The Antibiotic Prescribing Pathway for Presumed Urinary Tract Infections in Nursing Home Residents

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OBJECTIVES: Due to the high rates of inappropriate antibiotic prescribing for presumed urinary tract infections (UTIs) in nursing home (NH) residents, we sought to examine the antibiotic prescribing pathway and the extent to which it agrees with the Loeb criteria; findings can suggest strategies for antibiotic stewardship.

METHODS: Chart review of 260 randomly-selected cases from 247 NH residents treated with an antibiotic for a presumed UTI in 31 NHs in North Carolina. We examined the prescribing pathway from presenting illness to the prescribing event, illness work-up and subsequent clinical events including emergency department use, hospitalization, and death. Analyses described the decision-making processes and outcomes and compared decisions made with Loeb criteria for initiation of antibiotics.

RESULTS: Of 260 cases, 60% had documented signs/symptoms of the presenting illness and 15% met the Loeb criteria. Acute mental status change was the most commonly documented sign/symptom (24%). NH providers (81%) were the most common prescribers and ciprofloxacin (32%) was the most commonly prescribed antibiotic. Fourteen percent of presumed UTI cases included a white blood cell count, 71% included a urinalysis, and 72% had a urine culture. Seventy-five percent of cultures grew at least one organism with \( \geq 100,000 \) colony-forming units/milliliter and 12% grew multi-drug resistant organisms; 28% of antibiotics were prescribed for more than 7 days, and 7% of cases had a subsequent death, emergency department visit, or hospitalization within 7 days.


Key words: older adults; antibiotic stewardship; decision-making

Suspected urinary tract infections (UTIs) in adults aged 65 years and older account for over 8 million office visits and 1 million emergency department visits, with estimated annual treatment costs exceeding \$2.3 billion.\(^1\) Few clinical problems pose more decision-making difficulties than the management of suspected UTIs\(^3\)–\(^5\) due to the absence of a diagnostic gold standard exist and high rates of asymptomatic bacteriuria (ASB).\(^6,7\) Also, significant disagreement exists in clinical practice as to what constitutes a UTI.\(^8\)

Among nursing home (NH) residents, clinical decision-making for suspected UTIs poses additional challenges for a variety of reasons. For one, ASB rates are particularly high in NHs; as many as half of all positive urine cultures should be considered false positives for the presence of a UTI.\(^9\) Additionally, due to high rates of dementia among NH residents, nurses must often infer the presence of symptoms based on nonverbal responses.\(^10\) Further, NH providers, such as physicians, physicians’ assistants, and nurse practitioners, often make decisions with extremely limited information, not knowing an individual resident’s medical history\(^11,12\) or not examining a resident at the time of illness presentation.\(^13\) Although evidence shows that clinical prescribing guidelines for UTIs such as the Loeb criteria can reduce overprescribing in NHs,\(^14\) broader
evidence suggests that NH nurses and providers often deviate from guidelines when making antibiotic prescribing decisions.15–18 Instead of relying on evidence-based guidelines, they also consider non-specific signs and symptoms (e.g., lack of appetite) and non-symptom information such as a patient or family request for an antibiotic, which can foster overprescribing.19

Looking at the prescribing pathway (the resident’s presenting illness episode, the prescribing event, the work-up, and subsequent clinical events) may shed light on the importance of each component of the pathway and provide targets for future interventions. Previous studies have been small or focused on only one part of the process. A prospective study of 100 NH residents evaluated the association between three common diagnostic guidelines for UTI and positive urine cultures.20 The positive predictive value of these guidelines was around 60%. Another study yi

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As part of a larger study of antibiotic stewardship, we collected baseline data to understand the prescribing pathway for presumed UTIs. We examined documented clinical symptoms (particularly the Loeb criteria), prescriber characteristics, antibiotic prescribed, rates of antibiotic cessation after a negative urine culture, and subsequent clinical events. We aimed to determine the frequency of prescriptions that met Loeb criteria, and the rate of discontinuation of antibiotics in the face of negative urine cultures.

METHODS

Study Design

Part of a dissemination effort to improve infection management and antibiotic stewardship, we enrolled 31 community-based NHs in North Carolina. At baseline a team of geriatricians and research staff visited each NH between November 2014 and March 2015 to conduct medical record audits. In each NH, we randomly selected ten “new” cases of treated UTIs from the infection log for the preceding month. A new case was defined as an antibiotic prescription for a UTI without a prior prescription for a UTI in the past 21 days, assuming that antibiotics clustered closer together were a part of the same illness episode and not discrete UTIs. In NHs having fewer than ten cases in the month, all available cases were audited, yielding a total of 260 cases. The University of North Carolina Institutional Review Board approved the study. The funding source had no role in the study.

Population

To be eligible for study participation, cases had to be recorded on each NH’s infection control log and resulted in a systemic antibiotic for a presumed UTI. Residents were eligible regardless of whether they were in short-term rehabilitation or long-term stay beds.

Measures

We developed an abstraction form to record elements in the antibiotic prescribing pathway. The form included information about the presenting illness episode (i.e., vital and physical signs and symptoms), the prescribing event (i.e., the type and administrative route of the antibiotic, where it was prescribed and by whom), the illness work-up (i.e., laboratories, urinalysis and urine culture performed, and results), the antibiotic treatment course (i.e., when started and changed or stopped, and duration), and subsequent clinical events such as emergency department use, hospitalization, and death (in the 7 days after the antibiotic was initiated). Descriptive information about the NHs and residents also were obtained.

We created three variables from the existing data: (1) whether the presenting illness met the minimum set of prescribing criteria developed by Loeb et al.;21 (2) whether at least one of the Loeb criteria was present; and (3) whether the cultured organism met criteria for a multi-drug resistant organism per the Centers for Disease Control criteria.22 The Loeb minimum criteria for prescribing are consensus-guidelines and differ depending on whether or not a resident has an indwelling-catheter.14,21 In residents with indwelling urinary catheters, a patient is considered to have a UTI and a prescription is appropriate if the resident has either a fever (temperature >100°F) or new onset of costovertebral angle tenderness, symptoms of rigors, or new symptoms of delirium. If there is not a catheter, a patient is considered to have a UTI and a prescription is appropriate if the resident has either pain or difficulty with urination alone, or fever and at least 1 of the following: new or increased urinary urgency, new or increased urinary frequency, new or increased suprapubic pain, new costovertebral angle tenderness, frank hematuria, or new or worsened urinary incontinence. We conducted an additional analysis using the Loeb criteria but considering fever to be ≥99.0°F instead of >100°F, based on recent evidence.23 A positive urine culture was considered ≥10,000 colony-forming units/milliliter (CFU/mL) in those with a documented catheterized specimen and ≥100,000 CFU/mL for all other specimens. In terms of determining multi-drug resistance, if results of the urine culture and sensitivities were not present, the results were presumed to be negative.

Data Analysis

Descriptive statistics were used to examine the data, including frequencies, percentages, means, and standard deviations (SD). When the classification of symptom and signs was unclear, two authors (CK and PS) reviewed the data and assigned them to existing categories if they independently agreed on their reassignment. CK summarized the 17 cases of emergency department use, hospitalization, and death. All quantitative analyses were conducted using SPSS (Armonk, NY: IBM Corporation), version 23.

RESULTS

Of the 31 NHs, most were for-profit (77%) with a mean size of 109 licensed beds (Table 1). During the 1-month

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Table 1. Nursing Home and Resident Characteristics

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>n (%) or Mean (SD)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nursing home characteristics (n = 31)</td>
<td></td>
</tr>
<tr>
<td>Nursing home chain</td>
<td>15 (48%)</td>
</tr>
<tr>
<td>Long-term care medical practice&lt;sup&gt;a&lt;/sup&gt;</td>
<td>16 (52%)</td>
</tr>
<tr>
<td>For-profit</td>
<td>24 (77%)</td>
</tr>
<tr>
<td>Has a dementia unit</td>
<td>5 (16%)</td>
</tr>
<tr>
<td>Number of licensed beds per site, mean ± SD</td>
<td>109 ± 44</td>
</tr>
<tr>
<td>Staff turnovers in prior 5 years (number per site), mean ± SD</td>
<td></td>
</tr>
<tr>
<td>Administrator</td>
<td>2 ± 1</td>
</tr>
<tr>
<td>Director of Nursing</td>
<td>3 ± 2</td>
</tr>
<tr>
<td>Infection control nurse</td>
<td>2 ± 2</td>
</tr>
<tr>
<td>Medical Director</td>
<td>2 ± 1</td>
</tr>
<tr>
<td>Licensed nurses (RN/LPN)</td>
<td>29 ± 14</td>
</tr>
<tr>
<td>Resident characteristics (n = 247)&lt;sup&gt;b&lt;/sup&gt;</td>
<td></td>
</tr>
<tr>
<td>Age, mean ± SD (range)</td>
<td>81 ± 13 (36–110)</td>
</tr>
<tr>
<td>Gender, female</td>
<td>184 (75%)</td>
</tr>
<tr>
<td>On hospice</td>
<td>15 (6%)</td>
</tr>
<tr>
<td>Routine catheter use&lt;sup&gt;c&lt;/sup&gt;</td>
<td></td>
</tr>
<tr>
<td>Indwelling urinary catheter</td>
<td>37 (15%)</td>
</tr>
<tr>
<td>Routine intermittent catheterization</td>
<td>5 (2%)</td>
</tr>
<tr>
<td>Suprapubic catheter</td>
<td>13 (5%)</td>
</tr>
<tr>
<td>None of the above</td>
<td>179 (73%)</td>
</tr>
<tr>
<td>Unknown</td>
<td>13 (5%)</td>
</tr>
</tbody>
</table>

SD = Standard Deviation; RN = Registered Nurse; LPN = Licensed Practical Nurse.

<sup>a</sup>Long-term care medical practice is a group of medical providers including physicians, physician assistants, and nurse practitioners who provide care only to NH residents.

<sup>b</sup>Thirteen residents had repeated treated UTIs, resulting in a total of 260 cases of treated UTIs.

<sup>c</sup>Excludes ileal conduit.

study period, we reviewed 247 NH residents’ charts who received 260 antibiotic prescriptions for a presumed UTI, with 13 (5%) residents receiving two prescriptions. Most of the NH residents who received an antibiotic were female (75%) and without routine catheter use (73%), with a mean age of 81 (range 36–110).

Presenting Illness

Overall, 60% of cases had at least one sign or symptom documented in the 48 hours prior to when the prescription was written. Of the 156 cases with documented signs or symptoms, 106 (68%) met at least one Loeb criterion (41% of all cases; see Table 2). In total, 15% met all Loeb criteria, and when we used a less stringent criterion for fever (temperature ≥99.0°F), 17% met criteria. The most common urinary tract-specific sign or symptom was dysuria (9%). Overall, acute mental status change was the most commonly documented sign or symptom (24%).

Prescribing Event

NH residents most commonly received oral antibiotics (87%), with ciprofloxacin being the most common prescription (32%), followed by trimethoprim-sulfamethoxazole (14%) and nitrofurantoin (11%) (see Table 3). Most antibiotics were prescribed by NH providers (81%). A minority were prescribed during a hospital inpatient stay (10%), emergency department visit (6%), or visit to a medical office (2%). Of the 194 prescriptions for whom the prescriber’s profession was noted, physicians prescribed 52%, nurse practitioners prescribed 38%, and physician’s assistants prescribed 10%.

Illness Work-Up

As part of the illness work-up, white blood cell (WBC) counts, urinalyses, and urine cultures were performed for some residents. Fourteen percent of cases had a WBC count performed within the 3 days prior to the antibiotic (Table 4). Of the 71% who had a urinalysis performed, 10% were negative and 29% were both nitrite and leukocyte esterase positive. Similarly, 72% of cases had a urine culture performed, of which 3% were unknown, 3% resulted no bacteria, and 4% were polymicrobial. Ninety percent of cultures resulted some type and quantity of organism; 78% were considered positive cultures based on our definition. Forty three of the 70 cases missing a urine culture (29%) were prescribed outside of the NH. Cases diagnosed in the NH were more likely to have an available urine culture (P < .001). Escherichia coli (E. coli) resulted most often (34%), followed by Klebsiella (14%) and Proteus (12%) as shown in the Figure 1. Multi-drug resistant
organisms (MDROs) were found in 12% of positive cultures, and only 7% of organisms were pan-sensitive. Although not statistically significant, residents with indwelling catheters were more likely to be diagnosed with an MDRO than those without (8% vs. 4%, P = .43).

### Antibiotic Treatment Course

Antibiotics were prescribed for a mean duration of 7.5 days (SD ± 2.7); 28% of prescriptions exceeded 7 days, 3% exceeded 10 days, and 9% were for 5 days or less. In total, 2% of prescriptions were discontinued, and 19% were changed during the illness course. On average, residents with indwelling catheters received antibiotics for 8.4 days (95% CI 7.5, 9.2) as compared to those without who received antibiotics for 7.2 days (95% CI 6.9, 7.5) (P = .008).

### Clinical Events

Of the 247 cases, 7% had a subsequent clinical event. Two died during the course of treatment, eleven required hospitalization, and 4 required emergency department visits. Of the 11 hospitalized residents, 5 (45%) had 1 or more of the signs or symptoms found in the Loeb criteria and three (27%) met the Loeb criteria. One of the residents who died was on hospice.

### DISCUSSION

In this study of 31 NHs, examination of the prescribing pathway for presumed UTIs revealed a number of findings that may explain why adherence to infection control guidelines continues to be poor and how each contributes to inappropriate antibiotic prescribing and the development of MDROs. As a sign of potentially inappropriate prescribing, non-specific signs or symptoms, particularly acute mental status change, were more commonly found with antibiotic prescriptions than the expert-derived Loeb criteria for antibiotic initiation. Most antibiotics were prescribed by NH providers, and antibiotics were rarely discontinued before the course was completed. In addition, for a sizeable minority, the duration of antibiotics appeared overly long (i.e., 28% were prescribed for more than 7 days, which exceeds the recommended duration of 3–7 days.24,25 Among positive cultures, 92% grew an organism resistant to at least one antibiotic.

Diagnostic errors along the pathway appear to play a significant role in inappropriate antimicrobial prescribing in NH residents, as also seen in hospitalized patients.26 One of the early components of the prescribing pathway, namely signs and symptoms of the presenting illness, may represent an important area for intervention. Despite the existence of evidence-based guidelines for more than a decade, we continue to see that most presumed UTIs fail to meet minimum requirements for appropriate antibiotic initiation.10,18,27,28 Admittedly, one possible explanation for apparent inappropriate prescribing may be the lack of documentation in NH charts. Even when symptoms were documented, however, non-specific signs and symptoms such as confusion or agitation (which have shown little to no association with UTIs29) were documented most frequently. Clinical assessments in the NH are particularly difficult. Due to high rates of dementia, nurses often must coax symptom reports from residents.10 Nurses may note a patient’s non-specific change in condition, which then sets off a cascade of information gathering and further decisions, including calling a physician, that result in an antibiotic prescription. NH providers often have limited familiarity with a NH resident,11,12 and may not examine a resident with a suspected UTI until days after an antibiotic has been initiated. In one study, only 29% of residents with unstable vital signs were examined by a provider on-site or prior to transfer to a hospital.13 Further, although 78% of providers awaited culture results prior to prescribing, we found limited evidence that residents were reassessed to determine if the clinical picture warranted a UTI diagnosis or if the culture results represented ASB. Given the non-specific nature of many signs or symptoms used to diagnose suspected UTIs, a period of watchful waiting with non-pharmacologic management, such as timed toileting and oral hydration, with a reassessment of the resident’s status may provide a reasonable approach to reducing unnecessary prescribing.

Most of these prescriptions (81%) were written by NH providers, whose knowledge of appropriate antibiotic prescribing is historically low.30 To address this issue, efforts such as clinical decision support systems (CDSS) or peer evaluations may improve prescribing quality and safety in NHs. A systematic review of CDSS in NHs to improve medication safety31 found that CDSS use for residents with renal insufficiency significantly increased appropriate medication orders by 20%.32 A recent RCT of 47 primary care practices testing a CDSS to support
prescribing for upper respiratory infections also found antibiotics prescribing fell significantly compared to controls. Consequently, work is needed to see if targeted CDSS for the diagnosis of infections can be translated from the outpatient setting to the NH.

Other steps in the pathway may benefit from potential interventions. In hospital settings, discontinuing reflexive urine cultures has reduced antibiotic prescribing. A similar intervention may be useful in NHs. Even when faced with negative cultures, providers apparently failed to reexamine their diagnosis and discontinue antibiotics—despite the fact that discontinuing antibiotics, when no infection is present, reduces antibiotic resistance. Outpatient treatment guidelines for UTIs in the United Kingdom explicitly state that antibiotics should be discontinued if a culture yields no bacterial count. Yet our providers discontinued antibiotics in only 4 of 6 cases with no bacterial growth. Standing orders for antibiotic cessation when urine cultures are negative may reduce antibiotic use.

Infectious Disease Society of America (IDSA) recommends that nitrofurantoin, trimethoprim-sulfamethoxazole, fosfomycin, or pivmecillinam be used before a fluoroquinolone (e.g., ciprofloxacin or levofloxacin) or beta-lactam. Fluoroquinolones are discouraged because of antimicrobial resistance and the duration of treatment is also sometimes inappropriate. Our study shows continued frequent prescribing of fluoroquinolones (40% of cases) and less-frequent prescribing of recommended first-line treatments (25% of cases). The Infectious Disease Society of America (IDSA) recommends that nitrofurantoin, trimethoprim-sulfamethoxazole, fosfomycin, or pivmecillinam be used before a fluoroquinolone (e.g., ciprofloxacin or levofloxacin) or beta-lactam. Fluoroquinolones are discouraged because of side effects ranging from neurologic toxicity to cardiac abnormalities, and cross-resistance between ciprofloxacin and levofloxacin is nearly universal.
Administration released an updated warning around fluoroquinolones, stating the serious side effects seen outweigh the benefits of use. In addition, providers in our study prescribed overly long courses of antibiotics, over a quarter prescribed for more than 1 week, although guidelines recommend durations longer than 7 days in only a minority of cases. One explanation for the longer duration of treatment may be that NH providers are concerned about complicated UTIs such as CAUTIs. The duration for antibiotic treatment for CAUTIs is somewhere between 5 and 14 days per various guidelines. We found the duration of antibiotics was a little over a day longer between those with and without catheters (both averaged over 7 days), suggesting that possible concerns over complicated infections were less likely. IDSA recommends the shortest effective duration of antibiotic treatment possible, noting that shorter duration does not increase treatment failure nor increase adverse clinical events in older women with UTIs.

The generalizability of our findings is limited by geographic locale and number of NHs. The study NHs, however, did not differ significantly from all NHs nationally. In addition, because we relied on the infection log to identify eligible residents, we cannot comment on any cases or residents that failed to make the infection log. Also, for a few variables, we had more than 10% missing data, rendering uncertain the true frequency of some findings, such as prescriber type. Certain data elements for sepsis, such as heart rate and respiratory rate were not included in the chart abstraction, making it difficult to determine UTI versus more complicated infections such as pyelonephritis, which might argue for a longer duration of antibiotic treatment. Lastly, we did not look for resolution, continuation, or development of symptoms after the antibiotic.

**CONCLUSION**

This study revealed that non-specific signs and symptoms were most often found in cases of presumed UTIs; that NH providers represented the largest group of antibiotic prescribers; that prescribers often failed to discontinue antibiotics even when clinically reasonable; and that non-recommended antibiotics were often prescribed and for overly long durations. Three ways to improve guideline adherence are: (1) providing educational activities that specifically target NH nurses and NH providers to reduce the over-reliance on non-specific signs/symptoms and urine culture results, (2) encouraging antibiotic cessation in the face of negative cultures, and (3) improving the antibiotic choice and shortening the antibiotic duration. Implementing antibiotic stewardship programs offers an evidence-based way to increase the appropriate use of antibiotics.

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