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## **Original Research Article**

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# Blood Stream Infection in Pediatric Patients of a Tertiary Care Hospital: A Bacteriological and Antimicrobial Profile

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#### ABSTRACT

#### Keywords

Blood stream infection, *Staphylococcus*, *Klebsiella*, Antimicrobial profile.

**Article Info** 

Accepted: 22 February 2017 Available Online: 10 March 2017 Blood stream infections are very common in the pediatric age group and these are one of the common causes of morbidity and mortality in neonates and children. In developing countries, the rate of blood stream infections in children is about 20– 50% and it is estimated that one in five neonates is suffering from and blood infection. The present study was undertaken to determine the etiological agents causing blood stream infection and their antibiotic susceptibility patterns in pediatric patients. A total of 3255 samples with suspected septicaemia were processed aerobically. Culture positivity was 4.4%. *Coagulase negative Staphylococci* (53.65%) was the commonest isolate followed by *K. Pneumoniae* (35.48%). Around 27.27% of MRSA and 12.9% of ESBL producers were isolated in our study. All GPCs were 100% sensitive to vancomycin and teicoplanin and GNBs were 100% sensitive to colistin. Repeated isolation of doubtful pathogens confirms true bacteraemia. Specific antibiotic utilisation strategies like antibiotic restriction, combination therapy and antibiotic escalation and de-escalation may help to decrease or prevent the emergence of resistance.

### Introduction

Blood stream infections are very common in the pediatric age group and these are one of the common causes of morbidity and mortality in neonates and children (Meremkwer *et al.*, 2005). In developing countries, the rate of blood stream infections in children is about 20–50% and it is estimated that one in five neonates is suffering from blood infection (Anuradha *et al.*, 1995; Elster *et al.*, 2005).

Blood stream infection is caused by a variety of Gram-positive as well as Gram-negative bacteria, and sometimes yeasts (Gomaa *et al.*, 2001). The spectrum of causative organisms changes over times and varies from region to region. This is due to the changing pattern of antibiotic use and changes in lifestyle (Shrestha *et al.*, 2008).

Several risk factors have been identified both in the neonates and children, which make them susceptible to infections. The risk factors for neonatal septicemia include premature rupture of membrane, prolonged rupture, prematurity, urinary tract Infection, poor maternal nutrition, low birth weight, birth asphyxia and congenital anomalies (Meremkwer *et al.*, 2005). While in children are serious injury, chronic antibacterial therapy, malnourishment, chronic medical problems, and immuno-suppressants therapy constitute the risk factors for septicemia. Polymicrobial sepsis occurs in high risk patients and is associated with catheters, gastrointestinal diseases, neutropenia and malignancy (Enrione *et al.*, 2007).

Children with septicemia present with fever, difficulty in breathing, tachycardia, malaise, refusal of feeds or lethargy (Nwadioha et al., 2010). They can have serious consequences shock, multiple organ like failure, disseminated intravascular coagulation, etc. Thus, the blood stream infections constitute one of the most serious situations and, as a result, timely detection and identification of blood stream pathogen is important (Mackie and MacCartney). Clinical assessment using a combination of symptoms and signs is a useful guide for the provisional diagnosis of septicemia. But bacteriologic culture to isolate the offending pathogen remains the mainstay of definitive diagnosis of septicemia (Meremkwer et al., 2005).

The choice of empirical antimicrobial requires the knowledge of the epidemiology of common pathogens in the given setting, which constantly changes, necessitating periodic review and their sensitivity profile. Furthermore, this knowledge would help us formulate an antibiotic policy. The present study was designed and implemented to determine the etiology of blood stream infections in pediatric patients and to examine antibiotic susceptibility patterns of the isolated organisms.

### **Materials and Methods**

The study was conducted in the department of Microbiology, Raja Rajeswari Medical

College and Hospital, Bengaluru from January 2014 to September 2015.

Under aseptic precautions, paired blood samples were collected from a total of 3255 patients with suspected blood stream infections. One milliliter (neonates) and 5 ml (children) blood was collected and inoculated into 10 and 50 ml of brain heart infusion broth (1:10 dilution) respectively. The culture bottles were incubated at 37°C aerobically and periodic subcultures were done day 2, day 4 and finally on day 7 onto blood agar and Mac Conkey's agar. Plates were examined for the growth of bacteria. All positive cultures were identified by their characteristic appearance on their respective media, Gram staining reaction and were confirmed by the pattern of biochemical reactions using the standard methods (Mackie and MacCartney).

### Antibiotic susceptibility testing

The antibiotic susceptibility test was done by Kirby-Bauer disk diffusion method. Zone sizes were measured and interpreted according to CLSI standards. Drug resistant strains in primary screening were further processed for the detection of extended spectrum beta lactamases (ESBL) in gram negative bacterial isolates and methicillin resistance in *Staphylococcus aureus* strains.

The isolates were considered as Methicillin Resistant Staphylococcus aureus (MRSA) if the zone of inhibition was less than 21 mm and Methicillin Sensitive Staphylococcus aureus (MSSA) if it was  $\geq 22$  mm. For Coagulase negative Staphylococcus (CONS), if the zone of inhibition was less than 24 mm considered Methicillin resistant as *Coagulase-negative Staphylococcus* (MRCONS) and if it was  $\geq 25$  mm Methicillin sensitive Coagulase negative Staphylococcus (MSCONS) (CLSI, 2014).

ESBL producers were detected by combination disk method using cefotaxime (30  $\mu$ g) and cefotaxime/clavulanate (30/10  $\mu$ g) (Himedia Mumbai, India). An increase of 5 mm in the zone of inhibition in a disk containing clavulanate compared to the drug alone was considered as positive for ESBL producers.

### Statistical analysis

Data was entered into a computerized Excel (Microsoft Excel 2009) spread sheet, and subsequently it was analyzed using SPSS software. Descriptive statistics (means and percentages) were used wherever necessary.

#### **Results and Discussion**

A total of 3255 blood samples from paediatric cases with suspected history of sepsis were received to the department of microbiology during the period from January 2014 to September 2015. Out of which 4.4% (144/3255) were culture positive (paired sample), while 95.6% (3111/3255) were sterile. Table 1 shows age wise distribution of samples, of which children between 1month and 1 year (66%) constituted majority. The most common isolate in this study was gram organisms which accounts for positive 56.94% (82/144)and gram negative organisms were 43.05% (62/144). Among gram positive organisms, the predominant isolate was Coagulase negative Staphylococci (53.65%) followed by Staphylococcus aureus (25.60%). Among gram negative bacilli, the predominant isolate was K. pneumoniae (35.48%) followed by Escherichia coli (24.19%), Pseudomonas aeruginosa (16.12%) and Non fermenting gram negative bacilli (9.67%) (Table 2).

The antibiotic susceptibility pattern of gram positive organisms and gram negative organisms were detailed in the tables 3 and 4 respectively. Septicaemia is a clinical syndrome associated with considerable morbidity and mortality. The timely detection of bacteraemia can have a profound influence on the final clinical outcome. Culture positivity in our study was 4.4%, which is close to study done by Thacker *et al.*, (6.97%). The difference in positivity could be due collection of single sample in patients at our institution. Male preponderance was found in our study which is similar to that of Tallur *et al.*, (2000).

In the present study gram positive and gram negative organisms accounted for 56.94% and 43.05% respectively, which is in concordant with that of Mikulska *et al.*, (2014) (55% and 45%). Among gram positive and gram negative organisms, *Coagulase negative Staphylococcus* (53.65%) and *Klebsiella pneumoniae* (35.48%) respectively were the most common isolates, similarly was observed in Premalatha *et al.*, (2014) (21.21% and 36.36%) study.

Our study showed Coagulase negative *Staphylococci* was the most common blood culture isolate, which are often skin colonizers and appear in blood culture as common contaminants during the time of sample collection. Skin disinfection with 70% isopropyl alcohol at the time of venepuncture can limit this potential contaminant species to a large extent.

In this study, the proportion of MRSA was 27.27%. The occurrence of MRSA is more common because of indiscriminate use of higher antibiotics as an emergency empirical therapy. Vancomycin remains the drug of choice for the Gram positive bacterial isolates. Klebsiella pneumoniae was more imipenem, sensitive meropenem, to ertapenem, gentamicin and amikacin, and resistant to ampicillin, amoxy-clay, cephalosporins and ciprofloxacin, which is similar to study by Premalatha et al., (2014).

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Age	Male (%)	Female (%)	Total
0-1 Month	26	15	41
1 Month-1 Year	36	30	66
1 Year- 18 Years	18	19	37
	80 (55.5)	64 (44.5%)	144

## **Table.1** Age and gender wise distribution of isolates

#### Table.2 Distribution of isolates

Name of the isolate	Total no.	Percentage
Gram positive organisms	82	56.94
Coagulase negative	4.4	52.65
Staphylococcus	44	55.05
Staphylococcus aureus	21	25.60
Streptococcus species	11	13.41
Enterococcus species	06	7.31
Gram negative organisms	62	43.05%
Klebsiella pneumoniae	22	35.48
Escherichia coli	15	24.19
Pseudomonas aeruginosa	10	16.12
Non –fermenting gram negative rods	06	9.67
Salmonella typhi	02	3.22
Klebsiella oxytoca	01	1.61
Citrobacter koseri	01	1.61
Citrobacter freundii	01	1.61
Enterobacter species	01	1.61
Aeromonas species	01	1.61
Proteus mirabilis	01	1.61
Providencia species	01	1.61
Total no. of isolates	144	

#### Table.3 Antibiotic susceptibility pattern of gram positive organisms

Organism	Р	Amp	Am	Ox	Cn	Cx	Cip	G	Sxt	Do	Е	CD	Lz	Va	Те
			с												
MRSA (6)	0	0	0	0	0	0	33.33	100	66.66	83.33	83.33	100	100	100	100
MSSA (15)	33.33	80	100	100	100	100	66.66	100	86.66	86.66	86.66	100	100	100	100
MSCONS	25	25	100	100	100	100	75	79.54	75	90.90	75	79.54	100	100	100
(44)															
Streptococcus	100	100	100	100	100	100	81.81	-	-	-	100	100	100	100	100
<b>spp</b> (11)															
Enterococcus	50	83.33	100	-	0	0	50	50	-	-	-	-	100	100	100
spp (6)															

P-Penicillin, Amp-Ampicillin, Amc-Amoxy-clavunate, Ox-oxacillin, Cn-cephalexin,

Cx-Cefoxitin, Cip- Ciprofloxacin, G- Gentamicin, Sxt- Cotrimoxazole, Do- Doxycycline,

E- Erythromycin, CD- Clindamycin, Lz- Linezolid, Va- Vancomycin, Te- Teicoplanin.

Organism																				
	Amp	Amc	Ŀ	Cn	Cxm	Caz	Ctx	CX	At	Pit	Ert	Imp	Mrp	Ŀ	Ak	T0b	Cip	Cot	$T_g$	CI
K. pneumoniae (22)	0	20	-	22	22	25	40	60	33	57	77	80	77	87	80	100	40	44	100	100
E. coli (15)	16	20	-	-	16	20	16	40	50	80	66	100	83	50	83	100	16	16	100	100
P. aeruginosa (10)	-	-	100	-	-	75	-	-	100	-	-	100	50	100	66	100	75	-	100	100
NFGNB (6)	-	-	0	-	-	80	-	-	-	70	-	-	80	-	-	85	-	-	100	100
S. typhi (2)	100	-	-	-	-	-	100	-	100	100	100	100	100	50	50	-	50	100	100	100
K. oxytoca (1)	0	30	-	20	28	28	40	70	35	60	80	88	76	92	80	100	40	42	100	100
Citrobacter freundii (1)	0	0	0	0	0	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Citrobacter koseri (1)	0	0	0	0	0	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Enterobacter spp. (1)	0	0	0	0	0	100	100	0	100	100	100	100	100	100	100	-	100	100	100	100
Aeromonas spp. (1)	0	-	0	-	-	-	-	-	100	100	100	100	100	100	100	-	100	100	100	100
Proteus mirabilis (1)	100	10 0	100	10 0	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100	100
Providencia spp. (1)	0	0	0	0	0	0	0	0	100	100	100	100	100	100	100	100	100	0	100	100

# **Table.4** Antibiotic susceptibility pattern of gram negative organisms

Amp- Ampicillin, Amc- Amoxy-clavunate, Pi- Piperacillin, Cn - Cephalexin, Cxm- Cefuroxime, Caz- Ceftazidime, Ctx- Cefotaxime, Cx- Cefoxitin, At-Aztreonam, Ptz- Piperacillin- tazobactam, Ert- Ertapenem, Imp- Imipenem, Mrp- Meropenem, G- Gentamicin, Ak- Amikacin, Tob- Tobramycin, Cip- Ciprofloxacin, Sxt- Cotrimoxazole, Tg-Tigecycline, Cl- Colistin

Rate of ESBL producers in our study is 12.9% which agrees with study done by Prabhash *et al.*, (2010) 15.15%. All gram negative isolates were 100% sensitive to colistin and tigecycline, so was observed in Premalatha *et al.*, (2014) study.

In conclusion, it is evident from this study that among gram positive organisms *CONS*, and *S. aureus* and among gram-negative organisms *Klebsiella pneumoniae*, *E. coli* are the leading cause of paediatric sepsis. Also the most challenging observation was the resistance pattern of these isolates to multiple antibiotics. The main forces driving the increase in antimicrobial resistant bacteria are the inappropriate use of antibiotics. Specific antibiotic utilization strategies like antibiotic restriction, combination therapy and antibiotic escalation and de-escalation may help to decrease or prevent the emergence of resistance.

# Recommendation

Continuous monitoring of paediatric septicaemia and formulation of appropriate antibiotic policy is the need of the hour for surveillance and control of infection in the hospital.

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