

# Inadequate Double-Pseudomonal Coverage with Ciprofloxacin versus Aminoglycosides

Beaumont



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## Introduction / Rationale

- Pseudomonas aeruginosa* is an important gram-negative bacilli often implicated in serious hospital and healthcare-associated infections, is often antibiotic resistant, and is associated with a high mortality rate
- In the absence of acquired additional antibiotic resistance, there are limited antimicrobial agents with reliable activity against pseudomonas isolates
  - Antipseudomonal penicillins
  - Antipseudomonal cephalosporins
  - Monobactam
  - Fluoroquinolones
  - Carbapenems
  - Aminoglycosides
- Although controversial, combination antimicrobial therapy with two different classes is likely indicated in certain high-risk patients and severe infections to increase the likelihood of effective empiric antibiotic therapy
- Combinations utilized in our intensive care units for empiric double coverage of pseudomonas includes an antipseudomonal beta-lactam plus either ciprofloxacin or an aminoglycoside
- ICU antibiograms suggest ciprofloxacin to have a higher rate of resistance compared to aminoglycosides and beta-lactams at our institution
- It is unknown at our institution if ciprofloxacin or the aminoglycosides add additional coverage to patients with pseudomonas isolates resistant beta-lactams

## Objectives

- Determine the rate of ciprofloxacin resistance to pseudomonas isolates that are resistant to piperacillin-tazobactam and/or cefepime
- Determine the rate of aminoglycoside (gentamicin, tobramycin) resistance to pseudomonas isolates that are resistant to piperacillin-tazobactam and/or cefepime
- Compare the rates of resistance between ciprofloxacin and the aminoglycosides to pseudomonas isolates that are resistant to piperacillin-tazobactam and/or cefepime

## Methods

- Study Design
  - Single center, retrospective and prospective, observational chart review
  - Approved by the Institutional Review Board at Beaumont Health
  - Data will be collected from the electronic medical record

Figure 1: Eligibility Criteria

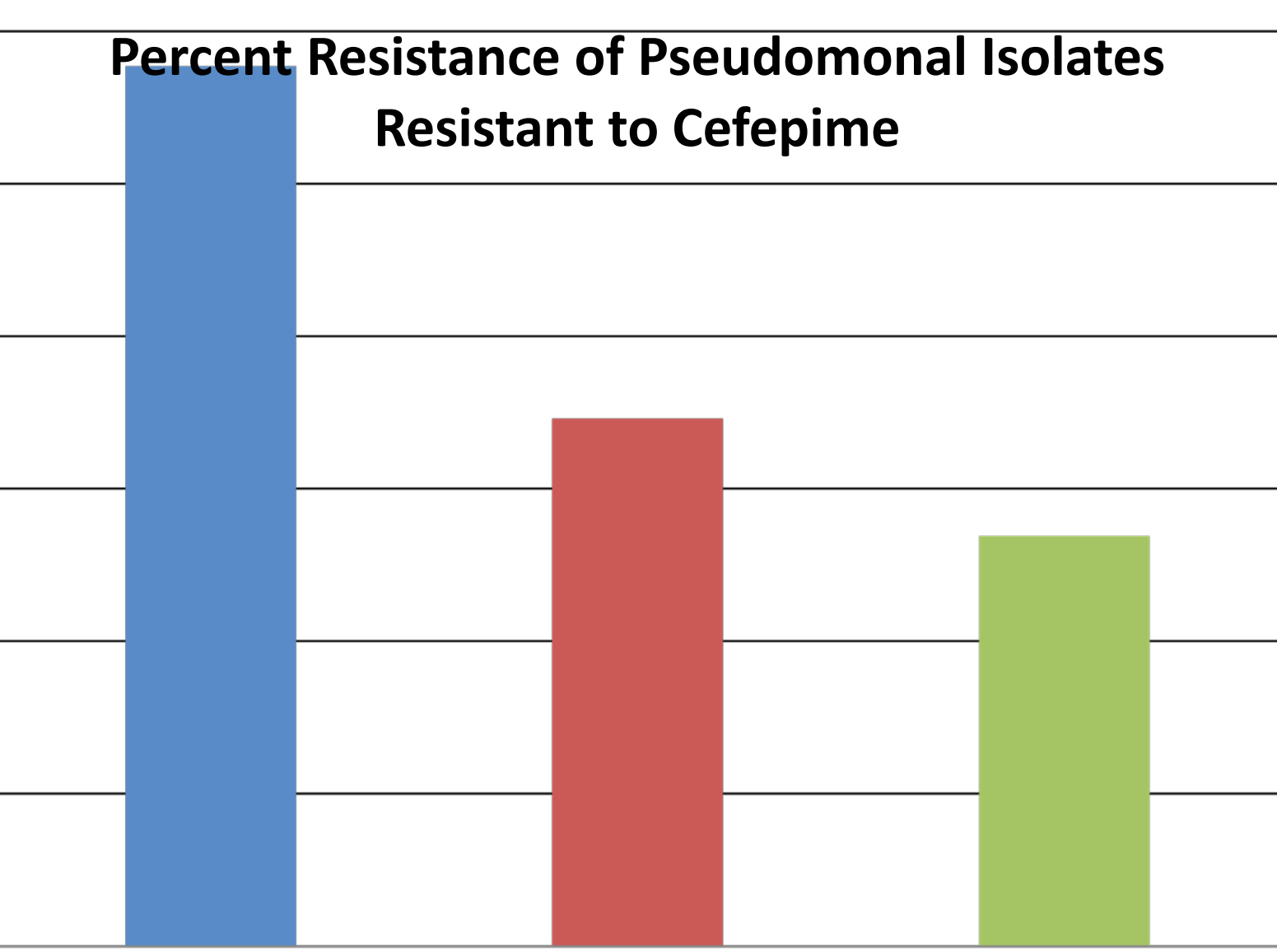
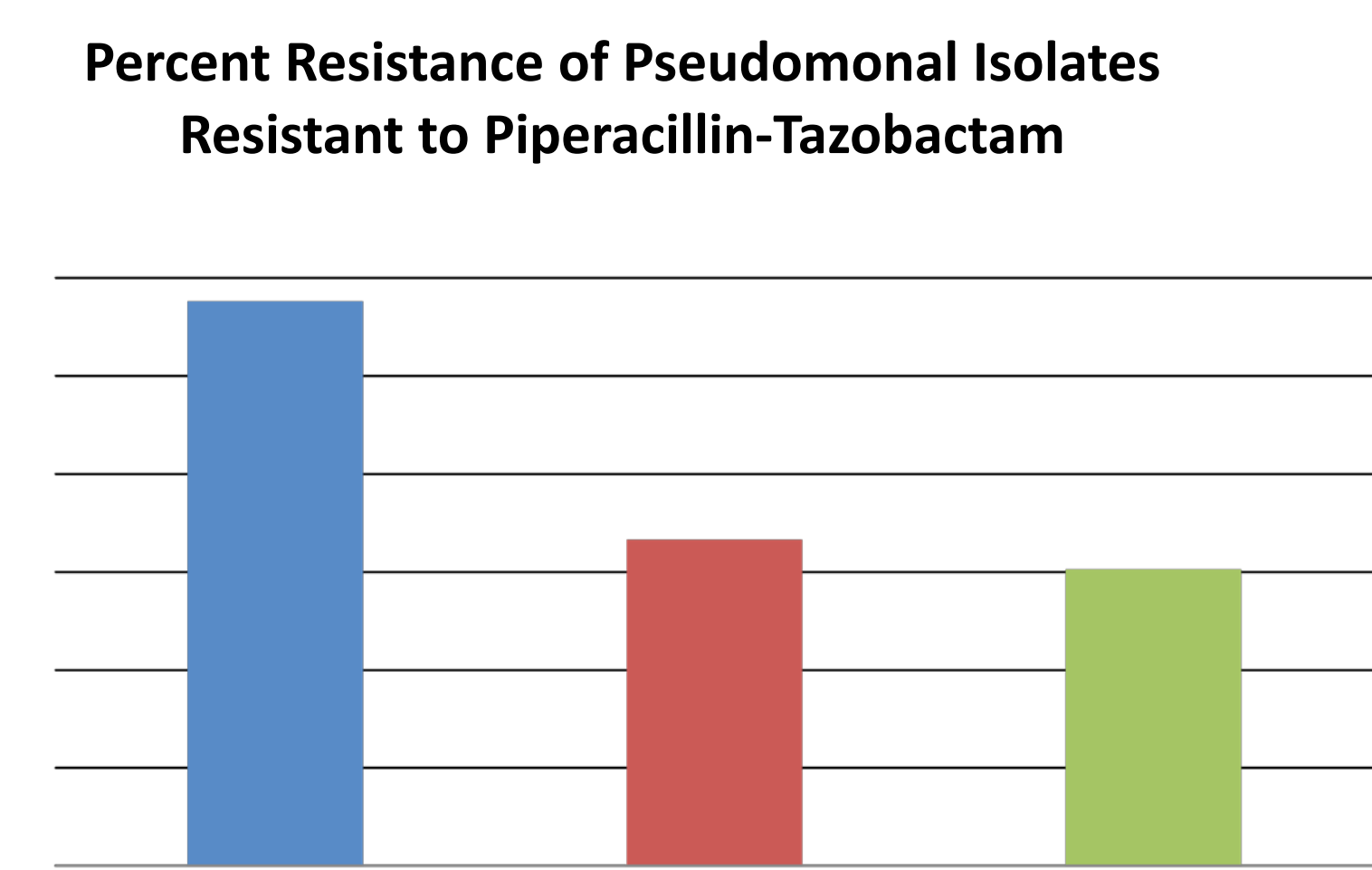
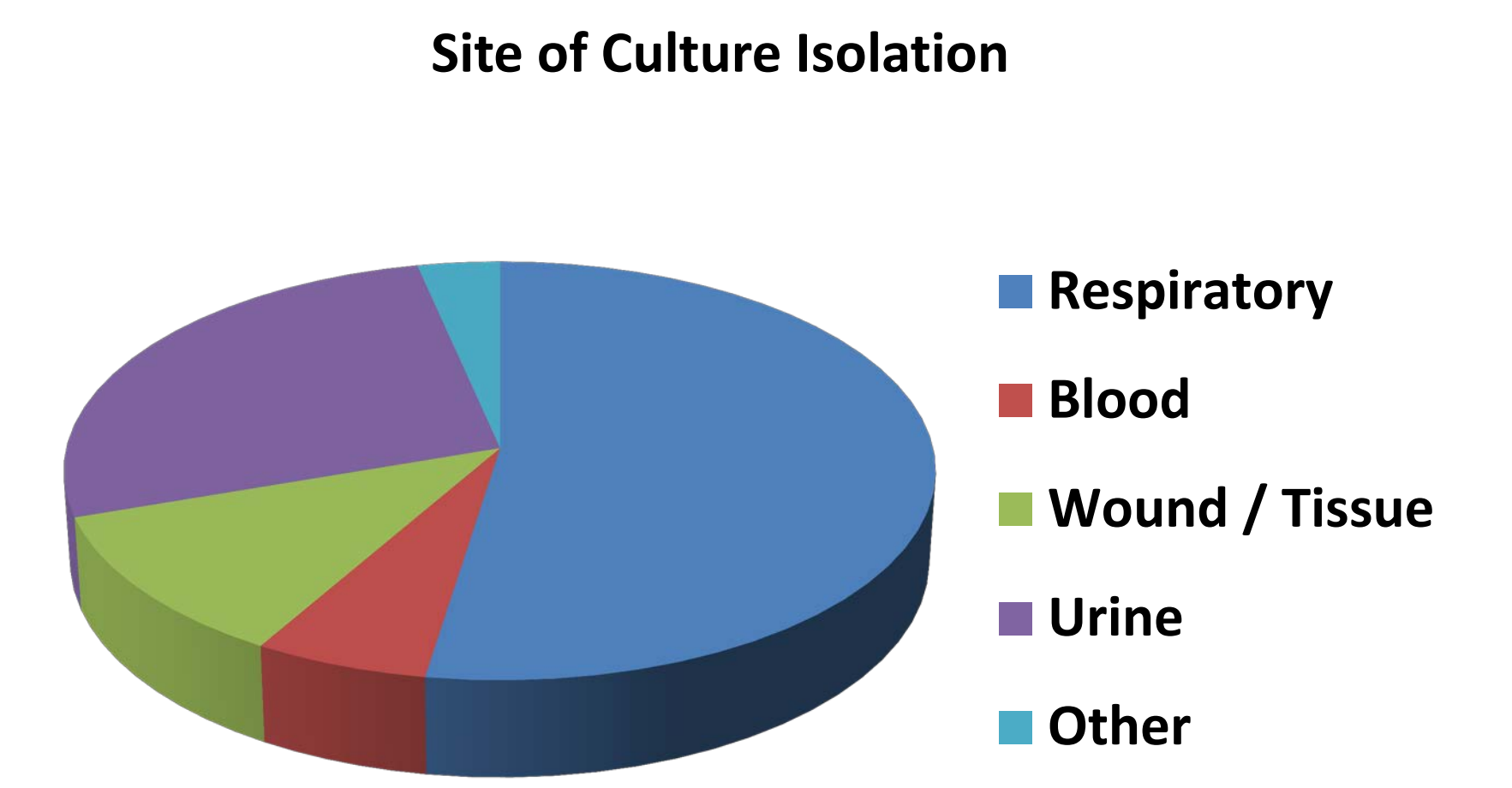
Inclusion Criteria	Exclusion Criteria
<ul style="list-style-type: none"><li>Adult patients (≥ 18 years)</li><li>Admission to any intensive care unit at Beaumont Hospital – Royal Oak from 1/1/2011 through 12/31/2012</li><li>Positive Pseudomonas culture from any source</li></ul>	<ul style="list-style-type: none"><li>Age &lt; 18 years</li><li>Susceptibility data not available</li></ul>

## Methods (continued)

<b>Table 1: Data Collection</b>	
<ul style="list-style-type: none"><li>Demographic information</li><li>ICU Location</li><li>Date of culture</li><li>Source of culture:<ul style="list-style-type: none"><li>Respiratory</li><li>Blood</li><li>Wound / Tissue</li><li>Urine</li><li>Other</li></ul></li></ul>	<ul style="list-style-type: none"><li>Antibiotic course of therapy at time of pseudomonas culture</li><li>ICU and hospital mortality</li><li>Antibiotic sensitivities:<ul style="list-style-type: none"><li>Cefepime</li><li>Ciprofloxacin</li><li>Gentamicin</li><li>Tobramycin</li><li>Piperacillin-tazobactam</li></ul></li></ul>

## Results

- Total of 324 pseudomonas cultures were isolated
- Total of 323 sensitivities were able to be evaluated
- Other isolation sites:
  - Cerebrospinal fluid
  - Bile fluid
  - Ascites fluid
  - Peritoneal fluid
  - Pelvic fluid
  - Lung autopsy



## Limitations

- Retrospective chart review
- Correlation between colonization and infection was absent
- Sensitivity data limited to piperacillin-tazobactam, cefepime, gentamicin and tobramycin
- Pseudomonal isolate speciation was not recorded

## Discussion / Conclusions

- The rate of ciprofloxacin resistance to pseudomonas isolates resistant to piperacillin-tazobactam or cefepime approached 58 percent
- The rate of aminoglycoside (gentamicin, tobramycin) resistance to pseudomonas isolates resistant to piperacillin-tazobactam or cefepime was between 27 and 34 percent
  - Gentamicin: 33-34%
  - Tobramycin: 27-30%
- There is roughly a 2-fold increase in ciprofloxacin resistance to pseudomonas isolates as compared to aminoglycoside resistance.
- When utilizing double gram(-) coverage for infections in the critically ill patient, aminoglycosides should be favored over ciprofloxacin in the absence of contraindications.

## References

- Tuon FF, Gortz LW, Rocha JL. Risk factors for pan-resistant Pseudomonas aeruginosa bacteremia and the adequacy of antibiotic therapy. Braz J Infect Dis 2012; 16:351.
- Park SY, Park HJ, Moon SM, et al. Impact of adequate empirical combination therapy on mortality from bacteremic Pseudomonas aeruginosa pneumonia. BMC Infect Dis 2012; 12:308.
- Bowers DR, Liew YX, Lye DC, et al. Outcomes of appropriate empiric combination versus monotherapy for Pseudomonas aeruginosa bacteremia. Antimicrob Agents Chemother 2013; 57:1270.
- Chamot E, Boffi El Amari E, Rohner P, Van Delden C. Effectiveness of combination antimicrobial therapy for Pseudomonas aeruginosa bacteremia. Antimicrob Agents Chemother 2003; 47:2756.
- Vardakas KZ, Tansarli GS, Bliziotis IA, Falagas ME. β-Lactam plus aminoglycoside or fluoroquinolone combination versus β-lactam monotherapy for Pseudomonas aeruginosa infections: a meta-analysis. Int J Antimicrob Agents 2013; 41:301.
- Paul M, Leibovici L. Combination antibiotic therapy for Pseudomonas aeruginosa bacteraemia. Lancet Infect Dis 2005; 5:192.
- Peña C, Suarez C, Ocampo-Sosa A, et al. Effect of adequate single-drug vs combination antimicrobial therapy on mortality in Pseudomonas aeruginosa bloodstream infections: a post Hoc analysis of a prospective cohort. Clin Infect Dis 2013; 57:208.

## Disclosures

Authors of this presentation have the following to disclose concerning possible financial or personal relationships with commercial entities that may have a direct or indirect interest in the subject matter of this presentation:

- All authors have nothing to disclose