Supplemental Digital Content 1. Summary of existing World Health Organization (WHO) guidance for empirical antibiotic therapy for meningitis in neonates and young infants

| Date | Document Title | Age group | First-line antibiotics | If no improvement 48 hours | Comments |
|------|---|--------------------|------------------------------------|----------------------------|---|
| 1999 | Conclusions from the WHO Multicenter Study | < 90 days | Third generation cephalosporin | Not discussed | Ampicillin and gentamicin for initial therapy of neonatal |
| | of Serious Infections in Young Infants ²¹ | | | | sepsis and anti-Staphylococcal agent for skin sepsis |
| 2000 | Management of the Child with a Serious | 1 week to 2 | Ampicillin + gentamicin or | Change to ceftriaxone or | one or Change antibiotics to organism sensitivities if known |
| | | | Ampicillin + chloramphenicol | 5 | |
| | Infection or Severe Malnutrition ²² | months | (if >37 weeks gestation) | Cefotaxime | |
| 2003 | Managing Newborn problems ²³ | < 1 week | Ampicillin + gentamicin | Replace ampicillin with | Repeat LP after 48 hours if no improvement. |
| | | | | cefotaxime | Change antibiotics to organism sensitivities if known. |
| 2005 | IMCI ²⁴ | 1 week to 2 | Benzylpenicillin + gentamicin Basi | Basic care guidelines* | Guidance covers all possible serious bacterial infections † |
| | | months | Benzyipeniciiini + gentamicin | | |
| 2005 | Pocketbook of Hospital Care for Children ²⁵ | < 2 months | Ampicillin/Ben Pen + gentamicin or | Not discussed | Document notes that ceftriaxone may cause biliary sludging |
| | | | Ampicillin + chloramphenicol or | | |
| | | | Ceftriaxone / Cefotaxime | | leading to jaundice |
| 2008 | IMCI Chart Booklet - Standard ²⁶ | < 2 months | Ampicillin + gentamicin | Basic care guidelines* | Guidance covers "possible serious bacterial infections" † |
| 2008 | IMCI for high HIV setting - Chart Booklet ²⁷ | < 2 months | Ampicillin + gentamicin | Basic care guidelines* | Guidance covers "very severe disease" † |
| 2008 | IMCI Adaptation Guide; Part 2c ²⁸ | 1 week to 2 months | Benzylpenicillin + gentamicin | Basic care guidelines* | Guidance covers "possible serious bacterial infections". † |
| | | | | | Cefotaxime or ceftriaxone if gentamicin unavailable or |
| | | | | | penicillin-resistant S. pneumoniae |

*Guidelines aimed at first-level care settings and recommend referral of all infants with possible serious bacterial infection.

† No separate guidelines for neonatal meningitis.

‡ If >37 weeks gestation

IMCI = Integrated Management of Childhood Illness