Review Article

Blood Donor Arm Cleaning – Another step towards preventing the contamination of blood products for transfusion purposes

ABSTRACT

Blood Establishments strive to provide blood and blood derivatives that are safe for transfusion. A recurrent concern is providing blood components that are free from bacterial contamination.—In fact, Beacterial and fungal contamination of blood products are—is nowadays a major apprehension when it comes to transfusion adverse events. Over the years, several disinfection products and protocols have been devised to mitigate the risks of such contaminants, nonetheless, sepsis is still the leading cause of transfusion reaction fatalities. This fact raises the question whether disinfection on its own is sufficient for preventing such outcomes.

Key words: Contamination, blood products, blood donor, disinfection, cleaning.

1. INTRODUCTION

Blood establishments developed several preventive measures to avert this unwanted entry of bacteria into the blood stream from happening. A very important measure in preventing this is, how the skin is disinfected before blood collection(1). In fact, linadequate or ineffective disinfection is one of the main sources of bacterial contamination of blood products(2). Over the years, several disinfection products and protocols have been devised, however, sepsis is still the leading cause of transfusion reaction fatalities(3). Several A number of factors could affect the efficacy of disinfection which increases the likelihood of contaminated blood products. It is being proposed that cleaning the skin prior to before blood donation increases the effectiveness of the disinfection process(4).

2. CAUSES OF FAILED DISINFECTION

2.1. Bacterial load on the skin

The skin harbours a diverse microbiome, some of which have a symbiotic relationship with the skin itself(5). It has been shown that the higher the bacterial load on the skin, the less effective skin disinfection is(6). The greater the bacterial load on the skin, the more contact time <u>is</u> required to eradicate most of the bacteria. Additionally, clumped bacterial cells on the skin are more difficult to eradicate <u>rather</u> than single bacterial cells(7). Micro-organisms can also be found in the deeper layers of the skin and on skin appendages such as sweat glands, sebaceous glands and hair follicles(8). Resident bacteria cannot be eradicated by disinfection and run the risk of contaminating blood products after venepuncture(9).

2.2. Dimpled phlebotomy sites

Dimpled phlebotomy sites tend to make skin disinfection less effective. Repeated blood donations result in scarring of the skin and the formation of dimples at the venepuncture sites. Research has shown that dimpled and scarred phlebotomy sites make disinfection less effective and lead to a higher risk of bacterial contamination of blood products(10).

2.3. Resistance to disinfection

Bacterial resistance to disinfectants can be innate or acquired(11). Innate resistance of bacteria to the disinfection processis due to structures that act as barriers. For example, bacterial spores are resistant due to the spore coat and cortex. <u>Gram Gram-negative</u> bacteria are surrounded by an outer membrane and mycobacteria have an outer waxy cell wall which makes the uptake of disinfectant more difficult(7).

Disinfectants are useful in reducing the number of skin flora but over time bacteria are known to develop a degree of resistance to the product(12). Resistance can be acquired by a mutation or by

amplification of an endogenous chromosomal gene(11). In fact, tThe minimum inhibitory concentration (MIC) and minimum bactericidal concentration (MBC) tend to be higher in places where disinfectants are routinely used, such as blood establishments and operating theatres (13). One of the found bacteria on the antecubital fossa is Staphylococcusaureusand it produces protein complexes known as adherens that will help it adhere to the skin(14). One of the reasons why Staphylococcus aureus has a higher MIC is due to mutations in the smr and norA, qacA, and qacB genes(11). Since the skin contains various bacterial species and strains, those that are resistant to a disinfectant will survive the process of disinfection, thus causing the risk of contaminating blood products(13). The World Health Organization recommends chlorhexidine (CHX) to be used as a skin disinfectant prior to venepuncture and it is one of the most widely used disinfectants in healthcare settings(15). When compared to other disinfection products, CHX offers the greatest residual activity which means its antimicrobial effect is maintained even after drying completely(16). Although 0.5% CHX is effective at eradicating a vast number of Gram-positive and Gram-negative bacteria, it is responsible for the low bacterial reduction rates and for the increasing microbial persistence(17).

2.4. Biofilm formation

Bacteria have the ability to form a dense layer and attach themselves firmly to the skin surface, thus they are difficult to remove by the disinfection process(7). The MIC of CHX required to inactivate bacteria in biofilms is 64 times higher than the MIC required to inactive free-floating bacteria. Once formed, bacteria produce a matrix which will further protect them from the action of disinfectants. An example of bacteria which can form biofilms is *Staphylococcus epidermidis* and it is one of the main contaminants of platelet units(18).

2.5. Low concentrations of disinfectants

When a disinfectant below the MBC is used, it is likely that it will will likely be ineffective when it comes to eliminating bacteria. Additionally, disinfectants at low concentrations may cause bacteria to exhibit the hormetic effect. This means that bacteria are stimulated to grow even more at low concentrations of a disinfectant(19).

3. THE CLEANING TECHNIQUE

The cleaning technique is divided into 3 stages: preparation, washing and drying(20). The first stage involves wetting the skin with water prior tebefore the application of soap. The second stage involves the application of liquid soap to the skin and rubbing for a few seconds to create a lather. This is followed by scrubbing the skin for at least 20 seconds(21). Friction is created through the action of scrubbing the skin with soap and water on the skin. This helps to lift off bacteria from the skin(4). Prolonged washing time i.e., more than 30 seconds, loosens resident bacteria from the skin but does not remove them. Therefore, rinsing with clean and running water is essential to remove the loosened resident bacteria by dilution(22). Drying ef—the skin with a clean towel is an important last stage but many tend to skip it(21). Drying the skin after cleaning makes it more difficult to acquire bacteria on the skin(23).

The uppermost layer and deeper layers of the skin harbour many resident and transient bacteria (24). Apart from causing diseases when transmitted from one person to another, they run the risk of contaminating blood products if proper cleaning of the antecubital fossa is not carried out(25). Cleaning the skin with soap and water is more effective than washing the skin with just water or no hand washing at all(26). Cleaning the skin with just water is ineffective because bacteria is are mixed with the oil on the skin. Oil and water are immiscible; therefore, soap is needed to be able to remove both oil and bacteria from the skin(4).

Soap is a detergent-based product that contains sodium or potassium hydroxide and esterified fatty acids. It also contains surfactant molecules which lower the surface tension between water and oil on the skin, making it easier for surfactant molecules to reach the skin surface and exert their effect. Surfactant molecules in soap can exist in two forms: as monomers when they are low in concentration, but they can also come together to form micelles when they are at a high concentration, also known as the critical micelle concentration. A surfactant monomer is amphipathic as it consists of a hydrophilic head and a lipophilic tail. When they are in the micelle shape, they come together in a way that the lipophilic tails point inwards, and the hydrophilic heads point outwards(27). Some bacterial cell membranes have a similar structure to that of surfactant monomers.

The cell membrane is also amphipathic as it consists of a phospholipid bilayer with an outer hydrophilic region and a middle lipophilic region. Like a micelle, the phospholipid hydrophilic heads in the bacterial cell membrane point outwards and the lipophilic tails point towards the centre. The presence of oil on the skin makes it easier for bacteria to adhere to the skin(28).

3.1. How cleaning works

When surfactant monomers come in contact with the skin through the act of cleaning, they insert themselves in the outer half of the bacterial cell membrane and arrange themselves in a similar waysimilarly arrange themselves. Surfactant monomers put mechanical stress on the bacterial cell membrane and bends it. Consequently, pores in the bacterial cell membrane form which leads to the breakdown of the bacterial cell. Hybrid micelles form containing both surfactant monomers and phospholipids from the bacterial cell membrane (27).

3.2. Why is cleaning necessary?

The presence of dirt on the skin increases the bacterial load, and as mentioned earlier, the bacterial load on the skin affects the effectiveness of disinfection. The higher the bacterial load on the skin, the less effective disinfection will be(6). Dirt on the skin can act as a barrier which prevents the disinfection product from reaching the surface of the skin and to-effectively perform it (21). Since cleaning reduces the initial bacterial load on the skin, it helps to increase the effectiveness of the disinfection process, which will remove the remaining bacteria(29). Cleaning also makes the disinfection process more effective by perforating the outer membranes of bacteria which will be furtherbroken-down during disinfection(4).

Bacteria on the skin can proliferate to colonise the skin surface and produce a biofilm(30). To increase their adherence to the skin and their proliferating capability, bacteria produce an extracellular matrix containing DNA, lipids, proteins and polysaccharides(31). Surfactant molecules disrupt the cohesive forces in the extracellular matrix and the bonds between bacteria and the skin, detaching them from the skin(30). Micelles can also trap the oil on the skin, which makes bacteria more likely to adhere to the skin. During rinsing, hybrid micelles are washed off; therefore, removing bacteria that were once present on the skin(28).

4. Conclusion

The presence of bacteria on the skin is one of the main causes of bacterial contamination of blood products. One of the reasons is inadequate skin disinfection before venepuncture. Bacteria may remain present on the skin even if skin disinfection is carried out properly. Cleaning of the skin also reduces the bacterial load through the action of detergent monomers. Therefore, to further reduce the bacterial load on the skin, cleaning of the antecubital fossa should be carried out, making skin disinfection more effective.

5. CONSENT

Not Applicable.

6. REFERENCES

- Suddock JT, Crookston KP. Transfusion Reactions. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2023 [cited 2023 Oct 12]. Available from: http://www.ncbi.nlm.nih.gov/books/NBK482202/
- Wagner SJ, Leiby DA, Roback JD. Existing and Emerging Blood-Borne Pathogens: Impact on the Safety of Blood Transfusion for the Hematology/Oncology Patient. Hematol Oncol Clin North Am. 2019 Oct;33(5):739–48.
- Vamvakas EC, Blajchman MA. Transfusion-related mortality: the ongoing risks of allogeneic blood transfusion and the available strategies for their prevention. Blood. 2009 Apr 9:113(15):3406–17.
- 4. Centers for Disease Control and Prevention. Show Me the Science How to Wash Your Hands [Internet]. 2021 [cited 2023 Oct 12]. Available from: https://www.cdc.gov/handwashing/show-methe-science-handwashing.html

- 5. Swaney MH, Kalan LR. Living in Your Skin: Microbes, Molecules, and Mechanisms. Infect Immun. 2021 Mar 17;89(4):e00695-20.
- 6. Arghittu A, Dettori M, Deriu GM, Soddu S, Manca PC, Carboni AA, et al. Controlling Infectious Risk in Transfusion: Assessing the Effectiveness of Skin Disinfection in Blood Donors. Healthcare (Basel). 2022 May 5;10(5):845.
- Centers for Disease Control and Prevention. Factors Affecting the Efficacy of Disinfection and Sterilization [Internet]. 2019 [cited 2023 Oct 12]. Available from: https://www.cdc.gov/infectioncontrol/guidelines/disinfection/efficacy.html
- 8. Callewaert C, Ravard Helffer K, Lebaron P. Skin Microbiome and its Interplay with the Environment. Am J Clin Dermatol. 2020 Sep;21(Suppl 1):4–11.
- 9. Pratt R, Hoffman P, Robb F. The need for skin preparation prior to injection: point counterpoint. British Journal of Infection Control. 2005 Aug 1;6(4):18–20.
- 10. McDonald CP. Bacterial risk reduction by improved donor arm disinfection, diversion and bacterial screening. Transfus Med. 2006 Dec;16(6):381–96.
- 11. Rozman U, Pušnik M, Kmetec S, Duh D, Šostar Turk S. Reduced Susceptibility and Increased Resistance of Bacteria against Disinfectants: A Systematic Review. Microorganisms. 2021 Dec 10;9(12):2550.
- Lu J, Guo J. Disinfection spreads antimicrobial resistance. Science. 2021 Jan 29;371(6528):474–474.
- 13. van Dijk HFG, Verbrugh HA. Resisting disinfectants. Commun Med. 2022 Jan 11;2(1):1-5.
- 14. Byrd AL, Belkaid Y, Segre JA. The human skin microbiome. Nat Rev Microbiol. 2018 Mar;16(3):143–55.
- 15. World Health Organization. Venepuncture for blood donaton. In: WHO Guidelines on Drawing Blood: Best Practices in Phlebotomy [Internet]. World Health Organization; 2010 [cited 2023 Nov 18]. Available from: https://www.ncbi.nlm.nih.gov/books/NBK138671/
- 16. Lim KS, Kam PCA. Chlorhexidine--pharmacology and clinical applications. Anaesth Intensive Care. 2008 Jul;36(4):502–12.
- 17. Celere MS, Ferreira O, Ubiali EMA, Julião FC, Fernandes AFT, de Andrade D, et al. Antimicrobial activity of two techniques for arm skin disinfection of blood donors in Brazil. Transfus Med. 2012 Apr;22(2):116–21.
- 18. Taha M, Kalab M, Yi QL, Landry C, Greco-Stewart V, Brassinga AK, et al. Biofilm-forming skin microflora bacteria are resistant to the bactericidal action of disinfectants used during blood donation. Transfusion. 2014 Nov;54(11):2974–82.
- Morales-Fernández L, Fernández-Crehuet M, Espigares M, Moreno E, Espigares E. Study of the hormetic effect of disinfectants chlorhexidine, povidone iodine and benzalkonium chloride. Eur J Clin Microbiol Infect Dis. 2014 Jan;33(1):103–9.
- Seal K, Cimon K, Argáez C. Hand Antisepsis Procedures: A Review of Guidelines [Internet]. Ottawa (ON): Canadian Agency for Drugs and Technologies in Health; 2017 [cited 2023 Oct 12]. (CADTH Rapid Response Reports). Available from: http://www.ncbi.nlm.nih.gov/books/NBK470488/
- 21. Centers for Disease Control and Prevention. When and How to Wash Your Hands [Internet]. 2023 [cited 2023 Oct 12]. Available from: https://www.cdc.gov/handwashing/when-how-handwashing.html

- 22. Michaels B, Gangar V, Schultz A, Arenas M, Curiale M, Ayers T, et al. Water temperature as a factor in handwashing efficacy. Food Service Technology. 2002;2(3):139–49.
- 23. World Health Organization. Normal bacterial flora on hands. In: WHO Guidelines on Hand Hygiene in Health Care: First Global Patient Safety Challenge Clean Care Is Safer Care [Internet]. World Health Organization; 2009 [cited 2023 Nov 18]. Available from: https://www.ncbi.nlm.nih.gov/books/NBK144001/
- 24. World Health Organization. Review of preparations used for hand hygiene. In: WHO Guidelines on Hand Hygiene in Health Care: First Global Patient Safety Challenge Clean Care Is Safer Care [Internet]. World Health Organization; 2009 [cited 2023 Oct 12]. Available from: https://www.ncbi.nlm.nih.gov/books/NBK144041/
- 25. Dubey A, Sonker A, Chaudhary R. Snake in the grass: A case report of transfusion reactions due to contaminated donor arm disinfectant. Asian J Transfus Sci. 2017;11(1):50–2.
- 26. Burton M, Cobb E, Donachie P, Judah G, Curtis V, Schmidt WP. The effect of handwashing with water or soap on bacterial contamination of hands. Int J Environ Res Public Health. 2011 Jan;8(1):97–104.
- 27. Woods Z. LifeCanvas Technologies. 2020 [cited 2023 Nov 17]. How do detergents dissolve lipid membranes? Available from: https://lifecanvastech.com/how-do-detergents-dissolve-lipid-membranes/
- 28. National Institute of Health. National Institutes of Health (NIH). 2009 [cited 2023 Nov 17]. Study Finds Unexpected Bacterial Diversity on Human Skin. Available from: https://www.nih.gov/news-events/news-releases/study-finds-unexpected-bacterial-diversity-human-skin
- 29. U.S. Department of Agriculture. Clean THEN Sanitize: A One-Two Punch to Stop Foodborne Illness in the Kitchen [Internet]. 2019 [cited 2023 Nov 17]. Available from: https://www.usda.gov/media/blog/2019/08/27/clean-then-sanitize-one-two-punch-stop-foodborne-illness-kitchen
- 30. Falk NA. Surfactants as Antimicrobials: A Brief Overview of Microbial Interfacial Chemistry and Surfactant Antimicrobial Activity. J Surfactants Deterg. 2019 Sep;22(5):1119–27.
- 31. Brandwein M, Steinberg D, Meshner S. Microbial biofilms and the human skin microbiome. NPJ Biofilms Microbiomes. 2016;2:3.