

## Understanding secondary bacterial infections associated with COVID-19 and influenza

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*Evaluating how a nasal colonization risk mitigation strategy can help improve patient outcomes*



# Housekeeping

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Introduction

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*Consultant*

*Disclosure: Gwen Borlaug, member of the Global Life Technologies Corp. Speaker's Bureau*



# Learning Objectives

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- Describe how secondary bacterial infections impact patients with respiratory viral infections such as COVID-19 and influenza.
- Describe the interactions among influenza virus, the nasal microbiome and *Staphylococcus aureus*.
- Identify nasal and skin decolonization as a potential strategy to mitigate risk of MSSA and MRSA secondary infections associated with respiratory viral infections.
- List the benefits of universal, house-wide nasal and skin decolonization for hospital patients.



# Topics

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- I. Secondary bacterial infection in viral respiratory disease
- II. Bacterial colonization: A risk factor for secondary bacterial infection
- III. Exploring risk mitigation strategies
- IV. Universal nasal decolonization: A strategy to prevent healthcare-associated *S. aureus* infections



# Secondary bacterial infections following viral respiratory disease

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## Section I



# Secondary bacterial infections (SBI)

**Bacterial co-infection** – a bacterial infection occurring simultaneously with onset of respiratory viral infections

**Secondary bacterial infection** – most commonly presents as bacterial infection (e.g. pneumonia), occurring after onset or in recovery phase of respiratory viral infections

These infections are associated with:

- greater severity of illness
- greater use of healthcare resources
- increased risk of death



# Secondary bacterial infection among COVID-19 patients

SBI in COVID-19 patients increase morbidity, mortality and antimicrobial resistance (AMR) threat



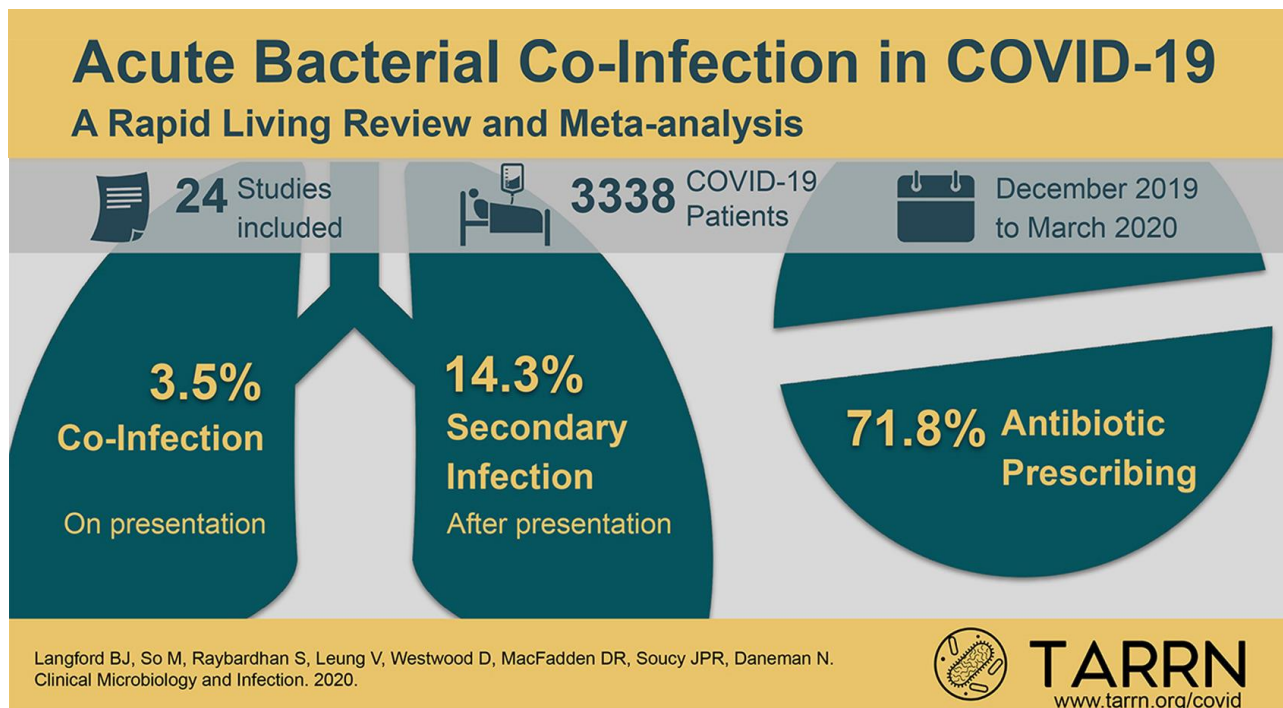
## Most prevalent pathogens:

*S. aureus*  
*S. pneumoniae*  
*N. meningitidis*  
*H. influenzae*  
*K. pneumoniae*

- Hospital acquired-infection in **13.5%** patients (Yang et al.)
- **50%** of non-survivor cases had SBI
- VAP in **31%** of patients requiring invasive respiratory support (Zhou et al.)
- **57.9%** of severely and critically ill patients developed secondary bacterial infections. (Zhang et al.)
- SBI developed at a median of **17** days after illness onset (Zhou et al.)



# Acute bacterial co-infections associated with COVID-19 illness



**4.9%**  
(2.6 to 7.1)  
**Co-infection**

**16%**  
(12.4 to 19.6)  
**Secondary Infection**

**UPDATE:** October 29, 2020  
38 studies

Accessed February 22, 2021



# Multicenter, Case-Control Study

## Risk Factors and Outcomes of Hospitalized Patients with Severe COVID-19 and Secondary Bloodstream Infections: A Multicenter, Case-Control Study

- 34% (128/375) of COVID-19 patients developed secondary BSI
- 91% caused by bacterial pathogens
  - *S. aureus*
  - *Enterococcus spp.*
  - *E. coli*
- 51% were hospital-associated
- Sources
  - Unknown
  - CLABSI
- Increased hospital stays and worse clinical outcomes
- Conclusions
  - Antimicrobial measures in hospitals need to be improved
  - Further studies need to develop prevention and treatment protocols



# Secondary bacterial infections among COVID-19 patients

Retrospective observational case series of patients with coronavirus disease 2019 (COVID-19) who developed secondary *S. aureus* bacteremia across 2 New York City hospitals.

42 hospital patients with secondary *S. aureus* bacteremia

55% and 67% died at 14 days and 30 days respectively following first positive blood culture

Independent risk factors for 14 day mortality included hospital onset bacteremia and age



The screenshot shows the Open Forum Infectious Diseases website. The header is orange with the journal name. Below it is a dark navigation bar with links for Issues, More Content, Publish, Alerts, and About. A small box on the right says 'All Open Forum Infectious Diseases'. The main content area has a yellow sidebar on the left with a journal cover image and a list of article contents: Abstract, METHODS, RESULTS, DISCUSSION, CONCLUSIONS, Acknowledgments, and References. The main text area on the right features the article title 'Staphylococcus aureus Bacteremia in Patients Infected With COVID-19: A Case Series' with an orange icon. Below the title is the author list: Jaclyn A Cusumano, Amy C Dupper, Yesha Malik, Elizabeth M Gavioli, Jaspreet Banga, Ana Berbel Caban, Devika Nadkarni, Ajay Obla, Chirag V Vasa, Dana Mazo, and a 'Show more' link. There is also a link to 'Author Notes'. The publication information states 'Open Forum Infectious Diseases, Volume 7, Issue 11, November 2020, ofaa518, https://doi.org/10.1093/ofid/ofaa518' and 'Published: 12 November 2020'. Below this are icons for PDF, Split View, Cite, Permissions, and Share. The article's abstract is displayed in a light blue box, starting with 'Background' and describing the study's findings on secondary bacterial infections in COVID-19 patients.

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**Staphylococcus aureus Bacteremia in Patients Infected With COVID-19: A Case Series** 

Jaclyn A Cusumano , Amy C Dupper, Yesha Malik, Elizabeth M Gavioli, Jaspreet Banga, Ana Berbel Caban, Devika Nadkarni, Ajay Obla, Chirag V Vasa, Dana Mazo ... [Show more](#)

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Open Forum Infectious Diseases, Volume 7, Issue 11, November 2020, ofaa518, <https://doi.org/10.1093/ofid/ofaa518>

Published: 12 November 2020 [Article history ▾](#)

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**Abstract**

**Background**

Previous viral pandemics have shown that secondary bacterial infections result in higher morbidity and mortality, with *Staphylococcus aureus* being the primary causative pathogen. The impact of secondary *S. aureus* bacteremia on mortality in patients infected with severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) remains unknown.



# Secondary bacterial infections associated with COVID-19: Risk



Endogenous colonization with flora harboring antimicrobial resistance

Data is suggestive of nosocomial transmission of hospital organisms in critically ill ventilated patients

## Colonization is a risk:

- COPD is comorbidity in severe COVID-19. COPD patients are colonized by bacterial pathogens even at the stable phase of the disease, making it likely that SBI infection occurs in patients already colonized with bacteria.

## Nosocomial acquisition likely:

- The median LOS of COVID-19 patients: 7 days (can reach  $\geq 14$  days)
- Risk of a hospital-associated pneumonia increases significantly the longer the hospitalization period.
- More than 90% of hospital-associated pneumonias are associated with mechanical ventilation, one of the therapeutics used in COVID-19 patients admitted in the ICU.



# SBI in COVID-19: *Statement from front-line intensive care experts*

The experts suggest closely monitoring the signs of **secondary infection**, especially in critically ill patients with COVID-19 who have been admitted to ICU > 48 h (expert opinion).

## Rationale

- Both long course of the disease and immunosuppressive state place the severe and critical COVID-19 patients at a high risk of secondary infection (including bacteria and fungus).
- The data on the epidemiology of secondary infection in COVID-19 patients are lacking.
- Based on the evidence from H1N1, **secondary infection** is **very common** in patients admitted to ICU > 48 h.
- Strategies for **preventing healthcare-acquired infections** should be effectively implemented, and multiple site samples (blood, sputum, etc.) should be routinely collected to monitor the signs of secondary infection.





The national standard infection ratio for central  
line-associated bloodstream infections increased:



in 2020 Q2 vs. 2019 Q2



Source: Patel PR, et al. *Infect Control Hosp Epidemiol*. 2021;doi:10.1017/ice.2021.108.



# Secondary bacterial infections associated with influenza

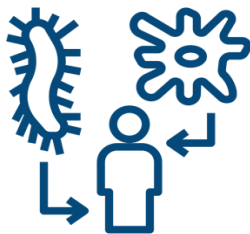
Influenza virus infects 5%–20% of the US population yearly, with 23,000 to 61,000 deaths annually.

Up to 75% of those infected with influenza that go on to acquire pneumonia are confirmed to have SBI

USA: influenza and pneumonia currently rank 8th overall as a cause of death, annually.

In seasonal epidemics, SBI associated with influenza is associated with:

- increases in hospital admissions
- more severe symptoms
- increases in mortality (mortality rates ranging from 11- to 15-fold higher than those of influenza alone)



## Most common causative pathogens

*Streptococcus pneumoniae*  
*Staphylococcus aureus*,  
*Haemophilus influenzae*





# Secondary bacterial infections during influenza pandemics

## 1918-1919 PANDEMIC | SPANISH FLU | H1N1 STRAIN



20-40%  
of the global  
population  
became ill



THREE  
waves of the  
pandemic  
occurred



675K  
people died  
in the United  
States



50 MIL  
people died  
across the  
world

>WWI

SPANISH FLU  
killed more  
people than  
World War I

>95% of deaths attributable to  
secondary bacterial pneumonia (est.)

***Streptococcus pneumoniae*** most  
frequently recovered etiologic agents

## 1957-1958 PANDEMIC | ASIAN FLU | H2N2 STRAIN



CHINA  
was the  
first country  
to identify  
the strain



MAY 1957  
vaccine production  
began with  
limited supply  
available



SCHOOL AGE  
children spread  
in classrooms &  
brought home  
to families



IMMUNITY  
to strain  
was rare in  
people younger  
than 65



70K  
deaths in the  
U.S. caused  
by this  
pandemic

1.5 million deaths worldwide

***Staphylococci*** assumed a novel  
prominence as the leading etiologic  
agent

Secondary **staphylococcal** infections  
continued to be seen through the  
second wave of the “Asian flu” in  
1960–1961



# Secondary bacterial infections during influenza pandemics

## 1968-1969 PANDEMIC | HONG KONG FLU | H3N2 STRAIN

**DEATHS  
PEAKED**

September  
1968–March 1969

**34K**

total deaths  
in the  
United States

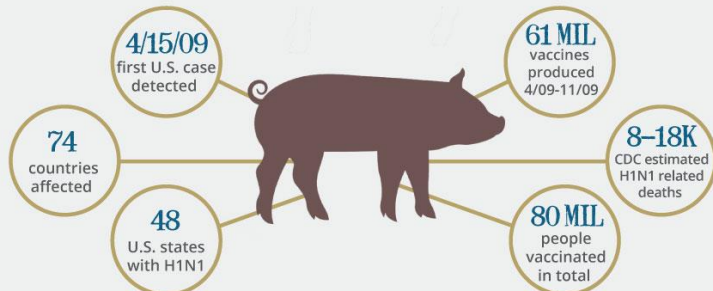
**MILDEST**

flu pandemic  
of the 20th  
Century

**VIRUS**

still  
circulates  
today

## 2009-2010 PANDEMIC | SWINE FLU | H1N1 STRAIN



SOURCES:  
Centers for Disease Control and Prevention  
archives.gov  
flu.gov



HEALTH SCIENCE CENTER  
TEXAS A&M UNIVERSITY  
transformingHEALTH

Higher risk age/group: elderly

A high correlation between pneumonia, especially **staphylococcal** pneumonia, and influenza infection was documented.

Higher risk age/group: children, teens and young adults

30-55% of case mortality associated with bacterial pneumonia

***S. pneumoniae*** most common bacteria identified

Pediatric ICU studies:

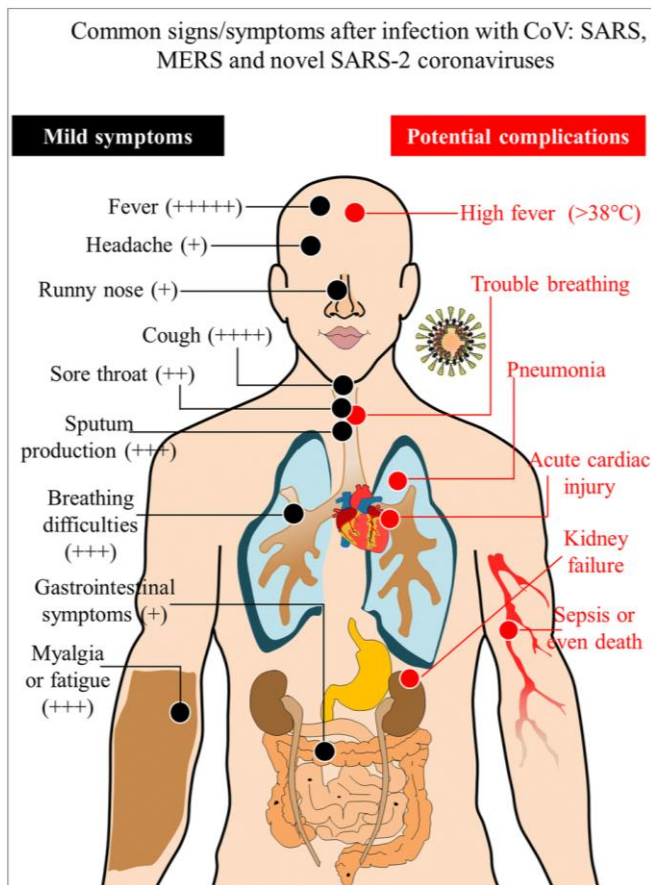
- 33% with SBI, ***S. aureus*/MRSA** most common (26%)
- 51% with SBI, ***S. aureus*** most common (35%)



Morris D. et al. Front. Microbiol 2017. 8:1041  
MacIntyre, C.R., et al. BMC Infect Dis 18, 637 (2018)  
Schwarzmann SW, et al. Arch Intern Med, 1971, vol. 127(pg. 1037-41)



# Secondary bacterial infections during previous coronavirus outbreaks



**SARS-CoV** (2002) and **MERS-CoV** (2012) caused severe pneumonia and death.

SARS-CoV: up to 30% of patients diagnosed with secondary bacterial infections (positively associated with disease severity)

Common etiologic agents:

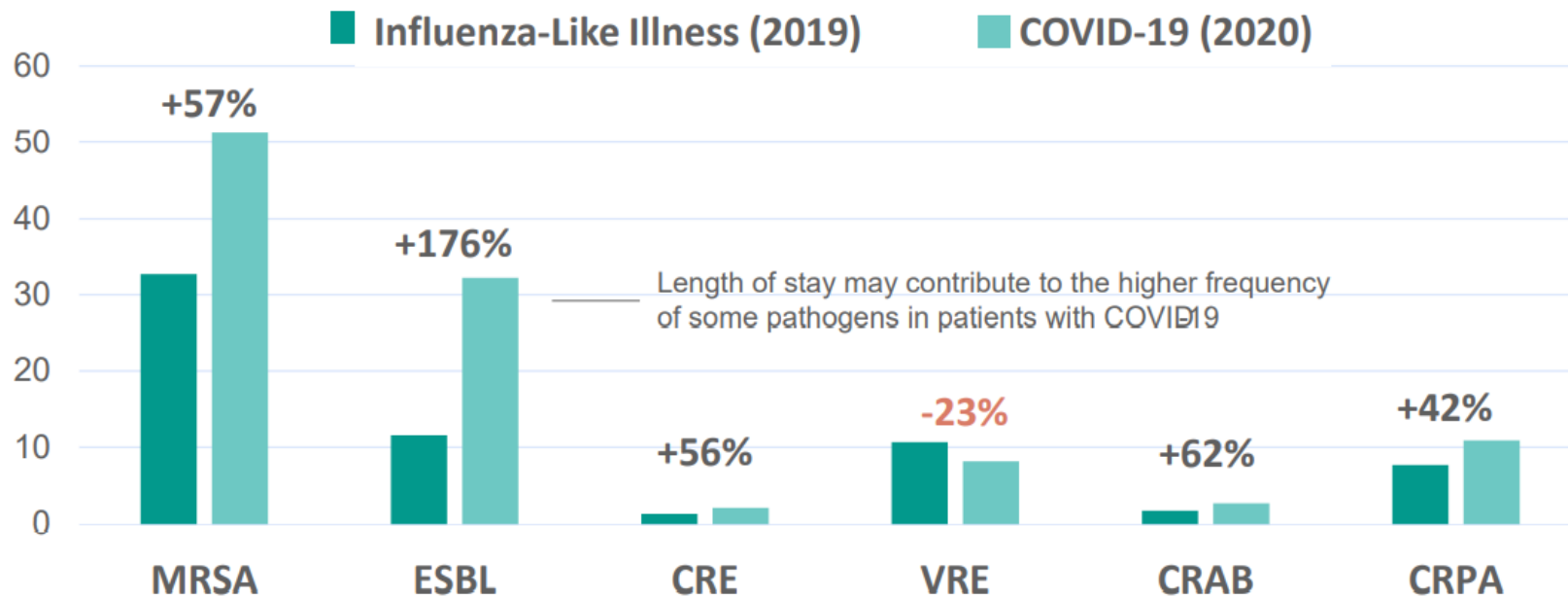
- SARS-CoV—**MRSA**, *Klebsiella*, *P. aeruginosa* and *Streptococcus*
- MERS—**MRSA**, others included carbapenem-resistant *Acinetobacter baumannii*, VRE and *S. pneumoniae*





# Antibiotic-Resistant Pathogens in Hospitalized Patients: Hospital-onset

Rate of hospital-onset resistant organisms per 10,000 discharges



The Intersection of Antibiotic Resistance (AR), Antibiotic Use (AU), and COVID-19 (hhs.gov)



Of the six pathogens, MRSA and ESBL contribute the most to total costs nationally due to their high burden despite having lower healthcare costs per case. The paper was published in *Clinical Infectious Diseases* in early 2021.





# Key Takeaways

- **Healthcare infection control is critical to fighting AR and SARS-CoV-2 infections**
  - No clear evidence that COVID-19 patients are more susceptible to bacterial/fungal infections—similar frequency as patients with influenza-like illness (ILI). **However, we are seeing sporadic outbreaks of AR infections in COVID units & higher rates of hospital onset infections**
  - COVID-19 creates perfect storm for AR infections **in healthcare settings:** length of stay, crowding, sick patients, antibiotic use, infection control issues
- **Antibiotic use fluctuated, appears stable but remains too high**
  - Hospitals: Spiked in early 2020 but flattened as pandemic continued
  - Outpatient, nursing homes: Significant drops from previous years
- Highlights continued importance of infection control and antibiotic stewardship—**both are dependent on the resiliency of these programs**



# **Bacterial colonization: a risk factor for secondary bacterial infection**

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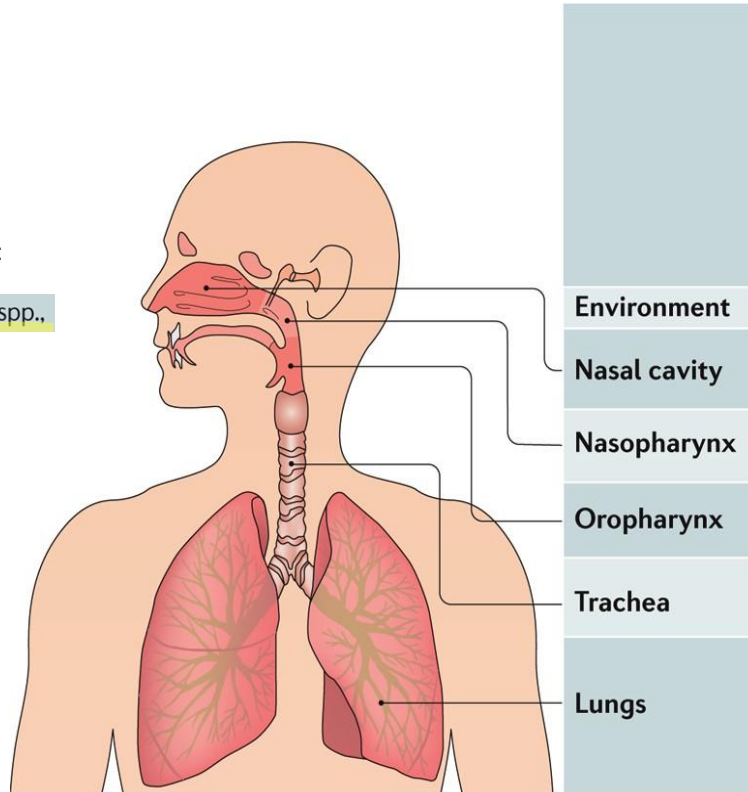
## Section II



# Colonization in respiratory tract

Anterior nares:  
reservoir

*Staphylococcus* spp.,



Viral infections are associated with increased colonization by potentially pathogenic bacteria (known as “pathobionts” or **opportunistic pathogens**).

*Staphylococcus* spp.,  
*Propionibacterium* spp.,  
*Corynebacterium* spp.,  
*Moraxella* spp. and *Streptococcus* spp.

*Moraxella* spp., *Staphylococcus* spp.,  
*Corynebacterium* spp.,  
*Dolosigranulum* spp., *Haemophilus* spp.  
and *Streptococcus* spp.

*Streptococcus* spp., *Rothia* spp.,  
*Veillonella* spp., *Prevotella* spp. and  
*Leptotrichia* spp.

*Prevotella* spp., *Veillonella* spp.,  
*Streptococcus* spp. and  
*Tropheryma whippelii*

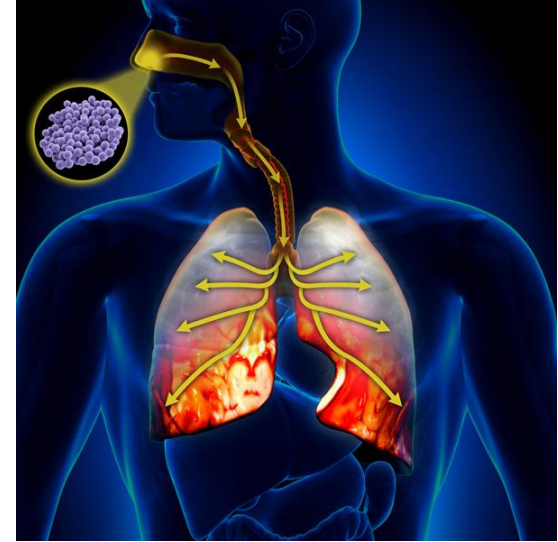


# Bacterial colonization in upper respiratory tract

**Bacterial colonization** of the upper respiratory tract (URT) is generally considered as the first step in the development of invasive bacterial infections, including secondary bacterial infections following respiratory viral infection.

Possible **mechanisms** by which influenza and other viral infections might predispose infected hosts to secondary bacterial pneumonia is by:

- fostering enhanced growth of pathogens; increasing nasal colonization ***S. aureus***
- facilitating the subsequent entry of large bacterial loads into the lower respiratory tract (LRT)

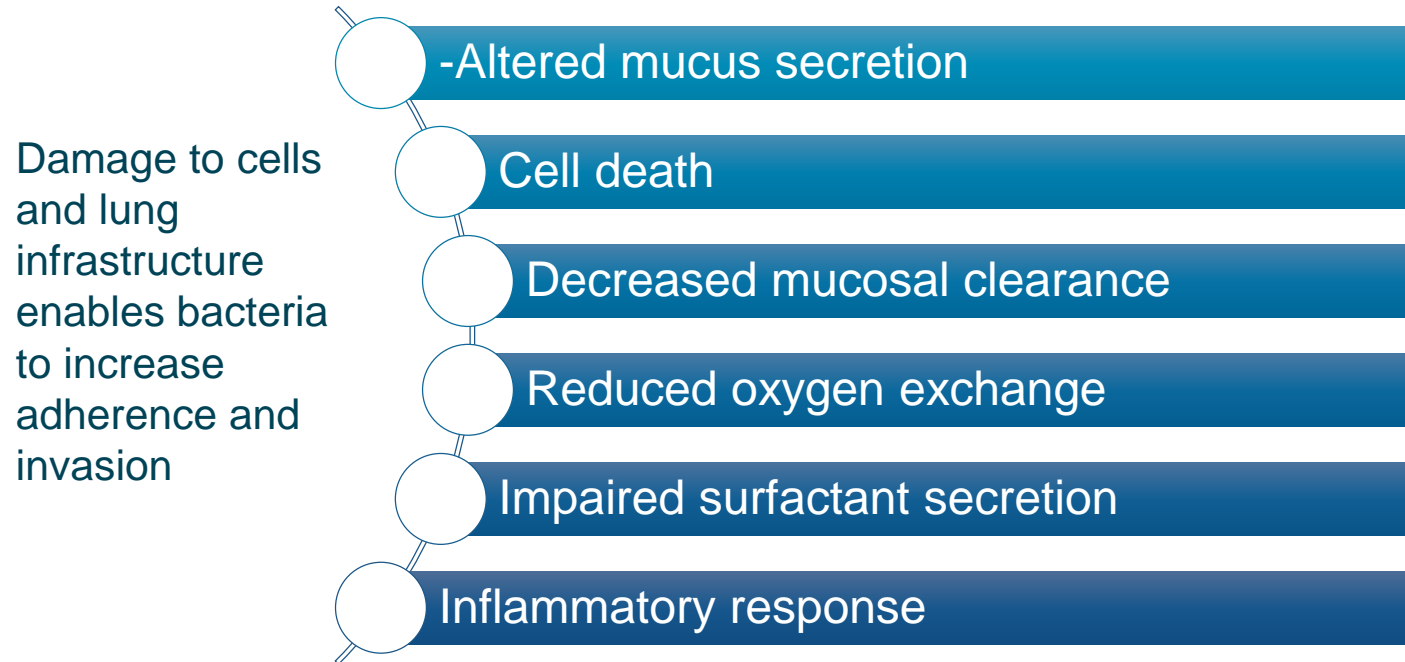


Influenza virus infection is believed to facilitate migration of bacteria from URT to LRT where pathogens can now cause serious disease.



# Mechanisms responsible for SBI with viral respiratory infections

Viral infections promote bacterial colonization of the airway through a variety of mechanisms/detrimental changes:





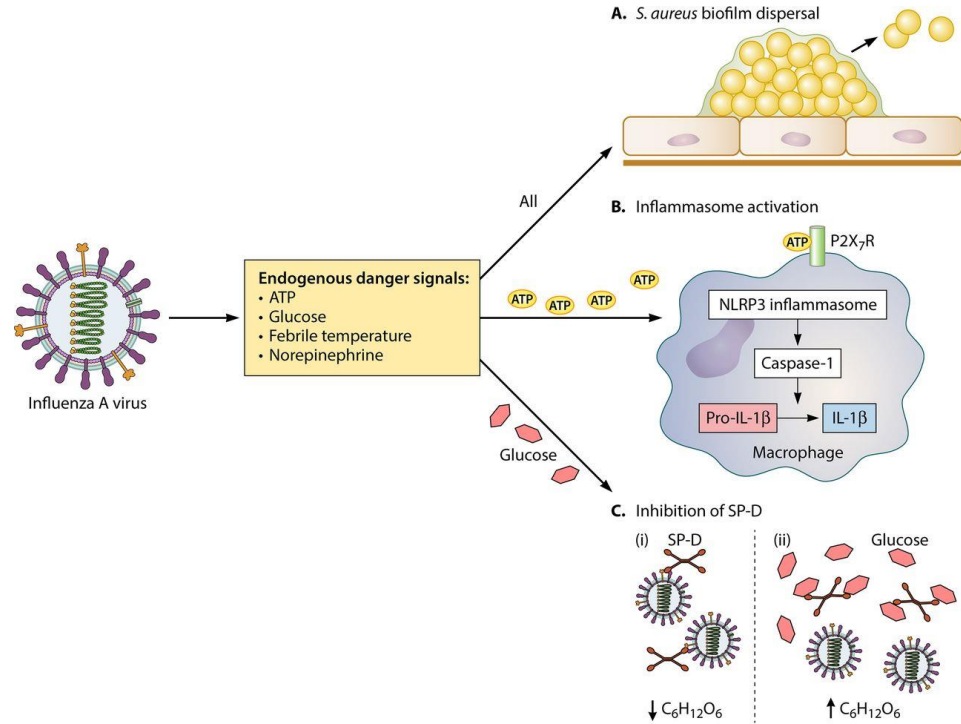
# Nasal colonization: *Staph aureus* and influenza

Nasal carriage of *S. aureus* is a significant risk factor for secondary staphylococcal pneumonia in influenza A virus (IAV)-infected hosts.

Persistent nasal carriers of *S. aureus* are predisposed to invasive disease, including secondary staphylococcal respiratory infection.

*S. aureus* may be aspirated from the nose into the lung, with the potential to cause respiratory infection in a host made susceptible by presence of IAV.

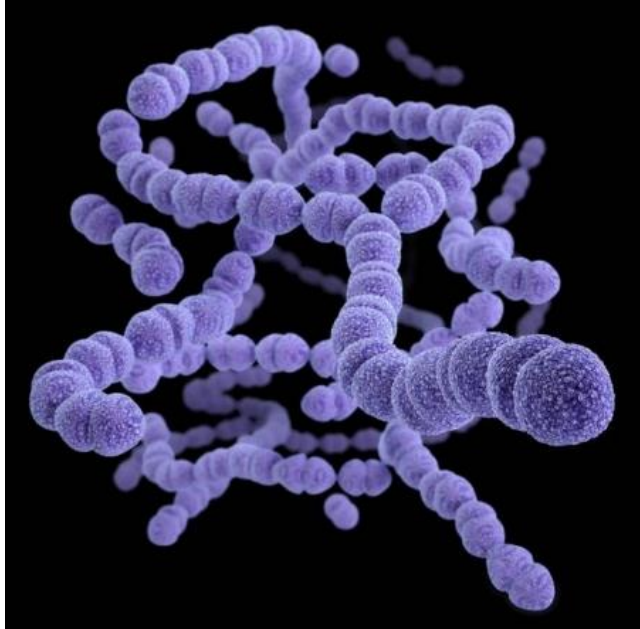
*S. aureus* biofilm dispersal from the nasal environment into the lung is another mechanism of potential inoculation.



**Roles of endogenous danger signals during influenza A viral infection.**



# *Streptococcus pneumoniae*



- Most common bacteria found in SBIs
- High mortality and morbidity during influenza epidemics and pandemics.
- Most common cause of community-acquired pneumonia and invasive disease (sepsis and meningitis) worldwide. (30% lab-confirmed cases of CAP involve bacterial-viral co-infection)
- 4 million cases of infection and 22,000 deaths annually in USA (2011 data\*)
- **Pneumococcal vaccination** has shown to reduce risk of secondary bacterial pneumonia.
- Vaccine implementation has successfully reduced pneumococcal disease, (**45% reduction in incidence** in those with influenza)



# Staphylococcus aureus



- *S. aureus* infection in the intensive care unit (ICU) most commonly manifests as sepsis, VAP, and infection of surgical sites and indwelling medical devices.
- ***S. aureus* nasal colonization has been identified as a major risk factor for the development of nosocomial staphylococcal infection.**
- 20–30% of the healthy population is persistently colonized by *S. aureus* and 60% are intermittently colonized
- Although vaccine development has lowered the mortality of other bacterial infections, all vaccination attempts aimed at preventing *S. aureus* invasive infections have failed in human trials.

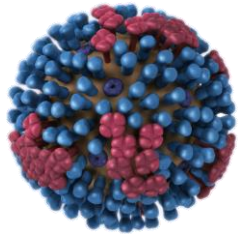




# Staphylococcus aureus and influenza



- Complicates influenza infection, increasingly so in more recent years/pandemics.
- Common cause of pneumonia, specifically necrotizing pneumonia caused by MRSA (30% mortality rate)
- MRSA currently accounts for 20%–40% of hospital-acquired and ventilator-associated pneumonias and 9% of community-acquired pneumonias.
- Increased intensive care admission, mechanical ventilation, and mortality have been described in children and young adults with influenza A and concomitant *S. aureus* infection compared to those with either influenza or *S. aureus* infection alone.





# Exploring risk mitigation strategies

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## Section III



# Mitigating risk

## Most effective risk mitigation strategies:

### #1 Influenza vaccine !!!

Pneumococcal vaccine

Haemophilus influenzae vaccine

Prompt antiviral treatment/prophylaxis



**Eventually:** SARS-CoV-2 vaccine





# Risk mitigation for staphylococcal SBI

## Modifiable risk factors

```
graph TD; A[Modifiable risk factors] --- B[Nasal colonization]; A --- C[Transmission/acquisition HAI];
```

Respiratory viral infection promotes nasal colonization. Preventive measures can be directed at reducing nasal colonization to mitigate the risk of subclinical aspiration of bacteria colonizing the nose.

### Nasal colonization

Nasal carriage of *S. aureus* is a significant risk factor for secondary staphylococcal pneumonia.

*Colonization—subsequent infection*

### Transmission/ acquisition HAI

Secondary infections can be acquired from the patient's environment i.e., hospital-acquired or nosocomial infections.

*Transmission—acquisition—subsequent infection*



# *S. aureus* nasal colonization, a risk factor for infections

*S. aureus* nasal colonization, a risk factor for infections.

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## *S. aureus* nasal colonization, a risk factor for

## Reference

Surgical site infections after orthopedic surgeries

[Kalmeijer et al., 2000](#); [Yano et al., 2000](#); [Weiser and Moucha, 2015](#)

*S. aureus* infections in HIV-infected patients

[Nguyen et al., 1999](#); [Sissolak et al., 2002](#)

Surgical site infections after cardiac surgeries

[Kluytmans et al., 1995](#); [Muñoz et al., 2008](#)

ICU-associated *S. aureus* infections

[Honda et al., 2010](#)

Bacteremia in nonsurgical patients

[Wertheim et al., 2004](#)

Recurrent furunculosis and impetigo

[Durupt et al., 2007](#); [Demos et al., 2012](#)

Catheter-related infections in dialysis patients

[Luzar et al., 1990](#); [Katneni and Hedayati, 2007](#)

Diabetic foot ulcer infections

[Dunyach-Remy et al., 2017](#)

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Nasal carriage of *S. aureus* is a significant risk factor for secondary staphylococcal pneumonia in influenza A virus (IAV)-infected hosts.



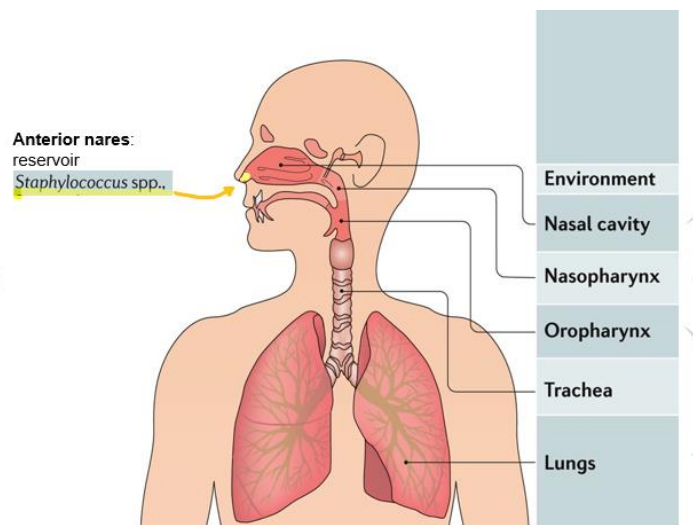


Can nasal decolonization be considered a supplemental risk mitigation strategy to prevent secondary staphylococcal pneumonia?



# Risk mitigation for staphylococcal SBI

## Nasal decolonization as a risk mitigation strategy??



## Pathogenesis of HAP and HCAP in non-intubated patients

- Micro-aspiration of contaminated nasal/oropharyngeal secretions into the lung in persons with compromised defense mechanisms

~ **MAYBE**

## Pathogenesis of VAP

- Aspiration of oropharyngeal or gastric contents that have been colonized by endogenous flora

~ **MAYBE**

## Pathogens from the environment / Acquisition

- hands or attire of healthcare workers
- pathogens attached to respiratory equipment

~ **YES, as source control**



# Risk mitigation for staphylococcal SBI

Should nasal and skin staphylococcal decolonization protocols be deployed in long-term care facility COVID-19 units as source control to mitigate transmission of MSSA and MRSA??



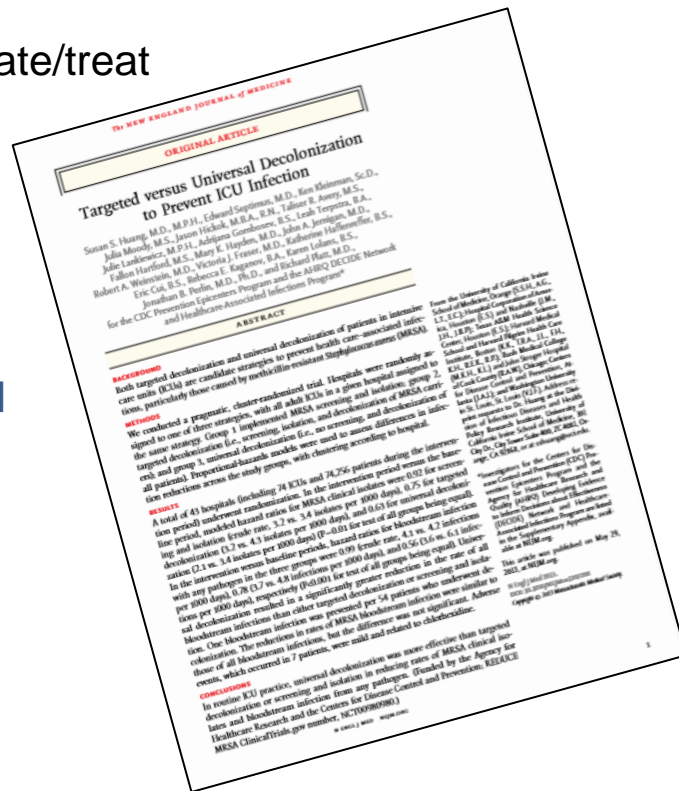
# **Universal nasal decolonization:**

## **A strategy to prevent hospital-onset *S. aureus* infections**

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### Section IV











# Recommendations for Mupirocin Use in Routine Decolonization

**2013 Huang study:** If this practice (universal decolonization) is widely implemented, vigilance for emerging resistance will be required.

**2006 CDC Guidelines for Managing Patients with MDRO:** Routine decolonization is not recommended, however, when decolonization does occur, mupirocin antibiotic susceptibility testing should be performed each time patients undergo mupirocin decolonization to avoid treatment failures.

**2009 CID mupirocin resistance article:** A strategy for monitoring the prevalence of resistance should be developed and implemented whenever mupirocin is to be routinely used.

**2013 ASHP guidelines:** When decolonization therapy (e.g., mupirocin) is used as an adjunctive measure to prevent *S. aureus* SSI, surveillance of susceptibility of *S. aureus* isolated from SSIs to mupirocin is recommended.



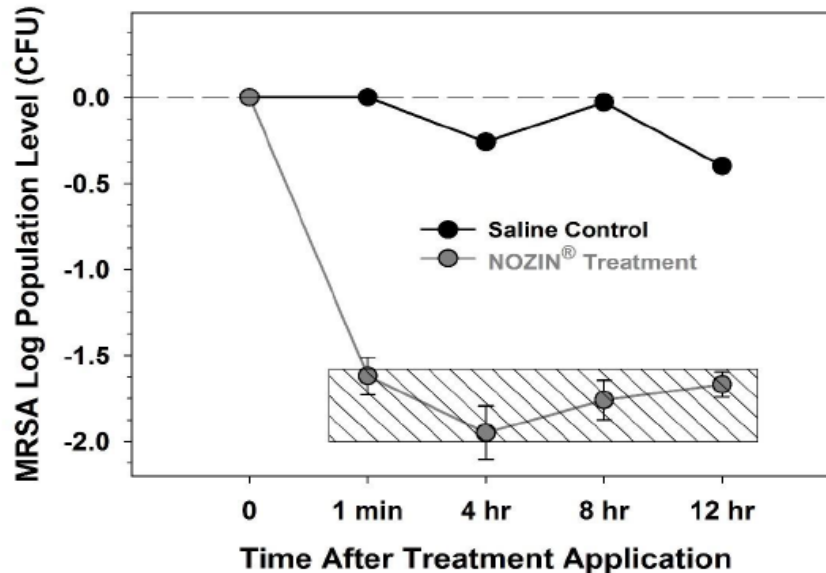
# Nasal Decolonization Agents

Benefits	Alcohol-based antiseptic	Antibiotic prophylactic (mupirocin)	Povidone iodine antiseptic
Effective vs. MRSA/MSSA – 99%	✓	✓	✓
Non-antibiotic--no reported resistance	✓	✗	✓
Effective day 1	✓	✗	✓
Easy to use	✓	✗	✓
Suitable for daily use (2x/day)	✓	✗	✗
Compliance assurance – pre-op	✓	✗	✓



# In vitro Activity of Alcohol-Based Nasal Antiseptic

Evaluation of antimicrobial persistence of alcohol-based nasal antiseptic intended for use to decolonize the human anterior nares. A standard skin explant model was used to evaluate bacteriostatic effect of the product at 20 percent of recommended dose vs. MRSA-ATCC #33592. The baseline surface inoculation and post-treatment levels (cfu/cm<sup>2</sup>) in six replicates for each time point were confirmed by direct count and compared to negative controls.





# In vivo Activity of Alcohol-Based Nasal Antiseptic

## Reduction of Staph Aureus Carriage by alcohol-based nasal antiseptic

Arm/Group Title	Alcohol-Based Nasal Antiseptic	Placebo
▼ Arm/Group Description:	Participants known to exhibit Staph aureus carriage by previous nasal swab screening and randomly assigned received application by nasal swab of alcohol-based nasal antiseptic (Nozin® Nasal Sanitizer®) at 0, 4 and 8 hrs.	Participants known to exhibit Staph aureus carriage by previous nasal swab screening and randomly assigned received application of placebo treatment with phosphate-buffered saline at 0, 4 and 8 hrs.
Overall Number of Participants Analyzed	20	19
Median (Inter-Quartile Range) Unit of Measure: Percent change in colonization		
	-99.8 (-100 to -80.6)	-13.5 (-69.3 to 149.4)



# Study

## Can a nasal and skin decolonization protocol safely replace contact precautions for colonized MRSA patients?

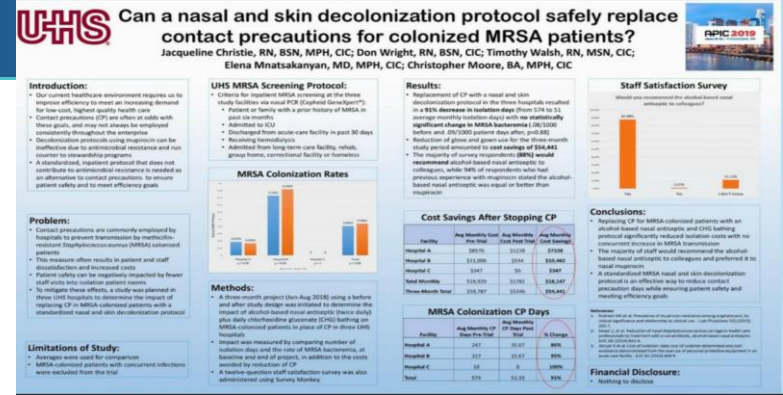
### Introduction:

Our current healthcare environment requires us to improve efficiency to meet an increasing demand for low-cost, highest quality health care

Contact precautions (CP) are often at odds with these goals, and may not always be employed consistently throughout the enterprise

Decolonization protocols using mupirocin can be ineffective due to antimicrobial resistance and run counter to stewardship programs

A standardized, inpatient protocol that does not contribute to antimicrobial resistance in needed as an alternative to contact precautions to ensure patient safety and to meet efficiency goals



### Conclusions:

Replacing CP for MRSA-colonized patients with an alcohol-based nasal antiseptic and CHG bathing protocol significantly reduced isolation costs with no concurrent increase in MRSA transmission

The majority of staff would recommend the alcohol-based nasal antiseptic to colleagues and preferred it to nasal mupirocin

A standardized MRSA nasal and skin decolonization protocol is an effective way to reduce contact precaution days while ensuring patient safety and meeting efficiency goals.



# Study

## Can a nasal and skin decolonization protocol safely replace contact precautions for MRSA-colonized patients?

“...mupirocin is unpleasant to use and must be consecutively dosed twice a day for 5 days to achieve a log kill indicative of nasal decolonization.”

“...patients find povidone iodine unpleasant due to its skin staining properties and odor.”

“...subsequent development of an alcohol-based nasal antiseptic offers enhanced effects when compared to PVI and mupirocin: it does not stain, is clean and well tolerated by patients, has a pleasant citrus odor, is suitable for self application and has no known mechanisms that contribute to microbial resistance...achieves a log kill consistent with decolonization after one application.”

American Journal of Infection Control 48 (2020) 922–924

Contents lists available at ScienceDirect

**American Journal of Infection Control**

journal homepage: [www.ajicjournal.org](http://www.ajicjournal.org)

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AJIC  
American Journal of  
Infection Control

Practice Forum

Can a nasal and skin decolonization protocol safely replace contact precautions for MRSA-colonized patients?

Jacqueline Christie RN, BSN, MPH, CIC \*, Don Wright RN, BSN, MPH, CIC, Jacalyn Liebowitz RN, BSN, MBA, DNP, NEA-BC, FACHE, CPHQ, Paul Stefanacci MD, FACS, MBA

Departments of Quality and Nursing, Universal Health Services, Inc., King of Prussia, PA

Check for updates

### Conclusion:

Replacing contact precautions for high-risk MRSA-colonized patients with an alcohol-based nasal antiseptic and CHG bathing significantly reduced isolation costs with no increase in MRSA bacteremia. In addition, the alcohol-based nasal antiseptic was preferred by staff when compared to nasal mupirocin.

\* Christie J, Wright D, et al *Am J Infect Control*, Vol. 48, Issue 8, p922–924, August 2020



## Effectiveness of an Alcohol-Based Nasal Antiseptic in Reducing MRSA Bacteremia in an Adult Intensive Care Population

### Conclusion:

Incidence of MRSA bacteremia in a 61-bed adult ICU was reduced from 0.24 infections per 1000 for approximately 12,000 patient days ( $p < 0.001$ ) by replacing mupirocin with a staphylococcal decolonization protocol of alcohol-based nasal decolonization in addition to CHG bathing.

### Effectiveness of an Alcohol-Based Nasal Antiseptic in Reducing MRSA Bacteremia in an Adult Intensive Care Population

Lauren Reeves <sup>(a1)</sup>, Lisa Barton <sup>(a1)</sup>, Michelle Nash <sup>(a1)</sup>, Jennifer Williams <sup>(a1)</sup> ... 

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#### Abstract

**Background:** Hospitalized patients are at an increased risk of invasive infection with *Staphylococcus aureus* when colonized with the bacteria on admission. Rates of methicillin-resistant *Staphylococcus aureus* (MRSA) bacteremia are directly correlated with overall patient acuity, placing patients in intensive care areas at greatest risk. Universal decolonization with nasal antibiotic ointments has been shown to reduce the incidence of invasive MRSA in critically ill patients; however, debate remains regarding the long-term efficacy of this strategy and the possibility of developing antimicrobial resistance. An alcohol-based nasal antimicrobial may be an effective alternative. This study evaluated the effectiveness of a twice daily alcohol-based product in reducing the rate of MRSA bacteremia in an academic tertiary-care adult intensive care setting. **Methods:** Our study was an observational design with retrospective and prospective cohorts each consisting of 61 critical care beds. The baseline incidence of MRSA bacteremia was determined from a 7-month period preceding the implementation of the nasal antimicrobial. At implementation, each admission received an electronic order for an alcohol-based nasal antiseptic that was applied twice daily during the intensive care stay. The primary outcome was the incidence of MRSA bacteremia in each group. MRSA bacteremia was defined by the CDC NHSN criteria after review by an infection prevention nurse. The  $\chi^2$  test was used to compare the rates between the 2 groups, and  $P < .005$  was considered significant. **Results:** The study periods contained similar patient days, with 12,475 in the retrospective group and 12,733 in the prospective group. The rate of MRSA bacteremia in the retrospective cohort was 0.2404 compared to 0 in the prospective cohort. This rate change was statistically significant, with  $P < .0001$ . **Conclusions:** The alcohol-based nasal antiseptic was effective in reducing healthcare-onset MRSA bacteremia in this intensive care population. This approach may be a safe and effective alternative to nasal antibiotic ointment that avoids antibiotic resistance risks.

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**Disclosures:** None



# Study

## Does Universal Nasal Decolonization with an Alcohol-Based Nasal Antiseptic Reduce Infection Risk and Cost?

### Results:

Compared with baseline, between April 2018 and March 2019, there was:

- a decrease in MRSA bacteremia from 3/1,000 patient-days to 0/1,000 patient-days
- a reduction in CP from 3.78 to 1.53/1,000 patient-days, a reduction in nasal screens from 3,874 to 605
- a reduction of all-cause (Gram-negative and Gram-positive) SSI after all surgical procedures from 3/4,313 procedures to 0/4,872 procedures.
- After accounting for the cost of the nasal antiseptic, the reduction in gowns, gloves and nasal screening tests resulted in \$104,099.91 costs avoided.

Poster number 567



### Does Universal Nasal Decolonization with an Alcohol-Based Nasal Antiseptic Reduce Infection Risk and Cost?

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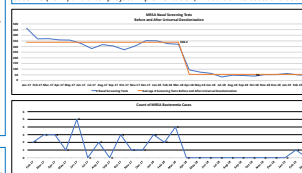


**Background:** Nasal decolonization with mupirocin to reduce infection risk, has been associated with mupirocin-resistant *Staphylococcus aureus* (SA). A community hospital identified two patients colonized with methicillin and mupirocin-resistant SA (MRSA), one scheduled for surgery, one for inpatient IV antibiotic therapy. Instead of mupirocin, an alcohol-based nasal antiseptic was applied to these patients twice daily for 5 days, resulting in a negative MRSA nasal screening test in both patients. Neither patient developed an infection during or after treatment. Building on this success, a plan was made to assess the impact of universal nasal decolonization to replace screening and contact precautions for MRSA colonized patients, and to reduce surgical site infections (SSI).

**Problem:**

- Contact precautions are commonly employed by hospitals to prevent transmission by methicillin-resistant *Staphylococcus aureus* (MRSA) colonized patients
- This measure often results in patient and staff dissatisfaction and increased costs
- Patient safety can be negatively impacted by fewer staff visits into isolation patient rooms
- Patient compliance with pre-operative nasal mupirocin is reportedly less than optimal
- Mupirocin-resistant *Staphylococcus aureus* is a documented ongoing concern
- Nasal antibiotic is not in alignment with antibiotic stewardship principles.

**Methods:** A 26-month before and after study, was initiated in April 2018. The project involved twice daily application of alcohol-based nasal antiseptic for all inpatients, and preoperatively for all surgical patients in addition to existing preoperative chlorhexidine bathing. No other practice change was made during this period. Assessment of impact was planned by comparing incidence of MRSA bacteremia and SSI at baseline (2017) and after project implementation, in addition to costs avoided with reduction of nasal screening and CP.



**Results:** Compared to baseline, between April 2018 and March 2019, there was a decrease in MRSA bacteremia from an average of 2.14 cases per month to an average of 0.08 cases per month, a reduction in CP from 3.78 to 1.53/1,000 patient days, a reduction in total annual nasal screens from 3,874 to 605, and a reduction in the incidence of all cause (Gram negative and Gram positive) SSI after all surgical procedures from 3/4,313 procedures to 0/4,872 procedures. After totaling the costs of the nasal antiseptic, use of gloves and gowns, and nasal screening tests – separate from treatment costs – our adoption of universal decolonization resulted in a savings of \$104,100.

**Conclusion:** House-wide application of alcohol-based nasal antiseptic in place of screening and contact precautions, resulted in a reduced incidence of both MRSA bacteremia and SSI for all types of surgical procedures, in addition to significant costs avoided.

	2017 CP (Current and Travel)	2019 CP (Universal Decolonization)
MRSA Screens	\$ 154,000	\$ 28,000
Gloves, Gowns, Supplies	\$ 96,100	\$ 91,100
Cost of Mupirocin	\$ 4,100	\$ 3,000
Cost of Treatment**	\$ 504,000	\$ 18,000
Total Cost of MRSA Bacteremia	\$ 758,200	\$ 139,100
Cost of Alcohol Based Nasal Antiseptic	\$ -	\$ 22,000
Costs of MRSA Isolation, Decolonization and Antisepsis during CP of the study period	\$ 758,200	\$ 161,100

\*Estimated cost of an ICU-attributable bloodstream infection of \$18,000  
 \*\*\$7,000-\$20,000 is based on several commonly cited sources \*\*

Financial Disclosure: Nothing to disclose

### Conclusion:

- House-wide application of alcohol-based nasal antiseptic in place of screening and contact precautions, resulted in a reduced incidence of both MRSA bacteremia and SSI for all types of surgical procedures, in addition to significant costs avoided.



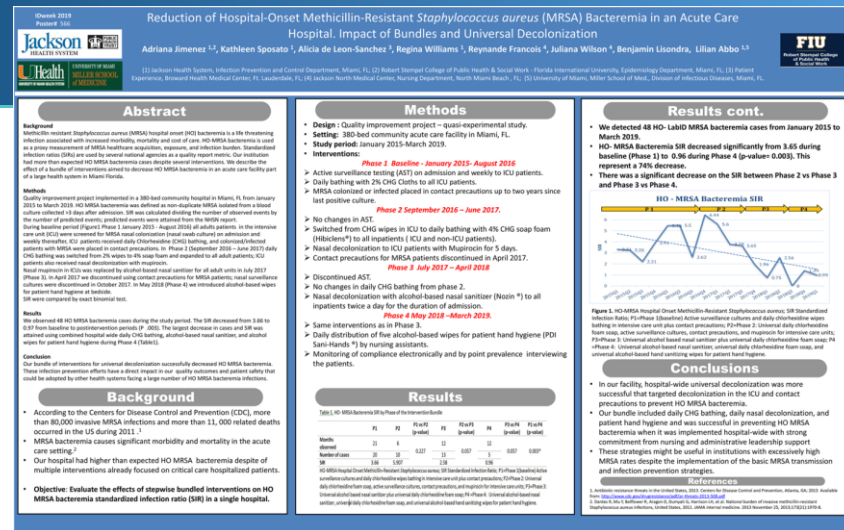
# Reduction of Hospital-Onset Methicillin-Resistant Staphylococcus aureus (MRSA) Bacteremia in an Acute Care Hospital: Impact of Bundles and Universal Decolonization

### Results:

The SIR decreased from 3.66 to 0.97 from baseline to post-intervention periods ( $P = 0.003$ ).










The largest decrease in cases and SIR was attained using combined hospital-wide daily CHG bathing, alcohol-based nasal sanitizer, and alcohol wipes for patient hand hygiene during Phase 4 (Table 1).

Our bundle of interventions for universal decolonization was successful in decreasing HO MRSA bacteremia.










# MRSA bacteremia reduction

 Reduction	Author (location)	ICU	Hospital- wide	CP/Isolation discontinued	Additional	Existing CHG
100%	Arden (Pinellas, FL)				No CHG added	
74%	Jimenez (Jackson, FL)				Bundled nasal decolonization, CHG bathing and patient hand hygiene	 in ICU
100%	Reeves (Methodist, TN)				No CHG added	




# Savings from replacing Contact Precautions

Cost Savings	Author (location)	Screening discontinued	Additional	Period
\$104K	Arden (Pinellas, FL)		Hospital wide.	12-month
\$64K	Deatherage (Marshall, CA)		Hospital wide.	12-month
\$1.4 mill	Whitaker (Tampa, FL)		Hospital wide.	12-month
\$223K	Landis (Frederick, MD)		Hospital wide. Includes savings from CP replacement, screening and SSI cost avoidance.	Annual
\$200K	Steigmeier (Wesley Chapel, FL)		High risk patient population.	12-month
\$430K	Christie* (UHS, PA)		MRSA colonized patients. Combined savings in 7 hospitals.	10-month



# Reduction in use of Contact Precautions for MRSA colonization

 Reduction	Author (location)
<b>88%</b>	Christie* (UHS, PA)
<b>60%</b>	Arden (Pinellas, FL)
<b>38%</b>	Steigmeier (Westley Chapel, FL)
<b>42%</b>	Whitaker (Tampa, FL)



## Key points

- The anterior nares are the primary sites of *S. aureus* carriage, which is a precursor to and the primary risk factor for development of *S. aureus* infections.
- Nasal and skin decolonization is a recognized strategy to reduce healthcare-associated *S. aureus* infections.
- Nasal colonization with *S. aureus* increases the risk of secondary staphylococcal infections among patients with influenza infection (and possibly COVID-19 patients?)
- Nasal and skin decolonization may have the potential to also reduce the risk of secondary staphylococcal pneumonia (currently no clinical evidence).



# Recommendations for reducing hospital-onset *Staphylococcus aureus* infections

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Centers for Disease Control and Prevention  
Society for Healthcare Epidemiology of America



# Centers for Disease Control and Prevention (2019)

## Strategies to Prevent Hospital-onset(HO) *Staphylococcus aureus* Bloodstream Infections in Acute Care Facilities

### 1. Prevent Healthcare-Associated Infections

Central line-associated bloodstream infections (CLABSI)

Surgical site infections (SSI)

BSI among hemodialysis patients

Ventilator-associated pneumonia (VAP)

\* <https://www.cdc.gov/hai/prevent/staph-prevention-strategies.html>  
<https://www.cdc.gov/infectioncontrol/pdf/guidelines/bsi-guidelines-H.pdf>  
<https://jamanetwork.com/journals/jamasurgery/fullarticle/2623725>  
[https://www.cdc.gov/dialysis/PDFs/Dialysis-Core-Interventions-5\\_10\\_13.pdf](https://www.cdc.gov/dialysis/PDFs/Dialysis-Core-Interventions-5_10_13.pdf)  
<https://www.cambridge.org/core/journals/infection-control-and-hospital-epidemiology/article/strategies-to-prevent-ventilator-associated-pneumonia-in-acute-care-hospitals-2014-update/2D8A9D3BFD8BC8A68E04906B5C2CEF66>





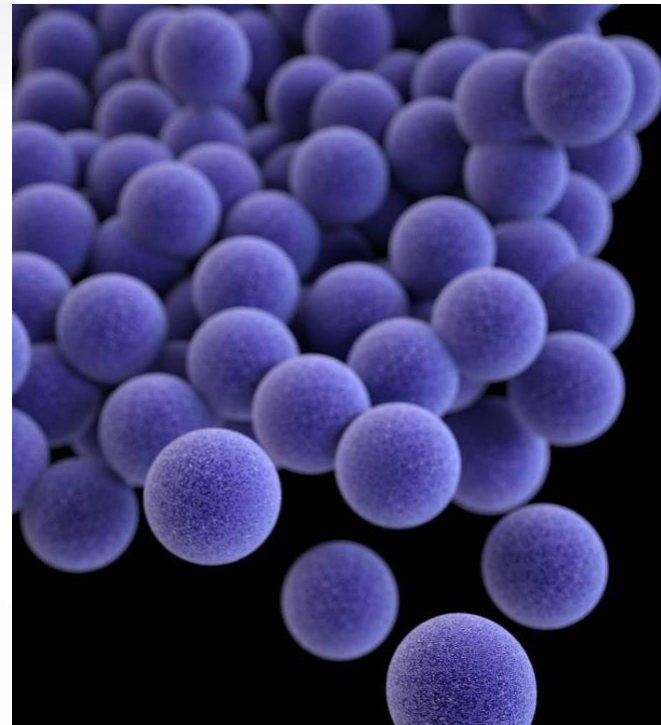
## Strategies to Prevent Hospital-onset *Staphylococcus aureus* Bloodstream Infections in Acute Care Facilities

### 2. Practice Source Control

**ICU patients:** Decolonize all patients with intranasal staphylococcal antibiotic/antiseptic plus topical CHG (core strategy).

**Non-ICU patients:** Decolonize patients with CVC or midline catheter with intranasal staphylococcal antibiotic/antiseptic plus topical CHG (supplemental strategy).

**Surgical patients:** For all patients undergoing high risk surgeries (e.g. cardiothoracic, orthopedic, and neurosurgery), unless known to be *S. aureus* negative, use an intranasal anti-staphylococcal antibiotic/antiseptic and CHG wash or wipes prior to surgery (core strategy).





## Strategies to Prevent Hospital-onset *Staphylococcus aureus* Bloodstream Infections in Acute Care Facilities



### 3. Prevent Transmission of MRSA

**Place MRSA colonized or infected patients** in private rooms and on contact precautions.

**Use dedicated patient-care equipment** (e.g. blood pressure cuffs, stethoscopes), and single use disposable items (e.g. single patient digital thermometer) whenever possible. If common use of equipment for multiple patients is unavoidable, clean and disinfect such equipment before use on another patient.

**Provide regular competency-based training** on use of PPE and monitor adherence.

**Place patients with excessive wound drainage on contact precautions** and in a private room regardless of MDRO status.



## Strategies to Prevent Methicillin-Resistant *Staphylococcus aureus* Transmission and Infection in Acute Care Hospitals: 2014 Update



**ICU:** MRSA decolonization can be targeted to MRSA-colonized persons or applied universally to populations deemed to be at high risk for infection.

(Level 1 Evidence: Provide universal decolonization to ICU patients when MRSA not effectively controlled).





# Benefits of universal decolonization programs

- Mitigates risk of infection to the colonized patient.
- Mitigates risk of acquisition in the non-colonized patient.
- More effective than targeted (screening) decolonization in reducing healthcare-associated infections caused by staphylococcal organisms.
- Decolonizes patients with MSSA in addition to those with MRSA.
- Provides source control, reducing contamination of HCP hands, the patient environment and equipment and thus decreasing risk of transmission to other patients.
- Costs less to decolonize an entire patient population at risk than to screen and place in contact precautions.
- Eliminates the need for contact precautions, thus improving both patient and staff satisfaction.



# Benefits of expanding to house-wide decolonization programs

- Eliminates the need to manage MSSA and MRSA colonized patients with contact precautions in all units.
- Inclusion of all patients facility-wide simplifies and improves compliance with the decolonization protocol and provides consistency across all units.
- Delivers cost savings when replacing "screen and isolate" protocols with universal decolonization.
- Improves patient flow and throughput.
- Increases patient and staff satisfaction throughout the facility.
- May be beneficial to general ward patients with respiratory viral illnesses as a strategy to prevent secondary bacterial infections.



# Resources for staphylococcal decolonization

Centers for Disease Control and Prevention. Strategies to prevent hospital-onset(HO) *Staphylococcus aureus* bloodstream infections in acute care facilities, 2019. <https://www.cdc.gov/hai/prevent/staph-prevention-strategies.html>

Society for Healthcare Epidemiology of America. Strategies to prevent methicillin-resistant *Staphylococcus aureus* transmission and infection in acute care hospitals: 2014 Update. 35(s2) Sept 2014:S108-S132. DOI: <https://doi.org/10.1017/S0899823X00193882>

Health Research and Educational Trust. Preventing surgical site infections: 2018 Update. <http://www.hret-hiin.org/Resources/ssi/18/surgical-site-infections-change-package.pdf>

World Health Organization. Global guidelines on the prevention of surgical site infection, 2016. <https://www.who.int/gpsc/ssi-prevention-guidelines/en/>

Bode et al. Preventing surgical-site infections in nasal carriers of *Staphylococcus aureus*. *N Engl J Med* 2010; 362:9-17.

Huang SS et al. Targeted vs. universal decolonization to prevent ICU infection. *N Engl J Med* 2013; 368 (24) 2255-65.

Huang SS et al. Chlorhexidine vs. routine bathing to prevent multidrug-resistant organisms and all-cause bloodstream infections in general medical and surgical units (ABATE infection trial): a cluster-randomized trial. *The Lancet* 2019; 393: 1205-1215.



## In summary...

...universal staphylococcal decolonization using alcohol-based nasal antiseptics is an evidence-based, cost-effective strategy, that, when used in addition to current infection prevention practices (HAI prevention bundles, hand hygiene, environmental cleaning and disinfection), mitigates risk of hospital-acquired MSSA and MRSA infections.

### Who benefits?

- ✓ ICU patients
- ✓ Surgical patients
- ✓ General ward patients with central/midline catheters
- ✓ Patients with respiratory viral infections who are at risk of acquiring SBI
- ✓ Healthcare personnel
- ✓ Chief financial officer
- ✓ Materials Management staff





**Questions?**

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