

1 **Supplementary Material**

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Group	Sequence	Physics	Characteristics	Utility
Spin Echo	T1-Weighted MRI	Longitudinal, spin-lattice relaxation	Low signal intensity of tissue with high water content, high signal intensity of high fat tissue (white matter) ¹	Good anatomical detail, macroscopic overview of tissue ¹
	T2-Weighted MRI	Transverse, spin-spin relaxation	High signal intensity of tissue with high water content (edema, infarction, inflammation), intermediate intensity of high fat tissue (white matter) ¹	Foundational overview of standard brain anatomy, assists in evaluation of brain pathology ¹
Inversion Recovery	Fluid-attenuated inversion recovery	Selective suppression of fluid (CSF) signal	Reduced intensity of CSF signal, high signal intensity of tissue edema/inflammation ²	Distinguishes subtle white matter abnormalities ²
Diffusion Weighted	Conventional Diffusion Weighted Imaging (DWI)	Measures the diffusion of water molecules	Increased signal intensity in areas with restricted diffusion of water ³	Allows mapping of the diffusion of water within the brain ³
	Diffusion Tensor Imaging (DTI)	Measures the anisotropic diffusion of water in the brain	Estimation of axonal (white matter) organization within the brain ⁴	Assessment of white matter tracts, deformations of white matter secondary to brain pathology
Functional MRI	Blood-oxygen-level-dependent (BOLD) imaging	Measures properties of hemoglobin to assess local changes in blood flow, with correlation to activity	Identification of regions of activity within the brain ⁵	Visualization of areas of increased cortical activity during rest or activation ⁵

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4 **Supplementary Table 1.** Overview of MRI sequence types utilized in the studies discussed
5 within this review.

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Structure	Function	System	Functional Symptoms
Superior Longitudinal Fasciculus (Right) ⁶	Connects the frontal, occipital, parietal, and temporal lobes. Visuospatial attention (goal-directed, stimulus driven) and motor control. ⁷	Visual, motor, cognitive, behavioral	Observed: Crewmembers with the largest spaceflight-associated balance disruptions had greater WM structure changes in the SLF ⁶ Potential: Visuospatial cognitive dysfunction (searching, shifting spatial attention, performing mental rotations, detecting patterns) ⁷
Inferior Longitudinal Fasciculus ⁶	Connects temporal and occipital lobes. Visual modality, object/face/place processing, emotional processing, visual memory. ⁸	Visual, cognitive, behavioral	Potential: Visual neglect, visual amnesia, visual hallucinations, prosopagnosia, visual hypo-emotionality ⁸
Inferior Occipitofrontal Fasciculus ⁶	Connects frontal and occipital/temporal lobes. Hypothesized role in spatial attention and cognition (non-dominant), goal-oriented behavior, social cognition, attention ⁹	Visual, cognitive, behavioral	Potential: Reduced verbal fluency, spatial neglect, anosognosia, potential for inability to identify facial emotions ⁹
R. posterior thalamic radiations ¹⁰	Posterior radiations connect thalamus and occipital cortex, contain optic radiations. ¹¹	Visual	Potential: Visual pathway deficits ¹¹
Calcarine, middle occipital, inferior occipital gyri ¹⁰	Location of the primary visual cortex and secondary visual association cortex ¹²	Visual, cognitive	Potential: Visual hallucinations, visual field cuts ¹²
Right fusiform gyrus ¹⁰	Facial and word recognition, within-category identification ¹³	Visual, cognitive, behavioral	Potential: Prosopagnosia, dyslexia ¹³
Inferior cerebellar peduncle ⁶	Connects the cerebellum to the spinal cord and medulla. Carries input from vestibular receptors (afferent and efferent) to help with posture, balance, coordination. ¹⁴	Cerebellar, vestibular	Potential: Vertigo and imbalance mimicking a peripheral vestibular issue, ocular tilt, spontaneous nystagmus ¹⁴
Middle cerebellar peduncle ⁶	Afferent pathway from pons to cerebellum. Controls the initiation, planning and timing of volitional motor activity ¹⁵	Cerebellar, motor	Potential: Cerebellar ataxia (limb, gait, speech) ¹⁵
Corticospinal Tract ⁶	Efferent motor information from the cerebral cortex to the body ¹⁶	Motor	Potential: Central motor weakness ¹⁶
White matter underlying primary motor and sensory cortices ⁶	Afferent and efferent motor and sensory information, sensory integration ^{17,18}	Motor, somatosensory, vestibular, cognitive	Potential: Impairment in voluntary movement, sensory ataxia, altered sensory perception of somatosensory input and vestibular equilibrium ^{17,18}
Inferior and posterior parietal lobe ⁶	Body spatial representation, central vestibular and proprioceptive processing, vertical upright perception ¹⁹	Somatosensory, vestibular, cognitive	Potential: Sensorimotor deficits, including deficits in perception and spatial memory, inaccurate reaching and grasping, inattention, deficits in trunk and limb orientation ^{19,20}
Thalamus ¹⁰	Relays information between cortex and subcortical areas. ²¹	Visual, somatosensory, cognitive, behavioral	Potential: Neuropsychological disturbances, motor weakness or sensory loss, deficits in episodic memory, executive function, processing speed, directed attention, working memory ²¹
Frontal poles ²²	Higher order cognition, emotion, goal directed behavior, task prioritization ²³	Cognitive, behavioral	Potential: Loss of empathy, difficulty re-prioritizing tasks ²³

Temporal Poles ²²	Semantic naming, facial recognition, integration of emotion with sensory information. ²⁴	Visual, cognitive, behavioral	Potential: Social withdrawal, failure to produce appropriate social signals or recognize them, self-centered, loss of empathy, unstable mood states. ²⁴
Supplementary Motor Region ^{25,26}	Self-initiated movement, stimulus-cued movement, speech production, temporal triggering. ^{27,28}	Motor, visual, cognitive, behavioral	Potential: Akinesia, speech deficits, executive working memory deficits ^{27,28}
Premotor Region ²⁵	Integration of visuospatial information for movement planning, both external and internal memory-based ²⁹	Visual, motor, cognitive, behavioral	Potential: Impaired performance of visually and verbally cued tasks, difficulty initiating memory cued tasks, motor apraxia ³⁰
Primary Sensorimotor Region ²⁵	Integration of sensory and motor skills needed for skilled movement ¹⁸	Motor, somatosensory	Potential: Impaired cross motor function and reflexes, impaired fine motor skills, disrupted learning of new motor tasks ¹⁸
Left Caudate Nucleus ²⁵	Voluntary movement, postural control, cognitive planning of adaptive goal-directed behavior. ³¹	Motor, somatosensory, cognitive	Observed: Decrement in postural control on Recovery from Fall/Stand Test, Dynamic Postural Stability Test. ²⁵ Potential: Gait disturbances, slowed walking speed. Impaired visual discrimination, alternation behavior, cognitive-driven strategy switching. ³¹⁻³³
Lower Extremity Primary Motor Area/Mid-Cingulate ²⁵	Motor execution. Processing of internal/external states (emotional states) and translation into motor commands. ³⁴	Motor, cognitive, behavioral	Potential: Impaired movement ³⁴

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Supplementary Table 2. Correlation between the structure/connectivity alterations seen with neuroimaging after spaceflight, their relationship to major CNS pathways, and the predicted sensorimotor/cognitive/behavioral symptoms associated with deficits within the pathway.

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