Antibiotic consumption and ventilator-associated pneumonia rates, some parallelism but some discrepancies

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Abstract: Ventilator-associated pneumonia (VAP) is a common infection in intensive care units (ICUs) but its clinical definition is neither sensitive nor specific and lacks accuracy and objectivity. New defining criteria were proposed in 2013 by the National Healthcare Safety Network (NHSN) in order to more accurately conduct surveillance and track prevention progress. Although there is a consistent trend towards a decrease in VAP incidence during the last decade, significant differences in VAP rates have been reported and are persistently lower in NHSN and other American reports (0.0 to 4.4 VAP per 1,000 ventilator-days in 2012) compared to the European Centre for Disease Prevention and Control (ECDC) data (10 VAP per 1,000 ventilator-days in 2014). In the United States, VAP has been proposed as an indicator of quality of care in public reporting, and the threat of financial penalties for this diagnosis has put pressure on hospitals to minimize VAP rates that may lead to artificial lower values, independently of patient care. Although prevention bundles may contribute for encouraging reductions in VAP incidence, both pathophysiologic and epidemiologic factors preclude a zero-VAP rate. It would be expected from the trend of reduction of VAP incidence that the consumption of antibiotics would also decrease in particular in those hospitals with lowest VAP rates. However, ICU reports show a steadily use of antibiotics for nosocomial pneumonia in 15% of patients and both ECDC and NHSN data on antibiotic consumption showed no significant trend. Knowledge of bacterial epidemiology and resistance profiles for each ICU has great relevance in order to understand trends of antibiotic use. The new NHSN criteria provide a more objective and quantitative data based VAP definition, including an antibiotic administration criterion, allowing, in theory, a more comprehensive assessment and a reportable benchmark of the observed VAP and antibiotic consumption variability.

Keywords: Antibiotic; intensive care unit (ICU); ventilator-associated pneumonia (VAP)

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Introduction

Ventilator-associated pneumonia (VAP) is one of the most common health care-associated infections (HAI) and accounts for 25% to 42% of all infections that occur in intensive care units (ICUs) (1-3). It can hit 10% to 25% of all ICU patients (4,5), being associated with longer duration of mechanical ventilation, longer ICU and hospital stays, increased healthcare costs and increased morbidity and mortality (6,7). VAP mortality rates range from 20% to 70% with healthcare costs of \$20,000 to \$40,000 per patient (4,8), making it a primary focus for risk-reduction strategies (9). In recent years preventive measures have been implemented and disseminated to reduce both its incidence

Variables	Criteria before 2013	Criteria after 2013
Respiratory status	New/worsening dyspnoea; purulent sputum; pulmonary infiltrates or consolidations on chest radiograph	≥0.20 increase in FiO ^{2^{†,‡}; ≥3 cmH₂O increase in PEEP^{†,‡}}
Infection/ inflammation	Temperature >38 °C; WBC ≥12,000 cells/mm ³ ; antimicrobial agent not required	Temperature <36 or >38 °C [§] ; WBC ≤4,000 or ≥12,000 cells/ $mm^{3\$}$; one or more new antimicrobial agent required and continued for ≥4 days [§]
Additional data	ETT/MV during the 48 h preceding the onset of pneumonia; positive culture (ETA $\ge 10^5$ CFU/mL; BAL $\ge 10^4$ CFU/mL) from appropriate specimen	Gram's staining of ETA or BAL showing ≥25 neutrophils and ≤10 epithelial cells per low-power field ¹ . Positive culture from: (I) ETA ≥10 ⁵ CFU/mL ^{\bullet} ; (II) BAL ≥10 ⁴ CFU/mL ^{\bullet}
+		+

 Table 1 VAP definition criteria evolution (10,17)

[†], must follow a period ≥ 2 days of stable or decreasing daily minimum PEEP or daily minimum FiO₂ and persist for ≥ 2 days; [‡], the presence of at least one of these respiratory parameters constitutes a VAC; [§], the presence of VAC plus one of these infection or inflammation parameters constitutes a IVAC; ¹, the presence of IVAC plus one of this parameter constitutes a possible pneumonia; [¢], the presence of IVAC with both a positive Gram's staining and a positive quantitative culture constitutes a probable pneumonia. WBC, white blood cell; ETT/MV, endotracheal intubation/mechanical ventilation; CFU, colony-forming unit; FiO₂, inspired fraction of oxygen; PEEP, positive end expiratory pressure.

and its consequences (10) and VAP has become a mean for tracking quality of care for ventilated patients (11).

VAP definition

Despite VAP epidemiological impact, it has no valid and reliable definition and even the most widely used VAP criteria are neither sensitive nor specific (6) and lack accuracy and objectivity (7,12,13), with documented poor correlation between the clinical diagnosis of VAP and histologic examination (14). Clinical scores have been applied but none was consistent in accurately identifying VAP (15). From 2002 onwards, the National Healthcare Safety Network (NHSN) criteria for VAP diagnosis include a combination of X-ray, signs or symptoms and laboratory data (16) (Table 1) in patients who were tracheally intubated and under mechanically ventilation at some time (with no minimal required amount of time) during the 48 h preceding the onset of pneumonia. Clinically, VAP can be subdivided into early and late onset: early onset VAP occurs within the first 4 days and late onset VAP after 4 days of mechanical ventilation (15,18). Both the subjectivity and variability inherent to chest X-ray technique and interpretation and to signs or symptoms recognition and proper documentation in medical records make these criteria unreliable to accurately identify VAP (6).

This is far from ideal to conduct surveillance and track prevention progress in an era of public reporting of HAI rates, benchmarking, inter-facility comparisons, and payfor-reporting and pay-for-performance programs. As an example, simply changing the microbiological method of diagnosis within the context of normal clinical practice influences VAP rates and antibiotic use. Morris *et al.* (19) compared the diagnosis of VAP in 53 patients with clinical suspicion using endotracheal aspirate (ETA) and bronchoalveolar lavage (BAL) and found that ETA overdiagnosed VAP compared with BAL (89% versus 21% of clinically suspected cases, P<0.0001). Using infection surveillance data of 643 patients, 110 of which with clinically suspected VAP, Morris *et al.* (19) modelled the potential effect on VAP incidence and antibiotic use of using exclusively ETA or BAL and found that changing from exclusive ETA to BAL diagnosis would decrease reported VAP incidence by 76% (95% CI, 67% to 87%) and antibiotic use by 30% (95% CI, 20% to 42%).

In 2013, the Centers for Disease Control and Prevention (CDC) brought together a multidisciplinary team of renowned experts to address the limitations of the NHSN definition and propose a new set of metrics (6). Therefore, a tiered definition algorithm was developed with three main targets: first, to extend surveillance to generic complications of mechanical ventilation (both infectious and non-infectious); second, to make surveillance as objective as possible; third, to enable an automatic enrollment of clinical records. It emphasizes the importance of preventing all complications of mechanical ventilation rather than pneumonia alone (20), introducing the concept of ventilator-associated event (VAE). VAE consists of ventilator-associated conditions (IVAC), infection-related ventilator-associated complications (IVAC) (a subset of

VAC with infectious signs), possible and probable VAP (IVAC with microbiological evidence of pneumonia) that are characterized based solely on quantitative (and so objective) criteria (*Table 1*). Such tiers and definitions are only suitable for potential use in public reporting, benchmarking or internal quality improvements initiatives and are not designed for use in the management of patients. As a practical example, they completely disregard radiographic criteria, not its role in daily practice, but its lack of objectivity and accuracy to constitute a useful tool in surveillance reports (17).

VAP rates surveillance

Significant differences in VAP rates have been reported in the literature (21).

The International Nosocomial Infection Control Consortium (INICC) conducted a cohort prospective multicentre surveillance study of device associated HAI in 703 ICUs in 50 countries from Latin America, Europe, Eastern Mediterranean, Southeast Asia, and Western Pacific World Health Organization regions (United States not included) from 2010 to 2015 (22). They found a pooled mean VAP rate (number of VAP over 1,000 ventilation-days) of 12.2 (95% CI, 12.0-12.4) with lower incidence in surgical cardiothoracic ICUs (6.2, 95% CI, 5.5-7.0) and higher incidence in neurologic ICU (19.2, 95% CI, 15.6-23.5). The pooled mean of ventilator used ratio (ventilator-days per 100 patient-days) was 37 (95% CI, 37-37), being lower in medical cardiac ICU (20, 95% CI, 20-20) and higher in trauma ICU (66, 95% CI, 65-66). A significant trend toward the reduction of VAP rates in INICC ICUs was observed since 2002, when surveillance reports presented a VAP rate of 24.1 (95% CI, 22.8-25.5), virtually twice the observed VAP rate in the 2010-2015 period.

Martin-Loeches *et al.* (5) prospectively analyzed almost 3000 ICU patients from 114 ICUs in Europe and South America during 2013 and 2014 and found an incidence of VAP of 8.8 per 1,000 ventilation-days that was statistically similar to the incidence of ventilator-associated tracheobronchitis (VAT) (10.2 per 1,000 mechanically ventilated days, P=0.48). Trauma patients had a higher proportion of VAP and VAT (18% and 19%) when compared to both medical (9% and 12%) and surgical (9% and 10%) patients, which had already been described in previous studies (23).

When comparing these results with the United States (US) surveillance data reported by the NHSN the

differences are quite significant, with much higher VAP rates in the INICC report, in some case more than 20 times (22,24). The most recent NHSN report (from 2012) found pooled VAP rates of 0.0 to 4.4 with pooled mean ventilator used ratio of 3 to 47 ventilator-days per 100 patient-days (24). Making a direct comparison by ICU-type we found in surgical cardiothoracic ICUs an INICC VAP rate of 6.2 (95% CI, 5.5-7.0) versus a NHSN VAP rate of 1.7 (95% CI, 1.5-1.9), in neurologic ICU an INICC VAP rate of 19.2 (95% CI, 15.6-23.5) versus a NHSN VAP rate of 3.0 (95% CI, 2.3-3.8) and in respiratory ICU an INICC VAP rate of 13.8 (95% CI, 11.9-15.8) versus a NHSN VAP rate of 0.7 (95% CI, 0.2-1.7) (22). Some differences may clearly reflect socioeconomic contrasts leading to differences in daily clinical practices, namely a possible irregular adherence to and compliance with the guidelines, low nurse-to-patient staffing ratios, lack of medical supplies or an insufficient number of experienced nurses or trained health care workers (22).

Deeper examining the NHSN reported data, there is a 13% to 19% per year reduction in VAP rates from 2002 to 2008 (25) and more than 50% of non-teaching medical ICUs in the United States are reporting VAP rates of zero since 2010, with higher mean rates in surgical units (2.5 VAP per 1,000 ventilator-days) versus medical units (1.0 VAP per 1,000 ventilator-days) (20,24,26,27). Enthusiasm fades and curiosity sets in when comparing these data with European results, where socioeconomic conditions and sophisticated prevention and care programs are similar to the United States. In 2014, the European Centre for Disease Prevention and Control (ECDC) described in patient-based surveillance, a pooled rate of 10.0 VAP episodes per 1,000 intubation-days that varied between 2.8 in the United Kingdom and 15.8 in Italy and Belgium, with ventilator used ratio between 32.8 ventilator-days per 100 patient-days in Luxembourg and 68.2 ventilator-days per 100 patient-days in Portugal (28). These rates remained stable compared with 2011 and 2012 (29) and represent a 25% reduction of pooled VAP rate since 2007 (30).

These significant differences between NHSN and INICC/ ECDC data raise the question to what extent the NHSN numbers reflect real improvements in care rather than surveillance biases (17). In the United States, VAP has been proposed as an indicator of quality of care in public reporting, and its prevention is a national patient safety goal (15). The threat of financial penalties for this diagnosis has put pressure on hospitals to minimize VAP rates (18) which may prompt surveyors to apply subjective criteria more strictly,

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leading to misleading lower rates, independently of patient care. Klompas (31) described 8 initiatives that hospitals might be considering applying to make VAP surveillance more rigorous but which will artificially lower VAP rates despite not materially improving prevention and care, regardless of how well intentioned those measures might be. Such initiatives included interpretation of clinical signs and chest X-ray as strictly as possible, require consensus between two or more infection preventionists and intensivists, require BAL cultures for diagnosis, set quantitative growth thresholds for BAL and ETA cultures, transfer out patients needing prolonged mechanical ventilation and admit uncomplicated postoperative mechanically ventilated patients (31). This makes evident the high variability and how artificial or manipulated the rates might be (32) and enables the fact that clinicians who track VAP in parallel with their hospitals' infection surveyors consistently find more cases than the surveyors do (17).

VAP rates and antibiotic consumption

Optimal antibiotic therapy is a major focus of VAP treatment (33). Both inappropriate initial antimicrobial treatment and delays in the administration of appropriate antibiotic therapy are associated with higher mortality and longer hospital stays in ICU patients (33,34). Due to these observations, a number of clinical practice guidelines for antibiotic therapy for VAP have evolved over the years (34,35). The optimal duration of antimicrobial treatment for VAP is unknown but there is a growing evidence that shorter antibiotic courses of 6–8 days can be effective and simultaneously minimize the consequences of antibiotic overuse in critical care, including antibiotic resistance, adverse effects and costs (35,36).

It would be expected from the evidence of benefit towards shorter courses of antibiotics associated with the reduction of VAP rate that the consumption of antibiotics would also decrease in particular in those hospitals with VAP rate of "zero". Ferreira *et al.* (2) evaluated the clinical impact of a good practice bundle in ICU patients and found a decreased occurrence of VAP (P<0.01), a significant reduction in hospital costs (P<0.05) and lower mortality (P<0.01) as well as significantly less days of antibiotic use (P<0.0001). Rosenthal *et al.* (37) demonstrated the effectiveness of a multidimensional approach for prevention of VAP in adult ICUs (VAP rate evolved from 22.0 to 17.2, P=0.0004) and a simultaneous pooled reduction in antibiotic consumption. Primary analysis revealed a significant increase in the ratio of patients free of antibiotics comparing both periods (36.2% to 40.9%, P=0.0002), with a differential decrease in consumption of ciprofloxacin (6.6% to 5.6%, P=0.0087) and ceftriaxone (17.7% to 12.1%, P=0.0001). When analyzed defined daily dose (DDD) per 1,000 patient-days (DDD/1,000 PD), a similar reduction in antibiotic prescription was noticed during intervention period (958.3 to 925.8 DDD/1,000 PD). It was found a reduction in the administration of five groups of antibiotics—cephalosporins, quinolones, lincosamides, glycopeptides, and aminoglycosides—and a rise of carbapenems and penicillins (38).

This data however is not consistent with ICU reports that show a steadily use of antibiotics for nosocomial pneumonia in 15% of patients (17). Fihman *et al.* (1) retrospectively analyzed a 5-year trend of ICU antibiotic consumption and resistance in bacteria causing VAP between 2007 and 2011 in a university-affiliated tertiary care institution in France. The frequency of early VAP episodes decreased significantly over the study period (P<0.011) but antibiotic consumption rose by more than one-third during this period (1,004 to 1,389 DDD/1,000 PD). This change though was not statistically significant (P=0.07) and is comparable to the median (range) antibiotic consumption of 1,254 (range, 348–4,992) DDD/1,000 PD reported by Hanberger *et al.* (39) in a European ICU multicenter surveillance program during 2005.

ECDC 2007 surveillance report showed that 77.4% of patients received at least one systemic antibiotic while in the ICU. Since information on dosage is not collected in the surveillance protocol, antimicrobial consumption was expressed as the number of ICU days with at least one antimicrobial administered (two different antimicrobials on one day =1 day)-67.5 antibiotic days per 100 patient days—as well as the total number of antimicrobial days (two different antimicrobials on 1 day =2 days) per 100 patient days in the ICU-131.8 antibiotic days per 100 patient days (30). During the biennium 2011-2012 the ECDC reported a prevalence of antimicrobial use among ICU patients of 56.5% and 37.3% of all antibiotics used to treat HAI were used in ICU patients (37.3%). Tracheal intubation was an independent risk factor for antibiotic consumption (adjusted odds ratio 2.1, 95% CI, 1.9-2.3) (29). Between 2011–2015, overall antibiotic consumption in the European hospital sector (expressed as DDD per 1,000 inhabitants and per day) showed no significant trend (40). There was also no significant trend observed in the consumption of antibiotics for treatment of patients

infected with serious multidrug-resistant (MDR) bacteria during this period at European level (1). In 2015, the European population-weighted mean consumption of antibiotics for systemic use in the hospital sector was 2,000 DDD/1000 PD, ranging from 1.0 in the Netherlands to 2.9 in Malta (40). However, no ECDC surveillance data specifically analyses the trend of antibiotic consumption in VAP patients.

Magill et al. (41) conducted a cross-sectional evaluation of antimicrobial use in US acute care hospitals, assessing it over a 2-day period in 10 American states between May and September 2011. Hospital antibiotic use prevalence was 49.9% (95% CI, 49.0-50.9%) with higher values in ICUs than in other locations (57.7%, 95% CI, 55.4-60.0% versus 48.6%, 95% CI, 47.6-49.6%; P<0.001). Baggs et al. (42) estimated trends in antibiotic use in more than 300 US hospitals from 2006 to 2012. Overall rates of antibiotic use did not change significantly during the study period. Antibiotic administration prevalence was 55.1% and the rate of antibiotic use for all study years was 755 days of therapy per 1,000 patient days (DOT/1,000 PD), rising from 732.5 DOT/1,000 PD in 2006 to 767.5 DOT/1,000 PD in 2012. Antibiotic consumption for all study years was 52% greater in ICUs compared to noncritical care locations (1,092 versus 720 DOT/1,000 PD, P<0.001). As its European counterparts, no CDC surveillance data specifically analyses the trend of antibiotic consumption in VAP patients.

Discussion

VAP has an undeniable clinical and epidemiological relevance and accounts for up to 50% of all antibiotics used in ICUs (1,20,43). It would then be expected that VAP rates and antibiotic consumption evolved similarly, which surprisingly does not happen. The described limitations in the definition of VAP contribute to this disagreement but are not the only intervening factors.

Particular attention should be taken in interpreting studies with surprisingly low VAP rates. Such reports must be carefully analyzed in order to correctly access how artificial or manipulated the rates might be. Although VAP prevention bundles may contribute for encouraging reductions in VAP episodes, one should not lightly believe that a 'zero-VAP rate' is feasible (32). VAP pathophysiology and epidemiology limit its ability to prevent all episodes of VAP (32). Prevention bundles are essentially effective in reducing late onset VAP, that accounts for up to 60% of VAP (44), while early onset VAP, in turn, is mainly related to the presence of an endotracheal tube and the resulting violation of natural defense mechanisms (15). VAP risk factors such as each patient host flora, severity of underlying disease, microaspiration of oropharyngeal content (5) (particularly important in trauma patients when a low Glasgow Coma Score is present but also relevant whenever tracheal intubation was unscheduled and/or the patient was nonfasted), grade of functional immunosuppression when critically ill, previous surgery and antibiotic exposure (15) are all endogenous to patients, cannot be fully addressed by prevention bundles and may be unavoidable consequences of caring for critically ill patients (17,18).

Besides variation in diagnostic and record practices, ICU characteristics and patient population, such as clinical severity and infection control guidelines may constitute surveillance biases and also affect the reported incidence of VAP (28).

The evaluation of antibiotic consumption is not uniform. The World Health Organization concept of DDD-the assumed maintenance dose per day in adults-can be used to standardize the comparisons of drug consumptions in various conditions (38) but its ratio per 1,000 patient-days is not used in all antibiotic surveillance reports; some authors presented antibiotic use by the ratio of patients treated with antibiotics and this difference in antibiotic quantification may contribute with confusing data. Using the DDD/1,000 PD ratio may itself constitute a bias because, although describing antibiotic pressure it does not describe accurately the number of patients treated and those not treated with antibiotic during each period (38): the exact same DDD/1,000 PD ratio may represent a small number of patients receiving a class of antibiotics for a long period or at a high dose as well as a large number of patients receiving small amounts of antibiotic or antibiotic over a short period (1,38). Moreover, this method is influenced by the length of stay of patients and may overlook antibiotic de-escalation (45).

Knowledge and understanding of bacterial epidemiology and resistance trends for each ICU have great relevance. In general, late onset VAP is caused by multi-drug resistant and more difficult to treat bacteria than early onset VAP (15). Data from ECDC (28), NHSN (46), Canadian (33) and French (1) ICUs all report different bacteria isolation rates and resistance patterns which implies distinct initial empiric antibiotic and diverse global and group-specific antibiotic use. ECDC data from 2014 (47) and 2016 (28) reported that the most frequently isolated microorganisms in VAP were *Pseudomonas aeruginosa* followed by *Staphylococcus aureus* [with an average percentage of methicillin-resistant *Staphylococcus aureus* (MRSA) of 43.0%], *Klebsiella spp.* and *Escherichia coli*. Fihman

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et al. (1) observed a significant increase in resistance rates to several classes of antimicrobial drugs in a retrospective 5-year trend analysis of VAP patients in a French tertiary-hospital ICU. Such resistance trend was more significant due to an increase in the isolation of AmpC-producing Enterobacteriaceae. Koulenti et al. (44) prospectively analyzed VAP patients from 27 ICUs in 9 European countries and isolated MDR organisms (MRSA, Pseudomonas aeruginosa, Acinetobacter baumannii and Stenotrophomonas maltophilia) in 53.7% of cases. Weiner et al. (46) described antimicrobial resistance patterns for VAPs reported to the NHSN from 2011 and 2012: the most isolated bacteria were Staphylococcus aureus (percentage of MRSA 46.1% in 2011 and 42.4% in 2012) and Pseudomonas aeruginosa (percentage of MDR 20.8% in 2011 and 19.9% in 2012). Craven et al. (7) examined the incidence and microbiological data from VAT and VAP in medical and surgical ICU patients and MDR pathogens were isolated in 50% of VAP patients, namely MRSA that was present in more than 20% of VAP patients. The emergence over the last decade of MDR bacteria certainly contributed to an increase in antibiotic consumption, which itself increases the selective pressure favoring the most resistant bacteria (1,46).

The new NHSN criteria provides a more objective and quantitative data based definition, that may theoretically, be advantageous. First, they can potentially reduce the methodological biases on IVAC rate calculation, providing trustworthy values for surveillance and benchmarking. Second, these new criteria provide clinicians with the capability of early recognizing and attending patients with mechanical ventilation-related problems that could be missed by VAP prevention or ancient surveillance protocols (17), thus taking the first step towards elucidating these events and creating opportunities to improve care. Third, the duty to include an antibiotic administration in IVAC definition, may supply reliable and comparable data on antibiotic consumption between ICUs (17). Quality indicators for infection control and antibiotic stewardship and/or surveillance programs will then have the possibility to become aware of IVAC-related antibiotic prescription, to identify outlier prescribers and to tailor antibiotic therapy, thereby reducing the risk of MDR bacteria emergence. All this may allow a more comprehensive assessment of the observed VAP and antibiotic consumption variability.

It is important though to realize that the new NHSN metrics are not free from potential biases. An IVAC can encompass both VAP and VAT clinical concepts since it does not use chest X-ray as a diagnostic criterion and it is impossible to clinically distinguish VAP from VAT without it. It will then be possible to observe identical IVAC rates between studies but with differences between VAT and VAP numbers, which in addition to being discrepant may not be real: some patients who were diagnosed as having VAP may in fact have VAT and vice versa (5). Lachiewicz et al. (48) assessed the effect of the NHSN definition (possible VAP) on VAP rates in a burn ICU over different time periods (from July 2011 to December 2012 they used the old definition; from January 2013 to June 2014 and from July 2014 to December 2015 the new criteria were applied) and found fewer VAP cases (rates from 4.47 to 1.03 and 0.55 VAP/1,000 ventilator days, respectively) with the new definition. No statistically significant differences were found between VAP groups regarding length of stay and mortality. However, when the ICU infection preventionists evaluated cases screened from July 2014 to December 2015 that did not meet possible VAP criteria, 18 VAP cases meeting the pre-2013 criteria were identified, resulting in an adjusted incidence of 4.96 VAP/1,000 ventilator days. Younan et al. (10) compared the outcomes of patients meeting the classic VAP and/or the possible VAP criteria in a cohort of ICU trauma patients. A significant difference was found between classic VAP (30.9%) and possible VAP (6.6%) incidence and only 5.8% of patients (n=1,116) met both definition criteria, with identical mortality. These results imply that the NHSN new surveillance definition may miss potentially clinically meaningful events that are important for driving infection prevention (10,48), due to its low sensitivity and positive predictive value for identifying VAP (11).

Conclusions

The epidemiological relevance and clinical impact of VAP are unequivocal. It is therefore essential to ensure that objective and uniform definitions are used. VAP management must focus not only on early suspecting, diagnosing and properly treating but also on reliably recording epidemiological data (regarding both VAP incidence and antibiotic doses and duration of therapy) in order to assess and benchmark the quality of care for each type of ICU patients as accurately as possible. Microbiology epidemiological data-including prescription habits, the infections treated, infection control measures and epidemic phenomena between ICUs-are keystone for diagnosis and for tailoring antibiotic therapy. Appropriate antibiotics in adequate doses followed by de-escalation based on microbiological results and the clinical response of the patient constitute the best weapon against the potential

for antibiotic overuse, the emergence of MDR bacteria, unnecessary adverse effects and potential toxicity. This must be the way to guarantee that there will be fewer discrepancies and more parallelism between VAP rate and antibiotic consumption in the near future.

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Footnote

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