

Long COVID: Further Advances in Our Understanding of the Role of Specific Chemokines (CCR5, CCR6, CCR9, and CCL3) in Pediatrics

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ABSTRACT

Introduction: ARS-CoV-2 infection (COVID-19) is a major pandemic that has

caused significant mortality and morbidity worldwide. About 80% of patients

had mild or moderate disease, and 5% of patients with severe disease

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developed severe disease. Long COVID-19, also known as post-COVID-19 syndrome, is a condition in which individuals continue to experience symptoms for weeks or months after recovering from COVID-19. There is a lack of sufficient data on the risk factors for post-COVID-19 sequelae in children. This study aimed to evaluate the long-term effects of severe COVID-19 on promotor methylation and expression genes, including *CCR5, CCR6, CCR9,* and *CCL3* in children.
Methods and Materials: Clinical data and blood samples from 94 long COVID patients and 25 healthy subjects were collected. Promotor methylation and mRNA expression of *CCR5, CCR6, CCR9,* and *CCL3* genes in these patients and control group were assessed through methylation-specific PCR and real-time

Results: Our result indicated that promotors of CCR5 (p = 0.01) and CCL3 (p = 0.006) in convalescence COVID-19 children were hyper-methylated compared to the healthy control group. Subsequently, *CCR5* and *CCL3* transcripts decreased compared to the control group (p = 0.01). In addition, *CCL3* transcript in children with convalescent COVID-19 decreased compared to the control group (p = 0.008). However, we did not observe any significant modification in the transcript levels of *CCR6* (p = 0.7) and *CCR9* (p = 0.46) in children with convalescent COVID-19 compared to all the control groups.

Conclusion and Discussion: The *CCR5* and *CCL3* promoter region DNA methylation and the subsequent decrease in the expression of these genes were possibly correlated with long-term COVID-19 occurrence in children. Our study revealed additional data on the SARS-CoV-2-mediated inflammatory response.

Keywords: CCR5 receptor antagonists, Chemokine CCL3, Methylation

PCR assay.

