Early-onset meningitis with delayed presentation: Is there a role for prevention?

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In their valuable study, Snoek and co-workers¹ reported 88 infants (\leq 3 days old, gestational age \geq 34 weeks) with early-onset sepsis (EOS) and/or meningitis (EOM) due to group B streptococcus (GBS, n = 81) or E.coli (n = 7). Cases came from the Dutch prospective nationwide cohort study (2018-2021). Actual treatment performed under Dutch guidelines was compared with hypothetical indications from NICE guidelines and the US early-onset sepsis calculator (EOSC). EOSC would have led to delayed antibiotic treatment in significantly more cases than Dutch and NICE guidelines. However, the proportion of uninfected neonates exposed to antibiotics should also be considered, given the potential long-term consequences of neonatal intestinal dysbiosis.² This question was unaddressed in the study, but one UK network reports high rates of neonatal antibiotic treatment (14%) with NICE guidelines (from which Dutch guidelines are adapted).3

Despite the very inclusive criteria for treatment under the Dutch guidelines, the overall percentage of culture-proven EOMs (13%) was higher compared with US areas adopting a maternal prenatal screening (\sim 6%).⁴ Most EOMs (90%) occurred in infants who were not started on treatment until after 24 h of age: we wonder if these EOMs with delayed presentation had been exposed to intrapartum antibiotic prophylaxis (IAP). Indeed IAP, by lowering colony counts in GBScolonized mothers, protects neonates from infection that would be acquired during passage through the birth canal.5 Thus, IAP-exposed neonates who are healthy-appearing at birth are less likely to develop GBS-EOS and EOM with delayed presentation. Identification and intrapartum treatment of GBS colonized women prevents almost all EOMs.

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Abbreviations: EOS, Early-onset sepsis; EOM, Early-onset meningitis; GBS, Group B streptococcus; EOSC, Ealry-onset sepsis calculator; IAP, Intrapartum antibiotic prophylaxis

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Authorship statement

All authors substantially contributed to the work. Prof. Berardi conceptualized the letter, drafted the initial manuscript, reviewed and edited it and supervised the study. Dr. Miselli contributed to conceptualization, drafted the initial manuscript and reviewed and edited it. Dr. Lugli, Bedetti and Zinani contributed to conceptualization, and reviewed and edited the manuscript.

All authors gave final approval of the version to be published and agree to be accountable for all aspects of the work.

Declaration of interests

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References

- Snoek L, Kassel MNV, Krommenhoek JF, et al. Neonatal early-onset infections: comparing the sensitivity of the neonatal early-onset sepsis calculator to the Dutch and the updated NICE guide-lines in an observational cohort of culture-positive cases. *eClinical-Medicine*. 2022;14. https://doi.org/10.1016/j.eclinm.2021.101270.
 Zeissig S, Blumberg RS. Life at the beginning: perturbation of the
- 2 Zeissig S, Blumberg RS. Life at the beginning: perturbation of the microbiota by antibiotics in early life and its role in health and disease. *Nat Immunol.* 2014;15:307–310.
- 3 Goel N, Cannell S, Davies G, et al. Implementation of an adapted sepsis risk calculator algorithm to reduce antibiotic usage in the management of early onset neonatal sepsis: a multicentre initiative in Wales, UK. Arch Dis Child Fetal Neonatal Ed. 2021. fetalneonatal-2020-321489.
- Schrag SJ, Farley MM, Petit S, et al. Epidemiology of invasive earlyonset neonatal sepsis, 2005 to 2014. *Pediatrics*. 2016; 138: e20162013.
- Berardi A, Spada C, Vaccina E, Boncompagni A, Bedetti L, Lucaccioni L. Intrapartum beta-lactam antibiotics for preventing group B streptococcal early-onset disease: can we abandon the concept of 'inadequate' intrapartum antibiotic prophylaxis? *Expert Rev Anti Infact Ther*. 2020;18:37–46.

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