

Role of adaptive and innate immunity in Restless Legs Syndrome

<u>**G. Benbir Senel**</u>¹, A. Kokoglu², A. Engin², M.Y. Gelmez²,

E. Cetin², D. Karadeniz¹, G. Deniz²

¹Istanbul University- Cerrahpasa, Cerrahpasa Faculty of Medicine, Neurology Department, Istanbul, Turkey

²Istanbul University, Aziz Sancar Institute of Experimental Medicine, Immunology Department, Istanbul, Turkey

RLS RLS Controls

+OSAS

RLS RLS Controls

+OSAS

RLS RLS

+OSAS

Controls

Introduction

Restless legs syndrome (RLS) is one of the most common sleep-related movement disorders. The underlying pathogenetic mechanisms are mainly dopaminergic dysregulation in the central nervous system and brain iron deficiency. Recent studies, however, suggest some other theories on RLS pathophysiology, such as peripheral neuropathy, small intestinal bacterial overgrowth, and immunologic alterations. In this study, we aimed to assess the potential role of innate and adaptive cells of the immune

system on RLS.

Methods

In this study, we prospectively enrolled patients with RLS admitting to our Sleep and Disorders Unit. All patients had detailed clinical evaluation and all had one-night polysomnography (PSG) in sleep laboratory to exclude any associated sleep disorders, such sleep apnea. The presence of medical diseases and/or use of any medications and/or substances were also set as the exclusion criteria. A sex- and age-matched healthy controls were also prospectively enrolled; detailed clinical evaluation and onenight PSG were also performed in all healthy subjects. Blood samples obtained from all participants were investigated for the following parameters: T, B, NK, NKT and ILC cell ratio; intracellular IFN-γ, IL-6, IL-10, IL-13 cytokines in T, B, NK cells; CD8+ T and NK cell cytotoxic activity (which were analyzed by flow cytometry); IFN-γ, TNF-α, IL-2, IL-4, IL-6, IL-10 and IL-13 levels (which were analyzed with ELISA).

Results

We observed that the ratio of ILC-1 subset (Figure 2) and IL-13+CD4+ T cells (Figure 3) were





Conclusion

Our results showed that innate rather than adaptive immunity is altered in RLS.

References

1. Abbas, A.K., A.H. Lichtman, and S. Pillai, Cellular and molecular immunology E-book. 2014: Elsevier Health Sciences.

- 2. O'Brien, K.L. and D.K. Finlay, Immunometabolism and natural killer cell responses. Nat Rev Immunol, 2019. 19(5): p. 282-290.
- 3. Zhu, J., H. Yamane, and W.E. Paul, Differentiation of effector CD4 T cell populations (*). Annu Rev Immunol, 2010. 28: p. 445-89.
- 4. Manconi, M., et al., *Restless legs syndrome*. Nat Rev Dis Primers, 2021. 7(1): p. 80.
- 5. Earley, C.J., et al., Connectome and molecular pharmacological differences in the dopaminergic system in restless legs syndrome (RLS): plastic changes and neuroadaptations that may contribute to augmentation. Sleep Med, 2017. 31: p. 71-77.
- 6. Besedovsky, L., T. Lange, and J. Born, Sleep and immune function. Pflugers Arch, 2012. 463(1): p. 121-37.
- 7. Weinstock, L.B., A.S. Walters, and P. Paueksakon, Restless legs syndrome--theoretical roles of inflammatory and immune mechanisms. Sleep Med Rev, 2012. 16(4): p. 341-54.

Acknowledgements

The present work was supported by the Research Fund of Istanbul University (Project no: 37726) and Directorate of Presidential Strategy and Budget (Project no: 36288).

https://esleepeurope.eu/



https://esleepeurope.eu/