Brain Regions Guide

Explore the areas of the brain associated with each CBS Health task
A. Document Overview
B. Brain Networks Behind the Cambridge Brain Sciences Tests
C. Summary Table of the Brain Regions Specific to Each Test
D. Detailed Supporting Data
E. References
A. Document Overview

This document identifies the brain regions that have been linked with performance on each Cambridge Brain Sciences test. The information comes from studying how performance on the tests is affected by factors such as injuries or disease, as well as studying healthy brains using functional brain imaging technologies like functional magnetic resonance imaging (fMRI) and positron emission tomography (PET).

When evaluating the core cognitive abilities measured by the Cambridge Brain Sciences tests, brain regions are recruited to perform a specific task (e.g. manipulate items in memory), rather than deal with a specific type of information (e.g. words vs. numbers), ensuring that our abstract computerized tests are suited to target particular cognitive abilities that apply across a wide range of contexts. Furthermore, all tests (and indeed, most behaviors) require more than one function, such that performance requires a network of brain regions rather than a single region. “It is simply very hard to be precise about the function of a region when that region is important in such a diversity of behavior” (Duncan & Owen, 2000). Nonetheless, research has revealed brain regions that are associated with the cognitive abilities targeted by each test.
B. Brain Networks Behind the Cambridge Brain Sciences Tests

In a landmark study published in *Neuron* (see Hampshire et al., 2012), we found that not only did performance tend to cluster into three cognitive domains—reasoning, short-term memory, and verbal ability—but each domain recruited distinct brain networks.

These networks within the frontal, parietal, and temporal cortices contribute to multiple tests. That is, any given test can recruit from multiple networks to varying degrees—it is not the case that any test or real-world activity activates one specific brain region, but rather, mixes and matches from brain regions suited to the various requirements and stages of a complex task.

The short-term memory (STM) network (red) and reasoning network (blue).

The mapping of each test to each network, based on performance data. The numbers are called factor loadings, and indicate the correlation between the observed test score and the latent “cognitive domain” score associated with a specific brain network.
### C. Summary Table of the Brain Regions Specific to Each Test

Brain regions that correspond with each task. See Section D for detailed explanations.

<table>
<thead>
<tr>
<th>Domain</th>
<th>Task</th>
<th>Outcome</th>
<th>Associated Brain Regions</th>
</tr>
</thead>
</table>
| MEMORY   | Monkey Ladder | Visuospatial Working Memory | • Prefrontal cortex / mid-dorsolateral prefrontal cortex  
• Premotor cortex  
• Posterior parietal cortex                                                                 |
| MEMORY   | Spatial Span  | Spatial Short Term Memory | • Right mid-ventrolateral area  
• Parieto-occipital regions                                                                  |
| MEMORY   | Token Search  | Working Memory            | • Frontal and temporal lobes  
• Amygdalo-hippocampal region  
• Mid-ventrolateral frontal cortex  
• Mid-dorsolateral cortex  
• Premotor cortex                                                                       |
| MEMORY   | Paired Associates | Episodic Memory  | • Left dorsolateral prefrontal cortex, ventral and anterior left prefrontal cortex  
• Ventral region of the parietal cortex  
• Occipitotemporal region  
• Cerebellum                                                                         |
<table>
<thead>
<tr>
<th>Domain</th>
<th>Task</th>
<th>Outcome</th>
<th>Associated Brain Regions</th>
<th>Example Brain Region Images</th>
</tr>
</thead>
</table>
| REASONING | Rotations | Mental Rotation | • Intraparietal sulcus  
• Medial superior precentral cortex | ![Images] |
| Polygons | Visuospatial Processing | | • Intraparietal sulcus  
• Right dorsolateral prefrontal cortex  
• Right hemisphere | ![Images] |
| Odd One Out | Deductive Reasoning | | • Anterior frontal cortex, anterior cingulate  
• Anterior insula / frontal operculum  
• Inferior frontal sulcus  
• Presupplementary motor area  
• Intraparietal sulcus | ![Images] |
| Spatial Planning | Planning | | • Frontal lobe  
• Mid-dorsolateral frontal cortex  
• Caudate nucleus and thalamus  
• Lateral premotor area  
• Anterior cingulate | ![Images] |
<table>
<thead>
<tr>
<th>Domain</th>
<th>Task</th>
<th>Outcome</th>
<th>Associated Brain Regions</th>
<th>Example Brain Region Images</th>
</tr>
</thead>
<tbody>
<tr>
<td>VERBAL ABILITY</td>
<td>Grammatical Reasoning</td>
<td>Verbal Reasoning</td>
<td>• Frontal operculum&lt;br&gt;• Posterior temporal lobe&lt;br&gt;• Superior parietal lobe&lt;br&gt;• Dorsal prefrontal cortex&lt;br&gt;• Ventral prefrontal cortex</td>
<td><img src="image1.png" alt="Image" /></td>
</tr>
<tr>
<td></td>
<td>Digit Span</td>
<td>Verbal Short Term Memory</td>
<td>• Mid-ventrolateral prefrontal cortex&lt;br&gt;• Left temporo-parietal lobe&lt;br&gt;• Basal ganglia</td>
<td><img src="image2.png" alt="Image" /></td>
</tr>
<tr>
<td>CONCENTRATION</td>
<td>Feature Match</td>
<td>Attention</td>
<td>• Mid-ventrolateral frontal cortex&lt;br&gt;• Right inferior frontal gyrus</td>
<td><img src="image3.png" alt="Image" /></td>
</tr>
<tr>
<td></td>
<td>Double Trouble</td>
<td>Response Inhibition</td>
<td>• Right prefrontal cortex&lt;br&gt;• Dorsolateral frontal cortex&lt;br&gt;• Left inferior frontal gyrus&lt;br&gt;• Dorsal striatum</td>
<td><img src="image4.png" alt="Image" /></td>
</tr>
</tbody>
</table>
D. Detailed Supporting Data

Monkey Ladder

Monk Ladder involves the cognitive function of working memory, but also requires visuospatial functions. Imaging data has revealed the brain regions involved in working memory using nonverbal stimuli with a spatial component. Across studies using tasks of this type, the prefrontal, premotor, and posterior parietal cortex are found to be involved (see Owen et al., 2005, for further details). The mid-dorsolateral prefrontal cortex, in particular, plays a role in working memory regardless of whether or not there is a spatial component (Owen et al., 1998).

Spatial Span

Spatial Span is a spatial short-term memory task that requires holding a spatial sequence in memory for a short time. An early imaging study (Owen et al., 1999) using Spatial Span found increased blood flow in the right mid-ventrolateral area during the task. In Spatial Span, information does not need to be manipulated in memory, so additional brain regions involved in similar working memory tasks (e.g., Token Search) are not involved in this simpler task. Damage to the parieto-occipital regions of the brain causes impairment in visuospatial tasks like Spatial Span, but those with frontal lobe damage tend to be more impaired if there is a planning component, rather than pure memorization.
Token Search

Token Search combines several cognitive abilities: short-term memory, working memory, and a strategy/planning component. Patients with damage to the frontal lobe are impaired even on easy puzzles, despite performing well on simpler short-term memory tests. This implies that the frontal lobe is particularly important for the strategy component of Token Search. Patients with lesions in the temporal lobe or amygdalo-hippocampal region, on the other hand, are impaired only on more difficult puzzles, implying that these regions are more responsible for the memory component of the task. Imaging studies confirm that the mid-ventrolateral frontal cortex, mid-dorsolateral cortex, and premotor cortex are all activated during this complex task (Owen et al., 1996).

Paired Associates

Paired Associates is an episodic memory task in which two items—such as object identity and location—must be paired in memory, then later recalled. The left dorsolateral prefrontal cortex is related specifically to encoding memories and using executive processes to create an organizational structure, which is required for performance in Paired Associates, and has also been observed in realistic recall of object locations (Hayes et al., 2004). More ventral and anterior left prefrontal cortex regions appear to be recruited for general episodic memory encoding (Fletcher, Shallice, & Dolan, 1998a). During retrieval of the encoded memories, the ventral prefrontal cortex and the ventral region of the parietal cortex are activated specifically during tasks with cued recall (as in Paired Associates; Fletcher et al., 1998b). Furthermore, Gould et al. (2006) confirmed that lateral and medial prefrontal and parietal cortices, as well as occipitotemporal and cerebellar regions, were associated with encoding and retrieval of object-location pairs in Paired Associates in both healthy controls and people with Alzheimer’s disease.
The Rotations test requires visuospatial processing, and specifically the ability to mentally rotate objects. A review of the literature (Zacks, 2008) found that mental rotation activates the intraparietal sulcus and adjacent regions—areas that contain spatially mapped representations, implying that a representation of objects is simulated and rotated in the brain in an analog manner.

However, the medial superior precentral cortex is also activated—this is a motor area, implying that at least in some cases, a simulation of moving the object occurs in the brain. Damage to the brain can impair mental rotation; for example, patients with Parkinson’s disease perform poorly, which extends to real-world difficulty with tasks such as learning and navigating routes (Uc et al., 2007).

In general, visuospatial tasks activate the frontoparietal network, and particularly the intraparietal sulcus region of both hemispheres. Disruptions to the right parietal cortex specifically may have consequences for visuospatial function, such as visual judgement about angles (Sack et al., 2007). Polygons requires quick visuospatial processing, object recognition, and reasoning to determine if the shapes are the same. Compared to some similar tasks, memory is less applicable to Polygons, as there is no delay between presentation of the shape and making a judgment. One study (Pollman & von Cramon, 2000) found the right dorsolateral prefrontal cortex activated when performing goal-directed visual search, but was not related to memory, suggesting this area is unique to visuospatial processing. Damage to the brain can impair visuospatial function and attention, leading to poor Polygons performance. In one study (Lee et al., 2008), nearly half of patients with acute right hemisphere strokes were impaired in an interlocking polygons task, and extreme cases can lead to neglect of one side of the visual field, leading to failure to attend to one side of the shape.
Odd One Out is a test of deductive reasoning—a core cognitive ability often included in tests of fluid intelligence. Engaging in difficult deductive reasoning results in a characteristic pattern of activity in several regions of the brain, including the anterior frontal cortex, anterior insula / frontal operculum, inferior frontal sulcus, anterior cingulate, presupplementary motor area, and intraparietal sulcus.

One study (Woolgar et al., 2010) verified that these specific areas play a role in fluid intelligence by confirming that damage to these regions—but not outside of them—predicts poor performance in reasoning tests, including items similar to Odd One Out.

Spatial Planning requires the brain’s reasoning and forward-thinking abilities. Spatial working memory is also required in order to hold plans in memory long enough to execute. The frontal lobe is known for its involvement in these higher executive functions. Early imaging studies (e.g. Owen et al., 1996) found that the mid-dorsolateral frontal cortex was activated in various versions of Spatial Planning. The caudate nucleus and the thalamus were involved only in more difficult puzzles. A subsequent study (Dagher et al., 1999) added the lateral premotor and anterior cingulate areas to the network responsible for the visual processing and anticipation of movement required for spatial planning.
Grammatical Reasoning

Grammatical Reasoning primarily involves two cognitive processes: verbal-based reasoning to determine what the sentence should be describing, and comparison of this internally-generated answer with the image on the screen. One study (Drummond et al., 2003) found that engaging in these tasks recruited brain regions associated with rehearsal and language processing within the frontal operculum and posterior temporal lobe language areas, and the superior parietal lobe visuospatial processing region was also active. In more difficult problems, additional brain regions in the dorsal and ventral prefrontal cortex are recruited. Interestingly, regions associated with working memory do not seem to be involved, as previously thought. This reliance on verbal and reasoning abilities over working memory is confirmed by imaging and behavioral data in the context of the other eleven tests (Hampshire et al., 2012).

Digit Span

Digit Span requires verbal (versus spatial) short-term memory. The frontal cortex is involved in tasks that require memorizing and recalling verbal information. More specifically, the mid-ventrolateral prefrontal cortex, primarily in the right hemisphere, is activated during verbal short-term memory tasks (Owen et al., 2000). This area of the brain is involved in tasks that require active, conscious retrieval and reproduction of stored information. Because Digit Span is a straightforward memorization task that does not require manipulation of items in memory (as in backwards digit span tasks and CBS working memory tasks, such as Token Search), it does not recruit additional brain regions involved in more complex tasks, such as the dorsolateral prefrontal cortex. Rather, the ventrolateral frontal cortex’s general role is to enable the low-level encoding strategies required in Digit Span, such as rehearsal and initiation of intentional retrieval of information from memory. Furthermore, a study of stroke survivors (Geva et al., 2021) indicated that damage to the left temporo-parietal and basal ganglia structures consistently disrupts performance on Digit Span.
Feature Match

Feature Match is a perceptual test that requires the cognitive ability of tuning attention and monitoring for differences. The mid-ventrolateral frontal cortex is involved in functions targeted by Feature Match, such as deliberate, focused control of thought and action (Owen & Hampshire, 2009). Within this network, the right inferior frontal gyrus (IFG) in particular responds when a desired feature that is being monitored for (such as a difference) is found (Hampshire et al., 2009). This region adapts to activate in response to currently-relevant information—that is, it responds selectively to items that are most relevant to the task at hand, rather than any specific stimulus.

Double Trouble

Double Trouble is a test of focused attention and response inhibition. The right prefrontal cortex, and in particular the dorsolateral region, is involved in tests that require sustained focused attention, such as Double Trouble. Damage to these regions, common in cases of traumatic brain injury, may be responsible for the problems with attention that often follow injury, and impair performance on tasks like Double Trouble (Manly et al., 2003; Dimoska-Di Marco, 2011). In addition to attention, Double Trouble involves the cognitive ability of response inhibition—the ability to suppress making responses to task-irrelevant information (e.g., the meaning of a word, rather than its colour). This can be differentiated from simple attention or effort. Brain regions involved in this function, which is particularly prominent in Double Trouble’s double-Stroop design, include the left inferior frontal gyrus (Taylor et al., 1997). The dorsal striatum has also recently been implicated as important for cognitive control, but not cognitive effort (Robertson et al., 2015).
E. References


Note: brain regions are approximate, based on available research using tasks identical to or similar to CBS Health tasks. Brain illustrations created with BioRender.com.