

Emerging trends of nosocomial *Citrobacter* species surgical wound infection: concern for infection control

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Abstract

Background

Surgical wound infection is a common problem among patients who undergo operation. Several factors play important role in this infection process including endemic nosocomial infection without proper infection control measures.

Objectives

To study the occurrence of the pathogens in post-operative wound infections, their antibiotic resistance patterns, and comparison with published reports.

Methods

Various specimens obtained from the surgical wound during a period of four months were processed for bacteriological culture in the Department of Microbiology, Kathmandu Medical College, Kathmandu. Antibiotic susceptibility test was performed by Kirby-Bauer disk diffusion test for pathogens isolated. The relevant literatures were searched and compared with the present study.

Results

Among 79 culture positive cultures, *Citrobacter* sp. (n=23) was most frequently isolated from surgical wound infection. Twenty strains were multi-drug resistant. In comparison with other studies, this study highlights the emergence multi-drug resistant *Citrobacter* sp. as a leading cause of surgical wound infection. *E. coli*, *Staphylococcus aureus*, *Acinetobacter* sp., *Klebsiella pneumoniae*, *Pseudomonas aeruginosa*, *Proteus* sp., and *Enterobacter* sp. were also isolated from 19, 13, 10, 8, 2, 2, and 2 cultures, respectively. More than 50% of these pathogens were resistant to most of the β -lactam antibiotics tested and most of them were multi-drug resistant while these pathogens showed variable level of resistance to fluoroquinolones and amino glycosides.

Conclusion

The frequent isolation of multi-drug resistant nosocomial strains of *Citrobacter* sp. in surgical wound infection is a remarkable trend. This pathogen and their resistant genes could be endemic to the institution and can cause difficult-to-treat infection if infection control committee is not revitalized and infection control strategies are not implemented.

Key words

Citrobacter sp., Multi-drug resistant, surgical wound infection.

Introduction

Numerous patients are admitted in a hospital for surgical procedures. The incision site is often complicated by nosocomial wound infection which gives psychological and emotional stress to the patient and to the entire family, significantly delays the hospital discharge, increases the financial burden and raises the morbidity and mortality¹. The surgical wound infection is common in Nepal and worldwide. The rate of surgical wound infection in Tribhuvan University Teaching Hospital (TUTH), Nepal and the post-operative cesarean wound infection in Patan Hospital was 4% and 2.76%, respectively^{2,3}. Nosocomial wound infection is also a common problem in other hospitals of Nepal⁴.

Among the pathogens causing nosocomial wound infection coagulase positive *Staphylococcus aureus* is common and the treatment is often complicated by the isolation of Methicillin-resistant *S. aureus* (MRSA) and Vancomycin Resistant *S. aureus* (VRSA)^{2,3,5-7}. Vancomycin resistance significantly raises the hospital stay to 18.1 days and cost as high as \$ 27,190⁸. Gram negative pathogens like, *Escherichia coli*, *K. pneumoniae*, *Proteus sp*, *Enterobacter sp*, *P. aeruginosa*, *Acinetobacter baumannii*, and *Citrobacter sp.* also

causes nosocomial wound infection². The commonest Gram negative organisms isolated were *P. aeruginosa*, *E. coli*, and *K. pneumoniae* in wound infection^{2,9}. The antibiotic resistance in these pathogens is increasing. The emergence of resistance in these pathogens to third generation cephalosporins, carbapenems, tigecycline, colistin and rifampicin has marked the therapeutic deadlock¹⁰⁻¹². The rates of isolation of multi-drug resistant superbugs are increasing causing havoc in health care settings.

Here, we studied the prevalence of the pathogens in post-operative wound infections, their antibiotic resistance patterns and compared the trends of pathogen occurrence with published reports.

Methods

This laboratory research was conducted in Department of Microbiology, Kathmandu Medical College in collaboration with Department of Dermatology, Kathmandu University School of Medical Science, Dhulikhel, Nepal.

Bacterial isolates: The bacterial isolates that were isolated in the Department of Microbiology from pus, wound swab, and debrided tissues cultures from post-operative patients admitted in Kathmandu Medical College Teaching Hospital during the period of January

to March 2010 were studied. The pathogens were identified using standard phenotypic methods¹³.

Antibiotic susceptibility tests: Disk Diffusion Test was performed to assess the in-vitro activity of different antibiotics to the isolated pathogens. All of the antibiotic disks ($\mu\text{g}/\text{disk}$) were purchased from Hi-Media Laboratories Pvt. Ltd., India except Cefoxitin and Cefazolin which were purchased from Oxoid, UK. The resistant, intermittent, and susceptible zone size was interpreted based on the guidelines of manufacturer which is based on CLSI guidelines. The intermittent zone of resistance was interpreted as resistant. Multi-drug resistant (MDR) isolates were defined as the isolates that were resistant to two or more than two antibiotics tested.

Comparison of pathogens isolated (with published reports): Literature search was performed using the google search engine. The key words used were, surgical wound infection, pathogens, and antibiotic resistance. Three main relevant literatures^{2,9,14} were identified and the data from these studies were compared with the preset study.

Results

Among 79 pathogens isolated from different specimen sources within four months period, *Citrobacter sp.* was recovered from 23 cultures (Table 1). Similarly, *E. coli*, *S. aureus*, *Acinetobacter sp.*, *K. pneumoniae*, *P. aeruginosa*, *Proteus sp.*, and *Enterobacter sp.* were isolated from 19, 13, 10, 8, 2, 2, and 2 cultures, respectively.

More than 50% of the isolates isolated were resistant to most of the β -lactam antibiotics tested. Strains of *Citrobacter sp.* (n=5), *E. coli* (n=3), and *Acinetobacter sp.* (n=5) were tested for β -lactam and β -lactam inhibitor combination (sulphaperazone+sulbactam) and all isolates of *Citrobacter sp.* and *E. coli* were susceptible while few strains (n=2) of *Acinetobacter sp.* were resistant to this combination. Imipenem resistance was noted for one *E. coli* isolate. Fluroquinolones (Ofloxacin, Ciprofloxacin, and Norfloxacin) were insensitive to most strains of *E. coli*, *Acinetobacter sp.*, and *P. aeruginosa* while they were sensitive to other pathogens. Most pathogens were susceptible to the amino glycosides tested.

Among 23 isolates of *Citrobacter sp.* 20 showed MDR phenotype. Similarly, MDR phenotype of *E. coli*, *S. aureus*, *Acinetobacter sp.*, *K. pneumoniae*, *P. aeruginosa*, *Proteus sp.*, and *Enterobacter sp.* noted were 17, 6, 9, 6, 2, 2, and 2 in number, respectively.

Among seven most frequently recovered pathogens from wound infection, *S. aureus* was the most frequently

isolated in three published literatures (Table 2). However, this study identified *Citrobacter* sp. as a commonest pathogen isolated. *Citrobacter* sp. although less common, they were also isolated among top seven pathogens in tow of those studies. *E. coli* was the second most common in this study and two other studies. The frequency of distribution of other pathogens varied among different studies.

Discussion

Nosocomial surgical wound infection in patients attending to hospital is increasing and multidrug resistant pathogens are being isolated. *S. aureus*, *E. coli*, are *K. pneumoniae* were common pathogen to cause wound infection². This study highlighted the emergence of MDR *Citrobacter* sp. as a leading cause of wound

Table 1: Pathogens isolated from surgical wound infection and their antibiotic resistance patterns

Pathogens (n)	Resistant isolates (no. of isolates tested)																				
	A	AC	PC	CX	CI	KZ	CP	SC	TB	CB	CA	VA	FOX	OF	CF	NX	G	AK	CO	T	I
<i>Citrobacter</i> sp. (23)	15(17)	--	--	--	13(20)	--	17(20)	0(5)	3(4)	--	--	--	--	5(21)	2(22)	4(22)	9(20)	5(18)	12(20)	--	0(1)
<i>E. coli</i> (19)	13(15)	1(1)	2(3)	--	10(19)	--	13(16)	0(3)	1(2)	3(3)	3(3)	--	--	8(19)	9(18)	9(17)	5(14)	3(19)	6(14)	--	1(2)
<i>S. aureus</i> (13)	--	8(12)	--	3(4)	--	0(5)	--	--	--	--	--	0(8)	0(2)	--	3(13)	--	1(7)	0(12)	3(6)	3(11)	0(2)
<i>Acinetobacter</i> sp. (10)	3(4)	--	3(4)	--	3(8)	--	3(3)	2(5)	5(5)	--	--	--	--	6(7)	6(10)	6(6)	4(4)	7(10)	1(5)	0(2)	--
<i>K. pneumoniae</i> (8)	7(8)	--	--	--	3(7)	--	5(7)	--	--	--	--	--	--	1(7)	2(8)	2(8)	1(7)	1(8)	4(8)	--	--
<i>P. aeruginosa</i> (2)	2(2)	--	--	--	2(2)	--	2(2)	--	0(1)	0(1)	--	--	--	2(2)	2(2)	2(2)	1(2)	1(2)	2(2)	--	--
<i>Proteus</i> sp. (2)	2(2)	--	--	--	2(2)	--	2(2)	--	--	--	--	--	--	0(2)	0(2)	0(2)	1(2)	2(2)	1(1)	--	--
<i>Enterobacter</i> sp. (2)	1(1)	--	1(1)	--	0(2)	--	1(1)	--	1(1)	1(1)	--	--	--	0(2)	0(2)	0(1)	0(2)	0(1)	--	--	--

“n”-total no. of isolates recovered; “--”, antibiotics not tested; A, Amoxycillin (10 µg); AC, Amoxy-clavulenic acid (20/10 µg); PC, Piperacillin (100 µg); CX, Cloxacillin (30 µg); CI, Ceftriaxone (30 µg); KZ, Cefazolin (30µg); CP, Cephalexin (30 µg); SC, Suphoperazone+Salbactam (75/10 µg); TB, Tobramycin (10 µg); CB, Carbenicillin (100 µg); CA, Ceftazidime (30 µg); VA, Vancomycin (5 µg); FOX, Cefoxitin (30 µg); OF, Ofloxacin (5 µg); CF, Ciprofloaxcin (5 µg); NX, Norfloaxin (10 µg); G, Gentamicin (30 µg); AK, Amikacin (10 µg); CO, Sulphamethoxazole+Trimethoprim (23.75/1.25 µg); T, Tetracycline (30 µg); I, Imipenem (10 µg)

Table 2: Frequency of pathogens isolated from wound infection in different hospitals and this study

S.N.	TUTH, 2003 (2)	TUTH, 2003 (14)	MHRC, 2007 (9)	This study
1	<i>S. aureus</i>	<i>S. aureus</i>	<i>S. aureus</i>	<i>Citrobacter</i> sp.
2	<i>P. aeruginosa</i>	<i>E. coli</i>	<i>E. coli</i>	<i>E. coli</i>
3	<i>E. coli</i>	Coagulase negative <i>Staphylococcus</i> sp.	<i>P. aeruginosa</i>	<i>S. aureus</i>
4	<i>K. pneumoniae</i>	<i>K. pneumoniae</i>	<i>S. pyogenes</i>	<i>Acinetobacter</i> sp.
5	<i>Acinetobacter</i> sp.	<i>Proteus vulgaris</i>	<i>K. pneumoniae</i>	<i>K. pneumoniae</i>
6	<i>Citrobacter</i> sp.	<i>Proteus mirabilis</i>	Coagulase negative <i>Staphylococcus</i> sp.	<i>P. aeruginosa</i>
7	<i>Proteus vulgaris</i>	<i>Citrobacter</i> sp.	--	<i>Proteus</i> sp.

Number in the parenthesis indicate the reference; MHRC, Medicare Hospital and Research Center.

infection at the point of time studied. *Citrobacter* sp. ranked 6th and 7th frequently isolated pathogen in wound infection^{2,14} while none was isolated among 200 specimens processed for bacteriological culture⁹. Among 197 isolates studied, 2 isolates of *Citrobacter* sp. were isolated and none of them were MDR. Most of the isolates of *Citrobacter* sp. isolated in this study were resistant to β -lactam antibiotics and were susceptible to sulphoperazone-sulbactam, fluoroquinolones and amino glycosides. However *Citrobacter* sp. susceptible to sulphoperazone-sulbactam combination were also seen in patients presented with pulmonary disease¹⁵. Among 23 isolates of *Citrobacter* sp., 20 isolates (86.95) showed MDR phenotype. This study is the first study to highlight the emergence of MDR *Citrobacter* sp. in the hospital in Kathmandu as major etiology of surgical wound infection. *E. coli* was the second most common pathogen recovered from surgical wound infection and other pathogens like, *S. aureus*, *Acinetobacter* sp., *K. pneumoniae* were also common.

Although, *Acinetobacter* sp. and *P. aeruginosa*, were less frequently isolated in wound infection these are notorious pathogens to cause myriad of nosocomial infections. Carbapenem, colistin, tigecycline and rifampin in combination with these antibiotics are used to treat the multi-drug, pan-drug resistant strains of these pathogens however the resistant strains to all of these antibiotics have already emerged^{12,16,17}. Most strains of these pathogens were resistant to β -lactam antibiotics, amino glycosides, and fluoroquinolones but were still susceptible to imipenem. However, one isolates of *E. coli* was resistant to imipenem. Imipenem although not widely used in clinical practice in Nepal, overuse of this antibiotic may select imipenem resistant strains and these strains could increase and spread.

Nosocomial pathogens are mostly normal flora of the skin, present in the hospital environment (water, air cooler), carried on innate objects (stethoscope, IV drips, IV stands, bed linens) and hands of medical staff¹⁵. The breach in the skin due to any interventions (surgery, intravenous line, catheters etc) or inappropriate patient care might help the pathogen to lodge in patient's body and gain access to the site of infection. The pre-existing illness like, metabolic diseases, immunosuppression, malignancy, and malnutrition have been described as predisposing factors for surgical wound infection. An emergency procedure, length of pre-operative hospitalization, antibiotic prophylaxis, duration of operation, wound class, and wound contamination also significantly contributes to the wound infection¹⁸.

Conclusion

This report depicts the reverse trends of pathogen isolation and increasing antibiotic resistance in nosocomial *Citrobacter* sp. isolated in surgical wound infection. This pathogen is ubiquitous in nature (food, soil, water) and a colonizer of human gastrointestinal tract and can cause difficult-to-treat nosocomial wound and other infections if infection control strategies like, disinfection of wards, contact precaution against infectious versus colonized patients and judicious use of antibiotics are practiced.

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