


Original Article

Blood culture procedures and practices in the neonatal intensive care unit: A survey of a large multicenter collaborative in California

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Abstract

Objective: To describe variation in blood culture practices in the neonatal intensive care unit (NICU).

Design: Survey of neonatal practitioners involved with blood culturing and NICU-level policy development.

Participants: We included 28 NICUs in a large antimicrobial stewardship quality improvement program through the California Perinatal Quality Care Collaborative.

Methods: Web-based survey of bedside blood culture practices and NICU- and laboratory-level practices. We evaluated adherence to recommended practices.

Results: Most NICUs did not have a procedural competency (54%), did not document the sample volume (75%), did not receive a culture contamination report (57%), and/or did not require reporting to the provider if <1 mL blood was obtained (64%). The skin asepsis procedure varied across NICUs. Only 71% had a written procedure, but ≥86% changed the needle and disinfected the bottle top prior to inoculation. More than one-fifth of NICUs draw a culture from an intravascular device only (if present). Of 13 modifiable practices related to culture and contamination, NICUs with nurse practitioners more frequently adopted >50% of practices, compared to units without (92% vs 50% of units; $P < .02$).

Conclusions: In the NICU setting, recommended practices for blood culturing were not routinely performed.

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Clinician confidence in the reliability of blood cultures is critical to the appropriate diagnosis and management of a bloodstream infection.¹ This is particularly true in the neonatal intensive care unit (NICU) setting in which non-culture-based biomarkers are not reliably predictive for bacterial sepsis.² Nonadherence to recommended practices may lead to false-negative or false-positive blood culture results that have important clinical impacts. These include delays in correct diagnosis, inappropriate antibiotic use with attendant adverse events (including potentially deleterious changes to the microbiome³), prolonged hospitalization, excess diagnostic or therapeutic procedures (eg, echocardiography, specialty consultations, catheter removal, among others), and direct and indirect economic costs.⁴

False-positive results due to contamination⁴ during the blood draw itself and less commonly, false-negative culture results due to inadequate sample volume,⁵ occur frequently in the

NICU. Both scenarios are problematic in view of the tendency toward conservative management which often follows, given the particular vulnerability of these infants.⁶ A national “standard” of <3% contamination rate⁷ (developed mainly for adults, in whom rates average 0.6–12.5%⁴) has recently been changed to 1% by the US Clinical and Laboratory Standards Institute, given the clinical impacts noted here.⁸ Several studies in neonatal populations demonstrate that contamination occurs at rates between 2.6% and 18%.¹ Improved adherence to available recommendations can lower contamination rates in the NICU⁹ and therefore predictably improve antimicrobial stewardship.

Nurses draw most blood cultures in NICUs, yet no recent data are available regarding adherence to contemporary practice standards for this critical procedure, which is very frequently used in this population. Several factors associated with NICU policy development, staff training and bedside processes directly affect the reliability of a blood culture result. We surveyed NICUs participating in an antibiotic stewardship collaborative in California on unit- and nurse-level blood culture practices and conformity with available recommendations to identify areas for practice improvement.

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Methods

We performed this survey as part of an antimicrobial stewardship collaborative managed by the California Perinatal Quality Care Collaborative (CPQCC) in partnership with the RAND Corporation and the Lundquist Institute at Harbor-UCLA Medical Center. CPQCC is a statewide network of >130 NICUs that work to improve the quality of clinical care through targeted and formal quality intervention projects (cpqcc.org). This stewardship project is known as the Optimizing Antibiotic Stewardship in California NICUs (OASCN, cpqcc.org/improvement/projects/OASCN). OASCN began with 31 NICUs and hosted learning sessions via a secure Zoom televideo platform link every 2 weeks from early March 2021 through February 2022. All clinical staff in participating NICUs (including physicians, nurses, pharmacists, and trainees) were invited to participate. We identified site leaders for each NICU as the administrative leads responsible for encouraging NICU staff to participate in the learning sessions and other OASCN activities. Approximately 60% of the site leaders were neonatologists; 30% were clinical nurse specialists, nurse managers, or educators; and the rest were neonatal nurse practitioners (NNPs) and pharmacists. More details on OASCN, including our use of didactics, real-time polls, feedback, a reference library, and an associated expert faculty panel, are provided in the Supplementary Material (online).

We performed a standalone web-based survey of OASCN NICUs regarding practices related to blood culturing and prevention of contamination. Implemented using a Qualtrics platform (qualtrics.com), it included 19 questions (~10-15 minutes to fully complete) related to policy, practice, documentation, written procedures, competency requirements and laboratory practices for reporting early results and contamination data (Supplementary Material online). We developed the survey after a review of the literature and published guidelines for reducing blood culture contamination, as well as related concerns expressed by providers during prior OASCN learning sessions and a session in April 2021 focused specifically on this topic. In mid-October 2021, 6 months later, a link to the survey was sent by email to the site leaders at each of the 29 OASCN NICUs participating at that time. We asked site leaders to distribute the survey to the person most knowledgeable about blood culture practice in their NICU. Up to 5 reminders were sent to those who had not yet responded. On November 18, 2021, OASCN held a nursing office hour on this topic separate from the usual sessions. On February 3, 2022, survey results were reported back to the collaborative as were practice improvements relayed by participants. We performed descriptive analyses of survey responses for all participating NICUs and by selected characteristics of the group. We also evaluated responses in relation to available national recommendations regarding blood culture practice in the NICU setting, when applicable. This work was approved by the RAND Human Subjects Protection Committee, the Stanford Panel on Medical Human Subjects and the John F. Wolf Human Subjects Committee of The Lundquist Institute, in accordance with the Declaration of Helsinki.

Results

Survey responses were received from 28 of the 29 OASCN NICU site leaders. The characteristics of the responding units are detailed in Table 1. The distribution of NICU size, California Children's Services (CCS) level, American Academy of Pediatrics (AAP) level of care, and hospital owner type were similar to that of the CPQCC overall (data not shown).

Table 1. Characteristics of Participating OASCN NICUs

Feature	No. (%) ^a
NICU size^b	
First tertile	8 (29)
Second tertile	8 (29)
Third tertile	12 (43)
CCS level^c	
Community	18 (64)
Regional	6 (22)
Intermediate	2 (7)
Non-CCS hospital	2 (7)
AAP level of care^d	
Level II	4 (14)
Level III	20 (72)
Level IV	4 (14)
Hospital owner	
Nonprofit	16 (57)
Government	6 (21)
Investor owned	6 (21)
Nurse practitioners work in the NICU	
Yes	12 (43)
No	16 (57)
Freestanding childrens hospital	
Yes	1 (4)
No	27 (96)

Note. OASCN, Optimizing Antibiotic Stewardship in California NICUs; NICU, neonatal intensive care unit; CCS, California Children's Services; CPQCC, California Perinatal Quality Care Collaborative; AAP, American Academy of Pediatrics.

^aTotals may not add to 100% due to rounding.

^bBased on total admissions of all CPQCC NICUs in 2020 in which tertile 1 was ≤229 admissions, tertile 2 was 230–365 admissions, and tertile 3 was 366–1,188 admissions. In OASCN, tertile 1 was 278 admissions, tertile 2 was 279–361 admissions, and tertile 3 was 362–877 admissions.

^cCCS is a state program that provides diagnostic and treatment services, medical case management, and physical and occupational therapy services to children aged < 21 years with selected medical conditions. CCS is administered as a partnership between county health departments and the California Department of Health Care Services (dhcs.ca.gov/services/ccs/Pages/default.aspx).

^dThe AAP levels of care are Level I, II, III, IV and commonly known as wellborn nursery, special care, NICU, and regional NICU, respectively (AAP comm fetus and newborn. *Pediatrics* 2012;130:587–597). This study did not include units with only level I care availability.

Survey responses are shown in Table 2. Notably, a large percentage of NICUs did not adhere to certain commonly accepted procedures associated with obtaining blood cultures and/or facilitate accurate interpretation of the results. These included having a standard order set (46% did not), mandating documentation of the volume obtained and site of draw (75% and 32% did not, respectively), having a nursing competency on this procedure (54% did not). Also, 64% did not require reporting to the provider if <1 mL was obtained. Furthermore, 75% of units were not aware of or did not receive a routine contamination report from the laboratory. Of the 7 laboratories that did provide such a report, 6 reported only to the department of infection prevention and control. Also, of those units that received reports, less than half had a standard process to follow up with the staff member who drew the contaminated specimen.

Table 2. Responses to OASCN Nursing Survey Questions

Feature	Yes, No. (%)	No, No. (%)	Unknown, No. (%)
Standard order set for culture? ^a	12 (43)	13 (46)	3 (11)
CHG-containing prep for skin asepsis?	23 (79)	5 (18)	...
Alcohol cap used on devices?	20 (71)	7 (25)	1 (4)
Required to report draw site? ^b	18 (64)	9 (32)	1 (4)
Must volume be recorded?	6 (21)	21 (75)	1 (4)
Written procedure for culturing?	20 (71)	6 (21)	2 (7)
Needle change before inoculate into bottle?	24 (86)	3 (11)	1 (4)
Prep bottle before inoculation?	25 (89)	2 (7)	1 (4)
Official blood culture competency?	12 (43)	15 (54)	1 (4)
Use of blood diversion? ^c	4 (14)	22 (79)	2 (7)
Minimum volume specified in unit?	25 (89)	3 (11)	...
Notify MD if <1 mL obtained?	9 (32)	18 (64)	1 (4)
Two samples drawn routinely?	4 (14)	24 (86)	...
Culture processed in your hospital?	21 (75)	7 (25)	...
Prelim positives called to unit?	25 (89) ^d	1 (4)	2 (7)
Routine contamination report?	7 (25) ^{e, f}	6 (21)	15 (54)

Note. OASCN, Optimizing Antibiotic Stewardship in California NICUs; CVC, central venous catheter; EMR, electronic medical record; NICU, neonatal intensive care unit.

^aAmong all respondents, if a vascular device (central venous catheter, peripheral IV, arterial line) was present the nurse would draw a blood culture via venipuncture (only) in 14 survey responses, vascular device (only) in 6 responses, vascular device and venipuncture in 4 responses, and CVC or venipuncture in 4 responses.

^bIf reporting was required it was documented in the EMR in 13 survey responses, bottle only in 5 respondents, bottle and the EMR in 6 responses, and was unknown by 3 respondents.

^cProcedure in which a small volume is "diverted" before use of a subsequent volume for blood culture.

^d23 laboratories call 24 hours and 7 days per week; 1 laboratory calls from 7:00 A.M. to 4:00 P.M. only; the other calls at an "unknown" time.

^e5 laboratories reported to infection prevention and control, 1 reported to the NICU physician, 1 reported to an unknown location.

^fAmong units with a contamination report, 3 (43%) followed up with the drawer.

Our review of national guidelines for blood cultures in the NICU setting revealed the lack of a consistent approach for most generally accepted principles of disinfection and/or reporting (Tables 3 and 2). Nonetheless, adherence was variable for the most widely accepted recommendations in our sample. Nearly one-fifth of responders indicated that a culture would be preferentially drawn from a pre-existing vascular device only (if present) rather than via peripheral venipuncture. Approximately 11% of NICUs did not change needles prior to inoculation and 7% did not disinfect the bottle prior to injection. Nearly one-third of units were not required to document the site of the blood draw. Also, 5 sites (18%) did not use a chlorhexidine-containing product for skin asepsis, regardless of weight or gestational or chronological age.

As noted in Table 4, of the 13 modifiable processes associated with obtaining a blood culture and prevention of contamination, NICUs with NNPs more frequently adopted >50% of the practices, compared to units without NNPs (92% vs 50% of units; $P < .02$). Nonetheless, nearly all of the other processes were also more common in units with NNPs such as having a standard order set for blood culture (6 of 12 [50%] vs 6 of 16 [38%]), having a unit blood culture competency (7 of 12 [58%] vs 5 of 16 [31%]), and a requirement to notify the provider if <1 mL of blood was obtained

(5 of 12 [42%] vs 4 of 16 [25%]), among others. Subgroup analyses by other NICU characteristics were also limited by small sample sizes (data not shown).

Discussion

In our cohort, we identified nonadherence to and quantified the variability between units for several of the most recommended blood culture practices. Most of our sites did not have certain structural processes in place such as a written neonatal-specific blood culture procedure, a requirement for nurses to complete a skills competency for blood drawing, or routine laboratory-based reporting of contamination rates. These systems-based interventions present opportunities for more direct awareness of the importance of blood culture by local leadership and for those who actually draw the blood. Development of such policies should be multidisciplinary and include those directly involved with sampling and ongoing education, and the laboratory. For example, contamination reports alert units to a potential process problem, and competencies and order sets ensure standardization of the procedure itself. Despite advancing diagnostic technology,¹⁰ blood culture remains the gold standard for identifying bacteremia in patients in whom sepsis is suspected. All of these factors are important for patient care such that real-time clinical interpretation of this important test is not undermined or deemed suspect in any way by perceived or real lapses in any specific process or procedure.

Our data significantly expand upon older surveys that queried only a limited range of culture practices and/or a subset of contemporary recommendations. In 2000, among 34 level III and IV NICUs in the United States asked about late-onset sepsis evaluations, 79% used an iodine-containing product for skin asepsis, and if present, a central venous catheter (CVC) was the preferred site for culture in >90%.¹¹ In 2010, in a survey of >700 AAP Section on Perinatal Pediatrics members, 82% reported drawing <1 mL of blood, 69% use iodine-based skin asepsis, and among patients with a central line, 75% drew a routine blood culture from that line.¹² In a national survey of level III–IV NICUs in 2014, only 27 (54%) of 50 used chlorhexidine prior to placing a peripheral intravenous line, and 24 (48%) of 50 used it to scrub a catheter hub prior to blood sampling. Also, 32 (64%) restricted its use by age or weight-based criteria.¹³ Our data confirm great variation and provide more detail on a larger group of practices known to impact blood culture reliability.

Nearly one-fifth of our surveyed NICUs reported no use of a chlorhexidine-containing product for skin asepsis, contrary to most guidelines (Table 2). Insufficient disinfection of the skin (or catheter hub or connector) site, including friction and allowing to dry, is an important source of bacterial contamination of the sample, as is poor overall technique by the individuals drawing the specimen.⁴ Notably, in 2012, the FDA changed its recommendation from "do not use" chlorhexidine to "use with care" in premature or infants <2 months of age. This is reflected in a 2022 Society for Healthcare Epidemiology of America White Paper in which the authors concluded that CHG-containing products were superior to iodine-based products and could be used safely on neonates of any gestation and age, with appropriate attention to using the minimum amount necessary and with removal of excess solution.¹⁴

Sample volume is the most important factor related to identifying bacteremia in neonates. For NICU patients, it is generally recommended to obtain at least 1 mL. This finding is supported by clinical and in vitro data,^{15,16} but opinions on the most appropriate

Table 3. Published Organizational Recommendations for Blood Culturing

Organization	Recommendations
APIC ^a (2012)	Tincture of iodine and chlorhexidine preferred over povidone-iodine; if aged <2 months, 70% isopropyl alcohol “acceptable” Top of bottle should be sterile Label should include date, time, site Prefer specimen from venipuncture, not indwelling device Blood diversion per institutional policy
Lippincott Manual of Nursing Practice ^b	No chlorhexidine if skin disorder or aged <2 months Replace needle, disinfect blood culture bottle top Document time, date, site, volume Use “most distal” site Draw at least 1 mL of blood
CLSI ^c (2022)	Chlorhexidine not for infants aged <2 months Disinfect top of bottle prior to inoculation Prefer specimen from venipuncture, not indwelling device Collect no more than 1% of blood volume (pediatric)
WHO Best Practices ^d (2010)	Avoid povidone iodine, Avoid chlorhexidine if aged <2 months Label with name, date, time

Note. APIC, Association for Professionals in Infection Control; CLSI, Clinical Laboratory Standards Institute; WHO, World Health Organization.

^awww.apic.org/Resource/TinyMceFileManager/2016/IPs_Guide_to_the_Lab_012016.pdf.

^bProcedures—Blood culture sample collection, assisting, neonatal (www.lww.com).

^c*Principles and Procedures for Blood Cultures, Second Edition*. CLSI guideline M47. Clinical and Laboratory Standards Institute; 2022.

^dWHO guidelines on drawing blood: best practices in phlebotomy (https://apps.who.int/iris/bitstream/handle/10665/44294/9789241599221_eng.pdf?sequence=1).

volume vary to a degree.^{17,18} In one study, sample size of <1 mL most commonly occurred in ill, very low-birth-weight infants >7 days of age,⁵ possibly an indirect result of concerns of causing anemia and technical difficulty. When we presented our survey results to the collaborative, we learned that concerns about inadequate volume in the blood culture samples led some clinicians to doubt the veracity of a negative result, as others have reported as well.¹ Many of our sites drew routine cultures from indwelling catheters, if present. Due to the known technical difficulties in obtaining blood from critically ill neonates, indwelling catheter samples may be the only way to obtain an adequate sample volume. If drawn from a CVC to maximize volume, it is essential to label the bottle with the site of draw (eg, CVC or peripheral) to help clinicians differentiate CLABSI from simple bacteremia for healthcare-associated infection reporting and clinical purposes.

Certain individual components of the blood culture process minimize contamination and others maximize yield. Others have suggested that a bundled approach with several interventions further enhances the reliability of blood culture results, similar to that which has become standard for the prevention of various healthcare-associated infections.¹⁹ These may include a combination of specific techniques for drawing the sample, skin and bottle-top disinfection prior to the draw, choice of culture site, standardized procedures with ongoing training, obtaining sufficient sample volume, and even use of a sterile process, among others.²⁰ Of interest, diversion of blood, during which a small volume is “diverted” before use of a subsequent volume for the blood culture and which was

Table 4. Responses to OASCN Nursing Survey Questions with Respect to Presence of Neonatal Nurse Practitioners (NNPs) in the NICU

Feature ^a	12 Units With NNPs, No. (%)	16 Units Without NNPs, No. (%)
Standard order set for culture? ^b	6 (50)	6 (38)
CHG-containing prep for skin asepsis?	12 (100)	11 (69)
Alcohol cap used on devices?	9 (75)	11 (69)
Required to report draw site?	8 (67)	10 (63)
Must volume be recorded?	4 (33)	22 (13)
Written procedure for culturing?	9 (75)	11 (69)
Needle change before inoculate into bottle?	11 (92)	13 (81)
Prep bottle before inoculation?	12 (100)	13 (81)
Minimum volume specified in unit?	11 (92)	14 (88)
Official blood culture competency?	7 (58)	5 (31)
Use of blood diversion? ^c	0	4 (25)
Notify MD if <1 mL blood obtained?	5 (42)	4 (25)
Routine contamination report? ^d	3 (25)	4 (25)

Note. OASCN, Optimizing Antibiotic Stewardship in California NICUs; NICU, neonatal intensive care unit; CHG, chlorhexidine gluconate; MD, medical doctor.

^aOf the 13 modifiable processes associated with obtaining a blood culture and prevention of contamination, NICUs with nurse practitioners more frequently adopted >50% of the practices, compared to units without (92% vs 50% of units; $P < .02$).

^bAmong units with a standard order set, 3 (50%) and 1 (17%) of units with and without NNPs designated a site of draw, respectively.

^cProcedure in which a small volume is “diverted” before use of a subsequent volume inoculation.²¹

^dAmong units with a contamination report, 1 (33%) and 2 (50%) of units with and without NNPs followed up with the drawer, respectively.

adopted by a small minority of units in our study, has been shown to be effective in minimizing contamination in adults.²¹ However, this procedure has not been recommended by any major recommending body for any age group. No such bundle has been specifically proposed or validated. We believe such a bundle would be a very useful tool for NICUs.

In our study, the presence of NNPs increases the likelihood of useful interventions around blood culturing being implemented therefore potentially facilitating antibiotic stewardship. Most NNPs work in level III and IV units, but more than one-third practice in level I and II units.²² A 2018 national survey of 171 NNPs revealed that >80% were involved in quality improvement and policy development beyond direct patient care.²² In addition, NNPs report that individual autonomy and collaboration with physicians are vital components of instituting change,²³ which are critical to implementation of unit-level best practices. Our data were not of sufficient granularity to probe causality between the overrepresentation of NNPs among NICUs with more adherence to guidelines. It may indeed reflect initiative by the NNPs to work with their unit leadership to institute recommended blood culture processes. It may also be a spurious finding resulting from greater resources and therefore capability to institute process change. Although NNP-containing units skewed toward a higher number of patients than those without NNPs, the distributions of other characteristics were similar, such as CCS level, AAP level of care, and number of neonatologists. NNPs have been shown to provide clinically effective care.²³ Further work to better characterize how NNPs may

contribute to quality improvement regarding blood culturing is clearly needed.

This study had several limitations. Our survey methodology did not include audit of responses as reported by the site leader. For example, more than half of respondents did not know whether a laboratory contamination report was provided. However, it is highly unlikely that action items following such a report would not be known to the site leader (if the report existed) because two-thirds of the leaders were neonatologists practicing in the unit. Our work (performed within the larger OASCN collaborative stewardship goals) did not include a prospective, formal, quality-improvement program around blood culturing, but follow-up evaluations and commentary suggested benefit following our feedback sessions. Lastly, our survey sample size was relatively small and ultimately included only ~20% of all NICUs in the State of California. The characteristics of our group are nonetheless similar to the CPQCC as a whole.

Fundamental individual components of a successful blood culturing process for neonates are well known: process standardization, chlorhexidine for asepsis in most infants, correct needle management, alcohol for bottle preparation, obtaining adequate volume, preparation of catheter hub if an indwelling catheter is used for sampling, ongoing education with competency requirements, and contamination monitoring with feedback.^{24,25} Several reports of formal quality initiatives focused on NICU blood culturing have shown that such efforts can improve bedside processes, reduce blood culture contamination, increase sample volumes, and minimize error in a sustainable way regardless of the level of neonatal care.^{9,26,27} No single national standard that addresses all meaningful components of blood drawing, training, and contamination reporting in the NICU is available. Such a document would be a useful tool for this vulnerable population. The practice variation we identified may, in part, reflect this reality.

Trust in the blood culture result is paramount to good clinical care, and better antimicrobial stewardship, yet this requires adherence to best practices and understanding by clinicians of its predictive value. For example, a common clinical challenge in the NICU is the ill baby presumed to have “culture-negative sepsis” leading to empiric broad-spectrum antibiotic therapy.¹ If all processes designed to increase reliability of this test were implemented, and clinicians were convinced of this at the unit level, this “diagnosis” (and therapy) may be mitigated over time.²⁸ In 2017, antibiotic use in CPQCC NICUs varied regardless of the CCS level or rate of proven early- or late-onset sepsis.²⁹ The Centers for Disease Control and Prevention (CDC) National Healthcare Safety Network now strongly encourages reporting of antimicrobial use, including for patients in NICUs. As the neonatal community continues to move to reduce antibiotic exposure,³⁰ strict adherence to and monitoring of best practices for obtaining a blood culture should be an important part of the process.

Supplementary material. To view supplementary material for this article, please visit <https://doi.org/10.1017/ice.2023.33>

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References

- Klingenberg C, Kornelisse RF, Buonocore G, Maier RF, Stocker M. Culture-negative early-onset neonatal sepsis—at the crossroad between efficient sepsis care and antimicrobial stewardship. *Front Pediatr* 2018;6:285.
- Cantey JB, Lee JH. Biomarkers for the diagnosis of neonatal sepsis. *Clin Perinatol* 2021;48:215–222.
- Wong E, Lui K, Day AS, Leach ST. Manipulating the neonatal gut microbiome: current understanding and future perspectives. *Arch Dis Child Fetal Neonatal Ed* 2022;107:346–350.
- Doern GV, Carroll KC, Diekema DJ, *et al.* A comprehensive update on the problem of blood culture contamination and a discussion of methods for addressing the problem. *Clin Microbiol Rev* 2020;33:e00009–19.
- Woodford EC, Dhudasia MB, Puopolo KM, *et al.* Neonatal blood culture inoculant volume: feasibility and challenges. *Pediatr Res* 2021;90:1086–1092.
- Donn SM. Medical liability, risk management, and the quality of health care. *Semin Fetal Neonatal Med* 2005;10:3–9.
- Clinical and Laboratory Standards Institute. *Principles and Procedures for Blood Cultures: Approved Guideline, vol. 46, no. 31.* Wayne, PA: CLSI; 2007. CLSI document M47-A.
- Clinical and Laboratory Standards Institute. *Principles and Procedures for Blood Cultures; Second Edition.* Wayne, PA: CLSI; 2022. CLSI document M47.
- Allen E, Cavallaro A, Keir AK. A quality improvement initiative to reduce blood-culture contamination in the neonatal unit. *Pediatr Qual Saf* 2021;6:e413.
- Peri AM, Harris P, Paterson DL. Culture-independent detection systems for bloodstream infection. *Clin Microbiol Infect* 2022;28:195–201.
- Rubin L, Sanchez P, Siegel J, *et al.* Evaluation and treatment of neonates with suspected late-onset sepsis: a survey of neonatologists’ practices. *Pediatrics* 2002;110:e42.
- Kerur B, Salvador A, Arbeter A, Schutzman DL. Neonatal blood cultures: survey of neonatologists’ practices. *World J Pediatr* 2012;8:260–262.
- Johnson J, Bracken R, Tamma P, Aucott SW, Bearer C, Milstone AM. Trends in chlorhexidine use in US neonatal intensive care units: results from a follow-up national survey. *Infect Control Hosp Epidemiol* 2016;37:1116–1118.
- Muller M, Bryant KA, Espinosa C, *et al.* SHEA neonatal intensive care unit (NICU) white paper series: practical approaches for the prevention of central-line-associated bloodstream infections. *Infect Control Hosp Epidemiol* 2022;4:1–46.
- Lancaster DP, Friedman DF, Chiotos K, *et al.* Blood volume required for detection of low levels and ultralow levels of organisms responsible for neonatal bacteremia by use of Bactec Peds Plus/F, Plus Aerobic/F medium, and the BD Bactec FX System: an in vitro study. *J Clin Microbiol* 2015;53:3609–3613.
- Schelonka RL, Chai MK, Yoder BA, Hensley D, Brockett RM, Ascher DP. Volume of blood required to detect common neonatal pathogens. *J Pediatr* 1996;129:275–279.
- Huber S, Hetzer B, Crazzolaro R, Orth-Höller D. The correct blood volume for paediatric blood cultures: a conundrum? *Clin Microbiol Infect* 2020; 26:168–173.
- Sunararajan S. Ideal blood inoculant volume for neonatal sepsis evaluation: an alternative approach. *Pediatr Res* 2021;90:930–933.
- Institute for Healthcare Improvement website. <https://www.ihl.org/resources/Pages/ImprovementStories/WhatIsaBundle.aspx>. Accessed February 21, 2022.
- Lamy B, Dargere S, Arendrup MC, Parienti J-J, Tattevin P. How optimize use of blood cultures for the diagnosis of bloodstream infections? a state of the art. *Front Microbiol* 2016;7:697.
- Povroznik MD. Initial specimen diversion device utilization mitigates blood-culture contamination across regional community hospital and acute-care facility. *Am J Med Qual* 2022;37:405–412.
- Snapp B, Reyna B. Role of the neonatal nurse practitioner in the community hospital. *Adv Neonatal Care* 2019;19:402–408.

23. Parker LA. Neonatal nurse practitioners in interdisciplinary care of high-risk infants. *NeoReviews* 2010;11:c24–c31.
24. Garcia RA, Spitzer ED, Beaudry J, Beck C, Diblasi R, Gilleeny-Blabac M. Multidisciplinary team review of best practices for collection and handling of blood cultures to determine effective interventions for increasing the yield of true-positive bacteremias, reducing contamination, and eliminating false-positive central line-associated bloodstream infections. *Am J Infect Control* 2015;43:1222–1237.
25. Clinical practice guideline: prevention of blood culture contamination. *J Emerg Nurs* 2018;44:285.e1–285.e24.
26. Singh MP, Balegar VKK, Angiti RR. The practice of blood volume submitted for culture in a neonatal intensive care unit. *Arch Dis Child Fetal Neonatal Ed* 2020;105:600–604.
27. Hamilton LZ, Gillet HE, Smith-Collins A, Davis JW. A sterile collection bundle intervention reduces the recovery of bacteria from neonatal blood culture. *Biomed Hub* 2018;3:1–7.
28. Cantey JB, Baird SD. Ending the culture of culture-negative sepsis in the neonatal ICU. *Pediatrics* 2017;140:e20170044.
29. Schulman J, Benitz WE, Profit J, *et al.* Newborn antibiotic exposures and association with proven bloodstream infection. *Pediatrics* 2019;144:e201191105.
30. Kahn DJ, Perkins BS, Barrette CE, Godin R. Reducing antibiotic use in a level III and two level II neonatal intensive care units targeting prescribing practices for both early- and late-onset sepsis: a quality improvement project. *Pediatr Qual Saf* 2022;7:e555.