

receiving a lipoglycopeptide without insurance, the variable direct cost avoidance was \$4,560 per case, or \$1,060 per day.

**Conclusion.** The use of lipoglycopeptides offers patient convenience and financial benefits, warranting its consideration for use in the ED at tertiary academic medical centers.

**Disclosures.** All authors: No reported disclosures.

### 1879. A Point Prevalence Study of Antibiotic Utilization in 61 Geographically Diverse Acute Care Hospitals (2017)

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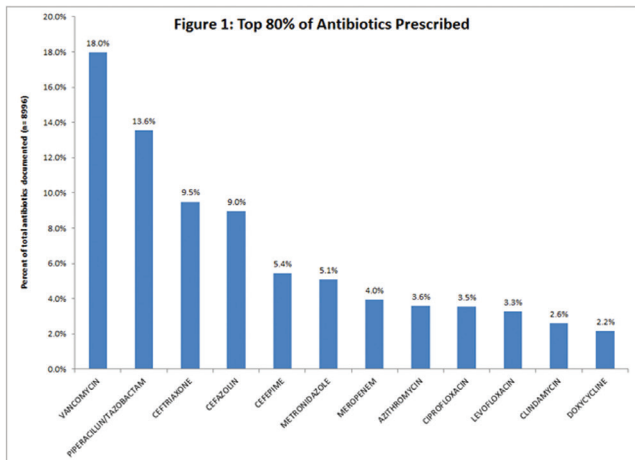
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**Background.** Antibiotic utilization for geographically diverse areas can be difficult to obtain. The purpose of this study was to characterize patterns of US antibiotic use over a defined period to provide comparative data for benchmarking and to assist with identifying antibiotic stewardship opportunities.

**Methods.** Data were obtained as part of a larger study evaluating antibiotic time out practices. Participating institutions submitted de-identified patient-level antibiotic use data from a single day (between October 16, 2017 and November 17, 2017). Indication, expected duration, and antibiotic stop dates were documented. Antibiotics were classified by American Hospital Formulary Service (AHFS) therapeutic category and evaluated to identify duplicate anti-anaerobic, anti-MRSA, and AHFS classes. Hospital teaching status and US Census region were recorded.

**Results.** A total of 6,184 courses of therapy (8,996 individual antibiotics) were evaluated from 61 hospitals. Sixty-four percent of therapy courses submitted were from academic medical centers. Distribution by census region was Midwest (44.7%), Northeast (15.11%), South (23.2%), and West (16.9%). Over half (53.7%) of therapy was empiric and 33.4% was directed. Sixty-six percent of courses did not include a stop date within the electronic medical record. Twelve drugs comprised 80% of total antibiotic use. Percentage of antipseudomonal use was similar across regions, but anti-MRSA therapy was higher in the South and Midwest. Duplicate  $\beta$ -lactam therapy and duplicate anti-anaerobe therapy were identified in 1.5% of total courses (each). Duplicate anti-MRSA therapy occurred in 0.29% of therapy courses. Three percent of patients developed a *Clostridium difficile* infection during their hospitalization.

**Conclusion.** Vancomycin and piperacillin-tazobactam were the most common antibiotics used which is consistent with other analyses, but anti-anaerobe use as a percentage of overall use was higher than expected. Duplicate anti-anaerobe and  $\beta$ -lactam therapy is less frequent, but still represents an opportunity for stewardship. Antipseudomonal and anti-MRSA agents represent two key categories for stewardship given the high percentage of use. The addition of a stop date to the antibiotic order presents an opportunity to improve overall utilization.



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### 1880. Does the Label Matter: Initial Use of Newly Approved Antimicrobial Agents in Community Hospitals Without Robust Antimicrobial Stewardship Programs

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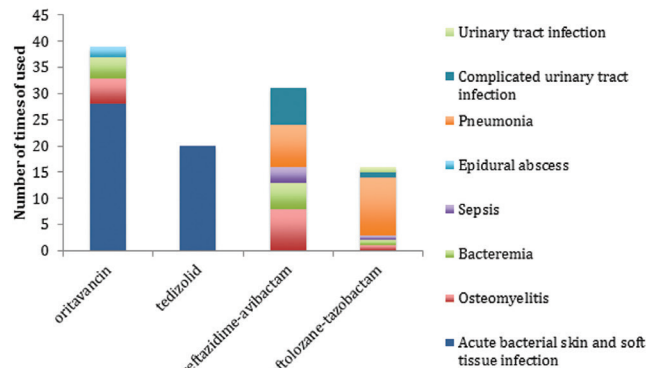
**Background.** Antimicrobial agents are often used for indications not approved by the Food and Drug Administration (FDA). While this can be appropriate at times, they may be used in settings with no published experience. Judicious use of new agents is critical for conserving their utility. This study characterized initial use after FDA approval of select antimicrobial agents in community hospitals without robust antimicrobial stewardship programs (ASP).

**Methods.** Initial use of systemic antimicrobials approved by the FDA since 2014 was retrospectively reviewed at 6 community hospitals (50–350 beds). Up to 10 charts of first use were reviewed per drug at each hospital. Time from FDA approval to first administration was measured for the following agents: ceftazidime-avibactam, ceftolozane-tazobactam, dalbavancin, isavuconazonium, oritavancin, and peramivir. Clinical indications, prescribing service, and empiric uses were recorded.

**Results.** Mean time from FDA approval to first administration ranged from 12 (tedizolid) to 26 months (dalbavancin). Of frequently used agents (Figure 1), adherence to initial FDA indications ranged from 7% (ceftolozane-tazobactam for complicated urinary tract infection) to 100% (tedizolid for acute bacterial skin and soft-tissue infection). Pneumonia (35%) and osteomyelitis (35%) were the most common off-label indications for ceftazidime-avibactam. Pneumonia was the most common off-label indication for ceftolozane-tazobactam (78%). The most common off-label indications for oritavancin were osteomyelitis (14%) and bacteremia (11%). Infectious Diseases was the main prescribing service for all agents (range 74–95%). Use of Gram-positive agents was mostly empiric whereas Gram-negative agents were targeted against specific pathogens.

**Conclusion.** Newly approved antimicrobial agents were used at these six community hospitals within 1–2 years after FDA approval. Agents with primarily Gram-positive activity were more often used for FDA approved indications. Given frequent use of novel Gram-negative agents for pneumonia, there is need for early trials to determine their role for this indication. In the meantime, ASPs should consider off-label indications such as pneumonia when developing local criteria for use.

**Figure 1. Clinical Indications of the Top Antimicrobials**



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### 1881. Empiric Pseudomonal Monotherapy vs. Combination Therapy for Community-Onset Pneumonia in Older Adults

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**Background.** Patients with pseudomonal pneumonia have a poor prognosis; therefore, IDSA guidelines recommend empiric antipseudomonal combination therapy when *Pseudomonas* is suspected, at least until treatment can be adjusted based on susceptibilities. However, combination antipseudomonal therapy is controversial. This study compared all-cause 30-day mortality in older patients who received antipseudomonal monotherapy (PMT) or antipseudomonal combination therapy (PCT) for the treatment of community-onset pneumonia.

**Methods.** This population-based cohort study used data from over 150 Veteran Health Administration hospitals. Patients were classified as low, medium, or high risk of drug-resistant pathogens according to a published rule. Patients were assigned to PCT or PMT groups based on antibiotics received in the first 48 hours of hospital admission. Separate multivariable logistic regression models were constructed to determine whether the choice of PCT or PMT was associated with 30-day mortality, after accounting for divergent baseline characteristics. Adjusted odds ratios (aORs) and 95% confidence intervals (95% CI) were calculated for the overall, low, medium, and high-risk groups.

**Results.** Of the 31,027 patients who met study criteria, 23% received PCT and 77% received PMT. Patients belonged to low (59%), medium (24%), and high (18%) risk groups. 30-day mortality was 18% overall, and increased among the groups: low (13%), medium (21%), and high (36%). Patient age (median of 78 years), race (>80% white), and sex (>98% male) were similar for patients receiving PCT and PMT. The unadjusted mortality difference between PCT and PMT was most pronounced in the

low-risk group (18% vs. 8%, 10% absolute risk difference), followed by the medium (24% vs. 18%, 6% difference) and high (39% vs. 33%, 6% difference) risk groups. PCT was associated with higher 30-day mortality than PMT overall (aOR, 1.54; 95% CI, 1.43–1.66), and in all three groups: low (aOR, 1.69; 95% CI, 1.50–1.89), medium (aOR, 1.30; 95% CI, 1.14–1.48), and high (aOR, 1.21; 95% CI, 1.04–1.40).

**Conclusion.** Older adults who received empiric combination antipseudomonal therapy for community-onset pneumonia fared worse than those who received monotherapy. Empiric combination antipseudomonal therapy should not be routinely offered to all patients suspected of having pseudomonal pneumonia.

**Disclosures.** All authors: No reported disclosures.

### 1882. Resistance (R) Trends in Gram-Negative Bacilli (GNB) to Fluoroquinolones (FQ), [Ciprofloxacin (CIP), Levofloxacin (LEV), Moxifloxacin (MOX)], Trimethoprim-Sulfamethoxazole (TMP/SMX) and Nitrofurantoin (NFT) over a 7-Year Period: Pre- and Post-Implementation of FQ Restriction at a Tertiary Care Veterans Affairs Medical Center (VAMC)

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**Background.** Unrestricted use of FQ has been associated with emergence of drug resistance in GNB. Active surveillance measures are essential components of antibiotic stewardship practice. The aim of this study was to evaluate antibiotic utilization, assess the impact of FQ restriction on resistance to FQ, TMP/SMX, and NFT, and identify quality improvement measures to prevent emergence of resistance in GNB.

**Methods.** A retrospective records review and comparison of antimicrobial susceptibility data for GNB isolated at the Detroit VAMC over a 7-year period, pre- and post-FQ restriction, implemented in 2013. Susceptibility testing was performed by reference broth micro-dilution methods in a central laboratory. Antibiotic usage data were obtained from pharmacy computer records from 2011 to 2017. Antibiotic use in inpatients was expressed as days of therapy/1,000 patient-days (DOT/PD) and as number of prescriptions filled for outpatients. Data were analyzed using Pearson correlation coefficient score.

**Results.** In 2016–2017, the most common GNB isolated in our institution were *E. coli* ( $n = 303$ ), *Klebsiella pneumoniae* ( $n = 100$ ), and *P. aeruginosa* ( $n = 70$ ). Inpatient: During 2011–2012, DOT/1,000 PD for CIP, LEV, and MOX were 34, 10, and 5 respectively, that dropped to 14, 5, and 3 during 2014–2017, post-FQ restriction initiated in 2013. Outpatient: During 2013–2017, outpatient CIP and MOX prescriptions decreased from 1936 to 781 and from 478 to 86, respectively; however, prescriptions for LEV, TMP/SMX, and NFT increased from 33 to 128, 680 to 1,074, and 95 to 322, respectively. Overall: resistance to CIP, LEV, and MOX had increased by 8% in *E. coli* (14–22%) and by 7% in *P. aeruginosa* (10–17%) during 2015–2017; FQ-R in *K. pneumoniae* and NFT-R in *E. coli* stayed low at 7% and 2%, respectively. Also, isolates of TMP/SMX-resistant *E. coli* and NFT-resistant *K. pneumoniae* increased from 20% to 27% and 40% to 51%, respectively.

**Conclusion.** Use of FQ among outpatients was still high (781 scripts in 2017) despite a 4-year restriction, resulting in high FQ-R in *E. coli* and *P. aeruginosa*; a concomitant increase in TMP/SMX and NFT resistance was noted, attributed to a compensatory increased use of these agents during the study period ( $P < 0.05$ ). Reversal of resistance trends may take a few years. Antimicrobial stewardship activities need to be enhanced in both ambulatory and inpatient settings in order to achieve optimal results.

**Disclosures.** All authors: No reported disclosures.

### 1883. Acute Kidney Injury in Patients With Pneumonia on Concomitant Anti-Methicillin-Resistant *Staphylococcus aureus* and Anti-Pseudomonal $\beta$ -Lactam Therapy

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**Background.** Empiric antibiotic treatment of serious and healthcare-associated pneumonia (PNA) often includes coverage of methicillin-resistant *Staphylococcus aureus* (MRSA) and *Pseudomonas aeruginosa* (PSA). Recent publications suggest that patients treated with the combination of vancomycin (V) and piperacillin-tazobactam (PT) have a greater risk of acute kidney injury (AKI) than those treated with V alone, or V in combination with another  $\beta$ -lactam, such as cefepime (C). There is a paucity of data regarding the risk of AKI in other regimens that provide MRSA and PSA coverage, such as linezolid (L)-PT or LC. We examined the incidence of nephrotoxicity in patients who received combination antibiotic therapy for PNA.

**Methods.** A retrospective cohort analysis of eligible adult patients ( $\geq 18$  years) admitted from July 1, 2014 to June 30, 2017 who received  $\geq 48$  hours of combination therapy was conducted. Patients were excluded if their baseline serum creatinine was  $\geq 1.4$  mg/dL, on renal replacement therapy, or if diagnosed with cystic fibrosis. The primary outcome was incidence of AKI as defined by RIFLE criteria. Comparisons

between the groups were analyzed by chi-squared test. To identify variables associated with AKI in a multivariable analysis, a repeated measures, mixed-effects logistic regression was utilized.

**Results.** There were 185 patient encounters included in the analysis. RIFLE-defined AKI occurred in treatment groups as follows: VPT 31/98 (31.6%); VC 5/50 (10.0%); LPT 4/12 (33.3%); and LC 4/25 (16.0%). There was a significant difference in rates of AKI among the four groups ( $P = 0.019$ ). In pooled analyses, no difference was identified between patients receiving V or L ( $P = 0.73$ ); however, patients who received PT had a higher incidence of AKI compared with those that received C ( $P = 0.002$ ). In logistic regression analyses, independent predictors of AKI were receipt of PT vs. C (odds ratio [OR] 3.2, 95% confidence interval [CI] 1.3–8.0) and SOFA score  $\geq 9$  (OR, 4.5; 95% CI 1.6–12.7).

**Conclusion.** No differences in AKI incidence were found between patients receiving vancomycin or linezolid; however, patients receiving piperacillin-tazobactam and those with SOFA scores  $\geq 9$  had a higher rate of AKI.

**Disclosures.** All authors: No reported disclosures.

### 1884. Assessment of Potential Antimicrobial-Related Harms in Hospitalized Adults With Common Infections

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**Background.** Recent data suggest antibiotic-related harm occurs in 1 in 5 hospitalized patients. The purpose of this study was to critically evaluate potential adverse drug-events (ADE) associated with antimicrobial administration in hospitalized family medicine (FAM) patients.

**Methods.** Retrospective cohort of adults receiving antimicrobial therapy for respiratory, urinary, and skin infection on an inpatient FAM ward between January 2017 and March 2018. Primary endpoint: potential ADEs up to 30-day post-therapy, identified using inpatient and outpatient electronic medical records. ADEs were classified as mild, moderate, or severe; Naranjo scores were used to classify causality. Other endpoints included risk associated with ADE, subsequent 30-day readmissions, and infections due to multidrug-resistant organisms up to 90-days post-therapy.

**Results.** 1,499 antibiotic days were assessed in 150 hospitalized adults. Fifty-four patients with at least one potential ADE (68 total) were identified. By Naranjo score, 10 (6.7%) patients had “probable” antibiotic related ADEs (score 5–8), all others were “possible” (score 1–4). Excluding patients with diarrhea receiving concomitant laxatives, 36 patients (24%) suffered from 50 potential ADEs, approximately 3.33 per 100 antibiotic days (Table 1). Thirteen (9.3%) had serious ADEs; 6 were receiving concomitant medications which may have contributed to harm, primarily nephrotoxins (5/6). Alteration of antimicrobial therapy was attributed to ADEs in 12/54 cases (22.2%) while 6 (11.1%) led to 30-day hospital or emergency department (ED) revisits. ADEs were not associated with any specific antimicrobial. Patients with ADEs were more likely to have ED/hospital revisits (OR = 2.42 [1.16–5.05]) and receive more total antibiotic days (11 [6–15] vs. 8 days [6–12 days],  $P = 0.036$ ) compared with those who did not.

**Conclusion.** One in four hospitalized FAM patients receiving antimicrobials experienced potential ADE. While varying in nature and severity, antimicrobial ADEs contribute to serious harm. These findings underscore need for improved awareness and judicious use.

|  | n (%)<br>(n=150) | Naranjo score<br>(range) |
|--|------------------|--------------------------|
| Patients experiencing only mild/moderate ADE | 41 (27.3)        | 3 (1–7)                  |
| Gastrointestinal                             |                  |                          |
| o Any diarrhea                               | 24 (16)          | 3 (1–7)                  |
| o Diarrhea without laxative                  | 6 (4)            | 4 (3–7)                  |
| o Nausea/vomiting                            | 7 (4.7)          | 3 (1–5)                  |
| Cardiac                                      |                  |                          |
| o QTc prolongation no event                  | 8 (5.3)          | 4 (3–4)                  |
| Renal (risk)*                                | 9 (6)            | 3 (2–5)                  |
| Transaminitis                                | 2 (1.3)          | 4 (2–6)                  |
| Altered mentation/neuropathy                 | 3 (2)            | 4 (3–5)                  |
| Patients experiencing severe ADE             | 13 (8.7)         | 4 (3–7)                  |
| Hematologic                                  | 5 (3.3)          | 4 (3–6)                  |
| Renal injury/failure*                        | 4 (2.7)          | 4.5 (3–6)                |
| C. difficile diarrhea                        | 1 (0.7)          | 7                        |
| Neurotoxicity                                | 2 (1.3)          | 4                        |
| Anaphylaxis/hives                            | 1 (0.7)          | 4                        |
| Cardiac event                                | 2 (1.3)          | 4 (2–6)                  |
| Subsequent infection with MDRO**             | 7 (4.7)          | n/a                      |

\*AKIN criteria by serum creatinine increase

\*\*Multi-drug resistant organisms: MRSA, VRE, ESB, CRE, MDR *Pseudomonas* or *Acinetobacter* spp without prior colonization

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