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### MULTIVARIATE HURST EXPONENT ESTIMATION IN FMRI. APPLICATION TO BRAIN DECODING OF PERCEPTUAL LEARNING.

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#### ABSTRACT

So far considered as noise in neuroscience, irregular arrhythmic field potential activity accounts for the majority of the signal power recorded in EEG or MEG [1,2]. This brain activity follows a power law spectrum  $P(f) \sim 1/f^{\beta}$  in the limit of low frequencies, which is a hallmark of scale invariance. Recently, several studies [1, 3-6] have shown that the slope  $\beta$  (or equivalently Hurst exponent H) tends to be modulated by task performance or cognitive state (eg, sleep vs awake). These observations were confirmed in fMRI [7–9] although the short length of fMRI time series makes these findings less reliable. In this paper, to compensate for the slower sampling rate in fMRI, we extend univariate wavelet-based Hurst exponent estimator to a multivariate setting using spatial regularization. Next, we demonstrate the relevance of the proposed tools on resting-state fMRI data recorded in three groups of individuals once they were specifically trained to a visual discrimination task during a MEG experiment [10]. In a supervised classification framework, our multivariate approach permits to better predict the type of training the participants received as compared to their univariate counterpart.

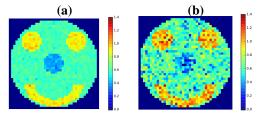
*Index Terms*— fMRI, scale-free brain activity, Total variation regularization, multisensory learning.

#### 1. INTRODUCTION

Scale-free brain activity is the prominent part of brain signals recorded in eletro and magneto-encephalography (EEG/MEG), which can be easily captured by measuring the linear slope  $\beta$ of the log-log plot of the power spectrum  $P(f) \sim 1/f^{\beta}$ . In contrast to some oscillatory regimes which are only observed in response to external stimulations (eg  $\gamma$  oscillations beyond 30 Hz), scale-free activity is a persistent brain dynamics in the limit of low frequencies (0.1 Hz up to 3 Hz), observed both at rest and during task performance as well as in various cognitive states (eg, sleep) [1,3,5,6,11]. It has been shown that scale-free brain activity is functionally associated with neural excitability [2], hence supporting the observation that the  $\beta$ parameter decreases (or the slope becomes flatter) when an individual is engaged in a task as compared to rest. The functional relevance of scale-free brain dynamics has also been illustrated in fMRI even though this imaging technique is less rich temporally. Several fMRI studies [7–9, 12] have actually shown modulations in scale-free brain activity between rest and task and between healthy subjects and Alzheimer's diseased patients [13].

To date, the most commonly used approaches for estimating the slope  $\beta$  or equivalently the Hurst exponent H are threefold: (i) linear regression performed in the frequency domain on the log-scale Welch periodogram; (ii) Detrended Fluctuation Analysis that performs linear regression in the temporal domain [7] and (iii) linear regression from the logscale diagram in the wavelet domain [14]. The statistical performances of these estimators have been thoroughly compared in [15] where it has been shown that wavelet-based analysis is more efficient (no bias, smaller variance). Also, these approaches are univariate: they perform scale-free analysis voxelwise (fMRI) or sensorwise (M/EEG) whereas the neuroimaging data are multivariate. Here, our goal is to develop a multivariate wavelet-based Hurst exponent estimator that accounts for spatial correlation. Our approach is specifically dedicated to fMRI where the signals are short in time and where spatial correlation does exist. As illustrated on synthetic signals in Fig. 1, Hurst exponent estimates become less accurate when the signals are shorter. Hence, by taking local information into account, we hope counterbalancing the shortness of fMRI time series and recovering more accurate Hurst exponents. Our approach will be compared to the univariate wavelet-based estimator in a supervised classification setting. Of note, the proposed methodology cannot enter in a classical GLM analysis since, in contrast to functional connectivity, no seed-based time series can serve as specific regressor for whole brain analysis.

The paper is organized as follows. In Sec. 2, we summarize our learning MEG experiment [10] where three groups of individuals (V/AV/AVn) were engaged in different types of training to a complex visual task. These participants were then scanned in resting-state fMRI to test whether intrinsic



**Fig. 1**. Univariate estimation of Hurst exponents from spatially correlated synthetic signals as illustrated by the smiley. (a) The signal length is 4096 samples in each pixel. (b) The same signals are considered but truncated to the first 514 time points. Shorter signals yield less accurate estimates.

brain dynamics captures differences between training types. In Sec. 3, the multivariate wavelet-based Hurst exponent estimator is presented. In Sec. 4, we assess the performance of our approach on rs-fMRI data by computing the prediction accuracy of supervised classifiers that perform binary classification (eg, V-AV) from Hurst exponents as input features.

#### 2. MULTI-SENSORY LEARNING PARADIGM

In a recent study [10, 16], we tested whether scale-free brain dynamics measuring in MEG was modulated by perceptual learning. For doing so, 36 participants were submitted to a difficult visual discrimination task which consisted of identifying between two random dot kinematograms (one red, one green) which one was moving coherently for about 1 s. The coherence level corresponded to the rate of dots moving in the same direction: it was varied from 15 % to 95 % to make the task more or less difficult. The experiment lasted about 1h30 along which rest and task blocks alternated. A pre-training block was used to calibrate each individual's performance, namely to assess his perceptual threshold corresponding to the coherence level associated with 75 % of correct responses. An individual training period (4 task and rest blocks) was then performed at coherence levels around each individual's perceptual threshold ( $\pm 10\%$ ,  $\pm 20\%$ ).

The 36 individuals were split in three groups of equal size. Each group was submitted to a specific training type: either purely visual (V), or auditory-visual (AV) where congruent acoustic textures were delivered over headphones to make the learning process easier, or finally visual with acoustic white noise (AVn) to serve as control with respect to AV. Finally, a post-training block identical to the pre-training one was used to measure the improvement in behavioral performance in each individual. In [16], we have shown that the Hurst exponents were decreased after training and that the this decrease was negatively correlated with improvements in behavorial performance in the left inferior temporal cortex, visual motion area (MT) and right inferior parietal cortex. Rs-fMRI data were then acquired on the same individuals.

#### 3. SCALE-FREE PARAMETER ESTIMATION

#### 3.1. Scale-free modeling

Expanding on the classical modeling of scale-free dynamics by a power-law decrease of the sole power spectrum, quantifying self-similarity amounts to modeling signals with scalefree dynamics as fractional Gaussian noises (fGn), that is stationary Gaussian stochastic processes consisting of the fractional integration (of parameter H - 1/2) of a white (i.e., delta-correlated) Gaussian process. The sole parameter Hgoverns the entire covariance structure and thus, with Gaussianity, completely defines fGn. More precisely, self-similar parameter H quantifies the algebraic decrease of the correlation function: H = 1/2 indicates the absence of correlation, H < 1/2 betrays negative correlation and H > 1/2 marks long range positive correlation. Parameters H and  $\beta$  can be related as  $\beta = 2H - 1$ . While the classical definition of fGn implies 0 < H < 1, it can be theoretically extended to > 1 [17], while preserving the original intuition beyond fGn: the larger H, the longer term the covariance and the more structured the fGn.

#### 3.2. Scale-free analysis

Exponent  $\beta$  was classically estimated by linear regressions in a log of the power spectrum versus log of frequency plots. Alternatively, time domain approaches such as detrended fluctuation analysis [18] rely on linear regressions. It is now well-accepted that multiscale representations such as wavelet transforms are well-suited for the analysis of scale-free dynamics [14, 19].

Let  $\psi_0(t)$  denote a reference pattern referred to as the mother wavelet, the discrete wavelet coefficients  $d_X(j,k)$  are defined on a dyadic grid (scale  $a = 2^j$  and time  $t = k2^j$ ) as:  $d_X(j,k) = \int X(t)2^{-j}\psi_0(2^{-j}t-k)dt$ . In practice, we used Daubechies wavelets for  $\psi_0$  with a number of vanishing moment  $N_{\psi} = 2$ . It can be shown that for self-similar processes:  $S_X^d(j,q) \equiv \frac{1}{n_j} \sum_{k=1}^{n_j} |d_X(j,k)|^q \simeq K_q 2^{jqH}$ , where  $n_j$  is the number of  $d_X(j,k)$  available at scale j. This thus permits a robust and efficient estimation of H, often using q = 2 in analogy to Fourier spectrum [14], by performing weighted linear regression in a log scale diagram ( $\log_2 S_X^d$  vs  $\log_2 2^j = j$ ). This amounts to finding the minimizer of

$$f(H) = \sum_{j=j_1}^{j_2} n_j \|\log_2 S_X^d(j,2) - (2jH + c)\|_2^2.$$
(1)

where c is an intercept. Scales are linked to frequencies by  $f = 2^{-j} \frac{F_c}{\delta}$  where f is the frequency, j is the scale,  $\delta$  is the sampling period,  $F_c$  is the center frequency of the wavelet in Hz (equal to  $\frac{3}{4}$  for Daubechies wavelets). The scaling range  $(j_1, j_2)$  comprises the set of scales over which scale invariance is statistically valid.

#### 3.3. Multivariate Hurst exponent estimation

Let us denote I the number of voxels in the brain mask  $\Omega$  computed from fMRI data preprocessings ( $I \sim 5 \, 10^4$ ). Starting from Eq. (1), the data consistency cost function thus reads:

$$F(H) = \sum_{i \in \Omega} \sum_{j=j_1}^{j_2} n_j \|\log_2 S^d_{X_i}(j,2) - (2jH_i + c_i)\|_2^2$$
(2)

To incorporate spatial correlation information in Hurst exponent estimation and thus retrieve closer parameter values in neighboring voxels, we consider the regularized criterion

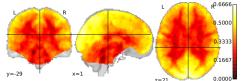
$$J_{\lambda}(H) = F(H) + \lambda G(H) \tag{3}$$

where F is given in Eq. (2) and G is a spatial regularization term.  $\lambda$  is the regularization parameter making the trade-off between data consistency and confidence in prior knowledge. Hereafter, we consider an isotropic total variation penalty term:  $G(H) = \sum_{i \in \Omega} \|\nabla H_i\|_2$  so as to preserve discontinuities between smooth regions of Hurst exponents. This choice makes  $J_{\lambda}$  convex but non-smooth. The minimization of Eq. (3) thus relies on a proximal gradient descent algorithm [20]. The calibration of  $\lambda$  was done on a discrete grid of parameter values as explained in the next section. Our implementation was done in Python and is available in Github<sup>1</sup>. It relies on the nilearn for convex optimization [20].

#### 4. RESULTS IN FMRI DATA ANALYSIS

We performed univariate and multivariate wavelet-based Hurst exponents estimation on 36 individuals. The scaling range  $(j_1, j_2)$  was set to (3, 7) which correspond to a frequencies ranging from 0.005 Hz up to 0.1 Hz, the typical interval where scale-free dynamics have been reported in [7–9]. The grouplevel median map of Hurst exponents (univariate estimator) is shown in Fig. 2 and reflects the gray-white matter contrast: Hurst exponents are above 0.5 mostly in the cortical regions. Conversely, the Hurst exponents associated with voxels located in the white matter the are significantly lower than 0.5. Given the existing spatial correlation on this map, we hypothesize that our multivariate wavelet-based estimator might deliver enhanced maps of Hurst exponents.

To compare multivariate and univariate Hurst exponent estimators, we pursued two objectives: (i) try to distinguish V, AV and AVn groups on the basis of of their rs-fMRI scale-free properties (H maps) and see how the performances of the supervised classifier are impacted by the estimator choice (see subsection 4.1); (ii) assess how the estimator choice affects the statistical significance when computing group-level differences using two-sample t-tests (see subsection 4.2).



**Fig. 2.** Median value of Hurst exponents estimated on real rs-fMRI data from the 36 individuals using the univariate wavelet-based estimator.

#### 4.1. Supervised classification

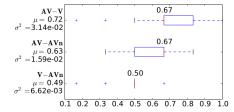
The supervised classifier we used relies on logistic regression as loss function to predict binary outcomes and makes group comparisons such as (AV -V) or any other pair. We thus split the group pair of 24 individuals (taken out of 36) in training and test sets. The training set was composed of subjects randomly chosen in both groups with the same occurrence proportions. The input features entered in the classifier were the individual brain maps of Hurst exponents computed either using the univariate or multivariate wavelet-based estimator. Regarding the univariate approach, we performed the analysis either on spatially unsmoothed fMRI images or on their spatially smoothed version with a Gaussian filter (FWHM = 6 mm). Next, we tested the accuracy of the classifier on unseen subjects to predict the training type they underwent. The evaluation of the classifiers was done by cross-validation, which consists of a stratified-shuffle split loop with 100 iterations and a test fold size of 30% of the whole dataset. Our implementation was based on the scikit-learn package [21].

So far, parameter  $\lambda$  has been set by hand to the value ( $\lambda = 3$ ) providing the best average classification accuracy across all group comparisons. This might induce slight overfitting. However, cross validation cannot be helpful in the present context since  $\lambda$  is not involved in the classification algorithm but instead in the multivariate estimation (3).

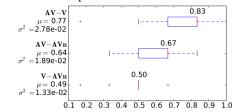
Fig. 3 shows the prediction accuracies<sup>2</sup> for the 3 investigated comparisons. We observed similar trends: the AV training type is easier to discriminate from others and the discrimination between V and AVn groups remains at the chance level. In contrast, we achieved up to 83 % of right classification for discriminating V from AV. This result was obtained using either multivariate inference or univariate inference on filtered data. We then tested the statistical significance of such prediction accuracy differences between methods using paired t-tests. The p-values are reported in Tab. 1. Regarding the **AV-V** comparison, we only found a significant difference between univariate approaches which is due to the smoothing effect. In contrast, for the **AV-AVn** comparison, we demonstrated that our multivariate estimation scheme provided significantly better predictions.

<sup>&</sup>lt;sup>1</sup>https://github.com/JFBazille/ICode.

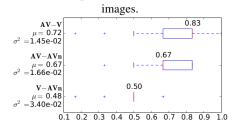
<sup>&</sup>lt;sup>2</sup>Box plots show the median and quartiles over 100 stratifications.



(a) Univariate Hurst exponent estimation from raw fMRI images.



(b) Univariate Hurst exponent estimation from smoothed fMRI



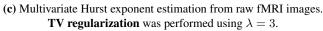


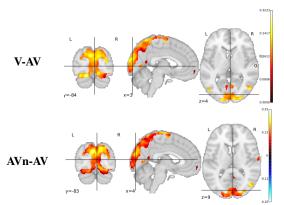
Fig. 3. Performance of the classifiers taking wavelet-based Hurst exponent estimates as input features to discriminate groups (AV-V, AV-AVn and V-AVn comparisons).

**Table 1**. Statistical comparisons between estimation methods regarding their prediction accuracy when performing classifications. Significant p-values for paired Student-t tests appear in bold font.

p-val.	AV -V	AV - AVn	V -AVn
Multiv. vs Univ. raw	0.37	$10^{-4}$	0.7
Multiv. vs Univ. smoothed	0.1	0.006	0.32
Univ. raw vs Univ. smoothed	0.006	0.22	0.53

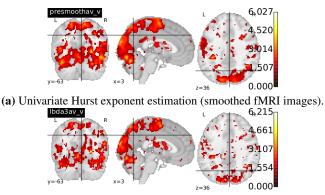
#### 4.2. Statistical comparisons: two-sample Student *t*-tests

We noticed that the best prediction was achieved for the (AV - V) comparison. Hence, individuals in the AV group are suspected to reflect very specific Hurst exponent behavior. To further investigate this issue, we computed the group-level differences (V-AV) and (AVn-AV) of averaged Hurst exponents and reported these parameter differences over the statistically significant voxels in Fig. 4. Clearly, the Hurst exponents in AV group are significantly lower than those in V and AVn groups, especially in the visual and parietal cortices. This indicates that neural excitability (decrease in H) was enhanced by multisensory training. This result is consistent with our previous observation [16] that after training the decrease in H computed from source reconstructed MEG time series was more significant in parieto-occipital cortices.



**Fig. 4**. Map of the mean group-level differences of univariate Hurst exponent estimates computed from smoothed fMRI images. Only significant voxels are reported (as in Fig. 5(b)).

Last, Fig. 5 shows different maps of  $(-\log_{10} p_{val})$  from two sample Student *t*-tests associated with the **AV-V** comparison. The multivariate Hurst exponent estimator computed with TV regularization ( $\lambda = 3$ ) provided slightly more significant peaks as compared to the univariate ones. Next, irrespective of the the estimation procedure, most of the significant voxels are located in the occipito-parietal and motor areas, the latter being not specifically expected to bring discriminative information between the V and AV groups.



(**b**) Multivariate Hurst exponent estimation ( $\lambda = 3$ ).

**Fig. 5.** Maps of  $-\log_{10} p_{\text{val}}$  from two-sample Student *t*-tests performing the comparison of Hurst exponent estimates between **AV** and **V** groups.

#### 5. CONCLUSION

In this paper, we proposed a multivariate wavelet-based Hurst exponent estimator specifically dedicated to fMRI data to compensate for the shortness of these time series. We demonstrate its relevance on resting-state fMRI data acquired after a learning experiment. We showed that this estimator outperformed its univariate competitors in a supervised classification task for predicting the training type the individuals underwent, especially for the (**AV-AVn**) comparison.

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