

## Context

- Numerous studies in the literature demonstrated that electroencephalography (EEG) associated with appropriate signal processing methods can bring valuable information on normal and impaired brain networks [1].
- EEG have been largely employed to investigate AD-related alterations in terms of functional connectivity assessment
- The proposed measures in the literature share two main drawbacks:
  - They do not address the problem of EEG non-stationarity
  - They do not consider EEG signal as a multidimensional time series
  - They quantify the spatial relationship between EEG signals without considering the complete alterations due to AD : reduction of complexity



## Epoch-based Entropy (EpEn)

- EpEn measures the information content or the disorder of multivariate EEG signals at two levels simultaneously, using a Hidden Markov Model :
  - at the time level : it quantifies the information content on piecewise stationary epochs of signals over time;
  - at the spatial level: it quantifies the functional connectivity in terms of the heterogeneity of piecewise stationary epochs between multi-channel EEG signals.
- Each observation  $z$  in a given epoch  $S_i$  is considered as a realization  $Z_i$  of a random variable  $Z$  that follows a given observation probability distribution  $P_i(z)$  modeled by the Gaussian mixture.

$$H^*(Z_i) = - \sum_{z \in S_i} P_i(z) \cdot \log_2 P_i(z) \quad \text{EpEn}(Z) = \frac{1}{N} \sum_{i=1}^N H^*(Z_i)$$

- In [2], the efficiency of EpEn was investigated in terms of its robustness to noise, its sensitivity to sampling frequency and to variations of hyper-parameters. We showed that EpEn (AUC=0,90) is more efficient for AD detection compared to correlation dimension (AUC=0,80) and Shannon entropy (AUC=0,74) used for complexity assessment.

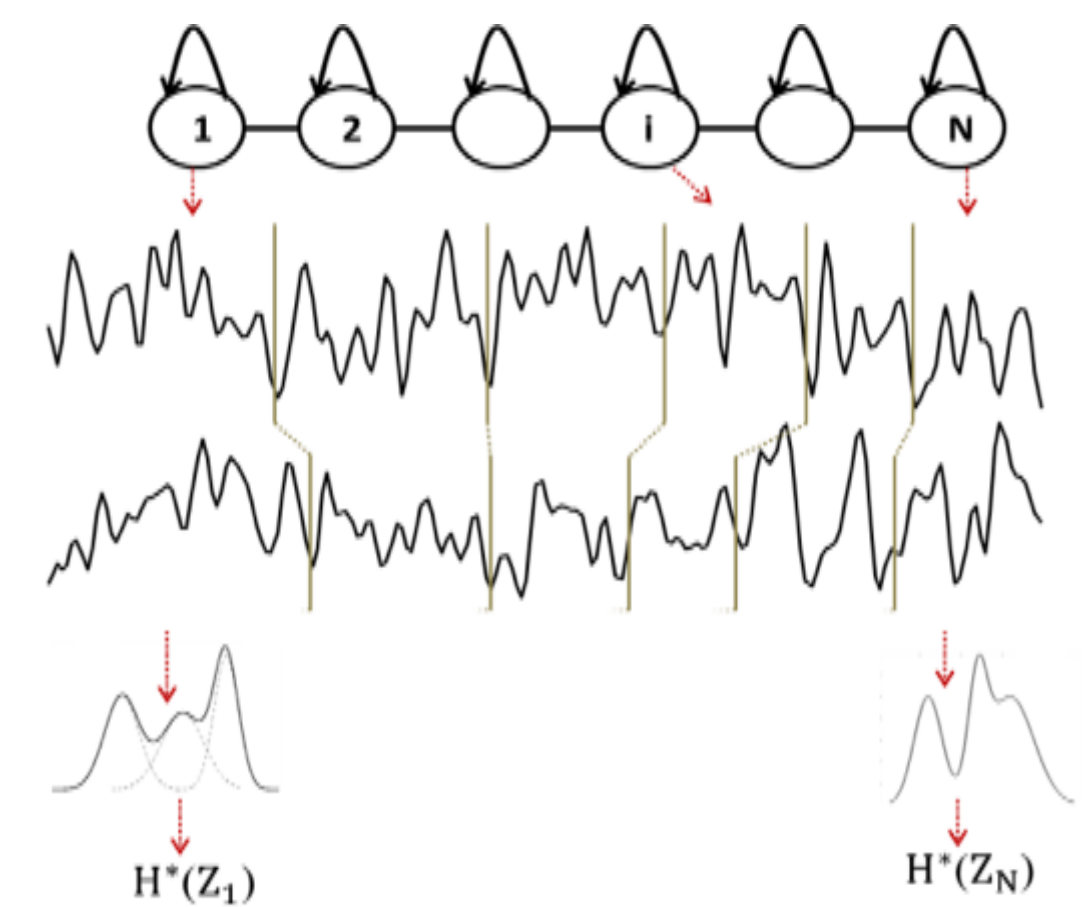


Fig. 1. EEG signal modeling with HMM

## EpEn as a functional connectivity measure

- We compared EpEn to four functional connectivity measures [3].
- Study population: 22 subjects with subjective cognitive impairment (SCI), 28 Mild AD patients (AD), and 22 patients with mild cognitive impairment (MCI).
- Recording at Charles-Foix hospital (France) with a Deltamed EEG acquisition system using 30 electrodes.

Table 1. Best performance with SVM classifier when discriminating AD from SCI with each EEG feature, considering a combination of two brain regions and

AD vs. SCI	Coherence	Phase synchrony	Granger causality	Mutual Information	EpEn
<b>Accuracy</b>	70%	72%	74%	70%	98%
<b>Sensitivity (AD)</b>	59.1%	63.6%	54.5%	89.3%	95.5%
<b>Specificity (SCI)</b>	78.6%	78.6%	89.3%	45.5%	100%

Table 2. Best performance with SVM classifier when discriminating AD from MCI with each EEG feature, considering a combination of two brain regions.

AD vs. MCI	Coherence	Phase synchrony	Granger causality	Mutual information	EpEn
<b>Accuracy</b>	68 %	64 %	58 %	74 %	100 %
<b>Sensitivity (AD)</b>	60.7 %	85.7 %	60.7 %	96.4 %	100 %
<b>Specificity (MCI)</b>	77.3 %	36.4 %	54.6 %	45.5 %	100 %

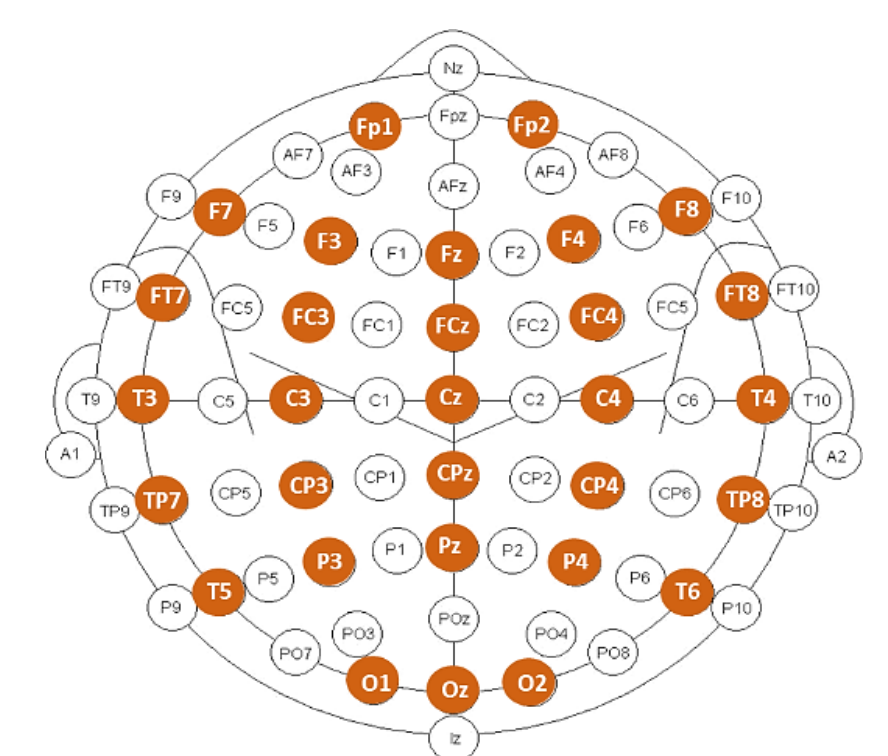


Fig. 2. Distribution of the 30 electrodes on the scalp

## Conclusion

- The statistical measure is by far a more reliable and discriminant feature for AD detection, on our experimental data.
- The originality of this statistical measure lies on the fact that it allows a better estimation of the spatio-temporal characteristics of EEG signals merged into a single figure.
- The structure of the HMM is suitable for modeling neural dynamics underlying the observed multidimensional EEG time series.

[1].H. Hampel, N. Toschi, C. Babiloni, F. Baldacci, et al. Alzheimer Precision Medicine Initiative (APMI). Revolution of Alzheimer Precision Neurology. Passageway of Systems Biology and Neurophysiology, J Alzheimers Dis., 64(s1):S47-S105, 2018.

[2] N. Houmani, F. B. Vialatte, G. Dreyfus, Epoch-based entropy for early screening of Alzheimer's disease, Int. J. of Neural Systems, Vol. 25, Issue 8, 2015, 1550032.

[3] N. Houmani, M. Abazid, K. De Santiago, J. Boudy, et al., EEG signal analysis with a statistical entropy-based measure for Alzheimer's disease detection, open access book, Advances in Signal Processing: Reviews, Book Series, Vol. 2 published by IFSA Publishing, S.L., accepted, 2021.