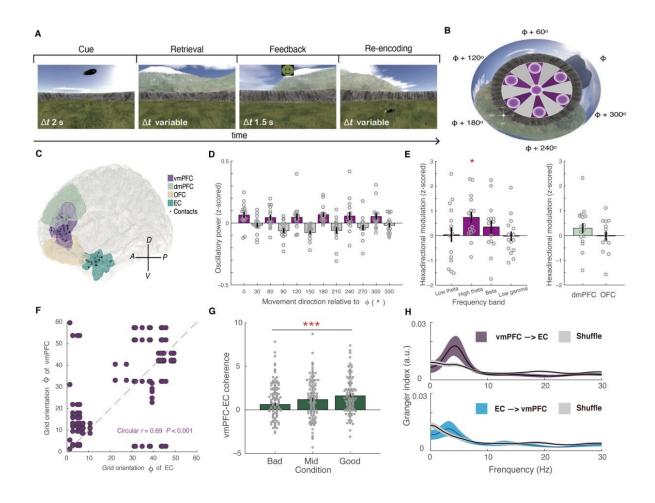


Certain brain rhythms coordinate cognitive map in human spatial navigation

October 27 2021



(A) Object-location memory task; (B) Schematic depiction of firing fields for one grid cell (purple circle). More fields are crossed during six moving directions which are aligned with the grid (purple), translating to stronger brain activity. (C) Depiction of all electrode contacts in mPFC and EC (black circles) relative to two control regions (dmPFC and OFC). (D) Theta power was higher during movements aligned with the grid axes as compared to misaligned movements.



Purple, aligned; gray, misaligned. (E) Hexa-directional modulation is unique to theta oscillation and mPFC. (F) mPFC and EC show similar grid orientations. (G) Theta coherence is significantly stronger in better spatial memory performance trials. (H) Granger causality analyses show significant influence from mPFC to EC (top) but not in the reverse direction (bottom) during memory retrieval. The gray lines denote 95% confidence intervals of the null distribution. Credit: Dong Chen

Chinese researchers have reported that slow rhythmic fluctuations in brain activity—theta oscillations in the medial prefrontal cortex (mPFC)—entorhinal cortex (EC) circuit—play an essential role in enabling spatial computations to guide human navigation behavior.

Led by Dr. Wang Liang from the Institute of Psychology of the Chinese Academy of Sciences (CAS), the researchers found that theta oscillations in the mPFC and EC exhibited grid-like representations that coordinated with each other and predicted navigational performance.

The study provides novel insights into the neural signature of the cognitive map and the functional role of grid codes during human spatial navigation.

The <u>human brain</u> creates a cognitive map of the surrounding environment to efficiently support spatial memory and guide flexible behavior in physical space. Place cells in the hippocampus and grid cells in the EC are the cellular components of the neuro navigation system. Grid cells have been identified in human EC and mPFC with a crucial role in conceptual space.

Individual neurons never fire in isolation and are not able to regulate complex cognitive behaviors, such as spatial navigation. Instead, synchronized firing neurons induce neural oscillations and form a



functional network.

Theta oscillations—slow, periodically fluctuating waves of neuronal activity—have an essential link to the grid cell assembly that has been well established using local field potential recordings. In 2018, the researchers revealed theta-based grid-like representations in human EC during spatial navigation.

In the current study, the researchers recorded mPFC and EC neural activity in drug-resistant epilepsy patients using depth electrodes invasively implanted in the brain by neurosurgeons, while patients played an object-location memory task in a virtual environment.

The patients were asked to remember the spatial location of eight objects and were tested on memory performance. They were able to utilize feedback information to improve their behavioral performance.

The researchers found that <u>theta oscillations</u> in the human ventromedial prefrontal cortex, rather than in the orbital frontal cortex or dorsal mPFC, exhibited a six-fold rotationally symmetric modulation as a function of movement direction.

"This modulation pattern provides direct evidence that the mesoscopic fingerprint of grid cells occurs in human mPFC," said Dr. Wang.

Furthermore, the researchers also found that grid orientations appeared to be consistent between the EC and mPFC, suggesting that these regions coordinated information processing to generate a coherent cognitive map for the encoding and retrieval of spatial memories.

Theta synchronization was correlated with this inter-regional consistency of grid representations and can predict the accuracy of spatial performance.



This study sheds light on the neural mechanisms underlying navigation and memory in humans, providing new insights into how mPFC–EC neural circuits at the mesoscopic level deal with <u>grid</u> information.

Many neurodegenerative diseases even in the early stage are associated with spatial disorientation and spatial memory deficits, such as Alzheimer's disease. Therefore, it is crucial to understand the underlying neuronal mechanisms. This study may help identify novel biomarkers for such neurological disorders.

This work was published in *Science Advances* on Oct. 27.

More information: Dong Chen et al, Theta oscillations coordinate gridlike representations between ventromedial prefrontal and entorhinal cortex, *Science Advances* (2021). <u>DOI: 10.1126/sciadv.abj0200</u>. <u>www.science.org/doi/10.1126/sciadv.abj0200</u>

Provided by Chinese Academy of Sciences

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