# ORIGINAL ARTICLE

# What should be the antibiotic prescription protocol for burn patients admitted in the department of burns, plastic and reconstructive surgery

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#### Key words

Antibiotic sensitivity; Bacteremia; Blood culture; Tissue culture

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#### Abstract

This is a prospective study with the aim to determine specific patterns of burn wound bacterial colonisation and antimicrobial resistance profiles. There is a high incidence of infections and septicaemia in post-burn patients, which, in turn, are associated with high morbidity and mortality, a fact that compelled us to undertake this study. The study was conducted over a period 11 months, from 1 August 2014 to 30 June 2015, in 50 burn patients admitted in our burn unit. Wound cultures were taken after 72 hours of admission from all the patients, and then, empirical systemic antibiotics were administered. For wound cultures; 1 cubic cm tissue was taken and placed in aerobic and anaerobic culture vials and transported to the microbiology lab under all aseptic precautions as soon as possible. At the time of fever any time after 72 hours of admission, 16 ml of blood was drawn under all aseptic precautions. Both aerobic and anaerobic blood culture vials were filled with 8 ml of blood each and transported to the microbiology lab. The results of culture and sensitivity reports of 50 patients were recorded. The data obtained was analysed using appropriate statistical analytical tests. The most common organism responsible for bacteraemia is Pseudomonas (43%). Most of the strains of organisms isolated were resistant to commonly used antibiotics in the hospital; Pseudomonas was found 100% resistant to a combination of ampicillin + sulbactum, ceftriaxone and was most often sensitive to imipenem, amikacin and vancomycin. Methicillin-resistant Staphylococcus aureus (MRSA) was also found resistant to commonly used antibiotics like ceftriaxone, ampicillin + sulbactum and ceftazidime + calvulanic acid. Linzolid and vancomycin were effective in 83% and 100% cases, respectively. We conclude that similar institution-specific studies should be conducted, and such studies will be helpful in providing useful guidelines for choosing effective empirical therapy that will have a great impact on the prevention of infection and its complications in burn patients because of bacteraemia.

#### Introduction

Most of the burn victims who survive including the initial 24 hours after burns succumb to infection of the burnt area and its complications (1). Various factors responsible are disruption of the skin barrier, a large cutaneous bacterial load, the possibility of the normal bacterial flora becoming opportunistic pathogens and severe depression of the immune system. All these factors contribute towards the sepsis in a burn victim (2). Despite

## **Key Messages**

- Every burn centre should have their own antibiotic prescription protocol
- Antibiotic protocols should be based on individual and periodic burn wound, blood culture and antibiotic sensitivity studies

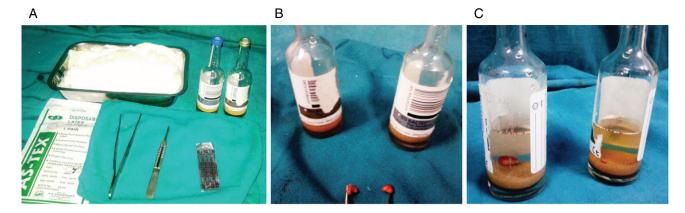


Figure 1 (A) Requirements for obtaining tissue culture, (B) burn wound tissue obtained for culture and (C) tissue culture bottles containing tissues for aerobic and anaerobic cultures.

various advances in infection control measures, management of burn septicaemia still remains a big challenge, and septicemia continues to be the leading cause of death in burn patients (3-5). Approximately 73% of all death within the first 5 days post-burn have been shown to be directly or indirectly caused by septic processes (6). The common bacteria isolated from burn patients include Pseudomonas aeruginosa, Staphylococcus aureus, Klebsillea and various coliform bacilli (7-11). Nosocomial outbreaks of infection in burn units are often because of multi-drug resistant bacteria (12,13). Gram-negative bacteraemias have been associated with a 50% increase in predicted mortality for patients with bacteraemia (14). Systemic antimicrobial treatment must be thoughtfully considered in the care of the burn patient to prevent the emergence of resistant organisms. The burn wound will always be colonised with organisms until wound closure is achieved, and administration of systemic antimicrobials will not eliminate this colonisation but rather promote the emergence of resistant organisms. If antimicrobial therapy is indicated to treat a specific infection, it should be tailored to the specific susceptibility patterns of the organisms as soon as this information is available. Systemic antimicrobials are indicated to treat documented infections, such as pneumonia, bacteremia, wound infection and urinary tract infection (UTI). Empirical antimicrobial therapy to treat fever should be strongly discouraged because burn patients often have fever secondary to the systemic inflammatory response to burn injury. In recent decades, the antimicrobial resistance of bacteria isolated from burn patients has increased (15). It is, therefore, essential for every burn unit to determine its specific pattern of burn wound microbial colonisation, time-related changes in predominant flora and antimicrobial resistance profiles. This would allow early management of septic episodes with proper empirical systemic antibiotics before the results of microbiological cultures become available, thus improving the overall infection-related morbidity and mortality in burn patients.

## Materials and methods

The study was conducted over a period of 11 months, from 1 August 2014 to 30 June 2015, in 50 burn patients admitted

Table 1 Incidence of wound infection and bacteraemia

Blood/tissue culture	Number of cases	Percentage
Positive	23	46
Negative	27	54
Total	50	100

Table 2 Bacteria isolated from tissue and blood culture (n = 23)

Bacteria	Number of cases	Percentage				
Pseudomonas aeruginosa	10	43				
MRSA	6	26				
Staphylococcus aureus	3	13				
Escherichia coli	2	9				
Proteus	1	4				
Klebsiella	1	4				
Total	23	100				

MRSA, methecilline-resistant S. aureus.

to our burn unit. In order to minimise the bias in our observations, many patients were excluded from the study. Patients on immunosuppression therapy, those with known malignancies and those with burn more than 80% of total body surface area (TBSA) have more chances of bacteremia because of their immunocompromised state. Patients who reported to hospital 48 hours after sustaining a burn injury as they could have acquired infection before admission or may have started systemic antibiotics were also excluded from the study. None of the patients in the study group was given prophylactic systemic antibiotics for first 72 hours; however, topical antiseptic silver sulphadiazine was used. Wound cultures were taken 72 hours after admission from all the patients, and then, empirical systemic antibiotics were administered. For wound cultures, 1 cm<sup>3</sup> of tissue was taken and placed in aerobic and anaerobic culture vials and transported to the microbiology lab under all aseptic precautions as soon as possible (Figure 1(A)-(C)). At the onset of fever any time after 72 hours of admission, 16 ml of blood was drawn using all aseptic precautions. Aerobic and anaerobic blood culture vials were filled with 8 ml each and transported to the microbiology lab. The results of the culture

#### Table 3 Sensitivity of bacteria isolated to the antibiotics

										Antibic	otics											
	Am	p+sbm	An	nx + clv	(	Ctx	n	ncn	Ct	z+clv		lzd	V	'nn	g	mc	а	ımk	i	mp	(	Cpf
Bacteria	N	%	N	%	N	%	N	%	N	%	N	%	Ν	%	N	%	N	%	Ν	%	N	%
Pseudomonas P. aeruginosa N = 10	0	0	8	80	0	0	×	×	2	20	×	×	5	50	2	20	6	60	8	80	6	60
MRSA $N = 6$	0	0	1	17	1	17	0	0	0	0	5	83	6	100	4	67	4	67	×	×	3	50
Staphylococcus aureus $N=3$	1	33	0	0	1	33	3	100	2	67	3	100	2	67	2	67	2	67	1	33	2	67
Escherichia coli N=2	0	0	1	50	2	100	×	×	2	100	×	×	1	50	2	100	2	100	2	100	2	100
Proteus $N = 1$	0	0	1	100	×	×	×	×	1	100	×	×	×	×	1	100	1	100	1	100	1	100
Klebsiella $N = 1$	0	0	0	0	0	0	×	×	1	100	×	×	×	×	0	0	1	100	1	100	1	100

amk, amikacin; Amp, ampicilline; Amx, amoxicilline; clv, clavulunate; cpf, ciprofloxacin; ctx, ceftriaxone; Ctz, ceftizidime; gmc, gentamycin; imp, imipenem; lzd, linezolid; mcn, methcilline; sbm, sulbactum; vmn, vancomycin; x, not checked.

 Table 4
 Relationship of bacteraemia with the percent of TBSA burnt and mortality

Bacteremia	Mortality					
4	0					
5	0					
14	2					
	4 5					

TBSA, total body surface area.

The significant correlation of percentage of burn with bacteremia and mortality has been observed with P value = 0.001; Spearman correlation coefficient = 0.51.

and sensitivity reports of 50 patients were recorded. The data obtained was analysed by using a one way analysis of variance (ANOVA).

### Results

The mean age of patients was  $21.91 \pm 11.54$  years. It is clear that infections are a severe problem among burn patients. The wound and blood cultures were positive in 23 patients with similar bacterial growth pattern in our study, which shows that the wound infection is the cause of sepsis in burn patients. The incidence of bacteremia is nearly 46% and more so in more severe burn patients (Table 1). The most common organism responsible for bacteraemia is Pseudomonas (43%) followed by methicillin-resistant S. aureus (MRSA) (26%); the other organisms isolated include S. aureus (13%), Escherichia coli (9%), proteus (4%) and kleibsella (4%) (Table 2). Most of the strains of organisms isolated were resistant to commonly used antibiotics in the hospital. Pseudomonas was found to be 100% resistant to a combination of ampicillin + sulbactum, ceftriaxone and was most often sensitive to imipenem, amikacin and vancomycin. MRSA was also found to be resistant to commonly used antibiotics like ceftriaxone, ampicillin + sulbactum, ceftazidime+calvulanic acid. Linzolid and vancomycin were effective in 83.33% and 100% cases, respectively (Table 3). Table 4 shows that the more the burnt TBSA, the higher the chance of bacteraemia and mortality.

## Discussion

In burn patients, bacteraemia develops as a result of damage to the skin (external barrier) or the respiratory tract and digestive tract (internal barrier) of the body. Bacteraemia is one of the criteria for the diagnosis of sepsis. Sepsis is very lethal for burn patients because it increases the production of inflammatory mediators and cytokines and causes their interaction that predisposes the development of multiple organ failure (MOF). MOF, at present, is the main cause of mortality in burn patients (4,5). Infection, and its complications, remains the leading cause of morbidity and mortality and continues to be the most challenging concern for the burn team. The infection and pathogen responsible for infection differs from hospital to hospital all over the world. In our study, 23 (46%) patients had positive blood cultures during the course of hospital stay, and 27 patients had sterile blood cultures. These observations are in accordance with those of Santucci et al. (16), who found the culture positivity of blood to be 49%. We observed that most of the patients showed culture positivity in the second week. These observations are in accordance with Vostrugina et al. (17) who, in their study, had found a mean time of  $16 \pm 11$ days, and Zorgani et al. (18), who, in their study, had found a majority of positive blood cultures in the first 2 weeks. In the present study, we observed Pseudomonas in 10 (43%) patients as the most common organism isolated from positive blood cultures followed by MRSA in 6 (26%), S. aureus in 3 (13%) E. coli in 2 (9%), kleibsella in 1(4%) and proteus in 1 (4%) cases. Our observations are in accordance with Nagoba et al. (19), who, in their study, had found Pseudomonas in 53.8% of cases as the most common organism isolated in sepsis patients followed by S. aureus in 38.4%. Yildirim et al. (20), who, in their study, had found Pseudomonas in 40.4% to be the most common organism followed by S. aureus 29.3%. Zorgani et al. (18), who, in their study, had found Pseudomonas in 41% of cases followed by S. aureus in 28%. Songa et al. (21) had found pseudomonas in 45.7% as the most commonly isolated organism from burn patients. The sensitivity and resistance pattern of P. aeruginosa observed in our study revealed 100% resistance to ampicillin and ceftriaxone, 80% resistance to gentamycin and ceftazidime + calvullinc and was found to be 80 % sensitive to imipenem and amoxyclave. MRSA was 100% resistant to ampicillin, amoxyclave and ceftazidime and was found to be 67%, 83% and 67% sensitive to gentamycin, linzolid and amikacin, respectively. S. aureus, klebseilla, MRSA and Pseudomonas were resistant to the most commonly used antibiotic in our hospital, for example, ceftriaxone, ampicillin sulbactam. Linzolid was effective against MRSA

and *S. aureus* in 83% and 100% of the cases, respectively. Amikacin was effective against *S. aureus*, Pseudomonas, MRSA in 67%, 60% and 67% of the cases, respectively. Most of the organisms were resistant to commonly used antibiotics. Our observations were in accordance with Yildirim *et al.* (20), Dhar *et al* (22), Khan *et al.* (23) and Vostrugina *et al.*, (17) who also demonstrated similar sensitivity and resistance pattern in their studies. Vostrugina *et al.*, (17) in their study, had observed a higher mortality in bacteraemic patients, which is consistent with our results. A strong correlation between total body surface area burnt and bacteraemia was observed in our study. Vostrugina *et al.* (17) also observed that bacteraemic patients had a larger body surface area burnt.

We believe better outcomes can be achieved in terms of reducing resistance development, which can be achieved through antibiotic and/or antiseptic stewardship. However, we are in favour of the use of topical antibiotic/antiseptic agents based on the previous culture and sensitivity pattern of the burn wards after taking the wound and blood specimen for culture and sensitivity. Once the microbiological agent sensitivity to the particular antibiotic/antiseptic agent has been confirmed, and that particular antimicrobial agent should be prescribed, which is the most scientific way to fight against the microbes and the development of resistance.

### Conclusion

We conclude that similar institution-specific studies should be conducted as such studies will be helpful in providing useful guidelines for choosing effective empirical therapy that will have a great impact on the prevention of infection and its complications in burn patients because of bacteraemia.

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