Long-range alpha-synchronisation as control signal for BCI: A feasibility study.

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ABSTRACT

Shifts in spatial attention are associated with variations in alpha-band (α, 8–14 Hz) activity, specifically in inter-hemispheric imbalance. The underlying mechanism is attributed to local α-synchronisation, which regulates local inhibition of neural excitability, as well as fronto-parietal synchronisation reflecting long-range communication. The direction-specific nature of this neural correlate brings forward its potential as a control signal in brain-computer interfaces (BCI). In the present study we explored whether long-range α-synchronisation presents lateralised patterns dependent on voluntary attention orienting, and whether these patterns can be picked up at single-trial level to provide a control signal for active BCI. We collected electroencephalography (EEG) data from a cohort of healthy adults (n = 10) while performing a covert visuospatial attention (CVSA) task. The data show lateralised patterns of α-band phase coupling between frontal and parieto-occipital regions after target presentation, replicating previous findings. This pattern, however, was not evident during the cue-target orienting interval, the ideal time window for BCI. Furthermore, decoding the direction of attention trial-by-trial from cue-locked synchronisation with support vector machine (SVM) was at chance level. The present findings suggest EEG may not be capable of detecting long-range α-synchronisation in attentional orienting on a single-trial basis, and, thus, highlight the limitations of this metric as a reliable signal for BCI control.

INTRODUCTION

A few decades ago, imagining an interface between the human brain and a computer was closer to science fiction than to scientific achievement. Nowadays, brain-computer interfaces (BCIs) can read out brain activity, extract features from that activity, and convert those features into outputs used for monitoring, controlling devices, or even modifying cognitive states (Blankertz et al. 2016). One significant challenge of BCIs is finding reliable control signals from brain activity with sufficiently high signal-to-noise ratio (SNR) at a trial-by-trial level to allow successful classification. Ideally, the appearance of the target brain activity should depend on endogenous mental states that the user can control at will. The use of non-invasive, cost-effective, and light-weight neuroimaging devices can, in turn, facilitate transfer to real applications. For now, EEG is the most viable candidate to achieve this.

Some EEG-based BCIs have used motor imagery as a control signal (e.g., imagined right/left-limb movement; Padfield et al. 2019), whereas others have used neural correlates of covert visuospatial attention (CVSA) (van Gerven and Jensen 2009; Treder et al. 2011; Tonin et al. 2013). Here, we will concentrate on the latter. In human behaviour, CVSA is used to direct processing resources to relevant locations in the environment whilst disengaging from irrelevant locations (Pashler 1999; Foster and Awh 2019). CVSA can be manipulated with the Posner cueing protocol (Posner 1980), which shows a robust effect on behavioural performance: higher accuracy and faster reaction times for targets appearing at the cued (attended) location compared to targets appearing in un-cued, putatively unattended locations (Posner 1980).

Shifts in CVSA produce changes in oscillatory activity in the alpha-band (α, 8–14 Hz) at parieto-occipital regions (Klimesch 1999; Foster et al. 2017; van Diepen et al. 2019). Typically, α-power presents a direction-specific imbalance when attention is covertly oriented to either the left or right visual field, revealing its potential as a control signal for BCI implementations (Rihs et al. 2007; Thut et al. 2006, see Astrand et al. 2014b for a review). This neural correlate is thought to correspond to a late stage in the brain processes involved in CSVA shifts. First, cueing information is integrated through sensory pathways in a bottom-up fashion, reaching higher visual areas in the parietal cortex (e.g., intraparietal sulcus) and eventually frontal regions (e.g., frontal eye fields) (Petersen and Posner 2012). From there on, top-down modulation shifts attention to the corresponding hemifield, where it is maintained during target anticipation (Simpson et al., 2012). The mechanism involved in this

top-down modulation is thought to involve long-range α-synchronisation between frontal and posterior cortex, which eventually leads to classical interhemispheric imbalances in α-power observed in the visual cortex (Sauseng et al. 2005; Doesburg et al. 2009, Lobier et al. 2018). This long-range synchronisation is a potential mechanism to increase the fidelity and effectiveness of communication throughout the brain (Clayton et al. 2018) among occipital, parietal, and frontal regions (Sadaghiani and Kleinschmidt 2016). Synchronising excitability cycles between distant neural populations increases the likelihood of spikes from one region discharging post-synaptic potentials during a specific (excitable) phase of the other (Fries 2005, 2015). Despite the overwhelming evidence supporting these models of visual processing, there is still debate on their temporal dynamics, lateralisation patterns and individual-level variability. Further research on these aspects may provide deeper insight into the exact mechanisms of long-range α-synchronisation within the fronto-parietal network (FPN).

The present study addressed whether long-range α-synchronisation in the FPN presents direction-specific lateralised patterns (i.e., contralateral to ipsilateral differences) regarding the attended location, and if such patterns emerge in single-trial dynamics with sufficient signal strength to make for a reliable control signal in BCI. To the best of our knowledge, BCIs based on attention have only used α-power as a control signal, despite the evidence of links between long-range α-synchronisation and behavioural performance, using group-level analyses (Sauseng et al. 2005; Doesburg et al. 2009; Doesburg et al. 2016). We addressed this by measuring EEG during CVSA using a Posner task and examining the time course of long-range α-phase synchronisation between cue onset and target appearance. This time window is essential for BCI since it is the period during which participants lateralise their attention covertly. Our approach was to replicate the analysis pipeline of Sauseng et al. (2005), reproduce the group effect at the target-locked window, and extend our analysis to the cue-to-target window, more relevant for BCI. We finally assessed the classification of attentional loci at the single-trial level using long-range α-phase synchronisation as a proof of concept for BCI.

MATERIALS AND METHODS

Participants

We used data from a previous, unrelated study (Torralba et al 2016). The dataset consisted of 15 participants (mean age = 22, SD = 3, 7 female). All participants provided informed consent and had a normal or corrected-to-normal vision. The study was ran in accordance with the Declaration of Helsinki and the experimental protocol approved by the local ethics committee CEIC Parc de Salut Mar (Barcelona, Spain). Five participants who presented equivalent detection and discrimination rates for stimuli appearing at cued and un-cued locations were discarded from the analysis, leaving a total of 10 participants.

Task

Before the experimental session, the participant's EEG activity was recorded during a five-minute recording at rest with eyes closed to extract the individual α frequency (IAF) used in the analyses. In the experimental session, participants performed a modified version of the Posner cueing task (see **Figure 1A**). The trial started with the onset of a central fixation cross, placed between two placeholder squares located 20º of visual angle left and right off centre, vertically shifted 20º of visual angle below the fixation cross (see **Figure 1A**). After 200 ms fixation period, a central auditory cue (100 ms duration) indicated the likely target location through either high pitch (2000 Hz) or low pitch (500 Hz) tones, the mapping was randomised across subjects. Participants should covertly attend to the indicated side, without moving their eyes, during a jittered inter-stimulus interval (ISI; 2000 ± 500 ms). The use of a jittered ISI was employed in order to avoid participants falling into a constant rhythmic pattern. Next, the target (a Gabor grating tilted 45º left or right, 50 ms duration) appeared briefly inside one of the placeholders, with 75% validity regarding the cued location. The grating contrast was adjusted individually, as described below. A noise pattern of equal overall luminance as the target was presented at the alternative placeholder, with the exact timings as the target. Participants were asked first to indicate if they had detected the target (yes/no detection) and subsequently the target's tilt (left/right discrimination). Both answers were made by keypress, in an un-speeded fashion, and with response mapping (top-bottom) orthogonal to the attention manipulation and varied from trial to trial. An inter-trial interval of 1000 ms followed the response, and a new trial began. Unless otherwise noted, the EEG analyses were done

on validly cued trials that responded correctly. On average, 289.9 ± 11.3 trials from each participant were employed for the EEG analysis.

The Gabor gratings used as stimuli were 0.002 cycles per degree, with a size of 3.35º, and were embedded in white noise. The contrast was adjusted individually using a preliminary threshold titration procedure in which thresholds for both sides (left and right) were independently adjusted to a 70% detection rate when cued (in the attended location). Stimuli were presented on a 21" CRT screen with a refresh rate of 100 Hz and a resolution of 1024 x 768 pixels. The experiment was implemented in MATLAB R2015b (MATLAB, RRID: SCR_001622) using the Psychophysics Toolbox (Psychophysics Toolbox, RRID: SCR_002881).

EEG recording

EEG recordings were obtained from 64 Ag/AgCl electrodes positioned according to the 10-10 system with AFz as ground and nose tip as reference. Impedances were kept below 10 kΩ. The employed system was an active actiCHAmp EEG amplifier from Brain Products (Munich, Germany). The signal was sampled at 500 Hz and processed in MATLAB 2020 and 2015 (MATLAB, RRID: SCR_001622) using custom functions and the FieldTrip toolbox (FieldTrip, RRID: SCR_004849).

EEG Pre-processing

In order to remove eye movements, blinks, and muscle artefacts, Independent Component Analysis (ICA) was carried out (Makeig et al. 1995). Additionally, manual artefact rejection was applied to discard trials where any EOG components had an amplitude higher than 50 µV. Defective channels were repaired using neighbours calculated by triangulation and splines for interpolating channel data. Following these steps, the data was demeaned and notch filtered at 50 Hz to exclude line noise. Next, fifth-order high-pass and sixteenth-order low-pass IIR Butterworth filters were employed to limit the signal between 16 and 45 Hz (Sauseng et al. 2005). The filtering was done forward and backwards (two-pass), which resulted in zero phase lag.

Determination of the individual α frequency (IAF)

The frequency of interest used in analyses of this study was adjusted for each participant depending on the individual α frequency (IAF) extracted from the five-minute recording (eyes

closed) previous to the experiment (see above). The IAF was determined based on the presence of a single peak (in other words, a local maximum) within the considered frequency band of interest (5-15 Hz) on the power spectrum density (PSD). A spectrogram was extracted for each parieto-occipital electrode (P7, P5, P3, P1, Pz, P2, P4, P6, P8, PO3, PO4, POz, PO9, PO10, O1, Oz, O2) using the Welch method (segments of 1000 ms with a 10% overlap, a Hanning taper to avoid spectral leakage and 0.25 Hz frequency resolution). The power spectrum was averaged across electrodes for each participant and normalised by the mean power from 1 to 40 Hz.

Time-frequency analysis

We performed long-range synchronisation analyses in two time windows. The first was time-locked to the target onset (target-locked) to replicate Sauseng et al. (2005) methods and validate our analysis pipeline. The second was time-locked to the cue onset (cue-locked) to estimate long-range α-phase synchronisation during covert visuospatial attention shifts.

Following Sauseng et al. (2005), for the target-locked analysis we used two windows of 200 ms: a pre-target (-200 to 0 ms) and a post-target window (200 ms to 400 ms). The latter excludes the interval 0 to 200 ms, most affected by the phase resetting effect of target presentation. For the cue-locked analysis, we used the cue-to-target time window between 500 ms and 1500 ms post-cue and divided it into five consecutive and non-overlapping 200 ms windows. By analysing from 500 ms onwards we avoid the event related potential (ERP) caused by cue presentation and allow endogenous attention shift to build up, a process which takes a few hundreds of milliseconds (Foxe and Snyder 2011). The cue-locked analysis period ends at 1500 ms, which was the minimum possible duration of the cue-to-target interval (duration of 2000 ± 500 ms, see methods). All epoched data was mirror-reflected to avoid edge artefacts (Cohen 2014) when performing the time-frequency analysis. Afterwards, data were trimmed, and reflected edges were removed.

We computed the Fourier coefficients using 5-cycle Morlet wavelets (Grossmann and Morlet 1984) with 16 logarithmically spaced frequencies ranging from 2.6 Hz to 42 Hz. For the analysis centred on the individual α frequency (IAF), we only used wavelets within the upper α-band (9.54 – 14.31 Hz) (Sauseng et al. 2005), whereas, for the exploratory analysis, we used the whole frequency range. This difference allowed us to conduct a hypothesis-driven analysis using the upper α-band to replicate Sauseng et al. (2005) and an exploratory

180 analysis to explore further long-range α-phase synchronisation in other frequency bands
181 beyond the IAF.

Connectivity measures

Three regions of interest (ROI) were defined for the analyses: A fronto-medial (FM) ROI (Fz,
FC1, FC2) and two symmetric posterior regions; the parietal left (PL) ROI (P5, P7, PO3, O1)
and the parietal right (PR) ROI (P6, P8, PO4, O2). To infer connectivity between each of
these parietal ROIs and the FM location, we used Phase Locking Value (PLV) (Lachaux et
al. 1999). This metric reports the consistency of phase differences between two locations
across multiple trials and is not affected by power differences. Mathematically, the PLV is
expressed as the absolute value of the average complex unit-length phase differences:

$$190 \quad PLV(x, y) = \left| \frac{1}{n} \sum_{k=1}^{n} e^{i\left(\varphi_x(k) - \varphi_y(k)\right)} \right| \tag{1}$$

where *n* corresponds to the total number of trials indexed by *k* and φ_x, φ_y correspond to the
phases at electrodes *x* and *y,* respectively. PLV was calculated according to equation (1)
using the phases for every combination of individual electrode pairs of the FM-PR and FM-
PL networks. Then, these values were averaged, resulting in a time series of PLV FM-PR
and FM-PL networks for each of the frequencies of interest and condition (*attended left* and
attended right) trials. Subsequently, the PLV time series were collapsed as either ipsilateral
(FM-PL network and attend left; FM-PR and attend right) or contralateral (FM-PR network
and attend left; FM-PL and attend right). Therefore, for each participant and frequency of
interest, two time series of PLV were obtained (contra- and ipsilateral PLV).

Classification

The trial classification was performed using Support-Vector Machines (SVM). We selected
the FM-PR and FM-PL connectivity for the metric used as input to the SVM. *Attended right*
and *attended left* labels of each trial were provided as ground truth for the algorithm. The
main goal of the classifier was to infer where the participant was attending on each trial based
on the long-range α-phase synchronisation in the left and right fronto-parietal networks. Note
that PLV is computed across trials, and SVM aims to classify on a single-trial basis, so PLV
was also calculated across time points (Cohen 2015). As a validation step, we repeated the
target-locked analysis employing this metric (i.e., cross-time PLV) before proceeding with the
cue-locked classification attempt.

We divided the cue-locked interval ranging from 500 to 1500 ms in bins of 200 ms, yielding

five values for FM-PR connectivity and five for FM-PL connectivity. The resulting ten values

were used for the SVM input to perform the optimisation and classification of the trials. Note

that for the classification, we only used the data from the participants that achieved a

significant difference in PLV values between parietal left and right ROIs in the pre-target

window of the target-locked analyses. For each participant, trials were split into a training

(80%) and testing (20%) set of trials to avoid overfitting. Then, the training set was subdivided

into sub-training (80%) and validation sets (20%).

Our initial approach was to use a linear kernel for the classification. However, after evaluating

the option through cross-validation of the validation set and obtaining a negative result (i.e.,

classification was not better than chance level), we decided to use instead of a Gaussian

kernel (i.e., Radial Basis Function). In order to select the most suitable and efficient values

for classifying *attended left* and *attended right* trials from the validation set, we optimised the

parametric space of the SVM, comprised of margin and gamma (γ) parameters, and explored

the parametric landscape ranging from 10^{-6} to 10^{3} in steps of 10 for both constants.

225   Inter-hemspheric power imbalance exploratory analysis

Besides calculating the long-range α-phase coupling, we also computed the inter-

hemispheric α-power imbalance at parietal regions both at the individual and at group-level

as a reality check. To extract the α-power during the task, we selected the epoch from -1.5

to 3 s in cue-locked trials by convolving the EEG signal with a set of complex Morlet wavelets

(Grossmann and Morlet 1984) of 6-cycles (n_c). The frequencies of the wavelets ranged from

IAF - 1 Hz to IAF + 1 Hz in 1 Hz steps. For instance, an IAF peak of 10 Hz would have a

bandwidth ranging from 8.33 Hz to 11.67 Hz. The power in each of the two symmetric ROIs

at posterior regions (PL and PR ROIs; same used in the connectivity analysis) was averaged,

and the power imbalance was computed in terms of the lateralisation index of α-power,

according to the formula proposed by Thut et al. (2006):

$$Lateralization\ Index = \frac{\alpha(PR\ ROI) - \alpha(PL\ ROI)}{mean\ of\ \alpha(PL+PR\ ROI)} \tag{2}$$

where α(PL ROI) and α(PR ROI) are the average of α-power over left and right regions of

interest, respectively. Equation (2) leads to smaller (negative) values in the case of α-activity

was more prominent over the left hemisphere than the right (α(PL ROI) > α(PR ROI)) and to

larger (positive) values for the opposite pattern (α(PL ROI) < α(PR ROI)). According to theory and previous findings, values of LI reflecting attention directed to the right visual field should be larger than LI values reflecting leftward directed attention.

Finally, we also checked whether there was any relationship between the α-power imbalance and the contra-ipsi difference of PLV for each attended location. We explored the correlations between α-lateralisation indexes and the effect in PLV contra-ipsi differences at the pre-target (-200 to 0 ms) and post-target (200 to 400 ms) windows using Pearson correlations.

Statistical analyses

A one-tailed nonparametric Monte Carlo permutation test was computed to determine significant differences in PLV between networks for each attended location (Mostame et al. 2019). For each participant, the *attended right* or left labels were randomly assigned to trials, and surrogate PLVs were calculated from the resulting dataset. This process was repeated 10,000 times (iterations) to create a null distribution of PLV values. The obtained p-value corresponded to the proportion of surrogate iterations with a contra-ipsi difference larger than the actual measured value (one-tailed test). This process was performed on every time window defined in the previous section. For the group analysis, the procedure was equivalent, but surrogate PLV distributions were averaged across participants before the statistical test.

For the statistical assessment of the α-power imbalance over time between *attended left* and *attended right* trials, we performed a cluster-based permutation test procedure (100,000 randomisations) for each participant and at the group-level (one-tailed permutation test) (Maris and Oostenveld 2007; Meyer et al. 2021). We assessed that lateralisation indexes for attended-right and attended-left trials were significantly two different distributions by applying a one-tailed t-test (independent samples) with α-level = 0.05 for each participant. At group-level, we performed a one-tailed paired t-test with the mean lateralisation indexes for *attended right* and *attended left* trials for each participant with α-level = 0.05. Correlations between α-power imbalance and the contra-ipsi difference of PLV were corrected for multiple comparisons by applying the False Discovery Rate (FDR) of Benjamini and Hochberg (Benjamini and Hochberg 1995).

RESULTS

Behavioural results

As expected, behavioural results showed that the detection rate was superior for cued (attended) trials 0.68 ± SEM=0.02 compared to un-cued (unattended) ones 0.46 ± 0.04 (see, **Figure 1B**). The pattern on each hemifield was equivalent to the overall pattern: on the left hemifield attended = 0.68 ± 0.03 and unattended = 0.47 ± 0.03; for the right hemifield attended = 0.67 ± 0.03, and unattended = 0.44 ± 0.06.

Target-locked long-range α synchrony

As described in the analyses section, long-range synchrony was estimated using PLV between a frontal ROI and each of two lateralised parietal ROIs. Here, we describe the target-locked analysis, mainly carried out to reproduce Sauseng et al. (2005). **Figure 2** shows the group-level connectivity analysis of the upper α-band (9.54 - 14.31 Hz). Phase coupling is depicted as the mean across the pre-target window (-200 to 0 s) and the post-target window (200 to 400 ms), as well the temporal course (from to -500 to 500 ms). Regarding the left fronto-parietal network (**Figure 2A left**), PLV was consistently higher when attention was directed right (contralateral) than left (ipsilateral) in both pre-target and post-target windows, although the PLV difference did not reach significance in the pre-target window. Regarding the right network (**Figure 2A right**), PLV was stronger when attention was directed left (contralateral) than right (ipsilateral) in the post-target window, whereas the pre-target window does not show this difference. This pattern generally replicates Sauseng et al. (2005) results, as indicated by the dashed lines in **Figure 2A** representing the mean phase-coupling from their study. Lower panels in **Figure 2A** display the temporal course of phase coupling to provide a time resolved illustration of the phase-coupling effect. For the attend right condition, PLV values in the left network should be higher than PLV values for the *attended left*. The inverse pattern should hold in the right network. Moreover, **Figure 2B** presents the PLV with side of attention collapsed as contra- and ipsilateral with respect to the corresponding network. Individual PLV values, marked as black dotted lines, exhibit a consistent contra- to ipsilateral increase in the post-target window. Group-level statistical analysis further showcased a significant difference limited to this time window (200 to 400 ms, $p < 0.001$).

In the single-subject analyses, 8 out of 10 participants showed contralateral PLV increases, in which 5 of them were significant (P04, $p < 0.001$; P05, $p < 0.001$; P06, $p < 0.05$; P07, $p < $

299 0.001; and P10, $p < 0.01$; see **Figure 2-1**). The lack of significant differences in the pre-target
300 window is consistent with individual phase coupling, as no clear contra-ipsi trends emerged.
301 Five participants had higher contralateral PLV values in this time window, 3 of which exhibited
302 a significant behaviour (P04, $p < 0.05$; P05, $p < 0.001$; and P10, $p < 0.05$; see **Figure 2-1**).

Cue-locked long-range α synchrony

The primary aim of this study was to explore the cue-to-target interval before target
presentation (500 ms to 1500 ms after cue onset) to ascertain whether attention-based long-
range connectivity during the orienting period could be a reliable signal for BCI control.
Considering that the cue indicates the hemifield to which participants should voluntarily
lateralise their attention, differences in contralateral and ipsilateral connectivity may
potentially emerge in this time window. So far, we have seen that this attention shift had
significant consequences on behaviour and target processing (post-target connectivity). At
the group level, however, no significant difference between contralateral and ipsilateral
connectivity in the upper α-band was found in any of the five 200 ms time windows considered
in the cue to target period (see **Figure 3A**). At the individual level, only one participant (P05)
showed a significant increase in the contralateral PLV in four out of five-time windows (see
Figure 3-1). This participant also showed a significantly higher contralateral connectivity in
both the pre-target and post-target time windows of the target-locked analysis.

We chose the upper α-band a priori given prior findings of Sauseng et al. (2005) as well as
the effects in the target-locked analyses of our own data. However, we conducted additional
analyses to explore other frequencies (between 2.4 and 42 Hz) in search of differences
between contralateral and ipsilateral PLV (see, **Figure 3B)**. Values were collapsed as the
difference between both measures (contra-ipsi) and z-scored. Over time, neither clear trends
across frequencies nor apparent increases are observed in either contralateral or ipsilateral
connectivity. Individual results of the exploratory analysis show the same results and do not
present relevant PLV patterns in any participant aside from upper α-band findings in P05 (see
Figure 3-2).

Classification

The results are hardly promising in generalising the use of long-range connectivity for BCI
control. However, BCI protocols are often very sensitive to individuals. Here, we intended to
seek proof-of-concept evidence. With this goal in mind, we attempted single-trial classification

from selected participants datasets, as either *attended right* or *attended left*, according to cue-
locked connectivity patterns. To maintain statistical independence between dataset selection
and test, participant selection was based on the target-locked data. We selected the 3
participants (P04, P05 and P10) for whom we found significant connectivity differences in the
pre-target time window of the target-locked analysis, and their data were used for the
classification. Please note that this participant selection of the pre-target window, from -200
to 0 ms from target onset, is aligned differently than the cue-locked analysis window, which
considers times up until -500 ms from target presentation. The total number of trials for each
participant was 272 (P04), 310 (P05), and 338 (P10).

As described in the methods section, we carried out a validation of cross-time PLV in the
target-locked window to understand whether this metric could replicate group-level
differences between contra- and ipsilateral networks found through cross-trial PLV. These
results can be seen in **Figure 4A**. Statistical analysis showed no significant differences
between contra- and ipsilateral scenarios in either time window. Individual values also
remained non-significant (see **Figure 4-1**). Considering the large parametric landscape of
SVM implementations we optimised the gamma and margin parameters of a Gaussian kernel
(see **Figure 4B)**. From a qualitative perspective, no clear maximum validation accuracy
values emerge from the landscape, although quantitative analysis identified a margin of
5.0119×10^{-6} and a γ value of 2.51×10^{-2} to have the most optimal validation set of the outcome.
The lack of an clear minimum suggests that the model may be unable to classify individual
trials regardless of the parametric values.

Ten-fold cross-validation was carried out independently on all participants' trials to reduce
the inter-individual variability and improve the classification accuracy. Single trials predicted
as either *attended right* or *attended left* were contrasted with the actual cue direction in each
trial. Classification outcomes of each participant are shown in **Figure 4C**, which resulted in
virtually chance-level sorting (0.548, 0.565, and 0.467, respectively). Confusion matrices
display the distribution of each class, revealing the skewed distribution of values towards
attended left labels, which is far from the ideal clustering along the diagonal of the matrix.

Inter-hemispheric power imbalance

As a reality check on the dataset, we addressed whether there was a difference in the α-
power inter-hemispheric imbalance between *attended left* and *attended right* trials. We
performed the cue-locked analysis at the group level, using the Lateralization Index (LI)

described by Thut et al. (2006) (see **Figure 5A).** On average, the lateralisation index was
significantly different between *attended right* and *attended left* in the expected direction ($p =$
.01, Cohen's d = -0.8356). At the individual level, 7 out of the 10 participants showed a
significant difference in lateralisation index between the two attention conditions ($p = $ *<.001*;
see **Figure 5-1**). We also performed a time-resolved version of this analysis within the said
window. The cluster-based permutation test (**Figure 5B**) showed significance within two time
periods, from 0.66 to 0.82 s and 1.34 to 1.5 s. At the individual level, only for one participant
(P01), the cluster-based permutation test revealed a significant cluster over time from 0.6 to
1 s (see **Figure 5-1**). Generally, these results are consistent with the results of previous
studies (e.g., Tonin et al. 2012; Thut et al. 2006), at least at the group level. It is more
challenging to compare single-subject data with other studies, as it usually is not reported or
statistically analysed.

Finally, we explored the potential correlation between α-power inter-hemispheric imbalance
measured with the lateralization index and α-phase coupling for each attended location(see
Figure 5 C-D). In the pre-target window (**Figure 5C**), the correlations for *attended right* ($r = $ -
0.25, $p > 0.05$) and *attended left* ($r = -0.13$, $p > 0.05$) did not reach significance. Neither did
the correlations for *attended right* ($r = -0.44$, $p > 0.05$) and *attended left* ($r = -0.42$, $p > 0.05$)
at the post-target (**Figure 5D**) window. A visual inspection indicated that participants showing
an effect in PLV contra-ipsi differences are below the correlation fit in pre-target and post-
target windows, suggesting that those participants have a more negative effect in PLV contra-
ipsi differences.

DISCUSSION

The present study addressed the relationship between shifts in visuospatial attention and the lateralisation of α-band coherence between frontal and parietal sites, with the intention to assess their feasibility as a control signal in BCI. Previous studies, using group-averaged multi-trial analyses, found increased long-range α-synchronisation in the hemisphere contralateral to the attended side of space, and suggested that it reflects top-down mechanisms of visual attention (Sauseng et al., 2005; Doesburg et al., 2009). We reasoned that if contra- to ipsilateral differences in synchronisation would emerge as a result of endogenous top-down mechanisms, they should be present following cue presentation as participants shift their attention. This hypothesis stems from instructing participants to shift their attention laterally in expectancy of target appearance engages frontoparietal visual processing pathways (Corbetta and Shulman 2002; Hopfinger et al. 2000; Asplund et al. 2010). Here, we sought proof of concept that long-range neural synchronisation engaged in this network could be used for BCI control on a trial-by-trial basis.

In attention orienting protocols, the cue-to-target period offers the possibility of implementing a BCI control harnessing covert attentional orienting in individual trials due to lateralisation of attention in anticipation of the target appearance. This would open the possibility of designing active BCI systems controlled by the user's voluntary decision to attend left or right covertly. Therefore, our study employed long-range α-synchronisation in the frontoparietal network (FPN) as means to investigate whether this brain measure could potentially discriminate attended locations of the left/right visual field.

We found significant group-level differences in contra- to ipsilateral long-range α-synchronisation around target onset time in direct replication of Sauseng et al. (2005). These results help confirm the importance of lateralised long-range α-synchrony along the FPN during orienting and especially reveal the potential of EEG to grasp these effects, at least during target processing at the group level. However, similar differences in synchrony were not observed during the post cue time window, which was the time of interest for BCI purposes. We extended the cue-locked analysis to other frequencies outside the α-band, with equally negative results. Finally, given the high individual variability of single-trial analysis outcomes, we attempted to classify the individual trials of selected participants for whom significant synchronisation differences following cue presentation were found, a benchmarking process that nevertheless rendered chance-level classification. Below, we

discuss how these results may be influenced by various methodological aspects (e.g., different time windows, classifier's input metric) and how they fit into state-of-the-art literature. Please note that because the focus of our study was on single-trial analysis, the sample size was relatively small for the group analyses (n = 10). Although this sample size was sufficient to confirm previous findings, the negative results of the group analyses should be interpreted with caution until a more extensive study is conducted.

Fronto-parietal network synchronisation characterises visuospatial attention

A result from our study is that long-range α-synchronisation within the FPN was associated with visuospatial attention orienting, in line with its putative role in this cognitive process (Jensen et al. 2015; Sacchet et al. 2015; Doesburg et al. 2009; Siegel et al. 2008). We observed a spatially distributed difference in upper α-coherence, with increased synchrony in the FPN pathway contralateral to the attended hemifield. According to current attention theories, the mechanism underlying this finding may relate to an initial attentional modulation stemming from frontal areas such as the frontal eye fields (FEF) and the intraparietal sulcus (IPS) (Corbetta and Shulman 2002; Kastner and Ungerleider 2000; Helfrich et al. 2018), leading to a state of α-band desynchronisation in the relevant area of the visual cortex (i.e., contralateral to attended hemifield) in anticipation of relevant information (Capotosto et al. 2009; Marshall et al. 2015). This explanation aligns with the well-established evidence that contralateral α-power suppression (also reproduced in our results) enables visual stimuli processing in the attended location (Doesburg et al. 2009; Thut et al. 2006; Yamagishi et al. 2003; Babiloni et al. 2006; Foxe and Snyder 2011; Klimesch et al. 2007; Lange et al. 2013), and that cyclic phase-dependent inhibition in low-level visual cortex dictates behavioural performance (i.e., reaction times) (Haegens et al. 2011; Klimesch 2012; Jensen et al. 2014; Samaha et al. 2015; VanRullen 2016). Both accounts fit with the idea that local α-power and long-range α-synchronisation may have separate roles in attention and perception (Bonnefond et al. 2017; Palva and Palva 2007, 2011; Sadaghiani and Kleinschmidt 2016).

Our results of the increased contralateral synchronisation within the FPN replicate the work of Sauseng et al. (2005) and validate our methodology and analysis pipeline (e.g., time-frequency analysis, synchronisation metric), setting the ground for the intended proof of concept test regarding transference to BCI. However, lateralised frontoparietal connectivity patterns in attentional and perceptual disposition as well as the role of α power/phase remain

challenged in the literature (Ruzzoli et al. 2019; van Diepen et al. 2019; Antonov et al. 2020). Lobier et al. (2018) found that α-synchronisation was associated with visuospatial attention but revealed distinct lateralisation patterns regarding the visual system and top-down attentional networks. They showed stronger ipsilateral synchronisation within the visual system (in line with Siegel et al. 2008; Doesburg et al. 2009) but no consistent lateralisation in long-range networks, suggesting their different involvement in visuospatial attention. A study by D'Andrea et al. (2019) found a modulation of frontoparietal α-β cross-frequency synchronisation during attention orienting, but not in α-synchronisation alone. Further, this cross-frequency connectivity pattern was strongly associated with right hemisphere frontal dominance (in line with Heilman and van den Abell (1980) and Zago et al. (2017)). This finding agrees with previous evidence of the crucial role of the right FEF in top-down attentional modulation (Esterman et al. 2015; Hung et al. 2011; Silvanto et al. 2006), supported by evidence using TMS (e.g., Capotosto et al. 2009). In light of this evidence and our results, the exact relationship between contralateral frontoparietal α-synchronisation and shifts in attention orienting is still unclear. Positive findings, however, such as the ones in the present study using a target-locked analysis, represent a basis for exploring earlier time windows capable of shedding light on the mechanism underlying FPN α-synchronisation.

Correlations between long-range α-synchronisation and individual reaction times in visuospatial tasks suggest this neural correlate may be observable at a single-subject level (Lobier et al., 2018). Despite this, significant group-level target-locked dynamics of increased synchrony did not transfer to all individuals in our study. The observed variability may be partially explained by individual differences in anatomical tracts of attentional relevance (e.g., superior longitudinal fasciculus) (Marshall et al., 2015). Findings employing MRI suggest that volumetric differences in these structures impact local visual cortex oscillations, leading to variability in EEG traces (Marshall et al. 2015; D'Andrea et al. 2019). However, this variability of individual results is challenging to set in the perspective of previous research simply because published studies do not report single-subject statistics. Ultimately, the outcomes of this study leave an incomplete understanding as to whether there is a true group effect that does not extend to all individuals or, contrarily, whether individual effects of specific participants are large enough to induce a group-level finding in previous research.

Lateralized patterns of α-synchronisation appear in target-locked but not cue-locked analysis

Long-range α-synchronisation presented contralateral increases at the post-target (200 to 400 ms, with t = 0 as target appearance) and the pre-target window (-200 to 0 ms). However, only the former time window resulted significant. This result is slightly different from Sauseng et al. (2005), who observed significant contralateral synchronisation increases within the FPN network at both time windows. However, the numerical differences were in the same line in both studies, and the different outcomes of some statistical significance tests may be just due to a lack of statistical power. Another potential explanation for the absence of significant findings at the pre-target window may be the difference in experimental paradigms. The task employed here had a long post-cue interval ranging from 2000 to 2500 ms (jittered between trials). Sauseng et al. (2005), on the other hand, had a shorter delay between cue and target appearance of 600-800 ms, thereby perhaps leading to a more concentrated time window where the attention shift may have taken place. If participants shifted attention at varying times from cue onset up to target appearance, this might explain why we were unable to capture the effect in anticipatory visuospatial attention.

In cueing paradigms, bottom-up integration of cue information through sensory pathways precedes top-down modulation of visuospatial attention (Simpson et al., 2011). The temporal course of this cognitive operation is thought to begin only after 150 ms of visual cueing in voluntary attention and includes frontal regions approximately after 350 ms. Furthermore, from 400-500 ms onwards, frontal and parietal regions are thought to be solely involved in attentional shifting and target discrimination (Simpson et al. 2011). Thus, if the FPN does present direction-specific synchronisation, we anticipated this would appear from 500 ms after cue onset onwards. Contrary to what we expected, we did not observe any significant contra- to ipsilateral differences in the cue-target time windows (500 to 1500 ms after cue onset). Previous studies employing a similar time window showed lateralisation patterns in parietal regions in α and β bands (Siegel et al. 2008; Pantazis et al. 2009) and frontoparietal lateralisation in low and high-frequency bands (Green and McDonald 2008; Gregoriou et al. 2009). Therefore, we extended our cue-locked analysis to other frequencies, achieving no significant contra- to ipsilateral differences. Note that PLV values were averaged across 200 ms windows, and this excludes, to a certain extent, the confound of frontal and parietal regions having different activation over time. Altogether, despite the evidence across multiple

frequencies of synchronisation in the cue-target time window, we did not find patterns of lateralised cue-locked connectivity within or outside the α-band.

Our negative results in the cue-locked analysis may align with the notion that late periods after cue onset are associated with direction-specific activity in parieto-occipital regions but not in frontal regions (e.g., FEF) (Doesburg et al. 2009; Simpson et al. 2011). Long-range α-synchronisation may, therefore, initially shift attention (shortly after cue presentation) and later (close to target presentation) maintain it at the directed hemifield favouring perception and releasing lateralised patterns (Lobier et al. 2018; Kastner and Ungerleider 2000; Hopfinger et al. 2000; Grent-'t-Jong and Woldorff 2007). This idea resonates with the essential question formerly posed by Sauseng et al. (2005), debating whether frontal involvement in long-range α-synchronisation is a causative or consequential correlate of posterior activation. Furthermore, it motivated the exploration of cue-locked intervals where bottom-up and top-down processing may have elicited stronger effects on α-band synchronisation.

Finally, to ensure participants correctly lateralised their attention during the cue-target interval, we carried out a reality check by calculating the α-power imbalance using the lateralisation index during this period (Thut et al. 2006). There was a clear difference in the averaged lateralisation index during the time course between 500 and 1500 ms at group-level. We further employed this metric (i.e., lateralisation index) to perform an exploratory analysis on its relationship with the difference in synchronisation between contra- and ipsilateral networks. Considering lateralised local α activity and lateralised long-range α-synchronisation are both relevant in successful attention orienting, we hypothesised that these two mechanisms would have a significant positive correlation. Therefore, individuals with high lateralisation index values should also present lateralised synchronisation within the FPN. In contrast to our expectations, there was no significant correlation between these two metrics, neither at the pre-target nor the post-target time windows.

Ultimately, we did not observe a significant increase in contralateral long-range α-synchronisation in the five 200 ms bins following cue onset. This time frame offered potential as it occurs much before target appearance and could be robustly employed in a covert visuospatial BCI decoder. By expanding our analysis to several frequencies and carrying out the aforementioned reality checks, we conclude that PLV measured from EEG may not serve as a reliable metric in capturing direction-specific synchronisation from frontal to posterior

541 regions, despite this evidence being present in parietal to occipital synchrony (Doesburg et
542 al. 2009).

EEG estimates of long-range α-synchronisation may not serve as a reliable control signal for BCI

The use of long-range α-synchronisation to decode attentional direction yielded chance level
results. We employed 200 ms time bins of contralateral and ipsilateral FPN connectivity as
input in an SVM classifier. Non-linear SVMs are widely employed in decoding cognitive neural
correlates of behavioural states. Furthermore, they outperform other classifiers such as
artificial neural networks, non-linear Bayesian estimators, and recurrent reservoir networks
(Astrand et al. 2014a). Prior work using SVMs, mainly centred around primate models and
invasive recordings, successfully decoded the attentional spotlight from frontal sites (Gaillard
et al. 2020; Tremblay et al. 2015; Esghaei and Daliri 2014). Although these modalities (i.e.,
LFP, intracranial-EEG) have a high signal-to-noise ratio (SNR), the objective of the present
study is to perform a BCI proof of concept and, thus, a more realistic and suitable imaging
method must be employed. This same principle can be utilised to discard other modalities
such as fMRI, where the temporal resolution is too low for real-time implementations, or MEG,
where the equipment is expensive and requires a magnetically shielded room limiting its
potential transfer out-of-lab applications. Contrarily, EEG is an affordable imaging modality
with a straightforward setup which provides high temporal resolution and portability. However,
the inconvenience of using EEG is low spatial resolution and low SNR. Despite this, decoders
have been commonly employed in BCI design employing parieto-occipital changes in α-band
activity to predict covert visuospatial attention tasks (Tonin et al. 2013; Treder et al. 2011).
The integrated approach between frontal and parieto-occipital attentional decoding, however,
has not been attempted. Here, we found that cue-locked synchronisation enclosed in the FPN
α-band is insufficient to determine the attentional location. This may be due to an inherent
lack of connectivity in the cue-target interval, or else more likely the inability of EEG to register
synchronisation patterns due to the aforementioned limitations.

Another potential cause why cue-locked FPN connectivity was not sufficient for classifying
single trials may be due to the change in PLV calculation from average to single-trial.
Standard cognitive research employs multiple trials to estimate consistent findings on
electrophysiological markers (M/EEG). BCIs, need by design to perform estimates robustly
and accurately in a single-trial fashion and thus require a trade-off between spatial (i.e.,

single-channel decoding is preferred) and temporal resolution. PLV is a measure of
consistency across multiple trials and cannot serve as a single-trial control signal. Therefore,
we computed PLV across time points within the same realisation. This new measure is also
referred to in the literature as the inter-site phase clustering (ISPC) and may represent a
different underlying process than that captured by classic PLV (Cohen 2015). This prompts
the question of whether long-range α-synchronisation is incapable of decoding the attended
location, or rather the single-trial nature of IPSC over time is responsible for this.

In sum, long-range α-synchronisation within the FPN estimated with EEG may not serve as
a control signal for BCI. This limitation may be due to incomplete information on neural
correlates due to the lack of cross-frequency analysis or the computational techniques
surrounding ISPC over time.

CONCLUSION

We found direction-specific contralateral patterns of upper α-synchronisation (i.e., PLV) within
the FPN following target appearance in a covert visuospatial task. This finding, however, did
not extend to pre-target or cue-target time windows. The modulatory role of α-synchronisation
in anticipatory attention through frontal, parietal and occipital regions suggests that PLV may
not constitute a reliable metric for this top-down visual processing. Furthermore, chance-level
classification resulting from using this metric in an SVM indicates that long-range α-
synchronisation computed with EEG may not be a suitable control signal for BCI.

CONFLICT OF INTEREST

593 Authors report no conflict of interest

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FIGURES

A

B

Figure 1. Experimental design and response rates. (A) Schematic trial representation. A black fixation cross in the middle of the screen and two squares (to-be-attended locations) at the bottom left, and bottom right positions were displayed continuously. At the beginning of each trial, participants were instructed to gaze at the fixation cross. After 200 ms (fixation period), an auditory cue appeared for 100 ms (cue period) indicating which hemifields participants must attend (75% validity). After a jittered interstimulus interval of 2000 ± 500 ms, a target appeared at the targeted location during 50 ms (target period). Participants had to report first if they had seen the target (detection task), and after 1000 ms, the location of the target (left/right discrimination task) during 1500 ms. An intertrial interval (ITI) of 1000 ms followed, and a new trial began (Adapted from Torralba et al. 2016). **(B) Response rates for detected and discriminated trials (HITS) related to attended and unattended trials.** Black lines over violin plots represent the mean value. Both overall performance (top) and right/left hemifields (bottom) are shown. White dots indicate individual values (adapted from Torralba et al. 2016).

Figure 2. Target-locked results. (A) Target-locked results of the phase-coupling for *attended left* (light blue) and *attended right* (dark blue) in FM-PL and FM-PR networks. The lower panels depicts the cross-trial average time course (± shaded SEM) of PLV in both conditions (*attended left* and *attended right*). Upper panels present the binned violin plots (mean and median) of the pre-target window (-200 to 0 ms) and the post-target window (200 to 400 ms); *$p < 0.05$. **(B) Target-locked results collapsed as either ipsilateral (FM-PL network and *attended left*; FM-PR and *attended right*) or contralateral (FM-PR network and *attended left*; FM-PL and *attended right*).** The lower panel shows the cross-trial average time course (± shaded SEM) of PLV in ipsilateral (light grey) and contralateral (dark grey) conditions. The upper panel exhibits the distribution of individual PLV with a violin plot, superimposed by the mean and the contra- to ipsilateral differences between individual PLV; ***$p < 0.001$.

Figure 3. Cue-locked results. (A) Group-level results of upper-alpha PLV. Upper panel shows phase coupling for ipsilateral (light grey) and contralateral (dark grey) sides in time-windows of 200 ms from the cue-locked interval (500 ms to 1500 ms after cue presentation). Lower panel shows mean and standard error of the mean (SEM) of the PLV values. **(B) Exploratory analysis of PLV differences.** Group-level temporal evolution of the z-scored difference between contralateral and ipsilateral PLV for each frequency band (2.4 - 42 Hz with 16 logarithmic steps). Z-score values range from -0.03 to 0.03.

Figure 4. Classification outcomes. (A) Cross-time PLV reality check. Replication of results from Fig. 2 calculating PLV across time points rather than across trials. (B) Optimisation results of gamma and margin parameters of the Gaussian kernel SVM. Ten-fold validation accuracies with varying margin values (x-axis) and gamma values (y-axis). Inset shows a detailed view of the z-axis. (C) Confusion matrices of the classification outcomes for each participant. Y-axis represents ground truth labels (*attended right* or *attended left*) and x-axis represents the classifier's outcomes. Percentages represent the fraction of correctly classified trials of each condition (i.e., each row sums to 100%). Under the percentage is the gross number of classified trials.

Figure 5. Lateralisation index reality check. (A) Averaged lateralization index for *attended left* (light blue) and *attended right* (dark blue; *$p > 0.05$). White dots denote individual scores and horizontal line indicates the group mean. **(B) Lateralisation index (mean ± SEM) over time.** Solid lines and shaded areas represent mean and standard error of the mean (SEM) interval, respectively. Dots on in the x-axis denote the significant difference over time between *attended left* (light blue) and *attended right* (dark blue) via cluster-based permutation test. **(C-D) Lateralisation indexes and the difference of contra- to ipsilateral PLV for *attended left* (light blue) and *attended right* (dark blue) at the pre-target window (C) and the post-target window (D).** At the pre-target the correlations for *attended right* ($r = -0.25$, $p > 0.05$) and *attended left* ($r = -0.13$, $p > 0.05$) did not reach significance and neither did the correlations for *attended right* ($r = -0.44$, $p > 0.05$) and *attended left* $r = -0.42$, $p > 0.05$) at the post-target window. Crosses denote participants with a significant effect in PLV contra-ipsi differences at the pre-target window (-200 to 0 ms; P04, P05, and P10) and the post-target window (200 to 400 ms; P04, P05, P06, P07, and P10). Dots represent the rest of the participants.

EXTENDED FIGURES

Figure 2-1. Individual results of target-locked PLV index. Violin plots represent the phase locking values (PLV) averaged over the pre-target (-200 to 0 ms, t = 0 as target appearance) and post-target time window (200 to 400 ms). Ipsilateral (FM-PL network and *attended left*; FM-PR and *attended right*) or contralateral (FM-PR network and *attended left*; FM-PL and *attended right*) scenarios are exhibited as either light grey or dark grey, respectively. *$p < 0.05$, **$p < 0.01$, ***$p < 0.001$.

Figure 3-1. Individual results of upper-alpha cue-locked PLV analysis. Violin plots represent the phase locking values (PLV) averaged over the five time windows (500 to 700, 700 to 900, 1100 to 1300, and 1300 to 1500 ms; t = 0 as cue appearance). Ipsilateral or contralateral scenarios are exhibited as either light grey or dark grey, respectively. *$p < 0.05$.

912
913 **Figure 3-2. Individual results of cue-locked exploratory PLV analysis.** Differences of contra- to
914 ipsilateral PLV are represented over frequencies (2.4 – 42 Hz in 16 logarithmic steps) as a percentage
915 of change regarding the corss-frequency mean of each individual.

916
917 **Figure 4-1. Individual results of target-locked cross-time PLV.** Violin plots represent the phase
918 locking values (PLV) obtained by calculating PLV as consistency throughout the pre-target (-200 to 0
919 ms) and post-target (200 to 400 ms) time windows. Ipsilateral or contralateral scenarios are exhibited
920 as either light grey or dark grey, respectively.

Figure 5-1. Individual results of lateralisation index. Violin plots represent the averaged lateralised index for *attended left* (light blue) and *attended right* trials (dark blue) over the cue-locked time window. Shaded plots represent lateralisation over time (mean ± SEM). Dots on in the x-axis denote the significant differences over time between *attended left* and *attended right* via cluster-based permutation test. *$p < 0.05$, **$p < 0.01$, ***$p < 0.001$.