



POINT OF VIEW

To reduce the current rates of catheter-related bacteremia after the implementation of the Zero programs: This is the challenge[☆]

Reducir las tasas actuales de bacteriemia relacionada con catéter tras la implantación de los programas Zero: este es el reto

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Current scenario of vascular catheter-related bacteremia

Vascular catheter-related bacteremia (CRB) is associated to increased patient morbidity-mortality and healthcare costs.¹ Different measures have therefore been proposed to try to prevent it.^{2,3} A meta-analysis has shown the application of a series of measures to result in an important decrease in the incidence of CRB in many studies worldwide.⁴

The Bacteremia Zero project implemented in 2009 reduced the incidence of CRB,⁵ comprising bloodstream infection of unknown origin and bacteremia secondary to central venous catheter (CVC) placement, from 4.9 to 2.8 cases per 1000 days of CVC, and from 2.7 to 1.4 per 1000 days of CVC, respectively.⁵ These rates have been maintained over time, possibly with the help of implementation of the Pneumonia Zero,⁶ Resistance Zero⁷ and Urinary tract infection Zero projects.⁸ The measures for the prevention of CRB include adequate hand hygiene, optimum barrier measures, skin disinfection with chlorhexidine, preference for subcla-

vian access, withdrawal of needless catheters, and hygienic handling of catheters.⁶

During these 10 years of implementation of the Bacteremia Zero project, new evidence has emerged referred to certain non-proposed measures such as CVCs impregnated with antimicrobials, dressing impregnated with chlorhexidine, daily chlorhexidine bathing and capping with antiseptic for needle-free connectors. Such measures have been proposed by the clinical practice guides (CPGs) of the Society for Healthcare Epidemiology of America (SHEA) published in 2014,⁹ the UK Department of Health published in 2014,¹⁰ and the Asia Pacific Society of Infection Control (APIC) published in 2016¹¹ (Table 1). The challenge therefore arises to continue reducing the current CRB rates.

Catheters impregnated with antimicrobials

The American,⁹ British¹⁰ and Asian CPGs¹¹ recommend the use of catheters impregnated with antimicrobials if the CRB rates remain high. A meta-analysis published in 2018¹² including 25 randomized clinical trials (RCTs) and 9368 CVCs (4001 standard catheters, 2598 catheters impregnated with chlorhexidine / silver sulfadiazine, 1635 impregnated with silver, and 1134 impregnated with rifampicin and miconazole or minocycline) found CVCs impregnated with chlorhexidine / silver sulfadiazine (odds ratio [OR]=0.64; 95% confidence interval [95%CI]=0.40–0.96) or antibiotics

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Table 1 Recommendations of different scientific societies and new evidence on measures for the prevention of vascular catheter-related bacteremia (CRB) following implementation of the Bacteremia Zero project.

	CVCs impregnated with CSD or RM	Dressings impregnated with chlorhexidine	Daily bathing with chlorhexidine	Capping with antiseptic for needle-free connectors
American guides 2014 ⁹	(I) in patients > 2 months if the CRB rates remain high or in the case of a personal history of recurrent CRB or high risk of sequelae due to CRB	(I) in patients > 2 months. Doubtful benefit if daily chlorhexidine bathing is used	(I) in patients > 2 months. Doubtful benefit if dressing impregnated with chlorhexidine are used	(I)
British guides 2014 ¹⁰	(A) if the CRB rates do not decrease	(B)	Not analyzed	Not analyzed
Asian guides 2016 ¹¹	(IA) if the CRB rates do not decrease	(IB) in patients > 2 months if the CRB rates do not decrease	(IIB)	Not analyzed
Most recent meta-analyses	Meta-analysis published in 2018 ¹²	Meta-analysis published in 2019 ¹⁴	Meta-analysis published in 2019 ¹⁵	Meta-analysis published in 2017 ¹⁶

The level of evidence of each measure for preventing CRB according to each clinical practice guide is indicated in parentheses. CRB: vascular catheter-related bacteremia; CSD: chlorhexidine / silver sulfadiazine; CVC: central venous catheter; RM: rifampicin - minocycline.

(OR=0.3; 95%CI=0.25–0.95) to pose a lesser risk of CRB than standard catheters. However, there were no significant differences between CVCs impregnated with silver and standard catheters (OR=0.77; 95%CI=0.46–1.27).

Limitations for the use of impregnated CVCs comprise possible allergies to the antimicrobial (isolated cases) or infections caused by microorganisms resistant to the drug (in *in vitro* studies and animal models).¹³

Dressings impregnated with chlorhexidine

The American,⁹ British¹⁰ and Asian CPGs¹¹ recommend the use of dressings impregnated with chlorhexidine. A meta-analysis published in 2019¹⁴ including 11 RCTs and 10,796 catheters (CVCs, arterial catheters and tunneled catheters) concluded that impregnated dressings reduce the risk of CRB (OR=0.60; 95%CI=0.42–0.85). However, the studies posed the limitation of including different types of catheters (CVCs, arterial catheters and tunneled catheters) and catheter insertion sites associated with different CRB risks; furthermore, the protective effect for each catheter type and site was not determined.

Dressings impregnated with chlorhexidine also have the inconvenience of possible allergies or infections caused by microorganisms resistant to chlorhexidine. On the other hand, none of the RCTs of the meta-analysis reported the incidence of resistances; only three of the trials reported the incidence of contact dermatitis (approximately 5% and particularly affecting newborn infants with a weight of under 1 kg and less than four months of age); and no systemic reactions to chlorhexidine were reported.

Bathing of the patient with chlorhexidine

Bathing of the patient with chlorhexidine is advised by the American⁹ and Asian CPGs,¹¹ and is not analyzed in the British guides.¹⁰ A meta-analysis published in 2019¹⁵ involving 26 studies (8 RCTs and 18 observational studies) analyzed the effect of daily bathing with chlorhexidine. Eighteen studies used disposable sponges with 2% chlorhexidine, while 8 studies used chlorhexidine solutions (at a concentration of 4% in 5 studies, 2% in two studies, and 0.9% in one study). The incidence of CRB was seen to be lower with daily chlorhexidine bathing than when soap and water were used (incidence rate ratio=0.59; 95%CI=0.52–0.68).

The possible limitations referred to the appearance of allergies or infections caused by microorganisms resistant to chlorhexidine were not analyzed in the studies included in the meta-analysis.

Capping with 70% isopropyl alcohol for needle-free connectors

The American CPGs⁹ recommend this practice, while the British¹⁰ and Asian guides do not analyze its use.¹¹ A meta-analysis published in 2017¹⁶ involving 7 observational studies (with a before-after implementation design) recorded a decrease in CRB (incidence rate ratio=0.59; 95%CI=0.45–0.77) associated to the use of capping with antiseptic versus standard capping – though the appearance of allergies to the antiseptic was not analyzed.

Proposal for reducing the current vascular catheter-related bacteremia rates

As in the case of the Pneumonia Zero project, there are both mandatory measures and optional measures. The four mea-

asures for preventing CRB (impregnated CVCs, impregnated dressings, chlorhexidine bathing and capping with antiseptic) could be incorporated to the Bacteremia Zero initiative as optional measures. The current mandatory measures of the Bacteremia Zero project certainly must be maintained, in view of the outcomes they afford. Each individual center should decide the need to adopt optional measures, the types of measures, and the kind of patients in which they should be applied. Optional measures possibly are indicated in those units that despite adequate compliance with the mandatory measures present CRB rates (encompassing bacteremia of unknown origin and secondary to CVC) of over three episodes per 1000 days of CVC, in accordance with the quality standard proposed by the SEMICYUC in 2017 (https://www.bing.com/search?PC=WCU&FORM=WCU&GDF&q=indicadoresdecalidad2017_semicyuc_spa-1.pdf). We could start by adopting some of the mentioned measures, with a view to securing increased efficiency (I personally would recommend impregnated CVCs, which moreover represent the optional measure with the greatest supporting evidence according to the latest CPGs^{10–12}). Likewise with a view to securing increased efficiency, we could start by applying the chosen optional measure in concrete clinical scenarios: 1) patients at increased risk of CRB (immune depressed subjects, altered skin integrity); 2) vascular accesses posing an increased risk of CRB (jugular vein with tracheostomy or femoral vein)¹⁴; and 3) patients at an increased risk of suffering complications if CRB develops (recent implantation of heart valves or aortic prostheses). Depending on whether or not the quality standard is reached after adopting the optional measure in the selected clinical scenarios, we could decide application of the measure to the rest of the patients, or adopt other optional measures in the selected clinical settings.

Conflicts of interest

The authors declare that they have no conflicts of interest.

References

- [1]. Chaves F, Garnacho-Montero J, Del Pozo JL, Bouza E, Capdevila JA, de Cueto M, et al. Diagnosis and treatment of catheter-related bloodstream infection: Clinical guidelines of the Spanish Society of Infectious Diseases and Clinical Microbiology and (SEIMC) and the Spanish Society of Intensive and Critical Care Medicine and Coronary Units (SEMICYUC). *Med Intensiva*. 2018;42:5–36.
- [2]. Lorente L. Antiseptic measures during the insertion and manipulation of vascular catheters. *Med Intensiva*. 2019;43 Suppl 1:39–43.
- [3]. Fernández Moreno I, Píriz Marabaján M. Antisepsis in the handling of vascular access connections. *Med Intensiva*. 2019;43 Suppl 1:44–7.
- [4]. Ista E, van der Hoven B, Kornelisse RF, van der Starre C, Vos MC, Boersma E, et al. Effectiveness of insertion and maintenance bundles to prevent central-line-associated bloodstream infections in critically ill patients of all ages: a systematic review and meta-analysis. *Lancet Infect Dis*. 2016;16:724–34.
- [5]. Palomar M, Álvarez-Lerma F, Riera A, Díaz MT, Torres F, Agra Y, et al. Bacteremia Zero Working Group. Impact of a national multimodal intervention to prevent catheter-related bloodstream infection in the ICU: the Spanish experience. *Crit Care Med*. 2013;41:2364–72.
- [6]. Álvarez Lerma F, Sánchez García M, Lorente L, Gordo F, Añón JM, Álvarez J, et al. Sociedad Española de Medicina Intensiva; Sociedad Española de Enfermería Intensiva. Guidelines for the prevention of ventilator-associated pneumonia and their implementation. The Spanish Žero-VAPbundle. *Med Intensiva*. 2014;38:226–36.
- [7]. Montero JG, Lerma FÁ, Gallego PR, Martínez MP, Rocha LÁ, Gaité FB, et al. Scientific Expert Committee for Zero Resistance Project. Combatting resistance in intensive care: the multimodal approach of the Spanish ICU Žero Resistance program. *Crit Care*. 2015;19:114.
- [8]. Álvarez Lerma F, Olaechea Astigarraga P, Nuvials X, Gimeno R, Catalán M, Gracia Arnillas MP, et al. Grupo de Estudio ENVIN-HELICS. Is a project needed to prevent urinary tract infection in patients admitted to spanish ICUs? *Med Intensiva*. 2019;43:63–72.
- [9]. Marschall J, Mermel LA, Fakih M, Hadaway L, Kallen A, O’Grady NP, et al. Society for Healthcare Epidemiology of America. Strategies to prevent central line-associated bloodstream infections in acute care hospitals: 2014 update. *Infect Control Hosp Epidemiol*. 2014;35:753–71.
- [10]. Loveday HP, Wilson JA, Pratt RJ, Golsorkhi M, Tingle A, Bak A, et al. UK Department of Health. Epic3: national evidence-based guidelines for preventing healthcare-associated infections in NHS hospitals in England. *J Hosp Infect*. 2014;86 Suppl 1: S1–70.
- [11]. Ling ML, Apisarnthanarak A, Jaggi N, Harrington G, Morikane K, Thu le TA, et al. APSIC guide for prevention of Central Line Associated Bloodstream Infections (CLABSI). *Antimicrob Resist Infect Control*. 2016;5:16.
- [12]. Wang H, Tong H, Liu H, Wang Y, Wang R, Gao H, et al. Effectiveness of antimicrobial-coated central venous catheters for preventing catheter-related blood-stream infections with the implementation of bundles: a systematic review and network meta-analysis. *Ann Intensive Care*. 2018;8:71.
- [13]. Lorente L, Lecuona M, Jiménez A, Raja L, Cabrera J, Gonzalez O, et al. Chlorhexidine-silver sulfadiazine- or rifampicin-miconazole-impregnated venous catheters decrease the risk of catheter-related bloodstream infection similarly. *Am J Infect Control*. 2016;44:50–3.
- [14]. Wei L, Li Y, Li X, Bian L, Wen Z, Li M. Chlorhexidine-impregnated dressing for the prophylaxis of central venous catheter-related complications: a systematic review and meta-analysis. *BMC Infect Dis*. 2019;19:429.
- [15]. Musuza JS, Guru PK, O’Horo JC, Bongiorno CM, Korobkin MA, Gangnon RE, Safdar N. The impact of chlorhexidine bathing on hospital-acquired bloodstream infections: a systematic review and meta-analysis. *BMC Infect Dis*. 2019;19:416.
- [16]. Voor In ’t Holt AF, Helder OK, Vos MC, Schaffthuisen L, Sülz S, van den Hoogen A, et al. Antiseptic barrier cap effective in reducing central line-associated bloodstream infections: A systematic review and meta-analysis. *Int J Nurs Stud*. 2017;69: 34–40.