#### • Suspected early-onset sepsis

- if hospitalized in the NICU, or for patients < 4 weeks of age with communityacquired infections, including suspected community-acquired pneumonia and urinary tract infection
- Most likely organisms: Group B strep, Enterobacteriaceae, Enterococcus spp, Listeria, HSV
- First line: ampicillin + tobramycin
- If suspicion of meningitis: replace tobramycin with cefotaxime
  - IF **CSF gram stain** shows **gram positive** cocci or gram positive rods, **add gentamicin** 1 mg/kg/dose IV q8h, for synergy in case of GBS or Listeria.
- if suspicion of **HSV**, add **acyclovir**; or add it if presence of 1 or more of following:
  - Ill-appearing
  - Altered mental status
  - Hypothermia
  - Seizures
  - Presence of vesicles
  - Exposure to maternal genital HSV lesions
  - Elevated ALT
  - CSF pleocytosis (using standard neonatal reference ranges) with a negative gram stain + leukopenia or thrombocytopenia

#### • Suspected late-onset sepsis

- In patients with more than 3 days of hospitalization in NICU
- Most likely organisms: Group B strep, Staph aureus, Enterobacteriaceae, P. aeruginosa, Enterococcus spp, Listeria, C. albicans
- First-line: cefazolin + tobramycin
- Replace **cefazolin** by **vancomycin**, as per NICU Prescription Guide, in following situations:
  - Proven infection with a methicillin-resistant gram positive bacteria
  - Colonization with MRSA
  - Unstable neonates (hypotension, increase in apneas and bradycardias requiring intubation, significant metabolic acidosis)
  - Neonates less than 34 weeks corrected gestational age in presence of a central line
  - Neonates with intracranial devices (VP shunt, subgaleal shunt, external ventricular shunt)
- IF VAP or NEC also suspected, use piperacillin-tazobactam instead
- If meningitis suspected, use meropenem instead
- If invasive candidiasis suspected, add amphotericin B deoxycholate, or

consider adding it in presence of following clinical clues:

- If persistent hypoglycemia and thrombocytopenia
- If multi-organ involvement
- If no response to broad-spectrum antibiotics
- if suspicion of **HSV**, add **acyclovir**; or add it if presence of 1 or more of following:
  - Ill-appearing
  - Altered mental status
  - Hypothermia
  - Seizures
  - Presence of vesicles
  - Exposure to maternal genital HSV lesions
  - Elevated ALT
  - CSF pleocytosis (using standard neonatal reference ranges) with a negative gram stain + leukopenia or thrombocytopenia

### <u>Necrotizing enterocolitis</u>

- Most likely organisms: Enterobacteriaceae, Bacteroides spp., Other anaerobes, Enterococus spp
- Stage 1 (suspected): piperacillin-tazobactam
- stage 2 (definite) : piperacillin-tazobactam
  - o If *meningitis* suspected, use *meropenem* instead
- Stage 3 (advanced), with septic shock and/or meningitis suspected: meropenem
  - If above considerations no longer present, narrow down agent spectrum to piperacillin/tazobactam

#### • Suspected central-line associated bloodstream infection (CLABSI):

- For patients hospitalized in NICU
- Most likely organisms: CoNS, Staph aureus, Enterobacteriaceae, Enterococcus spp, P. aeruginosa
- First line: cefazolin + tobramycin
- Replace **cefazolin** by **vancomycin** as per NICU Prescription Guide, in following situations:
  - Proven infection with a methicillin-resistant gram positive bacteria
  - Colonization with MRSA
  - Unstable neonates (hypotension, increase in apneas and bradycardias requiring intubation, significant metabolic acidosis)
  - Neonates less than 34 weeks corrected gestational age in presence of a central line
  - Neonates with intracranial devices (VP shunt, subgaleal shunt, external ventricular shunt)

#### • Hospital-acquired pneumonia

- including ventilator-associated pneumonia and aspiration pneumonia in hospitalized patient
- refer to late-onset sepsis for other pneumonia indications in patients hospitalized for more than 3 days
- Most likely organisms: Staph aureus, Enterobacteriaceae, Pseudomonas, A. baumannii
- First line: piperacillin-tazobactam
- If severe: replace by meropenem

## Healthcare-associated intra-abdominal infection

- Including surgical site infection, peritonitis, and spontaneous intestinal perforation in NICU
- Most likely organisms: Enterobacteriaceae, Bacteroides spp., Other anaerobes, Enterococus spp., Pseudomonas spp.
- Mild-moderate disease: Piperacillin/tazobactam
- Severe (e.g. concomitant sepsis, septic shock): Meropenem

## Healthcare-associated urinary tract infection

- e.g. catheter-associated, procedure-related
- refer to late-onset sepsis for other UTI indications in patients hospitalized for more than 3 days
- o Most likely organisms: Enterobacteriaceae, Pseudomonas, Enterocococcus spp,
  - Rarely: Staph aureus, Candida spp
- First-line: piperacillin-tazobactam
- Empiric antifungal therapy is not recommended as catheter-associated UTI due to Candida usually resolves with urinary catheter removal

- Surgical site infections: head, neck, trunk, extremity wound
  - Most likely organisms: Group A strep, Staph aureus, Clostridium spp.
  - First line: cefazolin
  - Suspected MRSA: replace cefazolin by vancomycin if mild or moderate; add vancomycin if severe

• Surgical site infections: GI tract, perineum, genital tract wound

- Most likely organisms: Staph aureus, Streptococci, Enterobacteriaceae, Anaerobes
- First line: piperacillin-tazobactam

# • <u>Healthcare-associated meningitis (e.g. CSF shunt infection, post-neurosurgery):</u>

- Most likely organisms: Staph aureus, CoNS, P. acnes, Enterobacteriaceae, Pseudomonas
- First line: meropenem + vancomycin
- May switch to cloxacillin if oxacillin-sensitive staph isolated.