



Self-similarity and multifractality in human brain activity: A wavelet-based analysis of scale-free brain dynamics[☆]

Daria La Rocca^{a,b}, Nicolas Zilber^{a,b}, Patrice Abry^c, Virginie van Wassenhove^{a,d},
Philippe Ciuciu^{a,b,*}

^a CEA/DRF/Joliot, NeuroSpin, Université Paris-Saclay, F-91191 Gif-sur-Yvette, France

^b INRIA, Parietal Team, Université Paris-Saclay, France

^c Univ Lyon, Ens de Lyon, Univ Claude Bernard, CNRS, Laboratoire de Physique, F-69342 Lyon, France

^d Cognitive Neuroimaging Unit, INSERM, Université Paris-Saclay, F-91191 Gif-sur-Yvette, France

ARTICLE INFO

Keywords:

1/f power spectrum
MEG
Temporal structure
Oscillations
Resting-state
Infra-slow activity
Fractal
Multifractal
Wavelet

ABSTRACT

Background: The temporal structure of macroscopic brain activity displays both oscillatory and scale-free dynamics. While the functional relevance of neural oscillations has been largely investigated, both the nature and the role of scale-free dynamics in brain processing have been disputed.

New method: Here, we offer a novel method to rigorously enrich the characterization of scale-free brain activity using a robust wavelet-based assessment of self-similarity and multifractality. For this, we analyzed human brain activity recorded with magnetoencephalography (MEG) while participants were at rest or performing a visual motion discrimination task.

Results: First, we report consistent infraslow (from 0.1 to 1.5 Hz) scale-free dynamics (i.e., self-similarity and multifractality) in resting-state and task data. Second, we observed a fronto-occipital gradient of self-similarity reminiscent of the known hierarchy of temporal scales from sensory to higher-order cortices; the anatomical gradient was more pronounced in task than in rest. Third, we observed a significant increase of multifractality during task as compared to rest. Additionally, the decrease in self-similarity and the increase in multifractality from rest to task were negatively correlated in regions involved in the task, suggesting a shift from structured global temporal dynamics in resting-state to locally bursty and non Gaussian scale-free structures during task.

Comparison with existing method(s): We showed that the wavelet leader based multifractal approach extends power spectrum estimation methods in the way of characterizing finely scale-free brain dynamics.

Conclusions: Altogether, our approach provides novel fine-grained characterizations of scale-free dynamics in human brain activity.

1. Introduction

1.1. Scale-free brain activity

Macroscopic brain activity consists of a mixture of synchronized and desynchronized activity (He et al., 2010; Breakspear, 2017). The synchronization of neural oscillations has been hypothesized to mediate neural communication (Engel et al., 2001; da Silva, 2013; Buzsáki, 2010; Wang, 2010), and their coupling, to be involved in information processing (Fries, 2005; Lakatos et al., 2005; Jensen and Colgin, 2007;

da Silva, 2013; Buzsáki et al., 2012). However, the existence, the properties and the functional relevance of scale-free dynamics in brain processing remain an open debate. Scale-free dynamics have been reported in spontaneous brain activity (He et al., 2010) and in data collected with various neuroimaging techniques including fMRI, magnetoencephalography (MEG), electroencephalography (EEG) and local-field-potentials (LFP) (He et al., 2010; Foster et al., 2016). The presence of scale-free dynamics was demonstrated in the infra-slow frequency range (from 0.01 Hz to 1 Hz, He et al., 2010; Buzsáki and Mizuseki, 2014; He, 2014; Becker et al., 2018) and in the slow fluctuations of

[☆] Work supported by French grant MultiFracS ANR-16-CE33-0020 to PA, ANR Schubert ANR-0909-JCJC-071 to PC and the HUBBLE Enhanced—Eurotalents/#233 contract from the 7th PCRD to DLR. The authors are grateful to Biyu J. He (NYU, School of Medicine) for fruitful discussions on scale-free brain dynamics and to Omar Darwiche Domingues (INRIA Parietal) for his implementation of the WLBMF toolbox in Python.

* Corresponding author at: CEA/DRF/Joliot, NeuroSpin, Université Paris-Saclay, F-91191 Gif-sur-Yvette, France.

E-mail addresses: daria.la-rocca@inria.fr (D. La Rocca), nicolas.zilber@hotmail.com (N. Zilber), patrice.abry@ens-lyon.fr (P. Abry), virginie.van.wassenhove@gmail.com (V. van Wassenhove), philippe.ciuciu@cea.fr (P. Ciuciu).

<https://doi.org/10.1016/j.jneumeth.2018.09.010>

Received 28 April 2018; Received in revised form 6 September 2018; Accepted 6 September 2018

Available online 10 September 2018

0165-0270/© 2018 Elsevier B.V. All rights reserved.

power of narrow-band neuronal oscillations (Freeman, 2000; Linkenkaer-Hansen et al., 2001; Monto et al., 2008; Palva et al., 2013). The empirical work in both humans and animals has revealed that scale-free dynamics of brain activity were modulated by the levels of wakefulness (vs. sleep) (Weiss et al., 2009; He et al., 2010; Tagliazucchi et al., 2013; Dehghani et al., 2012), consciousness (vs. anesthesia) (He and Raichle, 2009; Barttfeld et al., 2015), aging and neurodegenerative diseases (Suckling et al., 2008) as well as task performance (Buiatti et al., 2007; He et al., 2010; He, 2011; Ciuciu et al., 2012; Monto et al., 2008; Palva et al., 2013; Lin et al., 2016).

The intuition behind the scale-free concept is that the relevant information in the temporal dynamics of a given signal is coded within the relations that tie together temporal scales, rather than in the power of neuronal oscillations in specific bands. Its origin remains however poorly understood. Brain activity recorded with MEG or EEG is most comparable to LFP, and slow dynamic fluctuations likely reflect the up and down states of cortical networks as opposed to spiking activity *per se* (Baranauskas et al., 2012). Hence, although fast neuronal activity or avalanches could endogenously produce scale-free infra-slow brain dynamics, a careful statistical assessment remains necessary to draw conclusions on the nature of observed scale-free dynamics (Bedard et al., 2006; Touboul and Destexhe, 2010; Dehghani, 2012). A temporal hierarchy of neural oscillators has been considered a possible source of scale-free brain dynamics (Penttonen and Buzsáki, 2003; He et al., 2010) as well as the spatial repartition of neural sources. Dendritic filtering (Destexhe et al., 1999; Werner, 2010; Buzsáki et al., 2012) or the resistive brain milieu constitute other tentative origins for scale-free dynamics (Dehghani et al., 2010). That the structural configuration of neural networks and their dynamics may be arguably topologically intertwined (Arenas et al., 2008; Werner, 2010; Chaudhuri et al., 2017) is also important to keep in mind. To better understand the origins and nature of scale-free brain dynamics, we thus propose to use a rich and robust statistical framework.

1.2. Scale-free dynamics modeling and assessment

Scale-free dynamics recorded with neuroimaging techniques have generally been quantified using a $1/f^\beta$ power spectrum model over a large continuum of frequencies. As a result, the empirical assessment has often used Fourier-based spectrum estimation. As an alternative, self-similarity provides a well accepted model for scale-free dynamics that encompasses, formalizes, and enriches the traditional Fourier $1/f^\beta$ spectrum modeling, with models such as fractional Brownian motion (fBm) or fractional Gaussian noise (fGn) (Novikov et al., 1997; He et al., 2010; Ciuciu et al., 2012, 2014). The self-similarity, or Hurst, parameter H matches the spectral exponent β , as $\beta = 2H - 1$ for fGn, and as $\beta = 2H + 1$ for fBm. In the context of brain activity, H indexes how well neural activity is temporally structured (via its autocorrelation). Additionally, although H has been estimated using Detrended Fluctuation Analysis (DFA) (Linkenkaer-Hansen et al., 2001; Buiatti et al., 2007; He, 2011; Hardstone et al., 2012; Palva et al., 2013; Barttfeld et al., 2015), it is now well-documented that wavelet-based estimators provide significant theoretical improvements and practical robustness over DFA, notably by disentangling true scale-free dynamics from non-stationary smooth trends (Veitch and Abry, 1999; Torres and Abry, 2003; Baykut et al., 2005; Ciuciu et al., 2012, 2014). This is further detailed in Section 2.2.2. For a review on statistically relevant estimation of the self-similarity parameter, interested readers are also referred to (Bardet et al., 2003).

Often associated with Gaussianity, self-similarity alone does not fully account for scale-free dynamics. The main reason is that self-similarity restricts the description of neural activity to second-order statistics (autocorrelation and Fourier spectrum) and, hence, to additive processes. Yet, it has been proposed that multiplicative processes may provide more appropriate descriptions of neural activity (Buzsáki and Mizuseki, 2014). Independently of, and in addition to self-similarity,

multifractality provides a framework to model these non-additive processes (Shimizu et al., 2004; Suckling et al., 2008; Van de Ville et al., 2010). Multifractality can be conceived as the signature of multiplicative mechanisms, or as the intricate combination of locally self-similar processes. For instance, if a patch of cortex (i.e. the anatomical resolution of MEG recordings) is composed of several small-networks each characterized by a single self-similar parameter H , the multifractality parameter M constitutes an index capturing the diversity of H s and their interactions within the patch. Qualitatively, the multifractality parameter M quantifies the occurrence of transient local burstiness or non Gaussian temporal structures, not accounted for by the autocorrelation function or by the Fourier spectrum (hence, neither by H nor β). To meaningfully and reliably estimate M , it has been theoretically shown that the wavelet-based analysis must be extended to wavelet-leaders (Wendt et al., 2007).

1.3. Goals and contributions

The goal of the present work is to produce a rich and reliable characterization of scale-free temporal dynamics in human brain activity, and to provide the field with a robust and reliable procedure to do so. This is made possible (i) by the combined use of self-similarity and multifractality as independent and complementary modeling paradigms, and (ii) by the recourse to the wavelet and wavelet-leader based assessment framework yielding improved performance and robustness to nonstationary trend procedures (Veitch and Abry, 1999; Torres and Abry, 2003; Baykut et al., 2005; Ciuciu et al., 2012, 2014). The present work investigates the existence and characterization of scale-free dynamics in human cortical activity recorded with MEG, and investigates the modulation of H and M by resting-state and task.

2. Material and methods

2.1. Material

2.1.1. Participants

Twenty-four right-handed participants (10 females; mean age of 22.1 ± 1.9 y.o.) took part in the study. All had normal or corrected-to-normal vision and normal hearing and provided a written informed consent prior to the experiment in accordance with the Declaration of Helsinki (2008) and the local Ethics Committee on Human Research at NeuroSpin (Gif-sur-Yvette, France).

2.1.2. Experimental design

The resting-state block lasted 5 min during which participants kept their eyes open while staring at a black screen. Participants could mind-wander freely. 5 min were selected to be sufficient for an accurate estimation of scale-free properties but not long enough for participants' cognitive state to drastically change. Resting-state activity was recorded prior to any exposure to task or stimuli. The task block lasted 12 min during which participants performed a visual motion coherence discrimination task (Zilber et al., 2014). In each trial (2.5 s), participants decided which of two intermixed (green and red) clouds of dots was most coherent. Responses were delivered by button press. The experiment was conducted in a darkened soundproof magnetic-shielded room. Participants were seated in upright position under the MEG dewar facing a projection screen placed 90 cm away. The refresh rate of the projector (model PT-D7700E-K, Panasonic Inc, Kadoma, Japan) was 60 Hz. Participants were explained the task and were in contact at all times with the experimenter via a microphone and a video camera. Stimuli were designed using Matlab (R2010a, Mathworks Inc.) with Psychtoolbox-3 (Pelli, 1997) on a PC (Windows XP).

2.1.3. MEG data acquisition

Brain activity was recorded in a magnetically shielded room using a 306 MEG system (Neuromag Elekta LTD, Helsinki). MEG recordings

were sampled at 2 kHz and band-pass filtered between 0.03 and 600 Hz. Four head position coils (HPI) measured participants' head position before each block; three fiducial markers (nasion and pre-auricular points) were used for digitization and for alignment with the anatomical MRI (aMRI) acquired immediately after MEG acquisition. Electrooculograms (EOG, horizontal and vertical eye movements) and electrocardiogram (ECG) were simultaneously recorded. Before each experiment, 5 min of empty room recordings were acquired for the computation of the noise covariance matrix used in solving the MEG inverse problem.

2.1.4. Anatomical MRI acquisition and segmentation

The T_1 weighted anatomical MRI (aMRI) was recorded using a 3-T Trio MRI scanner (Siemens Erlangen, Germany). Parameters of the sequence were: $FOV = 256 \times 240 \times 176 \text{ mm}^3$, voxel size: $1.0 \times 1.0 \times 1.1 \text{ mm}^3$; acquisition time: 7 min 46 s; repetition time $TR = 2300 \text{ ms}$; inversion time $TI = 900 \text{ ms}$; flip angle = 9° ; transversal orientation, echo time $TE = 2.98 \text{ ms}$ and partial Fourier 7/8. Cortical reconstruction and volumetric segmentation of participants' T_1 weighted aMRI was performed with *Freesurfer*¹ (RRID: nif-0000-00304). This included: motion correction, average of multiple volumetric T_1 weighted images, removal of non-brain tissue, automated Talairach transformation, intensity normalization, tessellation of the gray-white matter boundary, automated topology correction, and surface deformation following intensity gradients (Dale et al., 1999). Once cortical models were complete, deformable procedures could be performed including surface inflation (Fischl et al., 1999a) and registration to a spherical atlas (Fischl et al., 1999b). These procedures were adopted using MNE (Gramfort et al., 2014, RRID: scires.000118) to morph individuals' current source estimates onto the *Freesurfer* average brain for group analysis.

2.1.5. MEG data preprocessing

Data preprocessing was done in accordance with accepted guidelines for MEG research (Gross et al., 2013). Signal Space Separation (SSS) was performed using *MaxFilter* to remove external magnetic interferences and discard noisy sensors (Taulu and Simola, 2006). Ocular and cardiac artifacts (eye blinks and heart beats) were removed using Independent Component Analysis (ICA) on raw signals. ICA was fitted to raw MEG signals, and sources matching the ECG and EOG signals recorded by dedicated channels were automatically found and removed.² Then, for the sake of computational efficiency, we down-sampled the preprocessed MEG time series at $f_s = 400 \text{ Hz}$ before applying signal reconstruction as described in Section 2.1.6, since scale-free analysis was focused on the low frequency content.

2.1.6. Coregistration and MEG source reconstruction

The co-registration of MEG data with the individual's aMRI was carried out by realigning the digitized fiducial points with the multimodal markers visible in MRI slices. We used a two-step procedure to ensure reliable MEG-aMRI coregistration: using *MRILAB* (Neuromag-Elekta LTD, Helsinki), fiducials were aligned manually with the multimodal markers on the MRI slice; an iterative procedure realigned all digitized points (about 30 more supplementary points distributed on the scalp of the subject were digitized) with the scalp of the participant and the MEG coordinates using the *mne_analyze* tools within MNE (Gramfort et al., 2014, RRID:nlx_151346). Individual forward solutions were computed using a 3-layer boundary element model (Hämäläinen and Sarvas, 1989) constrained by the individual's aMRI. Cortical surfaces were extracted with *Freesurfer* (RRID: nif-0000-00304) and decimated to about 5120 vertices per hemisphere with 4.9 mm spacing. The

forward solution, noise and source covariance matrices were used to compute the noise-normalized dynamic statistical parametric mapping (dSPM) (Dale et al., 1999) inverse operator (depth = 0.8).³ The unitless inverse operator was applied using a loose orientation constraint on individuals' brain data (Lin et al., 2006) by setting the transverse components of the source covariance matrix to 0.4. Importantly, considering that taking the norm of source dipoles is a nonlinear transformation that may modify scale-free properties (Zilber et al., 2012), we only kept the radial components. Using the individual cortical parcellation based on the Destrieux atlas (138 labels in total) provided by *Freesurfer*, reconstructed time series in vertices belonging to the same cortical label were grouped and averaged to obtain a unique time series. In this procedure, the signs of time series within labels were flipped according to anatomical orientation of vertices in such a way that signed activations did not cancel out after averaging (this is a standard label averaging used by the MNE software).

2.2. Methods

2.2.1. Scale-free modeling: From Fourier spectrum to selfsimilarity and multifractality

Scale-free dynamics are classically modeled by a power-law decrease of the Fourier power spectrum $\Gamma(f)$ with respect to frequencies f : $\Gamma(f) \approx C|f|^{-\beta}$. Such power laws can be understood as the signatures of the more general and better theoretically framed concept of self-similarity (Samorodnitsky and Taqqu, 1994). In essence, self-similarity amounts to modeling scale-free dynamics in data as fractional Gaussian noise (fGn), a Gaussian stationary stochastic process, consisting of the fractional integration (with parameter $H - 1/2$) of a white (i.e., delta-correlated) Gaussian process. The sole parameter H , theoretically related to β as $\beta = 2H - 1$, governs the entire covariance structure and thus, together with Gaussianity, completely defines temporal dynamics. More precisely, the self-similar parameter H quantifies the algebraic decrease of the autocorrelation function: $H = 1/2$ indicates the absence of correlation, $H < 1/2$ corresponds to negative correlation and $H > 1/2$ marks long range positive correlation. While the classical definition of fGn implies $0 < H < 1$, it can be theoretically extended to $H > 1$ (with the recourse to the notion of generalized processes and tempered distributions; Samorodnitsky and Taqqu, 1994), while preserving the original intuition beyond fGn: the larger $|H - 1/2|$, the more structured the temporal dynamics of data (as illustrated in Supplementary Movie 1). Beyond the global control of temporal dynamics via the covariance function, Gaussian self-similarity also implies the absence of fluctuations in the regularity of local temporal dynamics. Such local regularity is often quantified via the Hölder exponent $h(t) > 0$ (Wendt et al., 2007). For Gaussian self-similar processes, such as fGn, $\forall t, h(t) \equiv H$.

The multifractal paradigm extends self-similarity by preserving a control of the global temporal dynamics via the covariance function, driven by H , while enriching it with possible fluctuations along time of the local regularity $h(t)$ (Wendt et al., 2007). Multifractal models, such as multifractal random walk (MRW), are thus essentially stationary non Gaussian processes, defined as the fractional integration (of parameter $H - 1/2$) of a white (i.e., delta-correlated) Gaussian process, whose amplitude is modulated by another independent process, whose covariance decreases logarithmically slowly, with an amplitude controlled by the multifractality parameter $M \geq 0$ (Bacry et al., 2001). Self-similarity parameter H preserves the intuitive interpretation of global and overall dependence and structure in the temporal dynamics of data, while the additional multifractal parameter M allows local and transient departures from Gaussianity, hence burstiness in temporal dynamics, via fluctuations along time of the local regularity (as illustrated

¹ <http://surfer.nmr.mgh.harvard.edu>

² https://github.com/mne-tools/mne-python/blob/master/tutorials/plot_artifacts_correction_ica.py

³ https://github.com/mne-tools/mne-python/blob/master/tutorials/plot_mne_dspm_source_localization.py

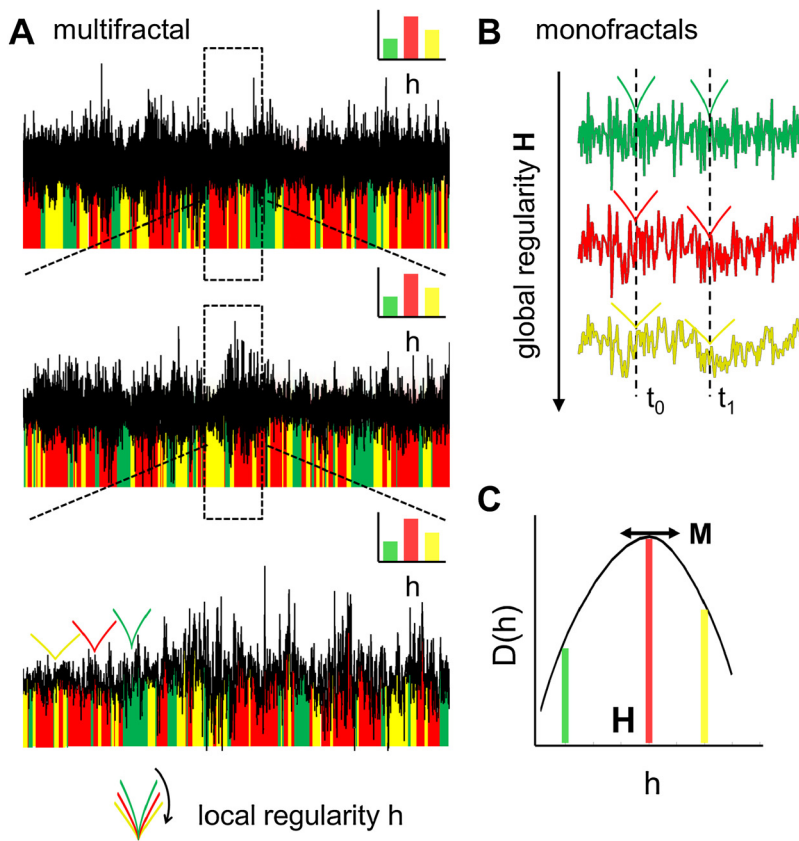


Fig. 1. Schematic introduction to multifractality. (A) Multifractal signals observed at three different time scales from coarse (top) to fine (bottom). Local temporal dynamics can be quantified by the Hölder exponent $h(t)$, a local regularity index. In this pedagogical example, $h(t)$ can only take three values: red, green and yellow. For multifractal signals, $h(t)$ is *per se* a very irregular function along time, with all possible h existing in any small subpart of the data. (B) For monofractal signals (with the same covariance function as the multifractal signals, hence same H), no fluctuations of local regularity are observed and the local h and the global H are everywhere identical. In these three examples of monofractal signals, the global regularity H of each signal increases from top to bottom. (C) The multifractality illustrated in (A) can be captured by a multifractal spectrum $D(h)$, quantifying by means of fractal (Hausdorff) dimension of the geometrical structure of time points that share the same local regularity $h(t) = h$. The most frequent Hölder exponent h , indicated in red, is closely related to the global self-similarity index H whereas multifractality M encompasses all local regularities h by reflecting the width of $D(h)$. Importantly, $D(h)$ remains the same for the whole signal and any subpart, it is hence scale invariant. (For interpretation of the references to color in this figure legend, the reader is referred to the web version of the article.)

in Supplementary Movie 2).

More technically, multifractal temporal dynamics imply that the fluctuations along time of local regularity are erratic, i.e., the function $h(t)$ is itself a very irregular function. Therefore, temporal dynamics are not well-described by the local function $h(t)$, but rather by a global function, the so-called multifractal spectrum $0 \leq D(h) < 1$. The multifractal spectrum, which consists of the fractal dimension of the set of points on the real line sharing the same regularity $h(t) = h$ (cf. [Wendt et al., 2007](#) for a technical definition), thus conveys a global information on the geometrical structure of $h(t)$, hence on temporal dynamics beyond the mere covariance function. These notions are pedagogically, hence qualitatively, illustrated on synthetic data in [Fig. 1](#).

While the multifractal spectrum $D(h)$ can theoretically consist of any shape, it is often efficiently approximated, for practical use, as a parabola controlled by H and M : $D(h) \simeq 1 - (h - H)^2/2M$. For Gaussian self-similar processes, $M = 0$ and $D(h) = \delta(h - H)$, with δ the Dirac-delta function. Parameters H and M thus provide independent and complementary characterization of scale-free dynamics in data ([Wendt et al., 2007](#)), with M adding the possibility to model burstiness in temporal dynamics by local departures from Gaussianity, while the global structure of temporal dynamics remains controlled by H .

2.2.2. Scale-free analysis: From spectral estimation to wavelet and wavelet-leader analysis

The scaling exponent β has classically been evaluated by means of spectrum estimation, i.e., by linear regressions in a log-log plot of estimated power spectrum versus frequency (as sketched in [Fig. 2](#)). In the present work, all Fourier spectra are estimated using the Welch periodogram procedure. Alternatively, time domain approaches such as detrended fluctuation analysis (DFA) ([Linkenkaer-Hansen et al., 2001](#)), also based on linear regressions, rely on quantifying the power of fluctuations in data increments computed at different lags (acting as scales). It is however now well-documented that multiscale representations, such as wavelet transforms, are well-suited for the

analysis of scale-free dynamics and achieve optimal and robust estimation performance cf. e.g., [Abry and Veitch \(1998\)](#), [Torres and Abry \(2003\)](#), [Veitch and Abry \(1999\)](#), and [Bardet et al. \(2003\)](#). 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: $d_X(j, k) = \int X(t)2^{-j}\psi_0(2^{-j}t - k)dt$. Under mild conditions on the choice of $\psi_0(t)$, it has been shown that for self-similar processes ([Abry and Veitch, 1998](#)):

$$S_X^d(j, q) \equiv \frac{1}{n_j} \sum_{k=1}^{n_j} |d_X(j, k)|^q \simeq K_q 2^{j\zeta(q)}, \quad \forall a = 2^j > 0, \quad \forall q > 0 \quad (1)$$

with $\zeta(q) = qH$, thus permitting a robust and efficient estimation of H by linear regressions ([Abry and Veitch, 1998](#)). It can further be shown that, with the particular choice $q = 2$, $S_X^d(j, q = 2)$, referred to as the wavelet spectrum, can be read as an estimator of the Fourier spectrum $\Gamma(f)$ ([Veitch and Abry, 1999](#)). Therefore, under elementary transformations, the Fourier and wavelet spectra can be mapped one onto the other, as $\Gamma(f) \simeq S_X^d(j = \log_2(f_0/f), q = 2)$, with f_0 a constant that depends on the choice of ψ_0 (and that can be well approximated for a large class of wavelets as $f_0 \simeq 3/4 \times f_s$, with f_s the sampling frequency) ([Abry and Veitch, 1998](#); [Veitch and Abry, 1999](#)). This is quantitatively illustrated in [Fig. 2](#). While both spectra yield equivalent information on the global temporal dynamics, it has been documented that the wavelet spectrum yields a more robust and more reliable estimate of H than Fourier spectrum does for β ([Veitch and Abry, 1999](#)). Notably, it was shown that the wavelet spectrum is less prone to bias induced by smooth trends or smooth non stationarity effects, than the classical Fourier spectrum, hence yielding robust estimates of the scale-free exponents.

For multifractal processes, when $M > 0$, the scaling exponents $\zeta(q)$ no longer follow the linear form qH , but rather consist of a concave function, which in first approximation and for practical purposes can be written as $\zeta(q) = qH - Mq^2/2$. The scaling exponents $\zeta(q)$ are further related to the multifractal spectrum $D(h)$ via a Legendre transform

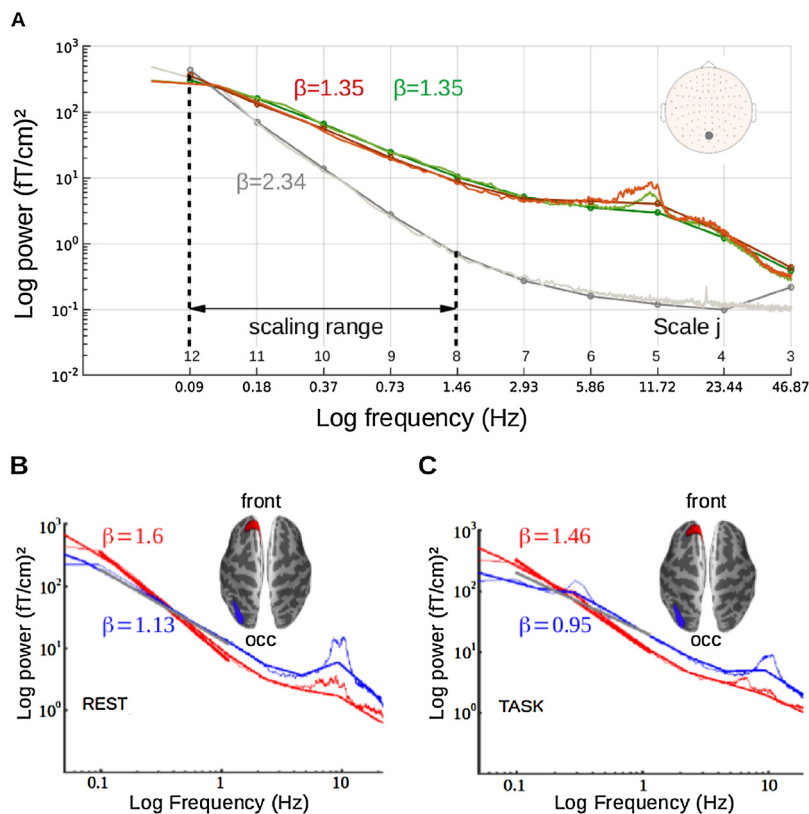


Fig. 2. Scale-free brain dynamics: Fourier vs. wavelet-based power spectra at rest and during task. (A) Group-average Fourier (thin lines) and wavelet (thick lines) power spectra computed in an occipital sensor (inset) for empty room (grey), resting-state (brown) and task (green) recordings. In both Fourier and wavelet log-power spectra, the linear fit indicates scale-free dynamics, and delineates the implicated range of scales ($j \in (8, 12)$) corresponding roughly from 0.1 Hz (i.e. $j = 12$) to 1.5 Hz (i.e. $j = 8$). The slopes quantify the scaling exponents β (of power spectra $1/f^\beta$) and the self-similarity index H . Human brain activity is characterized by a pink noise ($\beta \approx 1$) regime whereas empty room recordings correspond to brown noise ($\beta \approx 2$): hence, and importantly, this graph clearly shows that instrumental noise is not a spurious cause for observing scale-free dynamics in brain activity. (B and C) Group-average Fourier (thin lines) and wavelet-based (thick lines) power spectra computed in two frontal (red) and occipital (blue) cortical labels at rest (B) and during task (C). Larger β values correspond to steeper slopes, as shown in the frontal region (*front*, red label) compared to the occipital (*occ*, blue label). All plots clearly show that wavelet and Fourier spectra can be formally mapped one onto the other. (For interpretation of the references to color in this figure legend, the reader is referred to the web version of the article.)

(Wendt et al., 2007).

For more than one decade (Jaffard et al., 1998) it has been proved that a relevant estimate of parameter M requires to replace the wavelet coefficients with wavelet-leaders, defined as local suprema of the wavelet coefficients $d_X(j', k')$, across a local neighborhood $\lambda_{j,k} = [(k - 2)2^j + 1, (k + 1)2^j]$, for all finer scales $2^{j'} \leq 2^j$ (Wendt et al., 2007):

$$L_X(j, k) = \sup_{j' \leq j, 2^{j'} k' \in \lambda_{j,k}} |d_X(j', k')|. \quad (2)$$

Under mild restrictions, it has been shown that (Wendt et al., 2007):

$$C_1(j) \equiv \frac{1}{n_j} \sum_{k=1}^{n_j} \log_2 L_X(j, k) = H_0 + H_j, \quad (3)$$

$$C_2(j) \equiv \frac{1}{n_j} \sum_{k=1}^{n_j} (\log_2 L_X(j, k))^2 - C_1(j)^2 = M_0 - M_j. \quad (4)$$

These two fundamental relations show that parameters H and M can be estimated as linear regressions in diagrams $C_1(j)$ vs. j and $C_2(j)$ vs. j , respectively. This is illustrated in Fig. 3. To ease exposition, the functions $C_1(j)$ and $C_2(j)$ will hereafter be referred to as the wavelet-leader spectra.

2.2.3. Estimation set-up

In practice, in the following data analysis, spectral estimation was conducted using a standard Welch periodogram estimation procedure, with a classical Hamming windowing (window size of 8192 samples, roughly corresponding to 20 s, and 50% overlap in adjacent windows). Wavelet analysis was conducted using a fast pyramidal discrete wavelet transform algorithm, using Daubechies least asymmetric orthonormal wavelets with $N_w = 2$ vanishing moments (Mallat, 1998). Scale-free and multifractal analyses were performed using the state-of-the-art wavelet leader toolbox made publicly available⁴ and fully detailed in Wendt

et al. (2007), and now available as well in Python.⁵ It has been carefully checked that varying details in data analysis set-up leads to similar conclusions. For the sake of reproducibility, the Python scripts corresponding to the whole MEG data analysis we performed are made publicly available.⁶

3. Results: self-similarity and multifractality in human brain activity recorded with MEG

3.1. Assessing self-similarity in MEG data: range of frequencies, Fourier vs. wavelet power spectra

Fig. 2 reports the group-average Fourier and wavelet spectra in sensors and in cortical source estimations of the entire MEG data time series. As theoretically expected (cf. Section 2.2.2), the Fourier (thin lines) and the wavelet spectra (thick lines) superimposed very well, yielding consistent patterns across methods. Fig. 2 also shows that both spectra displayed power law behaviors over a broad range of frequencies ranging from roughly 0.1 Hz to 1.5 Hz. Importantly, Fig. 2A compares Fourier and wavelet spectra of human brain MEG data to those of empty-room MEG recordings. This formal comparison unambiguously showed that the spectra differed both in amplitude and in shape: the spectral exponent of human brain recordings was in the so-called pink noise regime ($1 \leq \beta \leq 2$) while empty-room recordings rather displayed brown noise temporal dynamics ($\beta \geq 2$). Thus, scale-free dynamics observed in MEG recordings through power spectrum analysis (both Fourier and wavelet) was not caused by instrumental or sensor noise, but rather resulted from macroscopic human brain activity.

Thus, and overall, Fig. 2A–C thus revealed that power law behaviors could be consistently observed during resting-state and during task, in

⁵ <https://github.com/neurospin/mfanalysis>

⁶ https://github.com/neurospin/dynacomf_mf_analysis

⁴ <http://www.ens-lyon.fr/PHYSIQUE/Equipe3/Multifractal/software.html>

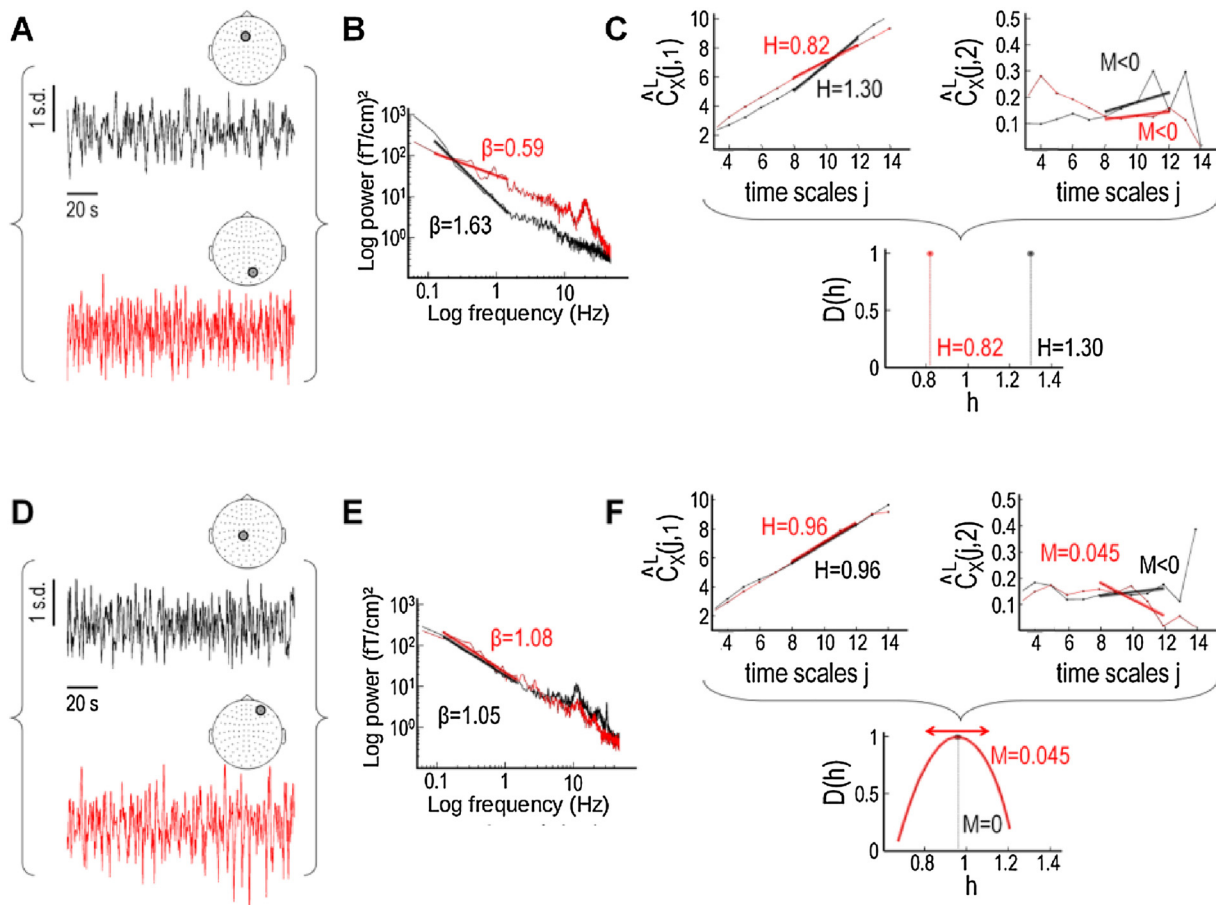


Fig. 3. Self-similarity and multifractality in spontaneous infraslow brain activity at rest. Normalized time series representing an individual's brain activity recorded in two different pairs of MEG sensors at rest (A–C) and during task (D–F). The pairs of sensors in rest and task conditions were selected in order to show two extreme examples where different self-similarity (but no multifractality) or different multifractality (but same self-similarity) can be observed, respectively. In (A–C), the black and red traces characterize frontal and occipital MEG sensors (magnetometers) at rest, respectively, as depicted in the topographies. In (D–F), the same colors are used to refer to central and right frontal MEG sensors (magnetometers) during task. In (A–C), the two signals show a difference in *self-similarity*. In (D–F), the two signals show differences in *multifractality* but no differences in self-similarity. Plots (B,E) correspond to periodogram based spectral estimation (cf. Section 2.2.3). Plots (C,F) correspond to wavelet-leader analysis (as detailed in Section 2.2.2). In particular, on top of panels (C,F), the quantities $C_1(j)$ and $C_2(j)$ defined in Eqs. (3) and (4) are computed as a function of time scale j . The H and M indexes are derived from these quantities using a regression analysis performed over the scaling range $(j_1, j_2) = (8, 12)$ which matched the (0.1, 1.5) Hz frequency range ($j = 8$ corresponds to 1.5 Hz and $j = 12$ corresponds to 0.1 Hz). The latter was used in panels (B,E) for estimating the β values. The multifractal spectra $D(h)$ are reported at the bottom of panels (C,F). (For interpretation of the references to color in this figure legend, the reader is referred to the web version of the article.)

the range of octaves $(j_1, j_2) = (8, 12)$. This range is associated with frequencies $\simeq(0.1, 1.5)$ Hz or, equivalently, with time scales $(2^j = 8/f_s, 2^j = 12/f_s) \simeq (0.66, 10)$ s.

To further provide an intuitive understanding of scale-free dynamics, we compared the Fourier and wavelet spectra of normalized MEG time courses (Fig. 3). We selected two different pairs of sensors for one participant at rest (Fig. 3A–C) and during task (Fig. 3D–F) in order to illustrate two extreme case examples. In Fig. 3A, the occipital sensor time series (red) appeared visually less structured over time (i.e., closer to white noise behavior) than the frontal sensor time series (black). This visual appreciation was quantified using a classic Fourier analysis showing a power law behavior at low frequencies ($P(f) \simeq 1/f^\beta$), with a scaling exponent β that was smaller for the occipital sensor (red trace) (Fig. 3B). Conversely, we found that the frontal sensor (black) showed a larger scaling exponent, hence a steeper slope, or equivalently, a stronger temporal autocorrelation, and was quantified by a stronger long-range dependency (Fig. 3B). The same analysis held when conducted from the wavelet spectrum (Fig. 3C), with very satisfactory matches for the estimated scaling exponents H and β according to $\beta = 2H - 1$.

3.2. Beyond self-similarity in MEG data by assessing multifractality

To demonstrate the interest of a multifractal description in brain activity, we first report the analysis of the same MEG time series recorded at rest already used in the previous Section (Fig. 3A). Multifractal analysis yielded negative estimates of the multifractality parameter (Fig. 3C): $M \leq 0$ indicating no multifractality at rest. This can be visualized by the δ -shape of the multifractal spectra $D(h)$, which only differed by their location on the h -axis, reflecting different self-similarity exponents H . In other words, for these time series, there was no additional information provided by using multifractal analysis.

We then repeated the same analysis on two MEG time-series collected during task (Fig. 3D). These case study time-series were chosen on purpose because they showed very similar Fourier spectra (Fig. 3E), hence displaying the same β s, and predictably, the same held true for the wavelet spectra (Fig. 3F) showing the same H s. Interestingly however, the frontal signal (red) appeared far more irregular and locally bursty than the central one (black). This was again quantified using multifractal analysis (Fig. 3F), which revealed that although H (top left) was identical in both frontal and central time series, the frontal time series was characterized by a positive $M = 0.045 > 0$ (hence,

displayed multifractality), while the central sensor did not ($M < 0$). The multifractal spectra for both time series thus summarized the two case study observations: whereas the location of their peaks coincided (same H), only one spectrum (red) showed a large parabola shape ($M > 0$).

These examples were chosen as pedagogical illustrations of the potential richness of scale-free temporal dynamics found in brain time series. Specifically, while the typical power spectrum analysis would conclude that these different time series share the same scale-free characteristics, multifractal analysis clearly showed differences in their temporal dynamics by quantifying the existence of transient and local irregularities observed in the frontal sensor (red) that did not exist in the central sensor data (black). Multifractality thus complements self-similarity in the characterization of scale-free dynamics in time series by quantifying local transient dynamics that are not well accounted for by the autocorrelation or by the Fourier spectrum.

3.3. Group-level analysis of scale-free brain dynamics

Having extended the framework for the assessment of scale-free brain dynamics to self-similarity H and multifractality M , we then proceeded with a comprehensive analysis of scale-free brain activity across all individuals ($n = 24$). For this, we assessed scale free activity in source reconstructed time series averaged within each cortical regions (see Section 2.1.6). The estimation of parameters H and M relied on the wavelet-leader multifractal formalism described in Section 2.2.2: i.e., MEG wavelet-leader spectra $C_1(j)$ and $C_2(j)$ were systematically computed on resting-state and task recordings separately for each cortical label and on a *per* individual basis. Results were then averaged across individuals to form $\bar{C}_1(j)$ and $\bar{C}_2(j)$, respectively. Importantly, since linear regression and group-level averaging were both linear after taking log in Eqs. (3) and (4), we could interchange them without impacting the results. For this reason, in what follows, we illustrated group-level values of $\bar{C}_1(j)$ and $\bar{C}_2(j)$ in log-scale diagrams from which we deduced the respective group-level H and M . The latter actually matched the group-level averages of subject-specific values of H and M , shown in the cortical maps of Figs. 4 and 5.

3.4. A fronto-occipital gradient of self-similarity

Fig. 4A reports the grand average $\bar{C}_1(j)$ for resting-state obtained in two cortical labels (one frontal in red, one occipital in blue). The self-similarity exponent H was found to be larger in the frontal label as compared to the occipital label. To systematically quantify this effect, the calculation of $\bar{C}_1(j)$ was conducted over the whole cortical surface. Using T -statistics, the null hypothesis $H = 0.5$ was tested at the group level. To account for multiple comparisons across the 138 labels covering the whole cortical surface, a correction was implemented using the false discovery rate (FDR) detection at $\alpha = 0.05$: $p_{\text{corrected}} < 0.05$. Fig. 4C reports the spatial distribution of statistically significant mean values of H ($H > 0.5$), yielding a key finding: the spatial distribution of estimated H s during rest revealed a fronto-occipital gradient, in which H significantly decreased from frontal ($H \simeq 1.2$) to occipital regions ($H \simeq 0.8$ – 0.9). This gradient was consistent with prior observations of scale-free activity observed in MEG and EEG recordings (Dehghani et al., 2010): larger H in frontal regions (i.e., steeper slopes for the spectra) would indicate stronger and longer temporal correlations, i.e. more-structured temporal dynamics, compared to occipital regions.

3.5. During task, an overall decrease of self-similarity accentuates the fronto-occipital gradient

In Fig. 4D, the spatial distribution of H significantly departed away from 0.5 during task yielding, by comparison to rest, another key finding: the decrease of H during task appeared to be global and almost significant everywhere over the cortical surface. Interestingly, the

anatomical fronto-occipital gradient at rest appeared to be further strengthened during perceptual task completion (cf. lateral views in Fig. 4D). We contrasted the H parameter estimates between rest and task using paired t -tests. FDR was applied to correct for multiple comparisons across cortical labels at $\alpha = 0.05$. In Fig. 4E, the statistical assessment of changes in H between rest and task confirmed our qualitative appreciation. Specifically, H was significantly diminished during task in numerous cortical regions including occipital, parietal, and primary motor cortices as well as right supplementary area (SMA) and ventrolateral prefrontal cortex (vlPFC) bilaterally. All these regions were previously shown to be essential in the perceptual task participants were engaged in Zilber et al. (2014).

3.6. Weak multifractality in resting-state

The group-average $\bar{C}_2(j)$ at rest for two cortical labels is illustrated in Fig. 5A. We found no multifractality ($M < 0$) in the frontal label (red) but found multifractality ($M = 0.017$) in the occipital label (blue). As previously done for estimates of H , we performed the analysis of multifractality at rest over the whole cortical surface. Using T -statistics, the null hypothesis $M = 0$ was tested at the group-level. The same FDR correction at $\alpha = 0.05$ was applied to correct for multiple comparisons across labels. Fig. 5C reports the spatial distribution of statistically significant mean values of M ($M > 0$). At rest, the presence of multifractality was confined to a few regions: the posterior superior temporal sulci, the occipital cortex, the right temporo-parietal junction and the frontal cortices, bilaterally. The observed values of M mostly ranged between 0.01 and 0.02, with the exception of the frontal poles which reached $M = 0.03$.

3.7. Multifractality: localized increase in regions engaged in the task

During task, the group-level wavelet-leader spectra $\bar{C}_2(j)$ (Fig. 5B) suggested an absence of multifractality in frontal regions ($M = 0$) but an increase of multifractality in occipital regions. M was increased by about 60% in occipital cortices, hence showing steeper slopes for $\bar{C}_2(j)$. A statistical assessment of multifractality over the whole cortical surface during task revealed a spatially extended set of cortical regions showing significant multifractality (Fig. 5D; mean values of M ($M > 0$)). Additionally, the multifractal parameter M took overall larger values compared to the distribution we had observed during resting-state. We notably found larger M values in cortical regions involved in the perceptual task participants were engaged in Zilber et al. (2014), namely: visual cortices (primary, secondary and visual motion region (hMT+)) as well as parietal cortices and the posterior superior temporal sulci.

When statistically contrasting the M parameter estimates between rest and task (paired t -tests, FDR correction for multiple comparisons across cortical labels at $\alpha = 0.05$), we found significant changes in M in brain regions including the right occipital cortices, SMA, the left temporo-parietal junction and posterior cingulate cortex (Fig. 5E). These changes corresponded to an increase in multifractality, though they remained limited in magnitude to a maximal increase of 0.01 (to be compared to an average value of the order of 0.02). This was a remarkable observation considering that the analysis was conducted over the whole cortical surface with no *a priori* restriction on the timing of the stimuli or cognitive operations implicated in the decision-making; rather, our analysis was performed over the whole time series.

A supplementary analysis of high-pass filtered data (cut-off frequency of 0.1 Hz) was performed to exclude any effect of artifacts overlapping on the infraslow frequency range (see Fig. S1). Furthermore, the passive Localizer recording, which was implemented to localize the hMT+ visual motion area in each individual (see (Zilber et al., 2014) for details), was analyzed to exclude the impact of a mere motor response in the modulation of scale-free parameters (see Figure S1B). Both analyses showed similar cortical distributions of parameters

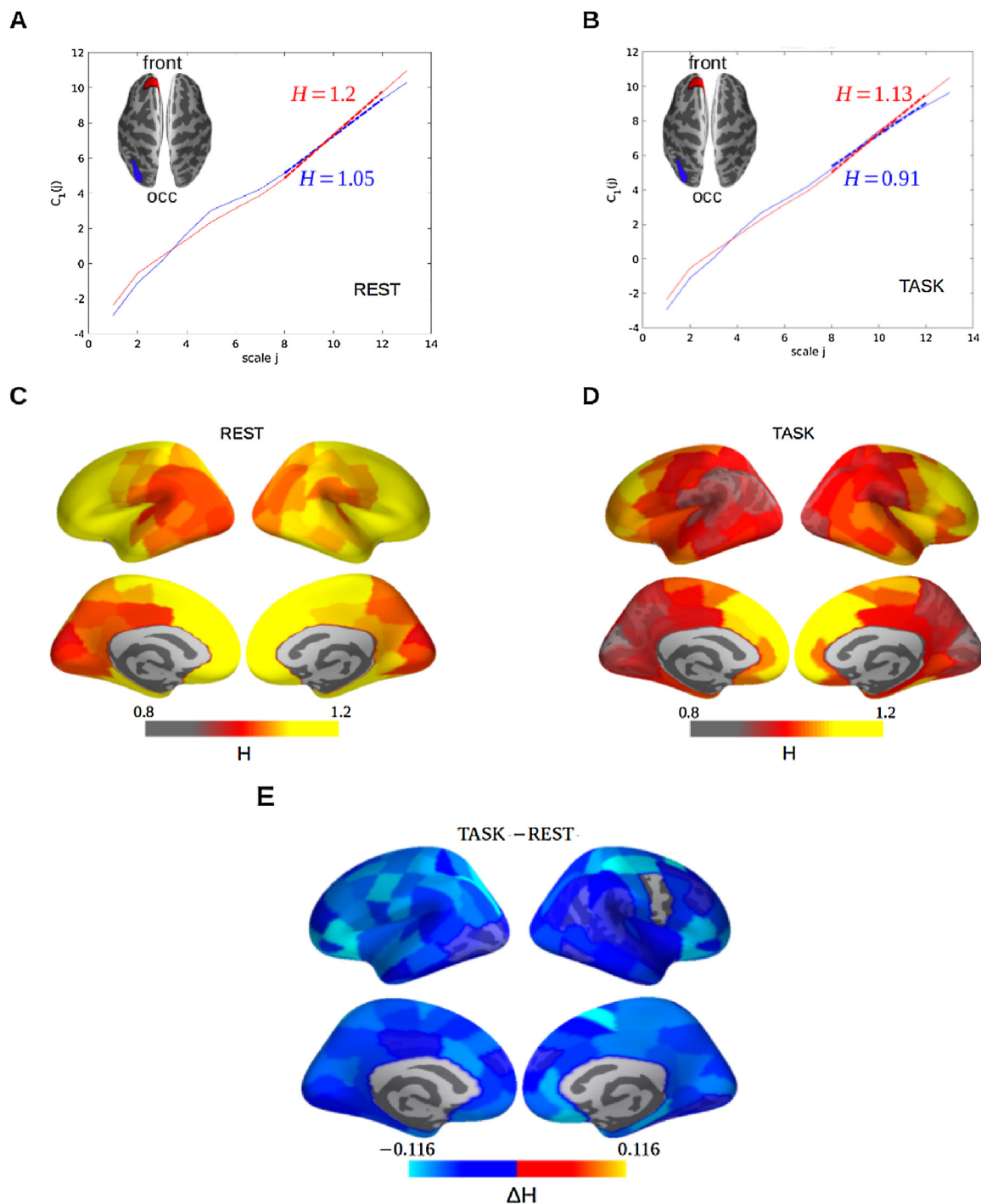


Fig. 4. Fronto-occipital gradient of self-similarity. For comparison with Fig. 2, we show group-average wavelet-leader structure functions $\overline{C}_1^*(j)$ in the same frontal (red) and occipital (blue) cortical labels during rest (A) and task (B) blocks. The linear fits were computed over the scaling range $8 \leq j \leq 12$ and matched the (0.1, 1.5) Hz frequency range used before for linear regression in the power spectra. The associated slopes provides estimates of group-level Hurst exponents H . (C and D) Group-average cortical maps (lateral and medial views on top and bottom, respectively, left hemisphere on the left) of Hurst exponents H at rest and during task, respectively. In both rest and task, a fronto-occipital gradient of self-similarity could be observed going from higher H in frontal regions to lower H in parieto-occipital regions). (E) Cortical maps contrasting H in task and resting-state testing the null hypothesis that $H_{\text{TASK}} = H_{\text{REST}}$. The statistical significance was assessed on a per label basis by computing a paired Student t -test and correcting for multiple comparisons with FDR at $\alpha = 0.05$. Estimates of H were smaller in task than in rest as shown by negative differences ($\Delta H = H_{\text{TASK}} - H_{\text{REST}} < 0$). This contrast indicated that, globally, self-similarity significantly decreased when participants performed a task as compared to when they rested. (For interpretation of the references to color in this figure legend, the reader is referred to the web version of the article.)

H and M to those shown in Figs. 4 and 5. Altogether, these results suggest that multifractal characterization of brain activity may capture relevant signatures of brain processing that are associated with task-relevant brain regions.

3.8. Covariation of self-similarity and multifractality from rest to task

So far, we reported significant differences for both H and M when contrasting rest and task, namely: while self-similarity H significantly decreased in task as compared to rest (Fig. 4E), multifractality M

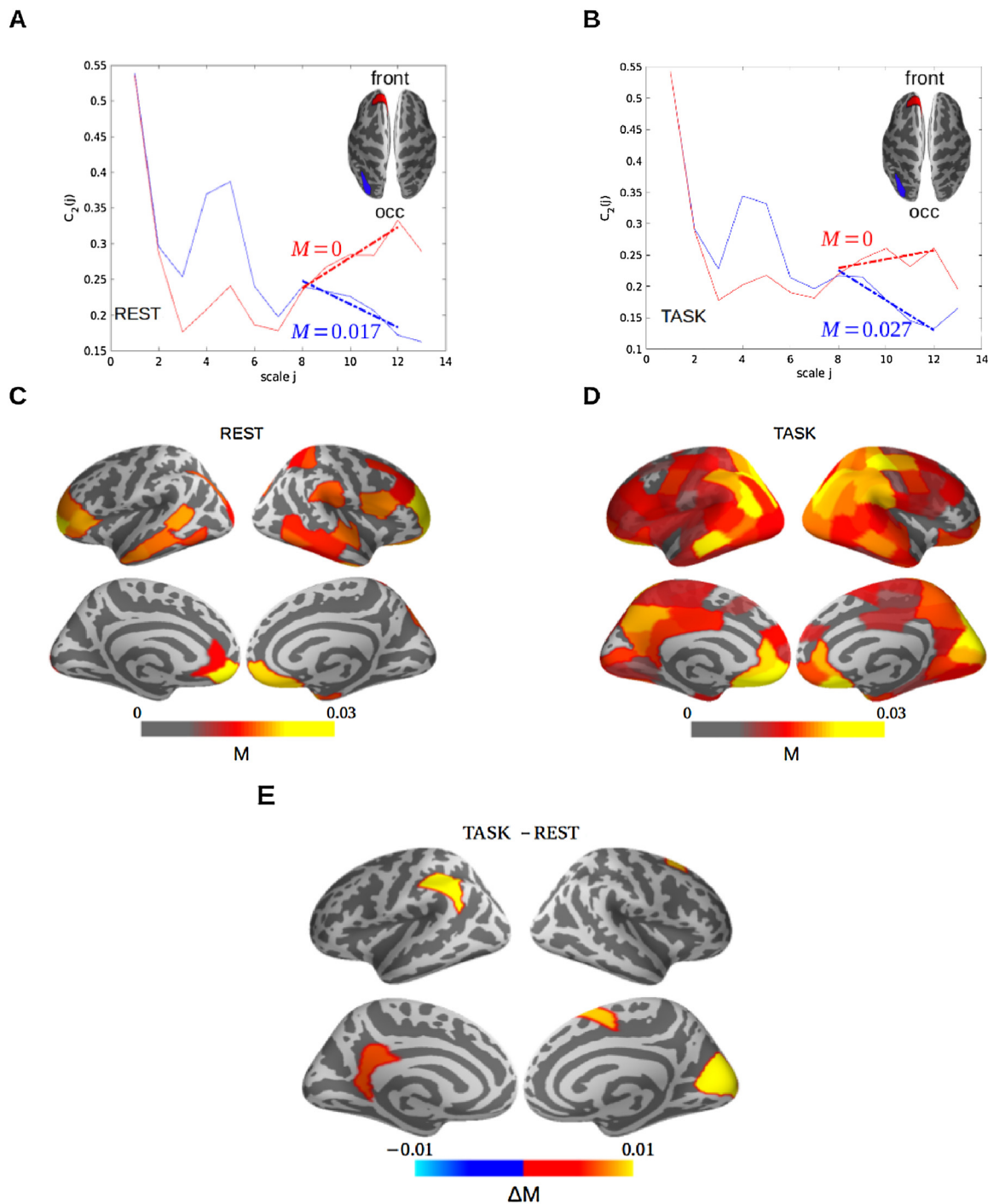


Fig. 5. Multifractal brain activity. Group-average wavelet-leader structure functions $\bar{C}_2(j)$ in the same frontal (red) and occipital (blue) labels at rest (A) and during task (B). The linear fits were computed over the scaling range $8 \leq j \leq 12$ and matched the (0.1, 1.5) Hz frequency range used before. The associated slopes provided estimates of the multifractal exponents M . (C and D) Statistically significant ($H_0: M = 0$) grand-average cortical maps of multifractal exponents M at rest and during task, showing sparser topographies than for self-similarity H , especially at rest. (E) Cortical maps contrasting M in task and resting-state testing the null hypothesis that $M_{TASK} = M_{REST}$. The statistical significance was assessed on a per label basis by computing a paired Student t -test and correcting for multiple comparisons with FDR at $\alpha = 0.05$. Estimates of M were bigger in task than in rest as shown by positive differences ($\Delta M = M_{TASK} - M_{REST} < 0$) in several regions involved in the task, notably visual, parietal and motor cortices. This contrast indicated that, locally, multifractality significantly increased when participants performed a task as compared to when they rested. (For interpretation of the references to color in this figure legend, the reader is referred to the web version of the article.)

significantly increased in task as compared to rest (Fig. 5E). Additionally, the changes in M were confined to a limited number of brain regions whereas the changes observed in H were more global, thereby yielding a global accentuation of the fronto-occipital gradient. Considering the possible overlap of cortical regions displaying both changes, we then asked to which extent the two characteristics of scale-

free dynamics may be related. We correlated the changes in H and M from rest to task on a label-by-label basis (Fig. 6). This analysis revealed that in some of the cortical regions showing task-related multifractality (Fig. 6A), there was a significant negative correlation between individual changes from rest to task of H ($\Delta H = H_{TASK} - H_{REST}$) and M ($\Delta M = M_{TASK} - M_{REST}$) (Fig. 6B).

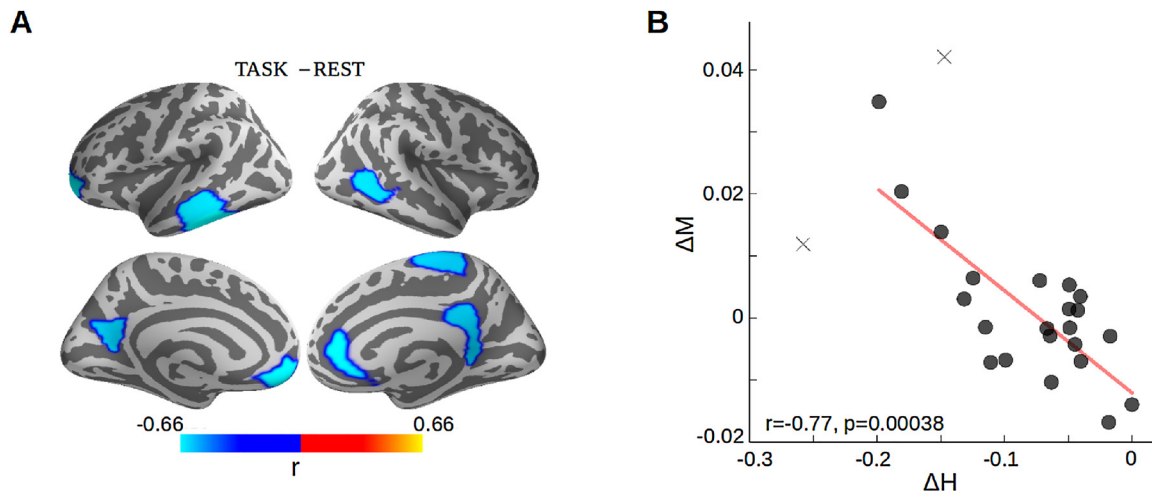


Fig. 6. Covariations of self-similarity and multifractality. Significant negative correlation between changes of H ($\Delta H = H_{\text{TASK}} - H_{\text{REST}}$) and M ($\Delta M = M_{\text{TASK}} - M_{\text{REST}}$). (A) Cortical source estimates associated with significant correlations (color-coded Pearson's r values) observed between ΔH and ΔM . The null hypothesis reads $r = 0$ in each cortical label. Statistical significance was assessed on a per individual and per label basis by computing a paired Student t -test. Corrections for multiple comparisons were performed using FDR at $\alpha = 0.05$. (B) Scatter plot of ΔH versus ΔM , averaged over all brain regions reported in (A). The significant negative correlation indicated concomitant local decreases of H (negative ΔH) and increase of M (positive ΔM). Each dots is an individual and crosses are outliers ($n = 24$). The outliers were automatically excluded as their distance to the mean was $> 2.5\sigma$ to the model.

These results constituted a particularly important finding: theoretically, H and M are independent parameters, which model very different aspects of scale-free dynamics. While self-similarity H provides insights on the temporal autocorrelation of brain activity, M informs on the burstiness of the signals. The observed negative covariation between self-similarity and multifractality is non-trivial and crucially suggests a potential coupling in the covariation of both indices. We discussed these findings further below.

4. Discussion

To briefly sum up, our key findings are: the existence of a fronto-occipital gradient of self-similarity in the human brain which increases during task as compared to rest. Second, and to the best of our knowledge, we observed for the first time multifractality on MEG data collected in a healthy human population and describe an anatomical distribution of multifractality during resting-state and task. Local changes in multifractality in task as compared to rest indicate a possible functional relevance of multifractal infra-slow dynamics in brain processing. Our empirical results raise several points of discussions and conclusions regarding the assessment of scale-free temporal brain dynamics. We discuss the main ones below.

4.1. Robust description of infraslow, scale-free dynamics, in human brain activity

Overall, our results support the notion that scale-free temporal dynamics constitute a signature of human brain activity as recorded with MEG. We showed that scale-free properties were neither induced by, nor to be confused with, instrumental or sensor noise considering that empty-room recordings did not display the same characteristics. Scale-free dynamics were observed in a range of frequencies corresponding to $0.1 \leq f \leq 1.5$ Hz. Such time scales are consistent with currently available data in the literature for the estimations of H or β (Monto et al., 2008; He et al., 2010; He, 2014; Becker et al., 2018), and typically characterize infra-slow neural dynamics (Wang, 2010; Buzsáki and Mizuseki, 2014; He, 2014). Scale-free temporal dynamics conceptually implies that, within the scaling range, no frequency plays a particular role. Conversely, all frequencies in that range contribute jointly and in a related manner to the described dynamics. Such relation is quantified by β , or in a richer framework which we propose here, by the joint

description of H and M parameters. Scale-free temporal dynamics thus correspond to arrhythmic signatures confined to infra-slow brain dynamics, which complement oscillatory activity typically seen at higher frequencies (> 2 Hz) (Wang, 2010; Buzsáki and Mizuseki, 2014; He, 2014). It is also noteworthy that while the concept of scale-free temporal dynamics theoretically implies the absence of any specific time scale, in practice, scale-free analyses have covered a finite range of frequencies, $0.1 \leq f \leq 1.5$ Hz.

We thus propose that the classical parameter β used to model scale-free dynamics as a power-law decay of the Fourier spectrum can be efficiently replaced by the self-similarity parameter H , which models the decay of the wavelet spectrum. While both exponents are theoretically equivalent and related ($\beta = 2H - 1$), it has been well documented in the scientific literature that H and wavelet analysis benefit from improved estimation performance (robustness to smooth non-stationarities) (Abry and Veitch, 1998; Veitch and Abry, 1999; Ciuciu et al., 2012, 2014). The larger H (or β) – i.e., the steeper the decay of power laws –, the more structured the temporal dynamics of the time series – i.e., the stronger the long range dependency quantified by the temporal correlations.

4.2. Multifractality: going beyond self-similarity

In addition to self-similarity, our results demonstrated the existence of multifractality while participants performed a task and, to a lesser extent, during resting-state. This specific result underline the richness of infraslow brain dynamics and of the usefulness of the framework we propose to characterize scale-free brain activity. Specifically, we showed that multifractality allows distinguishing time series that share the same global correlation structure (i.e., the same self-similarity) but different local transient structures and burstiness over time (i.e., multifractality). In other words, multifractality quantifies local scale-free temporal dynamics as transient departures from Gaussianity and support the recent mention that multiplicative processes should be taken into account in the assessment of macroscopic brain activity (Buzsáki and Mizuseki, 2014). Hence, our approach is key to proper scale-free modeling although it remains seldom discussed in the neuroscience literature (Shimizu et al., 2004; Suckling et al., 2008; Ciuciu et al., 2012). Additionally, and from a signal processing perspective, our results suggest that the multifractal random walk, which consists of a multifractal extension of fractional Gaussian noise (cf. Section 2.2.1 for

details), is a likely more accurate model to describe spontaneous brain activity in the infraslow regime (< 2 Hz).

4.3. Anatomical distribution of self-similarity and multifractality in resting-state activity

The spatial distributions of self-similarity and multifractality quantified at rest and during task were obtained using the theoretically robust and practically efficient wavelet-leader multifractal framework (Wendt et al., 2007). With this approach, we observed a fronto-occipital gradient of the self-similarity parameter H in resting-state. This observation was congruent with previous findings in the literature (Dehghani et al., 2010; Becker et al., 2018), but also extended them from scalp level to cortical source estimates. The fronto-occipital gradient corresponded to larger values of self-similarity in frontal regions and lower values in posterior regions. This pattern converges with the known distribution of temporal scales at which neural processing operate: a recent meta-analysis has notably showed a hierarchy of intrinsic time-scales going from slower dynamics in frontal to faster dynamics in sensory cortices (Murray et al., 2014). Comparable temporal hierarchies have been functionally described in the human visual system (Gauthier et al., 2012) and across brain systems (Hasson et al., 2015). These temporal hierarchies are functionally compatible with finer time scales needed for sensory sampling, and integrative processes over longer time scales occurring in frontal cortices for higher cognitive operations (Fuster, 2001; Miller and Cohen, 2001; Wood and Grafman, 2003). By indexing the anatomical distribution of temporal autocorrelation functions, the fronto-occipital gradient in H provides an alternative means to characterize the hierarchy of temporal scales in cortex.

Additionally, during resting-state, the presence of weak multifractality, naturally one order smaller than values of H , is consistent with well-behaved multifractal synthetic models (Wendt et al., 2007) or values reported for brain data (Shimizu et al., 2004, 2007; Suckling et al., 2008; Ciuciu et al., 2008, 2012, 2017; Weiss et al., 2009). The presence of M , especially during task performance, suggested that multifractality may be a relevant index for brain processing.

4.4. Global decrease of self-similarity from rest to task

By contrasting brain activity during engagement in a task against resting-state, we observed a general decrease of H over the whole cortex, suggesting an overall and global shortening of temporal autocorrelation during task performance. Additionally, the decrease in self-similarity was not uniform across brain regions, which contributed to the strengthening of the fronto-occipital gradient. In other words, relatively less short-time dynamics were found in frontal regions and more short-time dynamics were observed in posterior regions during task than during rest. The accentuation of the fronto-occipital gradient in H between rest and task is overall consistent with faster and richer dynamics deployed for the analysis of sensory information in cortical regions engaged in the task (Palva and Palva, 2012). This observation also converges with previous fMRI studies showing a lower regional H during task than during resting-state (He, 2011; Ciuciu et al., 2012) and stronger decreases of H with higher cognitive loads (Chang et al., 2012). The most salient differences of self-similarity were observed in regions involved in the task (occipital cortex, motor cortex, SMA and vPFC), *i.e.* the decrease in H was the largest in these regions. This observation is in line with the hypothesis that self-similarity may quantify neural excitability, with smaller values of self-similarity indexing higher levels of neuronal excitability in a given brain region (He et al., 2010; He, 2011; Palva and Palva, 2012; Palva et al., 2013).

4.5. Local increase of multifractality from rest to task

Although we found a large number of cortical labels showing a

significant presence of multifractality during task, contrasting task against rest revealed increases of M in only a small number of cortical regions. The relatively small changes of M in magnitude, the limited sample size (*i.e.* 24 individuals only) and the potentially large inter-individual variability may explain why only a fraction of cortical regions were reported as statistically significant in the paired t -test. Nevertheless, the presence of the highest M values in regions (occipitoparietal cortices, visual motion area, pSTS) involved in the visual motion discrimination task used here (Zilber et al., 2014) suggests that multifractality might be functionally relevant to cortical processing. The local changes of multifractality would be consistent with the notion that multifractality may reflect the combination of multiplexed self-similar processes, *i.e.* the superimposition of several self-similar processes associated with different neural populations within the same cortical patch (given the limits of the spatial resolution with MEG). As such, one working hypothesis for multifractality in brain processes is that it may index the number of neural processes within a cortical region employed in a given task. This working hypothesis will be actively investigated.

4.6. Covariation of self-similarity and multifractality from rest to task

We evidence an interesting covariation pattern in self-similarity and multifractality from rest to task: the MEG brain dynamics evolved from well structured and long term correlated global temporal dynamics (large H) with weak burstiness ($M \approx 0$, weak multifractality) at rest, to less structured global temporal dynamics (lower H , lesser long range dependence, or more power at the upper bound of the scaling range, *i.e.*, around 1Hz) during task performance, showing though much larger transient irregular and non Gaussian behaviors (larger M , multifractality). Let us emphasize that this covariation (decrease in H , increase in M) was non trivial and was not induced by the modeling nor by the analysis we undertook. This covariation thus constitutes a signature of the changes induced in brain dynamics when participants engaged in a perceptual discrimination task.

Our tentative explanation for this covariation is the following: the local decrease of temporal autocorrelation (H) suggests that neural populations in a given cortical region *and* at a large temporal scale (lower infra-slow *i.e.* ≈ 10 s) tend to operate more independently while, at the same time, the increase of temporal burstiness (M) in the same region suggests that the same neural populations may interact at finer temporal scales (higher infra-slow, *i.e.* ≈ 1 s). Distinct dynamic modes may thus take place as a function of task requirements: while neural excitability may be sufficient to detect the presence/absence of a stimulus in the environment (He et al., 2010; He, 2011; Monto et al., 2008), temporal multiplexing may be required for thorough analysis of sensory inputs. In other words, temporal multiplexing may occur when a certain level of neural excitability has been reached. However, both a multivariate approach and activity from specific neural populations (*e.g.* recorded via intracranial electrodes) should be explored to further investigate this hypothesis in future work.

5. Conclusions

Relying on the robust and efficient wavelet and wavelet-leader analysis framework, our present contribution showed that multifractality provides a fruitful paradigm to complement self-similarity in the modeling of scale-free temporal dynamics and infraslow macroscopic brain activity. We showed that spontaneous human brain activity at rest is well characterized by a strong self-similarity and weak multifractality, indicating a significantly globally-structured activity, with long range dependencies. The strength of this structured activity showed a fronto-occipital gradient. We showed that performing a task induced a non trivial (negatively correlated) local coupling of self-similarity and multifractality with an overall decrease of self-similarity (yet, strengthening of the fronto-occipital gradient) accompanied by a local increase of multifractality in task-relevant brain regions. Overall,

this pattern indicates less structured (or less correlated) temporal dynamics yet bursty occurrences of well-structured local scale-free patterns (not accounted for by self-similarity but well quantified by multifractality). Altogether, these observations support the hypothesis that (i) self-similarity, as indexed by parameter H , inversely reflects neural excitability, with large H corresponding to lower excitability and vice versa and that (ii) multifractality, indexed by M , might code for multiplexing of neural processes.

The present analysis of scale-free dynamics in brain temporal dynamics will be continued by exploring the benefits of using more refined analysis tools based on p -leaders (Leonarduzzi et al., 2017) or on multivariate models, rather than univariate, for self-similarity (Abry et al., 2018) and multifractality (Jaffard et al., 2018; Ciuciu et al., 2017).

Appendix A. Supplementary data

Supplementary data associated with this article can be found, in the online version, at <https://doi.org/10.1016/j.jneumeth.2018.09.010>.

References

- Abry, P., Veitch, D., 1998. Wavelet analysis of long-range-dependent traffic. *IEEE Trans. Inf. Theory* 44 (1), 2–15.
- Abry, P., Didier, G., et al., 2018. Wavelet estimation for operator fractional Brownian motion. *Bernoulli* 24 (2), 895–928.
- Arenas, A., Díaz-Guilera, A., Kurths, J., Moreno, Y., Zhou, C., 2008. Synchronization in complex networks. *Phys Rep* 469 (3), 93–153.
- Bacry, E., Delour, J., Muzy, J.-F., 2001. Multifractal random walk. *Phys Rev E* 64 (2), 2–5.
- Baranauskas, G., Maggolini, E., Vato, A., Angotzi, G., Bonfanti, A., Zambra, G., Spinelli, A., Fadiga, L., 2012. Origins of $1/f^2$ scaling in the power spectrum of intracortical local field potential. *J. Neurophysiol.* 107 (3), 984–994.
- Bardet, J.-M., Lang, G., Oppenheim, G., Philippe, A., Stoev, S., Taqqu, M.S., 2003. Semi-parametric estimation of the long-range dependence parameter: a survey. *Theory and Applications of Long-Range Dependence*. Birkhäuser, pp. 557–577.
- Bartfeld, P., Uhrig, L., Sitt, J., Sigman, M., Jarraya, B., Dehaene, S., 2015. Signature of consciousness in the dynamics of resting-state brain activity. *Proc. Natl. Acad. Sci. U.S.A.* 112 (3), 887–892.
- Baykut, S., Ozkurt, T., Erol, M., Akgul, T., 2005. The influence of a single-tone sinusoid over Hurst estimators. 13th European Signal Processing Conference.
- Becker, R., Van De Ville, D., Kleinschmidt, A., 2018. Alpha oscillations reduce temporal long-range dependence in spontaneous human brain activity. *J. Neurosci.* 38 (3), 755–764.
- Bedard, C., Kroeger, H., Destexhe, A., 2006. Does the $1/f$ frequency scaling of brain signals reflect self-organized critical states? *Phys. Rev. Lett.* 97 (11), 118102.
- Breakspear, M., 2017. Dynamic models of large-scale brain activity. *Nat. Neurosci.* 20 (3), 340–352.
- Buiatti, M., Papo, D., Baudonnière, P.-M., van Vreeswijk, C., 2007. Feedback modulates the temporal scale-free dynamics of brain electrical activity in a hypothesis testing task. *Neuroscience* 146 (3), 1400–1412.
- Buzsáki, G., Mizuseki, K., 2014. The log-dynamic brain: how skewed distributions affect network operations. *Nat. Rev. Neurosci.* 15 (4), 264–278.
- Buzsáki, G., Anastassiou, C.A., Koch, C., 2012. The origin of extracellular fields and currents – EEG, ECoG, LFP and spikes. *Nat. Rev. Neurosci.* 13 (6), 407–420.
- Buzsáki, G., 2010. Neural syntax: cell assemblies, synapses, and readers. *Neuron* 68 (3), 362–385.
- Chang, C., He, B.J., Duyn, J., 2012. Dependence of fMRI scale-free dynamics on cognitive load and temporal preprocessing. 18th Annual Meeting of the Organization for Human Brain Mapping 689.
- Chaudhuri, R., He, B.J., Wang, X.-J., 2017. Random recurrent networks near criticality capture the broadband power distribution of human ECoG dynamics. *Cereb. Cortex* 1–13.
- Ciuciu, P., Abry, P., Rabrait, C., Wendt, H., 2008. Log wavelet leaders cumulant based multifractal analysis of EVI fMRI time series: evidence of scaling in ongoing and evoked brain activity. *IEEE J. Sel. Top. Signal Process.* 2 (6), 929–943.
- Ciuciu, P., Varoquaux, G., Abry, P., Sadaghiani, S., Kleinschmidt, A., 2012. Scale-free and multifractal time dynamics of fMRI signals during rest and task. *Front. Physiol.* 3 Article 186.
- Ciuciu, P., Abry, P., He, B.J., 2014. Interplay between functional connectivity and scale-free dynamics in intrinsic fMRI networks. *Neuroimage* 95, 248–263.
- Ciuciu, P., Wendt, H., Combexelle, S., Abry, P., 2017. Spatially regularized multifractal analysis for fMRI data. *Engineering in Medicine and Biology Society (EMBC), 2017 39th Annual International Conference of the IEEE, IEEE 3769–3772*.
- da Silva, F.L., 2013. EEG and MEG: relevance to neuroscience. *Neuron* 80 (5), 1112–1128.
- Dale, A.M., Fischl, B., Sereno, M.I., 1999. Cortical surface-based analysis. I. Segmentation and surface reconstruction. *Neuroimage* 9 (2), 179–194.
- Dehghani, N., Bedard, C., Cash, S.S., Halgren, E., Destexhe, A., 2010. Comparative power spectral analysis of simultaneous electroencephalographic and magnetoencephalographic recordings in humans suggests non-resistive extracellular media. *J. Comput. Neurosci.* 29 (3), 405–421.
- Dehghani, N., Hatsopoulos, N.G., Hagaz, Z.D., Parkers, R.A., Greger, B., 2012. During wakefulness and sleep. *Scale-free Dynamics and Critical Phenomena in Cortical Activity*. pp. 21.
- Dehghani, N., 2012. Electromagnetic signature of human cortical dynamics during wakefulness and sleep signature électromagnétique de la dynamique corticale pendant l'éveil et le sommeil chez l'homme (Ph.D. thesis). Citeseer.
- Destexhe, A., Contreras, D., Steriade, M., 1999. Spatiotemporal analysis of local field potentials and unit discharges in cat cerebral cortex during natural wake and sleep states. *J. Neurosci.* 19 (11), 4595–4608.
- Engel, A.K., Fries, P., Singer, W., 2001. Dynamic predictions: oscillations and synchrony in top-down processing. *Nat. Rev. Neurosci.* 2 (10), 704–716.
- Fischl, B., Sereno, M.I., Dale, A.M., 1999a. Cortical surface-based analysis. II: Inflation, flattening, and a surface-based coordinate system. *Neuroimage* 9 (2), 195–207.
- Fischl, B., Sereno, M.I., Tootell, R.B., Dale, A.M., 1999b. High-resolution intersubject averaging and a coordinate system for the cortical surface. *Hum. Brain Mapp.* 8 (4), 272–284.
- Foster, B.L., He, B.J., Honey, C.J., Jerbi, K., Maier, A., Saalman, Y.B., 2016. Spontaneous neural dynamics and multi-scale network organization. *Front. Syst. Neurosci.* 10.
- Freeman, W.J., 2000. Mesoscopic neurodynamics: from neuron to brain. *J. Physiol. Paris* 94 (5–6), 303–322.
- Fries, P., 2005. A mechanism for cognitive dynamics: neuronal communication through neuronal coherence. *Trends Cogn. Sci.* 9 (10), 474–480.
- Fuster, J.M., 2001. The prefrontal cortex—an update: time is of the essence. *Neuron* 30 (2), 319–333.
- Gauthier, B., Eger, E., Hesselmann, G., Giraud, A.-L., Kleinschmidt, A., 2012. Temporal tuning properties along the human ventral visual stream. *J. Neurosci.* 32 (41), 14433–14441.
- Gramfort, A., Luessi, M., Larson, E., Engemann, D.A., Strohmeier, D., Brodbeck, C., Hamalainen, M.S., 2014. MEG and EEG data analysis with MNE-Python. *Neuroimage* 86, 446–460.
- Gross, J., Baillet, S., Barnes, G.R., Henson, R.N., Hillebrand, A., Jensen, O., Schoffelen, J.M., 2013. Good practice for conducting and reporting MEG research. *Neuroimage* 65, 349–363.
- Hämäläinen, M.S., Sarvas, J., 1989. Realistic conductivity geometry model of the human head for interpretation of neuromagnetic data. *IEEE Trans. Biomed. Eng.* 36 (2), 165–171.
- Hardstone, R., Poil, S.S., Schiavone, G., Jansen, R., Nikulin, V.V., Mansvelder, H.D., Linkenkaer-Hansen, K., 2012. Detrended fluctuation analysis: a scale-free view on neuronal oscillations. *Front. Physiol.* 3.
- Hasson, U., Chen, J., Honey, C.J., 2015. Hierarchical process memory: memory as an integral component of information processing. *Trends Cogn. Sci.* 19 (6), 304–313.
- He, B.J., Raichle, M.E., 2009. The fMRI signal, slow cortical potential and consciousness. *Trends Cogn. Sci.* 13 (7), 302–309.
- He, B.J., Zempel, J.M., Snyder, A.Z., Raichle, M.E., 2010. The temporal structures and functional significance of scale-free brain activity. *Neuron* 66 (3), 353–369.
- He, B.J., 2011. Scale-free properties of the functional magnetic resonance imaging signal during rest and task. *J. Neurosci.* 31 (39), 13786–13795.
- He, B.J., 2014. Scale-free brain activity: past, present, and future. *Trends Cogn. Sci.* 18 (9), 480–487.
- Jaffard, S., Lashermes, B., Abry, P., 2006. Wavelet leaders in multifractal analysis. In: Qian, M.T.V., Yueheng, X. (Eds.), *Wavelet Analysis and Applications*. Birkhäuser, Cambridge, pp. 219–264.
- Jaffard, S., Seuret, S., Wendt, H., Leonarduzzi, R., Roux, S., Abry, P., 2018. Multivariate multifractal analysis. *Appl. Comput. Harmon. Anal.* (in press).
- Jensen, O., Colgin, L.L., 2007. Cross-frequency coupling between neuronal oscillations. *Trends Cogn. Sci.* 11 (7), 267–269.
- Lakatos, P., Shah, A.S., Knuth, K.H., Ulbert, I., Karmos, G., Schroeder, C.E., 2005. An oscillatory hierarchy controlling neuronal excitability and stimulus processing in the auditory cortex. *J. Neurophysiol.* 94 (3), 1904–1911.
- Leonarduzzi, R., Wendt, H., Abry, P., Jaffard, S., Melot, C., 2017. Finite-resolution effects in p -leader multifractal analysis. *IEEE Trans. Signal Process.* 65 (13), 3359–3368.
- Lin, F.H., Belleveau, J.W., Dale, A.M., Hämäläinen, M.S., 2006. Distributed current estimates using cortical orientation constraints. *Hum. Brain Mapp.* 27 (1), 1–13.
- Lin, A., Maniscalco, B., He, B.J., 2016. Scale-free neural and physiological dynamics in naturalistic stimuli processing. *eNeuro* 3 (5) ENEURO-0191.
- Linkenkaer-Hansen, K., Nikouline, V., Palva, J., Ilmoniemi, R., 2001. Long-range temporal correlations and scaling behavior in human brain oscillations. *J. Neurosci.* 21 (4), 1370–1377.
- Mallat, S., 1998. *A Wavelet Tour of Signal Processing*. Academic Press, San Diego, CA.
- Miller, E.K., Cohen, J.D., 2001. An integrative theory of prefrontal cortex function. *Annu. Rev. Neurosci.* 24, 167–202.
- Monto, S., Palva, S., Voipio, J., Palva, J.M., 2008. Very slow EEG fluctuations predict the dynamics of stimulus detection and oscillation amplitudes in humans. *J. Neurosci.* 28 (33), 8268–8272.
- Murray, J.D., Bernacchia, A., Freedman, D.J., Romo, R., Wallis, J.D., Cai, X., Padoa-Schioppa, C., Pasternak, T., Seo, H., Lee, D., et al., 2014. A hierarchy of intrinsic timescales across primate cortex. *Nat. Neurosci.* 17, 1661–1663.
- Novikov, E., Nivkov, A., Shannahoff-Khalsa, D., Schwartz, B., Wright, J., 1997. Scale-similar activity in the brain. *Phys Rev E* 56 (10), 2387–2389.
- Palva, J.M., Palva, S., 2012. Infra-slow fluctuations in electrophysiological recordings, blood-oxygenation-level-dependent signals, and psychophysical time series. *Neuroimage* 62 (4), 2201–2211.
- Palva, J.M., Zhigalov, A., Hirvonen, J., Korhonen, O., Linkenkaer-Hansen, K., Palva, S., 2013. Neuronal long-range temporal correlations and avalanche dynamics are correlated with behavioral scaling laws. *Proc. Natl. Acad. Sci. U.S.A.* 110 (9),

- 3585–3590.
- Pelli, D., 1997. The VideoToolbox software for visual psychophysics: transforming numbers into movies. *Spat. Vis.* 10 (4), 437–442.
- Penttonen, M., Buzsáki, G., 2003. Natural logarithmic relationship between brain oscillators. *Thalamus Relat. Syst.* 2 (2), 145–152.
- Samorodnitsky, G., Taqqu, M., 1994. *Stable Non-Gaussian Random Processes*. Chapman and Hall, New York, USA.
- Shimizu, Y., Barth, M., Windischberger, C., Moser, E., Thurner, S., 2004. Wavelet-based multifractal analysis of fMRI time series. *Neuroimage* 22 (3), 1195–1202.
- Shimizu, Y., Umeda, M., Mano, H., Aoki, I., Higuchi, T., Tanaka, C., 2007. Neuronal response to Shepard's tones: an auditory fMRI study using multifractal analysis. *Brain Res.* 1186, 113–123.
- Suckling, J., Wink, A., Bernard, F., Barnes, A., Bullmore, E., 2008. Endogenous multifractal brain dynamics are modulated by age, cholinergic blockade and cognitive performance. *J. Neurosci. Methods* 174 (2), 292–300.
- Tagliazucchi, E., von Wegner, F., Morzelewski, A., Brodbeck, V., Jahnke, K., Laufs, H., 2013. Breakdown of long-range temporal dependence in default mode and attention networks during deep sleep. *Proc. Natl. Acad. Sci. U.S.A.* 110 (38), 15419–15424.
- Taulu, S., Simola, J., 2006. Spatiotemporal signal space separation method for rejecting nearby interference in MEG measurements. *Phys. Med. Biol.* 51 (7), 1759–1768.
- Torres, M.E., Abry, P., 2003. Comparison of different methods for computing scaling parameter in the presence of trends. In: *CD Memorias XIV Congreso Argentino de Bioingeniería y III Jornadas de Ingeniería Clínica (SABI 2003)*. Universitas, Córdoba, Argentina. . http://bioingenieria.edu.ar/grupos/ldnlys/meteorres/meteorres_files/SABI2003-074Torres.pdf.
- Touboul, J., Destexhe, A., 2010. Can power-law scaling and neuronal avalanches arise from stochastic dynamics? *PLoS ONE* 5 (2), e8982.
- Van de Ville, D., Britz, J., Michel, C.M., 2010. EEG microstate sequences in healthy humans at rest reveal scale-free dynamics. *Proc. Natl. Acad. Sci. U.S.A.* 107 (42), 18179–18184.
- Veitch, D., Abry, P., 1999. A wavelet based joint estimator of the parameters of long-range dependence. *IEEE Trans. Inf. Theory* 45 (3), 878–897.
- Wang, X.J., 2010. Neurophysiological and computational principles of cortical rhythms in cognition. *Physiol. Rev.* 90 (3), 1195–1268.
- Weiss, B., Clemens, Z., Bodizs, R., Vago, Z., Halasz, P., 2009. Spatio-temporal analysis of monofractal and multifractal properties of the human sleep EEG. *J. Neurosci. Methods* 185 (1), 116–124.
- Wendt, H., Abry, P., Jaffard, S., 2007. Bootstrap for empirical multifractal analysis. *IEEE Signal Proc. Mag.* 24 (4), 38–48.
- Werner, G., 2010. Fractals in the nervous system: conceptual implications for theoretical neuroscience. *Front Physiol* 1.
- Wood, J.N., Grafman, J., 2003. Human prefrontal cortex: processing and representational perspectives. *Nat. Rev. Neurosci.* 4 (2), 139–147.
- Zilber, N., Ciuciu, P., Abry, P., van Wassenhove, V., 2012. Modulation of scale-free properties of brain activity in MEG. In: *Proc. of the 9th IEEE International Symposium on Biomedical Imaging*. Barcelona, Spain. pp. 1531–1534.
- Zilber, N., Ciuciu, P., Gramfort, A., van Wassenhove, V., 2014. Supramodal processing optimizes visual perceptual learning and plasticity. *Neuroimage* 93 (Pt 1), 32–46.