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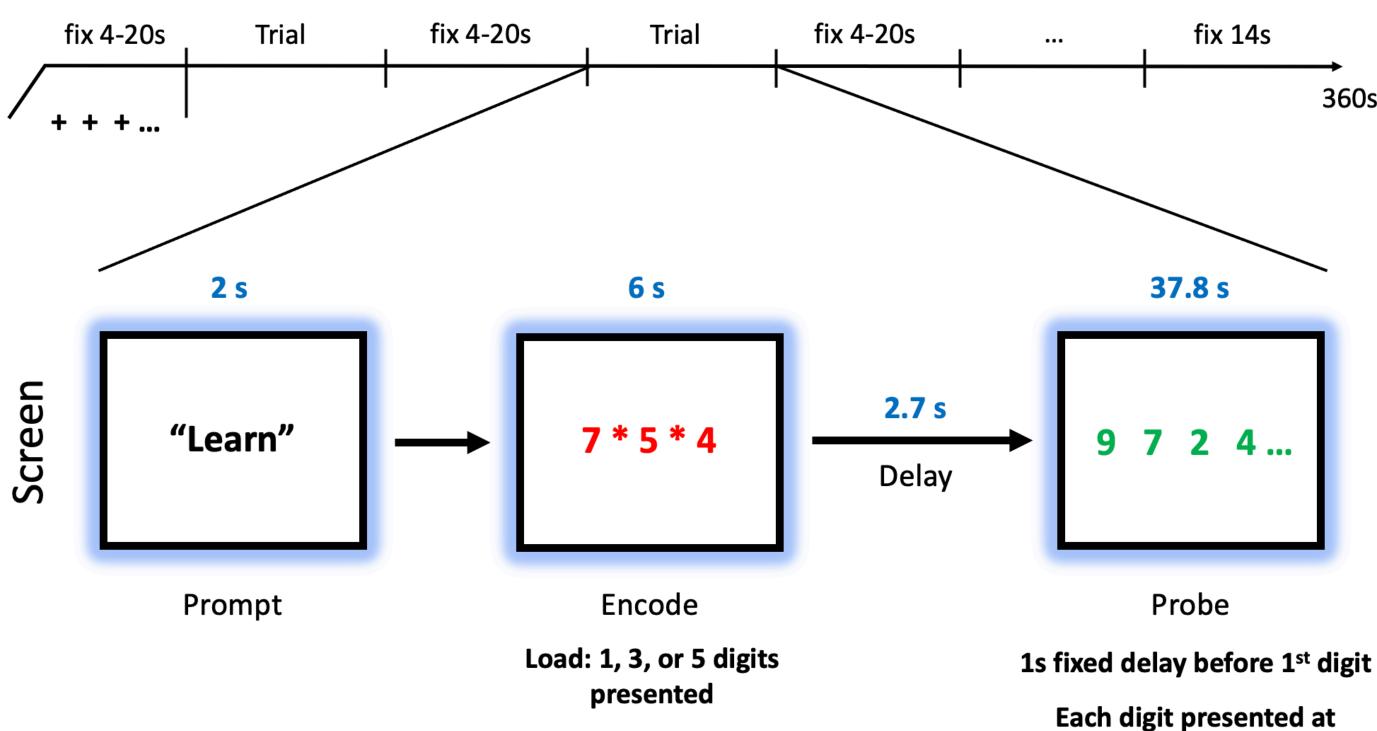
# Introduction

In cognitive neuroscience, fMRI investigation of the working memory through analysis of the Sternberg delayed recognition task has been a prominent research area for decades.<sup>[1]</sup> However, the field has yet to arrive at a consensus for a set of macroscale functional brain networks.<sup>[2]</sup> The present study utilized a version of the task called Sternberg Item Recognition Paradigm (SIRP).<sup>[1]</sup> We hypothesized the emergence of the Working Memory Big Five brain networks that were established in previous research.<sup>[3,4]</sup>

- **1. Response Network**: activates when an individual makes a response.<sup>[3,4]</sup>
- 2. Focus on Visual Features Network: is upregulated when perceived visual details are deemed important for attention.<sup>[3,4]</sup>
- **3. Initiation Network:** is elicited when restarting a required cognitive process after a short break.<sup>[3,4]</sup>
- 4. Internal Attention Network: shows activity when attention is focused on an internal representation such as working memory.<sup>[3,4]</sup>
- **5. Default Mode Network**: displays suppression until a response is made.<sup>[3,4]</sup>

# Methods

Patients with schizophrenia or schizoaffective disorder (n=87) and healthy subjects (n=79) completed the SIRP task in the fMRI scanner. They were asked to indicate if the probe digits were members of the memory set.



- Constrained Principal Component Analysis for fMRI (fMRI-CPCA) was applied on fMRI data to identify and characterize functional brain networks elicited by the task their associated hemodynamic responses (HDR).<sup>[5]</sup>
- Analysis of Variance was used to determine effects of conditions: group (healthy or schizophrenia), load (1, 3, or 5 digits), and post-stimulus time.



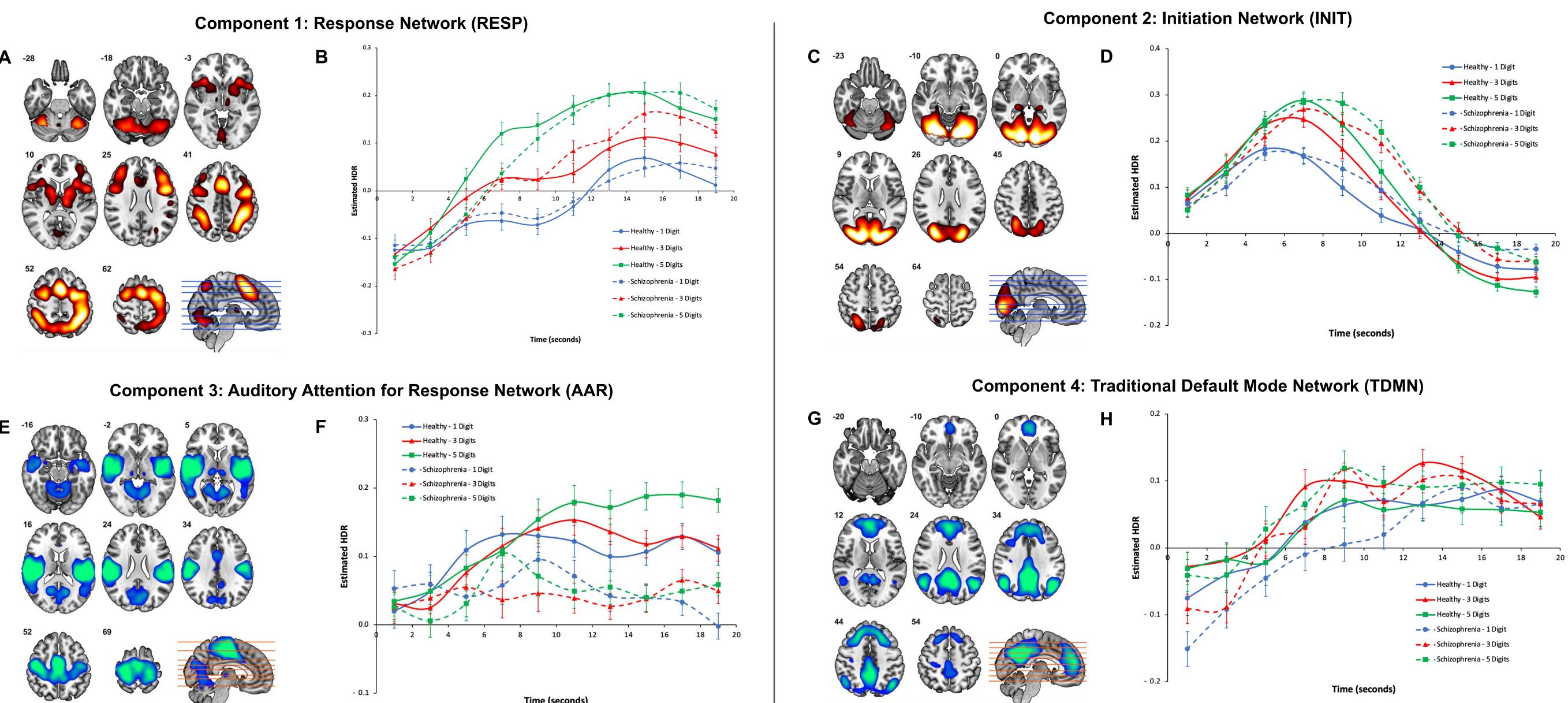
# Task-based Working Memory Functional Brain Networks Detectable by fMRI

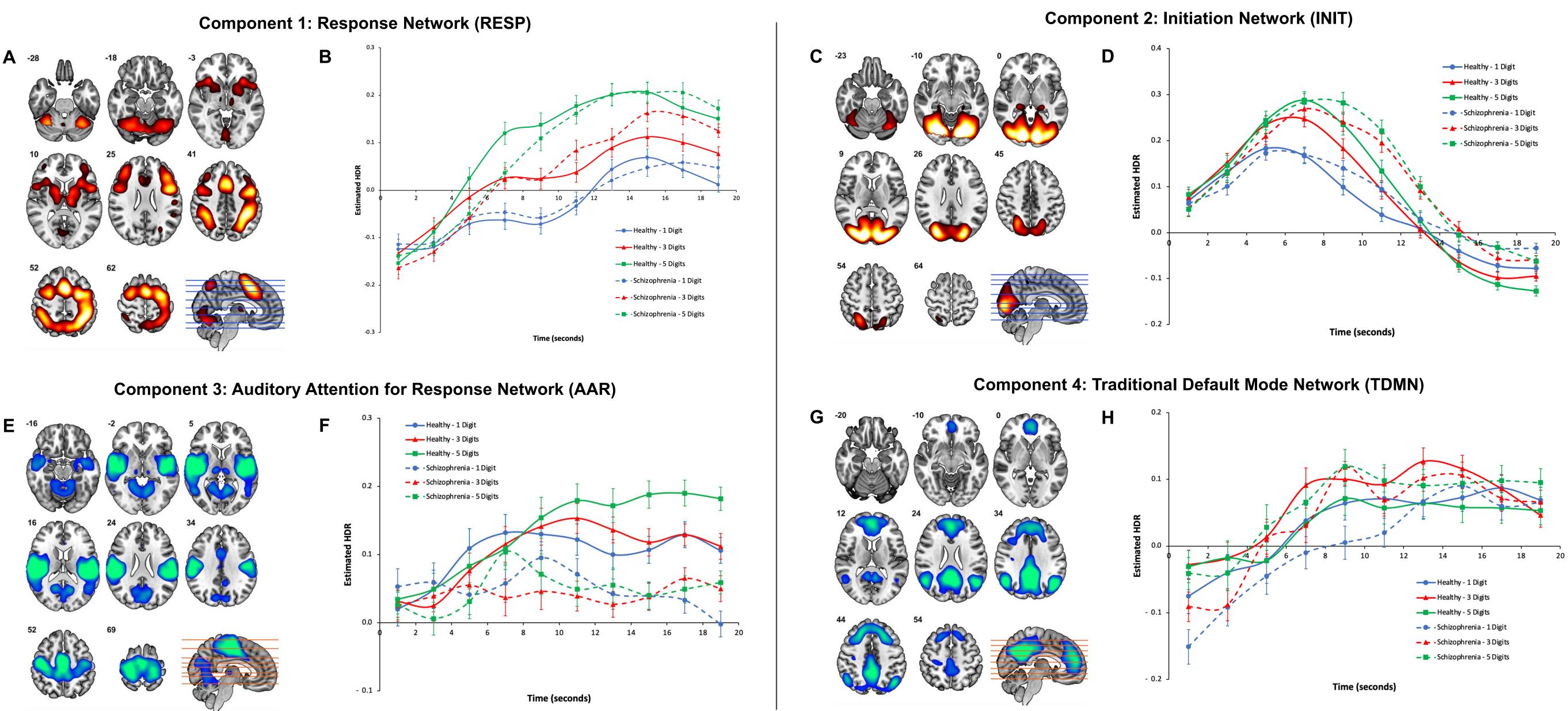
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1.1s with varying time in between (0.6 - 2.486s)

# Results

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# Discussion

- from auditory stimuli and towards visual details of digits on the screen.
- AAR network underwent deactivation when participants had to shift focus away • INIT and AAR networks have reduced activity in schizophrenia patients, implicating their dysfunction may underlie cognitive impairments in schizophrenia.
- Further studies can analyze a longer time window of the task trial to elicit all networks in the Big Five.
- Future research should investigate the effect of site difference, sex differences in brain networks, and neuropathological differences between schizophrenia and schizoaffective disorder.
- Meaningful correlations with symptoms can be established, which will contribute to the discovery and development of advanced treatment options.

### Four functional brain networks were extracted from the fMRI-CPCA analysis

Response Network and Initiation Network demonstrated load dependency: the more digits to be memorized, the higher the response. Initiation Network and Auditory Attention for Response Network showed significant difference in HDR between healthy and schizophrenia. Internal Attention Network was potentially merged into the extracted Response Network. Traditional Default Mode Network had no clear peaking or load dependency.

Figure 1. A,C,E,G: Dominant 10% of loadings for RESP, INIT, AAR, TDMN networks respectively (red/yellow = positive loadings, blue/green = negative loadings; A: threshold = 0.12, max = 0.22; C: threshold = 0.11, max = 0.36; E: threshold = -0.10, min = -0.18; G: threshold = -0.08, min = -0.16). Images are displayed in neurological orientation (left is left). B,D,F,H: Mean finite impulse response (FIR)-based predictor weights plotted over post-stimulus time by group and task condition for RESP, INIT, AAR, TDMN networks respectively. Positive predictor weights reflect increased signal in the networks. HDR = hemodynamic response.





### References



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1. Klabes, J., Babilon, S., Zandi, B., & Khanh, T. Q. (2021). The Sternberg Paradigm: Correcting encoding latencies in visual and auditory test designs. Vision, 5(2), 21. doi: 10.3390/vision5020021 2. Chai, W. J., Abd Hamid, A. I., & Abdullah, J. M. (2018). Working memory from the psychological and neurosciences perspectives: A Review. Frontiers in Psychology, 9. doi: 10.3389/fpsyg.2018.00401 Sanford, N., Whitman, J. C., & Woodward, T. S. (2020). Task-merging for finer separation of functional brain networks in working memory. Cortex, 125, 246-271. doi: 10.1016/j.cortex.2019.12.014 4. Sanford, N., & Woodward, T. S. (2021). Functional delineation of prefrontal networks underlying working memory in schizophrenia: A cross-data-set examination. Journal of Cognitive Neuroscience, 33(9), 1880-1908. doi: 10.1162/jocn a 01726

Metzak, P., Feredoes, E., Takane, Y., Wang, L., Weinstein, S., Cairo, T., ... Woodward, T. S. (2010) Constrained principal component analysis reveals functionally connected load-dependent networks involved in multiple stages of working memory. Human Brain Mapping, 32(6), 856–871. doi: 10.1002/hbm.21072

6. Keator, D. B., Van Erp, T. G., Turner, J. A., Glover, G. H., Mueller, B. A., Liu, T. T., . . . Potkin, S. G. (2016). The function Biomedical Informatics Research Network Data Repository. NeuroImage, 124, 1074-1079. doi: 10.1016/j.neuroimage.2015.09.003

