

NEUROLOGIST HANDOUT

Issues Specific to Myalgic Encephalomyelitis (per the ICC*)

Neurocognitive, sleep, autonomic and sensory disturbances, pain, headaches, and paresthesias are prominent neurological signs and symptoms. Cognitive impairments including slow processing of information, poor attention, word finding, and working memory are some of the most functionally disabling symptoms. 1, 2

Structural and functional abnormalities within the brain and spinal cord are consistent with pathological dysfunction of the regulatory centers and communication networks of the brain, CNS and ANS, and are essential for effective ongoing self-organization. 3

Reduced brainstem gray matter volume is consistent with insult to the midbrain at fatigue onset. Feedback control loops may suppress cerebral motor and cognitive activity, disrupt CNS homeostasis, and reset elements of the ANS. Brain stem injury and loss of homeostasis. Decreased gray matter volume in midbrain & pulse pressure suggest impaired cerebrovascular auto-regulation. Decreased white midbrain matter volume decreased with fatigue duration. 4, 29

Greater source activity and more parts of the brain are utilized in cognitive processing, which supports patients' perception of greater effort. 5, 6, 28

Reduced duration of uninterrupted sleep may explain reported unrefreshed sleep, pain and overwhelming fatigue. Prolonged sleep onset latency. Increased alpha intrusion into delta sleep 7

Decreased absolute cortical blood flow and further reduction in cerebral blood flow after exercise. Greater involvement of the brain correlates with greater severity 8, 9

Decreased hypoperfusion in brainstem distinguishes ME from depression. 10

Decreased cognitive functioning: prolonged reaction time 11

Increased plaque or hyperintensities in the white matter & tracts is consistent with demyelination or inflammation & increase risk of cerebrovascular events 12, 13

Reduced regional gray and white matter volumes are consistent with impaired memory and visual processing. 14, 15, 16

Decreased metabolism of glucose in the brain 17

Metabolism in brain stem differentiates ME from depression 8, 10

Greater effort is required - elevated source current & more regions of the brain are utilized in cognitive activity & fatiguing tasks: poor processing of auditory & spatial information, poor working memory. 1, 5, 6

Slower performance in visual imagery & motor tasks - ventral anterior cingulate cortex was active when controls made an error but not in patients. 15

Reduced blood flow in temporal lobes may contribute to memory and cognitive impairment & fatigue 18, 19

Elevated sensory signaling perceived by the brain as pain and fatigue. Musculoskeletal – (surface EEG scalp) CNS signals are altered when controlling voluntary muscle activities, especially when they are fatiguing. Poor and slower motor performance. 20, 21

Abnormal spatial and temporal symmetry of gait. 22

Cerebral spinal fluid - (spinal tap) increased opening pressure on lumbar puncture. Proteomes distinguish ME from post-treatment Lyme disease and controls. Spinal fluid increased lymphocytes and protein. 23, 24

IL-10 increased with granulocyte-macrophage (GM), colony-stimulating factor (CSF) suppression 24

Elevated lactate is consistent with reduced cortical blood flow, mitochondrial dysfunction & oxidative stress. Lateral ventricular: 297% vs. anxiety disorder & 348% vs. controls. 25

In CFS T1wSE was elevated in sensorimotor WM and decreased in the brainstem. 26

Neuroinflammation in Patients with CFS/ME: An ¹¹C-(R)-PK11195 PET Study. 27

Neuroinflammation in the dorsal root ganglia, (modulators of peripheral sensory information traveling to the brain) - Chaudhuri A. Abstract presentation at the Royal Society of Medicine Meeting 2009

Note: Many science documents refer to this patient population having CFS. While CFS is a broad vague term, studies listed here are chosen as they apply most closely to those who have ME as per the ICC.

More science information can be found at on the resources page at www.MEadvocacy.org

As with all support group files, this is prepared for informational purposes and is not to be considered medical advice. (8/20/2018)

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Resource citations

*Myalgic Encephalomyelitis International Consensus Primer for Medical Practitioners (Page 5)

http://sacfs.asn.au/download/me_international_consensus_primer_for_medical_practitioners.pdf

1. Objective evidence of cognitive complaints in CFS: a BOLD fMRI study of verbal working memory. *Neuroimage* 2005. www.ncbi.nlm.nih.gov/pubmed/15907308
2. Attention and verbal learning in patients with CFS. *J Int Neuropsychol Soc* 1998. www.ncbi.nlm.nih.gov/pubmed/9745235
3. CFS and the central nervous system. *J Int Med Res* 2008. www.ncbi.nlm.nih.gov/pubmed/18831878
4. A brain MRI study of CFS: evidence of brainstem dysfunction and altered homeostasis. *NMR Biomed* 2011. www.ncbi.nlm.nih.gov/pubmed/21560176
5. Functional neuroimaging correlates of mental fatigue induced by cognition among CFS patients and controls. *Neuroimage* 2007. www.ncbi.nlm.nih.gov/pubmed/17408973
6. EEG source analysis of CFS. 2010 www.sciencedirect.com/science/article/pii/S0925492709002406
7. Defining the occurrence and influence of alpha-delta sleep in CFS. *Am J Med Sci* 2007. www.ncbi.nlm.nih.gov/pubmed/17301585
8. Patients with CFS have reduced absolute cortical blood flow. *Clin Physiol Funct Imaging* 2006. www.ncbi.nlm.nih.gov/pubmed/16494597
9. Cerebral blood flow is reduced in CFS as assessed by arterial spin labeling. *J Neurol Sci.* 2011. www.ncbi.nlm.nih.gov/pubmed/21167506
10. Brainstem perfusion is impaired in CFS. *QJM* 1995. www.ncbi.nlm.nih.gov/pubmed/8542261
11. Influence of exhaustive treadmill exercise on cognitive functioning in CFS. *Am J Med* 1998. www.ncbi.nlm.nih.gov/pubmed/9790484
12. Neuroimaging in CFS. 1998; [www.amjmed.com/article/S0002-9343\(98\)00175-2/pdf](http://www.amjmed.com/article/S0002-9343(98)00175-2/pdf)
13. A chronic illness characterized by fatigue, neurologic and immunologic disorders, and active human herpes virus type 6 infection. *Ann Intern Med* www.ncbi.nlm.nih.gov/pubmed/1309285
14. Regional grey and white matter volumetric changes in ME (CFS): a voxel-based morphometry 3 T MRI study. *Br J Radiol* 2012. www.ncbi.nlm.nih.gov/pmc/articles/PMC3474083/
15. Gray matter volume reduction in CFS. *NeuroImage* 2005. www.sciencedirect.com/science/article/pii/S1053811905001394
16. Mechanisms underlying fatigue: a voxel-based morphometric study of CFS. *BMC Neurol* 2004. www.ncbi.nlm.nih.gov/pubmed/15461817
17. Brain positron emission tomography (PET) in CFS. *Amer J Med* 1998. www.ncbi.nlm.nih.gov/pubmed/9790483
18. Study of Cerebral Perfusion by NeuroSPECT in Patients with Chronic Fatigue Syndrome. In: Hyde BM, Goldstein J, Levine P, eds. *The Clinical and Scientific Basis of Myalgic Encephalomyelitis, Chronic Fatigue Syndrome*. 1992: 432-8.
19. The assessment of vascular abnormalities in late life CFS by brain SPECT: Comparison with late life major depressive disorder. *J CFS* 1995. www.tandfonline.com/doi/abs/10.1300/J092v01n01_05
20. Evidence for and pathophysiologic implications of hypothalamic-pituitary-adrenal axis dysregulation in fibromyalgia and CFS. *Ann NY Acad Sci* 1998. www.ncbi.nlm.nih.gov/pubmed/9629295
21. Altered central nervous system signal during motor performance in CFS. *Clin Neurophysiol.* 2004. www.ncbi.nlm.nih.gov/pubmed/15351380
22. Alterations of spatial-temporal parameters of gait in CFS patients. 1998. [www.jns-journal.com/article/S0022-510X\(97\)00200-1/fulltext](http://www.jns-journal.com/article/S0022-510X(97)00200-1/fulltext)
23. Distinct cerebrospinal fluid proteomes differentiate post-treatment lyme disease from CFS. *PLoS ONE* 2011. <http://journals.plos.org/plosone/article?id=10.1371/journal.pone.0017287>
24. Spinal fluid abnormalities in patients with CFS. *Clin Diagn Lab Immunol* 2005. www.ncbi.nlm.nih.gov/pmc/articles/PMC540195/
25. Ventricular cerebrospinal fluid lactate is increased in CFS compared with generalized anxiety disorder: an in vivo 3.0 T (1)H MRS imaging study. *NMR Biomed* 2009. www.ncbi.nlm.nih.gov/pubmed/18942064
26. Hyperintense sensorimotor T1 spin echo MRI is associated with brainstem abnormality in CFS. www.sciencedirect.com/science/article/pii/S2213158218302237
27. Neuroinflammation in patients with CFS/ME: An ¹¹C-(R)-PK11195 PET Study. 2014. www.ncbi.nlm.nih.gov/pubmed/24665088
28. EEG characteristics in patients with CFS. (Abnormal changes in cerebral functions localized at the right frontal and left occipital) 2016. www.ncbi.nlm.nih.gov/pmc/articles/PMC4734796/
29. Grey and white matter differences in CFS – A voxel-based morphometry study. 2018. www.ncbi.nlm.nih.gov/pmc/articles/PMC5633338/