

Clinical Policy: Neonatal Sepsis Management

Reference Number: CP.MP.85

Last Review Date: 07/19

[Revision Log](#)

See [Important Reminder](#) at the end of this policy for important regulatory and legal information.

Description

Through the increased incidence of intra-partum antibiotics, early-onset neonatal sepsis is occurring less frequently. However, it continues to be a common cause of neonatal morbidity and mortality. The CDC (Centers for Disease Control and Prevention) defines early onset sepsis as a blood or cerebrospinal fluid culture-proven infection occurring within the first seven days of life.

Group B Streptococcus (GBS) remains the leading cause of neonatal sepsis. More than half of GBS cases occur in infants of mothers with negative GBS cultures, emphasizing the need to remain vigilant for signs of sepsis in all newborns. These infants require comprehensive assessment and treatment, as well as discharge planning, in order to ensure timely treatment in an effort to reduce morbidity and mortality.²

Policy/Criteria

It is the policy of Health Plans affiliated with Centene Corporation that the management of neonatal sepsis is **medically necessary** at the indicated level of care for the following circumstances:

I. Episode Day 1

- A. Well-appearing infants who are on 48 hours of antibiotics pending blood culture results are appropriate for level II (rev code 172) nursery.
- B. Symptomatic infants are appropriate for level III (rev code 173) nursery with all the following:
 1. Hypotonia, lethargy, or poor oral feeding;
 2. Temp $\geq 100.4^{\circ}\text{F}$ or $\leq 96.8^{\circ}\text{F}$ ($\geq 38.0^{\circ}$ or $\leq 36.0^{\circ}\text{C}$);
 3. On 48 hours of antibiotics pending blood culture results or treatment of positive blood cultures.

II. Episode Day 2 and Subsequent

- A. Infants with negative cultures who are determined to require antibiotics beyond 48 hours may be appropriate for transitional care or level I nursery (rev code 171) once antibiotics are the only intervention necessitating continued stay and if outpatient antibiotics are inappropriate or unmanageable.
- B. Symptomatic infants are appropriate for level III (rev code 173) nursery when meeting the following criteria:
 1. Hypotonia, lethargy, or poor oral feeding; and
 2. Temp $\geq 100.4^{\circ}\text{F}$ or $\leq 96.8^{\circ}\text{F}$ ($\geq 38.0^{\circ}$ or $\leq 36.0^{\circ}\text{C}$); and
 3. On 48 hours of antibiotics pending blood culture results or treatment of positive blood cultures.
- C. Asymptomatic infants on 48 hours of antibiotics pending blood culture results for ≤ 2 days are appropriate for Level II (rev code 172) nursery.
- D. Asymptomatic infants with a positive blood culture and no other indications are appropriate for transitional care or level I nursery (rev code 171).

Once the culture and sensitivity results are known and antibiotic therapy is established, a medically stable infant should be transitioned to a lower level of care for treatment completion if no other indications exist that require the current level of care. Transitional care nursery should be considered if antibiotics cannot safely be administered at home or at home with home health care.

III. Discharge criteria, with or without home antibiotics, meets all, as applicable:

- A. Member is clinically stable;
- B. Home situation is assessed and deemed adequate;
- C. Parent or caretaker is agreeable with the plan of care;
- D. If going home with antibiotics, all the following are met:
 - 1. A home infusion company is contracted which is experienced in neonatal IV (intravenous) therapy or short-term intramuscular therapy;
 - 2. Secure IV access is in place if chosen;
 - 3. The responsible physician (neonatologist, primary care pediatrician) and back-up health care facility (neonatal intensive care unit [NICU], community hospital) should be clarified to the family and home care agency prior to discharge.

Background

I. Identification and treatment of mother during pregnancy and labor⁹

- A. Women with GBS isolated in the urine any time during the current pregnancy or who had a previous infant with invasive GBS disease should receive IV intrapartum antibiotic prophylaxis. Third trimester screening for GBS colonization is not needed in this population.
- B. Women with symptomatic or asymptomatic GBS urinary tract infection (UTI) detected during pregnancy should be treated according to current standards of care for UTI during pregnancy and should receive intrapartum antibiotic prophylaxis to prevent early-onset GBS disease.
- C. All other pregnant women, including those with a scheduled cesarean delivery, should be screened at 36 0/7 – 37 6/7 weeks gestation for vaginal and rectal GBS colonization.
- D. At the time of labor or rupture of membranes, intrapartum antibiotic prophylaxis should be given to all pregnant women whose vaginal-rectal cultures were positive for GBS colonization, including those undergoing cesarean delivery. If cesarean delivery is performed before onset of labor on a woman with intact amniotic membranes, prophylaxis need not be given.
- E. When screening results are not available at the time of labor and delivery, intrapartum antibiotics should be given to women who present in labor with a substantial risk of preterm birth, who have preterm prelabor rupture of membranes (PPROM), rupture of membranes ≥ 18 hours at term, or who present with intrapartum temperature $\geq 100.4^{\circ}$ F ($\geq 38.0^{\circ}$ C). If a woman has none of the risks above but has a history of GBS colonization in a previous pregnancy, it is reasonable to offer intrapartum antibiotic prophylaxis and/or discuss it as an option in a shared decision-making process with the provider.
- F. If intraamniotic infection is suspected, broad-spectrum antibiotic therapy that provides coverage for polymicrobial infections as well as GBS should replace the antibiotic that provides coverage for GBS prophylaxis specifically.

- G. In the absence of GBS UTI that is symptomatic, or with GBS present at levels $\geq 10^5$ colony forming units (CFU)/mL, antimicrobial agents should not be used before the intrapartum period to eradicate GBS genitoretal colonization because such treatment has not been shown to provide better maternal or neonatal outcomes.
- H. Obstetric interventions, when necessary, should not be delayed solely to provide 4 hours of antibiotic administration before birth.

II. Identification and Treatment of the newborn

- A. Infants born at ≥ 35 weeks' gestation should be managed according to risk, as determined by categorical risk assessment, multivariate risk assessment (the Neonatal Early-Onset Sepsis Calculator), or clinical condition.¹²
- B. Preterm infants born at ≤ 34 weeks' are at highest risk for early-onset sepsis (EOS) in the following circumstances: Infants born preterm because of cervical insufficiency, preterm labor, PROM, intraamniotic infection, and/or acute and otherwise unexplained onset of nonreassuring fetal status are at the highest risk of EOS and GBS EOD. A blood culture and initiation of empirical antibiotic treatment is recommended for infants with any of these risk factors.¹²
- C. Preterm infants born at ≤ 34 weeks' are at lower risk for EOS when meeting all of the following: 1) maternal and/or fetal indications for preterm birth (such as maternal preeclampsia or other noninfectious medical illness, placental insufficiency, or fetal growth restriction), (2) birth by cesarean delivery, and (3) absence of labor, attempts to induce labor, or any ROM before delivery.
- D. For lower risk preterm infants, initial approaches include (1) no laboratory evaluation and no empirical antibiotic therapy or (2) blood culture and clinical monitoring. For infants who do not improve after initial stabilization and/or those who have severe systemic instability, the administration of empirical antibiotics may be reasonable but is not mandatory.¹²
- E. Physicians should use their best judgment to determine when cerebrospinal fluid analysis should be performed in the absence of bacteremia, as culture-confirmed meningitis in the absence of culture-confirmed bacteremia is approximately 1 to 2 cases per 100,000 live births.¹⁰ However, the rate of meningitis is higher in preterm infants, and a lumbar puncture for culture and analysis of CSF should be considered in clinically ill infants when there is a high suspicion for GBS EOD, unless the procedure will compromise the neonate's clinical condition.¹²
- F. Therapy for a term infant at risk of EOS should include antimicrobial agents active against GBS (including IV ampicillin and gentamicin), as well as other organisms that might cause neonatal sepsis, such as *E. coli*, et al.¹⁰
- G. Early-onset GBS infection is diagnosed by blood or CSF culture. Common laboratory tests such as the complete blood cell count and C-reactive protein do not perform well in predicting early-onset infection, particularly among well-appearing infants at lowest baseline risk of infection.¹²

III. General Considerations

- A. Stable infants at 35 weeks gestational age or older who are treated for sepsis should be discharged the same day the antibiotics are discontinued.

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- B. For ruling out sepsis due to perinatal risk factors, 48 hours of antibiotic administration is considered appropriate pending culture results and evaluation of lab data.
- C. When blood cultures are sterile, antibiotic therapy should be discontinued by 36 to 48 hours of incubation unless there is clear evidence of site-specific infection.¹⁰

Reviews, Revisions, and Approvals	Date	Approval Date
Policy developed, Neonatologist reviewed	08/13	08/13
Renamed “Authorization Protocol” to Policy/Criteria and reformatted text to episode days and discharge criteria. Remaining information moved to background. Background updated regarding timing of lumbar puncture Updated nursery levels based on updated Interqual criteria nursery leveling Neonatologist reviewed	09/14	09/14
References reviewed and updated; converted to new template. Neonatologist reviewed	09/15	09/15
Clarified bullets under III.D. No criteria change	04/16	
References reviewed and updated. Minor wording changes for clarification.	08/16	09/16
References reviewed and updated.	09/17	09/17
References reviewed and updated.	07/18	07/18
Policy reviewed by neonatologist and pediatrician.	05/19	
Edits to background information regarding identification and treatment of the mother, and identification and treatment of the newborn, per new ACOG and AAP guidelines.	07/19	07/19
Edits to background information regarding identification and treatment of the newborn per new AAP guidelines.	10/19	

References

1. Centers for Disease Control and Prevention. Prevention of perinatal group B streptococcal disease—revised guidelines from CDC, 2010. *MMWR Recomm Rep.* 2010;59(RR-10):1–36.
2. Polin RA. Management of Neonates with Suspected or Proven Early-Onset Bacterial Sepsis. *Pediatrics.* 2012; 129; 1006.
3. Newton ER, Clark M. Group B streptococcus and preterm rupture of membranes. *Obstet Gynecol.* 1988;71(2):198–202pmid:3275914
4. Schuchat A, Zywicki SS, Dinsmoor MJ, et al. Risk factors and opportunities for prevention of early-onset neonatal sepsis: a multicenter case-control study. *Pediatrics.* 2000;105(1 pt 1):21–26pmid:10617699
5. Schrag SJ, Hadler JL, Arnold KE, Martell-Cleary P, Reingold A, Schuchat A. Risk factors for invasive, early-onset *Escherichia coli* infections in the era of widespread intrapartum antibiotic use. *Pediatrics.* 2006;118(2):570–576pmid:16882809
6. Martius JA, Roos T, Gora B, et al. Risk factors associated with early-onset sepsis in premature infants. *Eur J Obstet Gynecol Reprod Biol.* 1999;85(2):151–158pmid:10584628.
7. Polin RA, Watterberg K, Benitz W, Eichenwald, E. The conundrum of early-onset sepsis. *Pediatrics* 2014;133;1122; originally published online May 5, 2014. <http://pediatrics.aappublications.org/content/133/6/1122>

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8. Edwards MS. Clinical features, evaluation and diagnosis of sepsis in term and late preterm infants. In: UpToDate, Kaplan SL, Garcia-Prats JA (Eds), UpToDate, Waltham, MA. Accessed 07/1/19.
9. American College of Gynecologists and Obstetricians Committee on Obstetric Practice. ACOG Committee Opinion 782: Prevention of Group B Streptococcal Early-Onset Disease in Newborns. *Obstet Gynecol* 2019; 134:e19-40.
10. Puopolo KM, Benitz WE, Zaoutis TE and the committee on Fetus and Newborn, and Committee on Infectious Diseases. Management of Neonates Born at ≥ 35 0/7 Weeks' Gestation With Suspected or Proven Early-Onset Bacterial Sepsis. *Pediatrics* Dec 2018, 142 (6) e20182894.
11. Karen M. Puopolo, William E. Benitz, Theoklis E. Zaoutis, Committee on Fetus and Newborn, Committee on Infectious Diseases. Management of Neonates Born at ≤ 34 6/7 Weeks' Gestation With Suspected or Proven Early-Onset Bacterial Sepsis. *Pediatrics* Dec 2018, 142 (6) e20182896.
12. Karen M. Puopolo, Ruth Lynfield, James J. Cummings, Committee on Fetus and Newborn, Committee On Infectious Diseases. Management of Infants at Risk for Group B Streptococcal Disease. *Pediatrics* Aug 2019, 144 (2) e20191881.

Important Reminder

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. The Health Plan makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved. "Health Plan" means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan's affiliates, as applicable.

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Note: For Medicare members, to ensure consistency with the Medicare National Coverage Determinations (NCD) and Local Coverage Determinations (LCD), all applicable NCDs, LCDs, and Medicare Coverage Articles should be reviewed prior to applying the criteria set forth in this clinical policy. Refer to the CMS website at <http://www.cms.gov> for additional information.

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