"Risk Factors for Catheter Related Bloodstream Infection in Patients at a Hem dialysis Unit and Molecular Characterization of MDR Strains of Staphylococcal Species: A Prospective Study"

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Abstract:

Background: Catheter-related bloodstream infections are a major complication of central venous catheter use. This study examines risk factors associated with catheter-related bloodstream infections and its Molecular Characterization of MDR S.aureus strains.

Aim and Objectives: This study evaluates the risk factors, antimicrobial resistance and molecular characterization of Staphylococcal species for the development of bloodstream infections in patients undergoing hemo dialysis.

Methods: In this Prospective study, single-center study, chronic renal disease patients on hemo dialysis who presented with positive blood cultures during the study were included during a 1year study period. Quantitative blood cultures (QBC) and catheter tip cultures were performed for the diagnoses according to the standard procedures. Bio film production in catheters was detected by 'tissue culture plate' (TCP) method and molecular characterization of the gram positive resistant staphylococcal isolates was done. Antimicrobial susceptibility testing was done by Kirby-Bauer disc diffusion method as per CLSI guidelines and E-test.

Results: Catheter-related blood-stream infections (CRBSI) in Rama Medical College Hospital and Research Centre, Mandhana, Kanpur were 4.74 per 1,000 catheter days. Of the 3 isolates that caused CRBSI, 2 were MRCONS, 1 was MRSA. Of the 10 Staphylococcus spp, isolated 80% were bio film producers. Of the 2 S. aureus isolates 1 isolate(50%)harbor the mecA gene and identified as methicillin-resistant S. aureus (MRSA) and the remaining 1 (50%) isolates were methicillin-susceptible (MSSA.). All the Staphylococcal isolates including MDR strains were 100% sensitive to Vancomycin and Linezolid. The AO staining was more sensitive and Gram staining of catheters showed higher specificity. Conclusion: Antimicrobial resistance was significantly higher in bio film producing Staphylococcal species in this study. Infection prevention measures for bloodstream infections related to central venous catheter use should be intensified, as well as judicious use of this route for vascular access for hem dialysis.

Key words: CRBSI, MDR, MRSA, MRCONS, MSSA QBC, TCP, E-test

Introduction

Central venous catheters (CVSs) are indispensable in modern-day medical practice, particularly in intensive care units. Although such catheters provide necessary access, their use puts patients at risk for local and systemic infectious complications, including local site infection, catheter-related blood-stream infections (CRBSI), septic thrombophlebitis, endocarditic, and other metastatic infections resulting in increased morbidity and mortality [1,2,3] in patients on renal replacement therapy. (e.g., lung abscess, brain abscess, osteomyelitis, and endophthalmitis) [4].

Gram-positive bacteria are most frequently isolated in blood cultures of hemodialysis patients. The major virulence factor for these organisms is their ability to form bio film, which also confers resistance to antimicrobial treatment.

Material and Methods

To determine the chronic renal disease patients on hemodialysis who presented with positive blood cultures. We conducted a Prospective study for a period of 1 year from November 2019 to December 2020 at Rama Medical College Hospital and Research Centre, Mandhana, Kanpur. Quantitative blood cultures (QBC) and catheter tip cultures were performed for the diagnoses according to standard procedures. Bio film production in catheters was detected by 'tissue culture plate' (TCP) method and molecular characterization of the gram positive resistant staphylococcal isolates were done using the Qiagen DNA extraction kit. Logistic regression was used for statistical analysis. Antimicrobial susceptibility testing was done by Kirby-Bauer disc diffusion method according to the CLSI guidelines and E-test.

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Figure 1: Patient inserted with CVSs



Figure 2: Catheter Tip

Results

From the above, we concluded our results stating that the catheter-related blood-stream infections (CRBSI) in Rama Medical College Hospital and Research Centre, Mandhana, Kanpur was 4.74 per 1,000 catheter days.

Of the 3 isolates that caused CRBSI, 2 were *MRCONS*, 1 was *MRSA*. Of the 10 Staphylococcus *spp*, isolated 80% were bio film producers. Of the 2 *S. aureus* isolates 1 isolate(50%) harbor the mecA gene and identified as methicillin-resistant S. aureus (MRSA) and the remaining 1 (50%) isolates were methicillin-susceptible (MSSA.) Femoral vein catheters insertion site, was found to be statistically significant as risk factors for CRBSI. All the Staphylococcal isolates including MDR strains were 100% sensitive to Vancomycin and Linezolid.. The AO staining was more sensitive and Gram staining of catheters showed higher specificity.



Figure 3: BancTec



Figure 4: MSSA by E -test strip and AST method



Figure 5: M R cons by E-test strip



Figure 6: MR cons by AST



Figure 7: MRSA by E-test strip



Figure 8: MRSA by AST



Figure 9: Bio film formation



Figure 10: DNA Extraction Kit



Figure 11: DNA Reagents





Figure 12: Isolated DNA from Bacterial culture of *S. aureus*

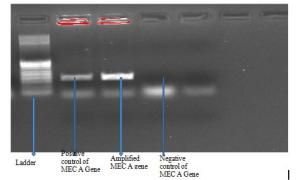


Figure 13: Amplified DNA with PCR for Mec A gene of S. aureus in this fig. Lane 1 is DNA ladder; Lane 2 is positive control of Mec A gene; Lane 3 is Amplified MecA gene; Lane 4 is Negative control for MecA gene

Discussion

In the present study Catheter-related blood-stream infections (CRBSI) in Rama Medical College Hospital and Research Centre, Mandhana, Kanpur was 4.74 per 1,000 catheter days which was similar to the study conducted by other author where CVC related bloodstream infection was 10.8 per 1000 (CVC-RBI)[5]. Gram-positive bacteria were isolated with the highest frequency which correlates with the study Fram D et al. [6]. Of the 3 isolates that caused CRBSI, 2 were MRCONS 1 was MRSA. Similarly other study show the highest organisms isolated by D F Fani et al. (2019) [7] of the 10 Staphylococcus spp. isolated 80% were bio film producers, of the 2 S. aureus isolates 1 isolate(50%) harbor the mecA gene and identified as methicillin-resistant S. aureus (MRSA) and the remaining 1 (50%) isolates were methicillin-susceptible (MSSA.) Permanent hemo dialysis catheters are associated with lower dialytic efficacy compared to arterio venous fistulae [8]. Moreover, there is an increased incidence of mechanical dysfunction [9], thrombosis [10] and bacteremia [11]. Related complications, especially infections, have unfavorable effects on morbidity, mortality and costs. Infections are the second most frequent cause of death in dialysis patients and therefore play a major role since they lead to about 1300 hospitalizations per 1000 patient years [12]. All the Staphylococcal isolates including MDR strains were 100% sensitive to Vancomycin and Linezolid. The AO staining was more sensitive and Gram staining of catheters showed higher specificity. Antimicrobial resistance was significantly higher in biofilm producing Staphylococcal species in this study.

Conclusion

Infection prevention measures for bloodstream infections related to central venous catheter use should be intensified, as well as judicious use of this route for vascular access for hemodialysis. Reducing exposure to the hospital environment through admission could contribute to a reduction in bloodstream infections in this population.

References

- S.I. Blot, P. Depuydt, L. Annemans, D. Benoit, E. Hoste, J.J. De Waele, *et al.* Clinical and economic outcomes in critically ill patients with nosocomial catheter-related bloodstream infections. Clin Infect Dis, 2005;41: 1591-1598
- D.G. Maki, D.M. Kluger, C.J. Crnich. The risk of bloodstream infection in adults with different intravascular devices: a systematic review of 200 published prospective studies. Mayo Clin Proc. 2006; 81:1159-1171
- D.K. Warren, W.W. Quadir, C.S. Hollenbeak, A.M. Elward, M.J. Cox, V.J. Fraser. Attributable cost of catheter-associated bloodstream infections among intensive care patients in a nonteaching hospital. Crit Care Med. 2006; 34:2084-2089

- O'Grady NP, Alexander M, Dellinger EP, Gerberding JL, Heard SO, Maki DG, Masur H, McCormick RD, Mermel LA, Pearson ML, Raad II, Randolph A, Weinstein RA Guidelines for the prevention of intravascular catheter-related infections. MMWR Recomm Rep. 2002; 51:1-29.
- FaridaSahli, RazikaFeidjel, RimaLaalaoui. Hemodialysis catheter-related infection: rates, risk factors and pathogens. Journal of Infection and Public Health. 2017; 10(4):403-408.
- Dayana Fram, Meiry Fernanda Pinto Okuno, Mônica Taminato, Vinicius Ponzio, Silvia Regina Manfredi, Cibele Grothe, Angélica Belasco, Ricardo Sesso, and Dulce Barbosa. Risk factors for bloodstream infection in patients at a Brazilian hemodialysis center: a case– control study. BMC Infect Dis. 2015; 15: 158.
- 7. Fani Delistefani et al. Risk factors for catheter-related infections in patients receiving permanent dialysis catheter. BMC Nephrology. 2019; 20:199.
- Ethier J, Mendelssohn DC, Elder SJ, Hasegawa T, Akizawa T, Akiba T, Canaud BJ, Pisoni RL. Vascular access use and outcomes: an international perspective from the dialysis outcomes and practice patterns study. Nephrol Dial Transpl. 2008; 23 (10):3219–26.
- Silva TN, Mendes ML, Abrao JM, Caramori JT, Ponce D. Successful prevention of tunneled central catheter infection by antibiotic lock therapy using cefazolin and gentamicin. Int Urol Nephrol. 2013; 45 (5):1405–13.
- Moss AH, Vasilakis C, Holley JL, Foulks CJ, Pillai K, Mcdowell DE. Use of a silicone dual-lumen catheter with a Dacron cuff as a long-term vascular access for hemodialysis-patients. Am J Kidney Dis. 1990; 16(3):211–5.
- Marr KA, Sexton DJ, Conlon PJ, Corey GR, Schwab SJ, Kirkland KB. Catheter-related bacteremia and outcome of attempted catheter salvage in patients undergoing hemodialysis. Ann Intern Med. 1997; 127(4):275–80.
- 12. D'Agata EMC. Antimicrobial-resistant, gram-positive bacteria among patients undergoing chronic hemodialysis. Clin Infect Dis. 2002; 35(10):1212–8.