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Review

Systematic review on the clinical presentation and management of the COVID-19 associated multisystem inflammatory syndrome in

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children (MIS-C)
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Supplementary

Table S1. Summary of the different countries at which patients presented (n = 646).

| Country | Frequency % (n) | References |
|--------------------------|-----------------|--------------------------------------|
| USA | 330 (51.1) | [6,11,13,14,20–22,24,27,29,30,35,38] |
| Turkey | 1 (0.15) | [36] |
| India | 26 (4.02) | [4,37,15] |
| Saudi Arabia | 1 (0.15) | [23] |
| Poland | 1 (0.15) | [25] |
| Spain | 31 (4.8) | [39] |
| UK | 138 (21.4) | [12,15,18,40] |
| Iran | 1 (0.15) | [32] |
| Italy | 11 (1.7) | [18,26,33] |
| Algeria | 1 (0.15) | [28] |
| Brazil | 1 (0.15) | [34] |
| Thailand and Netherlands | 21 (3.25) | [16] |
| France and Switzerland | 56 (8.68) | [30,31] |
| Chile | 27 (4.18) | [41] |
| Sum | 646 (100) | - |

| Total antibiotics used | Frequency | Percentage % | References |
|------------------------|-----------|-------------------|------------------------|
| 1 | 21 | 12 (n = 21/169) | [21,36,31] |
| 2 | 16 | 9 (n = $16/169$) | [20,21-24,26,28,32,38] |
| 3 | 4 | 2(n = 4/169) | [19,21] |
| 4 | 0 | 0 | - |
| 5 | 1 | 0.6 (n = 1/169) | [35] |
| 6 | 1 | 0.6 (n = 1/169) | [37] |
| Not specified | 126 | 75 (n = 126/169) | [11,13,29,34,41] |
| Total | 169 | - | - |

Table S2. Total number of antibiotics used in patients ($n^* = 169$).

(n): Changes indicate the prevalence in those who reported the specific findings.

| First-line antibiotics | Frequency | Percentage % | References |
|---|-----------|-------------------|------------------|
| Azithromycin | 4 | 2(n = 4/169) | [26,28,35,36] |
| Cephalosporins | 27 | 16 (n = 27/169) | [19,21,24,31] |
| • 1 st generation: cefazoline | 1 | 4(n = 1/27) | [21] |
| • 3 rd generation: ceftriaxone | 20 | 74 (n = $20/27$) | [21,24,31] |
| • 4 th generation: cefepime | 5 | 18 (n = 5/27) | [21] |
| • 5 th generation: ceftaroline | 1 | 4(n = 1/27) | [19] |
| Ciprofloxacin | 1 | 0.6 (n = 1/169) | [23] |
| Clindamycin | 2 | 1(n = 2/169) | [21] |
| Linezolid | 3 | 2(n = 3/169) | [22] |
| Meropenem | 2 | 1(n = 2/169) | [32] |
| Metronidazole | 1 | 0.6 (n = 1/169) | [20] |
| Piperacillin/tazobactam | 1 | 0.6 (n = 1/169) | [37] |
| Vancomycin | 2 | 1(n = 2/169) | [21,38] |
| Not specified | 126 | 75 (n = 126/169) | [11,13,29,34,41] |
| Total | 43 | - | - |

Table S3. Primary choice of antibiotics used $(n^* = 43)$.

(n): Changes indicate the prevalence in those who reported the specific findings.

| Second-line antibiotics | Frequency $(n = 22)$ | Percentage % | References |
|---|----------------------|------------------|------------------|
| Cephalosporins | 4 | 2(n = 4/169) | [21,22,38] |
| • 3 rd generation: ceftriaxone | 3 | 75(n = 3/4) | [22,38] |
| • 4 th generation: cefepime | 1 | 25 (n = $1/4$) | [21] |
| Ciprofloxacin | 1 | 0.6 (n = 1/169) | [20] |
| Clindamycin | 1 | 0.6 (n = 1/169) | [19] |
| Doxycycline | 2 | 1(n = 2/169) | [26,37] |
| Levofloxacin | 1 | 0.6 (n = 1/169) | [35] |
| Linezolid | 3 | 2(n = 3/169) | [21] |
| Meropenem | 2 | 1(n = 2/169) | [21,22] |
| Metronidazole | 3 | 2(n = 3/169) | [21,23,28] |
| Penicillin G | 1 | 0.6 (n = 1/169) | [24] |
| Vancomycin | 4 | 2 (n = 4/169) | [21,32] |
| Not specified | 147 | 87 (n = 147/169) | [11,13,29,34,41] |
| Total | 22 | - | - |

Table S4. Secondary choice of antibiotics used $(n^* = 22)$.

(n): Changes indicate the prevalence in those who reported the specific findings.

| Third-line antibiotics | Frequency $(n = 6)$ | Percentage % (n) | References |
|-------------------------|---------------------|------------------|------------------|
| Linezolid | 2 | 1 (n = 2/169) | [21,35] |
| Meropenem | 2 | 1 (n = 2/169) | [22,37] |
| Metronidazole | 1 | 0.6 (n = 1/169) | [21] |
| Piperacillin/tazobactam | 1 | 0.6 (n = 1/169) | [19] |
| Not specified | 163 | 96 (n = 163/169) | [11,13,29,34,41] |
| Total | 6 | - | - |

Table S5. Tertiary choice of antibiotics used $(n^* = 6)$.

*(n): Changes indicate the prevalence in those who reported the specific findings.

| Treatments | LOS in days: | Outcomes | Conclusion & caveats | References |
|-----------------------------|---------------|------------|--|------------|
| | $Mean \pm SD$ | | | |
| Antibiotic alone $(n = 2)$ | 3 | 100% | Lowest LOS with best outcome. Low patient | [23,36] |
| | | discharged | numbers to make any conclusive statement. | |
| Enoxaparin alone $(n = 3)$ | 5 ± 1 | 100% | Low LOS with best outcome. Low patient | [30] |
| | | discharged | numbers to make any conclusive statement. | |
| Aspirin alone $(n = 1)$ | N/A | 100% | No LOS Data available | [27] |
| | | discharged | | |
| Aspirin and (antibiotic or | 6 ± 1 | 100% | Moderate LOS with good outcome. Low | [30,37] |
| enoxaparin) $(n = 5)$ | | discharged | patient numbers to make any conclusive | |
| | | | statement. | |
| Triple therapy: aspirin and | 9 ± 1 | 100% | Highest LOS with good outcome. Low patient | [21] |
| antibiotics and enoxaparin | | discharged | numbers to make any conclusive statement. | |
| (n = 2) | | | | |
| Antibiotics and | 8 ± 3 | 100% | High LOS with good outcome. Low patient | [19,21,22] |
| enoxaparin (n = 8) | | discharged | numbers to make any conclusive statement. | |
| Other: inotropes, | N/A | 100% | No LOS data was available. | [15] |
| remdesivir, protease | | discharged | | |
| inhibitors, HCQ $(n = 1)$ | | | | |

Table S6. Outcomes of adjunct medications in the biologics and immunoglobulin therapy treatment group ($n^* = 22$).

(n): Changes indicate the prevalence in those who reported the specific findings.



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