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# A national survey of the diagnosis and management of suspected ventilatorassociated pneumonia

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## ABSTRACT

**Background:** Ventilator-associated pneumonia (VAP) affects up to 20% of patients admitted to intensive care units (ICU). It is associated with increased morbidity, mortality and healthcare costs. Despite published guidelines, variability in diagnosis and management exists, the extent of which remains unclear. We sought to characterise consultant opinions surrounding diagnostic and management practice for VAP in the UK. **Methods:** An online survey was sent to all consultant members of the UK Intensive Care Society (n=~1500). Data were collected regarding respondents' individual practice in the investigation and management of suspected VAP including use of diagnostic criteria, microbiological sampling, chest X-ray (CXR), bronchoscopy and antibiotic treatments.

**Results:** 339 (23%) responses were received from a broadly representative spectrum of ICU consultants. All respondents indicated that microbiological confirmation should be sought, the majority (57.8%) stating they would take an endotracheal aspirate prior to starting empirical antibiotics. Microbiology reporting services were described as qualitative only by 29.7%. Only 17% of respondents had access to routine reporting of CXRs by a radiologist. Little consensus exists regarding technique for bronchoalveolar lavage (BAL) with the reported volume of saline used ranging from 5 to 500 mL. 24.5% of consultants felt inadequately trained in bronchoscopy.

**Conclusions:** There is wide variability in the approach to diagnosis and management of VAP among UK consultants. Such variability challenges the reliability of the diagnosis of VAP and its reported incidence as a performance indicator in healthcare systems. The data presented suggest increased radiological and microbiological support, and standardisation of BAL technique, might improve this situation.

## INTRODUCTION

Ventilator-associated pneumonia (VAP) is a common nosocomial infection affecting up to 20% of patients admitted to intensive care units (ICUs).<sup>1–3</sup> VAP is associated with a 2–7-fold increased risk of death<sup>4 5</sup>; although the actual attributable mortality of VAP has proven difficult to determine with a wide range of

## **KEY MESSAGES**

- Despite guidelines, marked variation in the diagnosis and management of ventilator-associated pneumonia (VAP) continues to exist within the UK.
- Until a consensus approach is reached the use of VAP rates as a performance indicator in healthcare systems is unreliable.
- Increased radiological and microbiological support may improve the diagnosis of VAP.
- Better training in the performance of bronchoalveolar lavage (BAL) may lead to increased delivery of high-quality BAL and improved diagnosis of VAP.

estimates reported.<sup>1–3</sup> <sup>6–8</sup> Furthermore, VAP is associated with excess morbidity and healthcare costs, increasing hospital length of stay by an average of 4–9 days.<sup>1 5 6</sup> Despite the introduction of preventive measures aimed at reducing VAP rates it remains prevalent within ICUs. This has led to the incidence of VAP being adopted as a performance indicator in some healthcare systems. In turn, this has drawn further attention to the significant challenges the condition presents in terms of diagnosis and management. The lack of consensus as to diagnostic criteria or the 'gold standard' diagnostic test has hampered research development over recent years and more recently has led to discrepancies in clinical VAP rates and reported surveillance rates.9

Although guidelines have been published,<sup>10–12</sup> variability in the approach to diagnostic procedures and management of VAP exists and is poorly defined. We therefore sought to characterise consultant opinion surrounding diagnostic and management practice for VAP in the UK.

## **METHODS**

All consultant members of the UK Intensive Care Society were invited to take part in an online survey. A survey tool was developed





and piloted in a local ICU prior to distribution nationally. The survey collected data regarding demographics of the respondents' place of work and their individual practice in the investigation and management of suspected VAP including use of diagnostic criteria, microbiological sampling, chest X-ray (CXR), bronchoscopy and bronchoalveolar lavage (BAL) and antibiotic treatments.

The survey was hosted by the Survey Monkey website (http://www.surveymonkey.com). An email inviting participation was sent to all consultant members of the UK Intensive Care Society. All responses were anonymous in terms of the individual and their place of work. Both quantitative and qualitative data were collected for analysis. Reminder emails were sent to all consultant members 2 and 10 weeks after the initial invitation to encourage participation.

The survey questions are summarised in the online supplementary section.

Descriptive statistical data are presented as mean (SD), median (IQR) and percentages. Correlations were tested using Spearman's rank correlation coefficient.

## RESULTS

#### **Demographic data**

Responses were received from 339 of approximately 1500 consultants surveyed (23%). Only 266 respondents completed the survey in full (266/339=78.4%). Consultant experience in intensive care medicine varied with a median number of years practising of 14 (range 1–35). Respondents reported a median number of staffed level 3 beds (beds where patients may receive advanced respiratory support) per unit of 8 (range 2–40, IQR 6–12), with an average of 69.5% (SD±17) of beds estimated to be occupied by intubated, mechanically ventilated patients

at any one time. There was a wide case mix reported within the units. Mean case load is illustrated in figure 1. Responses were received from 28 consultants (8.3%) working in specialist units (21 specialist cardiothoracic and 7 specialist neurology/neurosurgery) where 100% of patients were from the relevant specialty.

## Diagnosis

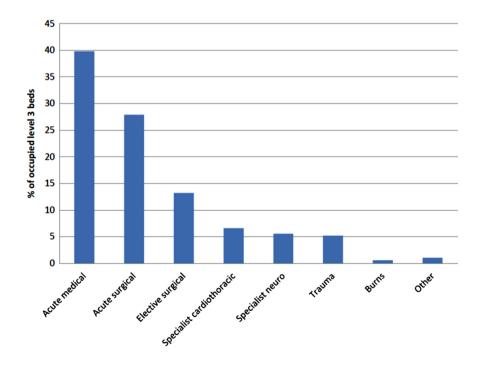
### Clinical criteria

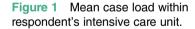
Respondents were asked to list criteria they considered to be mandatory for a diagnosis of VAP. A wide variety of responses were received. When respondents' answers were compared with the criteria set out in published guidelines for the diagnosis of VAP 60.6% (163/269) of respondents matched the Canadian Thoracic Society guidelines, 55.3% (149/269) the American Thoracic Society guidelines, 49.1% (132/269) the HELICS (Hospitals in Europe for Infection Control through Surveillance) criteria and 28.3% (76/269) the Guidelines from the British Society of Antimicrobial Therapy (figure 2). In total 33.1% (89/269) of respondents did not include CXR appearances within the list of criteria they considered mandatory for a diagnosis of VAP.

## Microbiological sampling and bronchoscopic technique

Respondents were asked to choose one option (from those shown in figure 3) which best reflected their current practice when suspecting a diagnosis of VAP. All respondents indicated that some form of microbiological confirmation should be sought with the majority of consultants, 58% (156/269) indicating that they would take an endotracheal aspirate (ETA) prior to starting empirical antibiotics.

In terms of experience in bronchoscopy 74.1% (200/270) of respondents reported that they personally





Guideline	Radiological Criteria	Clinical Criteria	Clinical Criteria
American Thoracic Society and Infectious Diseases Society of America	Presence of new or progressive radiographic infiltrates	AND at least 2 of 1. Fever > 38°C 2. White cell count >12,000/mm <sup>3</sup> or <4000/mm <sup>3</sup> 3. Purulent secretions	
British Society of Antimicrobial Chemotherapy	New or persistent infiltrate on CXR	AND / OR Purulent tracheal secretions	AND Increased O <sub>2</sub> requirement AND Core temperature >38.3*C AND Blood leucocytosis (> 10,000/mm <sup>3</sup> ) or leucopaenia (<4,000/mm <sup>3</sup> )
Association of Medical Microbiology and Infectious Disease Canada and Canadian Thoracic Society	One of the following abnormalities on CXR 1. Evidence of alveolar infittrates OR 2. Evidence of air bronchograms OR 3. New or worsening infittrates	AND 2 or more of 1. Temperature >38°C or < 36°C 2. Leukopaenia or leucocytosis 3. Purulent tracheal secretions 4. Decreased PaO <sub>2</sub>	
Hospitals in Europe Linked for Infection Control through Surveillance project (HELICS)	Two or more serial chest X-rays or CT-scans with a suggestive image of pneumonia for patients with underlying cardiac or pulmonary disease. In patients without underlying cardiac of pulmonary disease one CXR or CT scan is sufficient	AND at least one of the following 1. Fever >38 °C with no other cause 2. Leucopaenia (<4,000 WCC/mm3) or leucocytosis (>12,000 WCC/mm3)	AND at least one of the following 1. New onset of purulent sputum, or change in character of sputum (colour, odour, quantity, consistency) 2. Cough or dyspnoea or tachypnoea 3. Suggestive auscultation (rales or bronchial breath sounds), rhonchi, wheezing 4. Worsening gas exchange (eg O2 desaturation or increased oxygen requirements or increased ventilation demand)

Figure 2 Recognised criteria for diagnosis of ventilator-associated pneumonia (CXR, chest X-ray).

undertook bronchoscopy within the ICU. Only 27.1% (73/269) performed bronchoscopic sampling in cases of suspected VAP. The estimated number of bronchoscopies performed per month per unit varied widely with a rate per level 3 bed of 0-5.8 (mean 0.89, SD±0.82). There was little consensus in the reported technique for bronchoscopic sampling. The reported volume of saline used to perform a BAL ranged from 5 to 500 mL (median 20 mL, IQR 20-40) with only 9.6% (20/208) of respondents instilling>100 mL. In terms of the sampling site respondents were asked to identify which lobe/ segment they would lavage in a patient with diffuse bilateral shadowing on CXR. The most frequently cited locations were both lower lobes (27.3%, 57/209), right lower lobe (16.3%, 34/209), or the right middle lobe (13.4%, 28/209), while 22 respondents (10.9%) stated that they would lavage all lobes. 24.5% (49/200) of respondents who perform diagnostic bronchoscopy felt that they were not adequately trained in the procedure with 9.5% (19/200) stating they had received no training at all. The level of confidence in performing BAL was reported to be high, however, with a mean score of 8.3/10 (SD±1.7 on a linear scale of 0-10 where 0 represented no confidence and 10 represented 100% confidence). This compared to confidence levels of 4.7/10(SD±2.6) for recognition of a tumour, 1.8/10 (SD±3.0) for endobronchial biopsy and 8.4/10 (SD±1.5) for clearance of impacted mucus. There was no significant correlation between years of experience in intensive care medicine and confidence in performing BAL (r<sub>s</sub>=0.013, p=0.85).

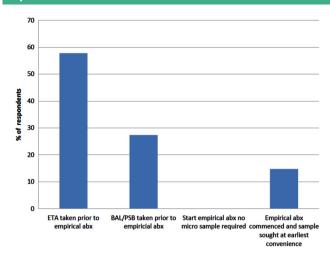
Data were collected about the availability of qualitative and quantitative microbiology services within respondents' hospitals (figure 4). Microbiology departments were said to issue qualitative reports only by 29.7% (80/269) of respondents, as compared with semiquantitative reporting (37.9%, 102/269) and quantitative reporting (18.6%, 50/269). 13.8% (37/269) of respondents were unsure of the nature of microbiology services available within their hospital.

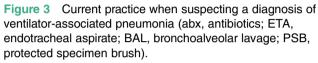
## Chest X-ray

Data were gathered in relation to the frequency of, and indications for, CXR in ventilated patients on ICU. In total, 76.7% (204/266) stated they only perform CXRs on ventilated patients when considered clinically indicated, 13.5% (36/266) reported requesting a CXR on patients routinely every 3–4 days, 5.3% (14/266) every other day, 2.6% (7/266) once a week and 1.9% (5/266) on a daily basis. The features reported as most likely to be considered an indication for a CXR in a ventilated patient were new signs on auscultation and a rise in required inspired oxygen concentration. These were followed in order of likelihood by; the presence of new purulent secretions, a new temperature, a new or increasing inotrope requirement, and a new rise in white cell count (WCC).

Data were collected regarding the level of reporting of CXRs on the ICU (figure 5). A total of 46.2% (123/266) stated that on their unit CXRs are reported by ICU staff, with interesting CXRs discussed with a radiologist in person or at an X-ray meeting (selected reporting). 18.8% (50/266) reported that CXR interpretation was done entirely by ICU staff (no reporting), 18.1% (48/266) by a radiologist when specifically requested to do so (reporting by request) and 16.9% (45/266) stated all

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CXRs on their unit were reported by a radiologist (full reporting).

Respondents were asked to indicate their level of confidence in interpreting CXRs performed on intubated, mechanically ventilated patients. The mean score was 8.4/10 (SD±1.1) on a linear scale of 0–10 where 0 represented no confidence and 10 represented 100% confidence. There was no significant correlation between confidence in interpreting CXRs and years of experience in ICU ( $r_s$ =0.079, p=0.199).

## Treatment

#### Pathogens

Respondents were asked to list what they considered to be the commonest organism causing VAP within their unit. The organism most frequently cited was *Pseudomonas aeruginosa* (30.1%, 81/269), followed by Enterobacteriaceae (35.7%, 96/269; including 'coliforms' (20.8%, 56/269),

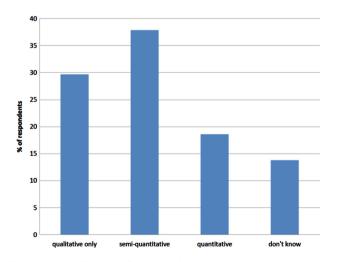


Figure 4 Availability of microbiology services within respondents' hospitals.

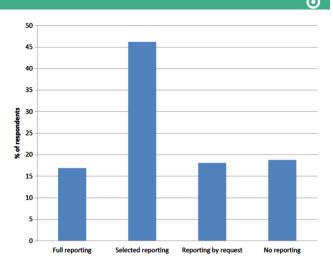


Figure 5 Level of reporting of chest X-rays on the intensive care unit.

*Escherichia coli* (10.8%, 29/269), *Klebsiella* spp (3.3%, 9/269) and *Enterobacter cloacae* (0.7%, 2/269)) and methicillinsensitive *Staphylococcus aureus* (10.8%, 29/269). Fungal and viral infections were mentioned infrequently (<1%).

#### Antibiotic therapy

A single agent antibiotic regimen was identified in the majority of cases as being the usual empirical antibiotic therapy (78.8%, 212/269). Piperacillin-tazobactam was the most frequently reported antibiotic, (69.9% (188/269) of respondents) followed by meropenem (24.2%, 65/269). Forty-three different single or double agent antibiotic regimens were listed as being usually prescribed empirical antibiotic therapy in cases of suspected VAP. The 15 most frequently cited antibiotic regimens are illustrated in figure 6. Median reported duration of antibiotic therapy for the treatment of VAP was 6 days (IQR 5–7). Prescribed courses ranged from 3 to 14 days in duration.

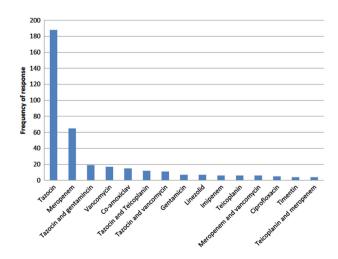


Figure 6 Reported empirical antibiotic selection in cases of suspected ventilator-associated pneumonia.

## DISCUSSION

The results of this study show that there continues to be wide variation in practice in relation to the investigation, diagnosis and treatment of VAP in the UK. Although the overall response rate was low, comparison with a PubMed search of surveys of ICU consultants over the past 10 years showed that responses were received from a broadly representative spectrum of consultants in terms of clinical experience and case load. This suggests that the data gathered are generally reflective of current UK consultant practice.<sup>13–16</sup>

It is well recognised that diagnosis of VAP is extremely difficult on clinical grounds alone.<sup>3 17</sup> There is significant overlap in the signs and symptoms associated with VAP and many other conditions affecting patients on ICU. Clinical criteria such as a new or persistent alveolar infiltrates on CXR in association with purulent tracheal secretions, increasing oxygen requirements, temperature  $>38^{\circ}$ C and WCC  $>10\ 000/\text{mm}^3$  or  $<4000/\text{mm}^3$  have been incorporated into guidelines and clinical scoring systems to aid in the diagnosis.<sup>10</sup> <sup>12</sup> <sup>17</sup> Despite such guidance surrounding the use of clinical criteria however, there was little consensus among respondents regarding those criteria considered mandatory for diagnosis, with only 60.6% of respondents listing criteria fulfilling at least one of the national guideline definitions for VAP based on clinical criteria. A total of 33.1% (89/269) of respondents did not consider a CXR to be required which is contrary to guideline recommendations stating that all patients with suspected VAP should have a CXR performed to look for the presence of new infiltrates.<sup>10–1</sup>

The radiological diagnosis of VAP is also challenging with low diagnostic sensitivity and specificity of CXR signs in ventilated patients leading to difficulties in interpretation. The only sign to correlate well with the presence of pneumonia is the air bronchogram but even the specificity of this is reduced when coexistent acute respiratory distress syndrome is present.<sup>18</sup> We are unaware of data comparing the diagnostic accuracy of radiologists and intensivists in the setting of suspected VAP, however data from the pneumonia literature generally suggests that interobserver variability in CXRs is high, and that radiologists probably provide greater diagnostic accuracy.<sup>19</sup> <sup>20</sup> Despite this only 16.9% of respondents had access to routine reporting of CXRs by a radiologist and confidence in interpreting CXRs was reported to be high with a mean score of 8.4/10.

In terms of microbiological sampling, the majority of respondents reported that their usual practice was to take an ETA for culture prior to starting empirical antibiotics. Colonisation of the proximal airways is common in intubated patients. Non-quantitative culture of pathogens from ETAs commonly reflects such colonisation rather than a pneumonic process. Qualitative ETAs are known to have high sensitivity and low specificity for the diagnosis of VAP, so while they may have a role in excluding VAP, their use for diagnostic purposes is associated with a high rate of false positives and excessive use of unnecessary antibiotics.<sup>3</sup> <sup>10</sup> <sup>21</sup> The suggestion that approximately 50% of respondents do not have access to quantitative or semiquantitative culture in decision-making may therefore have important consequences. Although there are few studies directly comparing the use of qualitative versus quantitative ETA, a diagnostic cut-off of >10<sup>6</sup> colony-forming units (CFU) per millilitre has been shown to significantly increase the specificity of ETA.<sup>3</sup> <sup>21</sup> Increased access to quantitative microbiology reporting may, therefore, reduce inappropriate antibiotic prescribing in this situation, however, an associated decrease in sensitivity, may put some patients at risk.

Similarly, non-quantitative BAL samples are hard to interpret whereas culture of pathogens at  $>10^3$  CFU/mL from protected specimen brush samples or at  $>10^4$  CFU/mL from BAL fluid appear to increase diagnostic accuracy.<sup>3</sup> <sup>22</sup> For example, pathogens cultured at  $>10^4$  CFU/mL from BAL fluid had sensitivity of 91% and specificity of 78% for the microbiological confirmation of infection in lung tissue, which in turn correlated with histological evidence of VAP.<sup>23</sup>

Considerable debate continues as to whether bronchoscopic sampling is indicated for the diagnosis of VAP.<sup>24-27</sup> Despite evidence showing that the sensitivity and specificity of bronchoscopic sampling are superior to ETA, benefits in terms of patient outcomes have been inconsistent in randomised trials. A recent meta-analysis has not shown differences in mortality, length of stay or antibiotic changes.<sup>28</sup> Bronchoscopic sampling is associated with a lower rate of positive culture among patients with suspected VAP compared with ETA.<sup>2</sup> Discontinuation of antibiotics in the face of negative BAL culture has been shown to be safe and result in lower antibiotic use and potentially fewer antibiotic-resistant pathogens.<sup>29</sup> This important potential benefit has not been adequately evaluated in the meta-analysis and reductions in antibiotic use, measured by an increase in antibioticfree days, is probably a more valid outcome measure than mortality for future VAP trials.

Relatively little is known about the general quality of alveolar sampling in suspected VAP. Our survey suggested huge variation in sampling techniques among the respondents who perform BAL. Guidelines for BAL generally recommend instillation of 100-240 mL of sterile saline to ensure sampling of the alveolar space.<sup>30-32</sup> The median instillate described in this survey was 20 mL, with only 9.6% (20/208) of respondents using  $\geq 100$  mL. This suggests that many BALs may not adequately sample alveolar tissue and may therefore be inappropriate for diagnosing pneumonia. The posterior segment of the right lower lobe is considered the most commonly involved segment in VAP.<sup>33</sup> Only 43.3% of respondents who perform BAL reported sampling the right lower lobe when CXR gives no indication of the most involved region of the lung. Finally, respondents' confidence in the ability to perform BAL seemed at variance with the fact that 24.5% described inadequate training in

bronchoscopy and 9.5% no training at all. The overall implication is that in a significant number of cases BAL is unlikely to provide optimal alveolar sampling.

Together, the data pertaining to CXRs, microbiological analysis and BAL suggest that an agreed optimal approach to the diagnosis of VAP is lacking in the UK. Few data are available to determine whether this situation is prevalent in other healthcare systems, but it seems likely that similar circumstances will be duplicated in at least some other countries. The clinical implications may be far-reaching. Several studies have shown that objective evidence for VAP is only obtained between 20% and 42% of patients in whom the condition was suspected on clinical grounds.<sup>34–36</sup> The overall trend for decision-making based on non-quantitative microbiology from ETA (or from small volume 'BAL' which has sampled the proximal airways) would tend to favour false-positive diagnoses and the use of unnecessary empirical antibiotics. Another important implication is that the heterogeneity of diagnostic approaches will impact on the reported incidence of VAP, significantly undermining the value of this index as a meaningful performance indicator in healthcare.

While our results suggest shortcomings in the diagnosis of VAP in current practice, several limitations must be considered in their interpretation. In particular we received a 23% response rate which is clearly too low to be entirely confident that our findings are representative. The low response rate in itself may potentially reflect a general apathy towards the subject of VAP generated by a lack of agreement with respect to the diagnostic and management approach.

Furthermore, in order to maintain anonymity and survey individual consultant practice we did not request the identity of individuals' place of work so we are unable to comment about differences in practice between units. Finally, the data collected are estimates and opinion and may not reflect an individual's actual practice.

#### CONCLUSION

There is wide variation in the diagnosis and management of VAP among UK ICU consultants. The data presented suggest that developing a standardised approach, incorporating increased radiological and microbiological support and delivery of standardised BAL protocols, might improve this situation. Until such time the use of reported VAP rates as a performance indicator within the UK healthcare system may be misleading.

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