

Bloodstream Infection in the Paediatric Cancer Patients: Bacteriological Profile and Drug Resistance Patterns in Shenzhen, China

Sandip Patil^{1,2}, Bruno S Lopes³, Hongyu Chen⁴, Lian Ma² and Feiqiu Wen^{1,2*}

¹Department of Haematology and Oncology, Shenzhen Children's Hospital, District Futian, Shenzhen, China

²Paediatric Research Institute, Shenzhen Children's Hospital, District Futian, Shenzhen, China

³Department of Medical Microbiology, School of Medicine, Medical Sciences and Nutrition, Aberdeen University, China

⁴Department of Laboratory Medicine, Shenzhen Children's Hospital, District Futian, Shenzhen, China

*Corresponding author: Feiqiu Wen, Department of Haematology and Oncology, Shenzhen Children's Hospital, District Futian, Shenzhen, China, Tel: +86-18938690333; Fax: +86-75583009888; E-mail: fwen62@126.com

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Abstract

Bloodstream Infections (BSI) remain a main cause of death in pediatric cancer patients. We present the first report on the microbiological profile of bacteremia and drug resistance patterns in Shenzhen, China. We examined the types of bloodstream infectious agents and their antimicrobial susceptibility pattern in the pediatric oncology patients, from Shenzhen Children's Hospital from January 2016 to October 2018. *Acinetobacter baumannii* (12.82%), *Klebsiella pneumoniae* (12.82%) and *Staphylococcus epidermidis* (34.61%) were highly prevalent in cancer patients with bloodstream infection. Extended-spectrum beta-lactamase-producing *Escherichia coli* and *Klebsiella pneumoniae* were also predominant. 29% of isolates were resistant to more than two class of antibiotics so call them as multi-drug resistant isolates.

Keywords: Multi-drug resistant bacterial infection; ESBL; Cancer

Letter to the Editor

Bloodstream Infections (BSI) remains a key cause of morbidity and mortality in patients undergoing cancer treatment [1]. Antimicrobial Resistance (AMR) pattern of pathogens causing BSI, significantly differ in developed and developing countries [2]. In this study, we focused on the trends of etiological agents causing BSI and determined the antimicrobial susceptibility pattern of bacteria isolated from pediatric oncology patients in Shenzhen, China for the same. A total of 607 non-repetitive isolates were collected by the Department of Haematology and Oncology, Shenzhen Children's Hospital during January 2016-October 2018. A total of 607 isolates recovered from all specimens includes blood, pus, urine, and catheter-associated secretions but our concerns with 78 (12.85%) isolate from a bloodstream infection. We studied total 78 (12.85%) isolates recovered from samples with BSI, comprising of 17 (2.8%), 40 (6.5%) and 21 (3.4%) isolates from the year 2016, 2017 and 2018 respectively. The Horasan and his co-worker reported BSI to be 21.4% in adult cancer patients which is higher than our observation [3]. In our study, 40 (51.28%) isolates were recovered from a female, while 38 (48.78%) were from male patients. The present study revealed the frequency of occurrence of Gram-negative bacteria to be about 47.43% (n=37) which was slightly lower than that of Gram-positive bacteria with 52.57% (n=41). Our observations are comparable with a published study conducted in Egypt [4]. All recovered isolates were confirmed by the API20 system in our clinical laboratory. During this period, blood samples were accounted for pathogenic bacteria *Staphylococcus epidermidis* 34.61% (n=27), followed by *Acinetobacter baumannii* 12.82% (n=10), *Klebsiella pneumoniae* 12.82% (n=10), *Pseudomonas aeruginosa* 10.25% (n=8), *Staphylococcus hemolyticus* and *Staphylococcus hominis* each 8.9% (n=7), *Escherichia coli* 5.12% (n=4), *Comamonas testosteroni* 3.8%

(n=3) and *Enterobacter cloacae* 2.5% (n=2). Antimicrobial susceptibility was determined by broth dilution method and breakpoint was determined as per CSLI standard. In addition, Extended-Spectrum Beta-Lactamase (ESBL) production was confirmed by using combination disc test. The test was performed by using Disc of both cefotaxime and ceftazidime, alone and in combination with clavulanic acid. The Gram-Negative pathogens were resistant to Amikacin, Ampicillin, Aztreonam, Ceftriaxone, Cefotetan, Cefazolin, Nitrofurantoin, Ampicillin-sulbactam, Trimethoprim, Tobramycin, and Piperacillin but sensitive to Ertapenem, Cefepime, Gentamicin, Imipenem, Levofloxacin and Ciprofloxacin (Figure 1). Gram-positive bacteria were resistant to the commonly used first-line of antibiotics (Figure 2). Antimicrobial susceptibility results showed that all *Acinetobacter baumannii* and *Klebsiella pneumoniae* isolates were resistant to carbapenem antibiotics. These multi-drug resistant pathogens are considered to be superbugs which may pose a serious threat to human health, especially in cancer patients. Such multi-drug resistant *Klebsiella pneumoniae* was reported from bloodstream infection in Tunisia [5]. We confirmed a rarely observed *Comamonas testosteroni* in bloodstream infection which showed 100% resistance to commonly used antibiotics. ESBL production was confirmed in two *Escherichia coli* and two *Klebsiella pneumoniae* isolates. The *Staphylococcus* species were resistant to Trimethoprim sulfamethoxazole followed by Nitrofurantoin, Oxacillin, Erythromycin, Tetracycline, Clindamycin, and Ciprofloxacin but sensitive to Quinupristin-dalfopristin, Moxifloxacin, Tigecycline, and Vancomycin (Figure 2). In this study, we evaluated the antimicrobial resistance pattern of Gram-positive and Gram-negative pathogens causing BSI, on the basis of phenotypic expression. *Acinetobacter baumannii*, *Klebsiella pneumoniae*, and *Staphylococcus epidermidis* were highly prevalent in the paediatric cancer patients with BSI. High microbial resistance was detected in causative agents of bloodstream infection in the paediatric patients with cancer. Our findings, bacteriological

profile and antimicrobial resistance pattern of BSI in the paediatric cancer patients, can help to determine the factors governing these infections, to antibiotic treatment policy and to improve the infection

control measures. We suggest that the continual monitoring of bloodstream infections is highly essential.

Bacteria	No. isolates	Drug-resistant isolates, %																	
		AMK	AMP	ATM	CAZ	CIP	CRO	CTT	CZO	ETP	FEP	GEN	IMP	LVX	NIT	SAM	SXT	TOB	TZP
<i>Acinetobacter baumannii</i>	10	100	100	100	100	0	100	100	100	N/D	0	0	0	0	100	10	20	0	N/D
<i>Klebsiella pneumoniae</i>	10	20	100	30	4	30	30	10	30	10	30	40	10	20	100	100	30	40	20
<i>Pseudomonas aeruginosa</i>	8	37.5	100	N/D	100	0	100	100	100	N/D	37.5	37.5	25	0	100	100	100	12.5	75
<i>Escherichia Coli</i>	4	0	0	0	25	50	50	100	100	0	0	0	0	100	0	75	100	50	100
<i>Comonas testosteroni</i>	3	100	100	100	0	0	100	100	100	N/D	100	100	0	0	100	100	100	100	100
<i>Enterobacter cloacae</i>	2	100	100	2	0	0	0	0	100	0	0	0	0	0	100	100	0	0	0

* AMK, Amikacin; AMP, Ampicillin; ATM, Aztreonam; CAZ, Ceftazidime; CIP, Ciprofloxacin; CRO, Ceftriaxone; CTT, Cefotetan; CZO, Cefazolin; ETP, Ertapenem; FEP, Cefepime; GEN, Gentamicin; IMP, Imipenem; LVX, Levofloxacin; NIT, Nitrofurantoin; SXT, Trimethoprim; FEP, Cefepime; SAM, Ampicillin-sulbactam; TOB, Tobramycin; TZP, Piperacillin

Figure 1: Antimicrobial Susceptibility of Gram-negative bacteria from a paediatric bloodstream infection.

Bacteria	No. Isolates	Drug-resistant isolates, %																
		CIP	CLI	ERY	GEN	LNZ	LVX	MXF	NIT	OXA	PEN	QDA	RIF	SXT	TC	TGC	VAN	
<i>Staphylococcus epidermidis</i>	27	14.8	29.6	46	25.9	7.4	11.11	0	96.2	59.25	0	0	11.11	100	37	0	0	
<i>Staphylococcus hemolyticus</i>	7	85.7	85.7	100	85.7	100	6	6	85.7	100	85.7	0	0	85.7	85.7	85.7	0	0
<i>Staphylococcus hominis</i>	7	0	14.28	57.14	14.28	71.42	0	0	71.42	14.28	2	0	14.28	100	57.14	0	0	

* CIP, Ciprofloxacin; CLI, Clindamycin; ERY, Erythromycin; GEN, Gentamicin; LNZ, Linezolid; LVX, Levofloxacin; Mxf, Moxifloxacin; NIT, Nitrofurantoin; OXA, Oxacillin; PEN, Penicillin; QDA, Quinupristin-dalfopristin; RIF, Rifampin; SXT, Trimethoprim-sulfamethoxazole; TC, Tetracycline; TGC, Tigecycline; VAN, Vancomycin.

Figure 2: Antimicrobial Susceptibility of Gram-positive bacteria from a paediatric bloodstream infection.

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