

# Ventilator-associated Pneumonia in Intensive Care Unit in a Tertiary Care Hospital

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## Abstract

**Background:** Ventilator-associated pneumonia (VAP) is the second most common cause of health care-associated infections with increased mortality.

**Aims and Objectives:** To determine the rate of VAP, causative pathogens, and its antimicrobial susceptibility pattern for early diagnosis and prompt treatment in reducing mortality in patients on mechanical ventilation in intensive care setting.

**Materials and Methods:** The study was done between September 2019 and November 2019 involving 180 patients on Mechanical ventilation. Microbiological processing, antimicrobial susceptibility testing, and enzyme detection for drug resistance done as per Clinical Laboratory Standards Institute guidelines.

**Results:** The rate of VAP ranged from 8% to 9%. The commonest pathogens encountered were *Klebsiella pneumoniae*, (38%) *Pseudomonas aeruginosa* (34%), *Acinetobacter baumannii* (20%), *Staphylococcus aureus*. (7%). Extended spectrum beta-lactamase production was seen in *K. pneumoniae*, (48%) *P. aeruginosa* (84%), *A. baumannii* (36%). Methicillin-resistant *S. aureus* accounted for 25%.

**Conclusion:** Early accurate diagnosis and treatment with appropriate antibiotics is the key in reducing mortality and morbidity in VAP in intensive care units.

**Key words:** Extended-spectrum  $\beta$ -Lactamases antibiotics, *Klebsiella pneumoniae*, Ventilator associated pneumonia

## INTRODUCTION

Ventilator-associated pneumonia (VAP) is the most common hospital-acquired infection with an incidence of 10–25% and accounts for 25% of all intensive care units (ICU) infections. VAP is associated with very high mortality which is highest in ICU mainly due to multidrug-resistant organisms such as *Acinetobacter* spp., Methicillin-resistant *Staphylococcus aureus*.<sup>[1]</sup> Hospital-acquired pneumonia prolongs the cost and duration of hospital stay.<sup>[2]</sup> Early-onset VAP results from aspiration of endogenous community-acquired organisms, for example., *Streptococcus pneumoniae*, *Haemophilus influenzae*,

and other organisms (aerobic gram-negative bacilli). Late-onset VAP is more severe and results usually from aspiration of gastric/oropharyngeal secretions and caused by potentially drug-resistant organisms such as MRSA and *Pseudomonas*.<sup>[2]</sup> The etiologic diagnosis of VAP is critical to the appropriate antibiotic therapy. Hospital personnel and the environment can be the microbial source, and because of the overuse of antimicrobial agents, they have become multiple drug resistance organisms. During these years, the use of invasive diagnostic and therapeutic methods has saved many lives but on the other hand, it can cause some life-threatening consequences due to persistence and resistance infections. Quantitative Endotracheal aspirate at a cutoff point of  $10^5$  cfu/mL is a practical diagnostic method in clinical suspected VAP. It can be easily and repeatedly performed to help clinicians in decision-making regarding antibiotic use.<sup>[3,4]</sup> Inappropriate or inadequate antimicrobial therapy for ventilator-associated pneumonia is associated with increased mortality and the emergence of multiple

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drug resistance, which emphasizes the importance of a microbiologically based approach in the treatment of VAP.<sup>[5]</sup> The aim of the study was to determine the rate of ventilator associated pneumonia, microbiological profile of endotracheal aspirates in a mechanically ventilated patients with pneumonia and its antimicrobial susceptibility pattern for early treatment of the patients.

## MATERIALS AND METHODS

The study is a cross-sectional study done from September 2019 to November 2019.

The study group involved patients who were mechanically ventilated in ICU in a tertiary care hospital mainly serving patients with diseases related to the nervous system.

Sample- Endotracheal aspirates were collected using the sterile sample collection kit. After proper handwashing and wearing sterile gloves before suctioning, the endotracheal secretions were collected from the endotracheal tube with the help of sterile mucous trap. The specimen collected was immediately transported to the laboratory within 1 h of collection. In case of Endotracheal secretion culture,  $>10^5$ – $10^6$  cfu/ml was considered significant for the diagnosis of VAP. Organisms causing VAP are isolated and recorded. Details of patients such as name, age, sex, hospital ID no., probable diagnosis, cause of putting patient on ventilator are recorded. About 100 samples of endotracheal aspirates were collected during the period. The samples are transported in a sterile container immediately to the laboratory. The samples were subjected to direct microscopic examination by performing gram staining.

The samples were inoculated into Mac Conkey agar, Blood agar, and Nutrient agar using Semi quantitative technique. Further processing, identification of pathogenic bacteria, and detection of its antimicrobial susceptibility pattern were done as per standard Clinical Laboratory Standard Institute guidelines.

The antibiotics used for Gram negative bacilli were Imipenem (10 mcg), Meropenem, Cefaperazone/Sulbactam (75/30 mcg), Piperacillin/Tazobactam (100/10 mcg), Amikacin (30 mcg), Gentamycin (10 mcg), Levofloxacin (5 mcg), Ofloxacin (5 mcg), Cephataxime (30 mcg), Ceftriaxone (30 mcg) Cefuroxime (30 mcg), Cefpodoxime (10 mcg), Ceftazidime (30 mcg), Cefaperazone (75 µg), Cefixime (5 mcg), and Amoxyclav (20/10).

The antibiotics used for Gram-positive cocci were Cefoxitin (30 mcg), Vancomycin (30 mcg), Linezolid

(30 mcg), Teicoplanin (30mcg), Cefaperazone/Sulbactam (75/30 mcg), Piperacillin/Tazobactam (100/10 mcg), Amikacin (30 mcg), Gentamycin (10 mcg), Levofloxacin (5 mcg), Ofloxacin (5mcg), Cephataxime (30 mcg), Ceftriaxone (30 mcg) Cefuroxime (30 mcg), Cefpodoxime (10 mcg), Amoxyclav (20/10).

### Detection of Methicillin Resistance among Gram-positive Organisms

The Clinical and Laboratory Standards Institute (CLSI) guidelines (2020) had recommended Cefoxitin disc diffusion method for the detection of MRSA. This is performed by using a 30 µg Cefoxitin disc and an inhibition zone diameter of  $\leq 21$  mm is reported as Methicillin-resistant and  $\geq 22$  mm is considered as Methicillin sensitive.

### Detection of Extended-spectrum $\beta$ -Lactamases (ESBL) among Gram-negative Organisms

ESBLs are defined as  $\beta$ -lactamases capable of hydrolyzing oxyimino-cephalosporins and are inhibited by  $\beta$ -lactamase inhibitors. Isolates showing a zone of inhibition  $< 22$  mm for Ceftazidime were tested for ESBL production as per CLSI criteria.

### Combined Disc Method

A combined disc method using Cefaperazone (75 mcg) and Cefoperazone/Sulbactam (75/30 mcg) performed for phenotypic confirmation of ESBL production, as recommended by the latest guidelines of CLSI. Organism was considered as ESBL producer if there was a more than 5 mm increase in zone diameter of Cefoperazone/Sulbactam disc and that of Cefaperazone disc alone.

## RESULTS

Among 180 samples of endotracheal aspirate, pathogens were isolated from 58 samples. The most common

**Table 1a: Sex distribution**

Male	112
Female	78

**Table 1b: Age distribution**

Age group	Diabetes
21–30	33
31–40	19
41–50	10
51–60	21
61–70	7

organism isolated was *Klebsiella pneumoniae* followed by *Pseudomonas aeruginosa*, *Acinetobacter baumannii*, and *S. aureus*.

## DISCUSSION

VAP is a common complication in patients requiring mechanical ventilation and it is the second most common

**Table 2: Rate of ventilator-associated pneumonia**

Month	Rate
September 2019	8.5 per 1000 ventilator days
October 2019	8 per 1000 ventilator days
November 2019	9 per 1000 ventilator days

**Table 3: Underlying condition of patient requiring mechanical ventilation**

Underlying condition of patient	Number	Percentage
Trauma	92	51
Cerebrovascular accident	46	26
Tumours	34	19
Others	8	4

**Table 4: Bacteria isolated from endotracheal aspirate**

Name of organism	Percentage
<i>Klebsiella pneumoniae</i>	22 (38)
<i>Pseudomonas aeruginosa</i>	20 (34)
<i>Acinetobacter baumannii</i>	11 (20)
<i>Staphylococcus aureus</i>	4 (7)
Coagulase negative <i>Staphylococci</i>	1 (1)

**Table 5: Gram-negative and Gram-positive infections**

Total organism	Gram-negative infection	Gram-positive infection
58	53 (92%)	5 (8%)

**Table 6: Antibiotic susceptibility pattern of Gram-negative bacteria**

Organism	IMP %	MRP %	CFS %	PIT %	AK %	G %	LE %	OF %	CTX %	CTR %	CAZ %	CPZ %	CXM %	CFM %	CPD %	AC %
<i>Klebsiella pneumoniae</i>	93	67	64	58	42	42	47	27	33	33	31	31	18	181	18	40
<i>Pseudomonas aeruginosa</i>	95	64	49	49	72	54	70	60	41	41	46	41	0	0	0	0
<i>Acinetobacter baumannii</i>	85	62	85	69	54	54	69	54	31	31	31	31	6	6	6	6

IMP: Imipenem, MRP: Meropenem, CFS: Cefepime/Sulbactam, PIT: Piperacillin/Tazobactam, AK: Amikacin, G: Gentamycin, LE: Levofloxacin, OF: Ofloxacin, CTX: Cephataxime, CTR: Ceftriaxone, CXM: Cefuroxime, CPD: Cefpodoxime, CAZ: Ceftazidime, CPZ: Cefepime, CFM: Cefixime, AC: Amoxycyclav

**Table 7: Antibiotic susceptibility pattern of Gram-positive bacteria**

Organism	VA %	LZ %	CFS %	PIT %	AK %	G %	LE %	OF %	CX %	CTX %	CTR %	CAZ %	CXM %	CFM %	CPD %	AC %
<i>Staphylococcus aureus</i>	100	100	75	75	88	75	88	88	75	63	63	63	63	63	63	63
Coagulase negative <i>Staphylococci</i>	100	100	100	50	50	100	100	100	100	50	50	50	50	50	50	50

CFS: Cefepime/Sulbactam, PIT: Piperacillin/Tazobactam, AK: Amikacin, G: Gentamycin, LE: Levofloxacin, OF: Ofloxacin, CTX: Cephataxime, CTR: Ceftriaxone, CXM: Cefuroxime, CPD: Cefpodoxime, CAZ: Ceftazidime, CPZ: Cefepime, CFM: Cefixime, AC: Amoxycyclav, VA: Vancomycin, LZ: Linezolid, CX: Cefoxitin

cause of hospital-acquired infection.<sup>16]</sup> In this study, both male and female patients of different age groups were affected [Table 1a and 1b]. The rate of VAP was found to be 8.5%, 8%, and 9.5% for 3 months, respectively [Table 2] which as even observed by Kalanuria *et al.*<sup>17]</sup> In this study the most common underlying condition of the patient requiring mechanical ventilation included trauma (51%) followed by cerebrovascular accident (26%), tumors (19%) and others (4%) [Table 3]. Vadivoo *et al.*<sup>18]</sup> in their study also documented the similar finding. In my study, the commonest organism isolated was *K. pneumoniae* (38%), followed by *P. aeruginosa* (34%), *A. baumannii*, (20%) *S. aureus* (7%), and Coagulase-negative *Staphylococci* (1%) [Table 4]. Payal Modi *et al.* had documented a similar finding in their study.<sup>19]</sup> In this study most of the infections are caused by Gram-negative bacteria accounting for 92% as compared to Gram-positive infections which accounted for 8% only [Table 5]. The Gram-negative bacteria such as *K. pneumoniae*, *P. aeruginosa*, and *A. baumannii* were 90% sensitive to Imipenem, Meropenem, 60–70% sensitive to Cefepime/Sulbactam, Piperacillin/Tazobactam and 40% sensitive to Aminoglycosides and Fluoroquinolones, and 30% sensitive to Cephalosporins [Table 6]. Gram-positive bacteria like *S. aureus* were 100% sensitive to Vancomycin, Linezolid, 80–90% sensitive to Cefepime/Sulbactam, Piperacillin/Tazobactam sensitive to Aminoglycosides and Fluoroquinolones [Table 7]. In my study ESBLs producers among *K. pneumoniae* were 48%, *P. aeruginosa* were 84%, *A. baumannii* were 36% [Table 8]. Higher prevalence of ESBLs were reported by Khanal *et al.*<sup>19]</sup> MRSA accounted for 25% in this study [Table 9].

Current guidelines for the management of VAP emphasize the principles of early, epidemiologically based, broad-spectrum antibiotics, microbiologically guided de-escalation of therapy, and the shortest possible course of effective

**Table 8: Total number of extended-spectrum  $\beta$ -lactamases**

Organism	Extended spectrum $\beta$ -lactamases (%)
<i>Klebsiella</i> species	48
<i>Pseudomonas</i> species	84
<i>Acinetobacter</i> species	36

**Table 9: Total number of Methicillin-resistant *Staphylococcus aureus***

Organism	Methicillin-resistant <i>Staphylococcus aureus</i>
<i>Staphylococcus aureus</i>	25%

antimicrobial treatment.<sup>[10,11]</sup> Early, accurate diagnosis is, therefore, fundamental in the management of patients with VAP.<sup>[12]</sup>

## CONCLUSION

As the problems of excessive antibiotic use and antimicrobial resistance continue to grow, an accurate microbiological diagnosis of VAP is likely to be the key element in ensuring appropriate antibiotic coverage for multidrug-resistant organisms followed by de-escalation based on microbiological culture results and clinical response of patient as well as for limiting the use and duration of empirically prescribed broad-spectrum antibiotics.

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