Frequent hospital readmissions for Clostridium difficile infection and the impact on estimates of hospital-associated C. difficile burden

Courtney R. Murphy
University of California - Irvine
Taliser R. Avery
Harvard University
Erik R. Dubberke
Washington University School of Medicine in St. Louis
Susan S. Huang
University of California - Irvine

Follow this and additional works at: https://digitalcommons.wustl.edu/open_access_pubs

Part of the Medicine and Health Sciences Commons

Please let us know how this document benefits you.

Recommended Citation
Murphy, Courtney R.; Avery, Taliser R.; Dubberke, Erik R.; and Huang, Susan S., "Frequent hospital readmissions for Clostridium difficile infection and the impact on estimates of hospital-associated C. difficile burden." Infection Control and Hospital Epidemiology. 33, 1. 20-28. (2012). https://digitalcommons.wustl.edu/open_access_pubs/785

This Open Access Publication is brought to you for free and open access by Digital Commons@Becker. It has been accepted for inclusion in Open Access Publications by an authorized administrator of Digital Commons@Becker. For more information, please contact vanam@wustl.edu.
Frequent Hospital Readmissions for *Clostridium difficile* Infection and the Impact on Estimates of Hospital-Associated *C. difficile* Burden

Courtney R. Murphy, MS; Taliser R. Avery, MSc; Erik R. Dubberke, MD, MSPH; Susan S. Huang, MD, MPH

**Objective.** *Clostridium difficile* infection (CDI) is associated with hospitalization and may cause readmission following admission for any reason. We aimed to measure the incidence of readmissions due to CDI.

**Design.** Retrospective cohort study.

**Patients.** Adult inpatients in Orange County, California, who presented with new-onset CDI within 12 weeks of discharge.

**Methods.** We assessed mandatory 2000–2007 hospital discharge data for trends in hospital-associated CDI (HA-CDI) incidence, with and without inclusion of postdischarge CDI (PD-CDI) events resulting in rehospitalization within 12 weeks of discharge. We measured the effect of including PD-CDI events on hospital-specific CDI incidence, a mandatory reporting measure in California, and on relative hospital ranks by CDI incidence.

**Results.** From 2000 to 2007, countywide hospital-onset CDI (HO-CDI) incidence increased from 15 per 10,000 to 22 per 10,000 admissions. When including PD-CDI events, HA-CDI incidence doubled (29 per 10,000 in 2000 and 52 per 10,000 in 2007). Overall, including PD-CDI events resulted in significantly higher hospital-specific CDI incidence, although hospitals had disproportionate amounts of HA-CDI occurring postdischarge. This resulted in substantial shifts in some hospitals’ rankings by CDI incidence. In multivariate models, both HO and PD-CDI were associated with increasing age, higher length of stay, and select comorbidities. Race and Hispanic ethnicity were predictive of PD-CDI but not HO-CDI.

**Conclusions.** PD-CDI events associated with rehospitalization are increasingly common. The majority of HA-CDI cases may be occurring postdischarge, raising important questions about both accurate reporting and effective prevention strategies. Some risk factors for PD-CDI may be different than those for HO-CDI, allowing additional identification of high-risk groups before discharge.

*Infect Control Hosp Epidemiol* 2012;33(1):20-28

Hospital length of stay has steadily decreased over the past 30 years, and increasingly, complex medical care is provided after discharge through home health and skilled nursing facilities. In turn, adverse events related to hospitalization, including hospital-associated infection, may increasingly be present after discharge and result in readmission. The costs and sequelae of hospital readmission have made it a target for hospital quality indicators and value-based purchasing. Currently, the Centers for Medicare and Medicaid Services reports hospitals’ rates of readmission following treatment for myocardial infarction, congestive heart failure, and pneumonia, prompting a national focus on preventing readmission. However, readmission rates for other important conditions, such as hospital-associated infections (HAIs), are not well studied despite national and state requirements for reporting hospital-specific rates of HAIs.

*Clostridium difficile* infection (CDI) is a common cause of diarrhea in healthcare settings and may be an important source of hospital readmissions. Hospital-associated *C. difficile* acquisition may not be evident until after hospital discharge, especially since the average hospital length of stay is 3–5 days and acquisition may initially be asymptomatic. In addition, risk factors related to medical care, such as predisposing antibiotics, may require time to deplete the normal intestinal flora and allow *C. difficile* to flourish and produce symptoms. The exact incubation period for CDI is unknown, but 3 studies found the incubation period to be less than 1 week. However, several studies have found patients may be at an increased risk for developing CDI up to 3 months after hospital discharge. One recent study found that among patients who developed CDI within 100 days postdischarge, 89% of patients developed CDI in the first 60 days.
and 85% occurred in the first month. To account for the range of incubation period, national guidance considers CDI occurring within 4 weeks of hospitalization as hospital-associated, and CDI occurring within 4–12 weeks of a hospitalization as potentially hospital associated. Concerns about CDI have been increasing in the United States. Hospitals’ incidence of CDI has been rising in the past decade. This has been associated with the emergence of a new epidemic strain, BI/NAP1/027, that produces 20-fold more toxin than other strains and is associated with high rates of colectomy and death. In fact, there is evidence in some hospitals that CDI prevalence may have surpassed that of methicillin-resistant Staphylococcus aureus (MRSA). With continued pressure to reduce hospital length-of-stay, the frequency of postdischarge CDI (PD-CDI)—defined here as community-onset CDI within 12 weeks after any hospitalization—may be increasing, along with the opportunity for prevention.

In response to rising CDI incidence, the Centers for Disease Control and Prevention (CDC) and the Society for Healthcare Epidemiology of America (SHEA) have recommended surveillance of healthcare-associated CDI (HA-CDI) rates, which includes both hospital-onset as well as community-onset healthcare-associated CDI. In addition, reporting of CDI rates has been legislated or is under legislative consideration in several states. Despite national guidance that postdischarge CDI events occurring within 4 weeks should be considered hospital associated and events between 4–12 weeks of discharge could potentially be hospital associated, hospitals performing CDI surveillance often do not track PD-CDI events. However, tracking postdischarge events may facilitate efforts to prevent readmissions and may be helpful for reporting hospital-specific CDI incidence. Moreover, patients requiring readmission for PD-CDI may not return to the original facility, suggesting that the incidence of HA-CDI may be significantly underestimated if PD-CDI events are not uniformly identified among hospitals.

We sought to identify CDI cases occurring at all hospitals in a large California county (population 3 million). We assessed the frequency of admission for new-onset CDI after a recent hospitalization and the impact of including PD-CDI events resulting in readmission on hospital-specific CDI incidence.

**METHODOLOGY**

**Description of Data Set**

We conducted a population-based retrospective cohort study to assess the frequency of postdischarge CDI events among adult inpatients in all 29 hospitals serving adults in Orange County, California, from January 1, 2000, to December 31, 2007. We used mandatory California hospital discharge data which provides line-item demographic and insurer information, ICD-9 codes (up to 25), and a unique identifier (record linking number) that allows patients to be tracked across hospital admissions. This data also includes a code
to indicate whether a given condition was present when the patient was admitted, known as the “present on admission” (POA) code, which has been used in California since 1996.34 We identified CDI cases using the ICD-9 diagnostic code 008.45 for pseudomembranous colitis. We defined 4 types of CDI cases: (1) hospital-onset CDI (HO-CDI) cases defined by POA = N (no); (2) PD-CDI cases defined by POA = Y (yes) with a history of hospitalization for any reason in the prior 12 weeks; (3) HA-CDI cases defined as the sum of HO-CDI and PD-CDI; and (4) community-associated CDI (CA-CDI) cases defined by POA = Y with no prior history of hospitalization in the previous 12 weeks. While we used 12 weeks for our primary analysis, we repeated all analyses using a 4-week cutoff for comparison. To reduce the chance that a code represented a past history of CDI without active infection during hospitalization, we limited cases with POA = Y to the first 3 coding positions. For POA = N cases, all coding positions were accepted. We excluded 932 cases of recurrent CDI, defined as cases occurring within 8 weeks of a previous CDI episode.9,22 Finally, we assessed the fraction of postdischarge events that occurred within 4 weeks of discharge. This study was approved by the Institutional Review Boards of the University of California Regents and the California Committee for the Protection of Human Subjects.

**Data Analysis**

**Patient Characteristics**

We collected demographic information for all patients in our cohort, including gender, age, race and ethnicity, and insurance type. We also assessed the proportion of hospitalized patients with select comorbidities using the Romano score34 and the proportion that had undergone surgery in the previous month. These characteristics were collected for all admissions and for those with CDI (HO-CDI, PD-CDI, and CA-CDI).

**Annual Incidence of CDI**

Annual CDI incidence across Orange County was determined for 2000–2007 and analyzed by χ² tests for trend. We identified all cases and subsets of CDI as defined above. Incidences of HO-CDI, PD-CDI, and HA-CDI were expressed per 10,000 admissions. CA-CDI incidence was expressed per 100,000 residents.

**Hospital Readmission for CDI**

We defined a PD-CDI readmission as a case with symptoms present on admission (POA = Y) that occurred within 12 weeks after a prior hospitalization for any reason, as described above. We calculated the percentage of all-cause readmissions that are due to PD-CDI. We excluded readmissions for recurrent CDI, which we defined as community-onset (POA = Y) cases readmitted within 8 weeks of a previous admission for CDI.22 We also determined how often patients readmitted for PD-CDI went to a different facility for their readmission.
Impact of Including Postdischarge CDI Readmissions in Hospital-Specific CDI Incidence

For each hospital, we determined the annual incidence of HO-CDI and HA-CDI for the years 2000–2007. Differences between annual HO-CDI and HA-CDI incidence were compared using paired t tests. We determined whether relative rankings by quartile of hospitals by CDI incidence were affected by inclusion of PD-CDI.

Identifying Individual and Hospital Predictors of CDI

We identified the primary admission diagnoses of admissions that were associated with HO-CDI and PD-CDI. For primary admission diagnoses associated with greater than 25 HO-CDI or PD-CDI events, we calculated the frequency of CDI compared to those without that primary admission diagnosis.

We performed bivariate analyses using χ² tests to identify individual and hospital level variables associated with the individual outcomes of HO-CDI and PD-CDI. For the PD-CDI outcome, we used characteristics from the PD-CDI (vs the index) admission and removed all hospitalizations that resulted in death, since these hospitalizations could not result in readmission. Individual variables included demographics, comorbidities, primary admission diagnosis, recent surgery, insurance type, year of hospital admission, and length of stay. Hospital variables included annual admissions, average length of stay, and hospital type (acute vs long-term care facility). Variables with P < .1 from bivariate testing were entered into a generalized linear mixed model which accounted for clustering by hospital (ProcGLIMMIX, SAS 9.2; SAS). Variables were retained at α = 0.05.

RESULTS

Patient Characteristics

Patients admitted with CDI were older, had more comorbidities, and were less likely to have undergone surgery in the past month compared to all hospitalized patients (Table 1). Among those with CDI, patients with HO-CDI and PD-CDI had similar distributions of age, race and ethnicity, and co-morbidities, but those with HO-CDI were more likely to be male and to have undergone surgery in the past month.

Annual Incidence of HO-CDI and HA-CDI

Annual incidence of HO-CDI in Orange County increased from 2000 to 2007, as shown in Figure 1 (P < .001 for test of trend). After including PD-CDI events, the annual incidence of HA-CDI increased 1.9-fold during the same period, from 28.7 to 52.2 per 10,000 admissions (χ², P < .001). By 2007, PD-CDI comprised the majority of HA-CDI cases (increasing from 46% in 2000 to 57% in 2007; P < .001 for test of trend). Restricting PD-CDI cases to the 4 weeks following discharge captured 73% of PD-CDI cases and resulted in similar annual incidences of HA-CDI, increasing 1.8-fold from 28.0 per 10,000 admissions in 2000 to 51.6 per 10,000 admissions in 2007 (χ², P < .001).

Frequency of New-Onset CDI as Reason for Hospital Readmission

Over 2000–2007, PD-CDI events resulting in readmission represented 1.8% (2,998 of 170,995) of all-cause readmissions within 12 weeks after discharge. When evaluating all admissions related to CDI occurring within 365 days of discharge, we found that the risk of readmission for CDI was higher in the first 12 weeks postdischarge, and highest in the first 4 weeks postdischarge (Figure 2). Of PD-CDI events occurring within 12 weeks of discharge, 58% (624 of 1071) occurred within the 4 weeks after discharge. After 12 weeks, the risk of readmission for CDI dropped to a stable, low level. Among PD-CDI cases readmitted within 12 weeks, 25% (746 of 2,998) were readmitted to a different hospital than the initial hospitalization.

Impact of Including CDI Readmissions on Hospital-Specific Rates

Figure 3 shows hospital-specific rankings according to CDI incidence for 2007, with and without including PD-CDI events (HO-CDI vs HA-CDI, respectively). The proportion of hospitals’ HA-CDI comprised by PD-CDI varied greatly (median 60% PD-CDI, range 0%–100%, for 2007). Hospital ranking by CDI incidence changed by a mean of 3 places after including PD-CDI events; only 5 of 30 hospitals did not change rank. Three hospitals became ranked in the worst quartile after including PD-CDI, including 1 hospital that had been ranked in the best quartile when PD-CDI events were excluded. Another 3 hospitals were no longer ranked in the worst quartile when PD-CDI events were included.

When restricting PD-CDI events to 4 weeks postdischarge, the proportion of HA-CDI comprised by PD-CDI similarly varied from 0%–100% (median 46%). Hospital rankings changed by a mean of 2.5 places, and 10 of 30 hospitals
changed quartile. Nine hospitals did not change rank; these hospitals all had zero CDI events.

**Identifying Individual and Hospital Predictors of CDI**

Primary admission diagnoses that occurred most often during HO-CDI and PD-CDI admissions are listed in Table 2. Several primary admission diagnoses were significantly associated with both HO-CDI and PD-CDI admissions on bivariate analysis, including septicemia, pneumonia, postoperative infection, and urinary tract infection.

Results from bivariate analysis (Table 3) were similar to those from multivariate analysis (Table 4). In multivariate analysis, HO-CDI and PD-CDI were both associated with increasing age, longer length of stay, Medicare insurance, recent surgery, comorbidities, select primary admission diagnoses (septicemia, postoperative infection, and pneumonia), and hospitals with a high percent of patients with a high comorbidity index. Non-white race, Hispanic ethnicity, and male gender were protective against PD-CDI but not HO-CDI.

**Annual Incidence of Community-Associated CDI**

Orange County’s incidence of CA-CDI also rose during 2000 to 2007. In this period, CA-CDI incidence increased 2.1-fold from 9.1 to 19.4 cases per 100,000 residents ($\chi^2, P < .001$), exclusive of PD-CDI cases. These rates were similar when PD-CDI was restricted to events within 4 weeks of discharge (9.5 to 19.8 cases per 100,000 residents; $\chi^2, P < .001$).

**DISCUSSION**

*Clostridium difficile* disease is a major cause of healthcare-associated infection and morbidity. Due to the known delay in presentation following antibiotic exposure, national guidelines consider cases up to 12 weeks following hospital discharge as potentially healthcare-associated and possibly preventable. Nevertheless, the majority of hospitals do not track postdischarge cases, and the impact of postdischarge cases has remained largely unknown. Remarkably, we found that PD-CDI cases within 12 weeks after hospital discharge accounted for the majority of HA-CDI and led to a 2-fold increase in HA-CDI incidence across hospitals in a large metropolitan county. These effects were largely driven by PD-CDI events within 4 weeks after hospital discharge. This finding illustrates the need to expand prevention and education strategies to include the postdischarge period and thereby reduce the frequency of PD-CDI events.

Inclusion of postdischarge CDI events substantially altered hospital-specific CDI incidence, but the impact varied widely by hospital. For example, PD-CDI cases accounted for all HA-CDI cases in one hospital and none of the cases in another. This suggests that tracking PD-CDI events may impact the validity of interfacility comparisons, since hospitals are affected differentially by including or excluding PD-CDI. These discrepancies could be magnified if only some, but not all, hospitals track PD-CDI. When we ranked hospitals by HO-CDI incidence, half the hospitals captured in the quartile with the highest HO-CDI incidence changed when PD-CDI was included. In fact, one hospital changed from the best quartile to the worst quartile when PD-CDI cases were captured. Further, changes in rank were similar when PD-CDI was restricted to events within 4 weeks postdischarge, with one-third of hospitals changing quartile after inclusion of PD-CDI. In addition, since 75% of patients with PD-CDI returned to the same hospital for readmission, hospitals may be able to track most PD-CDI cases by performing postdischarge surveillance for PD-CDI cases that readmit to their own facility. Additional notification of PD-CDI cases back to

<table>
<thead>
<tr>
<th>Primary admission diagnosis</th>
<th>N (%) with HO-CDI</th>
<th>OR</th>
<th>P value</th>
<th>N (%) with PD-CDI</th>
<th>OR</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td><em>Staphylococcus aureus</em> septicemia</td>
<td>31 (0.6)</td>
<td>10.09</td>
<td>&lt;.001</td>
<td>...</td>
<td>...</td>
<td></td>
</tr>
<tr>
<td><em>S. aureus</em> pneumonia</td>
<td>33 (0.7)</td>
<td>9.76</td>
<td>&lt;.001</td>
<td>...</td>
<td>...</td>
<td></td>
</tr>
<tr>
<td>Acute respiratory failure</td>
<td>117 (2.4)</td>
<td>8.56</td>
<td>&lt;.001</td>
<td>31 (0.4)</td>
<td>1.22</td>
<td>.007</td>
</tr>
<tr>
<td>Aspiration pneumonitis</td>
<td>99 (2.1)</td>
<td>6.55</td>
<td>&lt;.001</td>
<td>69 (0.8)</td>
<td>2.47</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Infection of vascular device</td>
<td>33 (0.7)</td>
<td>6.16</td>
<td>&lt;.001</td>
<td>26 (0.3)</td>
<td>2.65</td>
<td></td>
</tr>
<tr>
<td>Chemotherapy</td>
<td>30 (0.6)</td>
<td>5.43</td>
<td>&lt;.001</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><em>E. coli</em> septicemia</td>
<td>26 (0.5)</td>
<td>5.39</td>
<td>&lt;.001</td>
<td>26 (0.3)</td>
<td>2.94</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Septicemia</td>
<td>68 (1.4)</td>
<td>3.72</td>
<td>&lt;.001</td>
<td>184 (1.2)</td>
<td>3.20</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Postoperative infection</td>
<td>28 (0.6)</td>
<td>3.28</td>
<td>&lt;.001</td>
<td>26 (0.3)</td>
<td>1.67</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Acute renal failure</td>
<td>39 (0.8)</td>
<td>2.80</td>
<td>&lt;.001</td>
<td>47 (0.5)</td>
<td>1.85</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Pneumonia</td>
<td>98 (2.0)</td>
<td>1.69</td>
<td>&lt;.001</td>
<td>184 (2.1)</td>
<td>1.74</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Acute pancreatitis</td>
<td>31 (0.6)</td>
<td>1.51</td>
<td>.002</td>
<td>26 (0.3)</td>
<td>0.69</td>
<td>.8</td>
</tr>
<tr>
<td>Urinary tract infection</td>
<td>37 (0.8)</td>
<td>1.43</td>
<td>.19</td>
<td>96 (1.1)</td>
<td>2.04</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Hip fracture</td>
<td>...</td>
<td>...</td>
<td>...</td>
<td>31 (0.4)</td>
<td>1.76</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Cellulitis</td>
<td>...</td>
<td>...</td>
<td>...</td>
<td>45 (0.5)</td>
<td>1.69</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Colon diverticulitis</td>
<td>...</td>
<td>...</td>
<td>...</td>
<td>45 (0.5)</td>
<td>1.68</td>
<td>&lt;.001</td>
</tr>
</tbody>
</table>

**Note.** HO-CDI, hospital-onset CDI; PD-CDI, postdischarge CDI; OR, odds ratio.
### Table 3. Bivariate Analysis of Predictors of Hospital-Onset *Clostridium difficile* Infection (HO-CDI) and Postdischarge CDI (PD-CDI) Among All Adult Inpatients

<table>
<thead>
<tr>
<th>Individual variables (%)</th>
<th>HO-CDI</th>
<th>Non-HO-CDI</th>
<th>P value</th>
<th>PD-CDI</th>
<th>Non-PD-CDI</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>N</td>
<td>2,403</td>
<td>1,766,753</td>
<td>&lt;.001</td>
<td>3,077</td>
<td>1,725,165</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Age, years</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>18–&lt;40</td>
<td>6.7</td>
<td>28.7</td>
<td>&lt;.001</td>
<td>6.5</td>
<td>29.3</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>40–49</td>
<td>7.8</td>
<td>12.7</td>
<td></td>
<td>7.3</td>
<td>12.8</td>
<td></td>
</tr>
<tr>
<td>50–59</td>
<td>11.9</td>
<td>12.7</td>
<td></td>
<td>8.5</td>
<td>12.8</td>
<td></td>
</tr>
<tr>
<td>60–&lt;75</td>
<td>31.7</td>
<td>21.0</td>
<td></td>
<td>25.3</td>
<td>20.9</td>
<td></td>
</tr>
<tr>
<td>75+</td>
<td>41.9</td>
<td>24.9</td>
<td></td>
<td>52.4</td>
<td>24.2</td>
<td></td>
</tr>
<tr>
<td>Male gender</td>
<td>48.2</td>
<td>38.2</td>
<td>&lt;.001</td>
<td>39.4</td>
<td>37.9</td>
<td>.08</td>
</tr>
<tr>
<td>Race</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td>&lt;.001</td>
</tr>
<tr>
<td>White</td>
<td>80.9</td>
<td>80.3</td>
<td></td>
<td>87.7</td>
<td>80.2</td>
<td></td>
</tr>
<tr>
<td>Black</td>
<td>2.2</td>
<td>2.4</td>
<td></td>
<td>1.1</td>
<td>2.4</td>
<td></td>
</tr>
<tr>
<td>Asian</td>
<td>10.6</td>
<td>9.5</td>
<td></td>
<td>5.6</td>
<td>9.5</td>
<td></td>
</tr>
<tr>
<td>Other</td>
<td>6.3</td>
<td>7.8</td>
<td></td>
<td>5.6</td>
<td>7.9</td>
<td></td>
</tr>
<tr>
<td>Hispanic ethnicity</td>
<td>11.1</td>
<td>16.2</td>
<td>&lt;.001</td>
<td>9.5</td>
<td>16.4</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Medicare insurance</td>
<td>36.8</td>
<td>59.8</td>
<td>&lt;.001</td>
<td>29.9</td>
<td>60.6</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Medicaid insurance</td>
<td>92.0</td>
<td>89.5</td>
<td>&lt;.001</td>
<td>95.2</td>
<td>89.4</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Admit to acute hospital (vs LTAC)*</td>
<td>87.2</td>
<td>96.8</td>
<td>&lt;.001</td>
<td>96.3</td>
<td>96.8</td>
<td>.08</td>
</tr>
<tr>
<td>Admission year</td>
<td></td>
<td></td>
<td>&lt;.001</td>
<td>&lt;.001</td>
<td></td>
<td>&lt;.001</td>
</tr>
<tr>
<td>2000</td>
<td>10.2</td>
<td>11.7</td>
<td></td>
<td>7.9</td>
<td>11.6</td>
<td></td>
</tr>
<tr>
<td>2001</td>
<td>10.6</td>
<td>12.2</td>
<td></td>
<td>9.1</td>
<td>12.2</td>
<td></td>
</tr>
<tr>
<td>2002</td>
<td>11.7</td>
<td>12.4</td>
<td></td>
<td>9.6</td>
<td>12.4</td>
<td></td>
</tr>
<tr>
<td>2003</td>
<td>10.8</td>
<td>12.9</td>
<td></td>
<td>10.9</td>
<td>12.9</td>
<td></td>
</tr>
<tr>
<td>2004</td>
<td>11.8</td>
<td>12.7</td>
<td></td>
<td>12.2</td>
<td>12.7</td>
<td></td>
</tr>
<tr>
<td>2005</td>
<td>13.5</td>
<td>12.7</td>
<td></td>
<td>16.3</td>
<td>12.7</td>
<td></td>
</tr>
<tr>
<td>2006</td>
<td>16.3</td>
<td>12.5</td>
<td></td>
<td>17.6</td>
<td>12.6</td>
<td></td>
</tr>
<tr>
<td>2007</td>
<td>15.1</td>
<td>12.9</td>
<td></td>
<td>16.4</td>
<td>12.9</td>
<td></td>
</tr>
<tr>
<td>Length of stay &gt;5 days</td>
<td>96.7</td>
<td>27.4</td>
<td>&lt;.001</td>
<td>56.2</td>
<td>26.8</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Surgery</td>
<td>42.3</td>
<td>32.2</td>
<td>&lt;.001</td>
<td>25.7</td>
<td>30.5</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Comorbidities</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Diabetes</td>
<td>29.6</td>
<td>16.9</td>
<td>&lt;.001</td>
<td>24.7</td>
<td>16.6</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Cancer</td>
<td>15.1</td>
<td>7.8</td>
<td>&lt;.001</td>
<td>13.5</td>
<td>7.5</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Dementia</td>
<td>5.9</td>
<td>2.9</td>
<td>&lt;.001</td>
<td>5.8</td>
<td>2.8</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Ulcer</td>
<td>4.6</td>
<td>1.8</td>
<td>&lt;.001</td>
<td>2.7</td>
<td>1.7</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>AIDS</td>
<td>0.6</td>
<td>0.2</td>
<td>&lt;.001</td>
<td>0.4</td>
<td>0.2</td>
<td>.02</td>
</tr>
<tr>
<td>High comorbidity index*</td>
<td>64.5</td>
<td>27.1</td>
<td>&lt;.001</td>
<td>52.8</td>
<td>26.0</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Admitted to high-volume hospital*</td>
<td>61.9</td>
<td>61.1</td>
<td>&lt;.001</td>
<td>72.2</td>
<td>67.2</td>
<td>&lt;.001</td>
</tr>
<tr>
<td>Admitted to hospital with high length of stay*</td>
<td>85.4</td>
<td>77.2</td>
<td>&lt;.001</td>
<td>71.5</td>
<td>77.1</td>
<td>&lt;.001</td>
</tr>
</tbody>
</table>

**Note.** Data other than N and P shown as percentage. AIDS, acquired immune deficiency syndrome.

* LTAC, long-term acute care facility.

b Surgery indicates surgery during the current admission or within the previous 30 days.

Comorbidity index measured by Romano score.

d High volume, >10,000 annual admissions.

e High length of stay, >5 days.

transferring or recently discharging hospitals may also improve accuracy of CDI rates.

For prevention, patient characteristics may be utilized to identify populations at elevated risk for postdischarge CDI. We found that risk factors for HO-CDI and PD-CDI were often the same, including increasing age, higher length of stay, and overall poor health, including diabetes, cancer, and AIDS, in agreement with prior studies. In addition, prevention may be targeted at patients with specific primary admission diagnoses such as septicemia, postoperative infection, and pneumonia. These primary admission diagnoses all represent conditions likely to be treated with antibiotics, the main risk factor for CDI. The immediate postdischarge period should be considered an extension of the risk of CDI that begins during a hospital stay. This heightens the importance of educating high-risk patients before hospital discharge about the potential for postdischarge diarrhea and of identifying prophylactic solutions to prevent disease in the high-risk patient population.

In addition, we found that white and non-Hispanic patients
had a higher risk of PD-CDI. While we did not evaluate reasons for this difference, racial and ethnic disparities, including access to health care, have been well documented and may be magnified in the outpatient arena.\textsuperscript{36,37} Differences in access to outpatient care are likely to impact antibiotic use in these groups and their subsequent risk of PD-CDI. In addition, male gender was associated with lower risk of PD-CDI but not with HO-CDI. This difference may not be due

\begin{table}
\centering
\caption{Multivariate Analysis of Predictors of Hospital-Onset \textit{Clostridium difficile} Infection (HO-CDI) and Postdischarge CDI (PD-CDI) Among All Adult Inpatients}
\begin{tabular}{lcccc}
\hline
 & \multicolumn{2}{c}{HO-CDI vs all non-HO-CDI} & \multicolumn{2}{c}{PD-CDI vs all non-HO-CDI} \\
 & OR (95\% CI) & \textit{P} value & OR (95\% CI) & \textit{P} value \\
\hline
Age, years & & & & \\
18–40 & Reference & & Reference & \\
40–49 & 1.93 (1.29–2.89) & <.001 & 2.20 (1.53–3.17) & <.001 \\
50–59 & 1.82 (1.23–2.71) & <.001 & 2.21 (1.54–3.17) & <.001 \\
60–75 & 1.83 (1.28–2.61) & <.001 & 2.76 (1.99–3.82) & <.001 \\
75+ & 2.16 (1.51–3.09) & <.001 & 3.93 (2.85–5.44) & <.001 \\
Male gender & & & & \\
White & Reference & & Reference & \\
Black & 0.94 (0.61–1.45) & .01 & 0.49 (0.31–0.78) & .001 \\
Asian & 1.27 (1.06–1.52) & <.001 & 0.64 (0.53–0.77) & <.001 \\
Other & 0.79 (0.59–1.05) & <.001 & 1.04 (0.85–1.28) & <.001 \\
Hispanic ethnicity & & & & \\
& 0.94 (0.77–1.15) & .56 & 0.73 (0.62–0.86) & <.001 \\
Medicare insurance & & & & \\
& 0.61 (0.48–0.77) & <.001 & 0.70 (0.56–0.87) & <.001 \\
Admission year & & & & \\
2000 (reference) & & & & \\
2001 & 1.03 (0.81–1.33) & <.001 & 0.92 (0.75–1.10) & <.001 \\
2002 & 1.12 (0.88–1.43) & <.001 & 1.04 (0.86–1.27) & <.001 \\
2003 & 0.94 (0.73–1.21) & <.001 & 1.14 (0.94–1.38) & <.001 \\
2004 & 1.22 (0.96–1.55) & <.001 & 1.35 (1.12–1.62) & <.001 \\
2005 & 1.25 (0.99–1.58) & <.001 & 1.67 (1.40–1.99) & <.001 \\
2006 & 1.55 (1.24–1.94) & <.001 & 1.83 (1.53–2.18) & <.001 \\
2007 & 1.39 (1.10–1.74) & <.001 & 1.71 (1.43–2.04) & <.001 \\
Length of stay & 1.02 (1.02–1.02) & <.001 & 1.01 (1.01–1.01) & <.001 \\
Surgerya & 2.16 (1.91–2.43) & <.001 & 1.23 (1.12–1.35) & <.001 \\
Comorbidities & & & & \\
Diabetes & 1.36 (1.20–1.53) & <.001 & 1.14 (1.03–1.25) & <.001 \\
Cancer & 1.32 (1.13–1.55) & <.001 & 1.24 (1.09–1.42) & <.001 \\
Dementia & 1.57 (1.28–1.93) & <.001 & 1.03 (0.87–1.21) & <.001 \\
Ulcer & 1.92 (1.49–2.47) & <.001 & 1.31 (1.04–1.67) & <.001 \\
AIDS & 4.15 (2.02–8.50) & <.001 & 3.27 (1.60–6.65) & <.001 \\
High comorbidity indexb & 1.06 (1.02–1.11) & <.001 & 1.02 (1.01–1.03) & <.001 \\
Primary admission diagnosis & & & & \\
Chemotherapy & 7.08 (4.04–12.42) & <.001 & ... & ... \\
Staphylococcus aureus pneumonia & 5.13 (3.23–8.16) & <.001 & ... & ... \\
Infection due to vascular device & 3.05 (1.87–4.97) & <.001 & ... & ... \\
Septicemia & 2.70 (3.71–1.96) & <.001 & 3.55 (2.87–4.40) & <.001 \\
Postoperative infection & 2.36 (3.89–1.43) & <.001 & 2.39 (1.51–3.76) & <.001 \\
Acute respiratory failure & 2.25 (1.63–3.10) & <.001 & ... & ... \\
Pneumonia & 1.47 (1.93–1.11) & <.001 & 1.82 (1.54–2.16) & <.001 \\
Cellulitis & ... & ... & 2.41 (1.72–3.38) & <.001 \\
Colon diverticulitis & ... & ... & 2.37 (1.64–3.43) & <.001 \\
Urinary tract infection & ... & ... & 1.86 (1.48–2.33) & <.001 \\
Acute renal failure & ... & ... & 1.73 (1.25–2.39) & <.001 \\
\hline
\end{tabular}

\textbf{Note.} CI, confidence interval; OR, odds ratio.
\textsuperscript{a} Surgery indicates surgery during the current admission or within the previous 30 days.
\textsuperscript{b} Comorbidity index measured by Romano score.
to an increased postdischarge risk for women but may instead reflect men’s reduced tendency to seek care and therefore be hospitalized for PD-CDI. More research is needed to understand the reasons for differential risk in these groups in order to develop effective prevention strategies for the postdischarge setting.

Our study has several limitations. We did not capture PD-CDI cases treated in the outpatient setting, which may have led to an underestimate of PD-CDI incidence. Nevertheless, the focus on PD-CDI associated with rehospitalization ensured capture of the most serious cases. While errors present in administrative data may lead to an under- or overestimation of CDI incidence, previous studies indicate reasonable agreement between medical records and ICD-9 codes for identification of total CDI burden.

Nevertheless, without POA codes, ICD-9 codes alone have been variably successful at distinguishing between community versus healthcare-associated C. difficile disease. We anticipate that the POA code, which has been in standard use in California for over a decade, improves this determination. In fact, POA codes have proven useful in other diseases (pneumonia, myocardial infarction) for distinguishing between hospital versus community-onset disease. We also minimized the chances that a code represented a history of CDI only versus active disease by limiting diagnoses to the first 3 coded positions. Another limitation is that we did not account for certain known risk factors (such as antibiotic use) that were unavailable in administrative data. However, we included primary admission diagnoses that are frequently treated with antibiotics. Finally, we attributed a PD-CDI event to the most recent hospitalization within 12 weeks. This definition does not account for multiple hospital exposures during that period or for intervening nursing home admissions, which may also contribute to CDI acquisition. Further, since the incubation period for CDI is unknown, some cases occurring within our 12-week window may be due to exposures in the community, including outpatient antibiotic use, household pets, and contamination of food. Nonetheless, over half of PD-CDI cases detected in this study occurred within the first 4 weeks after discharge, a time window that is accepted by national guidance to most likely reflect healthcare-associated CDI events.

In summary, tracking PD-CDI cases doubled the incidence of HA-CDI in a large county. Since the majority of hospitals do not track PD-CDI cases, the frequency and impact of PD-CDI may be widely underestimated, resulting in missed opportunities to prevent readmissions. Importantly for public reporting purposes, including PD-CDI affected individual hospitals differently, leading to substantial changes in hospital rankings by CDI incidence. Uniform tracking of PD-CDI events would allow more accurate estimates of overall CDI incidence and more equitable hospital-to-hospital comparisons. We found that the vast majority of cases can be captured if hospitals track PD-CDI cases that return to the same facility. We also identified several patient characteristics that were associated with PD-CDI, suggesting that preventative strategies may effectively focus upon specific patient groups. Targeted education and prevention for CDI may become increasingly important to help hospitals lower their readmission rates.

ACKNOWLEDGMENTS

We thank Leah Terpstra and Kristen Elkins for their contributions.

Financial support. This study was funded by the Centers for Disease Control and Prevention Prevention Epicenters Program (1U01 CI000344, Platt).

Potential conflicts of interest. E.R.D. reports that he is a consultant for Merck, Optimer, Pfizer, and Sanofi-Pasteur and has received research support from Optimer and Merck. Support for E.R.D. came from National Institute of Allergy and Infectious Diseases and the Centers for Disease Control and Prevention (1 K23AI065806). All other authors report no conflict of interest. All authors submitted the ICMJE Form for Disclosure of Potential Conflicts of Interest, and the conflicts that the editors consider relevant to this article are disclosed here.

Address correspondence to Courtney R. Murphy, MS, Health Policy Research Institute, 100 Theory Drive, Suite 110, Irvine, CA 92617 (courtner@uci.edu).

REFERENCES

2. Yokoe DS, Avery TR, Huang SS. Surgical Site Infection Surveillance following Total Hip and Knee Arthroplasty Using California Administrative Data. Oral presentation, Society for Healthcare Epidemiology of America; April 1–4, 2011; Dallas, TX.


