

Compliance of Ventilator Associated Pneumonia Bundles and Incidence of Ventilator Associated Pneumonia in a Tertiary Care Surgical Intensive Care Unit: Prospective Observational Study

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ABSTRACT: Background: Ventilator-associated pneumonia (VAP) has a reported incidence ranging between 15% and 45 %. The mortality rate attributable to VAP is 27 - 43% when the causative agent was multidrug resistant. Length of stay (LOS) in the ICU is increased by 5 to 7 days and hospital (LOS) 2- to 3-fold in patients with VAP. VAP is defined as pneumonia occurring in a patient within 48 hrs or more after intubation and mechanical ventilation. Methods: At a tertiary care hospital, all adult patients over the age of 18 years admitted to the surgical ICU were screened for the risk factors for VAP from 01 Nov 2018 to 30 Apr 2020. The study population screened, while data was collected in only those requiring mechanical ventilation at admission or during the course of ICU stay and were prospectively followed for the occurrence of VAP until either discharge from the ICU or death. Only the first episode of VAP was evaluated. All mechanically ventilated patients had daily chest radiographs & white blood cell count along with hourly vital parameter charting and tracheal aspirate specimen / culture were sent on suspicion of a new infiltrate in the chest X-ray. Recording the compliance to VAP bundle were collected thrice a day in all ventilated patients during ICU admission.Results:The study cohort comprised of 621 surgical cases of which 232 patients required mechanical ventilation. The mean age of patients was 51.77 yrs with a male predominance (65.6%). 20 patients were excluded from the study as per exclusion criteria. The data collected was subjected to univariate analysis using Fisher's exact test & level of significance was set at P-value <0.05. It was found that 06 patients developed VAP with the incidence was 2.43%. The risk factors significantly associated with VAP in our study were presence of ETT, Tracheostomy, Head injury, Ryle's tube. The common organism isolated was Acinetobacter baumani. In 97.84% of patients atleast 5 of 9 studied VAP bundle components were adhered

to.**Conclusion:**The incidence of VAP noted in the surgical ICU over 1.5 years was 2.43%. More than 5 components of VAP bundles were adhered to in 97.84%.The difficulties in implementation of VAP bundles were related to non-availability of eqpt & patient specific nursing requirements.

Key-words: Ventilator associated pneumonia, mechanical ventilation, intubation, Acinetobacter baumani.

I. INTRODUCTION:

Ventilator-associated pneumonia (VAP) refers to pneumonia developed in patients who have been mechanically ventilated for duration of more than 48 hours [3]. The mortality rate attributable to VAP is 27% and has been reported to be as high as 43% when the causative agent was antibiotic resistant [4]. The length of stay in the intensive care unit is increased by 5 to 7 days [5] and hospital length of stay 2 to 3-fold in patients who develop VAP [6]. The risk of VAP is highest early in the course of hospital stay, and is estimated to be 3% per day during the first five days of ventilation, 2% per day during days 5-10 of ventilation and 1% per day thereafter [8]. Prevention is the key to limiting the morbidity and mortality while early & accurate diagnosis is fundamental in the management of patients with VAP [16]. VAP bundles were first introduced in 2004 by the Mayo clinic's [17] which were a set of recommendations found to have a significant effect in the prevention of ventilator associated pneumonia. This prospective observational study was to determine the incidence of ventilator associated pneumonia in the surgical ICU setting and compliance to the recommended ventilator associated pneumonia care bundle in our surgical ICU. This has neither been previously quantified nor studied at our institution.



II. MATERIALS AND METHODS:

After obtaining approval by the hospital ethics committee and informed consent, mechanically ventilated patients, admitted in the surgical ICU of tertiary care hospital were enrolled in to the study group. The data was prospectively collected at the surgical ICU of tertiary care teaching hospital and screened for risk factors of VAP between Nov 2018- Apr 2020. Although data was collected for all patients, those who required mechanical ventilation at admission or during the course of ICU stay were alone included in thestudy. Study group patients were prospectively followed for the occurrence of VAP until either discharge or death from the ICU. Data collected was analyzed using appropriate statistical analysis.VAP was defined as pneumonia occurring in a patient within 48 hrs or more after intubation with an endotracheal tube or tracheostomy tubes, which was not present before admission to the ICU. Only the first episode of VAP was evaluated. All mechanically ventilated patients had daily chest radiographs & white blood cell count along with hourly vital parameter charting. The tracheal aspirate specimen / culture was sent on suspicion of a new infiltrate in the chest Xray or new onset fever of $>101^{\circ}$ F.

The following basic data were collected at ICU admission: age, sex, premorbid conditions & concomitant diseases, presence of underlying malignancy, hospital-admission diagnosis were collected in all patients. Cause of ICU admission was classified as multiple injury, head injury, respiratory disease, neurologic disorder, cardiovascular disorder, intra-abdominal disorder, poisoning or miscellaneous. Other variables recorded include - admission source (ward or emergency room), having undergone surgery / emergency surgery. Specific medical care processes that were recorded as potential risk factors for the development of VAPinclude tracheostomy, dialysis, number of reintubations, presence of tube thoracostomy, sedative use, corticosteroids, inotropic drugs, presence and duration of central venous / arterial catheters, nasogastric tubes, type of nutritional support (parenteral or enteral feeding), duration of mechanical ventilation, and prior use of antibiotics. On ICU admission, the stringent implementation of the following components of the VAP bundle was ensured by all the staff after periodic suitable education in the ICU and the compliance noted as applicable. The education on VAP was conducted routinely.

(a) Elevation of head 30-40 degrees unless contraindicated

(b) Washing of hands before and after contact with each patient

(c) Use chlorhexidine mouthwash as part of daily mouth care

(d) Daily Sedation Vacation once a day

(e) Assessment for weaning from ventilator once a day

(f) Prophylaxis for DVT

(g) Prophylaxis for stress Ulcer

(h) Continuous removal of subglottic secretions using subglottic tubes

Daily checks were performed on implementation of the protocol once or thrice a day as required, while the patients are watched carefully for signs of fever with radiological patch, purulence in secretions, leukocytosis or leucopenia, culture positivity. Compliance to (a), (b), & (c) were noted thrice a day, while others noted once a day. All patients admitted to the surgical ICU were screened daily till discharge/ death from ICU.Patients requiring mechanical ventilation during the ICU stay were alone analyzed.Patient receiving palliative /terminal care/DNR orders. Patient admitted in the ICU and died within 24 hrs of ICU admission, Patients <18 years of age, Patients with pre-existing long term artificial airway devices were excluded from this study.

III. STATISTICAL ANALYSIS

Data analysis done by using SPSS version 17:0. After evaluating, the data were subjected to univariate analysis using Fisher' exact test to find the association between various parameters with occurrence of VAP. Level of significance was assumed at <5% ie P-value<0.05. Mann Whitney U test used to find the significance between median ICU stay with respect to occurrence of VAP and non-VAP as the data was skewed data and not following normal distribution.

IV. RESULTS:

The study cohort comprised of 621 patients comprising various post-surgical cases as well as direct surgical ICU admissions / transfers. 20 patients were excluded from the study as per exclusion criteria. Of the 601 patients enrolled in the study, 243 patients were mechanically ventilated and considered as a study group.



CONSORT DIAGRAM



The mean age of patients was 59.67 having a predominance of male population. There were no significant correlations between the age, sex and the incidence of VAP. Of the mechanically ventilated 243 patients, a total of 06 (six) patients developed VAPduring the ICU stay (incidence was 2.4 %) with the commonest causative organism being Acinetobacter baumani.

Age group	VAP			
	Present	Absent	Total	P-value
≤ 20	0	4	4	
21 - 40	0	47	47	
41 - 60	3	10	13	0.500
61 - 80	3	71	74	
> 80	0	5	5	

Table: Distribution of patients with respect to VAP and age (years)





Fig Age vs incidence of VAP

The total number of ventilator days amounted to 750 days and our incidence of VAP was 8 per 1000 ventilator days. The mean duration of ventilation was noted to be 12(±4.472) days for the VAP group and 2.84 (±3.49) days for the non VAP group. Longer duration of ventilation was found had a significantly higher incidence of VAP

and was statistically significant. The mean length of stay of the surgical ICU patient - 2.85 days. The mean duration of ICU stay, among ventilated patients in surgical ICU was 4.3 days. However, the average length of stay in the patients who developed VAP in our study was 16.5 days.

Table : Length of ICU stay				
VAD	LOS ICU stay			
VAP	Mean	SD		
Present (N=6)	16.5	4.03		
Absent (N=237)	4.5	3.77		
VAP	Median ICU stay	P-value		
Present (N=6)	nt (N=6) 11			
Absent (N=237)	3	< 0.001		

Post op neurosurgical cases formed the most common admission in our surgical ICU followed by post op oncology cases. Other admissions included cases pertaining to head injury, polytrauma, post op ortho, GI, urology, General surgery,

ENT etc. In our study, VAP occurred most commonly in head injury cases followed by one each in post operative neurosurgery, polytrauma and vascular case.

Table: Distribution of patients admitted in surgical ICU				
	VAP			
	Present	Absent	Total	
Head injury	3	24	27	
Poly trauma	1	8	9	
Post op Ortho	0	1	1	
Post op Neuro	1	83	84	



Post op GI	0	24	24
Post op Uro	0	2	2
Post op MDTC	0	65	65
Post op Gen Sur- gery	0	11	11
Post op Vascular	1	2	3
Post op ENT	0	1	1
Gen Surgery	0	3	3
ENT	0	1	1
GI	0	4	4
Neuro	0	4	4
MDTC	0	2	2
Vascular	0	2	2

Compliance with the VAP care bundle was defined as the percentage of mechanically ventilated patients for whom all eight components of the bundle are documented on the daily record. In our study it was found that the compliance to four of the major component of VAP bundles - head end elevation, oral/tracheal suction, chlorhexidine mouth wash, hand wash compliance were observed in almost every patients and the compliance rate was found to be >96%. The other components of VAP bundle - viz sedation vacation, assessment of weaning, stress ulcer prophylaxis was found to be > 90%. Compliance to VAP bundle components like DVT prophylaxis, subglottic tube due to some patient specific factors like immediate post op and non-availability of subglottic tube could not be observed effectively.

Compliance		VAP			Percentage of
		Present	Absent	Total	compnance
Head and elevation (> 30 degree)	Present	6	228	234	96.29%
	Absent	0	9	9	3.71%
Oral/Tracheal suction	Present	6	228	234	96.29%
	Absent	0	9	9	3.71%
Chlorhexidine mouth wash	Present	6	228	234	96.29%
	Absent	0	9	9	3.71%
Hand wash compliance	Present	6	228	234	96.29%
	Absent	0	9	9	3.71%

Table : Compliance of VAP bundles



Compliance		VAP			Percentage of
		Present	Absent	Total	compliance
Sedation vacation	Present	6	232	238	97.94%
	Absent	0	5	5	2.06%
Assessment wean- ing	Present	6	226	232	95.47%
	Absent	0	11	11	4.53%
DVT prophylaxis	Present	6	157	163	67.08%
	Absent	0	80	80	32.92%
Subglottic tube	Present	1	22	23	9.47%
	Absent	5	215	220	90.53%

V. DISCUSSION

The incidence of VAP reported in literature varies between 07-70% as studied in heterogeneous ICU settings. Our study was restricted to a surgical ICU in a tertiary care setting. The overall incidence found in our setting was 2.43% in a population studied over 18 months. In our study there was male predominance (67.48%) which was statistically insignificant (P- value 0.70). The average age group in our study was 59.6yrs. Presence of comorbidities like diabetes mellitus, asthma, renal failure, hypertension, COPD, stroke, jaundice did not play a significant role in the development of VAP. The average duration of ventilation in our study for non VAP patients 2.84 (\pm 3.49) days whereas it was 12 (\pm 4.47) days for VAP patients. The mean length of stay of the surgical ICU patient in our study was 2.85 days. However, among ventilated patients, the mean duration of ICU stay was 4.33days. The patients who developed VAP in our study had an ICU Length of stay of 16.5days while the non VAP patients had an ICU length of stay of 4.5days. This is in agreement with the studies which implicate increasing length of stay with increased incidence of VAP [6].

In our study, 03 out of 06 patients who developed VAP had severe head injury and the mortality rate was 100%.Our study found that the incidence of VAP was high in case of direct admission to the surgical ICU from the emergency room. This could be attributed to the prior status of the patient, likely aspiration prior to hospital admission, poor GCS and other co-morbidities. The most common organism associated with VAP in our study was AcinetobacterBaumanni followed by pseudomonas. Acinetobacter is a nonfermenting gram negative, aerobic coccobacillus found extensively in natural environment that has assumed an increasing importance in nosocomial infections in general and in VAP particular. Baribar et al. have reported 8.1% VAP cases caused by Acinetobacter [173].Various risk factors reported for the development of Acinetobacter VAP are ARDS, Large volume lung aspiration, head trauma and neurosurgery [174].

VI. COMPLIANCE

In our study we had a compliance to head end elevation in more than 96 % in a recording done thrice a day. In few cases the compliance was not carried out due to patient specific factors and occasionally lapses in nursing care. Although use of subglotic tubes is a recommended practice in the prevention of VAP the non-availability of tubes was a major factor in its use being minimal in our study. In our study the compliance to hand wash was adequate with almost 96 % compliance. Alcohol based hand rubs were the common method of compliance in our ICU. In our observational study, there was 96 % compliance to use of chlorhexidine mouth wash thrice a day. However Selective decontamination of the digestive tract (SDD) is not practiced in our ICU.

In our observational study the commonest organism isolated in suspected cases of VAP were Acinetobacter species and Pseudomonas. An earlier



study at our centre in 2010 showed an incidence of VAP of 32/1000 ventilated days. However, the present study was done in the surgical ICU of our hospital with aggressive implementation and education of the VAP bundles in this prospective trial. Our lower incidence of VAP could be attributed to the strict adherence of the VAP bundles [182].

Difficulties in implementation of VAP bundle: Patient related factors:

(a) Immediate post op

(b) Co-morbidities like coagulation disorders

(c) Sepsis induced coagulopathy

(d) Transportation and positioning of patients for imaging studies

Surgical factors:

(a) Patients with instructions contrary to VAP Bundle protocol

(b) Inability to wean

(c) Difficulty in oral hygiene and suctioning

Administrative factors:

- (a) Level of education among nursing staff
- (b) Non availability of subglottic tubes

VII. LIMITATIONS

We studied the incidence of only VAP and not hospital acquired pneumonia which did include non-ventilated patients. The second incidence of VAP if any was not studied. Our study was restricted to the patients admitted to the surgical ICU, where the expected co-morbidities were less compared to those in a medical or a general ICU and the average stay in ICU was only 2.85 days and mean duration of ventilation was 3.2 days. Hence the incidence of VAP could have been low. The severity scoring systems not taken into account and is a shortcoming in our study. The lack of availability of subglottic tube was a reason for poor compliance of this recommendation of the VAP bundle. Our sample was restricted to patients admitted in the surgical ICU and hence these results cannot be extrapolated to a general ICU. The aim of our study was to identify the incidence of VAP and not incidence of early onset or late onset of VAP.

VIII. CONCLUSION

VAP is a serious complication of mechanical ventilation. It is one of the most common of all hospital acquired infections which increases the morbidity and mortality as reported in literature. The incidence of VAP in our study was 6/243 ventilated patients which amounted to 2.4% of ventilated patients. The implementation of VAP bundle in our study was >90%. The strong adherence to VAP bundles which include - staff education & diligent implementation and improved nursing care probably had an important role in the low incidence of VAP in our study. However, the mortality attributable to VAP was 100% in our study. Strict adherence to the VAP bundles and incorporating these strategies to the bedside patient care of ventilated patients are likely to greatly decrease the incidence of VAP.

REFERENCES

- [1]. Klevens, Edwars, Richards, et al. Pub Health Rep 2007; 122:160-6.
- [2]. McEachern R Campbell GD Jr Hospital acquired pneumonia :Epidemiology, etiology and treatment. Infect dis clin North Am1998,12, 761-779.
- [3]. Tablan OC, Anderson LJ, Besser R, et al. Guidelines for preventing health-careassociated pneumonia, 2003: recommendations of CDC and the Healthcare Infection Control Practices Advisory Committee. MMWR Recomm Rep. 2004; 53(RR-3):1-36.
- [4]. Craven DE. Epidemiology of ventilator associated pneumonia. Chest. 2000;117 (4 suppl 2):186S-187S
- [5]. Safdar N, Dezfulian C, Collard HR, Saint S.Clinical and economic consequences of ventilator-associated pneumonia: a systematic review. Crit Care Med. 2005;33(10):2184-2193
- [6]. Kollef MH. The prevention of ventilator associated pneumonia. N Engl J Med.1999;340(8):627-634.
- [7]. American thoracic society. Hospitalacquired pneumonia in adults: diagnosis, assessment of severity, initial antimicrobial therapy, and preventive strategies. A consensus statement, American Thoracic Society,Nov 1995.Am J Respircrit care Med 1996;155:1711-1725. Trouillet JL, Chastre J VuagntA, Joly-Guillou ML,CombauxD, DombretMC, GilbertC.Ventilator associated pneumonia caused by potentially drug resistant bacteria. Am I Crit care med1998;157:531-539.
- [8]. Cook DJ,WalterSD,CookRJ,GriffithLE,GuyattGH, LeasaD,etal.Incidence of and risk factors for ventilated associated pneumonia in critically ill patients.Ann Intern med 1998;129:440.
- [9]. Johanson WG Jr,Pierce AK, SanfordJP, Thomas GD: Nosocomial respiratory infections with gram negative bacilli.The significance of colonization of the respiratory tract.Ann Intern Med 1972,77:701-06.
- [10]. Fabregas N, EwigS,Torres A, El-Ebiary M, Ramirez J, de La Bellacasa JP,

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BauerT,Cabello H:Clinical diagnosis of ventilator associated pneumonia revisite:comparitive validation using immediate post mortem lung biopsies,Thorax 1999, 54:867-73.Mediri, G U 1995.Diagnosis and differential diagnosis of ventilator associated pneumonia.Clin.chest med.16:61-93

- [11]. Ayala A,Perrin MM, Meldrum DR,Ertel W,Chaudry IH.Hemorrhage induces an increase in serum TNF which is not associated with elevated levels of endotoxin.Cytokine 1990;2:170-74.
- [12]. WunderinkRG,WoldenbergLS,Zeiss j, Day CM, Ciemins j, LacherDA.The radiologic diagnosis of autopsy-proven ventilator associated pneumonia.Chest 1992;101:458-63
- [13]. Fabregas N, EwigS, Torres A, El-Ebiary M, Ramirez J, de La BellacasaJP, BauerT, Cabello H:Clinical diagnosis of ventilator associated pneumonia revisite:comparitive validation using immediate post mortem lung biopsies, Thorax 1999, 54:867-73. Meduri GU, Mauldin GL, WunderinkRG, LeeperkvJr, Jones CB, Tolley E, MayhallG. Causes of fever and pulmonary densities in patients with clinical manifestations of ventilator-associated pneumonia. Chest 1994; 106:221-35.
- [14]. Am J RespirCrit Care Med 2005, 171:388-416. Guidelines for the management of adults with hospital-acquired, ventilatorassociated, and healthcare-associated pneumonia .Luna CM, Vujacich P, Niederman MS et al: Impact of BAL data on the therapy and outcome of ventilator-associated pneumonia Chest1997, 111:676-685. Iregui M, Ward S, Sherman G, Fraser VJ et al: Clinical importance of delays in the initiation of appropriate antibiotic treatment for ventilator associated pneumonia. Chest 2002, 122:262-268.
- [15]. Am J RespirCrit Care Med 2005, 171:388-416. Guidelines for the management of adults with hospital-acquired, ventilatorassociated, and healthcare-associated pneumonia Deeks JJ: Systematic reviews in health care: systematic reviews of evaluations of diagnostic and screening tests. BMJ2001, 323:157-162.Klompas M: Does this patient have ventilator-associated pneumonia? JAMA 2007, 297:1583-1593
- [16]. Dellinger RP, Levy MM, Carlet JM, Bion J et al :Surviving Sepsis Campaign guidelines for management of severe sepsis and septic shock. Crit Care Med 2004, 32:858-873

- [17]. Ventilator associated-pneumonia JAMA 2007;297:1616-1617
- [18]. Prevention of ventilator associated pneumonia:consultationNICE,Sept 2007,Ventilator associated pneumonia JAMA 2007;297:1616-17
- [19]. Haley RW, Hooton TM, Culver DH, Stanley RC, Emori TG, Hardison CD, Quade D, Shachtman RH, Schaberg DR, Shah BV, et al. Nosocomial infections in US hospitals, 1975–1976: estimated frequency by selected characteristics of patients. Am J Med 1981;70:947–959. Chastre J, Fagon JY. Pneumonia in the ventilator-dependent patient. In: Tobin MJ, editor. Principles and practice of mechanical ventilation. New York: McGraw-Hill; 1994. p. 857–890.
- [20]. Celis R, Torres A, Gatell JM, Almela M, Rodriguez-Roisin R, Agusti-Vidal A. Nosocomial pneumonia. A multivariate analysis of risk and prognosis. Chest 1988;93:318– 324. Torres A, Aznar R, Gatell JM, Jimenez P, Gonzalez J, Ferrer A, Celis R, Rodriguez-Roisin R. Incidence, risk, and prognosis factors of nosocomial pneumonia in mechanically ventilated patients. Am RevRespir Dis 1990;142:523–528
- [21]. Torres A, Aznar R, Gatell JM, Jimenez P, Gonzalez J, Ferrer A, Celis R, Rodriguez-Roisin R. Incidence, risk, and prognosis factors of nosocomial pneumonia in mechanically ventilated patients. Am RevRespir Dis 1990;142:523–528.
- [22]. Strausbaugh L. Nosocomial respiratory infections. In: Mandell GL, Bennett JE, Dolin R, editors. Principles and practice of infectious diseases.Philadelphia, PA: Churchill Livingstone; 2000. p. 3020–3027.
- [23]. Corley DE, Kirtland SH, Winterbauer RH, Hammar SP, Dail DH, Bauermeister DE, Bolen JW. Reproducibility of the histologic diagnosis of pneumonia among a panel of four pathologists: analysis of a gold standard. Chest 1997;112:458–465.
- [24]. Marquette CH, Copin MC, Wallet F, Neviere R, Saulnier F, Mathieu D, Durocher A, Ramon P, Tonnel AB. Diagnostic tests for pneumonia in ventilated patients: prospective evaluation of diagnostic accuracy using histology as a diagnostic gold standard. Am J RespirCrit Care Med 1995;151:1878– 1888.
- [25]. Rouby JJ, Martin De Lassale E, Poete P, Nicolas MH, Bodin L, Jarlier V, Le Charpentier Y, Grosset J, Viars P. Nosocomial bronchopneumonia in the critically ill. His-



tologic and bacteriologic aspects. Am Rev Respir Dis 1992;146:1059–1066.

- [26]. Kirtland SH, Corley DE, Winterbauer RH, Springmeyer SC, Casey KR, Hampson NB, Dreis DF. The diagnosis of ventilatorassociated pneumonia: a comparison of histologic, microbiologic, and clinical criteria. Chest 1997;112:445–457.
- [27]. Langer M, Cigada M, Mandelli M, Mosconi P, Tognoni G. Early onset pneumonia: a multicenter study in intensive care units. Intensive Care Med 1987;13:342–346.
- [28]. American Thoracic Society. Hospitalacquired pneumonia in adults: diagnosis, assessment of severity, initial antimicrobial therapy, and preventive strategies. A consensus statement, American Thoracic Society,November 1995. Am J RespirCrit Care Med 1996;153:1711–1725
- [29]. Vincent JL, Bihari DJ, Suter PM, Bruining HA, White J, Nicolas-Chanoin MH, Wolff M, Spencer RC, Hemmer M. The prevalence of nosocomial infection in intensive care units in Europe. Results of the European Prevalence of Infection in Intensive Care (EPIC) Study.EPIC International Advisory Committee. JAMA 1995;274:639–644.
- [30]. Chevret S, Hemmer M, Carlet J, Langer M. Incidence and risk factors of pneumonia acquired in intensive care units. Results from a multicenter prospective study on 996 patients. European Cooperative Group on Nosocomial Pneumonia. Intensive Care Med 1993;19:256–264
- [31]. Cross AS, Roup B. Role of respiratory assistance devices in endemic nosocomial pneumonia. Am J Med 1981;70:681–685
- [32]. Cook DJ, Walter SD, Cook RJ, Griffith LE, Guyatt GH, Leasa D, Jaeschke RZ, Brun-Buisson C. Incidence of and risk factors for ventilator-associated pneumonia in critically ill patients. Ann Intern Med 1998;129:433– 440
- [33]. Bell RC, Coalson JJ, Smith JD, Johanson WG. Multiple organ systemfailure and infection in adult respiratory distress syndrome.Ann InternMed 1983;99:293–298
- [34]. Andrews CP, Coalson JJ, Smith JD, Johanson WG. Diagnosis of nosocomial bacterial pneumonia in acute, diffuse lung injury. Chest 1981;80:254–258.
- [35]. Martin TR, Pistorese BP, Hudson LD, Maunder RJ. The function of lungand blood neutrophils in patients with the adult respiratory distresssyndrome. Implications for the

pathogenesis of lung infections.Am Rev Respir Dis 1991;144:254–262.

- [36]. Chollet-Martin S, Jourdain B, Gibert C, Elbim C, Chastre J, Gougerot-Pocidalo MA. Interactions between neutrophils and cytokines inblood and alveolar spaces during ARDS.Am J RespirCrit Care Med1996;154:594–601.
- [37]. Chastre J, Trouillet JL, Vuagnat A, Joly-Guillou ML, Clavier H, DombretMC, Gibert C. Nosocomial pneumonia in patients with acute respiratorydistress syndrome. Am J RespirCrit Care Med 1998;157: 1165–1172.
- [38]. Stevens RM, Teres D, Skillman JJ, Feingold DS. Pneumonia in an intensivecare unit.A 30-month experience. Arch Intern Med 1974;134:106–111.
- [39]. Fagon JY, Chastre J, Domart Y, Trouillet JL, Pierre J, Darne C, Gibert C. Nosocomial pneumonia in patients receiving continuous mechanicalventilation.Prospective analysis of 52 episodes with use of a protectedspecimen brush and quantitative culture techniques. Am Rev Respir Dis1989;139:877–884.
- [40]. Kollef MH, Silver P, Murphy DM, Trovillion E. The effect of late-onsetventilatorassociated pneumonia in determining patient mortality.Chest 1995;108:1655–1662.
- [41]. Daren et al The Attributable Morbidity and Mortality of Ventilator-Associated Pneumonia in the Critically III Patient.Am J Respir-Crit Care Med 1999 Vol 159.1249–1256
- [42]. Rello J, Ausina V, Ricart M, Castella J, Prats G. Impact of previous antimicrobial therapy on the etiology and outcome of ventilator-associated pneumonia. Chest 1993;104:1230–1235.
- [43]. Horan TC, Culver DH, Gaynes RP, Jarvis WR, Edwards JR, Reid CR.Nosocomial infections in surgical patients in the United States, January 1986–June 1992.National Nosocomial Infections Surveillance (NNIS) System. Infect Control HospEpidemiol 1993; 14:73–80.
- [44]. Spencer RC. Predominant pathogens found in the European Prevalence of Infection in Intensive Care Study. Eur J ClinMicrobiol Infect Dis 1996; 15:281–285.
- [45]. Bryan CS, Reynolds KL. Bacteremic nosocomial pneumonia. Analysis of 172 episodes from a single metropolitan area. Am Rev Respir Dis 1984;129:668–671
- [46]. Baker AM, Meredith JW, Haponik EF. Pneumonia in intubated trauma pa-



tients.Microbiology and outcomes.Am J RespirCrit Care Med 1996; 153:343–349.

- [47]. Antonelli M, Moro ML, Capelli O, De Blasi RA, D'Errico RR, Conti G, Bufi M, Gasparetto A. Risk factors for early onset pneumonia in trauma patients. Chest 1994;105:224–228.
- [48]. Rello J, Torres A, Ricart M, Valles J, Gonzalez J, Artigas A, Rodriguez- Roisin R. Ventilator-associated pneumonia by Staphylococcus aureus. Comparison of methicillinresistant and methicillin-sensitive episodes. Am J RespirCrit Care Med 1994;150:1545– 1549
- [49]. Singh N, Falestiny MN, Rogers P, Reed MJ, Pularski J, Norris R, Yu VL. Pulmonary infiltrates in the surgical ICU: prospective assessment predictors of etiology and mortality. Chest 1998;114:1129–1136
- [50]. Chastre J, Trouillet JL, Vuagnat A, Joly-Guillou ML, Clavier H, Dombret MC, Gibert C. Nosocomial pneumonia in patients with acute respiratory distress syndrome. Am J RespirCrit Care Med 1998;157: 1165–1172
- [51]. Delclaux C, Roupie E, Blot F, Brochard L, Lemaire F, Brun-Buisson C. Lower respiratory tract colonization and infection during severe acute respiratory distress syndrome: incidence and diagnosis. Am J RespirCrit Care Med 1997;156:1092–1098
- [52]. Markowicz P, Wolff M, Djedaini K, Cohen Y, Chastre J, Delclaux C, Merrer J, Herman B, Veber B, Fontaine A, et al. Multicenter prospective study of ventilator-associated pneumonia during acute respiratory distress syndrome. Incidence, prognosis, and risk factors. ARDS Study Group. Am J Respir-Crit Care Med 2000;161:1942–1948
- [53]. Hofkan G et al, Nosocomial pneumonia: the importance of deescalating strategy for antibiotic treatment of pneumonia in the ICU.Chest 2002;122:2183
- [54]. Ibrahim EH, Ward S, Sherman G, Kollef MH. A comparative analysis of patients with early-onset vs late-onset nosocomial pneumonia in the ICU setting. Chest 2000;117:1434–1442.
- [55]. Kirby BD, Snyder KM, Meyer RD, Finegold SM. Legionnaires' disease: report of sixtyfive nosocomially acquired cases of review of the literature. Medicine (Baltimore) 1980;59:188–205.
- [56]. Girod JC, Reichman RC, Winn WC, Klaucke DN, Vogt RL, Dolin R. Pneumonic and nonpneumonic forms of legionellosis.

The result of a common-source exposure to Legionella pneumophila. Arch Intern Med 1982;142:545–547

- [57]. Dore P, Robert R, Grollier G, Rouffineau J, Lanquetot H, Charriere JM, Fauchere JL. Incidence of anaerobes in ventilator-associated pneumonia with use of a protected specimen brush. Am J RespirCrit Care Med 1996;153:1292–1298.
- [58]. el-Ebiary M, Torres A, Fabregas N, de la Bellacasa JP, Gonzalez J, Ramirez J, del Bano D, Hernandez C, Jimenez de Anta MT. Significance of the isolation of Candida species from respiratory samples in critically ill, non-neutropenic patients. An immediate postmortem histologic study. Am J Respir-Crit Care Med 1997;156:583–590.
- [59]. Papazian L, Fraisse A, Garbe L, Zandotti C, Thomas P, Saux P, PierrinG, Gouin F. Cytomegalovirus. An unexpected cause of ventilator associated pneumonia. Anesthesiology 1996;84:280–287
- [60]. Marik PE, Careau P. The role of anaerobes in patients with ventilator associated pneumonia and aspiration pneumonia: a prospective study.Chest 1999;115:178–183.
- [61]. Ramsey PG, Fife KH, Kackman RC, Meyers JD, Corey L. Herpes simplex virus pneumonia: clinical, virologic, and pathologic features in 20 patients. Ann Intern Med. 1982; 97:813-820
- [62]. Bruynseels P, Jorens PG, Demey HE, et al. Herpes simplex virus in the respiratory tract of critical care patients: a prospective study. Lancet. 2003;362:1536-1541
- [63]. Heininger A, Jahn G, Engel C, Notheisen T, Unertl K, Hamprecht K. Human cytomegalovirus infections in nonimmunosuppressed critically ill patients. Crit Care Med. 2001;29:541-547.
- [64]. Park DR. The microbiology of ventilatorassociated pneumonia. Respir Care. 2005;50:742-763.
- [65]. Hanson LC, Weber DJ, Rutala WA, Samsa GP: Risk factors for nosocomial pneumonia in elderly. Am J Med 1992, 92:161-166.
- [66]. Langer M, Mosconi P, Cigada M, Mandelli M: Long-term respiratory support and risk of pneumonia in critically ill patients. Am Rev Respir Dis 1989, 140:302-305.
- [67]. Chevret S, Hemmer M, Carlet J, Langer M: Incidence and risk factors of pneumonia acquired in intensive care units. Results from a multicenter prospective study on 996 patients. European Cooperative Group on No-



socomial Pneumonia. Intensive Care Med 1993, 19:256-264

- [68]. Weber DJ, Rutala WA: Nosocomial infections associated with respiratory therapy. In Hospital Epidemiology and Infection Control 3rd edition. Edited by: Mayhall CG. Baltimore: Williams & Wilkins; 1996:748-758.
- [69]. Adair CG, Gorman SP, Feron BM, Byers LM, Jones DS, Goldsmith CE,Moore JE, Kerr JR, Curran MD, Hogg G, Webb CH, McCarthy GJ, Milligan KR: Implications of endotracheal tube biofilm for ventilator- associated pneumonia. Intensive Care Med 1999,25:1072-1076.
- [70]. Garibaldi RA, Britt MR, Coleman ML, Reading JC, Pace NL. Risk factors for postoperative pneumonia. Am J Med 1981;70:677–680.
- [71]. Cunnion KM, Weber DJ, Broadhead WE, Hanson LC, Pieper CF, Rutala WA. Risk factors for nosocomial pneumonia: comparing adult criticalcare populations. Am J RespirCrit Care Med 1996;153:158–162.
- [72]. Kollef MH, Wragge T, Pasque C. Determinants of mortality and multiorgan dysfunction in cardiac surgery patients requiring prolonged mechanical ventilation. Chest 1995;107:1395–1401
- [73]. Rello J, Sonora S, Jubert P, Artigas A, Rue M, Valles J: Pneumonia in intubated patients: role of respiratory airway care. Am J RespirCrit Care Med 1996, 154:111-115.
- [74]. Rello J, Diaz E, Roque M, Valles J: Risk factors for developing pneumonia within 48 hours of intubation. Am J RespirCrit Care Med 1999, 159:1742-1746.
- [75]. Sirvent JM, Torres A, El-Ebiary M, Castro P, de Batile J, Bonet A: Protective effect of intravenously administered cefuroxime against nosocomial pneumonia in patients with structural coma. Am J RespirCrit Care Med 1997, 155:1729-1734.
- [76]. Kollef MH: Ventilator-associated pneumonia: A multivariate analysis. JAMA 1993, 270:1965-1970
- [77]. Rello J, Ausina V, Ricart M, Puzo C, Quintana E, Net A, Prats G: Risk factors for infection by Pseudomonas aeruginosa in patients with ventilator associated pneumonia. Intensive Care Med 1994, 20:193-198.
- [78]. Atherton ST, White DJ. Stomach as source of bacteria colonising respiratory tract during artificial ventilation. Lancet 1978;2:968– 969.

- [79]. Hillman KM, Riordan T, O'Farrell SM, Tabaqchali S. Colonization of the gastric contents in critically ill patients. Crit Care Med 1982;10: 444–447.
- [80]. du Moulin GC, Paterson DG, Hedley-Whyte J, Lisbon A. Aspiration of gastric bacteria in antacid-treated patients: a frequent cause of postoperative colonisation of the airway. Lancet 1982;1:242–245.
- [81]. Donowitz LG, Page MC, Mileur BL, Guenthner SH. Alteration of normal gastric flora in critical care patients receiving antacid and cimetidine therapy. Infect Control 1986;7:23–26.
- [82]. George DL: Nosocomial pneumonia. In Hospital Epidemiology and Infection Control 3rd edition. Edited by: Mayhall CG. Baltimore: Williams & Wilkins; 1996:175-195.
- [83]. Leal-Noval SR, Marquez-Vacaro JA, Garcia-Curiel A, Camacho-Larana P, Rincon-Ferrari MD, Ordonez-Fernandez A, Flores-Cordero JM, Loscertales-Abril J: Nosocomial pneumonia in patients undergoing heart surgery. Crit Care Med 2000, 28:935-940.
- [84]. Spray SB, Zuidema GD, Cameron JL. Aspiration pneumonia; incidence of aspiration with endotracheal tubes. Am J Surg 1976;131:701–703.
- [85]. Sirvent JM, Torres A, Vidaur L, Armengol J, de Batlle J, Bonet A. Tracheal colonisation within 24 h of intubation in patients with head trauma: risk factor for developing early-onset ventilator-associated pneumonia. Intensive Care Med 2000;26:1369–1372
- [86]. Valles J, Artigas A, Rello J, Bonsoms N, Fontanals D, Blanch L, Fernandez R, Baigorri F, Mestre J. Continuous aspiration of subglottic secretions in preventing ventilator-associated pneumonia. Ann Intern Med 1995;122:179–186.
- [87]. Torres A, Gatell JM, Aznar E, el-Ebiary M, Puig de la Bellacasa J, Gonzalez J, Ferrer M, Rodriguez-Roisin R. Re-intubation increases the risk of nosocomial pneumonia in patients needing mechanical ventilation. Am J RespirCrit Care Med 1995;152:137–141.
- [88]. Brook AD, Sherman G, Malen J, Kollef MH. Early versus late tracheostomy in patients who require prolonged mechanical ventilation. Am J Crit Care 2000;9:352–359.
- [89]. Lesnik I, Rappaport W, Fulginiti J, Witzke D. The role of early tracheostomy in blunt, multiple organ trauma. Am Surg 1992; 58:346–349.



- [90]. Rodriguez JL, Steinberg SM, Luchetti FA, Gibbons KJ, Taheri PA, Flint LM. Early tracheostomy for primary airway management in the surgical critical care setting. Surgery 1990;108:655–659.
- [91]. Moore FA, Moore EE, Jones TN, McCroskey BL, Peterson VM. TEN versus TPN following major abdominal trauma: reduced septic morbidity. [Discussion p. 922– 913.] J Trauma 1989;29:916–922.
- [92]. Heyland DK, Cook DJ, Schoenfeld PS, Frietag A, Varon J, Wood G. The effect of acidified enteral feeds on gastric colonization in critically ill patients: results of a multicenter randomized trial. Canadian Critical Care Trials Group. Crit Care Med 1999;27:2399– 2406.
- [93]. Winterbauer RH, Durning RB, Barron E, McFadden MC. Aspirated nasogastric feeding solution detected by glucose strips. Ann Intern Med 1981;95:67–68.
- [94]. Strong RM, Condon SC, Solinger MR, Namihas BN, Ito-Wong LA, Leuty JE. Equal aspiration rates from postpylorus and intragastricplaced small-bore nasoenteric feeding tubes: a randomized, prospective study. J Parenter Enteral Nutr 1992;16:59–63.
- [95]. Montecalvo MA, Steger KA, Farber HW, Smith BF, Dennis RC, Fitzpatrick GF, Pollack SD, Korsberg TZ, Birkett DH, Hirsch EF, et al. Nutritional outcome and pneumonia in critical care patients randomized to gastric versus jejunal tube feedings. The Critical Care Research Team. Crit Care Med 1992;20:1377–1387.
- [96]. Craven DE, Kunches LM, Kilinsky V, Lichtenberg DA, Make BJ, McCabe WR. Risk factors for pneumonia and fatality in patients receiving continuous mechanical ventilation. Am Rev Respir Dis 1986;133:792–796
- [97]. Craven DE, Lichtenberg DA, Goularte TA, Make BJ, McCabe WR. Contaminated medication nebulizers in mechanical ventilator circuits. Source of bacterial aerosols. Am J Med 1984;77:834–838
- [98]. Andrews CP, Coalson JJ, Smith JD, Johanson WG. Diagnosis of nosocomial bacterial pneumonia in acute, diffuse lung injury. Chest 1981;80:254–258.
- [99]. Wunderink RG. Radiologic diagnosis of ventilator-associated pneumonia. Chest 2000;117:188S–190S
- [100]. Meduri GU, Belenchia JM, Estes RJ, Wunderink RG, el Torky M, Leeper KV Jr. Fibroproliferative phase of ARDS. Clinical

findings and effects of corticosteroids. Chest 1991;100:943–952.

- [101]. Villers D, Derriennic M, Raffi F, Germaud P, Baron D, Nicolas F, Courtieu AL. Reliability of the bronchoscopic protected catheter brush in intubated and ventilated patients. Chest 1985;88:527–530.
- [102]. Lambert RS, Vereen LE, George RB. Comparison of tracheal aspirates and protected brush catheter specimens for identifying pathogenic bacteria in mechanically ventilated patients. Am J Med Sci 1989;297:377–382.
- [103]. Baselski V. Microbiologic diagnosis of ventilator-associated pneumonia. Infect Dis Clin North Am 1993;7:331–357.
- [104]. Pugin J, Auckenthaler R, Mili N, Janssens JP, Lew PD, Suter PM: Diagnosis of ventilator-associated pneumonia by bacteriologic analysis of bronchoscopic and non-bronchoscopic "blind" broncoalveolar lavage fluid. Am Rev Respir Dis 1991, 143:1121-1129.
- [105]. Papazian L, Thomas P, Garbe L, Guignon I, Thirion X, Charrel J, Bollet C, Fuentes P, Gouin F: Bronchoscopic or blind sampling techniques for the diagnosis of ventilatorassociated pneumonia. Am J RespirCrit Care Med 1995, 152:1982-1991
- [106]. Torres A, Puig de la Bellacasa J, Xaubet A, Gonzalez J, Rodriguez-Roisin R, Jimenez de Anta MT, Agusti Vidal A: Diagnostic value of quantitative cultures of bronchoalveolar lavage and telescoping plugged catheters in mechanically ventilated patients with bacterial pneumonia. Am Rev Respir Dis 1989,140:306-310.
- [107]. Chastre J, Fagon JY, Soler P, Bornet M, Domart Y, Trouillet JL, Gibert C, Hance AJ: Diagnosis of nosocomial bacterial pneumonia in intubated patients undergoing ventilation: comparison of the usefulness of bronchoalveolar lavage and the protected specimen brush. Am J Med 1988, 85:499-506.
- [108]. el-Ebiary M, Torres A, Gonzalez J, de la Bellacasa JP, Garcia C, Jimenez de Anta MT, Ferrer M, Rodriguez-Roisin R: Quantitative cultures of endotracheal aspirates for the diagnosis of ventilator associated pneumonia. Am Rev Respir Dis 1993, 103:547-553.
- [109]. Marquette CH, Georges H, Wallet F, Ramon P, Saulnier F, Neviere R, Mathieu D, Rime A, Tonnel AB: Diagnostic aspirates with quantitative bacterial cultures in intubated patients with suspected pneumonia. Compar-



ison with the protected specimen brush. Am Rev Respir Dis 1993, 148:138-144.

- [110]. Marquette CH, Copin MC, Wallet F, Neviere R, Saulnier F, Mathieu D, Durocher A, Ramon P, Tonnel AB: Diagnostic tests for pneumonia in ventilated patients: prospective evaluation of diagnostic accuracy using histology as a diagnostic gold standard. Am J RespirCrit Care Med 1995, 151:1878-1888.
- [111]. Jourdain B, Novara A, Joly-Guillou ML, Dombret MC, Calvat S, Trouillet JL, Gibert C, Chastre J: Role of quantitative cultures of endotracheal aspirates in the diagnosis of nosocomial pneumonia. Am J RespirCrit Care Med 1995, 152:241-246.
- [112]. Cook D, Mandell L: Endotracheal aspiration in the diagnosis of ventilator-associated pneumonia. Chest 2000, 117:195-197.
- [113]. Valencia Arango M, Torres Marti A, InsaustiOrdenana J, Alvarez Lerma F, Carrasco Joaquinet N, HerranzCasado M, Tirapu Leon JP, Grupo de Estudio de la NeumoniaRelacionada con VentilacionMecanica; Grupo de Trabajo de EnfermedadesInfecciosas de la SEMICYUC: Diagnostic value of quantitative cultures of endotracheal aspirate in ventilator-associated pneumonia: a muticenter study. Arch Bronconeumol 2003, 39:394-399.
- [114]. Baughman RP, Thorpe JE, Staneck J, Rashkin M, Frame PT: Use of the protected specimen brush in patients with endotracheal or tracheostomy tubes. Chest 1987, 91:233-236.
- [115]. Richard C, Pezzano M, Bouhaja B, Rottman E, Rimailho A, Riou B, Auzepy P: Comparison of non-protected lower respiratory tract secretions and protected specimen brush samples in the diagnosis of pneumonia. Intensive Care Med 1988, 14:30-33.
- [116]. Villers D, Derriennic M, Raffi F, Germaud P, Baron D, Nicolas F, Courtieu AL: Reliability of the bronchoscopic protected catheter brush in intubated and ventilated patients. Chest 1988,4:527-530.
- [117]. Chastre J, Fagon JY: Invasive diagnostic testing should be routinely used to manage suspected pneumonia in mechanically ventilated patients. Am J RespirCrit Care Med 1994, 150:570-574.
- [118]. Croce MA: Diagnosis of Acute Respiratory Distress Syndrome and differentiation from ventilator-associated pneumonia. Am J Surg 2000:26-30.

- [119]. Baselski VS, el-Torky M, Coalson JJ, Griffin JP: The standardization of criteria for processing and interpreting laboratory specimens in patients with suspected ventilatorassociated pneumonia. Chest 1992, 102:571-579.
- [120]. Chastre J, Viau F, Brun P, Pierre J, Dauge MC, Bouchama A, Akesbi A, Gibert C: Prospective evaluation of the protected specimen brush for the diagnosis of pulmonary infections in ventilated patients. Am Rev Respir Dis 1984, 130:924-929.
- [121]. Sanchez-Nieto JM, Torres A, Garcia-Cordoba F, El-Ebiary M, Carrillo A, Ruiz J, Nunez ML, Niederman M: Impact of invasive and noninvasive quantitative culture sampling on outcome of ventilator- associated pneumonia. Am J RespirCrit Care Med 1998, 157:371-376.
- [122]. Heyland DK, Cook DJ, Marshall J, Heule M, Guslits B, Lang J, Jaeschke R: The clinical utility of invasive diagnostic techniques in the setting of ventilator-associated pneumonia. Chest 1999, 115:1076-1084.
- [123]. Shorr AF, Sherner JH, Jackson WL, Kollef MH: Invasive approaches to the diagnosis of ventilator -associated pneumonia: A metaanalysis. Crit Care Med 2005, 33:46-53.
- [124]. Niederman MS, Torres A, Summer W: Invasive diagnostic testing is not routinely manage suspected ventilator-associated pneumonia. Am J RespirCrit Care Med 1994, 150:565-569.
- [125]. Torres A, el-Ebiary M, Padro L, Gonzalez J, de la Bellacasa JP, Ramirez J, Xaubet A, Ferrer M, Rodriguez-Roisin R: Validation of different techniques for the diagnosis of ventilator-associated pneumonia. Am J Respir-Crit Care Med 1994, 149:324-331.
- [126]. Souweine B, Veber B, Bedos JP, Gachot B, Dombret MC, Regnier B, Wolff M: Diagnostic accuracy of protected specimen brush and bronchoalveolar lavage in nosocomial pneumonia: impact of previous antimicrobial treatments. Crit Care Med1998, 26:236-244.
- [127]. Safdar N, Abad C. Educational interventions for prevention of healthcare-associated infection: a systematic review. Crit Care Med 2008; 36: 933-940
- [128]. Salahuddin N, Zafar A, Sukhyani L, Rahim S, Noor MF, Hussain K et al. reducing ventilator-associated pneumonia rates through a staff education programme. J Hosp Infect 2004; 57: 223-227.
- [129]. Babcock HM, Zack JE, Garrison T, Trovillion E, Jones M, Fraser VJ, et al. An educa-



tional intervention to reduce ventilatorassociated pneumonia in an integrated health system: a comparison of effects. Chest 2004; 125: 2224-2231.

- [130]. Pittet D, Allegranzi B, Sax H, Dharan S, Pessoa-Silva CL, Donaldson L et al. Evidence-based model for hand transmission during patient care and the role of improved practices. Lancet Infect Dis 2006; 6: 641-652.
- [131]. Pratt RJ, Pellowe CM, Wilson JA, Loveday HP, Harper PJ, Jones SR et al. epic2: National evidence-based guidelines for preventing healthcare-associated infections in NHS hospitals in England. J Hosp Infect 2007; 65 (Suppl 1): 1-64.
- [132]. Crnich CJ, Safdar N, Maki DG. The role of the intensive care unit environment in the pathogenesis and prevention of ventilatorassociated pneumonia. Respir Care 2005; 50: 813-836
- [133]. Safdar N, Crnich CJ, Maki DG. The pathogenesis of ventilator-associated pneumonia: its relevance to developing effective strategies for prevention. Respir Care 2005; 50: 725-739
- [134]. SARI. Infection prevention and control building guidelines for acute hospitals in Ireland. 2009.
- [135]. Humphreys H, Johnson EM, Warnock DW, Willatts SM, Winter RJ, Speller DCE. An outbreak of aspergillosis in a general ITU. J Hosp Infect 1991; 18:167-177.
- [136]. Flynn PM, Williams BG, Hetherington SV, Williams BF, Giannini MA, Pearson TA. Aspergillusterreus during hospital renovation. Infect Control HospEpidemiol 1993; 14: 363-365.
- [137]. NDSC. National guidelines for the prevention of nosocomial invasive aspergillosis during construction / renovation activities. 2002.
- [138]. Stone PW, Pogorzelska M, Kunches L, Hirschhorn LR. Hospital staffing and health care-associated infections: a systematic review of the literature. Clin Infect Dis 2008; 47: 937-944.
- [139]. Hugonnet S, Chevrolet JC, Pittet D. The effect of workload on infection risk in critically ill patients. Crit Care Med 2007; 35: 76-81.
- [140]. Hugonnet S, Uçkay I, Pittet D. Staffing level: a determinant of late-onset ventilatorassociated pneumonia. Crit Care 2007; 11: R80.

- [141]. Pronovost PJ, Angus DC, Dorman T, Robinson KA, Dremsizov TT, Young TL. Physician staffing patterns and clinical outcomes in critically ill patients: a systematic review. JAMA 2008; 288: 2151-2162.
- [142]. Ram FS, Picot J, Lightowler J, Wedzicha JA. Non-invasive positive pressure ventilation for treatment of respiratory failure due to exacerbations of chronic obstructive pulmonary disease. Cochrane Database Syst Rev 2004; CD004104.
- [143]. Burns KE, Adhikari NK, Keenan SP, Meade M. Use of non-invasive ventilation to wean critically ill adults off invasive ventilation: meta-analysis and systematic review. BMJ 2009; 338: b1574
- [144]. Torres A, Gatell JM, Aznar E, el-Ebiary M, Puig de la Bellacasa J, González J et al. Reintubation increases the risk of nosocomial pneumonia in patients needing mechanical ventilation. Am J RespirCrit Care Med 1995; 152: 137-141.
- [145]. Kollef MH, Shapiro SD, Silver P, St John RE, Prentice D, Sauer S et al. A randomised, controlled trial of protocol directed versus physician-directed weaning from mechanical ventilation. Crit Care Med 1997; 25: 567-574.
- [146]. Marelich GP, Murin S, Battistella F, Inciardi J, Vierra T, Roby M. Protocol weaning of mechanical ventilation in medical and surgical patients by respiratory care practitioners and nurses: effect on weaning time and incidence of ventilator-associated pneumonia. Chest 2000; 118: 459-467.
- [147]. Brook AD, Ahrens TS, Schaiff R, Prentice D, Sherman G, Shannon W et al. Effect of a nursing-implemented sedation protocol on the duration of mechanical ventilation. Crit Care Med 1999; 27: 2609-2615.
- [148]. Quenot JP, Ladoire S, Devoucoux F, Doise JM, Cailliod R, Cunin N et al. Effect of a nurse-implemented sedation protocol on the incidence of ventilator-associated pneumonia. Crit Care Med 2007; 35: 2031-2036
- [149]. Hébert PC, Wells G, Blajchman MA, Marshall J, Martin C, Pagliarello G et al. A multicenter, randomized, controlled clinical trial of transfusion requirements in critical care. Transfusion Requirements in Critical Care Investigators, Canadian Critical Care Trials Group. N Engl J Med 1999; 340: 409-417.
- [150]. Shorr AF, Duh MS, Kelly KM, Kollef MH. Red blood cell transfusion and ventilatorassociated pneumonia: A potential link? Crit Care Med 2004; 32: 666-674.

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- [151]. 42. Bochicchio GV, Napolitano L, Joshi M, Bochicchio K, Shih D, Meyer W et al. Blood product transfusion and ventilator-associated pneumonia in trauma patients. Surg Infect (Larchmt) 2008; 9: 415-422.
- [152]. Siempos II, Ntaidou TK, Falagas ME. Impact of the administration of probiotics on the incidence of ventilator associated pneumonia: a meta-analysis of randomised control trials. Crit Care Med 2010; 38: 954-962.
- [153]. Resar R, Pronovost P, Haraden C, Simmonds T, Rainey T, Nolan T. Using a bundle approach to improve ventilator care processes and reduce ventilator-associated pneumonia. JtComm J Qual Patient Saf 2005; 31: 243-248.
- [154]. Hawe CS, Ellis KS, Cairns CJ, Longmate A. Reduction of ventilator-associated pneumonia: active versus passive guideline implementation. Intensive Care Med 2009; 35: 1180-1186.
- [155]. Shorr AF, O'Malley PG. Continuous Subglottic Suctioning for the Prevention of Ventilator-Associated Pneumonia; Potential Economic Implications. Chest 2001; 119:228-235
- [156]. Kollef MH, Skubas NJ, Sundt TM. A Randomized Clinical Trial of Continuous Aspiration of Subglottic Secretions in Cardiac Surgery Patients. Chest 1999; 116:1339-1346.
- [157]. Smulders K, van der Hoeven H, Weers-Pothoff I, et al. A Randomized Clinical Trial of Intermittent Subglottic Secretion Drainage in Patients Receiving Mechanical Ventilation. Chest 2002; 121:858-862.
- [158]. Boyce JM, Pittet D. Guideline for Hand Hygiene in Health-Care Settings.Recommendations of the Healthcare Infection Control Practices Advisory Committee and the HICPAC/SHEA/APIC/IDSA Hand Hygiene Task Force. Society for Healthcare Epidemiology of America/Association for Professionals in Infection Control/Infectious Diseases Society of America. MMWR Recomm Rep 2002;51 (RR-16): 1-45.
- [159]. Kress JP, Pohlman AS, O'Connor M, Hall JB. Daily interruption of sedative infusions in critically ill patients undergoing mechanical ventilation. N Engl J Med 2000; 342:1471-77.
- [160]. Geerts WH, Pineo GF, Heit JA, et al. Prevention of venous Thromboembolism: The Seventh ACCP Conference on Antithrom-

botic and Thrombolytic Therapy. Chest. Sep 2004; 126(3):338S-400S

- [161]. Collard HR, Saint S, Matthay MA, Prevention of Ventilator-Associated Pneumonia: An Evidence Based Systematic Review. Ann Intern Med 2003; 138:494-501.
- [162]. DeRiso AJ II, Ladowski JS, Dillon TA, Justice JW, Peterson AC. Chlorhexidine gluconate 0.12% oral rinse reduces the incidence of total nosocomial respiratory infection and nonprophylactic systemic antibiotic use in patients under-going heart surgery. Chest.1996;109:1556-1561.
- [163]. Koeman M, van der Ven AJ, Hak E, et al. Oral decontamina-tion with chlorhexidine reduces the incidence of ventilatorassociated pneumonia. Am J Respir Crit Care Med.2006;173: 1348-1355
- [164]. Vardakaz KZ, Siempos II, Falaqas ME Diabet Med 2007 Oct 24(10):1168-71
- [165]. S. Bernard, P. LeBlanc, F. Whittom et al., "Peripheral muscle weakness in patients with chronic obstructive pulmonary disease," American Journal of Respiratory and Critical CareMedicine, vol. 158, no. 2, pp. 629–634, 1998.
- [166]. E. F. M. Wouters, "Chronic obstructive pulmonary disease. 5: systemic effects of COPD," Thorax, vol. 57, supplement 12, pp.1067–1070, 2002.
- [167]. P. J. Barnes and B. R. Celli, "Systemic manifestations and comorbidities of COPD," European Respiratory Journal, vol.33, no. 5, pp. 1165–1185, 2009
- [168]. L. A. Callahan and G. S. Supinski, "Sepsisinduced myopathy," Critical Care Medicine, vol. 37, no. 10, pp. S354–S367, 2009
- [169]. Joshi N, Localio AR, Hamory BH. A predictive risk index for nosocomial pneumonia in the intensive care unit. Am J Med 1992;93:135–142.
- [170]. Li J, Kudsk KA, Gocinski B, Dent D, Glezer J, Langkamp-Hengen B, et al. Effects of parenteral and enteral nutrition on gutassociated lymphoid tissue. J Trauma 1995;39:44-51]
- [171]. Grau T, Bonet A, Minambres E, Pineiro L, Irles JA, Robles A, et al. The effect of Lalanyl-L-glutamine dipeptide supplemented total parenteral nutrition on infectious morbidity and insulin sensitivity in critically ill patients. Crit Care Med 2011;39: 1263
- [172]. Kollef MH, Von Harz B, Prentice D, Shapiro SD, Silver P, St John R, et al. Patient transport from intensive care increases the



risk of developing ventilator associate pneumonia. Chest 1997;112:765-73

- [173]. Acinetbacter-AA-Baribarj, correaH, MariscalD, GallegoM, VallesJ, RelloJ.Risk factors for infection by acinetobacterbaumanii in intubated patients with nosocomial pneumonia.Chest 1997;112:1050-4
- [174]. Husni RN, Goldstein LS, Arroliga AC, Hall GS, Fatica C, Stoller JK et al.Risk factor for an outbreak of multi-drug-resistant cinetobacter> nosocomialpneumoniaamong intubated patients. Chest. 1999; 115:1378-82
- [175]. AACN practice alert: ventilator-associated pneumonia. AACN Clin Issues.2005; 16(1):105-109]
- [176]. Kollef MH. Prevention of hospitalassociated pneumonia andventilatorassociated pneumonia.Crit Care Med. 2004;32:1396-1405
- [177]. Munro CL, Grap MJ. Oral health and care in the intensive care unit: state of the science. Am J Crit Care.2004;13:25-33.
- [178]. De Riso AJ Jr, Ladowski JS, Dillon TA, Justice JW, Peterson AC. Chlorhexidine gluconate 0.12% oral rinse reduces the incidence of total nosocomial respiratory infec-

tion and nonprophylactic systemic antibiotic use in patients undergoing heart surgery. Chest. 1996;109(6):1556-1561

- [179]. Donowitz LG, Page MC, Mileur BL, Guenther SH. Alteration of normal gastricflora in critical care patients receiving antacid and cimetidine therapy. Am JInfect Control. 1986;7:23-26.
- [180]. Cook D, Guyatt G, Marshall J, et al. A comparison of sucralfate and ranitidine for the prevention of upper gastrointestinal bleeding in patientsrequiring mechanical ventilation. Canadian Critical Care Trials Group.NEnglJ Med. 1998;338(12):791-797.
- [181]. Cook DJ, Fuller HD, Guyatt GH, et al. Risk factors for gastrointestinal bleeding in critically ill patients. Canadian Critical Care Trials Group.N Engl J Med. 1994;330(6): 377-381
- [182]. Col <u>Shivinder Singh</u>, Air Cmde <u>R. Chaturvedi</u>, Brig <u>S.M. Garg</u>, Col <u>Rashmi Datta</u>, Maj <u>Ambikesh Kuma</u>r. Incidenceof healthcare associated infection in the surgical ICU of a tertiary care hospital. MJAFI 2013;69:124-129