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ACCEPTED MANUSCRIPT

Fatigue in Multiple Sclerosis: is it related to cytokines and hypothalamic-pituitary-

adrenal axis?

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Abstract:

Background:

Fatigue is a common symptom of Multiple Sclerosis (MS) that diminishes the quality of life of patients, but its exact mechanism remains poorly understood. There is not a generally adopted scale to determine MS fatigue. Studies that investigated physiopathology of fatigue symptom have shown dysregulation of hypothalamic-pituitaryadrenal (HPA) axis. In the current study, we aimed to compare the results obtained with two separate scales, namely the Fatigue Severity Scale (FSS) and the Neurological Fatigue Index-Multiple Sclerosis (NFI-MS), and assess the relationship between fatigue and serum IL-1 β , TNF- α , IL-35, IL-2, IL-10, ACTH, cortisol, α -MSH, β -MSH, γ -MSH and CLIP (Corticotropinlike intermediate lobe peptide) in MS patients categorized as fatigued and non-fatigued on the basis of FSS scores. Methods:

For the study, a total of 54 (29 females, 25 males) patients diagnosed with RRMS including 26 with fatigue symptom (48.1%), and 26 healthy controls (13 females, 13 males) were enrolled. A FSS score ≥36 was

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