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Bacterial Isolates and Drug Susceptibility Pattern of Sterile Body Fluids at a Tertiary Care Hospital in Northern India

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Authors' contributions

This work was carried out in collaboration among all authors. All authors read and approved the final manuscript.

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ABSTRACT

Background: Body fluids obtained from sterile body sites are expected to be devoid of any pathogenic or commensal microorganisms. The cause of infected body fluids could be any pathological agents or skin contaminants harbored from intensive care units. This study identifies commonly isolated bacteria from the body fluid samples and their antibiotic sensitivity pattern.

Methods: All body fluid samples (except blood, cerebrospinal fluid and contaminated samples) received in Bacteriology section of Department of Microbiology in tertiary care centre in Northern India from November 2019 to May 2020 were included in the study. All microorganisms obtained on bacterial culture were subjected to identification by standard biochemical tests or MALDI-TOF-MS assay and antibiotic sensitivity testing by Kirby Bauer disc diffusion test.

Results: Out of 363 samples of body fluids, 113 (31.12%) showed bacterial growth on culture. Male predominance among patients with liver disease was deemed statistically significant in comparison to those without liver disease. Comorbidities like hypertension and encephalopathy among the patients with liver disease was statistically significant in comparison to those without liver disease. *Escherichia coli* was the most commonly isolated bacteria (20.35%) which was followed by

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Pseudomonas spp (15.92%) and *Klebsiella pneumoniae* (11.50%). Fifty one (45.13%) isolates were multidrug resistant. The isolation of MDR microorganisms from body fluid samples is statistically significant in cancer patients. Among MDR microorganisms, *Klebsiella pneumoniae* was deemed most resistant with presence of Extended spectrum beta lactamase (ESBL) character and also showed complete resistance to Carbapenems but all isolates were susceptible to Colistin. **Conclusion:** This study shows spectrum of bacterial isolates observed from body fluid samples received in our laboratory and helps in empirical treatment of patients based on antibiotic susceptibility patterns. It also highlights importance of adhering to antibiotic sensitivity results and infection control practices to prevent spread of multidrug resistant infection in hospital environment.

Keywords: Body fluids; multidrug resistance (MDR); chronic liver disease (CLD); MALDI-TOF-MS; antibiotic sensitivity pattern.

1. INTRODUCTION

Body sites are called sterile when they have no bacteria or other microorganisms as commensals in healthy condition. The infections caused in these sites could be either due to any pathological agents or skin contaminants harbored from intensive care units [1]. The body fluids obtained from sterile body sites are similarly expected to be devoid of any pathogenic or commensal microorganisms. Sterile body fluids include cerebrospinal fluid (CSF), synovial fluid, pericardial fluid, pleural fluid and peritoneal dialysate fluid. Most common causative agents of pathogenic microorganisms infecting the sterile body fluids are lactose fermenting gram negative Escherichia bacteria like coli. Klebsiella pneumoniae. Enterobacter species and Citrobacter species, non lactose fermenting gram negative bacteria like Acinetobacter species, Burkholderia cepacia, Pseudomonas species and gram positive organisms like Methicillin resistant Staphylococcus aureus and Enterococcus species which lead to morbid infections with poor outcome [2,3].

The morbidity and ability to cause life threatening infections has rendered these cases a medical emergency that demands early diagnosis and suitable treatment. Due to low inoculum of pathogenic bacteria and early administration of empirical antibiotics there were fewer chances of retrieving positive cultures [4]. The difficulty in diagnosing the causative microorganism of body fluid infections and need for effectively managing the patients admitted to the intensive care unit has made it the need of the hour to frame an antibiotic policy and to know the common pathogenic microorganisms with appropriate antibiotic sensitivity pattern [5].

As far as we know, there is limited knowledge regarding the microbiological characteristics,

bacterial spectrum and antimicrobial resistance in our geographical setting. The knowledge of the common pathogenic bacteria and their antibiotic susceptibility pattern is crucial for the clinicians to combat a range of infections and administer adequate antibiotics. So, this study conducted at a tertiary care centre in Northern India sheds light on the spectrum of bacterial isolates from the body fluid samples collected from outpatient department (OPD) and inpatient department (IPD) patients at our centre along with their antibiotic susceptibility patterns.

2. MATERIALS AND METHODS

This retrospective, observational, single centre study was conducted from November 2019 to May 2020 in the Bacteriology section of the Department of Microbiology at a tertiary care centre. A total of 363 non repeat samples of clinically suspected infected body fluids were collected aseptically and the samples included in the study consist of pleural fluid (110, 30.30%), pericardial fluid (20, 5.50%), ascitic fluid (181, 49.86%) and peritoneal dialysate fluid (52, 14.32%) samples.

Inclusion criteria: All the samples received from patients with clinically suspected body fluid infections, from the OPD and IPD patients at our centre, irrespective of age and gender were included.

Exclusion criteria: 1) Blood samples, 2) Cerebrospinal fluid samples, 3) Contaminated samples, 4) Body fluids with delay in transportation for more than 2 hours.

Sample processing: The samples were collected at a combined receiving station, sent to the bacteriology section of the Department of Microbiology and processed in our laboratory according to the standard protocols. The

samples were subjected to Grams' stain and culture. The culture media used were Blood agar, Mackonkey agar and Robertsons' cooked meat broth (RCM). Isolated colonies were observed on the Blood and Mackonkey agar plates and turbidity was observed in the RCM and the isolates were identified by Grams' stain and standard biochemical tests.

Antimicrobial susceptibility testing: The antibiotic sensitivity testing was performed for each of the bacterial isolates by Kirby Bauer Disc Diffusion method and Epsilometeric test method according to the guidelines of CLSI [6]. Antibiotic discs were used for Amikacin (30mcg), Ampicillin (10mcg), Ampicillin-Sulbactum (10/10mcg), Azetreonam (30 mcg), Ceftazidime (30mcg), Ceftriaxzone (30 mcg), Cefoperazone-Sulbactum (75/10 mcg), Cefoxitin (30 mcg), Ciprofloxacin (5 mcg), Clindamycin (2 mcg), Doxycycline (10 mcg), Ertapenem (10 mcg), Erythromycin (15 mcg), Gentamicin (10 mcg), Imipenem (10 mcg), Levofloxacin (5 mcg), Linezolid (30 mcg), Meropenem (30 mcg), Minocycline (30 mcg), Trimethoprim-Tigecycline (15 mcq). (1.25/23.75 sulphamethoxazole mca). Teicoplanin (30 mcg), Piperacillin-tazobactam Vancomycin (30 mcg) and (100/10 mcg), Colistin (0.016-256 mcg). Epsilometeric test strips were obtained from bioMérieux and used as per manufacturer's instructions. Briefly, inoculums were prepared for each bacterial isolate by adjusting the turbidity to 0.5 McFarland standard and spread on Muller-Hinton agar plates. Antibiotic discs and E-test strips were placed on the agar plates and incubated overnight at 37°C for 24 h. The zones of inhibition were measured and the isolates were sensitive, intermediate, classified as and resistant according to CLSI tables and guidelines [6].

Microbiological characteristics and drua resistance pattern were analyzed for all the samples included in the study. We further demonstrated the risk factors for isolation of which MDR microorganisms, among are Pseudomonas resistant to at least 3 antibiotic groups, carbapenemases-resistant and ESBLproducing Enterobacteriaceae, Enterococcus resistant to vancomycin, and MRSA [7,8]. We also assessed the risk factors for isolating bacterial body fluid infections in patients with and without chronic liver disease.

The statistical analysis for this study was performed by observing frequencies. Quantitative

variables were expressed as mean and standard deviation. In the analysis of risk factors for MDR, the comparison between groups for categorical variables was estimated by using χ 2 tests. The results were presented as 95% CIs. Statistical analysis was performed using the software program IBM SPSS Statistics version 20.0 (SPSS Inc.), with p < 0.05 considered as statistically significant.

3. RESULTS

3.1 Demographics

Out of the 363 samples of clinically suspected infected body fluids, 113 (31.12%) were found to be positive on bacterial culture. Among these culture positive patients a majority of 39 (34.50 %) were in the age group of 41 to 60 years followed by 21 to 40 years and 61-80 years which was 33 and 23 cases respectively. Body fluid samples from male patients were more predominant than samples from women, 74 (65.5%) patients were men and 39 (34.5%) patients were women respectively. Among the patients belonging to the age group of 41-60 years, male patients were predominant. The mean age of the patients suffering from liver disease in this study is 41.82 ± 20.80 , while the mean age among those without liver disease is 42.16 ± 21.33 and the male predominance among the patients with liver disease was deemed statistically significant in comparison to without liver disease, as seen in those Table 1. Out of the 113 culture positive samples. 63 (55.8%) were ascitic fluid samples, 38 (33.6%) were pleural fluid samples, 6 (5.3%) were peritoneal dialysate fluid samples, 4 (3.5%) were pericardial fluid samples and 2 (1.8%) were synovial fluid samples (Fig. 1). Escherichia coli was the most common isolated bacteria (20.35%) which was followed Pseudomonas spp (15.92%)and bv Klebsiella pneumoniae (11.50%) as observed in Fig. 2.

Among the clinical characteristics of the patients included, the patients were divided into two groups of those with liver disease and those without liver disease, as demonstrated in Table 1. Among the comorbidities taken into account among the patients included in our study group hypertension and encephalopathy among the patients with liver disease was statistically significant in comparison to those without liver disease.

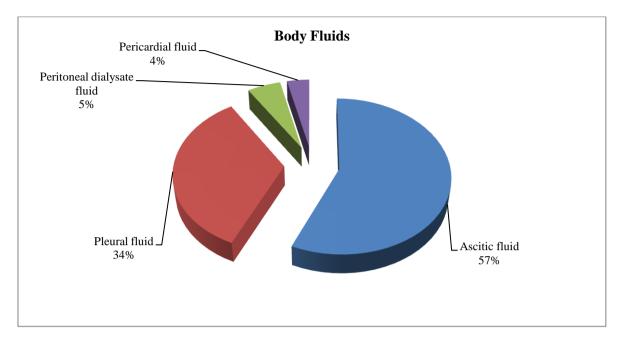


Fig. 1. Distribution of types of infected body fluid samples received in our laboratory (N= 113)

Table 1. Demographic characteristics of patients and risk factors for bacterial body fluid
infections with and without chronic liver disease (CLD) (N=113)

	With CLD (n=55)	Without CLD (n=58)	P-value
Demographics			
Age, years, mean (SD)	41.82 ± 20.80	42.16 ± 21.33	0.9318
Gender, male/female	43:12	31:27	0.006*
Comorbidities			
Diabetes mellitus, %	14	11	0.406
Chronic kidney	17	24	0.247
disease, %			
Heart disease, %	7	13	0.178
Hypertension, %	23	13	0.027*
Pleural effusion, %	16	26	0.084
COPD, %	4	7	0.390
Malignancy, %	6	11	0.231
Encephalopathy, %	21	5	<0.001*
Organ transplant, %	4	1	0.152
Post operative patients,	17	20	0.686
%			
Anemia, %	42	44	0.950
Other parameters			
Length of hospital stay,	22.38 ± 15.61	25.12 ± 19.08	0.407
mean (SD)			
Total leukocyte count,	15378.18 ± 10473.77	14076 ± 9262.49	0.485
mean (SD)			
SAAG ratio, mean (SD)	1.82 ± 0.905	1.52 ± 0.978	0.093
Death, %	16	18	0.822

* p-value <0.05 is significant

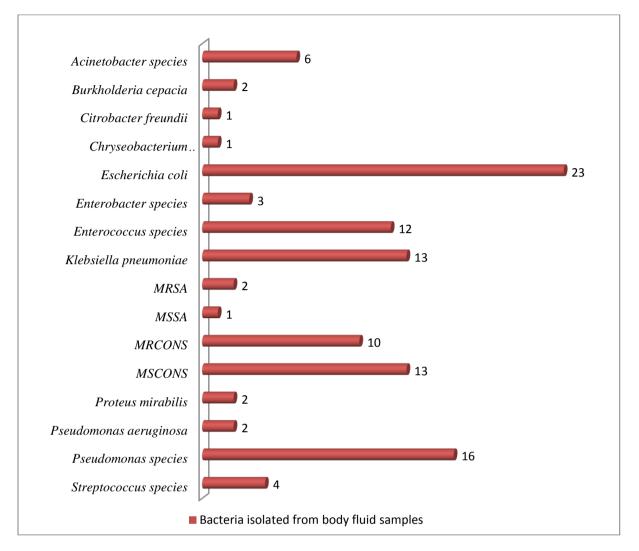


Fig. 2. Rate of bacteria isolated from the body fluid cultures performed in the laboratory (N=113)

MSCONS- Methicillin sensitive coagulase negative Staphylococcus MRCONS- Methicillin resistant coagulase negative Staphylococcus MSSA- Methicillin sensitive Staphylococcus aureus MRSA- Methicillin resistant Staphylococcus aureus

Multidrug resistance (MDR) was commonly encountered when reporting antibiotic sensitivity testing of the isolates. Fifty one (45.13%) isolates were Multidrug resistant. The mean age of patients suffering from body fluid infections with MDR microorganisms is 44.41 ± 18.33. Male predominance is observed in this study in cases of MDR infections of body fluids. The most common comorbidities encountered among the multidrua resistant isolates were anemia (78.43%), chronic liver disease (45.09%), chronic kidney disease (41.17%), post-operative hypertension patients (39.21%), (31.37%), pleural effusion (29.41%) and malignancy (23.53%). Table 2 demonstrates that the isolation of MDR microorganisms from body fluid samples is a statistically significant in the patients suffering from malignancies. The mean length of hospitalization in the patients is 25.24 19.62 days, the mean ± total leukocyte count and the mean serum ascites albumin gradient (SAAG) ratio of the samples from which the MDR microorganisms were isolated is 16221.57 ± 10400.81 per cubic mm and 1.59 ± 1.117 respectively.

3.2 Microbiological Characteristics

The microorganism most frequently isolated from the body fluid samples was Escherichia coli (20.35%) which was followed by Pseudomonas and Klebsiella pneumoniae spp (15.92%) (11.50%) as observed in Fig. 2. The microorganism mostly commonly isolated from ascitic fluid samples was Escherichia coli (14/23, 60.86%) followed by Klebsiella pneumoniae (9/13, 69.23%) and Pseudomonas spp (44.44%). The patients were grouped into two groups of patients with (55 patients) and without (58 patients) chronic liver disease, as demonstrated in the Table 1. Twenty three (41.81%) MDR microorganisms were isolated from the patients with chronic liver disease, while 28 (48.27%) MDR microorganisms were isolated from patients without liver disease. The most common microorganisms isolated among the patients with chronic liver disease were Escherichia coli (15. 27.27%) followed by Pseudomonas spp (10, 18.18%) and Klebsiella pneumoniae (6, 10.91%), of the above mentioned microorganisms 40%. 50% and 100% of the respective microorganisms Among were MDR. the gram positive microorganism. Methicillin Resistant Staphylococcus aureus was isolated in 2 (1.8%) and Enterococcus spp was isolated in 12 (10.6%) body fluid samples.

3.3 Multidrug Resistance

Fifty one (45.13%) isolates from body fluid samples were Multidrug resistant. Out of the most commonly isolated gram negative bacteria, Klebsiella pneumoniae seems to be the most resistant to most of the first line drugs for lactose fermenting gram negative bacteria: it is completely resistant to Ciprofloxacin, Ceftazidime Cefoperazoneand Ceftriaxzone. and Sulbactum, thus showing the presence of Extended spectrum beta lactamase (ESBL) character and showed Carbapenem resistance by complete resistance to Imipenem, Meropenem and Ertapenem and all isolates were susceptible to Colistin (Table 3).

The Acinetobacter spp isolates obtained were most susceptible to second line drugs like Minocycline, Tigecycline and drug of last resort Colistin, with 83.33% susceptibility to each drug, followed by Amikacin, to which only 50% of microorganisms were susceptible. The microorganism most resistant was to Ceftazidime, Ceftriaxzone and Cefoperazone Sulbactum, with 16.67% susceptibility to each isolate.

Demographics and risk factors	MDR microorganisms (n= 51/113, 45.10%)	P-value	95% CI
Demographics			
Age, years, mean (SD)	44.41 ± 18.33	0.268	39.25 – 49.56
Gender, male/female %	62.74/37.26	0.578	1.24 – 1.51
Comorbidities			
Diabetes mellitus, %	14 (27.45%)	0.216	1.60 – 1.85
Chronic liver disease, %	23 (45.09%)	0.491	1.41 – 1.69
Chronic kidney disease,	21 (41.17%)	0.327	1.45 – 1.73
%			
Heart disease, %	11 (21.56%)	0.328	1.67 – 1.90
Hypertension, %	16 (31.37%)	0.920	1.55 – 1.82
Pleural effusion, %	15 (29.41%)	0.122	1.58 – 1.84
COPD, %	6 (11.76%)	0.509	1.79 – 1.97
Malignancy, %	12 (23.53%)	0.022*	1.64 – 1.89
Encephalopathy, %	11 (21.56%)	0.741	1.67 – 1.90
Organ transplant, %	1 (1.96%)	0.248	1.94 – 2.0
Post operative patients, %	20 (39.21%)	0.184	1.47 – 1.75
Anemia, %	40 (78.43%)	0.599	1.10 – 1.33
Other parameters			
Length of hospital stay,	25.24 ± 19.62	0.426	19.72 – 30.75
mean (SD)			
Total leukocyte count,	16221.57 ± 10400.81	0.140	13296.29 -
mean (SD)			19146.84
SAAG ratio, mean (SD)	1.59 ± 1.117	0.446	1.27 – 1.90
Death, %	<u>19 (37.25%)</u>	0.132	1.24 – 1.51

 Table 2. Demographic characteristics of patients and risk factors for isolation Multidrug

 resistant microorganisms in patients with purulent infections (N=51)

* p-value <0.05 is significant

The *Pseudomonas spp* isolates isolated were most susceptible to second line drugs like Minocycline and Colistin, with 75% and 93.75% susceptibility respectively to each of these drugs, followed by Amikacin and Piperacillin -Tazobactam, to each of which the microorganism was 37.50% sensitive. The microorganism was most resistant to Azetreonam followed by Ceftazidime, with a susceptibility of 18.75% and 25% respectively (Table 3).

The gram positive cocci isolated from the samples in were grouped into Coagulase negative *Staphylococcus*, Coagulase positive

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Staphylococcus and Enterococcus spp. Of all the gram positive microorganisms isolated from samples Coagulase these positive and Coagulase negative Staphylococcus and Enterococcus isolates were most sensitive to Doxycycline, with 81.48%, 90.90% and 68.18%, respectively, followed by Amikacin among the Coagulase positive Staphylococcus and Coagulase Staphylococcus, negative with a susceptibility of 100% and 82.60% respectively, positive All the Coagulase Staphylococcus and Coagulase negative Staphylococcus were susceptible to Vancomycin and in case of Teicoplanin all Coagulase positive

Table 3. Percentage Sensitivity pattern for first and second line drugs in most commonly
isolated Gram negative bacilli

Antibiotics	<i>E. coli</i> % sensitivity	Acinetobacter species % sensitivity	<i>K. pneumoniae</i> % sensitivity	Pseudomonas species % sensitivity
Amikacin	73.91%	50.00%	15.38%	37.50%
Ceftazidime	8.69%	16.67%	0.00%	25.00%
Ceftriaxzone	0.00%	16.67%	0.00%	-
Ciprofloxacin	8.69%	33.33%	0.00%	-
Levofloxacin	-	-	-	37.5%
Cefoperazone- Sulbactam	34.78%	16.67%	0.00%	31.25%
Imipenem	60.87%	33.33%	0.00%	31.25%
Meropenem	60.87%	33.33%	0.00%	31.25%
Ertapenem	52.17%	-	0.00%	-
Colistin	100.00%	83.33%	100.00%	93.75%
Tigecycline	100.00%	83.33%	69.23%	-
Minocycline	95.65%	83.33%	69.23%	75.00%
Azetreonam	-	-	-	18.75%
Piperacillin-	-	-	-	37.5%
Tazobactam				

Table 4. Percentage Sensitivity pattern for first and second line drugs in most commonly isolated Gram positive cocci

Antibiotics	Coagulase positive Staphylococcus % sensitivity	Coagulase negative Staphylococcus % sensitivity	<i>Enterococcus</i> % sensitivity	
Ampicillin	-	-	25%	
Ampicillin- Sulbactum	66.67%	47.83%	41.67%	
Amikacin	100%	82.60%	-	
Clindamycin	33.33%	60.87%	-	
Cefoxitin	33.33%	56.52%	-	
Doxycycline	100%	73.91%	83.33%	
Erythromycin	0%	13.04%	-	
Gentamicin	0%	0%	33.33%	
Levofloxacin	0%	26.08%	16.67%	
Vancomycin	100%	100%	41.67%	
Teicoplanin	100%	91.30%	41.67%	
Linezolid	-	-	91.67%	
Minocycline	-	-	83.33%	

Staphylococcus were susceptible but 91.30% were susceptible among Coagulase negative Staphylococcus. Among the isolates of Enterococcus spp susceptibility to Linezolid and Minocycline was 91.67% and 83.33% respectively, thus proving that 15.7% of the Enterococci isolates were multidrug resistant (Table 4). The risk factors associated with the acquisition of multidrug resistance are increased length of stay in the hospital, patient comorbidities like immunosuppression, chronic liver disease, heart disease, diabetes mellitus and malignancy. Malignancy was a significant statistically risk factor for acquiring multidrug resistance, as described in Table 2.

4. DISCUSSION

The spectrum of pathogenic microorganism causing body fluid infections along with their antibiotic susceptibility patterns changes from one geographical region to another from one time to another. The increase in incidence of comorbidities, immunocompromised conditions and increase in length of hospital stay have combined to render ideally sterile ascitic and pleural fluids samples infected [9].

In this study, 31.13% of samples were bacterial culture positive, which was in agreement with the other studies were 30%, 31% positive cultures were noted [10,11].

The predominant microorganisms isolated was Escherichia coli (23), CONS (23), followed by (18). Klebsiella Pseudomonas species pneumoniae (13), Enterococcus species (12), Acinetobacter baumannii (6), Streptococcus species (4), Enterobacter species (3). Staphylococcus aureus (3), Burkholderia cepacia (2), Proteus mirabilis (2), Yeast like cells (2), Citrobacter freundii (1) and Chryseobacterium indologenes (1).

The most common organism isolated from pleural fluid was Escherichia coli (7) and these findings correlates with the findings of studies by Sharma et al, Sujatha et al and Evans et al where the most common causative agent of pleural fluid infections was Escherichia coli [10-While the second most common 12]. microorganism to cause pleural infections was Pseudomonas spp (6) in contrast to Acinetobacter spp in the study by Sharma et al and Klebsiella pneumoniae in studies by Sujatha et al and Evans et al.

This study represents the ability of the aerobic gram negative bacteria to predominantly cause purulent pleural fuid infections. In studies by Jain et al, Gupta et al and Mohanty et al a similar finding was observed that the gram negative bacteria were more commonly isolated from the purulent infections of pleural fluid [13-15]. The gram positive microorganisms isolated from infections consist mainly pleural fluid of Enterococcus species followed by Staphylococcus aureus. The isolation of aerobic gram negative bacteria or isolation of multiple pathogenic bacteria holds poor prognosis and thus arises need of rigorous antimicrobial therapy [5].

The gram negative bacteria most commonly isolated from ascitic fluid samples was Escherichia coli (14), followed by Klebsiella pneumoniae (9) and Pseudomonas spp (8) which is correlates with the studies conducted by Arroyo et al and Chawla et al which showed Escherichia coli as the most common microorganism causing peritonitis [16,17]. In a study by Harshika et al, non lactose fermenting bacteria were the most commonly isolated microorganisms from the ascitic fluid samples disagrees with the findings of this study but dissimilarity in the spectrum of pathogenic bacteria causing peritonitis also depends upon the geographical area of isolation [5].

Very few samples of pericardial fluid samples were obtained at our centre, which is only 4% of body fluid samples received in our laboratory, were culture positive. Out of the 4 culture positive pericardial fluid samples, one sample showed culture of Staphylococcus aureus as seen in a study by Reuter et al. [18]. Very rare cases of pericardial fluid infections are noted in literature and these infections can lead to life threatening complications like cardiac tamponade and constrictive pericarditis [5]. The most common underlying causes of pericardial infections include people undergoing haemodialysis, those from AIDS, those undergoing suffering chemotherapy and thoracic surgery [19,20].

The gram negative bacteria were almost completely susceptible to drug of last resort, Colistin, followed by good sensitivity for drugs like Minocycline and Tigecycline. A decreasing sensitivity of the gram negative bacterial isolates to carbapenems was noted, in contrast to good susceptibility to carbapenems seen in a studies conducted by Harshika et al. [5] and Sharma et al. [10]. The isolates in were relatively resistant to Gram positive bacteria isolated were highly sensitive to Vancomycin, Teicoplanin and Linezolid which is in agreement with the study by Sujatha et al. [11] but disagrees with the complete resistance to Gentamicin shown in among our isolates. The *Pseudomonas species* isolates were highly sensitive for Colistin and Minocycline. Only 37.5% isolates of *Pseudomonas species* were susceptible to Piperacillin- Tazobactam which is in contrast to a study conduct by Harshika et al. [5].

Thus, evaluation of the results of antimicrobial resistance to the isolates is suggestive of rapid emergence of multidrug resistance. Ability to attain the resistant genes from other resistant microorganisms has lead to rise of multidrug resistance through the years. Before seeking treatment at a tertiary care centre, most patients are subjected to unnecessary antimicrobial therapy that renders them resistant to most first line antibiotics given to treat a specific microorganism. Lack of proper antimicrobial stewardship program and no adherence to the antibiotic susceptibility testing has lead to increased antimicrobial resistance.

5. CONCLUSION

This study shows spectrum of bacterial isolates observed from the body fluid samples received in our laboratory and helps in guiding the empirical treatment of patients based on antibiotic susceptibility patterns. The increase in the incidence of multidrug resistance also points out the need to implement strict adherence to the antibiotic sensitivity. The spread of multidrug resistant and extensively drug resistant microorganisms can also be curtailed by strict adherence to infection control practices, educating the health care workers and patients about hand hygiene and make them aware about the morbidity of suffering from a multidrug resistant infection.

DISCLAIMER

The products used for this research are commonly and predominantly use products in our area of research and country. There is absolutely no conflict of interest between the authors and producers of the products because we do not intend to use these products as an avenue for

any litigation but for the advancement of knowledge. Also, the research was not funded by the producing company rather it was funded by personal efforts of the authors.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

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