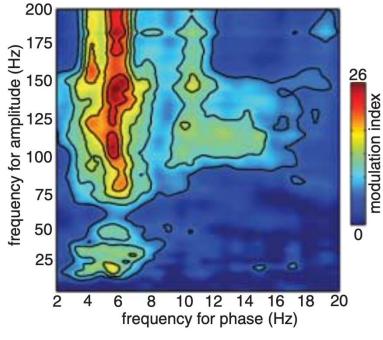
# Cross-frequency coupling

Instructor: Mark Kramer

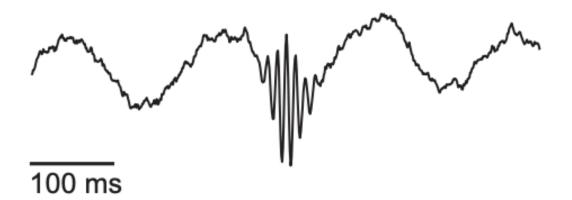
## Today

Cross-frequency coupling (one type of)

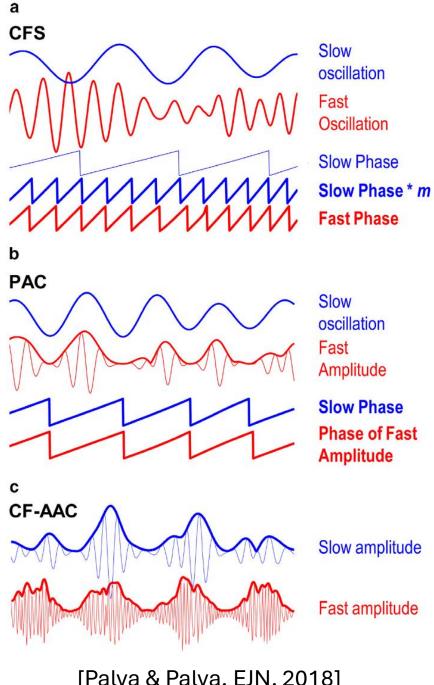
## Examples



[Canolty et. al., Science, 2006]



[Tort et. al., PNAS, 2008]



[Palva & Palva, EJN, 2018]

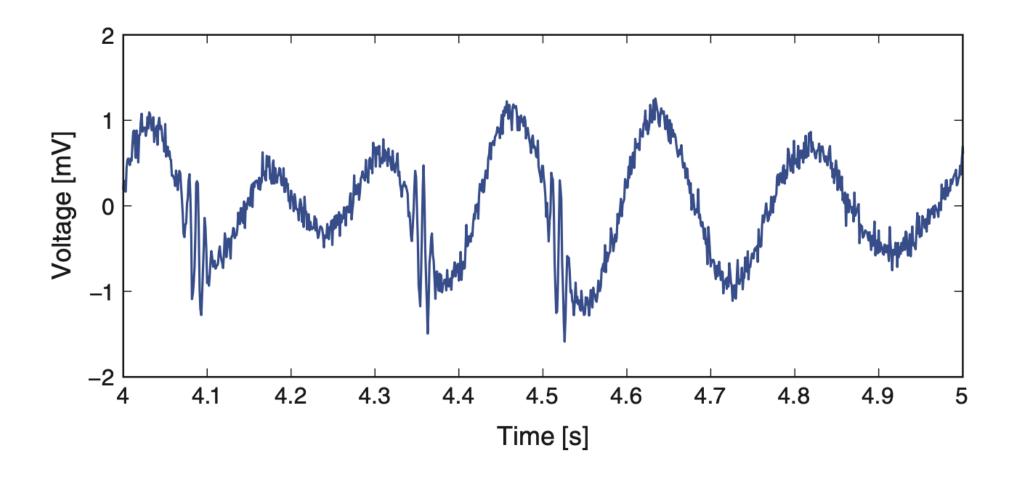
### Outline

**Data** 100 s of local field potential data sampled at 1000 Hz.

Goal Characterize the coupling between rhythms of different frequency.

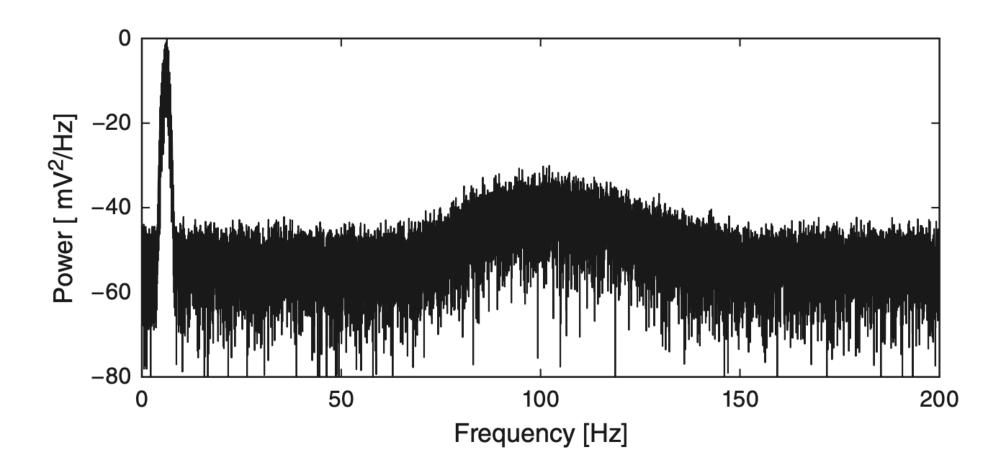
**Tools** Hilbert transform, analytic signal, instantaneous phase, cross-frequency coupling.

## Data



Q. How to make sense of these data?

## Spectrum



Q. What do you see?

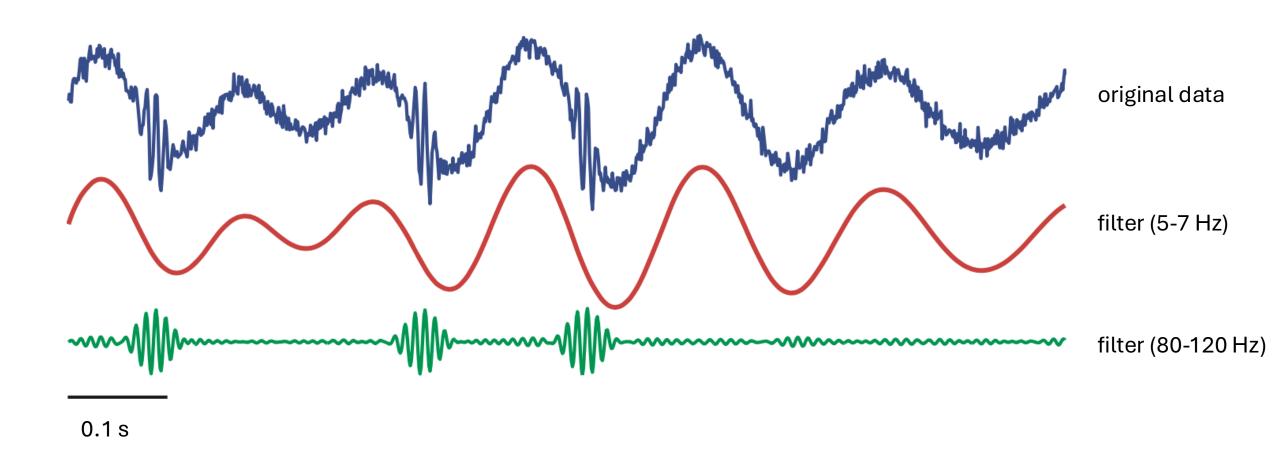
## CFC in three steps

#### **CFC** analysis steps

- 1. Filter the data into high- and low-frequency bands.
- 2. Extract the amplitude and phase from the filtered signals.
- 3. Determine if the phase and amplitude are related.

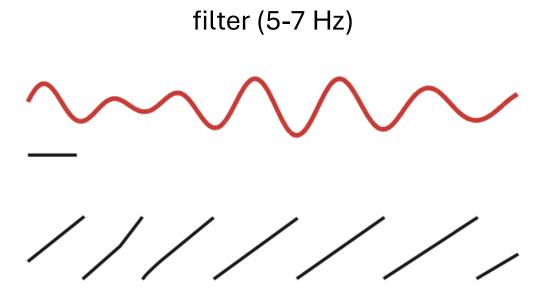
Let's perform each step ...

Filter the Data into High- and Low-Frequency Bands



**Q.** Why did we filter in these bands?

Extract the Amplitude and Phase from Filtered Signals



phase (of low frequency band)

filter (80-120 Hz)



amplitude envelope (of high frequency band)

Q. How?

$$y = H(x)$$

$$H(x) = \begin{cases} -\pi/2 \text{ phase shift if } f > 0, \\ 0 \text{ phase shift if } f = 0, \\ \pi/2 \text{ phase shift if } f < 0. \end{cases}$$

The Hilbert transform H(x) of the signal x produces a phase shift of  $\pm 90$  degrees for  $\mp$  frequencies of x.

**Define**: Analytic signal z

$$z = x + iy = x + iH(x)$$

<u>Impact</u>: remove negative frequencies from *z* 

**Q.** How?

**Q.** What does it do?

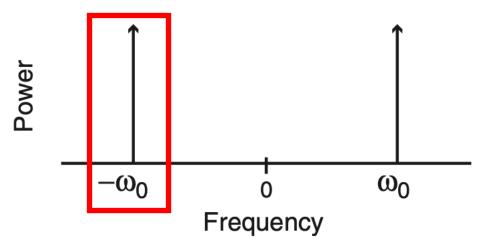
Ex.

$$x_0 = 2\cos(2\pi f_o t) = 2\cos(\omega_0 t)$$
 where  $\omega_0 = 2\pi f_o$ 

Euler's formula

$$x_0=e^{i\omega_0t}+e^{-i\omega_0t}$$
 positive frequency negative frequency

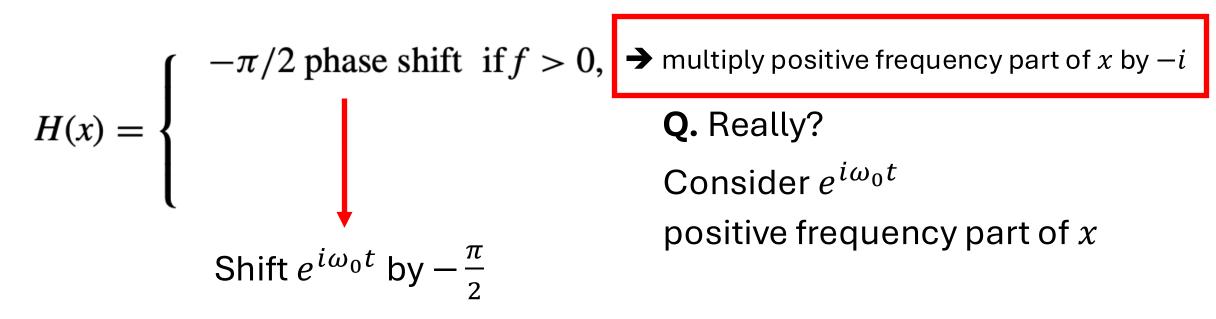
we usually ignore this one



Note: The spectrum has two peaks

Apply the Hilbert transform to  $x_0$ .

$$x_0 = e^{i\omega_0 t} + e^{-i\omega_0 t}$$



$$\rightarrow e^{i(\omega_0 t - \frac{\pi}{2})} \rightarrow e^{i\omega_0 t} e^{-i\pi/2} \rightarrow e^{i\omega_0 t} (\cos \frac{\pi}{2} - i \sin \frac{\pi}{2}) \rightarrow e^{i\omega_0 t} (-i)$$

Apply the Hilbert transform to  $x_0$ .

$$x_0 = e^{i\omega_0 t} + e^{-i\omega_0 t}$$

$$H(x) = \begin{cases} -\pi/2 \text{ phase shift if } f > 0, \\ 0 \text{ phase shift if } f = 0, \\ \pi/2 \text{ phase shift if } f < 0. \end{cases} \rightarrow H(x) = \begin{cases} -ix & \text{if } f > 0, \\ 0 & \text{if } f = 0, \\ ix & \text{if } f < 0. \end{cases}$$

So, 
$$x_0 = e^{i\omega_0 t} + e^{-i\omega_0 t}$$
  $y_0 = H(x_0) = -ie^{i\omega_0 t} + ie^{-i\omega_0 t}$  Euler's formula 
$$= 2\sin(\omega_0 t)$$

Hilbert Transform of  $x_0$  (a cosine function) is a sine function.

Analytic signal z

$$z = x + iy = x + iH(x)$$

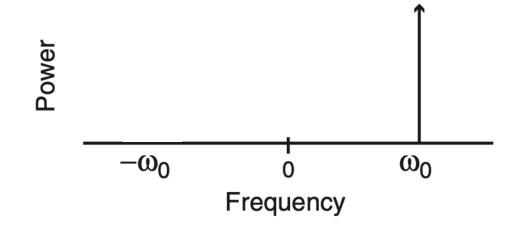
$$\uparrow \qquad \uparrow$$

$$2\cos(\omega_0 t) \quad 2\sin(\omega_0 t)$$

$$= 2\cos(\omega_0 t) + i 2\sin(\omega_0 t)$$

$$=2e^{i\omega_0t}$$

The analytic signal contains no negative frequencies



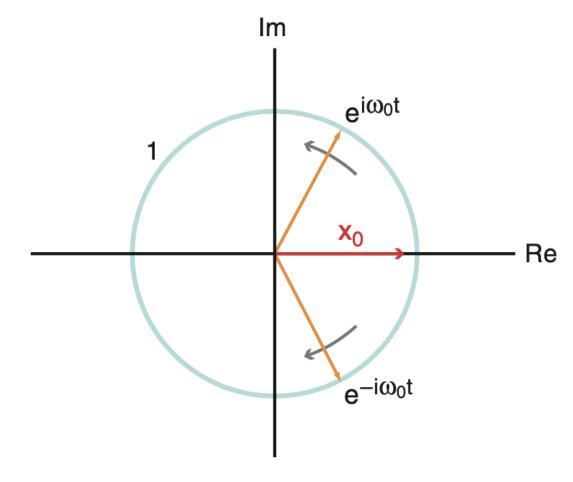
Original signal 
$$x_0 = 2\cos(2\pi f_0 t) = e^{i\omega_0 t} + e^{-i\omega_0 t}$$

Complicated (2 complex exponentials)

Analytic signal  $z_0 = 2e^{i\omega_0 t}$ 

Simple (1 complex exp)

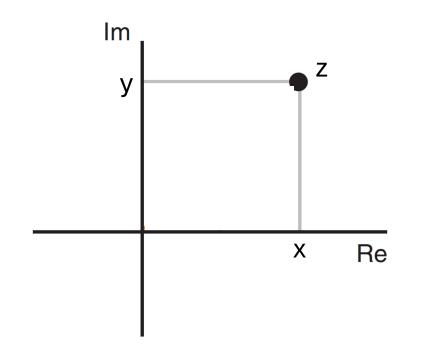
A point in the complex plane



<u>Analytic signal</u>

$$z = x + iy$$

A point in the complex plane



$$z(t) = A(t) e^{i \phi(t)}$$
amplitude phase

Get the **amplitude** and **phase** from the analytic signal

Ex.

$$z_0(t) = 2e^{i\omega_0 t} \qquad A(t) = 2 \qquad \phi(t) = \omega_0 t$$

## CFC in three steps

#### **CFC** analysis steps



1. Filter the data into high- and low-frequency bands.



2. Extract the amplitude and phase from the filtered signals.

3. Determine if the phase and amplitude are related.

Determine if the Phase and Amplitude are Related

Define the two-column vector

$$\begin{pmatrix} \phi(1) & A(1) \\ \phi(2) & A(2) \\ \phi(3) & A(3) \\ \vdots & \vdots \end{pmatrix}$$

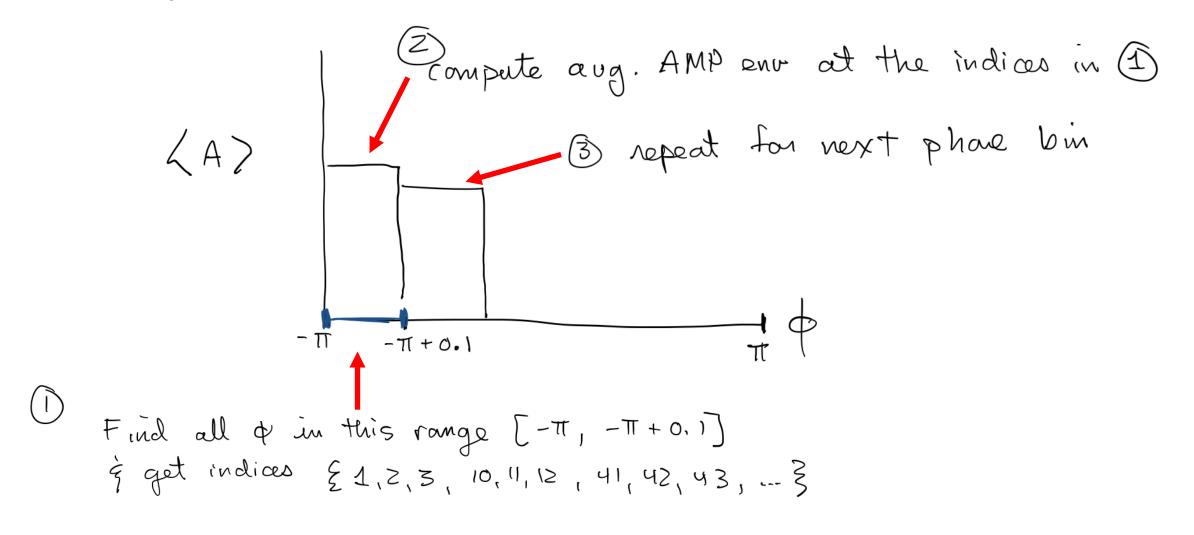
phase of low frequency band activity

amplitude of high frequency band activity

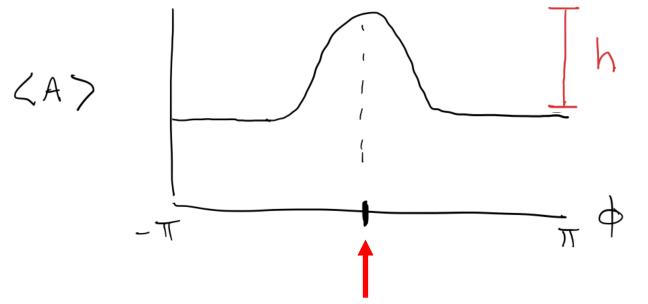
Make a histogram

Determine if the Phase and Amplitude are Related

Divide the phase into bins of size 0.1



If phase modulate amplitude



summarize extent of modulation

**Q.** What does no phase-amplitude coupling look like in the plot?

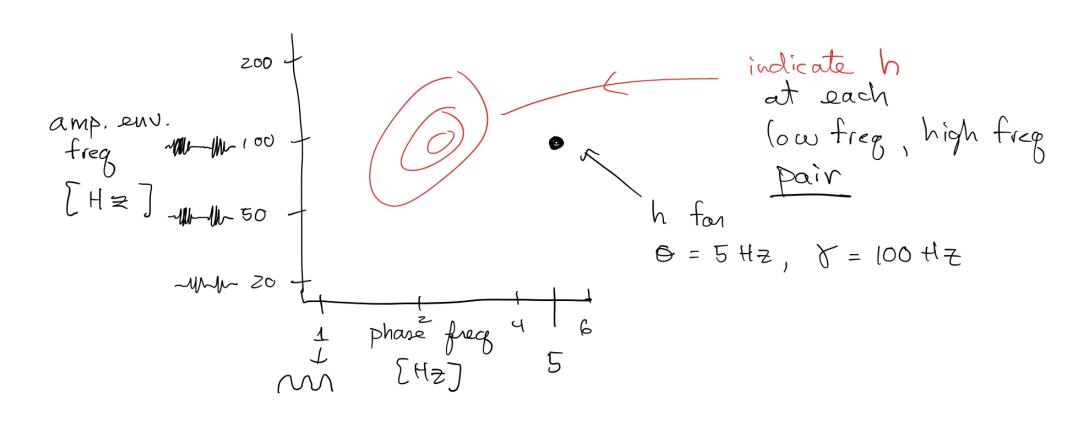
At this phase, amplitude envelope is big

→ phase modulates amplitude

## CFC - Step 4

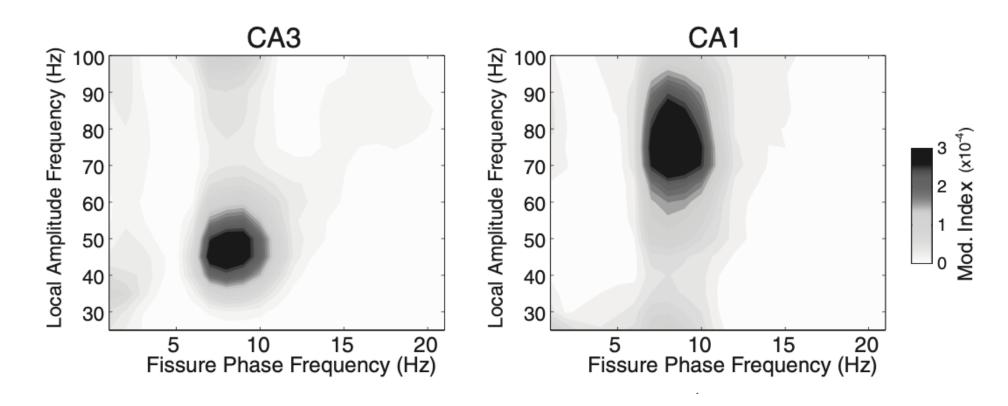
(optional): Repeat for other frequencies.

#### Summarize in a comodulogram



(optional): Repeat for other frequencies.

#### Summarize in a <u>comodulogram</u>

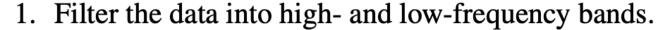


[Tort et al., J Neurophysiol, 2010]

## CFC in three steps

#### **CFC** analysis steps







2. Extract the amplitude and phase from the filtered signals.



3. Determine if the phase and amplitude are related.

### Python