Antimicrobial Susceptibility Pattern of Acinetobacter calcoaceticus-Acinetobacter baumannii Complex Isolated from Sputum in a Tertiary Care Hospital

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ABSTRACT

Introduction
Acinetobacter calcoaceticus-Acinetobacter baumannii (ACB) complex is one of the commonest cause of hospital acquired and ventilator associated pneumonia. Multidrug resistant Acinetobacter species have become a matter of huge concern. This study was done to find out the antibiotic susceptibility pattern of Acinetobacter calcoaceticus-Acinetobacter baumannii complex from sputum samples.

Methods
This descriptive cross-sectional study was carried out in Clinical Microbiology laboratory from July 2018 to Jan 2019 after ethical approval. Acinetobacter calcoaceticus-Acinetobacter baumannii complex was identified on the basis of its microscopy and morphological characteristics followed by biochemical tests. Antibiotic sensitivity test of isolated pathogens was done using Muller Hinton Agar by Kirby-Bauer method.

Results
Of the 384 culture positive sputum specimen, 76 (19.80%) were Acinetobacter calcoaceticus-Acinetobacter baumannii complex. Most of the isolates were resistant to commonly used antibiotics, 72.36% of the isolates were multidrug resistance and 3.95% isolates were resistant to tigecycline.

Conclusion
This study provides valuable information regarding prevalence of Acinetobacter calcoaceticus-Acinetobacter baumannii complex from sputum specimen. The alarming number of Multidrug resistance isolates is worrisome finding. Antibiotics like Tigecycline and Colistin which is still sensitive to isolates should be cautiously used only in MDR cases.

Keywords: Acinetobacter, ACB complex, antibiotic resistance, MDR, Nepal

INTRODUCTION

Acinetobacter calcoaceticus-Acinetobacter baumannii complex (ACB) organisms are aerobic, non-fermenting gram negative coccobacilli. Clinically ACB organisms mainly cause health-care associated infections but are also associated with community acquired infections.¹ The respiratory tract is an important site of colonization and is the most frequent site of infection.² The most common clinical condition associated with these microorganisms is hospital-acquired pneumonia (HAP), particularly for patients receiving mechanical ventilator assistance.³

Despite advances in health care and wide variety of antibiotics, life threatening infections caused by ACB complexes are considered as one of the major health problem. Emergence of infections caused by multidrug resistant (MDR) strains of ACB complex increase morbidity, mortality and impose an enormous burden on health care cost. Alarmingly some strains of ACB exhibits resistance to ‘last-resort’ drugs like tigecycline, Polymixin B and
Colistin sulfate. The resistance pattern of bacteria changes over time and varies from place to place. Therefore regular surveillance is needed to treat infections empirically and effectively.\(^1,6\)

This study aimed to determine the prevalence of ACB complex in sputum sample, their antibiotic susceptibility pattern and frequency of MDR ACB complexes in our setting.

METHODS

A hospital based descriptive cross sectional study was carried out in Clinical Microbiology Laboratory of Kathmandu Medical College and Teaching Hospital (KMCTH), Kathmandu Nepal from the month of July 2018 to Jan 2019. Ethical approval was received from Institutional Review Committee. All the sputum samples, appropriately collected, labelled, properly transported and processed for aerobic bacterial cultures. A total of 384 culture positive sputum isolates were included in this study. Samples received were processed according to standard microbiological procedures.\(^6\) Suspected colonies of ACB complex were further processed. Identification of ACB complex was done on the basis of colony character, gram’s staining and biochemical tests.\(^7\) Antimicrobial susceptibility of all isolates was determined by the standard Kirby Bauer disk diffusion method according to norms of Clinical Laboratory Standards Institute (CLSI). Antibiotics included were Amikacin (30µg), Ceftriaxone (30 µg), Ceftazidime (30 µg), Ciprofloxacin (5 µg), Pipracillin (30 µg), Piperacillin/Tazobactum (100/10 µg), Imipenem (10 µg), Tigecycline (15 µg), Colistin (20 µg) and Polymyxin B (300units).\(^8\) In this study, if the isolates were resistant to at least three classes of first line antimicrobial agents, they were regarded as MDR.\(^9\)

RESULTS

Out of 384 culture positive sputum sample a total of 76 (19.80%) ACB organisms were isolated, out of which 40 (52.7%) were from males and 36(47.4%) were from females. The mean age was 60.46 ranging from 16 to 98 years, with most growths from 61-80 years of age (44.74%). Prevalence of ACB complex among age group 61-80 was found to be statistically highly significant (P value< 0.001).

The antimicrobial susceptibility pattern of the ACB complex showed resistance to most of the antibiotics. Highest resistance was seen to Ceftriazone (86.84%), Pipracillin (84.21%) and Ciprofloxacin (81.58%). Resistance to Ceftazidime was (80.70%) and that of Amikacin was (77.63%). Highest sensitivity was seen to Colistin (100%), Tigecycline (94.74%). Over all sensitivity and resistance pattern is shown in Table 2.

Out of 76 ACB complex, 72.36% were MDR.

DISCUSSION

The prevalence and resistance of ACB complex to antibiotics has amplified considerably over the past few years. In this study, the prevalence of ACB complex among sputum isolates is 19.80%, which is similar to studies done by Nepal et al in 2016 which showed the prevalence of 17.33%, but our result is higher than the study done in 2013 by Mishra et al which showed prevalence of 11.23% and Tamang et al in 2005 reported prevalence as 3.9% from respiratory secretions. These findings suggest an increasing trend in the prevalence of ACB complex.\(^10-12\)

Though ACB complex is known to have low virulence but among gram negative bacteria they show resistance to majority of commonly used antibiotics. This study demonstrated higher prevalence of drug resistance among the ACB complex towards majority of the antibiotics. Our study reported high resistant to cephalosporins and Fluroquonolones. Ceftazidime resistance was seen in 80.7% which is similar to the studies done in Nepal by Amatya et al and Shrestha et al.\(^13,14\) This is probably due to the widespread use of third-generation cephalosporins in our setup. Some of the studies done in Nepal around the world also showed 100% resistance to Ceftazidime.\(^15,16\) Ciprofloxacin resistance was 81.58% in our study which is higher than other studies conducted in Kathmandu which showed resistance of 64.5% and 68.7%.\(^11,12\) The differences in the sample types, size, source, site of study, antibiotics usage, and

| Antibiotics          | Resistant (%) | Intermediate resistant (%) | Sensitive (%) |
|----------------------|--------------|---------------------------|--------------|
| Amikacin             | 77.63        | 2.63                      | 19.74        |
| Ceftriazone           | 86.84        | 2.63                      | 13.16        |
| Ceftazidime          | 80.70        | -                         | 19.30        |
| Ciprofloxacin        | 81.58        | -                         | 18.42        |
| Colistin             | 0.00         | -                         | 100          |
| Meropenem            | 61.84        | -                         | 35.53        |
| Pipracillin          | 84.21        | -                         | 15.79        |
| Pipracillin/Tazobactum | 73.68     | 1.32                      | 25.00        |
| Tigecycline          | 3.95         | 1.32                      | 94.74        |
the hospital infection control practices followed can influence the resistance pattern. Equally likely is that the data reflects an actual increase in the resistance to these antibiotics over the years as there is wide use of antibiotics available over the counter without specific laboratory tests in our country.

Once considered as drug of choice for *Acinetobacter* infections, Carbapenem resistant strains are reported worldwide.17,18 In our study Meropenem was resistant in 61.84%. Study conducted by Zelleri et al suggested carbapenem resistance is higher in places where they are used extensively.19 Carbapenems are used extensively in Nepal which explains increase in resistance to these antibiotics.

The frequency of MDR ACB complex in our study was 72.36% which is comparable to Amatya et al in 2018.13 Other parts of Southeast Asia have also shown increasing MDR trend.20 This current finding implies that only higher antibiotics like Colistin and Tigecycline as a treatment option to combat MDR ACB complex.

Antibiotics like tigecycline and colistin should be judiciously used only in MDR cases. Our study did not document any resistance to Colistin which was similar to previous studies done in Nepal,11,13,14 But 3.95% of isolates were tigecycline resistant. Tigecycline is one of the “last-resort” antimicrobial agents for antibiotic resistance in ACB infection. Though it has been used for only about 10 years, significant percentage of resistance has been observed in many studies done in Nepal and elsewhere.1,12,16

**CONCLUSION**

The present study establishes ACB complex as important isolate from sputum sample. With increase in prevalence of MDR ACB complex cautious use of antibiotics should be practiced. Irrational use of antibiotics, absence of antimicrobial stewardship program in hospitals, lack of surveillance and reporting system, failure to observe infection control practices like hand washing and barrier nursing could be some reasons for this problem. Although no isolate exhibited resistance to Colistin and few isolates were Tigecycline resistant screening test and MIC determination are recommended in monitoring the response to therapy and for early detection of impeding resistance among local strains and these antibiotics should only be used in MDR cases.

**CONFLICT OF INTEREST**

None declared.

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