

# Nonlinear Interaction Decomposition (NID): A Novel Framework for Studying Cross-Frequency Coupling in Human Brain

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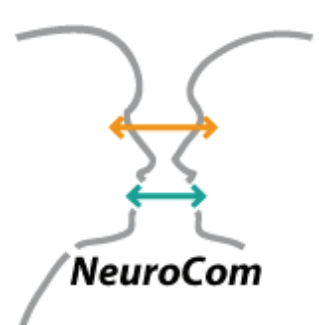
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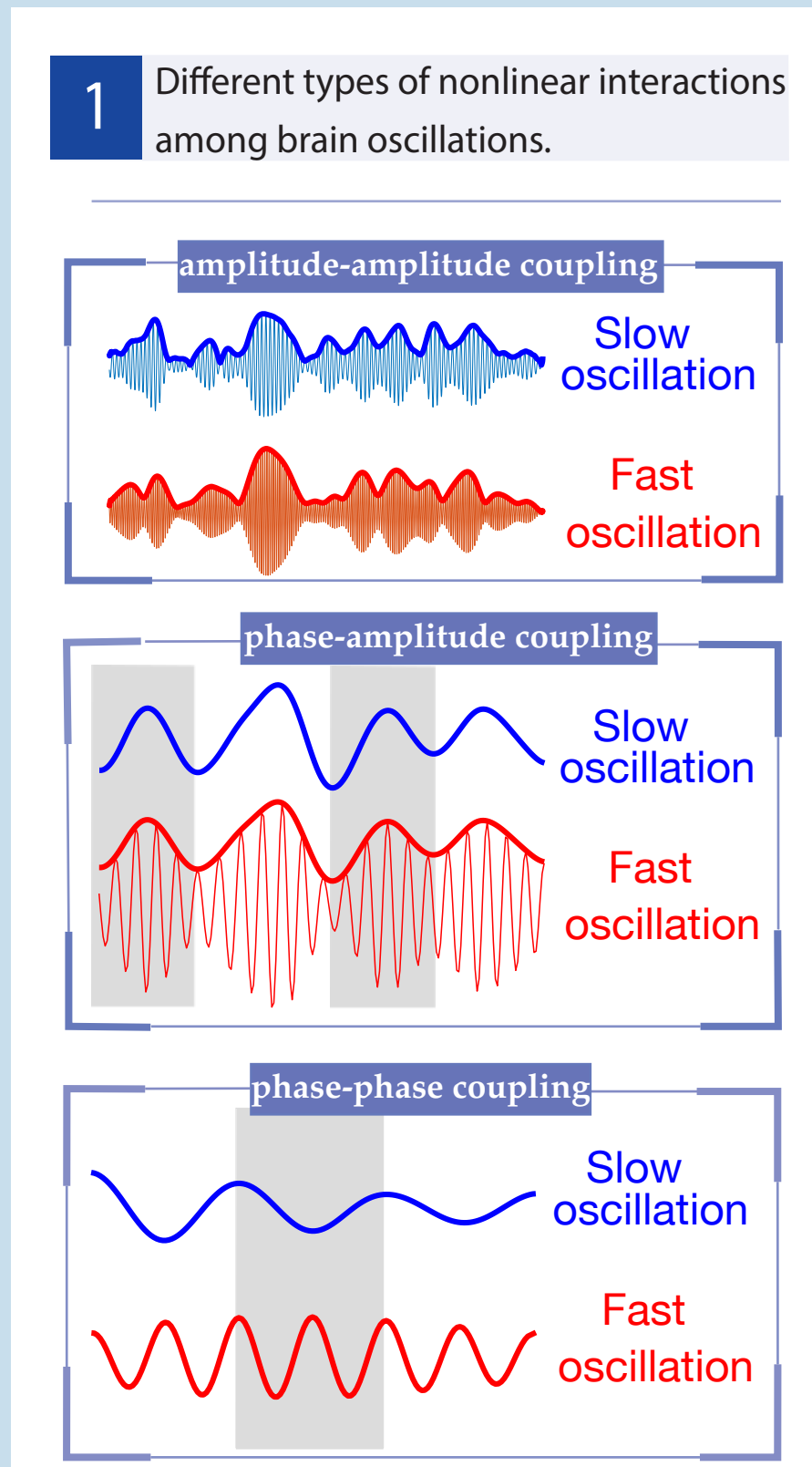


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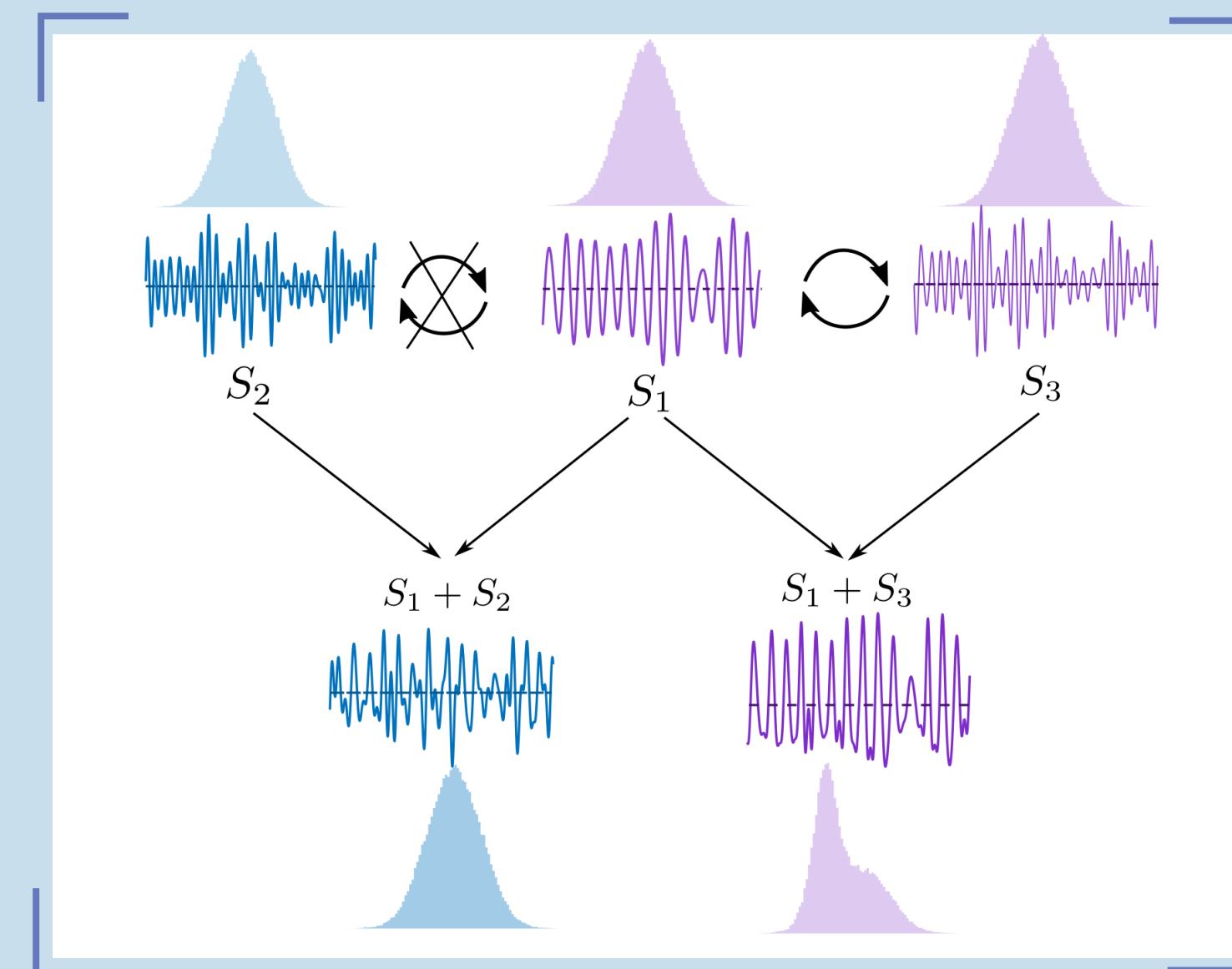
## Introduction

- A group of activated neurons that synchronize rhythmically produce oscillations (Singer, 1993).
- EEG/MEG record a mixture of all activities in the brain. Therefore, studying source signals from specific brain regions requires some kind of source separation method.



- Oscillatory neural populations interact within one frequency band (Engel and Fries, 2010), or across different frequency bands (Palva & Palva, 2018).
- Cross-frequency coupling enables integration of spectrally distributed information processing in the brain.
- Separation of cross-frequency coupled oscillations is challenging.
- We propose a novel method for extraction of nonlinearly interacting oscillations in the brain.

## NID's Main Idea



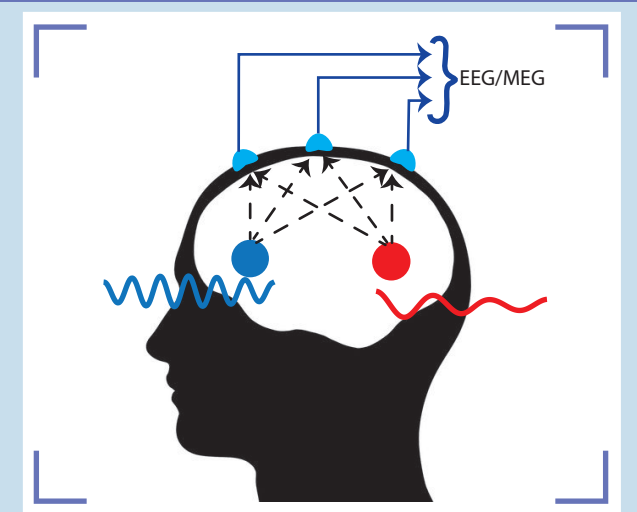
### WORKING PRINCIPLES:

- Narrow-band oscillations in the brain ( $S_1, S_2, S_3$ ) is approximately Gaussian distributed.
- Linear mixture of two independent narrow-band oscillations ( $S_1, S_2$ ) has Gaussian distributed.
- Linear mixture of phase-phase coupled oscillations ( $S_1, S_3$ ) has non-Gaussian distribution.

## Method

### Problem Formulation

- Source model:  $\mathbf{X} = \mathbf{P}^{(n)}\mathbf{S}^{(n)} + \mathbf{P}^{(m)}\mathbf{S}^{(m)} + \xi$   
multi-channel observed signal  $\xrightarrow{C \times T}$   $\xrightarrow{C \times N}$   $\xrightarrow{N \times T}$   $\xrightarrow{N \text{ activation patterns of the oscillations at } f_n}$   $\xrightarrow{N \text{ oscillations at } f_n}$  frequency  $f_m$
- Goal: extraction of oscillations and activations patterns at  $f_n$  and  $f_m$



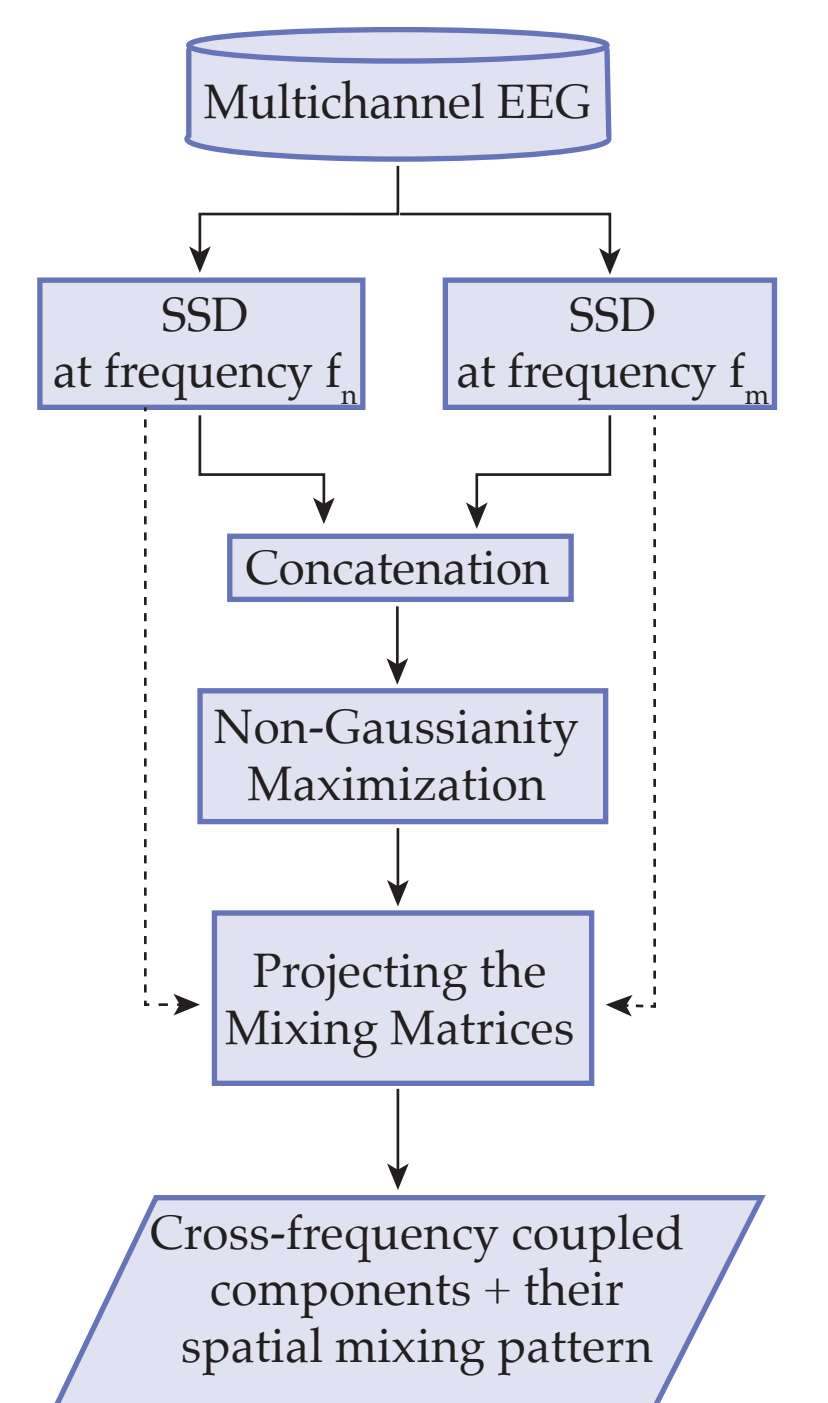
### Nonlinear Interaction Decomposition (NID)

- SSD decomposition:  $\mathbf{X}^{(n)} = \mathbf{A}_{ssd}^{(n)}\mathbf{X}_{ssd}^{(n)}$   
 $\xrightarrow{N \text{ SSD activation patterns of the oscillations at } f_n}$   $\xrightarrow{N \text{ SSD components at } f_n}$
- Concatenation:  $\mathbf{X}_{Aug} = [\mathbf{X}_{ssd}^{(n)}, \mathbf{X}_{ssd}^{(m)}]$
- Non-Gaussianity maximization:  $\mathbf{X}_{Aug} = \mathbf{A}_{NG}\mathbf{R}$   
 $\xrightarrow{2N \times T}$   $\xrightarrow{2N \times N}$   $\xrightarrow{\mathbf{W}_{NG} = [\mathbf{W}_{NG}^{(n)}, \mathbf{W}_{NG}^{(m)}]}$   $\xrightarrow{\mathbf{A}_{NG} = [\mathbf{A}_{NG}^{(n)}, \mathbf{A}_{NG}^{(m)}]}$
- NID activation pattern:  $\mathbf{P}_{NID}^{(n)} = \mathbf{A}_{ssd}^{(n)}\mathbf{A}_{NG}^{(n)}$
- NID source signals:  $\mathbf{S}_{NID}^{(n)} = \mathbf{W}_{NG}^{(n)}\mathbf{X}_{ssd}^{(n)}$

### Spatio-Spectral Decomposition (SSD)

- SSD is a method for extraction of oscillations at a specific frequency band (Nikulin et al, 2011). It finds the spatial filters maximizing the SNR at a defined frequency band.

### 2 Block-diagram of NID

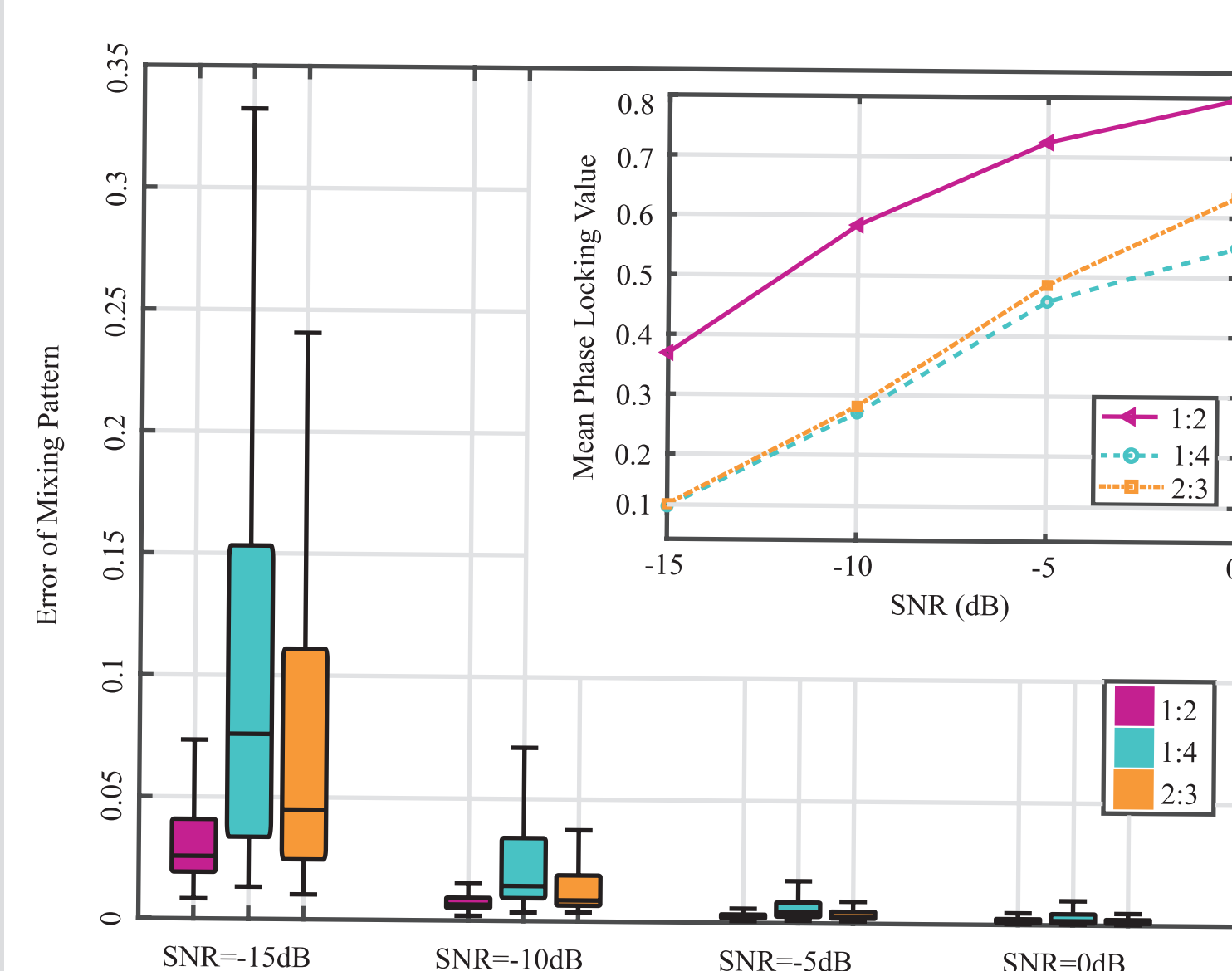


## Results

### Simulated Data

- Simulated data with 2 pairs of oscillations with phase-phase coupling at frequency  $f_n = n f_b$  and  $f_m = m f_b$  (denoted by n:m coupling).
- Error of mixing pattern is the dissimilarity between the recovered and original activation patterns.
- Figure 3: The low median error of extraction of mixing patterns and the high mean PLV of the extracted oscillations show the reliability of NID.
- Other additional simulations are done with 5 pairs of coupled source signals.
- We did the same simulations for extraction of amplitude-amplitude coupled oscillations and comparable results were achieved.

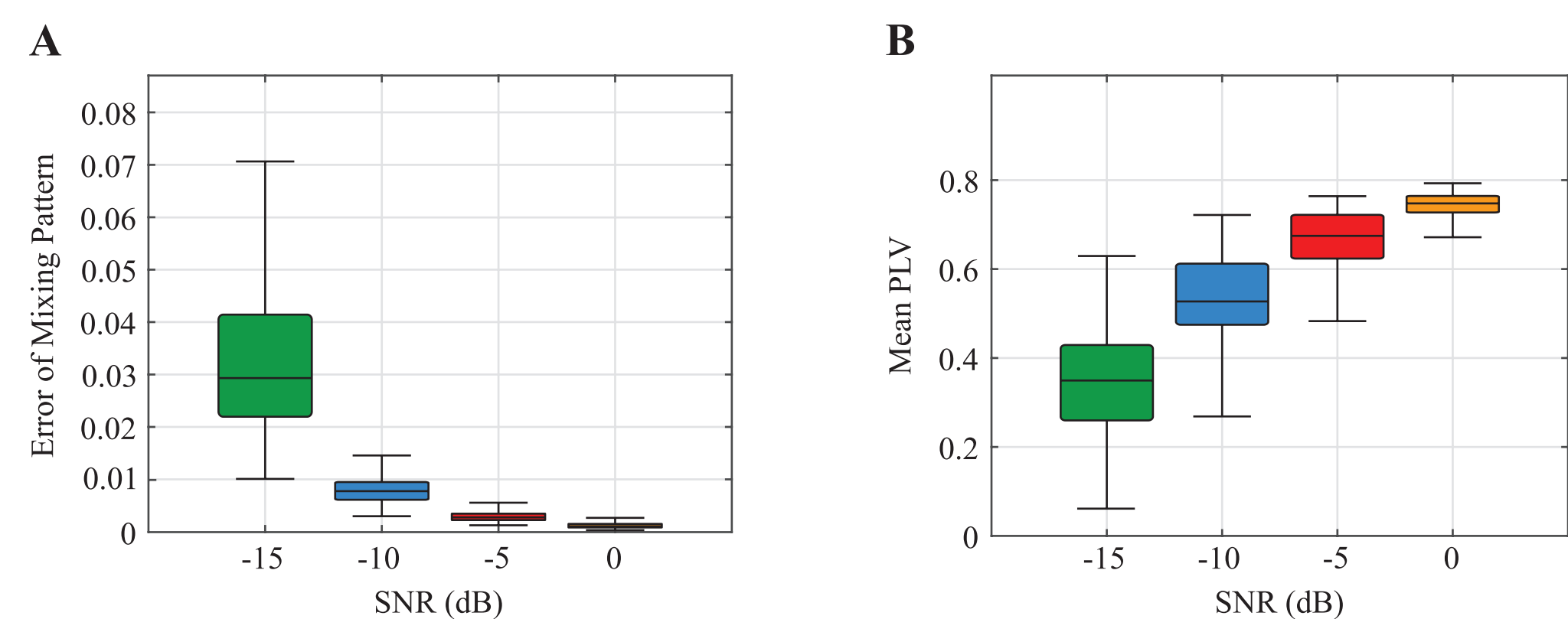
3 Main plot: Error of mixing pattern extraction for simulated data. Subplot: mean PLV of the extracted coupled source signals.



### Extension of NID for extraction of n:m:r coupling:

- The augmented matrix is built from concatenating the SSD components of the three frequency bands.
- Simulated EEG data with 2 triplets of source signals at 10Hz, 20Hz, and 30Hz.
- Figure 4: the median error of mixing patterns is small and the mean PLV is large for low SNRs.

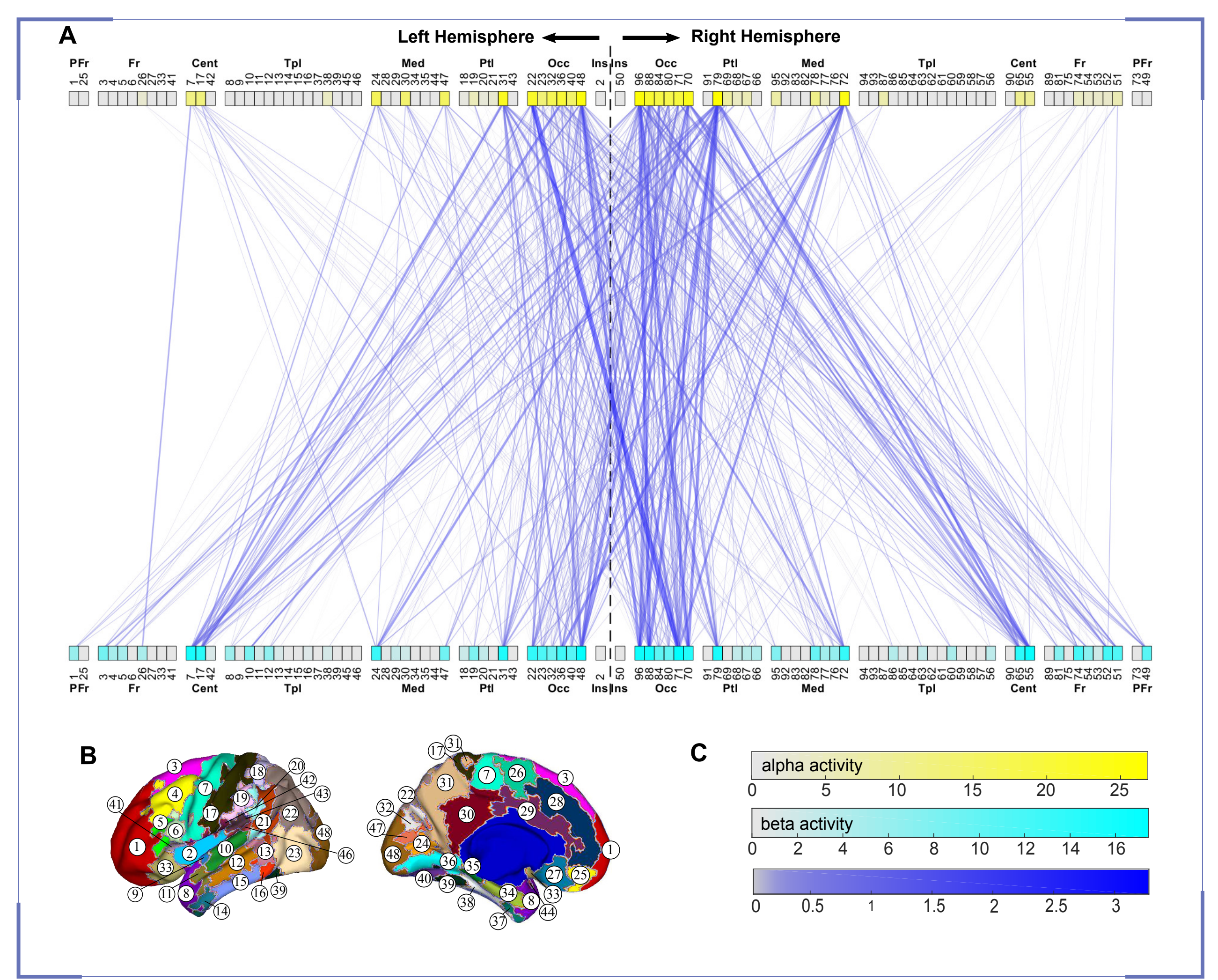
4 NID for extraction of triplets of coupled oscillations at 10Hz, 20Hz, and 30Hz. A: Error of mixing pattern extraction for simulated data. B: mean PLV of the extracted coupled source signals.



### Resting-state EEG

- Alpha-beta interaction of resting-state EEG data of 82 subjects from LEMON dataset (Babayan et al, 2018).
- Many diverse interactions are extracted by NID. A particularly interesting coupling is between alpha activity of occipital area and beta activity of motor area.

- A bipartite graph (panel A) illustrating the alpha-beta interactions between ROIs of Harvard-Oxford atlas (panel B), computed for 82 subjects.
- A connection between node r1 of the upper and node r2 of the lower part indicates that alpha oscillations in ROI r1 interact with beta-oscillations in ROI r2.
- The edge weights are proportional to the number of active voxels (across subjects) in the two ROIs. Color-codes in panel C.



## Conclusion

- Former methods maximize a contrast function based on the desired coupling. NID is based on the statistical properties of the source signals.
- NID is generalizable for detection of coupling of oscillations in multiple (>2) frequency bands. No other method can be generalized in this way.
- NID can extract a subset of coupled activities in the brain that can be used for connectivity analysis.
- Future work: investigating how brain networks of resting-state EEG are predictive of task performance.

## References

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