School on Nonlinear Dynamics, Complex Networks, Information Theory and Machine Learning in Neuroscience, 22-26 May 2023

### Nonlinear time series analysis

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Class 1: From dynamical systems to complex systems Class 2: Univariate time series analysis Class 3: Bivariate and multivariate analysis





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### Outline

- Uncovering similarities between lasers and neurons
- Analysis of regime transitions: EEG signals and vegetation fields
- Network based analysis of retina fundus images



## Data analysis methods allow to discover statistical similarities in very different systems



#### Data analysis method: ordinal analysis

$$\{\dots X_i, X_{i+1}, X_{i+2}, \dots\}$$

Possible order relations among three numbers (e.g., 2, 5, 7)



Bandt and Pompe: Phys. Rev. Lett. 2002

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#### The number of ordinal patterns increases as D!

1 \_ ~ 7 ~ 13 ~ 19 ~ ~ 2 8 14 20 3 / 9 / 15 / 21 / 4 10 16 22 5 11 17 23 6 12 18 24

A problem for short datasets.

U. Parlitz et al. / Computers in Biology and Medicine 42 (2012) 319-327

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#### Ordinal patterns can be defined using a lag

Example: climatological time series (monthly sampled)

- Consecutive months: [... $x_i(t), x_i(t+1), x_i(t+2)...$ ]
- Consecutive years: [... $x_i(t)$ ,... $x_i(t+12)$ ,... $x_i(t+24)$ ...]



Y. Zou et al. Phys. Rep. 787, 1 (2019)

From a time series, by counting the number of times the different patterns appear, we can calculate the set of "ordinal probabilities"





- A. Analyze the probabilities (are differences statistically significant?)
- B. Compute information theory measures (entropy, complexity)



Are the *D*! ordinal patterns equally probable?

#### Null hypothesis:

 $p_i = p = 1/D!$  for all  $i = 1 \dots D!$ 

• If at least one probability is not in the interval  $p \pm 3\sigma$  with  $\sigma = \sqrt{p(1-p)/N}$ 

and N the number of ordinal patterns:

We **reject** the NH with 99.74% confidence level.

Else, we fail to reject the NH with 99.74% confidence level.





# Example: chaotic time series generated with the Logistic map x(i+1) = r x(i)[1-x(i)] r=3.99



#### "Normal" and "Ordinal" bifurcation diagrams of the Logistic map



Pattern **210** is always forbidden; pattern **012** is more probable as r increases

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Ordinal analysis has been extensively used:

- To test if a model is good for the data
- To fit the parameters of a model
- To classify different types of data based on the probabilities of ordinal patterns

I. Leyva, J. M. Martinez, C. Masoller, O. A. Rosso, M. Zanin, "20 Years of Ordinal Patterns: Perspectives and Challenges", EPL 138, 31001 (2022).

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#### ECG signals: analysis of time series of inter-beat intervals





#### **Classifying ECG signals according to ordinal probabilities**



- Analysis of raw data (statistics of ordinal patterns is almost unaffected by a few extreme values)
- The probabilities are normalized with respect to the smallest and the largest value occurring in the data set.
   <u>U. Parlitz et al. Computers in Biology and Medicine 42, 319 (2012)</u>

## Sequence of inter-spike-intervals (ISIs) $\Rightarrow$ sequence of ordinal patterns



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# The analysis of the ordinal probabilities uncovers similarities in ISI sequences





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# Uncovering similarities between neurons and lasers... Interesting but relevant?

- Data centers, AI systems, HPC consume huge amounts of energy.
- Big concern in the context of climate change.
- The human brain processes huge amounts of information using only 19 Watts.
- Uncovering genuine similarities between neurons and lasers will allow to develop photonic neurons, able to process information as real neurons do, but
  - much faster,
  - with much less energy consumption.



European Centre for Medium-Range Weather Forecasts, Reading, UK





# Time series recorded in our lab show excitability, tonic spikes, and bursting. Similar to real neurons?



A. Aragoneses, S. Perrone, T. Sorrentino, M. C. Torrent and C. Masoller, "Unveiling the complex organization of recurrent patterns in spiking dynamical systems", Sci. Rep. 4, 4696 (2014).

C. Quintero-Quiroz, J. Tiana-Alsina, J. Roma, M. C. Torrent, and C. Masoller, "*Characterizing how complex optical signals emerge from noisy intensity fluctuations*", Sci. Rep. **6** 37510 (2016).

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### Main challenge







Understand how to mimic with lasers the way neurons encode and process information.



J. A. Reinoso, M. C. Torrent, and C. Masoller, "*Emergence of spike correlations in periodically forced excitable systems*", Phys. Rev. E. 94, 032218 (2016).

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### Single-neuron vs ensemble encoding

- Single-neuron encoding: slow because long spike sequences are needed to estimate the ordinal probabilities.
- Ensemble encoding: can be fast because, from the ISI sequences of all the neurons, few spikes per neuron can be enough to accurately estimate the probabilities.

$$\epsilon \dot{u}_{i} = u_{i} - \frac{u_{i}^{3}}{3} - v_{i} + a_{0} \cos(2\pi t/T) + \frac{\sigma}{k_{i}} \sum_{j}^{N} a_{ij}(u_{j} - u_{i}) + \sqrt{2D}\xi_{i}(t), \qquad i \neq j$$
  
$$\dot{v}_{i} = u_{i} + a.$$
  
$$k_{i} = \sum_{j} a_{ij}$$

M. Masoliver and C. Masoller, "*Neuronal coupling benefits the encoding of weak periodic signals in symbolic spike patterns*", Commun. Nonlinear Sci. Numer. Simulat. 88, 105023 (2020).

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## Ensemble encoding of a weak sinusoidal signal in the frequencies of occurrence of ordinal patterns



M. Masoliver and C. Masoller, Commun. Nonlinear Sci. Numer. Simulat. 88, 105023 (2020).

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# Laser-neuron comparison: encoding a weak periodic signal using spike rate code



Experiments modulating the laser current

Neuron model with the same input signal





 $\varepsilon \frac{dx}{dt} = x - \frac{x^3}{3} - y,$  $\frac{dy}{dt} = x + a + D\xi(t).$ 

J. Tiana-Alsina, C. Quintero-Quiroz and C. Masoller, "*Comparing the dynamics of periodically forced lasers and neurons*", New J. of Phys. 21, 103039 (2019) (2019). J. Tiana-Alsina, C. Masoller, "*Time crystal dynamics in a weakly modulated stochastic time delayed system*", Sci. Rep. 12, 4914 (2022).

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#### How about the temporal code?

### Ordinal analysis uncovers differences in spike timing.

Pulsed signal Sinusoidal 2.2 Diode 1.8 laser with 1.6 optical 1.4 1.2 feedback 0.8 10 20 30 40 10 20 30 40 50 60 50 60 70 80 70 80 Modulation Frequency [MHz] 0.07 Modulation Amplitude 0.02 0.06 FitzHugh-Nagumo 0.05 model 0.04 0.03 0.2 0.4 0.6 0.8 1.2 0.2 0.4 0.6 0.8 1 1 Modulation Frequency [Arb. u.]

Most probable pattern in color code

201

Probability

012

NSS

210

201

120 M

Probabilit

012 NSS

1.2

J. Tiana-Alsina, C. Quintero-Quiroz and C. Masoller, New J. of Phys. 21, 103039 (2019).

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#### Software

Python and Matlab codes for computing the ordinal pattern **index** are available here: U. Parlitz et al. Computers in Biology and Medicine 42, 319 (2012)



World length (wl): 4 Lag = 3 (skip 2 points) Result:

indcs= 3

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function indcs = perm\_indices(ts, wl, lag); m = length(ts) - (wl - 1) \* lag;indcs = zeros(m,1);for i = 1:wl - 1; st = ts(1+(i-1)\*lag: m+(i-1)\*lag);for j = i:wl - 1; indcs = indcs + (st > ts(1 + j\*lag : m + j\*lag));end indcs = indcs \* (wl - i); end indcs=indcs+1:

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#### Hands on activity: Ordinal analysis of the logistic map

$$x(i+1) = r x(i)[1-x(i)]$$

- 1. Test the "ordinal pattern" program with some "hand made" examples (e.g., the sequence [5,2,7] returns index 3).
- 2. For the logistic map with r=3.99, calculate the probabilities of the 6 D=3 ordinal patterns and plot the distribution.
- 3. Calculate the ordinal bifurcation diagram with r in (3.5,4).



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#### Analysis of eyes-closed eyes-open transition in EEG recordings of healthy subjects.



#### Eyes open



#### TABLE I. Description of the datasets used.

	DTS1	DTS2
Sampling rate (Hz)	256	160
Time task (seg)	120	60
Total points	30720	9600
Number of electrodes	16	64
Number of subjects	71	109

#### DTS1: Britbrain (Zaragoza) DTS2: Physionet

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### Permutation entropy: Shannon's entropy computed from ordinal probabilities



### **PE partially distinguishes EC-EO states**



C. Quintero-Quiroz, L. Montesano, A. J. Pons, M. C. Torrent, J. García-Ojalvo, C. Masoller, "*Differentiating resting brain states using ordinal symbolic analysis*", Chaos 28, 106307 (2018).

B. R. R. Boaretto, R. C. Budzinski, K. L. Rossi, C. Masoller, E. E. N. Macau, "Spatial permutation entropy distinguishes resting brain states", Chaos, Solitons & Fractals 171, 113453 (2023).

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# The spatial permutation entropy can be an early indicator of vegetation transitions



G. Tirabassi and C. Masoller, "*Entropy-based early detection of critical transitions in spatial vegetation fields*", PNAS 120, e2215667120 (2022).

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#### Network-based analysis of retina fundus images

- For the diagnosis of eye diseases & follow up of treatments.
- Biometric identity identification.
- The retina is a window to the brain.
- Opportunity to detect other diseases: alterations in retina network may reflect alterations in other arterial systems.







Advanced Biomedical Optical Imaging and Data Analysis



H2020-675512

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#### Data and image analysis steps

- 45 high resolution images (3504 × 2336 pixels)
  15 healthy subjects
  15 glaucoma
  - 15 diabetic retinopathy
- For every subject we had:
  - -fundus photography

-<u>manual</u> segmentation done by an expert ophthalmologist.



Steps:

- 1. Pre-process and un-supervisely, segment the images.
- 2. Extract network.
- 3. Compare networks obtained from different images.
- 4. Classify the images.

https://www5.cs.fau.de/research/data/fundus-images/

#### **Step 1: Pre-process and segmentation**









We adapted an *unsupervised* algorithm, originally developed for segmenting images of **cultured neural networks**.

Manual segmentation



D. Santos-Sierra, I. Sendiña-Nadal, I. Leyva et al. Cytometry Part A. 87, 513 (2015).

P. Amil, F. Reyes-Manzano, L. Guzmán-Vargas, I. Sendiña-Nadal, C. Masoller, "*Network-based features for retinal fundus vessel structure analysis*", PLoS ONE 14, e0220132 (2019).

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Step 2: extract the network (identification of the optical nerve, nodes and links and assign weights to the links).



# Steps 3 and 4: Compare the networks extracted from different images and classify the images.

- {p<sub>i,j</sub>}: distances between probability distributions that characterize the networks obtained from images i and j.
- We used nonlinear dimensionality reduction (*Isomap*) to reduce the set of 45x45 {p<sub>i,i</sub>} values to only two features.

Distance distribution to the central node in the *manual* segmentation



P. Amil et al, Network-based features for retinal fundus vessel structure analysis, PLoS ONE 14 e0220132 (2019).

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#### Performance of network features in the manual segmentation

#### Distribution of weights along the shortest path to central node

# Distribution of weighted degrees



P. Amil et al, Network-based features for retinal fundus vessel structure analysis, PLoS ONE 14 e0220132 (2019).

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#### In the automated segmentation



P. Amil et al, Network-based features for retinal fundus vessel structure analysis, PLoS ONE 14 e0220132 (2019).

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#### Take home messages

- Data analysis techniques allow us to uncover similarities in very different systems.
- Different methods provide *complementary* information, and can be adapted to study very different systems.
- "Surrogate" tests are needed to determine if the numerical values are statistically significant.
- Holger Kantz: "Every data set bears its own difficulties: data analysis is never routine".

#### References

- A. Aragoneses et. al, "Unveiling the complex organization of recurrent patterns in spiking dynamical systems", Sci. Rep. 4, 4696 (2014).
- J. A. Reinoso et. al, "Emergence of spike correlations in periodically forced excitable systems", Phys. Rev. E. 94, 032218 (2016).
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- C. Quintero-Quiroz et. al, "*Differentiating resting brain states using ordinal symbolic analysis*", Chaos 28, 106307 (2018).
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