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> Disclosures: Consultancy; Pfizer, GSK, Merck, PDI, BD, Germitec, GAMA All devices/methods discussed consistent with FDA and EPA regulations

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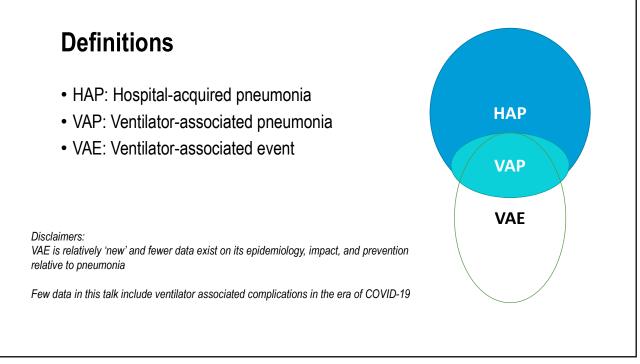
SCHOOL OF MEDICIN

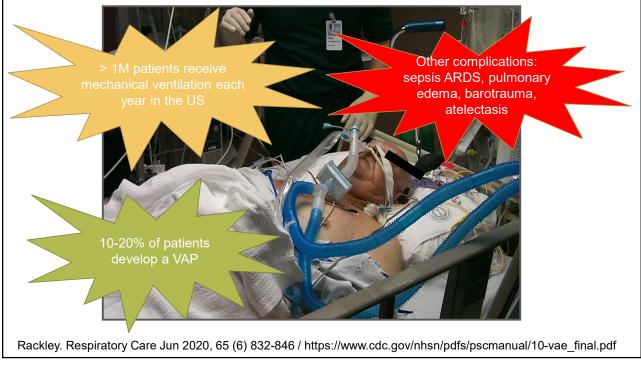
Overview	
 Ventilator associated events Surveillance Epidemiology Prevention Hospital Acquired Pneumonia Epidemiology Pathophysiology and Microbiology Diagnosis Prevention 	
Thanks to Sarah Lewis for slides	UNC SCHOOL OF MEDICINE

ESTIMATES OF HAIS OCCURRING IN ACUTE CARE HOSPITALS, US, 2011

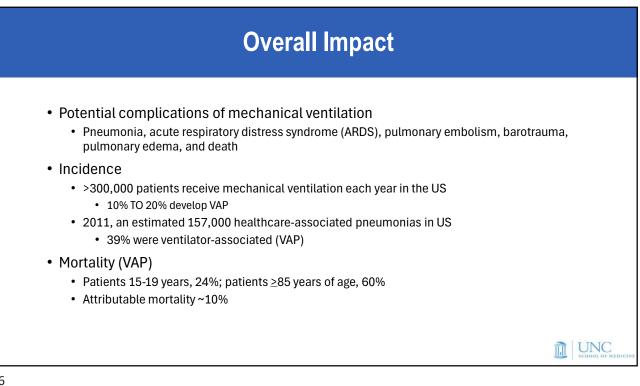
Major Site of Infection	Estimated Number (%)
Pneumonia	157,500 (21.8%)
Gastrointestinal illness	123,000 (17.0%)
Urinary tract infections	93,000 (12.9%)
Primary bloodstream infections	71,900 (10.0%)
Surgical site infections from any inpatient surgery	157,000 (21.7%)
Other types of infection	118,500 (16.3%)
Estimated total number of infections in hospitals	721,800

Magill SS, et al. New Engl J Med 2014;370:1198





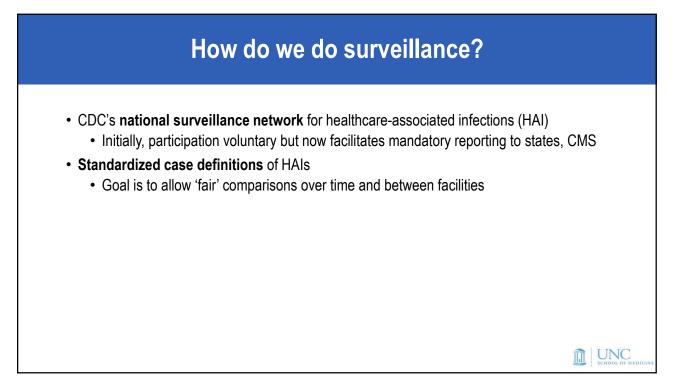


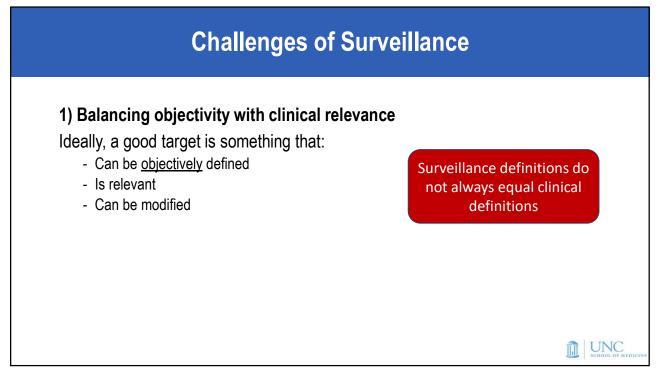


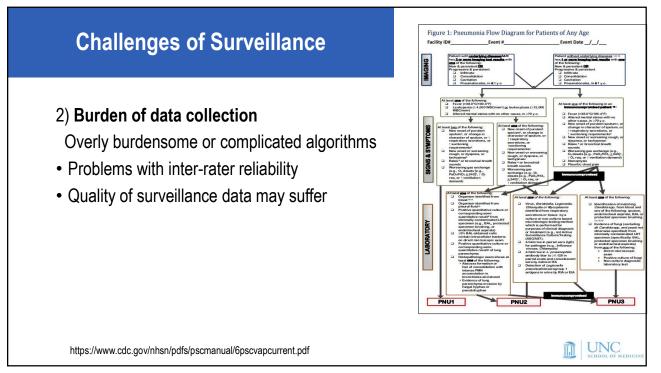
Type of Infection	Infections Identified in Survey	Surveyed Patients with Type of Infection	Estimated Infections in the United States*
	no.	% (95% CI)	no. (95% CI)
All health care-associated infections			
Pneumonia	110	24.3 (20.6-28.5)	157,500 (50,800-281,400)
Surgical-site infection	110†	24.3 (20.6-28.5)	157,500 (50,800-281,400)
Gastrointestinal infection	86	19.0 (15.6-22.8)	123,100 (38,400-225,100)
Urinary tract infection	65	14.4 (11.4-17.9)	93,300 (28,100-176,700)
Primary bloodstream infection	50	11.1 (8.4-14.2)	71,900 (20,700-140,200)
Eye, ear, nose, throat, or mouth infection	28‡	6.2 (4.2-8.7)	40,200 (10,400-85,900)
Lower respiratory tract infection	20	4.4 (2.8-6.6)	28,500 (6900-65,200)
Skin and soft-tissue infection	16	3.5 (2.1-5.6)	22,700 (5200-55,300)
Cardiovascular system infection	6	1.3 (0.5-2.7)	8,400 (1200-26,700)
Bone and joint infection	5	1.1 (0.4-2.4)	7,100 (1000-23,700)
Central nervous system infection	4	0.9 (0.3-2.1)	5,800 (700-20,700)
Reproductive tract infection	3	0.7 (0.2-1.8)	4,500 (500-17,800)
Systemic infection	1	0.2 (0.01-1.1)	1,300 (0-10,900)
Total			721,800 (214,700-1,411,000)
Infections in non-neonatal intensive care units			
Catheter-associated urinary tract infection	25	5.5 (3.7-7.9)	35,600 (9100-78,000)
Central-catheter-associated primary bloodstream infection	11	2.4 (1.3-4.2)	15,600 (3200-41,500)
Ventilator-associated pneumonia	35	7.7 (5.5-10.5)	49,900 (13,600-103,700)
Surgical-site infections attributed to Surgical Care Improvement Project procedures§	46	10.2 (7.6-13.2)	66,100 (18,700-130,300)
Hospital-onset infections caused by specific pathogens			
Clostridium difficile infection¶	56	12.4 (9.6-15.7)	80,400 (23,700-155,000)
MRSA bacteremia	7	1.5 (0.7-3.0)	9,700 (1700-29,600)

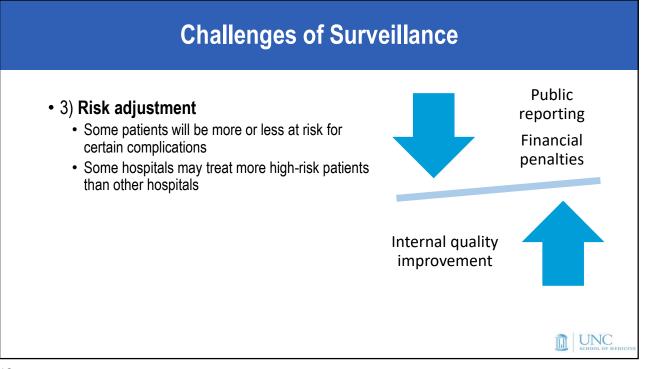


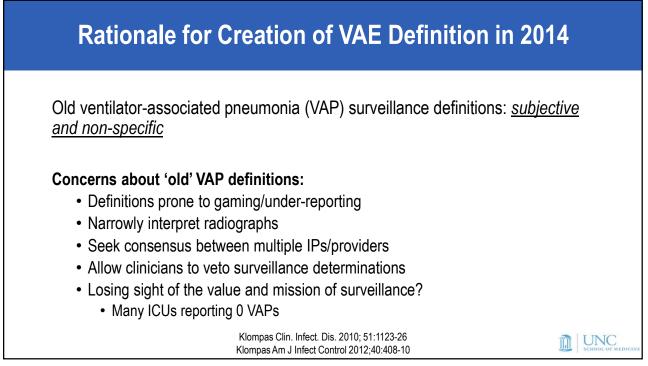


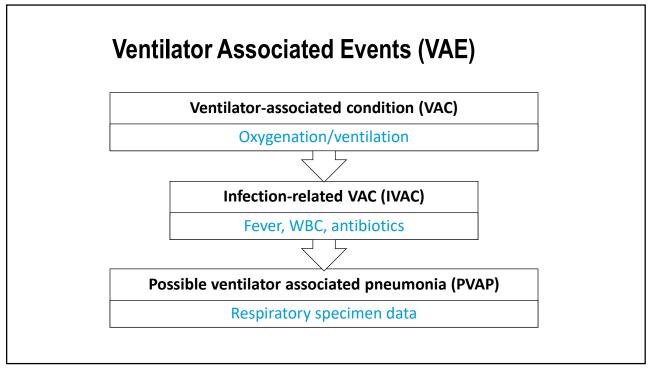


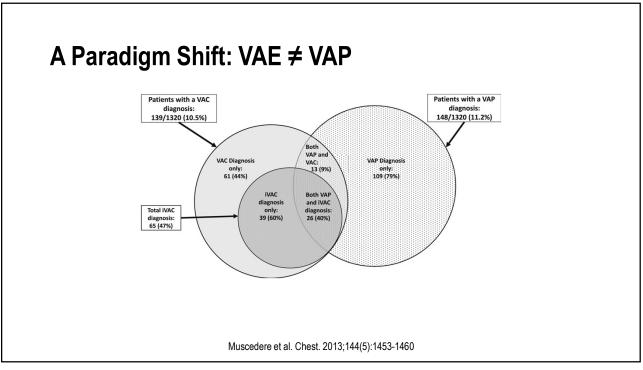












Infection-Related Ventilator-Associated Complication (n = 869)

189 (21.78)

260 (29.9) 213 (24.5)

124 (14.3)

83 (9.6)

381 (43.8)

240 (27.6)

12 (1.4)

95 (10.9)

44 (5.1)

42 (4.8)

4 (0.5)

8 (0.9)

30 (3.5)

137 (15.8)

23 (2.6)

1 (0.1)

9(1)

19 (2.2)

9 (1) 20 (2.3)

18 (2.1)

1 (0.1)

4 (0.5)

24 (2.8) 17 (2)

11 (1.3)

4 (0.5)

186 (21.4)

58 (6.7)

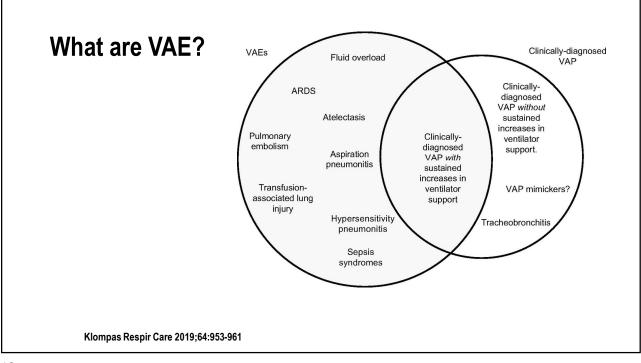
Ventilator-Associated Condition (n = 2,331)

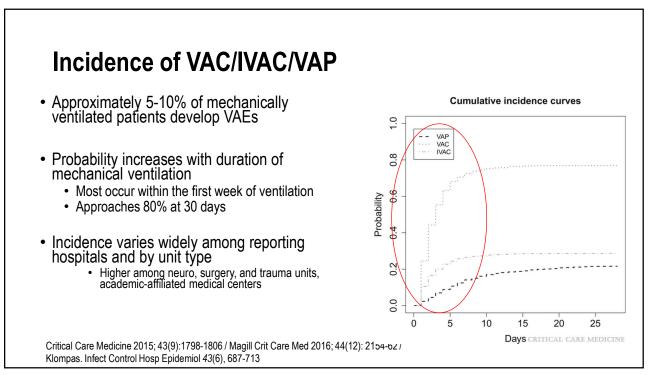
"Only VAC and IVAC ... are intended to be possible candidates for future use in public reporting, inter-facility comparisons, and payfor-performance programs. The VAC and IVAC definitions use criteria based on data anticipated to be available from most mechanically ventilated patients and less subject to manipulation or gaming. By contrast, the third definition tier, possible and probable VAP, was developed to be used only in internal quality improvement."

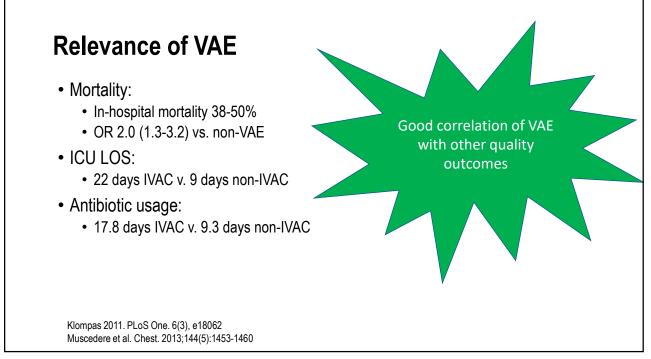
Magill et al. Clin Infect Dis 2013; 57(12):1742-46.

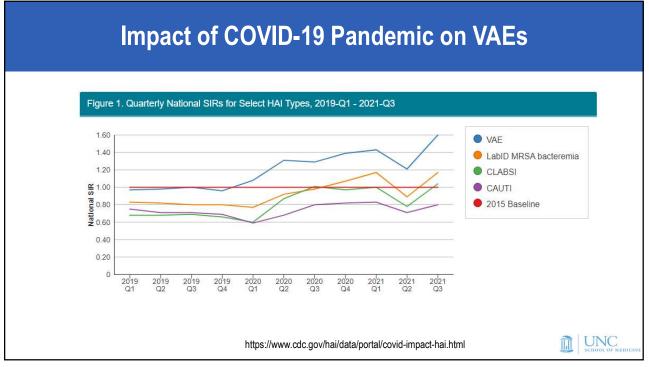
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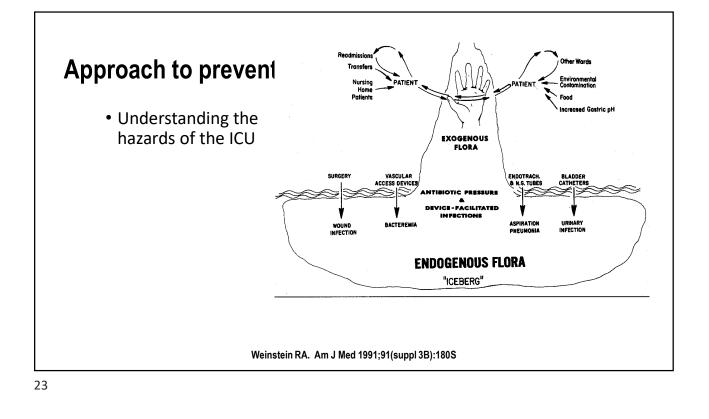
of etiologies per patie 818 (35.1) What are VAE? 726 (31.2) 445 (19.1) 214 (9.2) ≥ 4 128 (5.5) Nosocomial infections 637 (27.3) Ventilator-associated pneu 339 (14.5) Retrospective study- 3028 patients 1996-Tracheobronchitis 23(1) Bloodstream infection 173 (7.4) 2012 on mechanical ventilation >= 5 days Catheter-related infection 81 (3.5) Urinary infection 102 (4.4) 5 (0.2) Viral infection 10 (0.4) Surgical site infections latrogenic adverse events VAE are COMMON 41 (1.8) 322 (13.8) 37 (1.6) Pneumothorax • 77% of patients with at least 1 VAC Failure of planned extubatio 11 (0.5) cidental extubation 21 (0.9) · 29% of patients with at least 1 IVAC Self-extubation 71 (3) 14 (0.6) 52 (2.2) Venous puncture accident Atelectas Peripheral thrombosis 36 (1.5) There are many etiologies of VAE 9 (0.4) Pulmonary embolism Myocardial infarction 10 (0.4) · Infectious complications (not just pneumonia) Cardiac arrest 43 (1.8) 29 (1.2) common Cardioversi Gastrointestinal bleeding 26(1.1) · Non-infectious complications not directly related 5 (0.2) Acute mesenteric infa Intestinal pseudo-obstructio 2 (0.1) to mechanical ventilation also play role Transport 387 (16.6) Fluid resus 123 (5.3) Critical Care Medicine43(9):1798-1806, September 2015

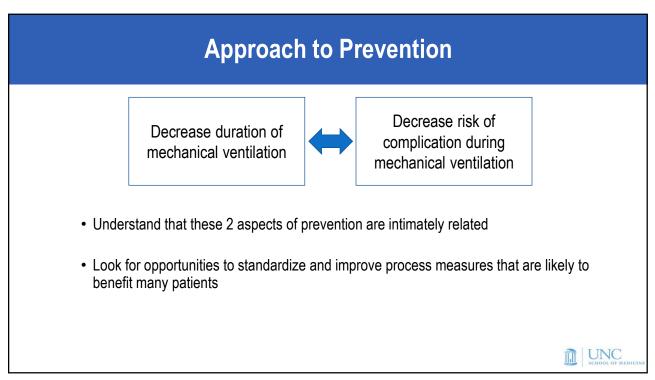


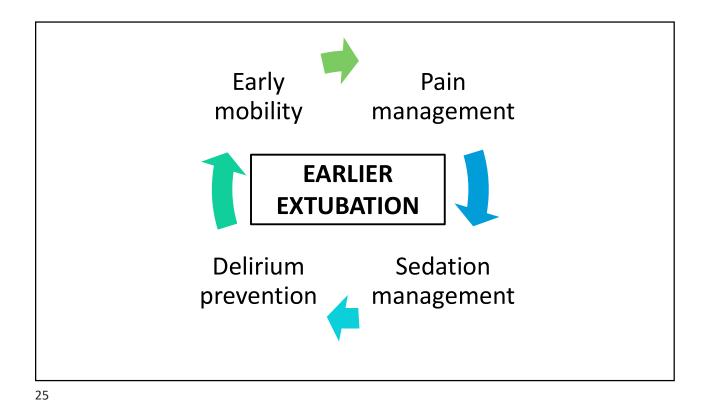


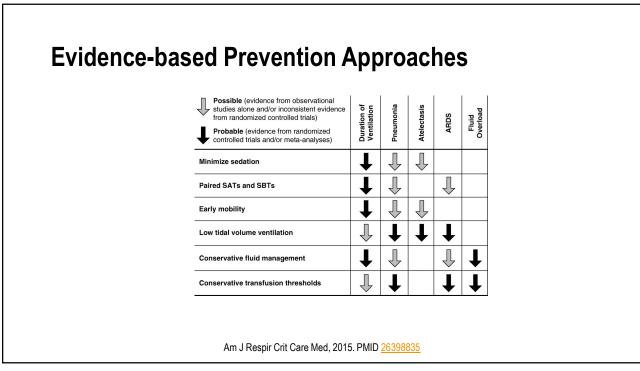


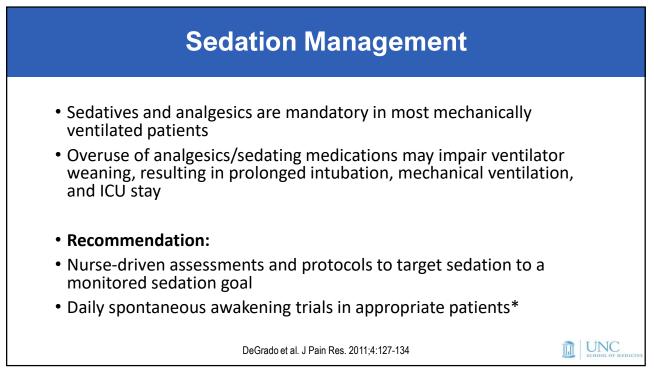


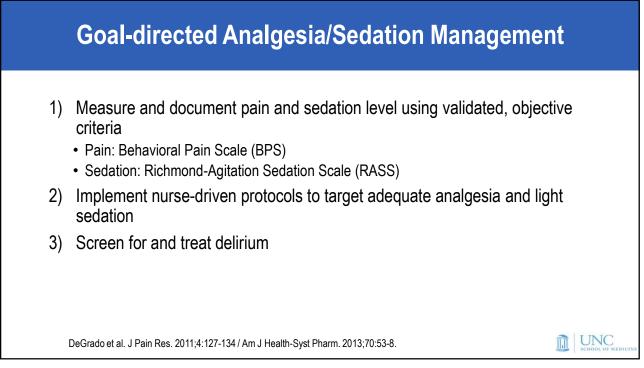


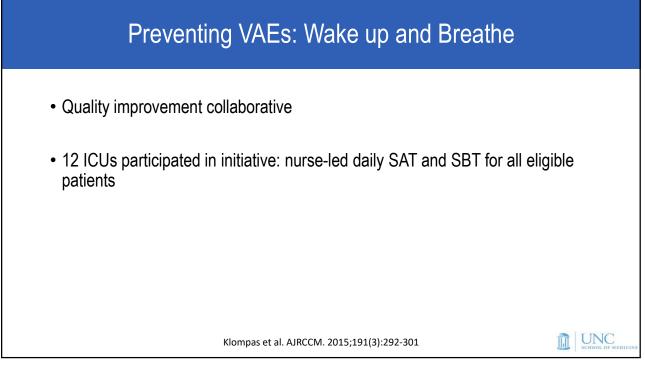


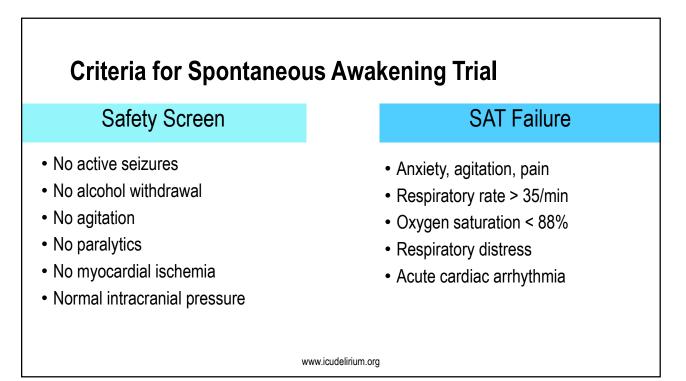


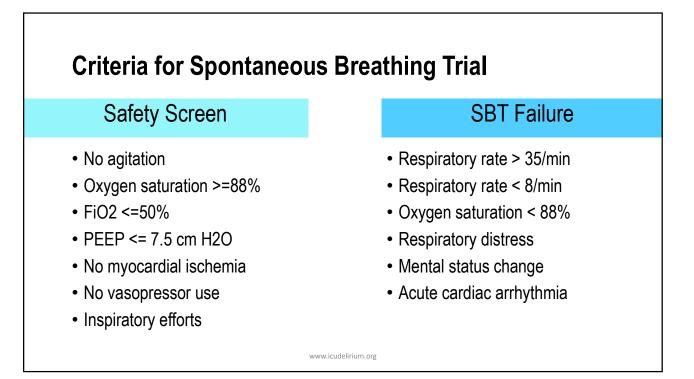


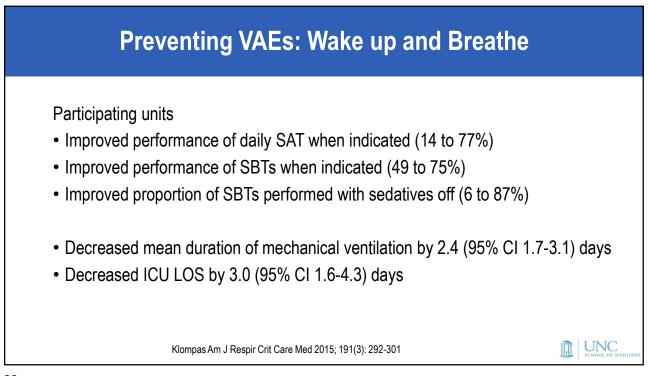


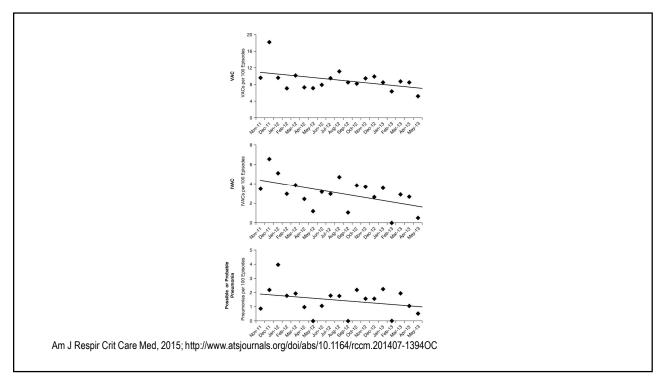


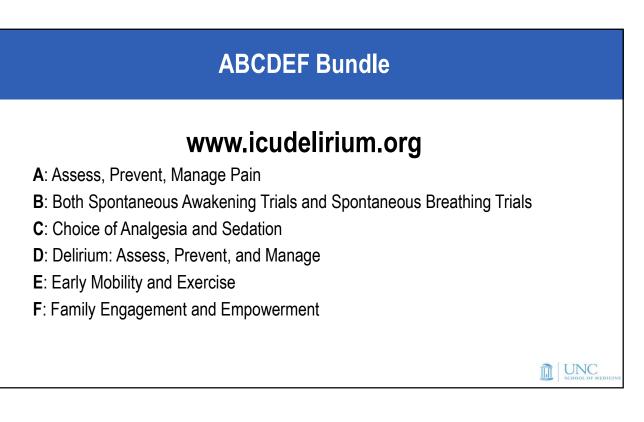






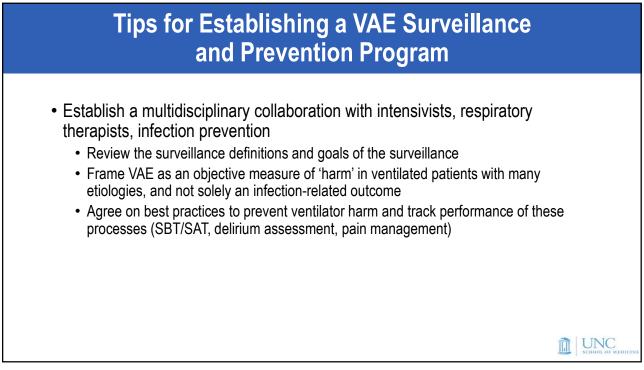


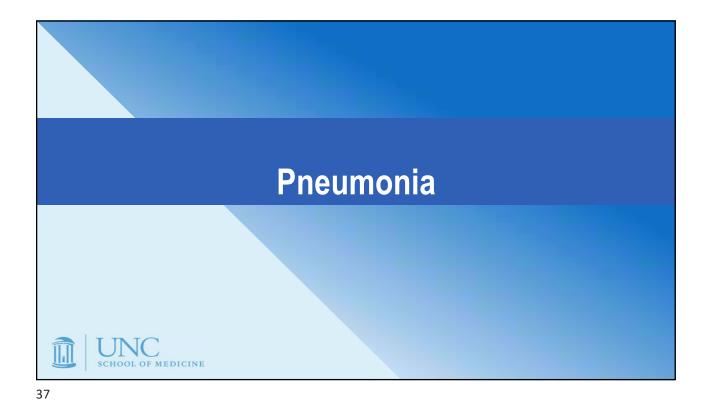




What VAEs Are and Are Not

	What They Are	What They Aren't	
Intent	Surveillance concept	Clinical diagnosis	
Surveillance	Objective and reproducible	Sensitive/specific for VAP	
Etiology	Many potential causes including non-infectious ones	Proxy for pneumonia	
Morbidity	Highly morbid	Not benign	
Prevention strategy	 Re-think prevention bundles: Minimize sedation Early mobility Low tidal volume ventilation Conservative fluid management 	Not fully preventable by traditional bundles	
	Michael Klompas Respir Care 2019;64:953-96	n m U	





Pneumonia and VAE Surveillance: Current State for Many IP programs

	PNEU	VAE
Surveillance	 Selectively performed on cases of BSI in patients with central venous catheters to determine if criteria met for secondary attribution (all programs) 	 Performed on all patients on mechanical ventilation > 4 days
Clinical relevance	 Poor correlation between clinical and surveillance definitions of pneumonia 	 Not specific for an individual clinical presentation – represents a large group of conditions
Prevention	Hand hygiene, avoid ventilation when possib elevate head of bed, minimize unnecessary	ole, early mobility, pain/sedation management, devices, antibiotic stewardship

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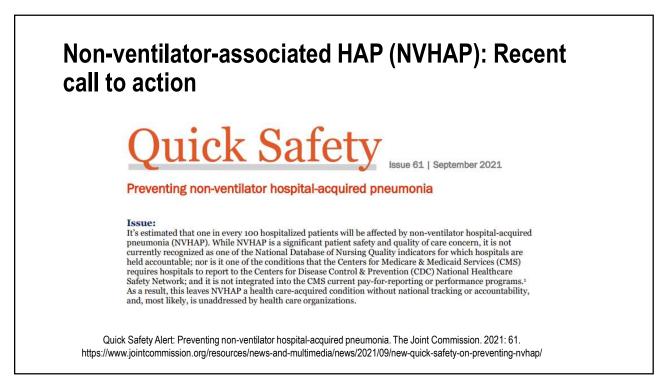
Non-ventilator- associated HAP (NVHAP): Recent call to action

Clinical Relevance

- 1% of all hospitalized patients develop NVHAP
- Crude mortality 15-30%
- Extends LOS up to 15 days
- · Increases antibiotic utilization
- · Increases risk for readmissions

Munro, S., Baker, D., Giuliano, K., Sullivan, S. Haber, J., Jones, B., Klompas, M. (2021). Nonventilator hospital-acquired pneumonia: A call to action: Recommendations from the National Organization to Prevent Hospital-Acquired Pneumonia (NOHAP) among nonventilated patients. *Infection Control & Hospital Epidemiology*, *42*(8), 991-996





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Non-ventilator- associated HAP (NVHAP): Recent call to action

Current Gaps

No current surveillance definition or methodology

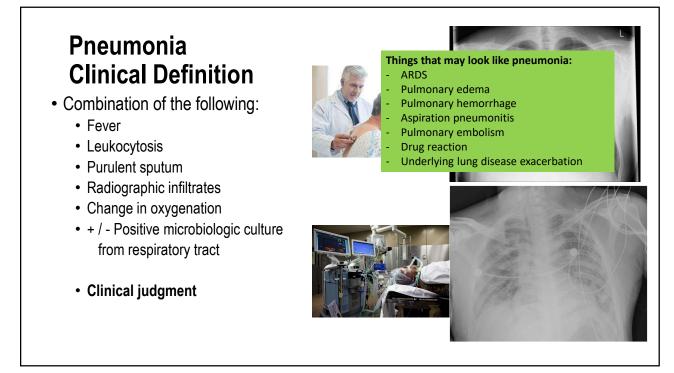
Big questions

- How can we improve the reproducibility, relevance, and efficiency of surveillance for HAP?
- Do we fully understand the mechanism of NVHAP to inform prevention strategies?
- What are the best-performing interventions to prevent NVHAP?

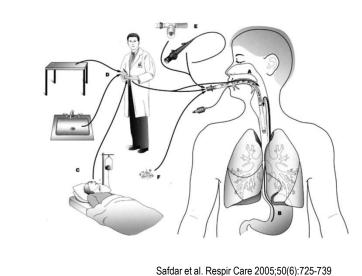
In absence of data

- Promote early mobility
- Screen for and manage dysphagia to reduce risk of aspiration
- Decrease risk of hospital transmission of respiratory viruses
- Perform regular oral care

Munro, S., Baker, D., Giuliano, K., Sullivan, S., Haber, J., Jones, B., . . . Klompas, M. (2021). Nonventilator hospital-acquired pneumonia: A call to action: Recommendations from the National Organization to Prevent Hospital-Acquired Pneumonia (NOHAP) among nonventilated patients. Infection Control & Hospital Epidemiology, 42(8), 991-996

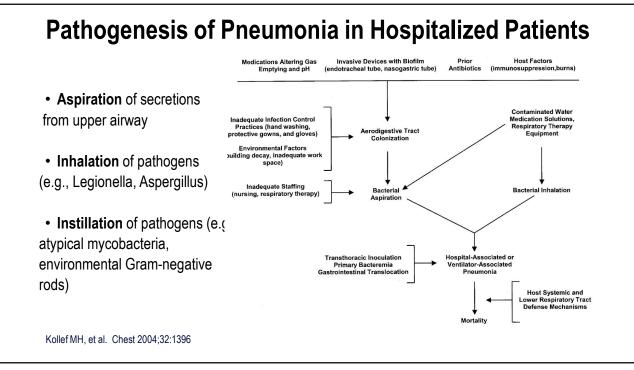


Healthcare-Associated Pneumonia (HAP) Pathogenesis



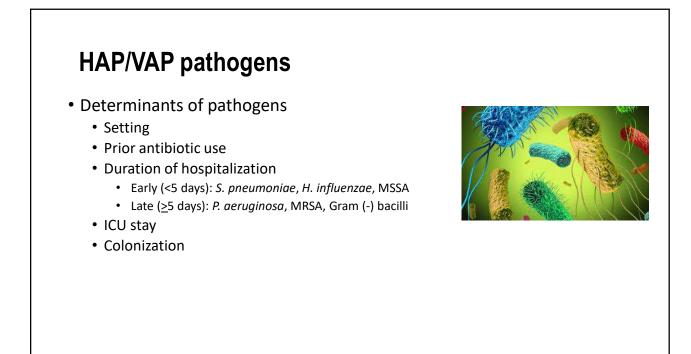
Aerodigestive tract colonization

- Colonization of the aerodigestive tract may occur **endogenously** (A and B) or **exogenously** (C through F)
- Exogenous colonization may result in primary colonization of the oropharynx or may be the result of direct inoculation into the lower respiratory tract during manipulations of respiratory equipment (D), during using of respiratory devices (E), or from contaminated aerosols (F).

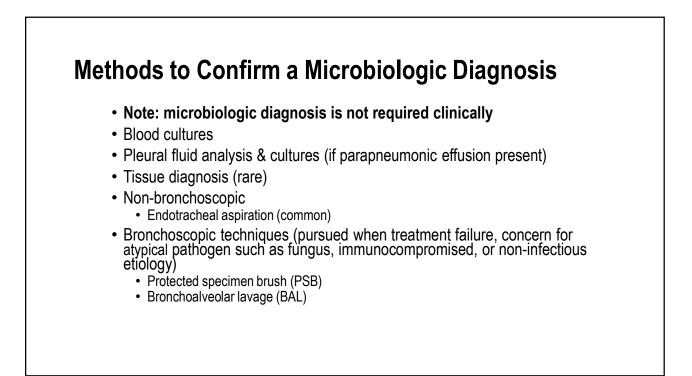


Host-related risk factors	Intervention-related risk factors
Medical history and underlying illness Male gender Extreme age Prior central nervous system disorder Immunocompromised Acute underlying diseases Emergent surgery Neurosurgery Thoracic surgery Cardiac surgery Burns Re-intervention Acute severity factors Organ system failure index of at least 3 Acute renal failure Acute respiratory distress syndrome ECMO, intra-aortic support Ulcer disease	Peri-operative transfusion of blood products Duration of the mechanical ventilation Reintubation Supine head position in patients receiving enteral nutrition Antibiotic therapy [®] Enteral nutrition Absence of subglottic secretion drainage [®] Intra-hospital transports Continuous sedation, use of paralytic agents Nasogastric tubes Tracheostomy Frequent ventilator circuit changes Intracuff pressure of less than 20 cm H ₂ O





Community acquired aspiration	Hospital acquired aspiration	Inhalational	Hematogenous
 Haemophilus influenzae Streptococcus pneumoniae Oropharyngeal streptococci and anaerobes 	 Oropharyngeal streptococci and anaerobes Enterobacteriaceae Pseudomonas 	 Fungi Legionella Viruses Mycobacteria 	 Staph aureus (common) Enterobacteriaceae (uncommon)



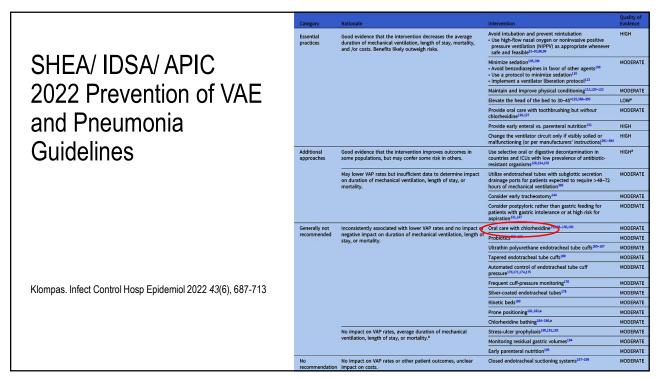
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Preventing HAP/VAP: An Important Target for Antimicrobial Stewardship

Pathogen	Incidence and resistance trends			
MRSA	Rate in VAP: 12-42%°			
	Rate of methicillin resistance is decreasing: 1.4–82% ^b			
Pseudomonas aeruginosa	Rate in VAP: 21–61% especially for the second episode of VAP ^a			
	MDR/XDR rates as high as 38–46% with 8–20% susceptible only to colistin [12–14]			
	Meropenem with >10% increase in resistance in North America with susceptibility across all classes of antimicrobials at 60–71% [10]			
Enterobacteriaceae	Rate in VAP: 5–19.1% with rising rates of resistance to all classes of antimicrobials ^a [9,10,13]			
	Rates of ESBL of 40% in Asia [9]			
Acinetobacter spp.	Rate in VAP: 4.8–36.5% (highest in Latin America and Asia) [9,10,13]			
	MDR rate as high as 80% and XDR 50% with 30% susceptible only to colistin [9,10,13]			
	Meropenem and doripenem with >10% increase in resistance [10], colistin-resistant cases reported [13]			

Abbreviations: ESBL, extended spectrum β -lactamases; MDR/XDR, multidrug resistant/extremely drug resistant; MRSA, methicillin-resistant *Staphylococcus aureus*; SA, *Staphylococcus aureus*; VAP, ventilator-associated pneumonia.

Guillamet CV, Kollef MH. Curr Opin Crit Care 2015;21:430-8



Category	Rationale	Intervention	Quality of Evidence
Essential	May lower VAP and/or PedVAE rates and have minimal risks of harm. Benefits likely outweigh potential risks.	Use non-invasive positive pressure ventilation in selected populations ^{62,205,206}	HIGH
	nami penena meny astrongi potential nata	Minimize the duration of mechanical ventilation	HIGH
		Use caffeine therapy to facilitate extubation ^{396,397}	HIGH
		Assess readiness to extubate daily	LOW
		Manage patients without sedation whenever possible ^{209,210}	LOW
		Avoid unplanned extubations and reintubations ²¹²	LOW
		Avoid reintubation by using nasal CPAP, non-invasive positive pressure ventilation (NIPPV), of high flow nasal cannula in the post-extubation period ³⁹⁶	HIGH
		Provide regular oral care with sterile water	LOW
		Change the ventilator circuit only if visibly soiled or malfunctioning ²⁵⁹ (or per manufacturer's instructions)	LOW
Additional	Unknown impact on VAP and VAE rates but risk of harm likely minimal. Reasonable to consider implementing if rates remain elevated despite essential practices.	Lateral recumbent positioning ²¹⁵	LOW
approaches		Reverse Trendelenberg positioning	LOW
		Closed/in-line suctioning systems ^{216,217}	LOW
		Oral care with maternal colostrum ²¹⁸	MODERATE
Generally not recommended		Regular oral care with an antiseptic or Biotene ²¹⁹	LOW
	May be harmful. Risk-benefit balance does not favor	Histamine-2 receptor antagonists ^{220,221}	MODERATE
		Prophylactic broad-spectrum antibiotics ^{222–225}	MODERATE
		Daily spontaneous breathing trials ^{398,399}	LOW
		Daily sedative interruptions	LOW
		Prophylactic probiotics or synbiotics ^{228,229}	LOW
	Not recommended because appropriate products are not available or approved for use in this population.	Endotracheal tubes with subglottic secretion drainage ports	NA
		Silver-coated endotracheal tubes	NA

Category	Rationale	Intervention	Quality of Evidence	
Essential practices	Interventions with minimal risk of harm and some data that they may	Avoid intubation if possible. Use non-invasive positive pressure ventilation for selected populations ²⁴⁰⁻²⁴²	MODERATE	
	lower VAP rates, PedVAE rates, and/	Assess readiness to extubate daily in patients without contraindications ⁷⁴⁴⁻²⁴⁸	MODERATE	1
	or duration of mechanical ventilation.	Take steps to minimize unplanned extubations and reintubations ²⁴⁹	LOW	ĺ
		Avoid fluid overload ^{251,253,254}	MODERATE	
		Provide regular oral care (i.e., toothbrushing or gauze if no teeth) ^{234,256,257}	LOW	
		Elevate the head of the bed unless medically contraindicated ²³⁴	LOW	
		Change ventilator circuits only if visibly soiled or malfunctioning ²⁵⁹ (or per manufacturer's instructions)	MODERATE	
		Prevent condensate from reaching the patient ^{234,266}	LOW	
		Use cuffed endotracheal tubes ^{262–264}	LOW	
		Maintain cuff pressure and volume at the minimal occlusive settings	LOW	
		Suction oral secretions before each position change	LOW	
Additional	Risk of harm likely minimal with some	Interrupt sedation daily ²⁶⁷	MODERATE	0
approaches	evidence of benefit in adult patients, but data in pediatric populations are limited. Reasonable to consider implementing if rates remain elevated	Utilize endotracheal tubes with subglottic secretion drainage ports for older pediatric patients expected to require >48 or 72 hours of mechanical ventilation ³⁹⁵	LOW	
	despite essential practices.	Consider early tracheostomy ²⁶⁸⁻²⁷⁰	LOW	1
recommended inadequat	Unknown impact on VAP rates and inadequate data on risks.	Prolonged systemic antimicrobial therapy for ventilator-associated tracheitis ⁷⁷	LOW	
	No impact on VAP rates. ^a	Selective oropharyngeal or digestive decontamination ²⁷⁴	LOW	
		Prophylactic probiotics ¹⁶³	LOW	
		Oral care with antiseptics such as chlorhexidine ^{280,284,285}	MODERATE	(
		Stress-ulcer prophylaxis ²⁸⁶⁻²⁸⁸	LOW	
	Lowers VAP rates in adults but no impact on duration of mechanical ventilation, length of stay, or mortality.	Silver-coated endotracheal tubes	LOW	
No recommendation	Limited data on pediatric patients, no impact on VAP rates or outcomes in adults, unclear impact on costs	Closed or in-line suctioning ²⁰³	LOW	

