Fundamentals of Analyzing Biomedical Signals

Interactions

# Measuring Interactions

# from Time Series

# Information-theory-based techniques

information-theory

**basic idea:** interaction  $\Leftrightarrow$  information flow

- the "stronger" the information flow, the stronger the interaction
- information flow from one system to another indexes directionality

characterize information with Shannon entropy  $H = -\sum_i p_i \log p_i$ 

p is the (normalized) probability for an event / state / amplitude /... to occur

estimate probability with  $p_i = \lim_{N \to \infty} \frac{N_i}{N}$ where *N* is the total number of events / states / amplitudes /...

# measuring interactions strength of interaction:

# information-theory mutual information

given systems X and Y, the mutual information is defined as:

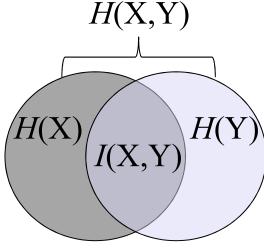
$$I(\mathbf{X}, \mathbf{Y}) = H(\mathbf{X}) + H(\mathbf{Y}) - H(\mathbf{X}, \mathbf{Y})$$

information generated by system X is characterized by the Shannon entropy:

$$H(\mathbf{X}) = -\sum_{i}^{N} p_{\mathbf{X}}(i) \log p_{\mathbf{X}}(i)$$

joint information is characterized by the Shannon entropy:

$$H(\mathbf{X}, \mathbf{Y}) = -\sum_{i,j}^{N} p_{\mathbf{X},\mathbf{Y}}(i,j) \log p_{\mathbf{X},\mathbf{Y}}(i,j)$$



# measuring interactions strength of interaction:

# information-theory relative entropy

relative entropy (also known as Kullback-Leibler divergence):

 $H_{\rm rel}(\mathbf{X}|\mathbf{Y}) = -\sum_{i}^{N} p_{\mathbf{X}}(i) \log \frac{p_{\mathbf{X}}(i)}{p_{\mathbf{Y}}(i)}$ 

- characterizes the similarity between the probability distributions.
- relative entropy is asymmetric:  $H_{rel}(X|Y) \neq H_{rel}(Y|X)$
- in general  $H_{\rm rel}$  is positive, and zero for identical systems

 $\rightarrow$  alternative definition of mutual information:

$$I_{\rm rel}(\mathbf{X}|\mathbf{Y}) = -\sum_{i,j}^{N} p_{\mathbf{X},\mathbf{Y}}(i,j) \log \frac{p_{\mathbf{X},\mathbf{Y}}(i,j)}{p_{\mathbf{X}}(i)p_{\mathbf{Y}}(j)}$$

characterizes *relative* difference between respective probability density distributions and the joint distribution density

S. Kullback, R.A. Leibler. On information and sufficiency. Ann. Math. Stat. 22, 79, 1951.

# measuring interactions strength of interaction:

# information-theory mutual information

properties of mutual information:

- symmetric: I(X, Y) = I(Y, X)
- I(X, Y) = 0 for independent (non-interacting) systems
- I(X, Y) = max for identical (fully synchronized) systems
- I(X, Y) increases monotonically with increasing coupling strength  $\rightarrow$  data-driven estimator for strength of interaction

disadvantages:

- only considers (single/joint) probability density distributions
- no information about dynamics
- can not explicitly distinguish between information exchange and joint information (e.g. due to common input or joint past)

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information-theory

mutual information

# measuring interactions strength of interaction:

extensions:

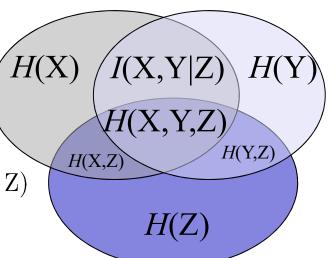
time-delayed mutual information (Kaneko, Physica D 23, 436, 1986)

#### partial (or conditional) mutual information

- (S. Frenzel & B. Pompe, PRL 99, 204101, 2007)
- part of mutual information of two random quantities that is not contained in a third one
- similar to partial correlation
- can also detect directionality\*

I(X, Y|Z) = H(X, Z) + H(Y, Z) - H(Z) - H(X, Y, Z)

\* K. Hlaváčková-Schindler, M. Paluš, M. Vejmelka & J. Bhattacharya. Causality detection based on information-theoretic approaches in time series analysis. Physics Reports, 441, 1-46, 2007



# measuring interactions *direction of interaction:*

# information-theory transfer entropy

- aim: characterize flow of information between systems X and Y
   idea: replace (static) probability density distributions by transition
   probability densities (cf. Entropies)
- given: time series *v*:  $v_1$ ,  $v_2$ , ...,  $v_N$  of some observable *x* and time series *w*:  $w_1$ ,  $w_2$ , ...,  $w_N$  of some observable *y*
- 1) incorporate time-dependence by relating previous samples  $v_i$  and  $w_i$  to predict the next value  $v_{i+1}$  (cf. N. Wiener),

2) consider generalized Markov condition (p = transition probability density):  $p(v_{i+1}|\mathbf{v}_i, \mathbf{w}_i) = p(v_{i+1}|\mathbf{v}_i)$ 

T. Schreiber, Measuring information transfer. Phys. Rev. Lett. 85, 461, 2000

# measuring interactions *direction of interaction:*

### information-theory transfer entropy

- 3) if systems X and Y independent  $\rightarrow$  Markov condition fulfilled
- 4) use relative entropy concept to quantify *incorrectness* of Markov condition; with this, transfer entropy is defined as:

$$T_{\mathrm{Y}\to\mathrm{X}} = \sum_{i}^{N} p\left(v_{i+1}, \mathbf{v}_{i}^{(k)}, \mathbf{w}_{i}^{(l)}\right) \log \frac{p\left(v_{i+1} | \mathbf{v}_{i}^{(k)}, \mathbf{w}_{i}^{(l)}\right)}{p\left(v_{i+1} | \mathbf{v}_{i}^{(k)}\right)}$$

(*l*,*k*) denote orders of Markov processes  $T_{X \rightarrow Y}$  defined in complete analogy

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transfer entropy

# measuring interactions *direction of interaction:*

properties of transfer entropy

- can detect direction of information flow since  $T_{Y \to X} \neq T_{X \to Y}$
- unbounded, needs suitable definition of directionality, e.g.

$$T := T_{\mathbf{Y} \to \mathbf{X}} - T_{\mathbf{X} \to \mathbf{Y}} \begin{cases} > 0 : \ \mathbf{Y} \text{ drives } \mathbf{X} \\ = 0 : \text{ no or symmetric bidir. coupling} \\ < 0 : \ \mathbf{X} \text{ drives } \mathbf{Y} \end{cases}$$

- depends on coupling strength → data-driven estimator for direction of interaction
- for Gaussian distributed data, transfer entropy equals Granger causality
- similar to conditional mutual information (replace system Z by e.g., past of system Y)

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measuring interactions *direction of interaction:* 

information-theory transfer entropy

Interactions

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extensions:

- multivariate (partial) transfer entropy
- various estimation techniques
- estimators for transient signals\* and delay-systems\*\*

\* H. Dickten, K. Lehnertz K. Identifying delayed directional couplings with symbolic transfer entropy. Phys Rev E 90, 062706, 2014 \*\* M. Martini, TA Kranz, T Wagner, K Lehnertz K. Inferring directional interactions from transient signals with symbolic transfer entropy. Phys Rev E 83, 011919, 2011

information-theory

#### measuring interactions strength and direction of interaction

how to estimate probability density distributions and the joint distribution densities from time series?

- counting (cumbersome)
- various binning techniques
- nearest neighbor estimators (e.g. Kozachenko-Leonenko)
- correlation sum (via phase-space embeddings)
- symbolization (e.g. based or permutation entropy\*)

### measuring interactions strength and direction of interaction

estimators based on the concept of symbolic dynamics and on symbolization

*symbolic dynamics:* modeling a smooth dynamical system by a *discrete space* consisting of infinite *sequences of symbols*, each of which corresponds to a state of the system, with the dynamics (evolution) given by the shift operator

symbolization: generate symbols via delay embedding

$$\mathbf{s}_i := (v_{i+(j_1-1)\tau}, v_{i+(j_2-1)\tau}, \dots, v_{i+(j_m-1)\tau})$$

where

$$v_{i+(j_1-1)\tau} \le v_{i+(j_2-1)\tau} \le \dots \le v_{i+(j_m-1)\tau}$$

 $\rightarrow$  symbol  $\hat{s}_i := (j_1, j_2, \dots, j_m)$ 

\* C. Bandt, B. Pompe. Permutation entropy: a natural complexity measure for time series. Phys. Rev. Lett., 88, 174102, 2002

an example

information-theory

#### strength and direction of interaction

embedded data: 
$$(3, 5, 9)$$
  $(10, 1, 6)$   
symbols  $\Rightarrow$   $(1, 2, 3)$   $(1, 3, 2)$ 

permutation entropy: 
$$H(m) = -\sum_{i=1}^{m!} \hat{s}_i \log \hat{s}_i$$
  
normalization:  $0 \le H = \frac{H(m)}{\log(m!)} \le 1$ 

 $H \rightarrow 0$  for deterministic systems,  $H \rightarrow 1$  for stochastic systems

Interactions

an example

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#### strength and direction of interaction

### information-theory

an example

given time series of systems X and Y:

- estimate permutation entropy from windowed data
- investigate changing tendency of permutation entropies

$$S(w_i) = \begin{cases} +1 : \text{ if } H(w_i) < H(w_{i+1}) \\ -1 : \text{ else} \end{cases}$$

- characterize in-step behavior of pairs of permutation entropies

$$\gamma := \sum_{i=1}^{N_w} S_{\mathbf{X}}(w_i) S_{\mathbf{Y}}(w_i)$$

-  $\gamma = 0$  for independent systems;  $\gamma \rightarrow 1$  for synchronized systems;  $\gamma$  increase monotonically with increasing coupling strength  $\rightarrow$  data-driven estimator for strength of interaction

an example

information-theory

 $( \land (k) \land (l) )$ 

#### measuring interactions

#### strength and direction of interaction

given time series of systems X and Y:

- for estimating probability density distributions and the joint distribution densities:

replace probabilities of data with probabilities of symbols count symbols ( $\rightarrow$  very fast)

- symbolic transfer entropy:

$$T_{\mathrm{Y}\to\mathrm{X}}^{\mathrm{S}} = \sum_{\hat{v}_{i+1}, \hat{\mathbf{v}}_{i}^{(k)}, \hat{\mathbf{w}}_{i}^{(l)}} p\left(\hat{v}_{i+1}, \hat{\mathbf{v}}_{i}^{(k)}, \hat{\mathbf{w}}_{i}^{(l)}\right) \log \frac{p\left(v_{i+1} | \mathbf{v}_{i}^{(\gamma)}, \mathbf{w}_{i}^{(\gamma)}\right)}{p\left(\hat{v}_{i+1} | \hat{\mathbf{v}}_{i}^{(k)}\right)}$$

- see properties of transfer entropy

- easy-to-use data driven estimator for direction of interaction

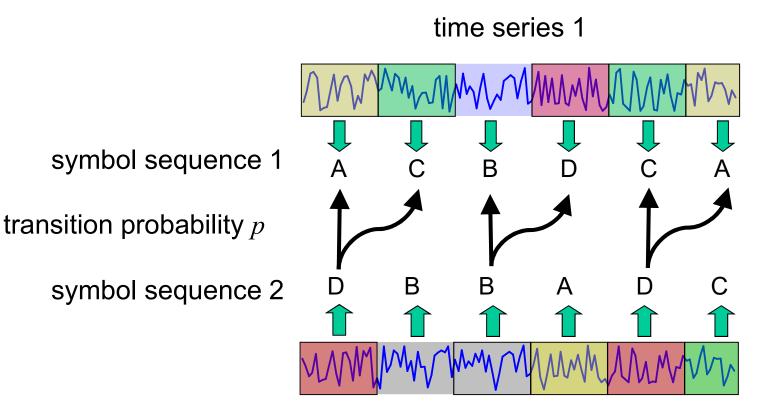
M. Staniek, K. Lehnertz. Symbolic Transfer Entropy. Phys. Rev. Lett. 100, 158101, 2008

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#### measuring interactions

#### strength and direction of interaction

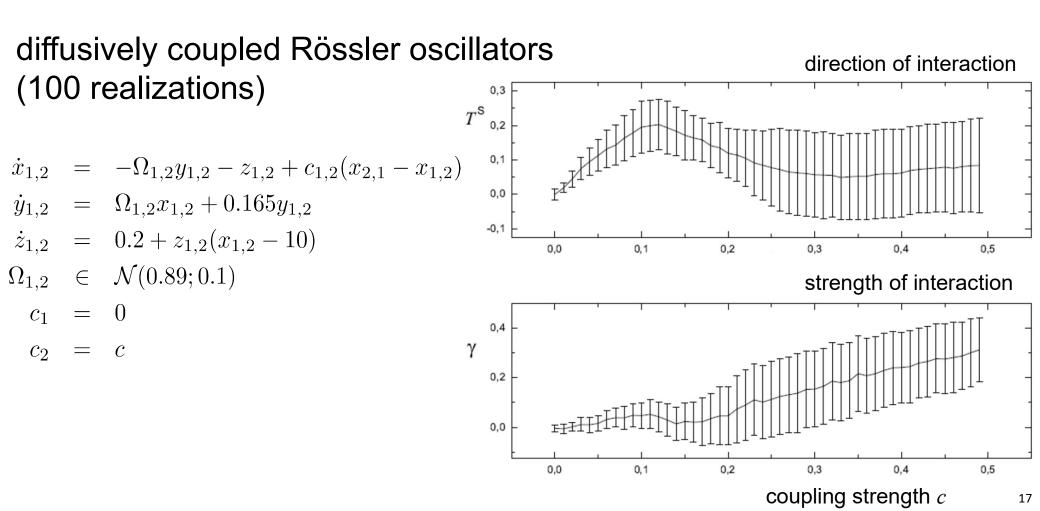


time series 2

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Interactions

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strength and direction of interaction

#### Fundamentals of Analyzing Biomedical Signals

measuring interactions

an example

# measuring interactions strength and direction of interaction

### information-theory

#### diffusively coupled Rössler-Lorenz oscillators (100 realizations of driver-responder system) direction of interaction $\dot{x}^{\mathrm{R}} = -\Omega^{\mathrm{R}}(y^{\mathrm{R}} - z^{\mathrm{R}})$ 0.8 $T^{S}$ $\dot{y}^{\mathrm{R}} = \Omega^{\mathrm{R}}(x^{\mathrm{R}} + 0.2y^{\mathrm{R}})$ 0.6 $\dot{z}^{\mathrm{R}} = \Omega^{\mathrm{R}} \left( 0.2 + z^{\mathrm{R}} (x^{\mathrm{R}} - 5.7) \right)$ 0,4 0,2 0.0 $\dot{x}^{L} = 10(y^{L} - x^{L})$ 2 3 strength of interaction $\dot{y}^{\mathrm{L}} = \Omega^{\mathrm{L}} x^{\mathrm{L}} - y^{\mathrm{L}} - x^{\mathrm{L}} z^{\mathrm{L}} + c(y^{\mathrm{R}})^{2}$ 0.4 $\dot{z}^{\mathrm{L}} = x^{\mathrm{L}}y^{\mathrm{L}} - \frac{8}{2}z^{\mathrm{L}}$ γ

0.2

0.0

0

2

3

coupling strength *c* 

 $\begin{array}{rccc} \Omega^{\mathrm{R}} & \in & \mathcal{N}(6,0.1) \\ \Omega^{\mathrm{L}} & \in & \mathcal{N}(28,1) \end{array}$ 

M. Staniek, K. Lehnertz. Symbolic Transfer Entropy. Phys. Rev. Lett. 100, 158101, 2008

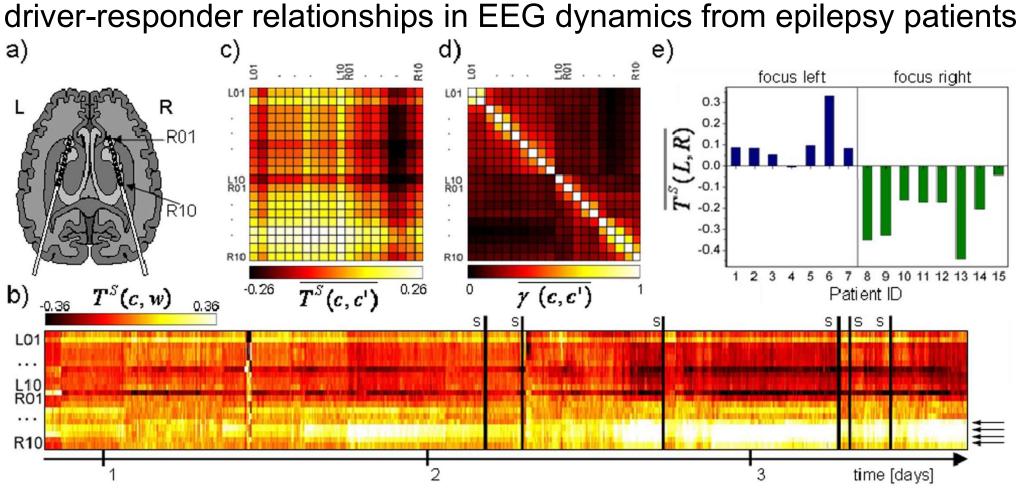
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### information-theory

### strength and direction of interaction

an example



M. Staniek, K. Lehnertz. Symbolic Transfer Entropy. Phys. Rev. Lett. 100, 158101, 2008

#### information-theory

#### strength and direction of interaction

permutation-entropy-based estimators

advantages

- easy-to-use, fast-to-calculate
- high robustness against noise (symbolization)

disadvantages

- symbolization may lead to loss of information
- require appropriate choice of embedding parameter
- choice of window-size, finiteness of available symbols
- "faster" system (eigen-frequency, noise) → driver (need reliable surrogate test for directionality)
- may be fooled by (unobserved) third system