



STUDY OF MULTIDRUG RESISTANT ACINETOBACTER IN INTENSIVE CARE UNIT (ICU) PATIENTS.

Microbiology

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ABSTRACT

Introduction- Acinetobacter had gained increasing attention as a nosocomial pathogen causing severe, life-threatening hospital infections, especially in intensive care units (ICUs). It is notorious to develop multi drug resistant. Management of multidrug-resistant Acinetobacter spp. infections is a great challenge for physicians and clinical microbiologists. **Objectives:** This study was designed to study the various species of Acinetobacter prevalence and antibiotic susceptibility patterns of Acinetobacter sp. as isolated from patients lodged in intensive care units (ICUs) of a tertiary care hospital in central India. **Material & methods:** The study was carried out in microbiology laboratory of a tertiary care hospital from July 2018 to July 2020. Samples were collected, processed and Acinetobacter isolates were identified by standard microbiological procedures. Antimicrobial susceptibility testing performed by kirby-bauer disc diffusion method as per CLSI guidelines. **Results:** Out of total 200 isolates of Acinetobacter, ACB complex 189 (94.5%) was found to be the most common species isolated. Maximum isolates were obtained from medicine intensive care unit. The most common infection was found to be pneumonia. Maximum isolates were resistant to Ceftriaxone, Ceftazidime, Ciprofloxacin etc and least resistance was seen in Imepnem, Piperacillin-Tazobactam, Amikacin.. All the strains were sensitive to Colistin. **Conclusion:** Acinetobacter has emerged as an important nosocomial pathogen showing high degree of resistance to most antimicrobials. Early identification and continued surveillance of their antibiogram is essential for preventing the spread of Acinetobacter in hospital environment.

KEYWORDS

Acinetobacter Sp., Intensive Care Units, Nosocomial Infections, Drug Resistance.

In recent year, Acinetobacter has emerged as an important nosocomial pathogen mainly affecting patients with impaired host defenses in Intensive care unit.¹ Acinetobacter spp. go so quickly from being unknown and unnamed organisms to become among the most dangerous pathogenic organisms of the present time. The history of this organism from being a saprophytic organism to superbug has developed to us to analyze the reason and mechanism of such evolution of "Acinetobacter".²

Acinetobacter species are ubiquitous in environment. They are usually free living saprophytes in soil and water.³ Up to 20% of healthy ambulatory adults exhibit cutaneous colonization by Acinetobacter and are the most common gram negative bacteria carried on the skin of hospital personnel.⁴

Acinetobacter baumannii has emerged as an important nosocomial pathogen, particularly for patients in intensive care units and with invasive indwelling devices. One of the most significant emerging multidrug-resistant (MDR) pathogens is Acinetobacter baumannii.⁵

They have been implicated in a variety of nosocomial infections, including bacteraemia, urinary tract infection, secondary meningitis, but their role as agents of nosocomial pathogen in pneumonia, particularly ventilator associated pneumonia in patients confined to hospital intensive-care units Acinetobacter have been isolated from various opportunistic infections, including skin and wound sepsis, septicaemia, pneumonia, endocarditis, meningitis and UTI.^{6,7}

Prior to the 1970s, it was possible to treat Acinetobacter infections with a range of antibiotics, including aminoglycosides, β -lactams.⁸ However, resistance to all known antibiotics has now emerged in Acinetobacter species thus leaving the majority of today's clinicians in great dilemma.⁹ Unfortunately, at this stage, very little is in the therapeutic pipeline and the new agents with activity against gram-negative organisms are all modifications of existing classes. Novel antibiotic targets and mechanisms of action are urgently required.¹⁰

MATERIAL AND METHODS

The study is being carried out in the Dept. of Microbiology in a tertiary care hospital from July 2018 to July 2020.

Clinical specimens like sputum, blood, pus, urine, CSF, peritoneal

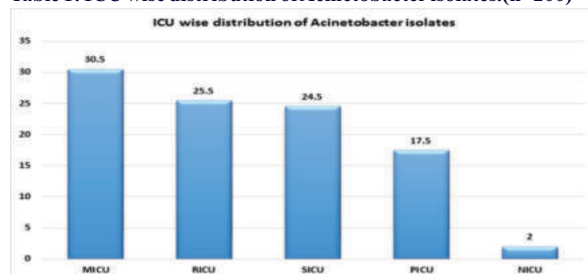
fluid etc. from patients with pneumonia, septicemia, wound sepsis, urinary tract infection, meningitis, peritonitis, endocarditis were received from patients of all age groups and both sex admitted in a various intensive care units (MICU, PICU, NICU, SICU) by standard collection procedures.¹¹ No specific exclusion criteria envisaged.

Acinetobacter were confirmed by catalase test, Nitrate test, OF test, citrate test, Growth at 44°C, growth on Macconkey. Further they were speciated by growth at 37°C, and 44 °C, Hemolysis, Gelatin hydrolysis, Arginine dihydrolase. Antimicrobial susceptibility testing was performed on Acinetobacter spp. by modified Kirby Bauer method as per the CLSI guidelines 2019.¹² MICs of Colistin were determined by the E-test method according to the manufacturer's guidelines (Himedia laboratories Pvt.Ltd.Mumbai).¹³

OBSERVATIONS AND RESULTS

This study was hospital based cross sectional observational study which was carried out during period of July 2018 to July 2020 in Department of Microbiology of a tertiary care center. A total of 200 isolates of Acinetobacter spp. obtained from various ICU were included in the present study.

Table 1: ICU wise distribution of Acinetobacter isolates.(n=200)



Out of total 200 Acinetobacter isolates, maximum were from Medical ICU (MICU) 61 (30.5%), followed by Respiratory ICU (RICU) 51 (25.5%), 49 (24.5%) from Surgery ICU (SICU), 35(17.5%) from pediatrics ICU (PICU) while least number of samples were from NICU 4 (2.0%) (Table 1).

Table 2: Infection wise distribution by Acinetobacter isolates

Sr. No	Infections	No of isolates (%)
1.	Pneumonia	91(45.5)
2.	Skin and soft tissue Infection	42(21.00)
3.	Septicemia	36(18.00)
4.	Urinary tract infection	12 (6.00)
5.	Ascites	12(6.00)
6.	Pleural effusion	4 (2.00)
7.	Meningitis	3 (1.5)
	Total	200

Table 2 shows, *Acinetobacter* isolates obtained from patients suffering from various infections. Out of total 200 isolates majority were of patients of pneumonia 91 (45.5%), followed by skin and soft tissue infection 42 (21%), septicemia 36 (18%). *Acinetobacter* was also found to be responsible for urinary tract infection 12 (6%), ascites 12 (6%), pleural effusion 4 (2%) and meningitis 3 (1.5%).

Table 3: Species wise distribution of *Acinetobacter* isolates

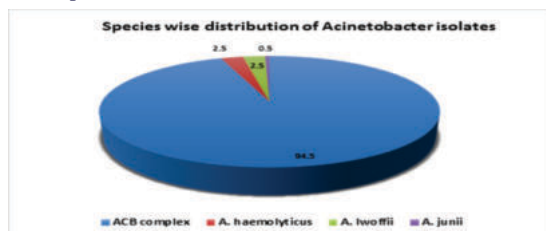


Table 3 shows that, out of total 200, ACB complex (*Acinetobacter calcoaceticus*–*Acinetobacter baumannii* complex) 189 (94.5%) was found to be the most common species isolated, followed by *A. lwoffii* and *A. haemolyticus* 2.5 % each, *A. junii* 0.5%. ACB complex isolates were significantly higher (94.5%) as compared to other species (5%). This was statistically significant with p value less than 0.001.

Table 4 : Antimicrobial Susceptibility pattern of *Acinetobacter* isolates

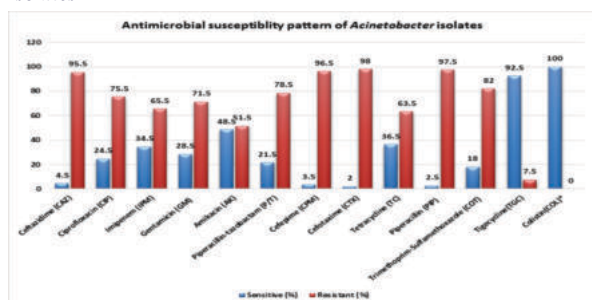


Table 10 shows that the maximum sensitivity of *Acinetobacter* was seen to Colistin (100%) followed by tigecycline (92.5%), and amikacin (48.5%) while Maximum resistance was observed to cefotaxime (98%), piperacillin (97.5%), cefepime (96.5%), ceftazidime (95.5%), Trimethoprim-Sulfamethoxazole (COT) (82%) and piperacillin-tazobactam (78.5%), Imipenem (65.5%).

DISCUSSION-

The incidence of *Acinetobacter* infection in the intensive care unit (ICU) is rising and causes a great concern to worldwide clinicians and intensivists due to their ability to develop resistance to multiple classes of antibiotics.

In the present study out of total 200 *Acinetobacter* isolates, maximum were from Medical ICU (MICU) 61 (30.5%), followed by Respiratory ICU (RICU) 51 (25.5%), 49 (24.5%) from Surgery ICU (SICU), 35(17.5%) from pediatrics ICU (PICU) while least number of samples were from NICU 4 (2.0%) (Table 2).

This study revealed nosocomial infection of *Acinetobacter* spp in MICU was 30.5% which was comparable with a study done by Neeta B Patwardhan,¹⁴ who found 34.5%. whereas infections in ICUs observed by Mathai AS¹⁵ was on lower side (10%).

In present study infection caused by *Acinetobacter* in RICU was 25.5%. In various Indian studies showed 29.28%,¹⁶ 34.8%,¹⁷ 41.8%¹⁸. There can be several reasons for this. The ICU mainly caters to medically critically ill patients. Accordingly, infectious diseases

(pulmonary or extrapulmonary) are the most common indication for admission in our ICU followed by respiratory failure, acute severe asthma, exacerbation of chronic obstructive lung disease and a few others.

Infection in SICU in our study was 24.5%. In a study from western Maharashtra 18.08%¹⁹ of *Acinetobacter* was responsible for nosocomial infection from patients admitted to the SICU. A Study done in north India reported 38.96%²⁰ and One such study by Nahar A et al. reported a prevalence of 33.7% in ICU patients.²¹ *Acinetobacter* infection contributed to 17.5% in PICU and 2 % from NICU while a study by shete et al showed 10.8 % of total septicemia cases.²²

In this study, 91 (45.5%) isolates from pneumonia patients. Mathai AS¹⁵ found *Acinetobacter* spp in 59 % of patients. Albert et al²³ isolated 45.00% *Acinetobacter* from VAP which was in concordance to our study whereas Ruiz-Santana et al²⁴ isolated 38.00%. skin and soft tissue infection due to *Acinetobacter* spp found to be 42 (21%). Joshi et al²⁵ reported 27.50%, While Dash et al²⁶ reported 56.9% respectively. 36 (18%) isolates of *Acinetobacter* spp were from septicemia cases. Various study ranges from 14-24% in various study²⁷⁻³⁰. 12(6%) isolates were isolated from UTI cases comparable findings of 5% and 8% were observed by Jaggi et al.²⁷ and Oberoi et al.³² respectively. Higher values of 31.60 % isolates were reported by Sands et al.³¹ There are reports of *Acinetobacter* meningitis in various study like Joshi et al²⁵ reported 0.8%, in 2013 Sinha et al²⁸ reported 0.71%, in 2014 Islahi et al²⁹ reported 2.17% isolates from meningitis.

In 200 isolates ACB complex (*Acinetobacter calcoaceticus*–*Acinetobacter baumannii* complex) 189 (94.5%) was found to be the most common species isolated, followed by *A. lwoffii* and *A. haemolyticus* 2.5 % each, *A. junii* 0.5%. Idomir et al³³ reported *A. baumannii* (87%) as commonest species isolated, followed by *A. calcoaceticus* (11.4%), *A. lwoffii* (0.93%) and *A. haemolyticus* (0.93%). Rit et al³⁴ reported *A. baumannii* (74.02%), *A. lwoffii* (14.2%), *A. haemolyticus* (7.79%) and *A. junii* (3.81%) in 2012. Species isolated by Tripathi et al³⁵ were *A. baumannii* (79.43%), *A. calcoaceticus* (12.14%), *A. lwoffii* (3.73%), *A. haemolyticus* (2.81%) and *A. junii* (1.89%).

Acinetobacter baumannii have proven to be among the most important pathogens frequently involved in serious hospital infection outbreaks.

The emerging resistance in today's world has created a major public health dilemma. Within the gram-negative bacilli, the non-fermentative bacilli occupy an important place and among them *Acinetobacter*.

In our study maximum sensitivity was found to be 100% to colistin and 92.5% tigecycline, study conducted by behera et al³⁷ and Khan ID et al³⁸ reported the same while islahi et al²⁹ noted lower sensitivity of 80.33% to colistin. Whereas 48.5% of *Acinetobacter* isolates were found to be sensitive to Amikacin which was comparable with study done by Tripathi et al³⁵ who noted 55.14% sensitivity to this drug.

carbapenems were considered as the gold standard for treating *Acinetobacter* sp. infections, but unfortunately, carbapenem resistance toward *Acinetobacter* sp. is becoming common.^{25, 36} we recorded 65.5% resistance to imipenem similar findings were reported by Kaur et al.⁴⁰

It is well known that this organism is rapidly developing resistance to various groups of antimicrobials including 3rd generation cephalosporins, fluororoquinolones, Aminoglycosides and carbapenems indicating high prevalence of MDR strains.^{22,28} In the present study, maximum resistance was seen to cefotaxime followed by piperacillin, cefepime and ceftazidime i.e 98%, 97.5%, 96.5% and 95.5% respectively, Which are quite comparable with various other studies.^{28,35,38}

Summary

Acinetobacter once considered non-virulent has been the documented cause of broad range of severe nosocomial infections. They cause a significant proportion of infections, especially in critically-ill patients receiving care in the intensive care unit (ICU) settings.. The iatrogenic factors i.e. overuse of broad spectrum antibiotics and high-risk patients with advanced age, history of alcohol intake, smoking, severity of illness, etc. Along with inherent and/or easily acquired mechanisms of

resistance makes this pathogen one of the most significant microbial challenges of the current era. Despite the majority of *A.baumannii* strains still being susceptible to carbapenems, many institutions around the world are faced with the challenging issue of pandrug resistance, leaving only limited therapeutic option. As a clinicians, microbiologists, and scientists must think broadly about approach to antimicrobial drug development, as novel targets will no doubt provide the most reward for afflicted patients. A combination of control measures is often required to eradicate these organism from the hospital environment. Continued surveillance in ICUs along with awareness of the need to maintain good housekeeping and control of the environment, including equipment decontamination, strict attention to handwashing and isolation procedures, and control of antibiotic usage, especially in high-risk areas, appears to be the combination of measures most likely to control the previously unabated spread of *Acinetobacter* spp. in hospitals.

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