LETTER TO THE EDITOR

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Understanding the Necessity to Recognize Multisystem Inflammatory Syndrome in Children (MIS-C) in Emerging Infectious Diseases based on the Lessons Learned from COVID-19: A Letter to the Editor



Mohammadreza Naghibi¹ and Rasoul Raesi^{2,3,*}

¹Pediatric Interventional Cardiology, Pediatric and Congenital Cardiology Division, Pediatric Department, Faculty of Medicine, Mashhad University of Medical Sciences, Mashhad-Iran

²Department of Health Services Management, School of Health, Mashhad University of Medical Sciences, Mashhad, Iran

³Department of Nursing, Torbat Jam Faculty of Medical Sciences, Torbat Jam, Iran

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*Address correspondence to this author at the Department of Health Services Management, School of Health, Mashhad University of Medical Sciences, Mashhad, Iran and Department of Nursing, Torbat Jam Faculty of Medical Sciences, Torbat Jam, Iran; E-mail: raesi.br881@gmail.com

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To The Editor,

In the wake of the COVID-19 pandemic, the medical community has been faced with a new challenge in the form of Multisystem Inflammatory Syndrome in Children (MIS-C), a rare but serious condition with potentially lifethreatening complications [1, 2]. We are writing to express our concerns about a new condition that has emerged during the COVID-19 pandemic known as MIS-C. As healthcare researchers, we believe it is crucial to raise awareness about MIS-C and its potential implications on pediatric health during this unprecedented global health crisis. As we navigate the complexities of emerging infectious diseases, it becomes increasingly crucial to recognize and understand the unique manifestations of MIS-C in pediatric populations. Drawing upon the lessons learned from the global response to COVID-19, this article delves into understanding the necessity of recognizing MIS-C in emerging infectious diseases.

Emerging infectious diseases can have a great impact, especially when it comes to the little ones [3]. Children have their unique ways of reacting to new infections [4]. This is where MIS-C comes into the spotlight, causing a stir in the realm of pediatric healthcare [5]. MIS-C is like a mystery package the immune system receives after a bout

of an infectious disease [6]. This inflammatory syndrome can appear with infections, leaving pediatricians confused by its varied presentations [7, 8]. MIS-C is a rare but serious condition that affects children and adolescents who have been infected with the novel coronavirus [9, 10]. The principal symptoms and signs of MIS-C are similar to those of Kawasaki disease (KD), suggesting a shared pathogenesis. Clinicians should suspect MIS-C if patients present with fever, inflammation of the heart, KD-like features (skin rash, conjunctivitis, oral mucosa changes, hand or foot edema), gastrointestinal symptoms (abdominal pain, vomiting, diarrhea), evidence of SARS-CoV-2 infection (by PCR or serology) or recent contact with COVID-19 patients, respiratory distress, joint pain, dysuria, skin erosions (including chilblains-like phenomenon affecting fingers and toes), swelling or erythema of the hands and feet, and altered consciousness (AVPU score below A) [11-14].

The exact cause of MIS-C is still not fully understood, but it is believed to be a delayed immune response to the virus [15]. This condition can be life-threatening and requires prompt recognition and treatment [16]. As healthcare providers and public health officials continue to learn more about MIS-C, it is important to implement

strategies for early detection and intervention to prevent severe outcomes in affected children [17]. One of the challenges in managing MIS-C is the lack of specific diagnostic criteria [18, 19]. The symptoms of MIS-C can overlap with other pediatric inflammatory conditions, making it difficult to differentiate and diagnose [20]. It is imperative for healthcare providers to be vigilant and consider MIS-C in children presenting with prolonged fever and systemic inflammation, especially if they have had a recent history of COVID-19 infection or exposure [21]. Furthermore, the long-term consequences of MIS-C are still unknown. Some children may experience ongoing cardiac issues or other complications even after the acute phase of the illness has resolved [22, 23]. This highlights the importance of long-term follow-up and monitoring of children who have experienced MIS-C to ensure optimal recovery and prevent future health complications [24]. In addition, the disproportionate impact of COVID-19 on children from marginalized communities raises concerns about the potential disparities in access to care for children with MIS-C [25]. Children from low-income families or minority populations may face barriers to healthcare services, leading to delays in diagnosis and treatment [26, 27]. It is essential for public health efforts to address these disparities and ensure equitable access to care for all children affected by MIS-C [28, 29].

As we continue to navigate the complexities of the COVID-19 pandemic, healthcare providers, researchers, and policymakers must collaborate and share information about MIS-C to improve our understanding of this condition and enhance our ability to effectively manage and prevent it. By prioritizing pediatric health and investing in research on MIS-C, we can better protect the well-being of our children and mitigate the long-term impact of this pandemic on future generations. In conclusion, the emergence of MIS-C as a complication of COVID-19 has highlighted the critical need for heightened awareness and recognition of this condition among healthcare providers, parents, and public health officials. As a rare but serious manifestation of SARS-CoV-2 infection, MIS-C can lead to significant morbidity and, in some cases, mortality if not promptly diagnosed and treated. The lessons learned from the COVID-19 pandemic emphasize that timely recognition of MIS-C is essential for improving outcomes in affected children, underscoring the importance of ongoing education and training for healthcare professionals regarding the clinical features and diagnostic criteria associated with this syndrome. Furthermore, the COVID-19 pandemic has demonstrated the necessity of robust surveillance systems and interdisciplinary collaboration to monitor and respond to emerging infectious diseases that can affect pediatric populations. The rapid global response to COVID-19 has provided valuable insights into the importance of data sharing, research, and public health initiatives aimed at understanding and mitigating the impact of such diseases. By applying these lessons to the recognition and management of MIS-C, healthcare systems can enhance their preparedness for future outbreaks, ensuring that children receive timely and appropriate care.

CONCLUSION

In conclusion, recognizing MIS-C as a significant consequence of emerging infectious diseases like COVID-19 is vital for protecting children's health. The pandemic has taught us the importance of early diagnosis, preventive management, and the need for comprehensive public health strategies that prioritize pediatric care. By fostering awareness, improving clinical practices, and strengthening health systems, we can better safeguard the well-being of children and respond effectively to future health crises.

AUTHORS' CONTRIBUTION

M.N.: contributed to the concept or design of the study. R.R.: wrote the manuscript.

LIST OF ABBREVIATIONS

KD = Kawasaki disease

MIS-C = Multisystem Inflammatory Syndrome in Children

CONSENT FOR PUBLICATION

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CONFLICT OF INTEREST

The authors declare no conflict of interest, financial or otherwise.

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REFERENCES

- [1] Hennon TR, Penque MD, Abdul-Aziz R, Alibrahim OS, McGreevy MB, Prout AJ. COVID-19 associated multisystem inflammatory syndrome in children (MIS-C) guidelines; a Western New York approach. Prog Pediatr Cardiol 2020; 101232. Online ahead of print.
 - http://dx.doi.org/10.1016/j.ppedcard.2020.101232
- [2] Zhao Y, Yin L, Patel J, Tang L, Huang Y. The inflammatory markers of multisystem inflammatory syndrome in children (MIS-C) and adolescents associated with COVID-19: A meta-analysis. J Med Virol 2021; 93(7): 4358-69. http://dx.doi.org/10.1002/jmv.26951 PMID: 33739452
- [3] Feldstein LR, Tenforde MW, Friedman KG, et al. Characteristics and outcomes of US children and adolescents with multisystem inflammatory syndrome in children (MIS-C) compared with severe acute COVID-19. JAMA 2021; 325(11): 1074-87. http://dx.doi.org/10.1001/jama.2021.2091 PMID: 33625505
- [4] Molloy EJ, Nakra N, Gale C, Dimitriades VR, Lakshminrusimha S. Multisystem inflammatory syndrome in children (MIS-C) and neonates (MIS-N) associated with COVID-19: Optimizing definition and management. Pediatr Res 2023; 93(6): 1499-508. http://dx.doi.org/10.1038/s41390-022-02263-w PMID: 36050390
- [5] Son MBF, Friedman K. COVID-19: Multisystem inflammatory syndrome in children (MIS-C) clinical features, evaluation, and diagnosis. 2021. Available from: https://www.uptodate.com/contents/covid-19-multisystem-inflamm atory-syndrome-in-children-mis-c-clinical-features-evaluation-and-

- diagnosis(accessed on 8-10-2024)
- [6] Hoste L, Van Paemel R, Haerynck F. Multisystem inflammatory syndrome in children related to COVID-19: A systematic review. Eur J Pediatr 2021; 180(7): 2019-34. http://dx.doi.org/10.1007/s00431-021-03993-5 PMID: 33599835
- [7] Levy M, Recher M, Hubert H, et al. Multisystem inflammatory syndrome in children by COVID-19 vaccination status of adolescents in France. JAMA 2022; 327(3): 281-3. http://dx.doi.org/10.1001/jama.2021.23262 PMID: 34928295
- [8] Blumfield E, Levin TL, Kurian J, Lee EY, Liszewski MC. Imaging findings in multisystem inflammatory syndrome in children (MIS-C) associated with coronavirus disease (COVID-19). AJR Am J Roentgenol 2021; 216(2): 507-17. http://dx.doi.org/10.2214/AJR.20.24032 PMID: 32755212
- [9] Simpson JM, Newburger JW. Multisystem inflammatory syndrome in children in association with COVID-19. Am Heart Assoc 2020; 49: 437-40.
- [10] Son MBF, Friedman K, Kaplan S, Sundel R, Randolph A. COVID-19: multisystem inflammatory syndrome in children (MIS-C) management and outcome. 2021. Available from: https://www.uptodate.com/contents/covid-19-multisystem-inflammatory-syndrome-in-children-mis-c-management-and-outcome(accessed on 8-10-2024)
- [11] Patel JM. Multisystem inflammatory syndrome in children (MIS-C). Curr Allergy Asthma Rep 2022; 22(5): 53-60. http://dx.doi.org/10.1007/s11882-022-01031-4 PMID: 35314921
- [12] Tolunay O, Çelik Ü, Arslan İ, et al. Multisystem inflammatory syndrome in children (MIS-C) associated with COVID-19: A case series experience in a tertiary care hospital of Southern Turkey. J Trop Pediatr 2021; 67(2): fmab050. http://dx.doi.org/10.1093/tropej/fmab050 PMID: 34028528
- [13] Jain S, Sen S, Lakshmivenkateshiah S, et al. Multisystem inflammatory syndrome in children with COVID-19 in Mumbai, India. Indian Pediatr 2020; 57(11): 1015-9. http://dx.doi.org/10.1007/s13312-020-2026-0 PMID: 32788432
- [14] Yasuhara J, Watanabe K, Takagi H, Sumitomo N, Kuno T. COVID-19 and multisystem inflammatory syndrome in children: A systematic review and meta-analysis. Pediatr Pulmonol 2021; 56(5): 837-48. http://dx.doi.org/10.1002/ppul.25245 PMID: 33428826
- [15] Greene AG, Saleh M, Roseman E, Sinert R. Toxic shock-like syndrome and COVID-19: Multisystem inflammatory syndrome in children (MIS-C). Am J Emergency Med 2020; 38(11): 2492.
- [16] Radia T, Williams N, Agrawal P, et al. Multi-system inflammatory syndrome in children & adolescents (MIS-C): A systematic review of clinical features and presentation. Paediatr Respir Rev 2021; 38: 51-7. PMID: 32891582
- [17] Aldawas A, Ishfaq M. COVID-19: multisystem inflammatory syndrome in children (MIS-C). Cureus 2022; 14(1): e21064. http://dx.doi.org/10.7759/cureus.21064 PMID: 35165539
- [18] Diorio C, Henrickson SE, Vella LA, et al. Multisystem inflammatory syndrome in children and COVID-19 are distinct

- presentations of SARS-CoV-2. J Clin Invest 2020; 130(11): 5967-75.
- http://dx.doi.org/10.1172/JCI140970 PMID: 32730233
- [19] Zhao Y, Patel J, Huang Y, Yin L, Tang L. Cardiac markers of multisystem inflammatory syndrome in children (MIS-C) in COVID-19 patients: A meta-analysis. Am J Emerg Med 2021; 49: 62-70.
 - http://dx.doi.org/10.1016/j.ajem.2021.05.044 PMID: 34082189
- [20] Palabiyik F, Akcay N, Sevketoglu E, Hatipoglu N, Sari EE, Inci E. Imaging of multisystem inflammatory disease in children (MIS-C) associated with COVID-19. Acad Radiol 2021; 28(9): 1200-8. http://dx.doi.org/10.1016/j.acra.2021.05.030 PMID: 34284918
- [21] Noval Rivas M, Porritt RA, Cheng MH, Bahar I, Arditi M. COVID-19-associated multisystem inflammatory syndrome in children (MIS-C): A novel disease that mimics toxic shock syndrome—the superantigen hypothesis. J Allergy Clin Immunol 2021; 147(1): 57-9. http://dx.doi.org/10.1016/j.jaci.2020.10.008 PMID: 33075409
- [22] Rostami-Maskopaee F, Ladomenou F, Razavi-Amoli SK, et al. Clinical characteristics and outcomes of the multisystem inflammatory syndrome in children (MIS-C) following COVID-19 infection in Iran: A multicenter study. PLoS One 2022; 17(9): e0274104.
 - http://dx.doi.org/10.1371/journal.pone.0274104 PMID: 36137147
- [23] Wu EY, Campbell MJ. Cardiac manifestations of multisystem inflammatory syndrome in children (MIS-C) following COVID-19. Curr Cardiol Rep 2021; 23(11): 168. http://dx.doi.org/10.1007/s11886-021-01602-3 PMID: 34599465
- [24] Consiglio CR, Cotugno N, Sardh F, Pou C, Amodio D, Rodriguez L. The immunology of multisystem inflammatory syndrome in children with COVID-19. Cell 2020; 183(4): 968-81.
- [25] Gottlieb M, Bridwell R, Ravera J, Long B. Multisystem inflammatory syndrome in children with COVID-19. Am J Emerg Med 2021; 49: 148-52. http://dx.doi.org/10.1016/j.ajem.2021.05.076 PMID: 34116467
- [26] Wangu Z, Swartz H, Doherty M. Multisystem inflammatory syndrome in children (MIS-C) possibly secondary to COVID-19 mRNA vaccination. BMJ Case Rep 2022; 15(3): e247176. http://dx.doi.org/10.1136/bcr-2021-247176 PMID: 35354564
- [27] Ebina-Shibuya R, Namkoong H, Shibuya Y, Horita N. Multisystem Inflammatory Syndrome in Children (MIS-C) with COVID-19: Insights from simultaneous familial Kawasaki Disease cases. Int J Infect Dis 2020; 97: 371-3. http://dx.doi.org/10.1016/j.ijid.2020.06.014 PMID: 32553716
- [28] Campanello C, Mercuri C, Derchi M, et al. Cardiovascular manifestations in multisystem inflammatory syndrome in children (MIS-C) associated with COVID-19 according to age. Children (Basel) 2022; 9(5): 583.
- http://dx.doi.org/10.3390/children9050583 PMID: 35626760
 [29] Lipton M, Mahajan R, Kavanagh C, et al. AKI in COVID-19-associated multisystem inflammatory syndrome in children (MIS-C). Kidney360 2021; 2(4): 611-8. http://dx.doi.org/10.34067/KID.0005372020 PMID: 35373052