Associationof Intravenous Immunoglobulins Plus Methylprednisolone vs Immunoglobulins Alone With Course of fever in Multisystem Inflammatory Syndrome in Children.

Naïm Ouldali, Julie Toubiana, Denise Antona, Etienne Javouhey et al. JAMA. doi:10.1001/jama.2021.0694.

Published online

February 1, 2021 IMPORTANCE: Multisystem inflammatory syndrome in children (MIS-C) is the most severe pediatric disease associated with severe acute respiratory syndrome coronavirus 2 infection, potentiallylife-threatening, but the optimal therapeutic strategy remains unknown. **OBJECTIVE:** To compare intravenous immunoglobulins (IVIG) plus methylprednisolone vs IVIG alone as initial therapy in MIS-C.

DESIGN, SETTING, AND PARTICIPANTS: Retrospective cohort study drawn from a national surveillance system with propensity score-matched analysis. All cases with suspected MIS-C were reported to the French National Public Health Agency. Confirmed MIS-C cases fulfilling the World Health Organization definition were included. The study started on April 1, 2020, and follow-up ended on January 6, 2021.

MAIN OUTCOMES AND MEASURES: The primary outcome was persistence of fever 2 days after the introduction of initial therapy or recrudescence of fever within 7 days, which defined treatment failure. Secondary outcomes included a second-line therapy, hemodynamic support, acute left ventricular dysfunction after first-line therapy, and length of stay in the pediatric intensive care unit. The primary analysis involved propensity score matching with a minimum caliper of 0.1.

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IVIG ALONE VS IVIG + STEROIDS FOR INITIAL THERAPY IN MIS-C

RESULTS: Among 181 children with suspected MIS-C, 111 fulfilled the World Health Organization definition (58 females [52%]; median age, 8.6 years [interquartile range, 4.7 to 12.1]). Five children did not receive either treatment. Overall, 3 of 34 children (9%) in the IVIG and methylprednisolone group and 37 of 72 (51%) in the IVIG alone group did not respond to treatment. Treatment with IVIG and methylprednisolone vs IVIG alone was associated with lower risk of treatment failure (absolute risk difference, -0.28 [95% CI, -0.48 to -0.08]; odds ratio [OR], 0.25 [95% CI, 0.09 to 0.70]; P = .008). IVIG and methylprednisolone therapy vs IVIG alone was also significantly associated with lower risk of use of second-line therapy (absolute risk difference, -0.22 [95% Cl, -0.40 to -0.04]; OR, 0.19 [95% Cl, 0.06 to 0.61]; P = .004), hemodynamic support (absolute risk difference, -0.17 [95% Cl, -0.34 to -0.004]; OR, 0.21 [95% CI, 0.06 to 0.76]), acute left ventricular dysfunction occurring after initial therapy (absolute risk

difference, -0.18 [95% CI, -0.35 to -0.01]; OR, 0.20 [95% CI, 0.06 to 0.66]), and duration of stay in the pediatric intensive care unit (median, 4 vs 6 days; difference in days, -2.4 [95% Cl, -4.0 to -0.7]).

CONCLUSIONS AND RELEVANCE : Among children with MIS-C, treatment with IVIG and methylprednisolone vs IVIG alone was associated with a more favourable fever course. Study interpretation is limited by the observational design.

EXPERT COMMENT



Severe MIS-C is a life threatening condition and can appear in postcovid period (2-3 months later acute Covid-19 infection). It should be differentiated from Kawasaki disease in children. Severe MIS-C can be associated with severe ventricular dysfunction with hemodynamic instability which is a predominant feature compare to coronary artery involvement in Kawasaki disease. Early Identification and aggressive management with IVIG and steroid as compare to IVIG shows more favorable course.

DR KAUSTABH CHAUDHURI

MD (PGI CHANDIGARH), PEDIATRIC CRITICAL CARE FELLOWSHIP (NUH, SINGAPORE) CONSULTANT, PEDIATRIC INTENSIVE CARE APOLLO HOSPITAL, KOLKATA, INDIA.

DR MANINDER S DHALIWAL

pedpearls@gmail.com

DR PIYUSH GUPTA IAP NATIONAL PRESIDENT

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With warm regards,

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Reference : Ouldali N, Toubiana J, Antona D, Javouhey E et al. French Covid-19 Paediatric Inflammation Consortium. Association of Intravenous Immunoglobulins Plus Methylprednisolone vs Immunoglobulins Alone With Course of Fever in Multisystem Inflammatory Syndrome <u>in Children.</u> JAMA. 2021 Feb 1:e210694. doi: 10.1001/jama.2021.0694 Epub ahead of print.