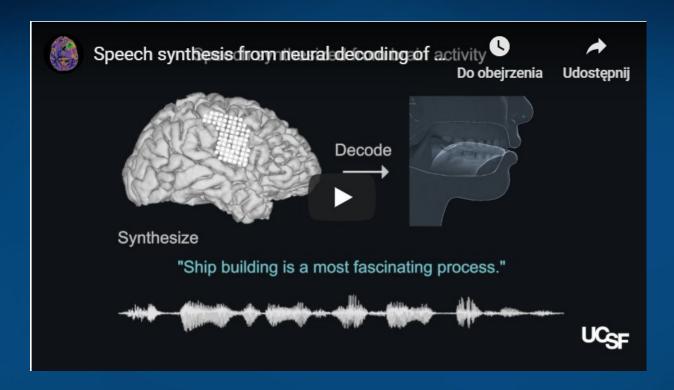
### Your brain is taking



Electrocorticographical signals may be used to train neural network and control a vocoder.



### Listing to thoughts



Patterns of cortical activations in higher order human auditory cortex allows for neural decoding of speech acoustic parameters, decoder is used to synthesize speech when a participant silently mimed sentences.

Pasley et al. (2012); G.K. Anumanchipalli, J. Chartier, E.F. Chang, Speech synthesis from neural decoding of spoken sentences. <u>Nature 24/4/2019</u>

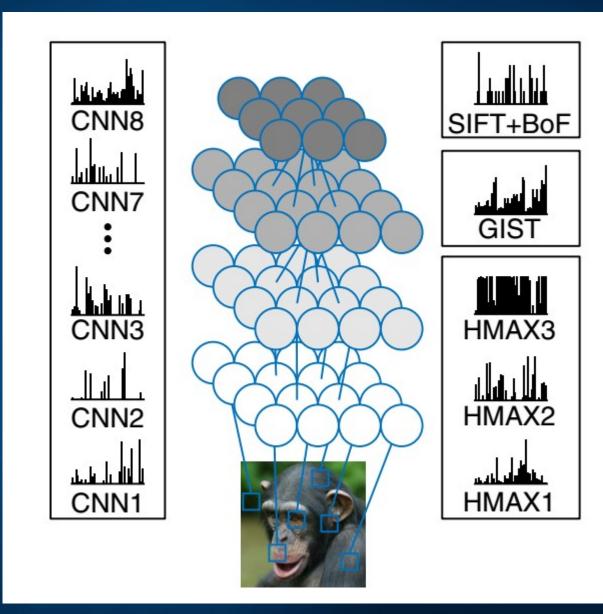
### Mental images from brain activity

Can we convert activity of the brain into the mental images that we are conscious of?

Try to estimate features at different layers.

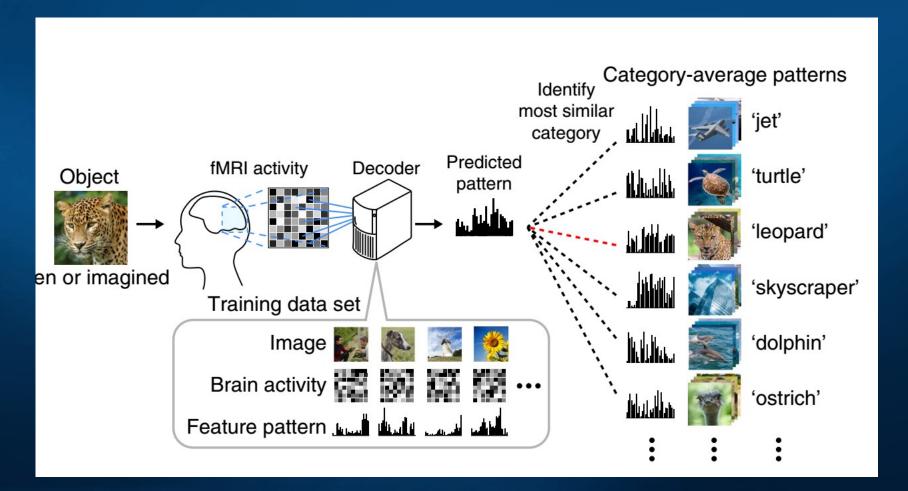
8-layer convolution network, ~60 mln parameters, feature vectors from randomly selected 1000 units in each layer to simplify calculations.

Output: 1000 images.



## Brain activity Mental image

fMRI activity can be correlated with deep CNN network features; using these features closest image from large database is selected. Horikawa, Kamitani, Generic decoding of seen and imagined objects using hierarchical visual features. Nature Comm. 2017.



### Neural screen

Features are discovered, and their combination remembered as face, but detailed recognition needs detailed recording from neurons – 205 neurons in various visual areas used.

L. Chang and D.Y. Tsao, "The code for facial identity in the primate brain". *Cell* 2017

DARPA (2016): put million nanowires in the brain!
Use them to read neural responses and 10% of them to activate neurons.

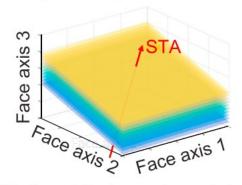
1. We recorded responses to parameterized faces from macaque face patches

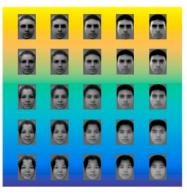


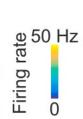
ML/MF

2. We found that single cells are tuned to single face axes, and are blind

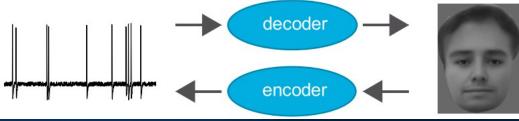
to changes orthogonal to this axis







3. We found that an axis model allows precise encoding and decoding of neural responses



## Brain networks

### Neuropsychiatric phenomics

2008: The

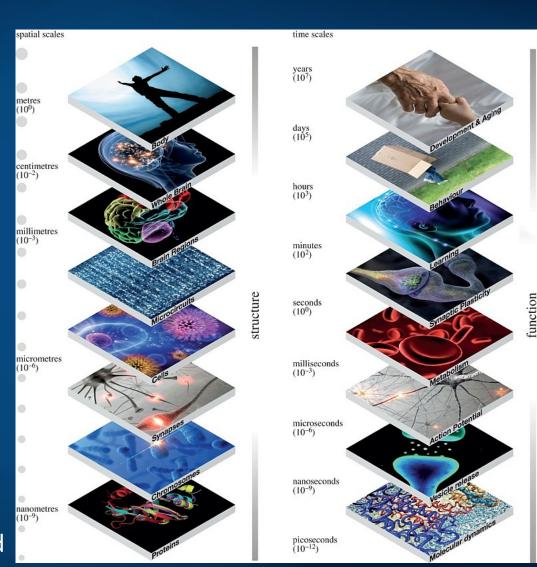
<u>Consortium for Neuropsychiatric</u> <u>Phenomics</u>

"... categories, based upon presenting signs and symptoms, may not capture fundamental underlying mechanisms of dysfunction" (Insel et al., 2010).

New approach: <u>RDOC NIMH</u>.

Description of organisms at different levels will help to answer different types of questions.

Network level is in the middle and can be connected to the mental level via computational models.



## RDoC Matrix for "cognitive domain"

Genes

Molecules

Cells

Circuits Physiology Behavior

Self-

**Paradigms** 

Construct/Subconstruct

55/104/45554255/104/455		XXX				,		Report	
Attention		Elements	Elements	Elements	Elements	Elements	Elements		Elements
Perception	Visual Perception	Elements							
	Auditory Perception	Elements							
	Olfactory/Somatosensory/Multimodal/Perception								Elements
Declarative Memory		Elements							
Language		Elements			Elements	Elements	Elements	Elements	Elements
Cognitive	Goal Selection; Updating, Representation, and Maintenance ⇒ Focus 1 of 2 ⇒ Goal Selection				Elements			Elements	Elements
	Goal Selection; Updating, Representation, and Maintenance ⇒ Focus 2 of 2 ⇒ Updating, Representation, and Maintenance	Elements							
	Response Selection; Inhibition/Suppression ⇒ Focus 1 of 2 ⇒ Response Selection	Elements							
	Response Selection; Inhibition/Suppression ⇒ Focus 2 of 2 ⇒ Inhibition/Suppression	Elements							
	Performance Monitoring	Elements	Elements		Elements	Elements	Elements	Elements	Elements
Working Memory	Active Maintenance	Elements	Elements	Elements	Elements	Elements			Elements
	Flexible Updating	Elements	Elements	Elements	Elements	Elements			Elements
	Limited Capacity	Elements	Elements		Elements	Elements			Elements
	Interference Control	Elements	Elements	Elements	Elements	Elements			Elements

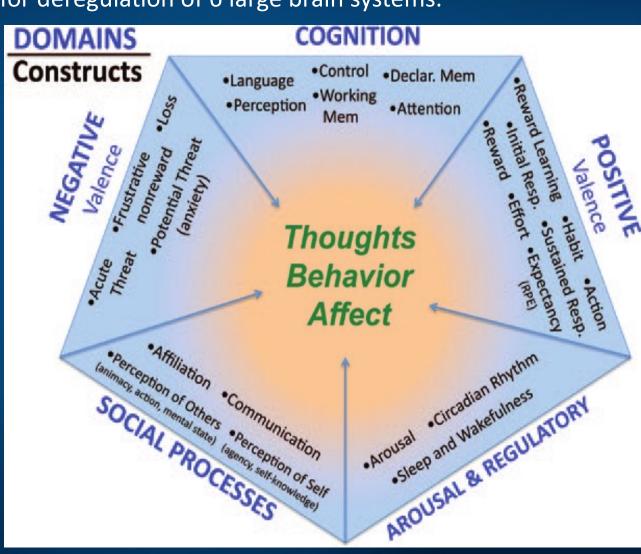


NIMH RDoC Matrix for deregulation of 6 large brain systems.

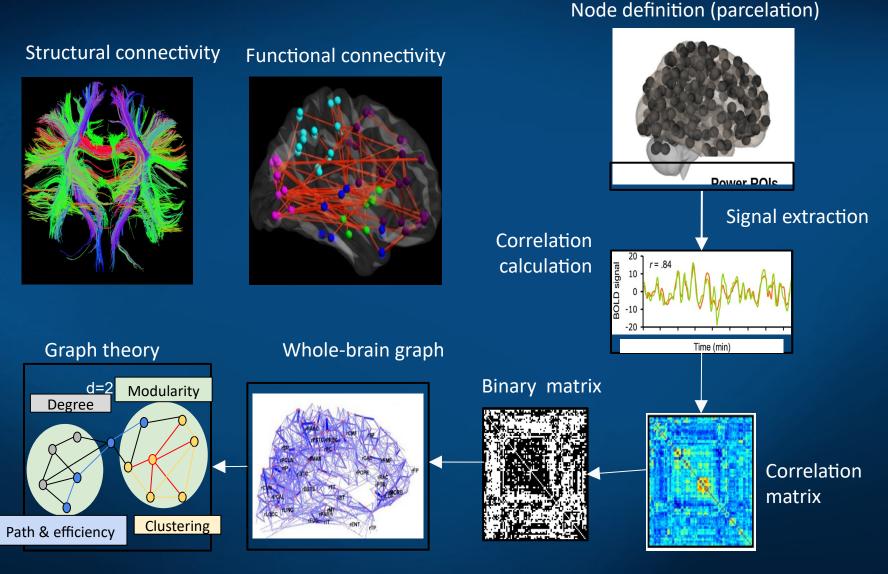
Psychological constructs are necessary to talk about mental states.

Sensorimotor systems added in Jan. 2019 as sixth brain system.

How are they related to physical processes?

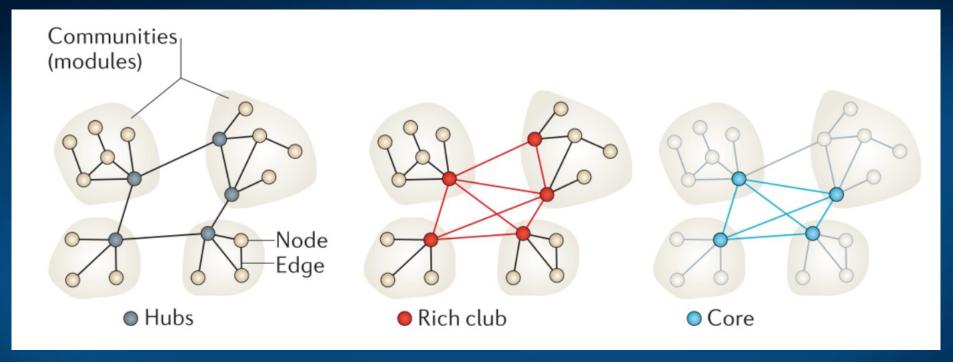


Human connectome and MRI/fMRI



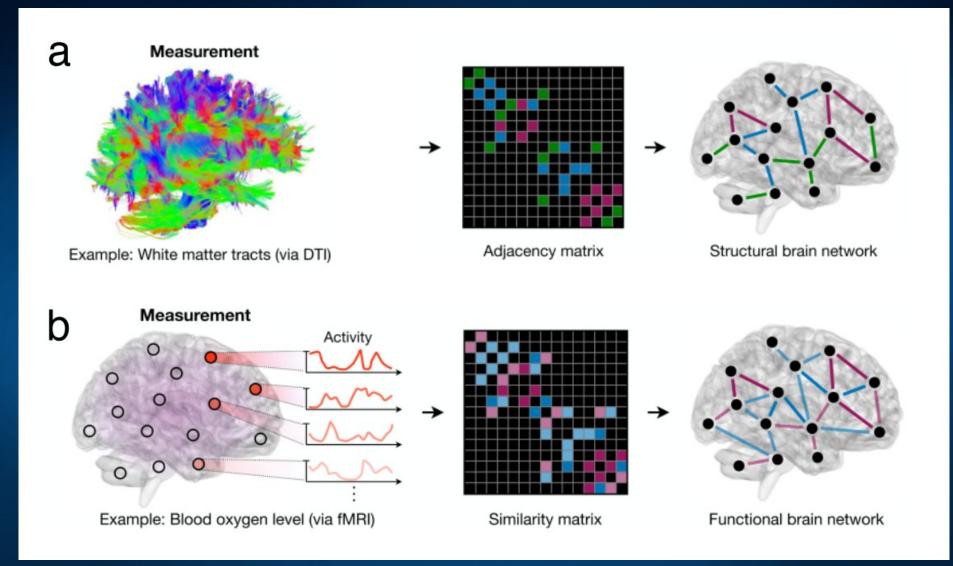
Many toolboxes available for such analysis.

### **Network Neuroscience**

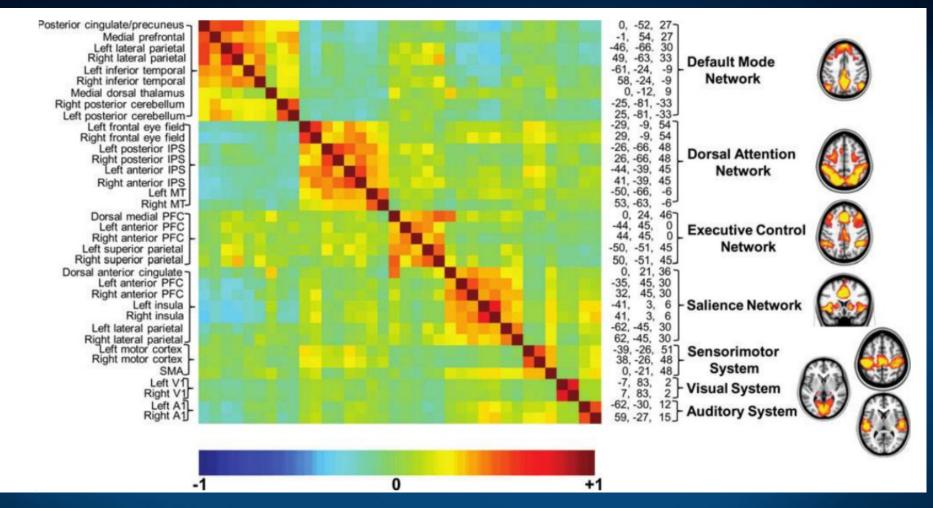


Network neuroscience is focused on identifying network structures, hubs, rich clubs and cores of the network. Hubs connect modules via long-distance connections. Hubs are also often densely interconnected forming so called 'rich club' or integrated core. New ways of quantification of various network structures are being developed.

Bullmore and Sporns (2012) The economy of brain network organization. Nature Reviews Neuroscience, 13(5):336.



Lynn and Bassett (2018) The physics of brain network structure, function, and control. arXiv:1809.06441.



Correlation matrix representing resting-state functional connectivity between selected brain regions Shows stronger connectivity for 7 large-scale brain networks: default mode (DM), dorsal attention (DAT), executive control network (FPN, CON), salience (SAL), sensorimotor (SOM), visual (VSN), auditory (ASN). Switching DMN  $\Leftrightarrow$  Salience  $\Leftrightarrow$  FPN

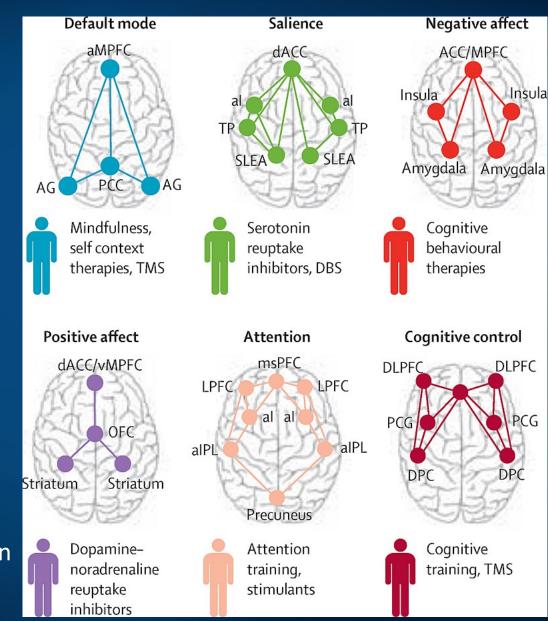
### Multi-level phenomics

Research Domain Criteria (RDoC) matrix is based on multi-level neuropsychiatric phenomics describing large brain systems deregulation, but links to behavior should be analyzed at the network level, where specialized functions are implemented. In AI:

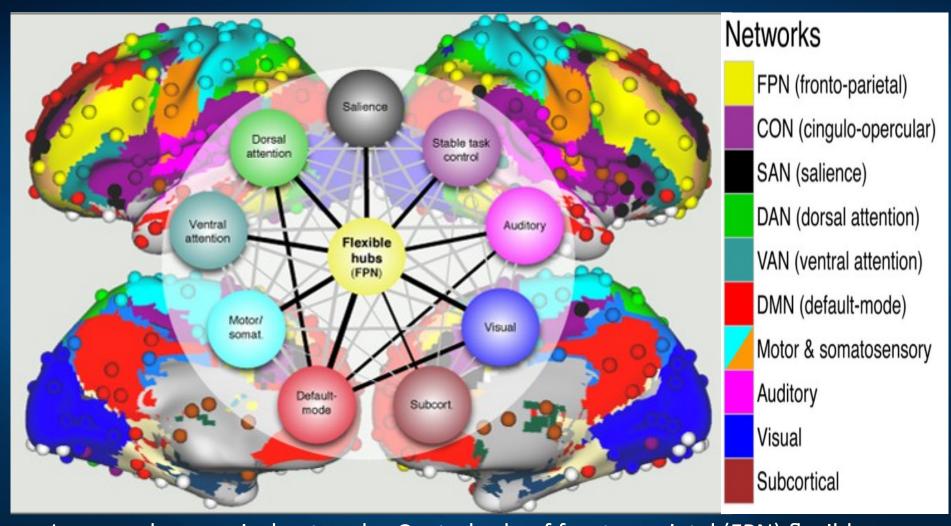
M. Minsky, Society of mind (1986)

Decompose brain network dynamics into meaningful components of activity related to various brain functions.

Include influence of genes, molecules, cells, circuits, physiology, behavior, self-reports on network functions.

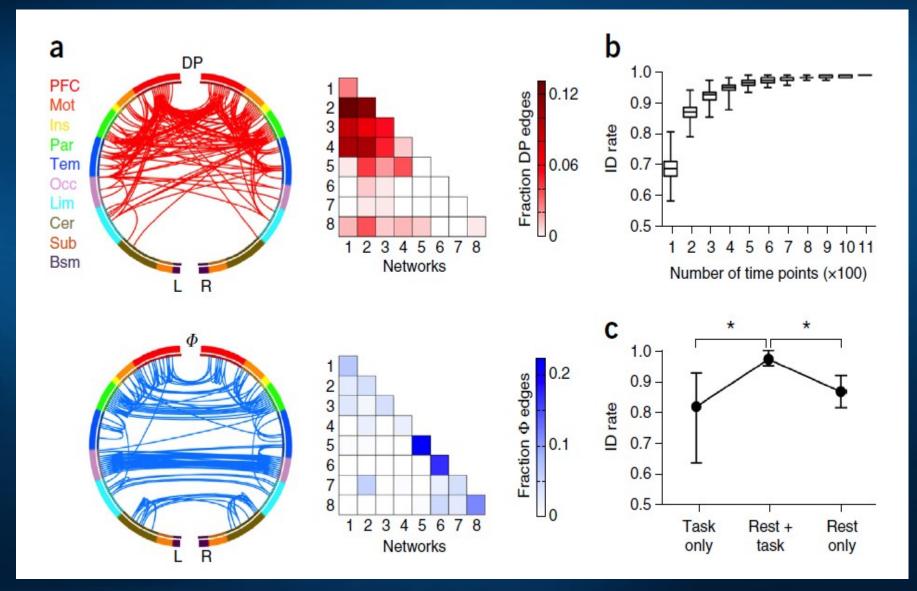


### Neurocognitive Basis of Cognitive Control



Large scale canonical networks. Central role of fronto-parietal (FPN) flexible hubs in cognitive control and adaptive implementation of task demands (black lines=correlations significantly above network average). Cole et al. (2013).

Finn et al. (2015), **Functional connectome fingerprinting**: identifying individuals using patterns of brain connectivity. Nature Neuroscience. Top: highly unique; Bottom: highly consistent connections.

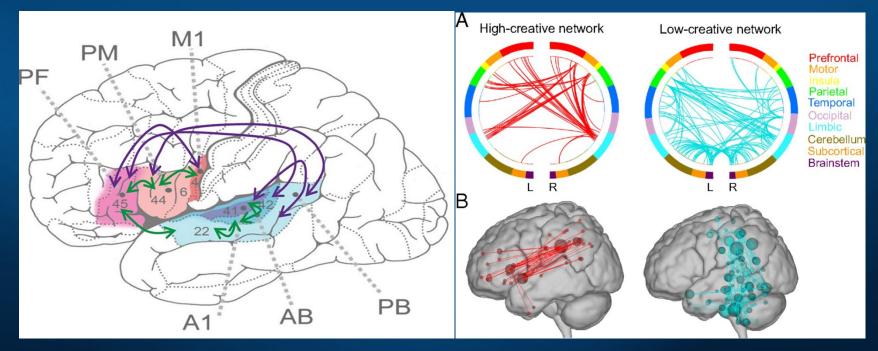


### Fluid nature



Development of brain in infancy: first learning how to move, sensorimotor activity organizes brain network processes, rather consistent.

<u>The Developing Human Connectome Project</u>: create a dynamic map of human brain connectivity from 20 to 44 weeks post-conceptional age, which will link together imaging, clinical, behavioral, and genetic information.

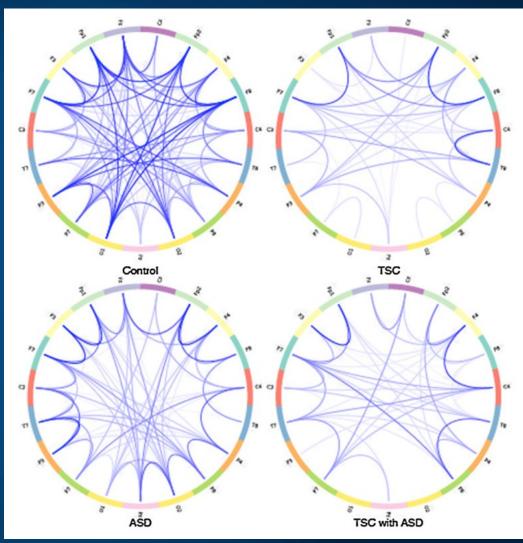


## ASD: pathological FC

Comparison of connections for patients with ASD (autism spectrum), TSC (Tuberous Sclerosis), and ASD+TSC.

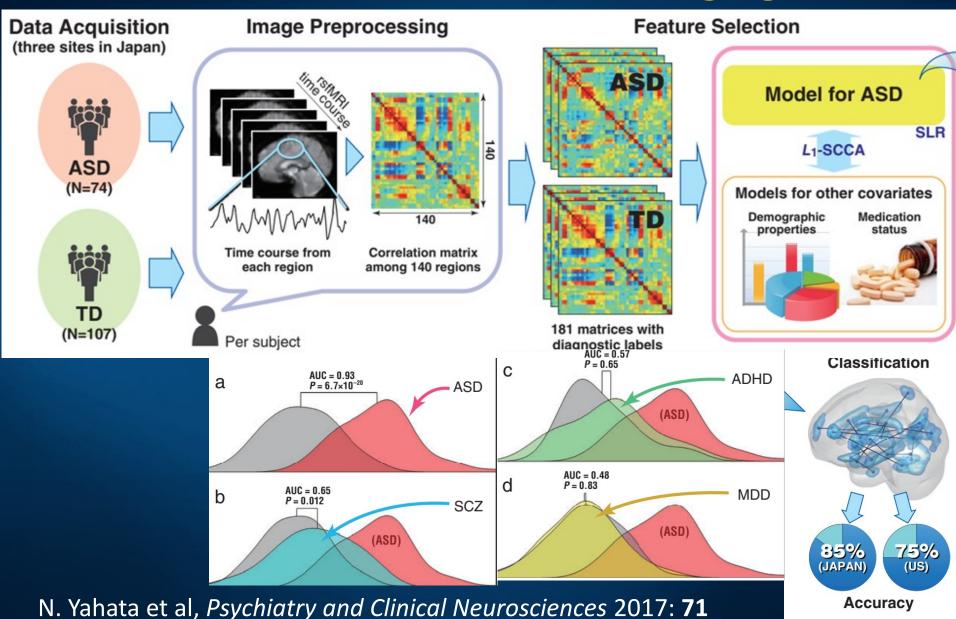
Coherence between electrodes.
Weak or missing connections
between distant regions prevent
ASD/TSC patients from solving
more demanding cognitive tasks.

Network analysis becomes very useful for diagnosis of changes due to the disease and learning; correct your networks!

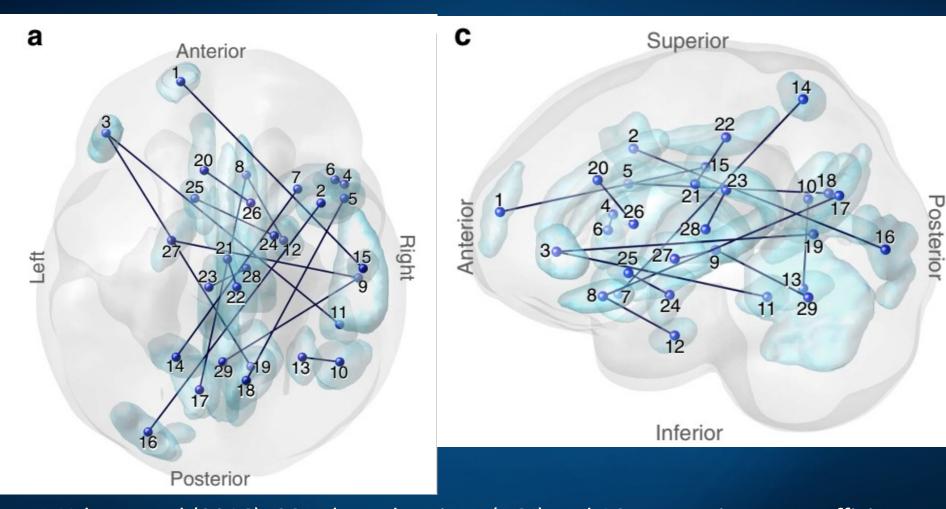


J.F. Glazebrook, R. Wallace, Pathologies in functional connectivity, feedback control and robustness. Cogn Process (2015) 16:1–16

## Biomarkers from neuroimaging

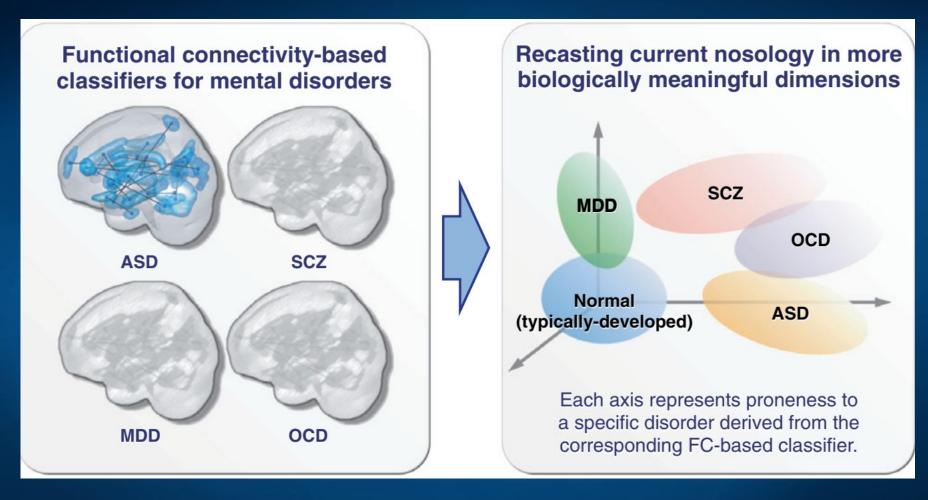


### Selected connections



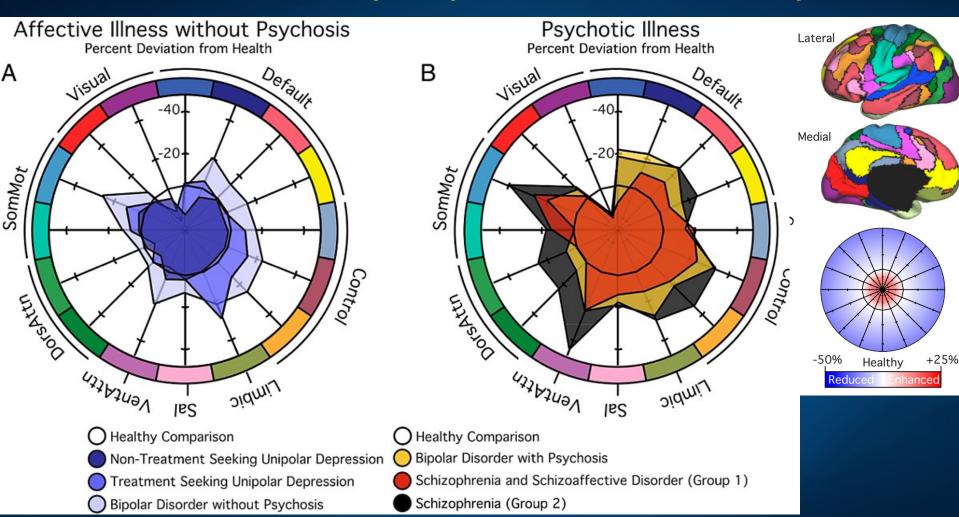
N. Yahata et al (2016): 29 selected regions (ROI) and 16 connections are sufficient to recognize ASD with 85% accuracy in 74 Japanese adult patients vs. 107 people in control group; without re-training accuracy was 75% on US patients.

### Biomarkers of mental disorders



MDD, deep depression, SCZ, schizophrenia, OCD, obsessive-compulsive disorder, ASD autism spectrum disorder. fMRI biomarkers allow for objective diagnosis. N. Yahata et al, *Psychiatry & Clinical Neurosciences* 2017; **71**: 215–237

### Connectivity in patients vs healthy



Regions determined based on the 17-network solution from Yeo et al. Control (health) = circle, % deviation in mean network connectivity shown.

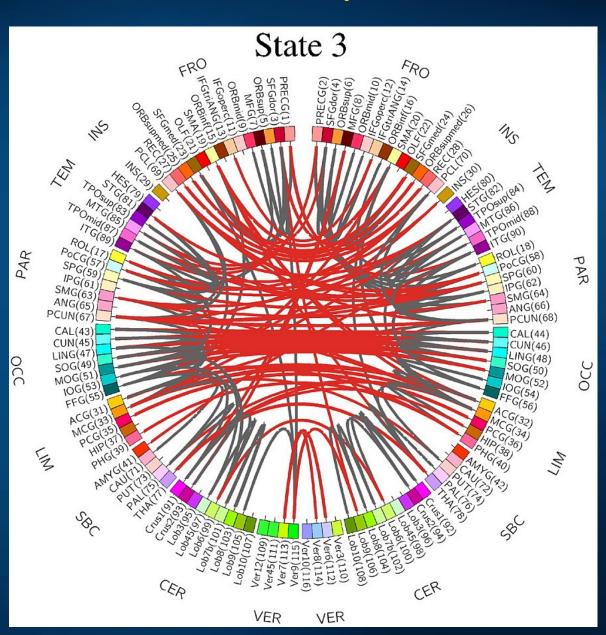
### Negative connections in MCI patients

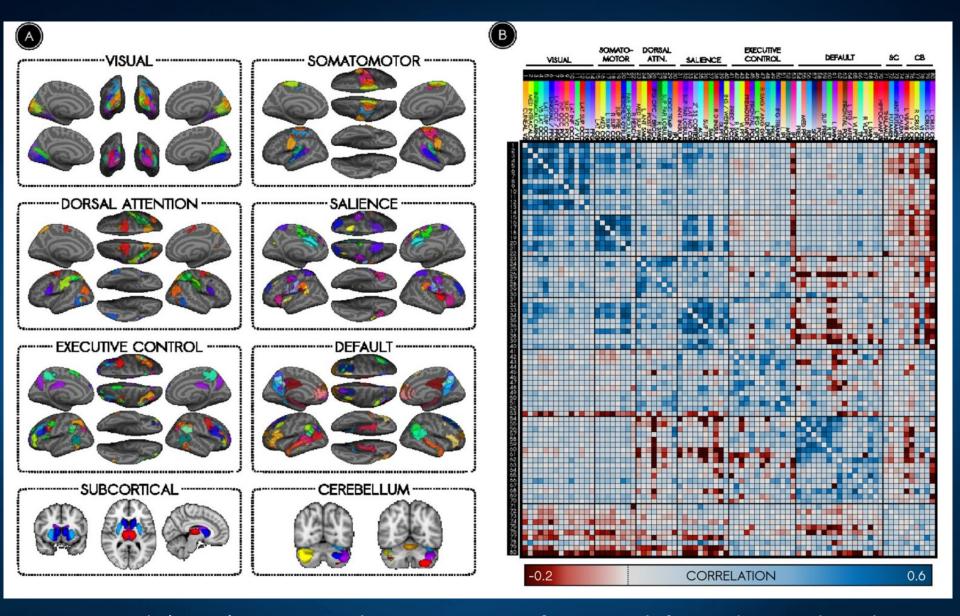
MCI patients (ADNI2), positive and negative functional connections in one of the 5 states of the Deep Auto-Encoder (DAE) + HMM models derived from the rs-fMRI time series.

Connections |W|>0.65.

Accuracy 72.6% with a sensitivity of 70.6% and a specificity of 75%.

Suk et al. Neuroimage (2016)

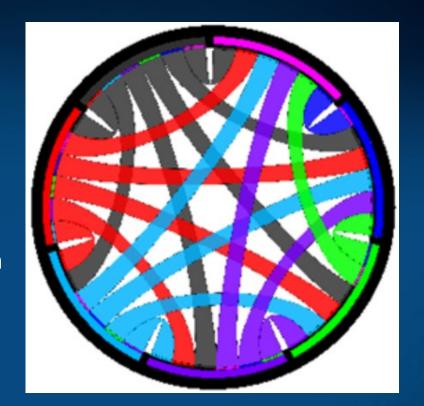


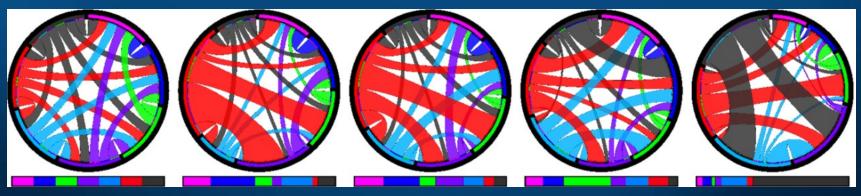


Ciric et.al. (2017). Contextual connectivity: A framework for understanding the intrinsic dynamic architecture of large-scale functional brain networks. *Scientific Reports* 7, 6537

### DMN time-averaged baseline.

Between-network allegiances (prob. that nodes are in the same community). Rim colors = canonical networks, rim length = greater allegiance to other networks, size of connections = strength of between-network allegiances. DMN1: weak within-network allegiance strong to DAT, SAL, and VIS.





VIS

SOM

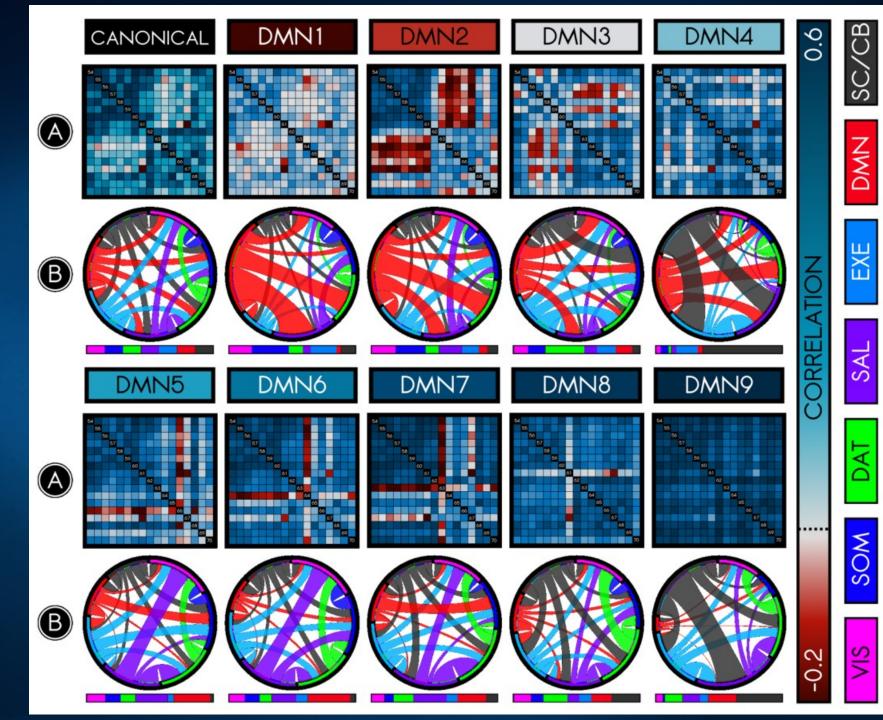
DAT

SAL

EXE

DMN

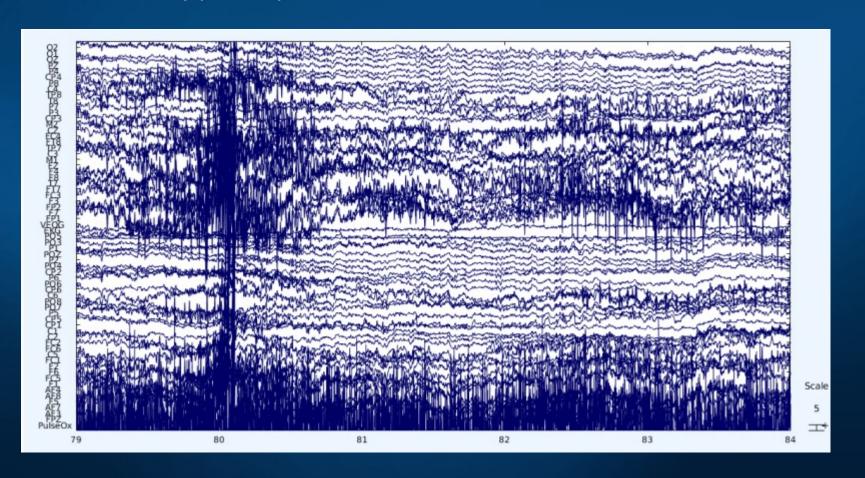
SC/CB



# Neurodynamics

### **EEG**

Brain networks from EEG? Technically difficult. Poor spatial resolution, only outer cortex, signals from the sensors are misleading, localization is necessery. Removal of artefacts is only partially automatic, it involves a lot of manual work.



### **EEG** for Brain Fingerprints

**fMRI** is too costly, difficult to standardize, to slow to follow dynamics. fMRI BFP are based on **V(X,t)** voxel intensity of fMRI BOLD signal changes, contrasted between task and reference activity or resting state. **EEG**: cheaper and better temporal resolution, use spatio-temporal maps, ERP maps/shapes, coherence, various phase synchronization indices for BFP.

- 1. Spatial/Power: direct localization/reconstruction of sources.
- **2. EEG microstates**, sequences & transitions, dynamics in ROI space.
- 3. Spatial/Synch: changes in functional graph network structure.
- **4. Frequency/Power**: ERS/ERD smoothed patterns E(X,t,f).
- 5. ERP global power maps: spatio-temporal averaged energy distributions.
- **6. EEG decomposition into components:** ICA, CCA, tensor, RP ...
- 7. Model-based: **The Virtual Brain**, integrating EEG/neuroimaging data.
- 8. Spectral fingerprinting (MEG, EEG), power distributions.

Neuroplastic changes of connectomes and functional connections are observed as a result of training to optimize brain processes.

### Model of reading & dyslexia

Emergent neural simulator:

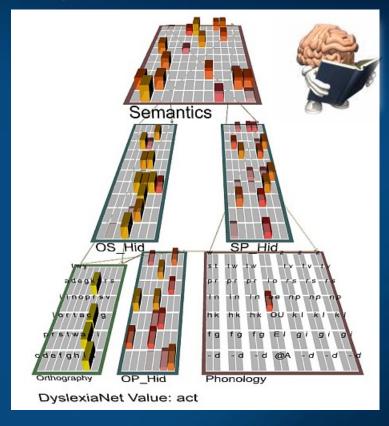
Aisa, B., Mingus, B., and O'Reilly, R. The emergent neural modeling system.

Neural Networks, 21, 1045, 2008.

3-layer model of reading:

orthography, phonology, semantics, or distribution of activity over **140 microfeatures** defining concepts.

In the brain: microfeature=subnetwork. Hidden layers OS/OP/SP\_Hid in between.



Learning: mapping one of the 3 layers to the other two.

Fluctuations around final configuration = attractors representing concepts.

How to see properties of their basins, their relations? Model in **Genesis**: more detailed neuron description.

### **Computational Models**

### Models at various level of detail.

 Minimal model includes neurons with 3 types of ion channels.

#### Models of attention:

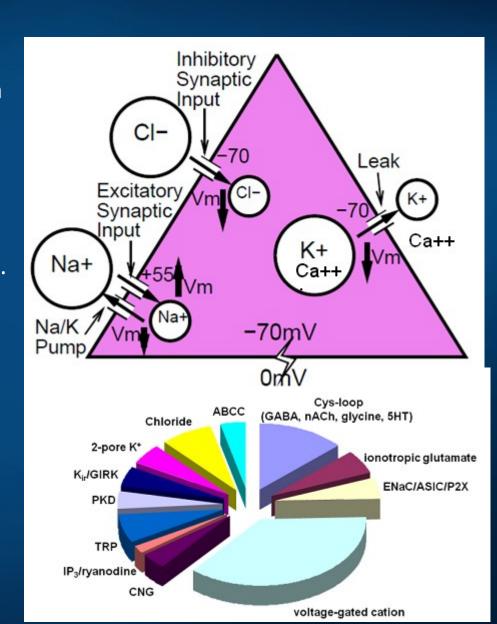
- Posner spatial attention;
- attention shift between visual objects.

#### Models of word associations:

sequence of spontaneous thoughts.

### Models of motor control.

Critical: control of the increase in intracellular calcium, which builds up slowly as a function of activation.
Initial focus on the leak channels,
2-pore K+, looking for genes/proteins.



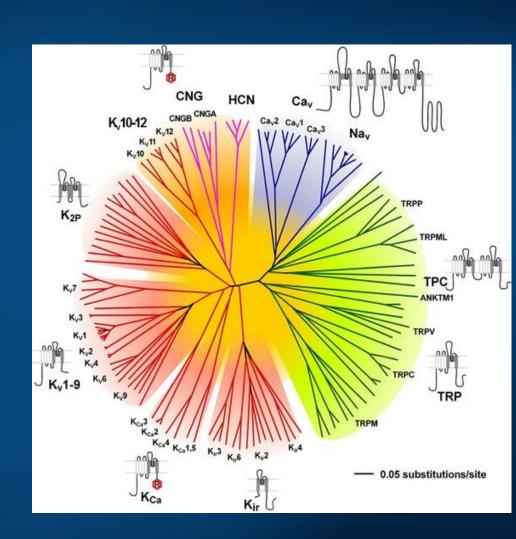
### Ion channels

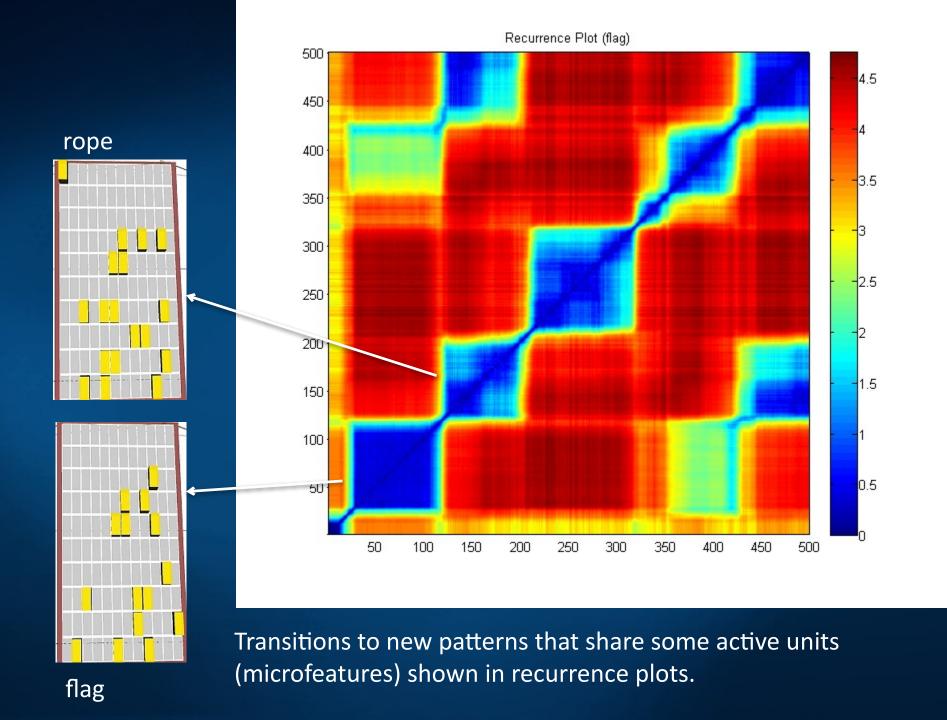
Hundreds of ion channels have been identified in neurons ...

Major challenge for computational neurosciences:

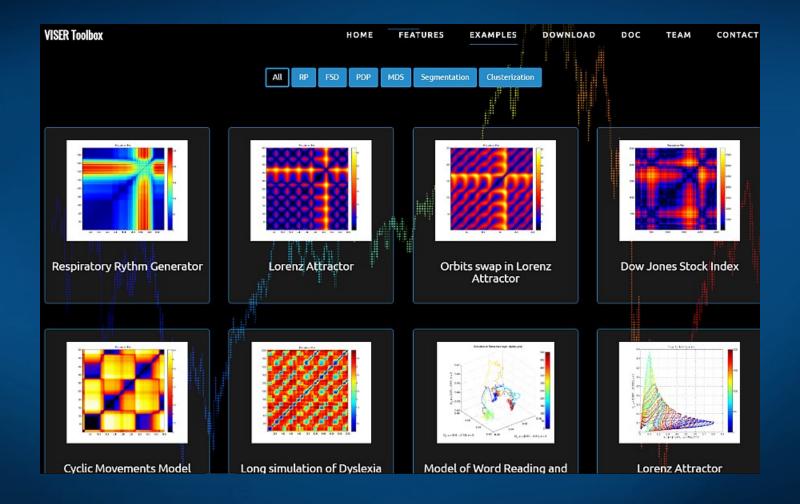
what happens with the nervous system when some of them are dysfunctional?

Leak channels regulate spontaneous transitions between attractor states.



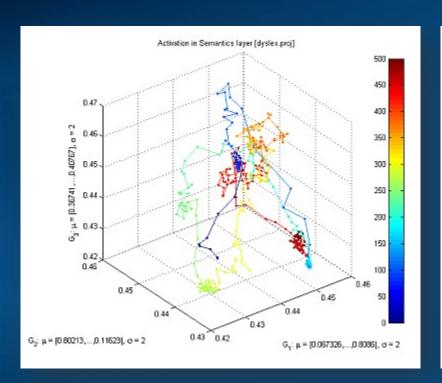


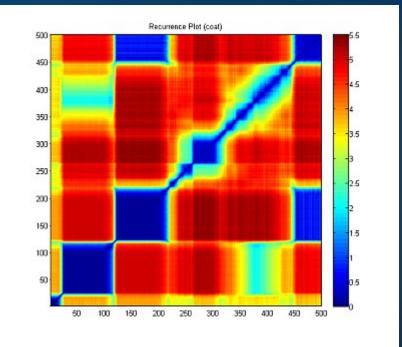
### Viser toolbox



Nasz <u>Viser toolbox</u> (Dobosz, Duch) do wizualizacji szeregów czasowych w wielu wymiarach różnymi technikami.

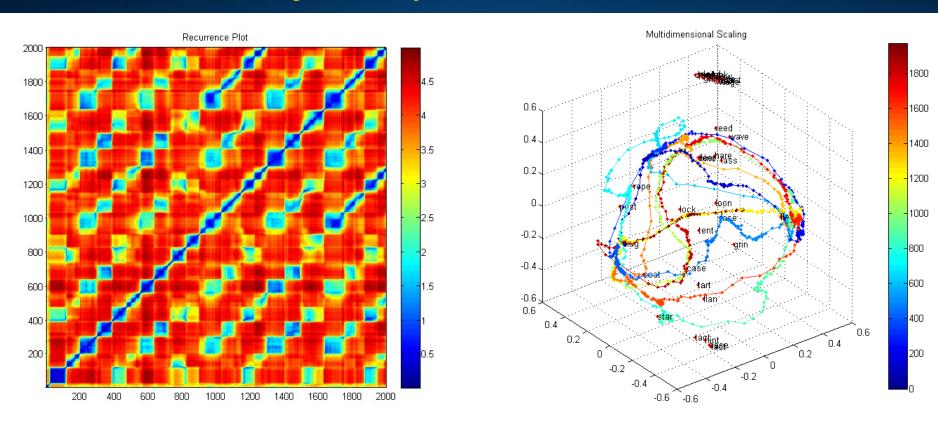
### **Fast transitions**





Attention is focused only for a brief time and than moved to the next attractor basin, some basins are visited for such a short time that no action may follow, corresponding to the feeling of confusion and not being conscious of fleeting thoughts.

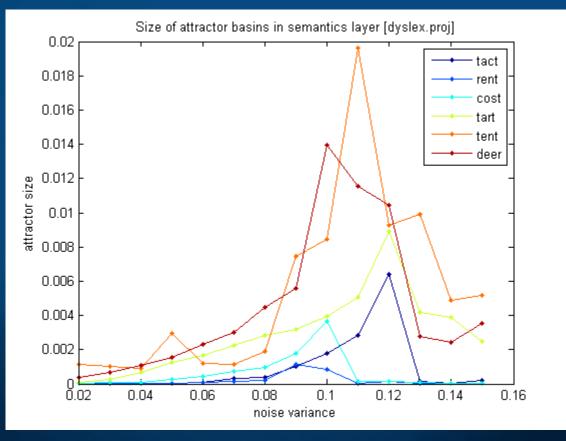
## Trajectory visualization



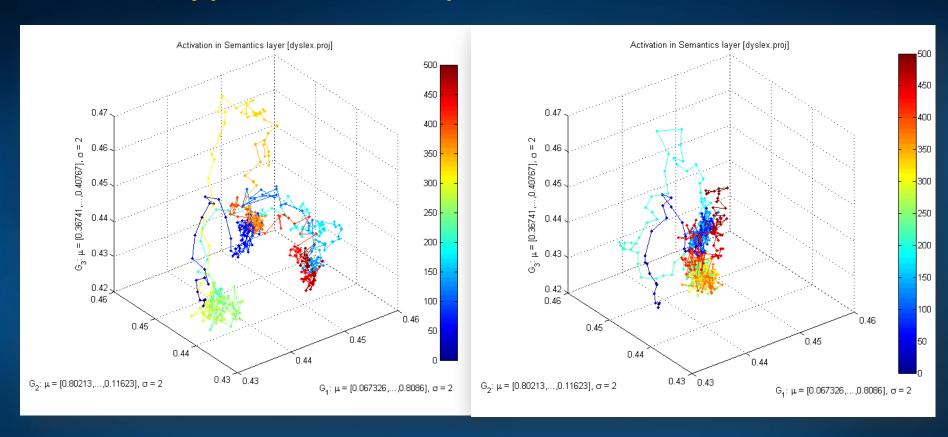
Recurrence plots and MDS/FSD/SNE visualization of trajectories of the brain activity. Here data from 140-dim semantic layer activity during spontaneous associations in the 40-words microdomain, starting with the word "flag".

### Depth of attractor basins

Variance around the center of a cluster grows with synaptic noise; for narrow and deep attractors it will grow slowly, but for wide basins it will grow fast. It may be used to estimate how strong states are entrapped in basins of attractors. Jumping out of the attractor basin reduces the variance due to inhibition of desynchronized neurons.

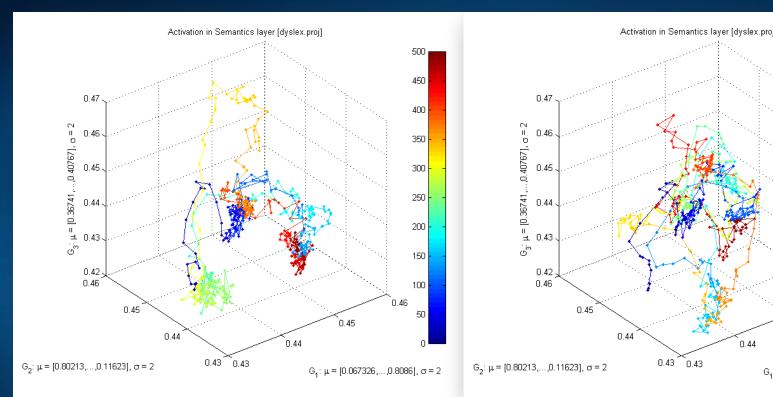


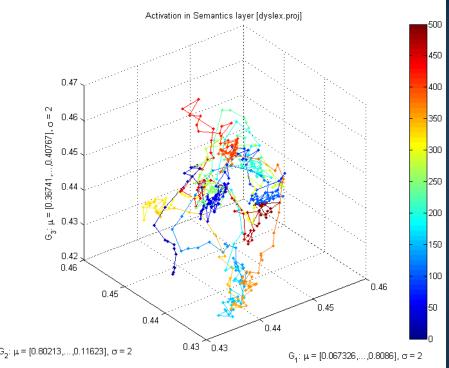
### Typical Development vs. Autism



All plots for the flag word, different values of b\_inc\_dt parameter in the accommodation mechanism. b\_inc\_dt = 0.01 & b\_inc\_dt = 0.005
b\_inc\_dt = time constant for increases in intracellular calcium building up slowly as a function of activation, controls voltage-dependent leak channels. http://kdobosz.wikidot.com/dyslexia-accommodation-parameters

# Typical Development vs ADHD



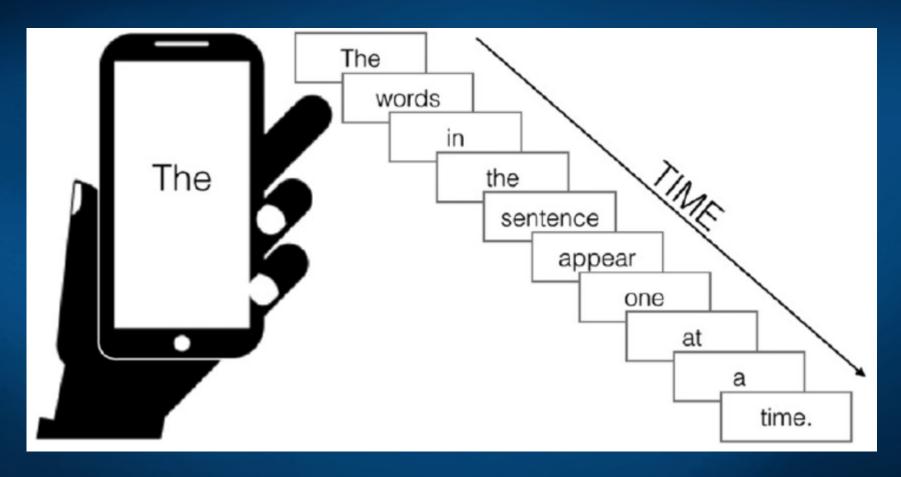


All plots for the flag word, different values of b\_inc\_dt parameter in the accommodation mechanism. b\_inc\_dt = 0.01 & b\_inc\_dt = 0.02.

b\_inc\_dt = time constant for increases in intracellular calcium which builds up slowly as a function of activation.

http://kdobosz.wikidot.com/dyslexia-accommodation-parameters

# Rapid Serial Visual Presentation

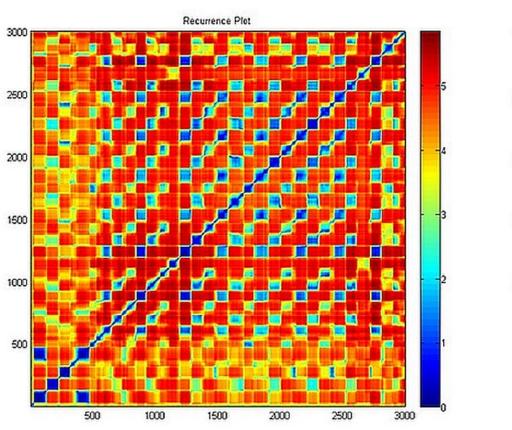


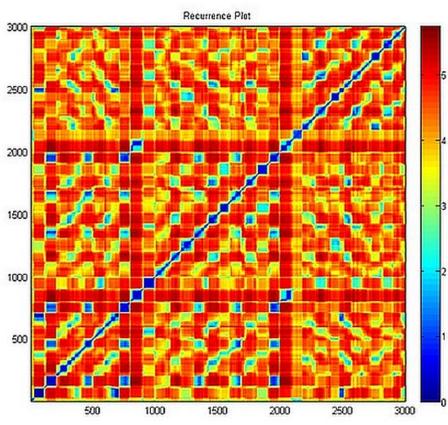
Any RSVP applications for fast reading.

Simulation: showing series of words, looking for attention/associations.

star => flea => tent => lock => tart => hind

# RSVP: typical brain

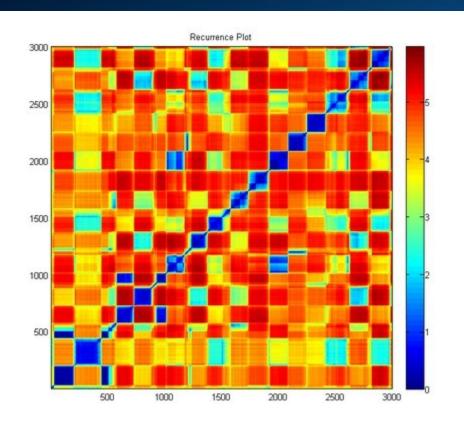


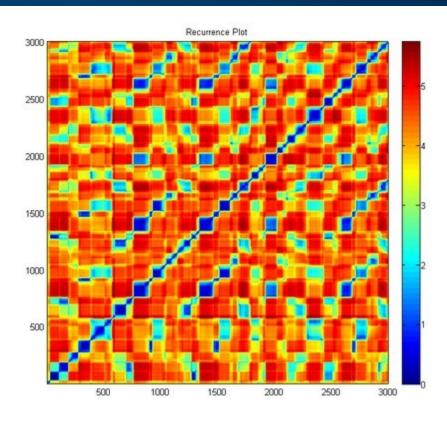


Normal speed associations, context=>understanding Some shallow microstates, no associations

too fast, speed 5x microstates get blurred, few associations

# **RSVP** simulations: HFA





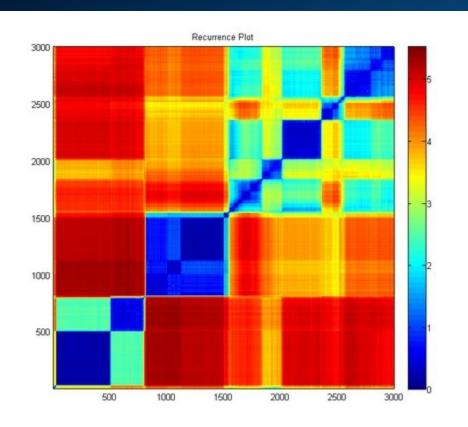
normal presentation long dwelling times

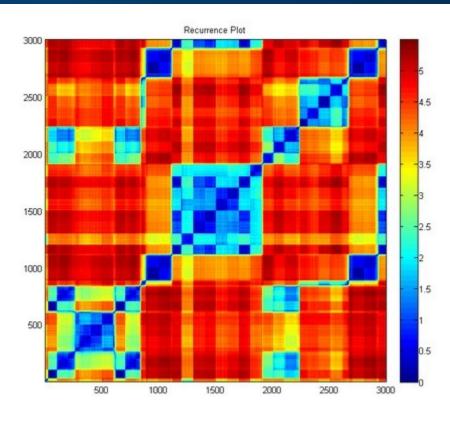
High functioning ASD case (HFA):

fast presentation

enforced quick resynchronization
more internal stimuli.

# RSVP simulations in deep autism

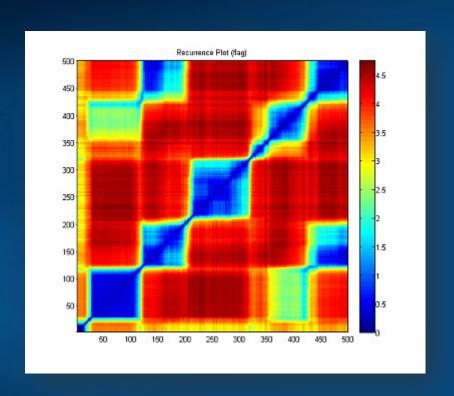


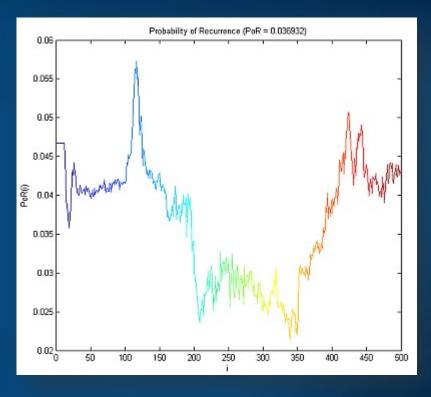


Normal speed skipping some words, no associations

fast presentation more internal states some associations arise

# Probability of recurrence





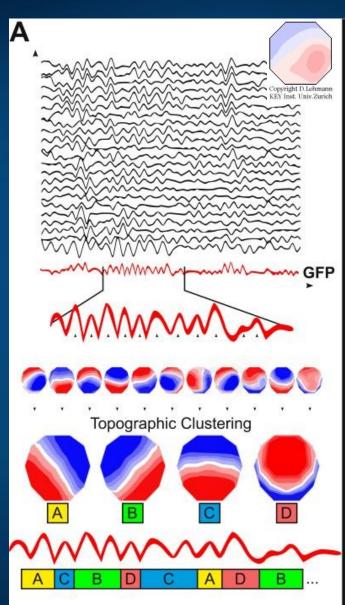
Probability of recurrence may be computed from recurrence plots, or from clusterization of trajectory points, allowing for evaluation how strongly some basins of attractors capture neurodynamics.

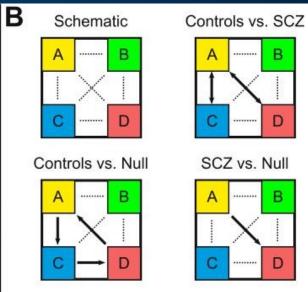
<u>Our Viser Toolbox</u> is used for all visualizations

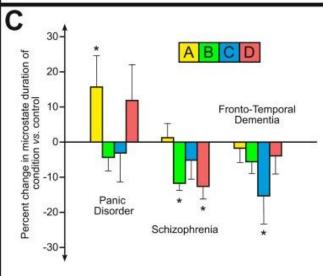
# Microstates in sensor space

Lehmann et al.
EEG microstate
duration and syntax
in acute, medicationnaïve, first-episode
schizophrenia.
Psychiatry Research
Neuroimaging, 2005

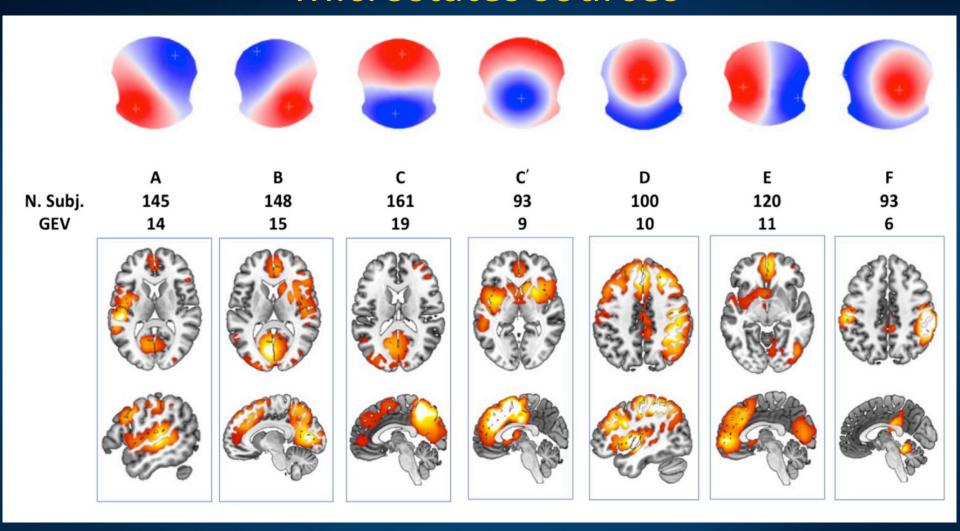
Khanna et al.
Microstates in
Resting-State EEG.
Neuroscience and
Biobehavioral
Reviews, 2015
4-7 states 60-150 ms
Symbolic dynamics.







## Microstates sources

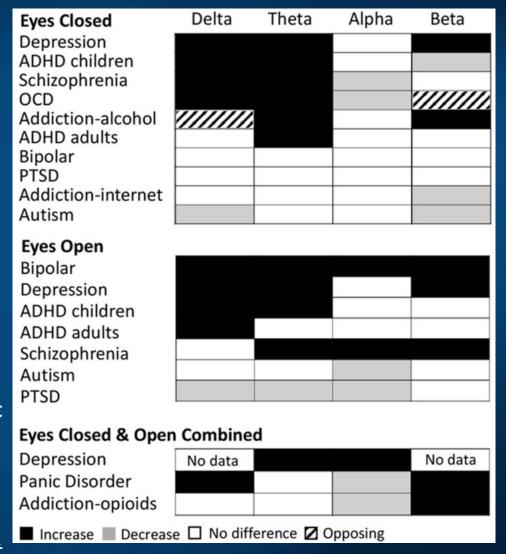


Michel, C. M., & Koenig, T. (2018). EEG microstates as a tool for studying the temporal dynamics of whole-brain neuronal networks: A review. *NeuroImage*, 180, 577–593. <a href="https://doi.org/10.1016/j.neuroimage.2017.11.062">https://doi.org/10.1016/j.neuroimage.2017.11.062</a>

# EEG bands and brain disorders

Differences in absolute power for each disorder (relative to control) for eyes closed condition (top), eyes open (middle) and eyes open and closed combined (bottom). White boxes indicate no change, black indicates an increase, and gray indicates a decrease. Hashed boxes - opposing results (contradictory).

Newson & Thiagarajan (2019).
EEG Frequency Bands in Psychiatric Disorders: A Review of Resting State Studies. Frontiers in Human Neuroscience, 12. <a href="https://doi.org/10.3389/fnhum.2018.0052">https://doi.org/10.3389/fnhum.2018.0052</a>



# Checkerboard reversal, 5 microstates

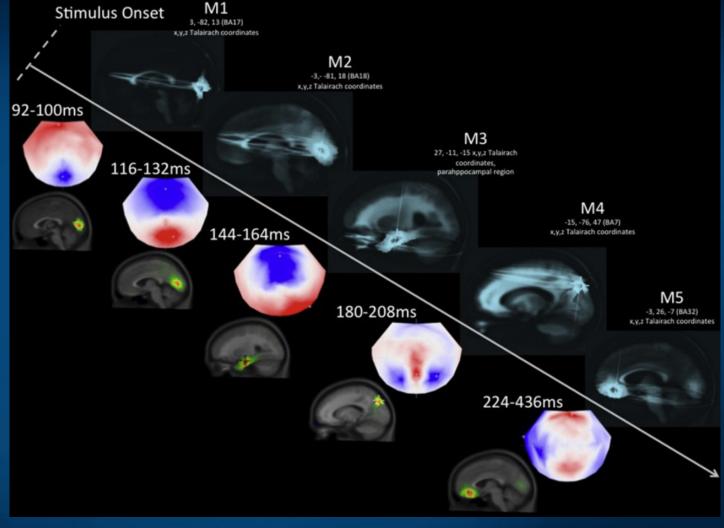
M1 => V1

M2 => V2

M3=>Parahippocampal

M4=>BA7, left PC, precuneus

M5=>dACC

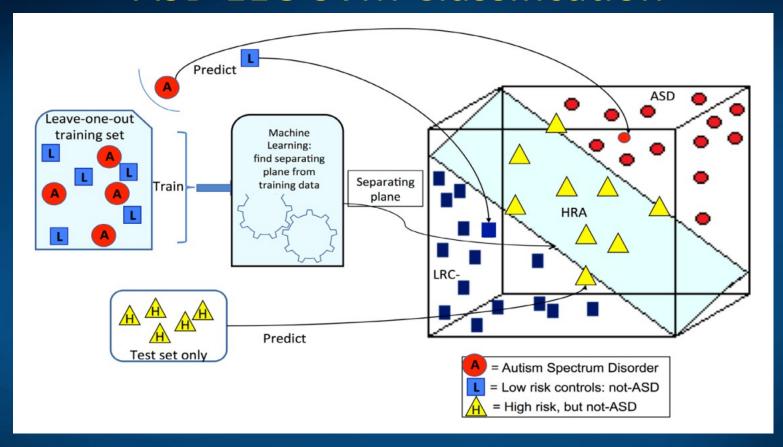


Cacioppo, S., Weiss, R. M., Runesha, H. B., & Cacioppo, J. T. (2014). Dynamic spatiotemporal brain analyses using high performance electrical neuroimaging: Theoretical framework and validation. *J. of Neuroscience Methods*, *238*, 11–34.

# Plan for action: 8-fold way.

- 1. Focus on neurodynamics. Include ion channels and other biophysical parameters for neurons/networks in your models.
- 2. Create simulation of normal functions, ex: attention shifts.
- 3. Catalogue all possible changes in biophysical parameters that lead to specific deregulation of normal behavior, ex: all types of ion channels.
- 4. Look for dysfunctional proteins related to biophysical parameters, ex: those proteins that build ion channels.
- 5. Use gene expression atlases to find correlations of proteins with mutations. Explain diversity of mutations and weak disease signals.
- 6. Predict changes in real brain signals: EEG/MEG, neuroimaging, intracranial.
- 7. Analyze existing neuroimaging data, functional and anatomical. Perform new experiments to verify proposed mechanisms leading to dysfunctions.
- 8. Propose close-loop therapies. Psychosomatic pain is a good target.

# **ASD EEG SVM Classification**



Wavelet decomposition, Recurrent Quantification Analysis, feature ranking and machine learning. Nonlinear features are critical to achieve good results, and their correlated with ASD depends on age.

# **EEG** early ASD detection

Bosl, W. J., Tager-Flusberg, H., & Nelson, C. A. (2018). EEG Analytics for Early Detection of Autism Spectrum Disorder: A data-driven approach. Scientific Reports, 8(1), 6828.

EEG of 3 to 36-month old babies, 19 electrodes selected from 64 or 128.

Daubechies (DB4) wavelets transform EEG signal into 6 bands.

7 features from Recurrence Quantitative Analysis (RQA): RP entropy, recurrence rate, laminarity, repetition, max/mean line length, trapping time.

In addition sample entropy and Detrended Fluctuation Analysis was used.

Nonlinear features were computed from EEG signals and used as input to statistical learning methods. Prediction of the clinical diagnostic outcome of ASD or not ASD was highly accurate.

SVM classification with 9 features gave high specificity and sensitivity, exceeding 95% at some ages. Prediction using only EEG data taken as early as 3 months of age was strongly correlated with the actual measured scores.

# EEG non-linear features

Features: not only structure, but also dynamics.

Nonlinear invariant measures of a time series and their physical interpretation, <u>recurrence quantification analysis</u> (RQA).

#### For example:

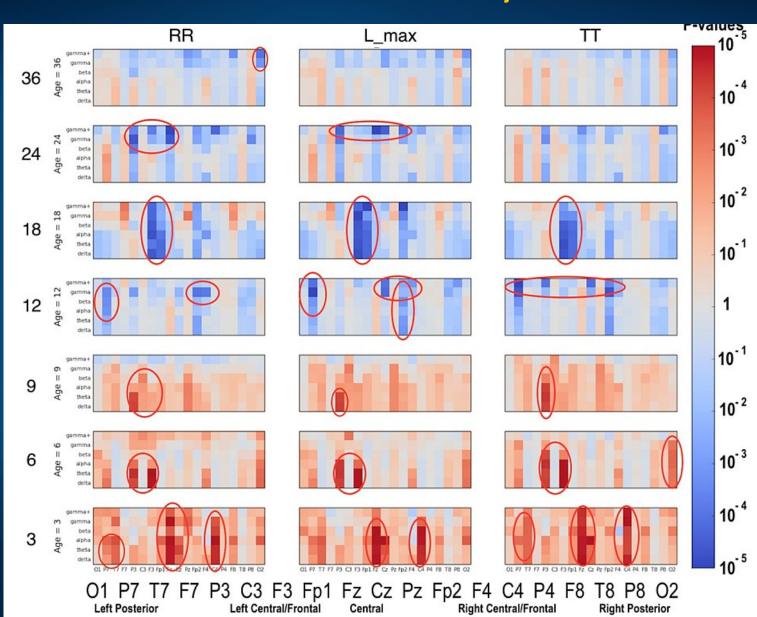
- Sample Entropy (SampE)
- 2. Entropy derived from recurrence plot (L\_entr).
- 3. Recurrence rate (RR), probability of recurrence.
- 4. Determinism (DET), repeating patterns in the system.
- 5. Laminarity (LAM), frequency of transitions between states.
- 6. Trapping time (TT), time in a given state.

# ASD vs Low Risk Healthy

RR = recurrence rate

L\_max = max line length, related to Lyapunov exponent

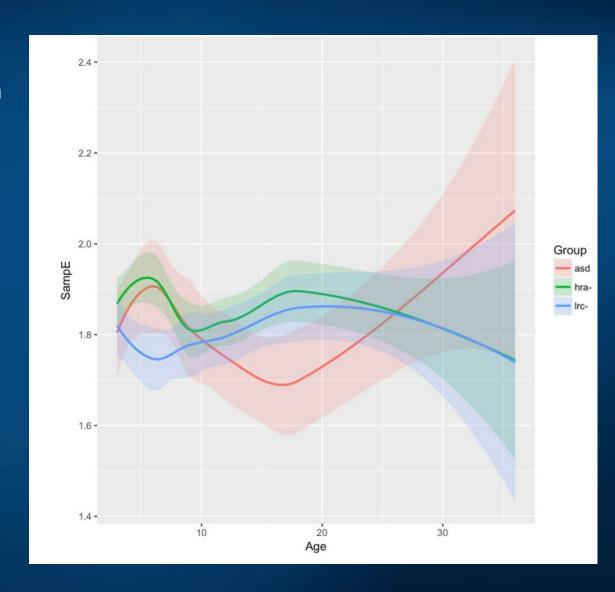
TT = trapping time



# **ASD EEG SVM Classification**

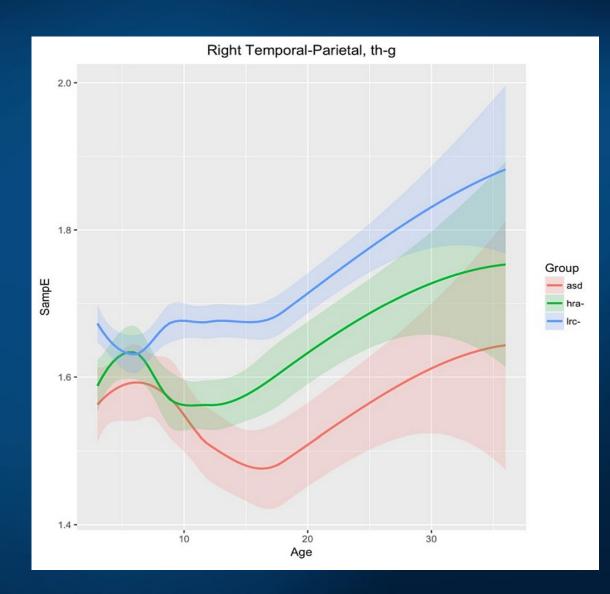
Developmental trajectories for SampE in the left temporal region (T7 sensor) in higher frequencies (beta+gamma) for ASD, LRC-, and HRA-

LRC low risk controls
HRA high risk for ASD
- no ASD



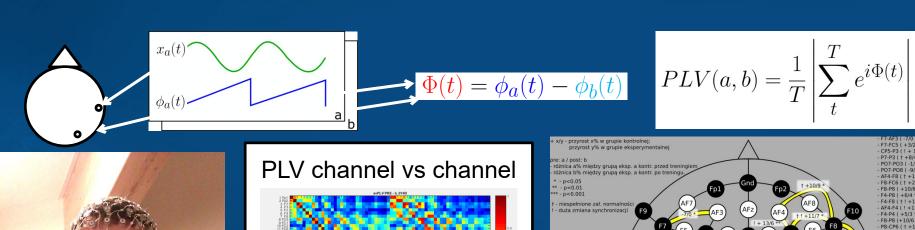
# **ASD EEG SVM Classification**

Developmental trajectories for SampE in the right temporal-parietal region (T8 +P4+P8 sensors) in frequencies theta through gamma for ASD, LRC-, and HRA-.

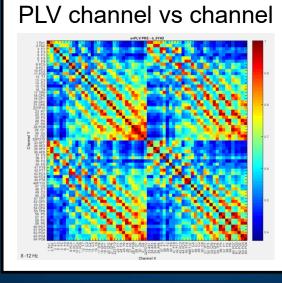


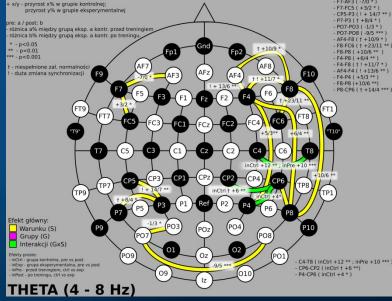
# Functional connectivity changes

Influence of brain games on functional connectivity: **Phase Locking Value** (Burgess, 2013; Lachaux 1999), phase differences between signals measured at each electrode. PLV => synchronization maps, info flow.



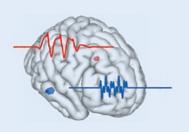






## EEG localization and reconstruction

**ECD** 



$$\widehat{d_j} = \operatorname{argmin} \parallel \phi - \sum_j \mathcal{K}_j d_j \parallel_{\mathcal{F}}^2$$

#### **Rotating dipole**

- Moving
- Rotating
- Fixed

He et al. Rev. Biomed Eng (2018)<sub>Sparse and Bayesian framework</sub>



$$\begin{split} \hat{\jmath} &= \underset{j}{\operatorname{argmin}} \parallel \mathcal{V} j \parallel_{1} + \alpha \parallel j \parallel_{1} \\ \text{S.T.} \parallel \phi - \mathcal{K} j \parallel_{\Sigma^{-1}}^{2} &\leq \varepsilon^{2} \end{split}$$

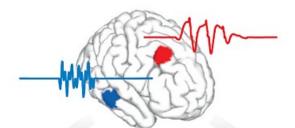
**IRES** 

Dipole model



#### Distributed model





#### MN ( $\ell_2$ ) family



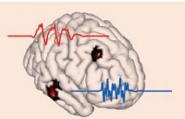
$$\begin{aligned} \hat{\mathbf{j}} &= \underset{j}{\operatorname{argmin}} \parallel \boldsymbol{\phi} - \mathcal{K} \boldsymbol{j} \parallel_{2}^{2} + \lambda \parallel \boldsymbol{j} \parallel_{2}^{2} \\ \hat{\mathbf{j}} &= \mathcal{T} \boldsymbol{\phi} = \mathcal{K}^{\mathsf{T}} \left( \mathcal{K} \mathcal{K}^{\mathsf{T}} + \lambda \boldsymbol{I} \right)^{\mathsf{T}} \boldsymbol{\phi} \end{aligned}$$

MN

- MN
   LORETA
- WMN



#### Nonlinear post hoc normalization



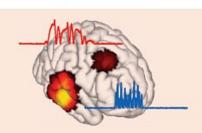
Beamforming and

scanning algorithms

$$\widehat{\boldsymbol{w}}_r = \underset{\boldsymbol{w}_r}{\operatorname{argmin}} \ \boldsymbol{w}_r^{\mathsf{T}} \boldsymbol{\mathcal{R}}_{\boldsymbol{\phi}} \boldsymbol{w}_r^{\mathsf{T}}$$

S.T. 
$$\begin{cases} \mathcal{K}_r^{\mathsf{T}} w_r = \xi_1 \\ w_r^{\mathsf{T}} w_r = 1 \end{cases}; \hat{j} = w^{\mathsf{T}} \phi$$

Beamformer (VBB)



$$\begin{split} \boldsymbol{\hat{f}}_{mn} &= \boldsymbol{T}_{mn} \boldsymbol{\phi} \\ \boldsymbol{\mathcal{S}}_{\boldsymbol{\hat{f}}} &= \boldsymbol{\mathcal{K}}^{\top} (\boldsymbol{\mathcal{K}} \boldsymbol{\mathcal{K}}^{\top} + \alpha \boldsymbol{I})^{\dagger} \boldsymbol{\mathcal{K}} \\ \boldsymbol{\hat{f}}_{sL} &= \boldsymbol{\hat{f}}_{mn} (\boldsymbol{\ell})^{\top} \left( [\boldsymbol{\mathcal{S}} \boldsymbol{\hat{f}}]_{\boldsymbol{\ell} \boldsymbol{\ell}} \right)^{-1} \boldsymbol{\hat{f}}_{mn} (\boldsymbol{\ell}) \\ &\text{sLORETA} \end{split}$$

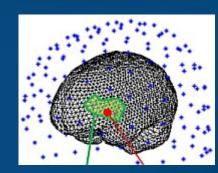
# Spatial filters

LCMV (Linearly Constrained Minimum Variance), classical reconstruction filter is a solution to the following problem:

$$=+, \approx, (\theta)\approx 1$$

LCMV has large error if:

- sources are correlated,
- SNR (signal to to noise ratio) is low, or
- forward problem is ill-conditioned.



Minimum variance pseudo-unbiased reduced-rank (MV-PURE, Piotrowski, Yamada, IEEE Transactions on Signal Processing **56**, 3408-3423, 2008)

$$W = \bigcap_{j \in \Upsilon} \underset{\hat{W} \in X_r}{\operatorname{arg \, min}} \left\| \hat{W}K(\theta) - I_l \right\|_{j}^{2}$$

where  $X_r$  is a set of all matrices of rank at most r, and set Y denotes all unitary norms. We use 15000 vertex FreeSurfer brain tessellation together with brain atlases that provide parcellation of the mesh elements into 100-240 cortical patches (regions of interest, ROIs).

# SupFunSim

SupFunSim: our library/Matlab /tollbox, direct models for EEG/MEG.

Provides many spatial filters for reconstruction of EEG sources: linearly constrained minimum-variance (LCMV), eigenspace LCMV, nulling (NL), minimum-variance pseudo-unbiased reduced-rank (MV-PURE) ...

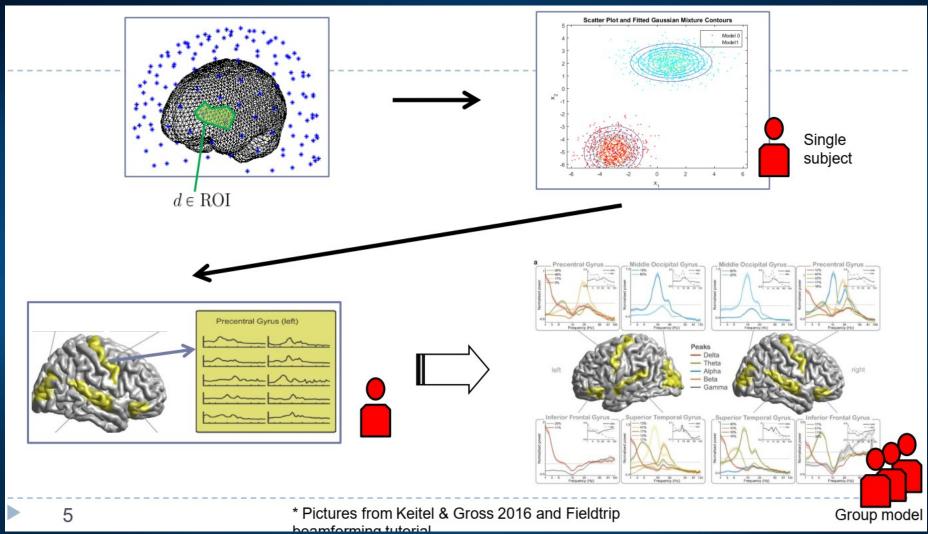
Source-level directed connectivity analysis: partial directed coherence (PDC), directed transfer function (DTF) measures.

Works with FieldTrip EEG/ MEG software. Modular, object-oriented, using Jupyter notes, allowing for comments and equations in LaTex.

```
A := H_{Src,R} := R^{-1/2}H \tag{34} B := H_{Src,N} := N^{-1/2}H \tag{35} \begin{bmatrix} 1 & \text{%file calculate}\_H\_Src.m \\ \text{function model} = \text{calculate}\_H\_Src(\texttt{MODEL}) \\ \text{model} = \texttt{MODEL}; \\ \end{bmatrix} \begin{bmatrix} \text{model.H\_Src}\_R = \text{pinv}(\text{sqrtm}(\text{model.R})) * \text{model.H\_Src}; \\ \text{model.H\_Src}\_N = \text{pinv}(\text{sqrtm}(\text{model.N})) * \text{model.H\_Src}; \\ \end{bmatrix} \begin{bmatrix} \text{end} \end{bmatrix}
```

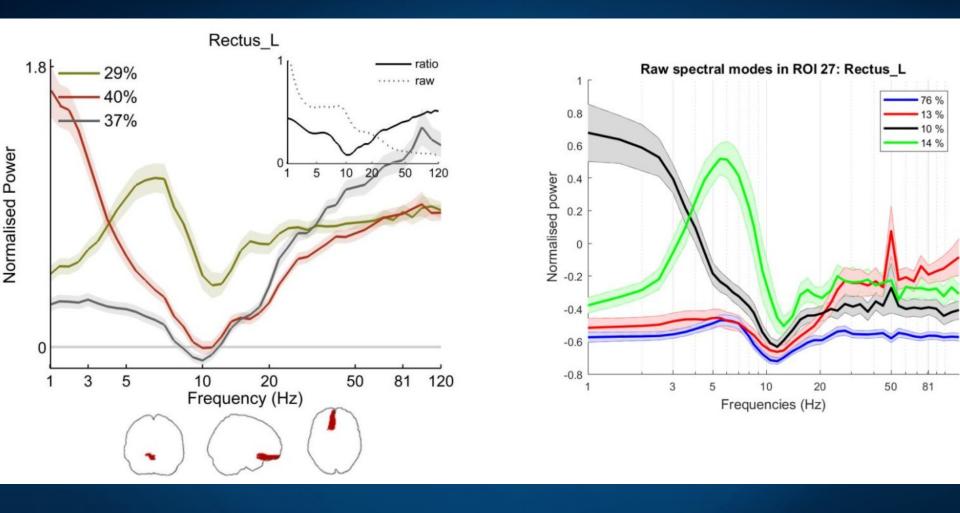
Rykaczewski, K., Nikadon, J., Duch, W., & Piotrowski, T. (2019). BioRxiv, 618694

# Spectral fingerprints

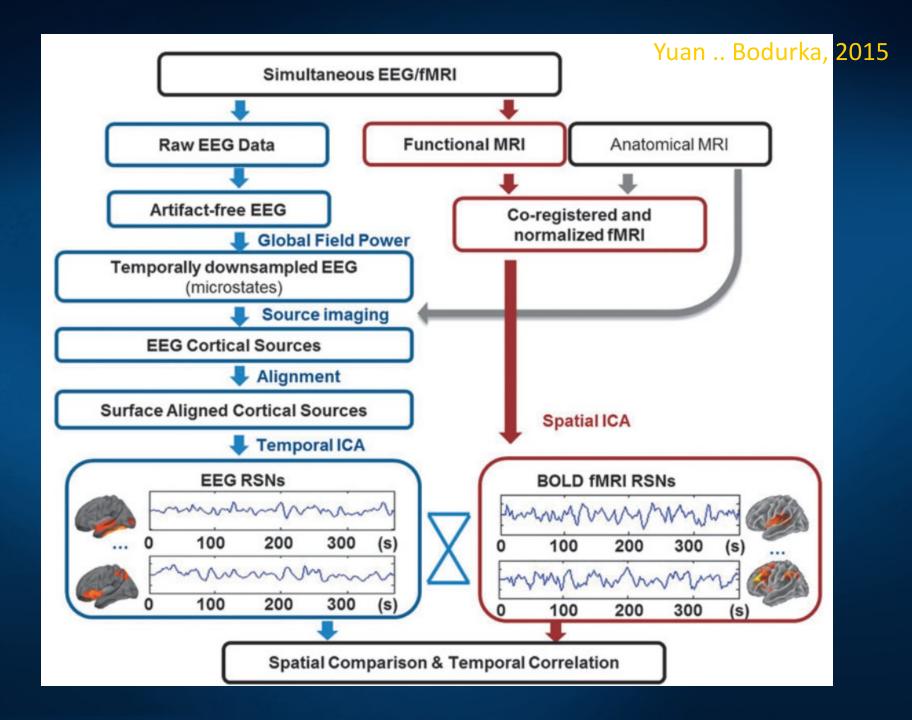


A. Keitel i J. Gross, "Individual human brain areas can be identified from their characteristic spectral activation fingerprints", *PLoS Biol* 14(6), e1002498, 2016

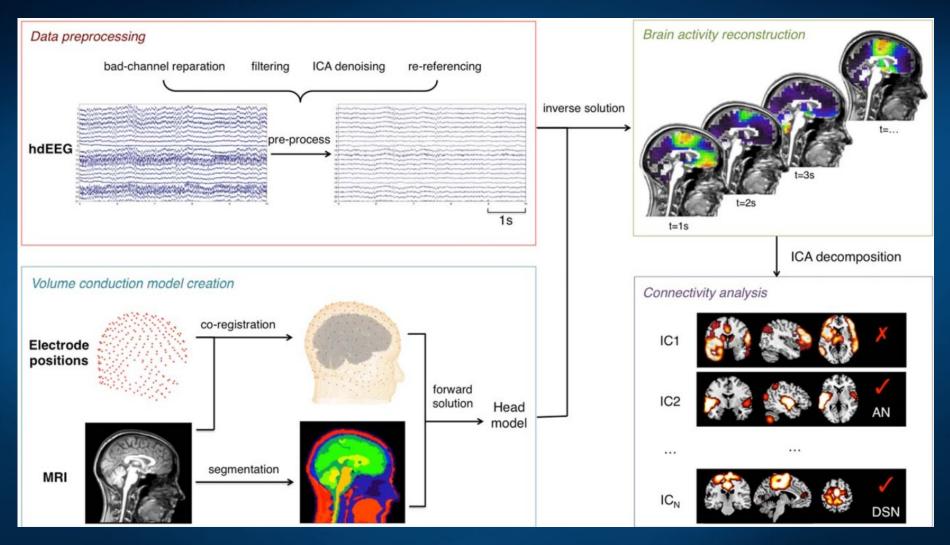
# Spectral fingerprints



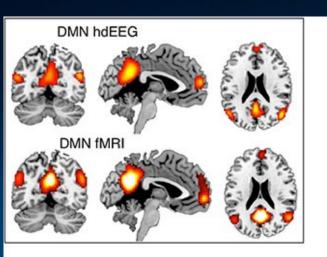
A. Keitel i J. Gross, "Individual human brain areas can be identified from their characteristic spectral activation fingerprints", *PLoS Biol* 14, e1002498, 2016

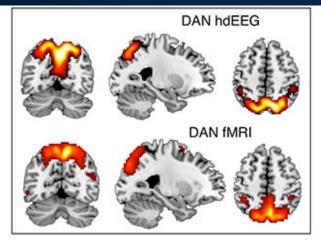


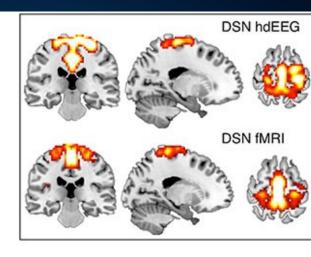
# 14 networks from BOLD-EEG

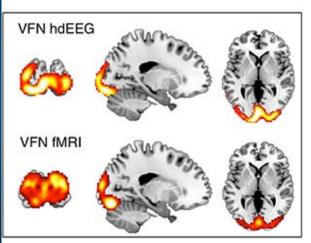


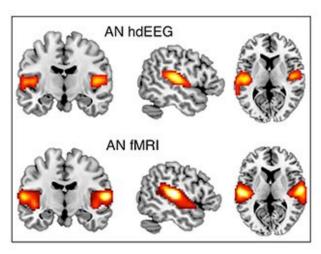
Liu et al. Detecting large-scale networks in the human brain. HBM (2017; 2018).

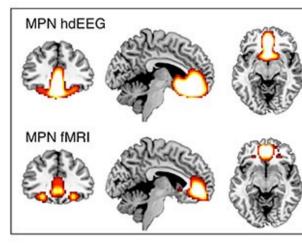








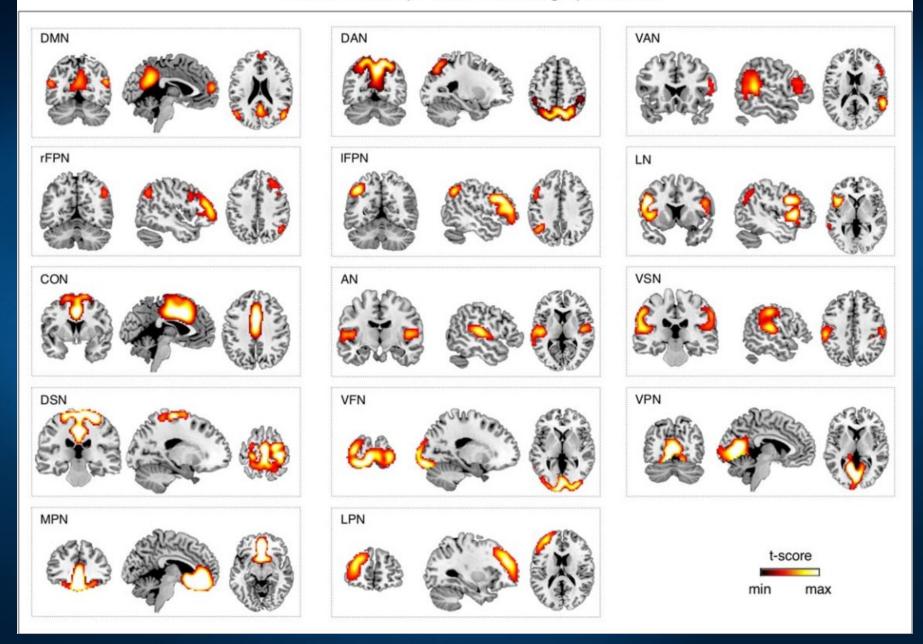




t-score min max

sICA on 10-min fMRI data (N = 24, threshold: p < 0.01, TFCE corrected). DMN, default mode network; DAN, dorsal attention network; DSN, dorsal somatomotor network; VFN, visual foveal network; AN, auditory network; MPN, medial prefrontal network.

#### EEG-RSN maps obtained using spatial ICA



# Plan for action - lessons from ASD

- 1. Focus on neurodynamics. Include ion channels and other biophysical parameters for neurons/networks.
- 2. Create simulation of normal functions, ex: attention shifts.
- 3. Catalogue all possible changes in biophysical parameters that lead to specific deregulation of normal behavior, ex: all types of ion channels.
- 4. Look for dysfunctional proteins related to biophysical parameters, ex: those proteins that build ion channels.
- 5. Use gene expression atlases to find correlations of proteins with mutations. Explain diversity of mutations and weak disease signals.
- 6. Predict changes in real brain signals: EEG/MEG, neuroimaging, intracranial.
- 7. Analyze existing neuroimaging data, functional and anatomical. Perform new experiments to verify proposed mechanisms leading to dysfunctions.
- 8. Propose close-loop therapies. Psychosomatic pain is a good target.

# Perspectives

- Many brain states are now linked to specific mental states, and can be transformed into signals that we can understand: motor intentions, plans, images, inner voices ...
- Some large-scale functional networks have reasonable (although still not perfect) interpretation, for example sensory networks, dorsal and ventral attention networks, executive control, motor networks.
- Individual differences and many psychological functions are directly linked to connectome and functional networks, including multistable properties.
- AI/ML draws inspirations from brain research, but also neural network models and learning algorithms (CNN, recurrence networks, reinforcement learning) help to interpret information processing in the brain.
- Many neurocognitive technologies are coming, helping to diagnose, repair and optimize brain processes.

# In search of the sources of brain's cognitive activity

Project "Symfonia", 2016-21











# My group of neuro-cog-fanatics





#### Join the global INCF community for

keynotes | panel discussions | posters | demos | socials

#### **Speakers**

Jan Bjaalie, University of Oslo
Rafal Bogacz, University of Oxford
Andrzej Cichocki, RIKEN CBS
Maureen Clerc, Inria
Carole Goble, University of Manchester
William Grisham, UCLA
Michael Hawrylycz, Allen Institute for Brain Science
Henry Kennedy, INSERM
Naomi Penfold, ASAPbio
Ariel Rokem, University of Washington
Frances Skinner, University of Toronto
Pedro Valdes-Sosa, Cuban Neuroscience Center,
University of Electronic Science and Technology China
Kirstie Whitaker, University of Cambridge

Alexander Woodward, RIKEN CBS

Jaroslaw Zygierewicz, University of Warsaw



#### **Session themes**

- Global brain projects: infrastructure interoperability and sustainability
- Data management and workflows in neuroscience
  - · Future of academic publishing
  - · Comparative and predictive connectomics
    - Brain Computer Interface (BCI)
- Neuroinformatics challenges in behavioral studies
  - · Building open science communities

# PP-RAI'2018

iii 18-19.10.2018 ♀ Poznań

Polskie Porozumienie na rzecz Rozwoju Sztucznej Inteligencji

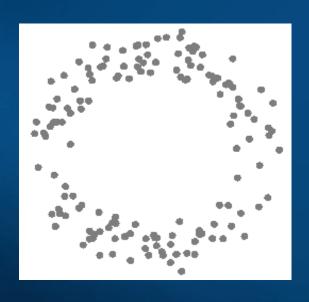
# PP-RAI'2019

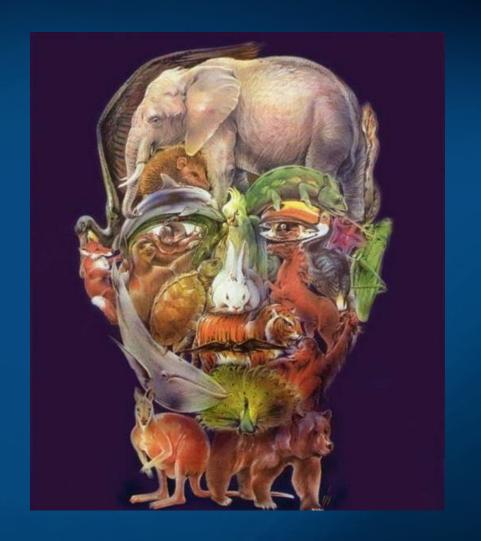
16-18.10.2019 — Wrocław

Polskie Porozumienie na rzecz Rozwoju Sztucznej Inteligencji

neuroinformatics2019.org

# Thank you for synchronization of your neurons





Google: W. Duch => talks, papers, lectures, Flipboard ...

