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Multimorbidity and Opioid Prescribing in Hospitalized Older Adults

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Abstract

Background: Multimorbidity and pain are both common among older adults, yet pain treatment strategies for older patients with multimorbidity have not been well characterized.

Objectives: To assess the prevalence and relationship between multimorbidity and opioid prescribing in hospitalized older medical patients with pain.

Methods: We collected demographic, morbidity, pain, and analgesic treatment data through structured review of the electronic medical records of a consecutive sample of 238 medical patients, aged \geq 65 years admitted between November 2014 and May 2015 with moderate-to-severe pain by numerical pain rating scale (range 4–10). We used the Cumulative Illness Rating Scale for Geriatrics (CIRS-G) to assess multimorbidity and cumulative illness burden. We examined the relationship between morbidity measures and opioid prescribing at hospital discharge using multivariate regression analysis.

Results: The mean age was 75 ± 8 years, 57% were female and 50% were non-White. Mean CIRS-G total score was 17 ± 6 , indicating high cumulative illness burden. Ninety-nine percent of patients had multimorbidity, defined as moderate-to-extremely severe morbidity in ≥ 2 organ systems. Sixty percent of patients received an opioid prescription at discharge. In multivariate analyses adjusted for age, race, and gender, patients with a discharge opioid prescription were significantly more likely to have higher cumulative illness burden and chronic pain.

Conclusion: Among older medical inpatients, multimorbidity was nearly universal, and patients with higher cumulative illness burden were more likely to receive a discharge opioid prescription. More studies of benefits and harms of analgesic treatments in older adults with multimorbidity are needed to guide clinical practice.

Keywords: hospitalization; illness burden; multimorbidity; older adults; opioids

Introduction

M ULTIMORBIDITY, defined as the presence of ≥ 2 comorbid illnesses, commonly afflicts community-dwelling older adults, involving about half of people aged 65–69 years and more than three quarters >85 years of age.^{1–3} Multimorbidity in older adults contributes to reduced functional status and lower quality of life, and increased inpatient days, rehospitalization, and mortality.^{4,5}

In community-dwelling older adults with multimorbidity, pain is among the most reported symptoms.⁷ For older adults living with pain,^{8–11} pharmacotherapy is the most common pain treatment they recieved.¹² However, the pres-

ence and severity of comorbid conditions affect analgesic safety in this population. For example, nonsteroidal antiinflammatory drugs (NSAIDs) incur greater risk in older adults with age- or disease-related declines in renal function.¹³ Studies of opioid-related adverse events in surgical inpatients suggest an association between older age, comorbidity severity, and opioid-related adverse events.^{14–16} Nevertheless, multimorbidity remains an understudied and under-recognized reality in pain treatment guidelines.² Despite the high prevalence of pain and multimorbidity in older adults, little is known about patterns of analgesic treatment for these patients, particularly in the medical inpatient setting.^{17,18}

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We undertook this study to describe the prevalence of multimorbidity and its relationship with opioid prescribing in hospitalized older adults with pain. We focused on opioid prescribing due to heightened national attention on opioid prescribing and the Centers for Disease Control and Prevention's (CDC) recommendation to prescribe opioids with caution in older adults.¹⁹ We hypothesized that patients with an opioid prescription at hospital discharge would have greater levels of morbidity than patients discharged without an opioid prescription.

Methods

Population and setting

We collected data from medical records of patients aged ≥ 65 years admitted to medicine services at an urban academic medical center from November 2014 to May 2015. The study methods and sampling criteria (Fig. 1) are reported elsewhere.²⁰ We included patients in the study if they had moderate-to-severe pain (pain score of ≥ 4 on a numerical pain rating scale [NPRS; 11 points scale] or ≥ 1 on the Checklist of Non-Verbal Pain Indicators [CNPI]^{21–23}) at admission (n=248) and did not die during hospitalization (n=10).

Trained research personnel collected data by structured chart abstraction from the electronic medical record (EMR). Ethical approval was obtained from the local institutional review board.

Independent measures

Demographic data included age at hospitalization, gender, race, ethnicity, marital status, insurance type, language preference, hospitalization length, and admission and discharge locations.

Pain. We abstracted admission and discharge pain scores—defined as NPRS, CNPI, and Verbal Descriptor

Scale (VDS)²⁴ pain scores documented within 24 hours of admission and 48 hours before discharge, respectively. To harmonize the three pain scales, all CNPI and VDS pain scores were converted to corresponding NPRS scores using a previously described algorithm.²⁰

Measures of morbidity. Multimorbidity has several features: (1) severity of the individual comorbid condition, (2) sum of high-severity comorbid conditions, and (3) combined severity of all comorbidity conditions resulting in overall illness burden. We chose the Cumulative Illness Rating Scale for Geriatrics (CIRS-G) to assess these three multimorbidity characteristics in our cohort.⁶ The scale assesses 14 body systems rated on a 5-point scale from 0 (no illness) to 4 (extremely severe, intractable illness). Although CIRS-G scores range from 0 to 56, very high scores are incompatible with life as they indicate multiorgan failure.⁵ The scale has good inter-rater reliability and validity in capturing overall illness burden for both inpatient and outpatient older adults.^{6,25}

Trained clinical personnel completed the CIRS-G scoring. The 1991 CIRS-G manual²⁶ was the main scoring reference due to its extensive validation^{6,27}; the 2008 CIRS manual⁵ was used to clarify ambiguous scenarios. Scorers conducted a structured assessment of each patient's admission and discharge notes. Scorers also reviewed the emergency department note, the last preceding primary care note, laboratory results, or keyword search results when needed. To ensure inter-rater reliability, the first 100 charts were double scored. All discrepancies were resolved in consultation with the senior author (C.S.R.).

We used various measures from the CIRS-G to indicate the three different aspects of multimorbidity. We used the *CIRS-G* score for each organ system to describe individual system-specific illness severity. We summed the number of organ systems with a score of ≥ 2 to assess multimorbidity severity.

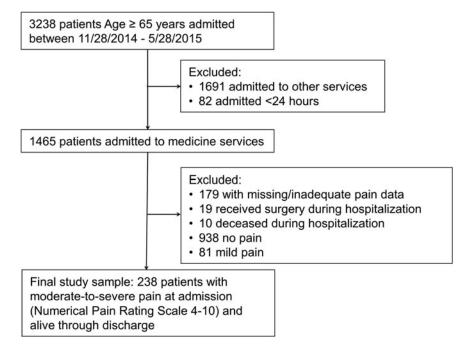


FIG. 1. Sampling flow chart of study participants.

We used the CIRS-G *total score* to characterize cumulative illness burden. We chose the term "burden" because it captures the overall impact of each comorbidity in a person, taking severity into account.²⁸

Outcome measures

Our major outcome of interest was opioid analgesic prescribing at hospital discharge. This variable included both new prescriptions resulting from the hospitalization and continued prescriptions to patients already prescribed an opioid before admission. Our secondary outcome assessed the receipt of higher dose opioids in the 24 hours before discharge. We converted all opioid doses to their morphine equivalent daily dose (MEDD), and we defined a *high dose* as MEDD \geq 50 mg based on the 2016 CDC guidelines.¹⁹

We also collected data on other analgesics (e.g., acetaminophen, NSAIDs, and adjuvants) prescribed at hospital discharge and on nonpharmacologic pain treatments (e.g., hot and cold application, repositioning) documented during hospitalization.

Statistical analysis

We compared differences in patients' characteristics with and without an opioid prescription at discharge using χ^2 tests for categorical variables and Student's *t*-tests for continuous variables. Multivariate logistic regression was used to estimate the odds ratio (OR) for characteristics associated with opioid prescription at discharge, and with high opioid dose. These models considered covariates of age, gender, race, and history of chronic pain. Analyses were conducted using SAS version 9.4 (SAS Institute, Inc., Cary, NC).

Results

Older adults admitted with moderate-to-severe pain (n=238) had a mean age of 74.9 ± 8.2 years, 56.7% were female, 49.6% were non-White, and 39.9% had a history of chronic pain (Table 1). Patients had a mean CIRS-G total score of 17.2 ± 5.5 , indicating high cumulative illness burden (Table 2).

Opioid prescriptions

Sixty percent (n=142) of patients received an opioid prescription at discharge (Table 1). Among patients receiving discharge opioids, 75.4% already had an opioid prescription before admission. In the bivariate analysis, patients were more likely to receive discharge opioid prescriptions if they had a chronic pain diagnosis (49.3% vs. 26.0%, p < 0.001), cancer diagnosis (44.4% vs. 22.9%, p < 0.001), English language preference (77.5% vs. 62.5%, p=0.012), and higher discharge pain scores (mean of 2.4 ± 2.7 vs. 1.4 ± 2.5 , p=0.002, Table 1). Among patients without a preadmission opioid prescription (n=118), the incident prescription rate was 14.7%.

Notably, opioid prescriptions were associated with prescription of other pharmacologic and nonpharmacologic pain treatments. Compared with patients not receiving opioids, those receiving discharge opioids were more likely to receive acetaminophen (69.7% vs. 33.3%, p < 0.001) and adjuvant analgesics (34.5% vs. 21.9%, p = 0.036) prescription at discharge and nonpharmacologic treatment order (56.3% vs. 35.4%, p = 0.002) during their hospitalization.

Among patients with discharge opioids (n=142), 35.2% received a high dose (MEDD \geq 50 mg) of opioids within 24 hours before discharge (Table 3). Compared with those

 TABLE 1. CHARACTERISTICS OF PATIENTS ADMITTED TO MEDICINE SERVICE WITH MODERATE-TO-SEVERE

 PAIN, OVERALL AND BY OPIOID PRESCRIPTION AT DISCHARGE

Characteristics, n (%) or mean±standard deviation	Overall (n=238)	Discharge opioid prescription		
		<i>No</i> (n=96)	<i>Yes</i> $(n = 142)$	р
Age in years	74.9 ± 8.2	75.5 ± 8.1	74.4 ± 8.3	0.30
Female	135 (56.7)	51 (53.1)	84 (59.2)	0.36
Race and ethnicity				
White	120 (50.4)	45 (46.9)	75 (52.8)	0.57
Asian	52 (21.9)	24 (25.0)	28 (19.7)	
Other	66 (27.7)	27 (28.1)	39 (27.5)	
Hispanic	26 (10.9)	15 (15.6)	11 (7.7)	0.06
Language and residence				
Preferred language of English	170 (71.4)	60 (62.5)	110 (77.5)	0.012
Admission from community	216 (90.8)	87 (90.6)	129 (90.8)	0.95
Discharge to community	147 (61.8)	61 (63.5)	86 (60.6)	0.64
Conditions in medical history				
Chronic pain	95 (39.9)	25 (26.0)	70 (49.3)	< 0.001
Depression	61 (25.6)	20 (20.8)	41 (28.9)	0.16
Cancer	85 (35.7)	22 (22.9)	63 (44.4)	< 0.001
Admission characteristics				
Admitted with existing opioid prescription	120 (50.4)	13 (13.5)	107 (75.4)	< 0.001
Opioid prescribed during admission	189 (79.4)	55 (29.1)	134 (70.9)	< 0.001
Hospital length of stay	5.7 ± 6.5	4.3 ± 4.0	6.6 ± 7.6	0.002
Admission pain score	6.3 ± 1.8	6.1 ± 1.8	6.5 ± 1.9	0.14
Discharge pain score	2.0 ± 2.7	1.4 ± 2.5	2.4 ± 2.7	0.002

Characteristics, n (%) or mean \pm standard deviation	Overall $(n=238)$	Discharge opioid prescription		
		<i>No</i> (n=96)	<i>Yes</i> (n = 142)	р
Cumulative illness burden (CIRS-G total score)	17.2 ± 5.5	15.8 ± 5.6	18.1 ± 5.3	0.002
System-specific illness severity (CIRS-G score by org	gan system, 0–4)			
Heart	1.36 ± 1.4	1.40 ± 1.3	1.33 ± 1.4	0.72
Vascular	1.92 ± 1.2	1.74 ± 1.1	2.04 ± 1.2	0.049
Hematologic	1.45 ± 1.2	1.24 ± 1.2	1.59 ± 1.2	0.027
Respiratory	1.50 ± 1.5	1.25 ± 1.4	1.66 ± 1.6	0.037
Eyes, ears, nose, and throat and larynx	0.82 ± 1.0	0.73 ± 1.0	0.87 ± 1.0	0.27
Upper gastrointestinal	1.24 ± 1.2	1.13 ± 1.2	1.31 ± 1.2	0.25
Lower gastrointestinal	1.13 ± 1.3	1.13 ± 1.2	1.12 ± 1.4	0.93
Liver	0.83 ± 1.2	0.95 ± 1.3	0.75 ± 1.2	0.21
Renal	0.76 ± 1.2	0.68 ± 1.1	0.81 ± 1.2	0.39
Genitourinary	0.96 ± 1.1	1.00 ± 1.1	0.93 ± 1.1	0.64
Musculoskeletal/integument	1.81 ± 1.2	1.35 ± 1.0	2.11 ± 1.2	< 0.001
Neurologic	0.95 ± 1.3	0.85 ± 1.3	1.01 ± 1.3	0.34
Endocrine/metabolic/breast	0.99 ± 1.1	0.95 ± 1.1	1.02 ± 1.2	0.62
Psychiatric	1.47 ± 1.3	1.42 ± 1.3	1.51 ± 1.3	0.57
Multimorbidity severity (number of organ systems wi	th CIRS-G score of \geq	2)		
Two or more systems	235 (98.7)	93 (96.9)	142 (100.0)	0.07
Three or more systems	226 (95.0)	88 (91.7)	138 (97.2)	0.07
Four or more systems	198 (83.2)	73 (76.0)	125 (88.0)	0.015

 TABLE 2. MEASURES OF MORBIDITY IN PATIENTS ADMITTED TO MEDICINE SERVICE WITH MODERATE-TO-SEVERE

 PAIN, OVERALL AND BY OPIOID PRESCRIPTION AT DISCHARGE

Each organ system scored 0-4 for illness severity: 0=none, 1=mild, 2=moderate, 3=severe, 4=extremely severe. CIRS-G, Cumulative Illness Rating Scale for Geriatrics.

receiving an MEDD <50 mg, those who received a high dose were younger (mean age 71.7 ± 7.4 vs. 75.9 ± 8.4 years, p=0.004) and were less likely to be discharged to the community (48.0% vs. 67.4%, p=0.024). Patients were more likely to receive a high dose if they had a history of chronic pain (64.0% vs. 41.3%, p=0.01), depression (42.0% vs. 21.7%, p=0.011), or metastatic cancer (36.0% vs. 15.2%, p=0.005) and had higher discharge pain scores (mean of 3.4 ± 2.7 vs. 1.9 ± 2.5 , p=0.002).

Multimorbidity and illness burden

Multimorbidity was nearly universal in this cohort: 99% of patients had moderate-to-extremely severe morbidity in ≥ 2 organ systems, 95% in ≥ 3 systems, and 83% in ≥ 4 systems (Table 2). The most common morbidities were vascular (affecting 72.7%), musculoskeletal (63.5%), psychiatric (52.1%), and upper gastrointestinal (47.9%) (data not shown). Although liver (24.0%) and renal (21.9%) systems had the lowest prevalence of morbidity, they still affected a large proportion of patients.

Relationship between measures of morbidity and opioid prescription

Cumulative illness burden was significantly higher in patients with discharge opioid prescriptions (CIRS-G total score 18.1 ± 5.3 vs. 15.8 ± 5.6 , p=0.002, Table 2). Multimorbidity *severity* was also significantly higher in the opioidprescribed group: 88% had ≥ 4 body systems with high levels of illness severity compared with 76% in those without a prescription (p=0.015). Patients with discharge opioid prescriptions had higher average scores for illness severity in vascular, hematopoietic, respiratory, and musculoskeletal systems (Table 2). Patients with a high opioid dose had significantly higher cumulative illness burden (CIRS-G total score 19.6 ± 5.0 vs. 17.2 ± 5.3 , p=0.01, Table 3).

In regression analysis, the odds of receiving discharge opioids was 1.6 times greater among individuals with chronic pain (OR 2.59, 95% confidence interval [CI] 1.45–4.61, p=0.001) and was also higher in patients with higher cumulative illness burden (with each point increase in CIRS-G total score, OR 1.08, 95% CI 1.03–1.14, p=0.003) after adjusting for age, gender, and race (Table 4). This association persisted even if cancer patients (n=85) were excluded from the analysis (OR 1.07, 95% CI 1.01–1.14, p=0.019). Furthermore, although the number of patients receiving a high MEDD dose was low (n=50), the odds of receiving a higher dose was similarly elevated with chronic pain and with incremental increases in CIRS-G total score (data not shown).

Discussion

Few studies have described opioid prescribing to hospitalized older adults in the context of multimorbidity.^{16,18,29,30} This is the first study to assess the relationship between comorbidity severity and burden and opioid use in hospitalized older adults. In this cohort of older medical inpatients with moderate-to-severe pain, most patients entered (50.4%) and left the hospital with an opioid prescription (59.7%). Multimorbidity was nearly universal and cumulative illness burden was strikingly high. Discharge opioid prescriptions were associated with higher cumulative illness burden and a history of chronic pain. The association with illness burden persisted even after adjustment for age, gender, race, and chronic pain, and after exclusion of cancer patients.

MULTIMORBIDITY AND OPIOIDS IN OLDER ADULTS

Change to minimize = p(0) or	Opioid Dose		
Characteristics, n (%) or mean \pm standard deviation	MEDD < 50 (n = 92)	$MEDD \ge 50 \ (n = 50)$	р
MEDD dose, mg	13.9±13.9, range 0–45	158.1±111.7, range 60–448	< 0.001
Age, years	75.9 ± 8.4	71.7 ± 7.4	0.004
65–74	44 (47.8)	36 (72.0)	0.006
75+	48 (52.2)	14 (28.0)	
Female	60 (65.2)	24 (48.0)	0.046
Race and ethnicity			
White	43 (46.7)	32 (64.0)	0.14
Asian	21 (22.8)	7 (14.0)	
Other	28 (30.4)	11 (22.0)	
Hispanic	7 (7.6)	4 (8.0)	0.93
Language and residence			
Preferred language of English	68 (73.9)	42 (84.0)	0.17
Admission from community	87 (94.6)	42 (84.0)	0.037
Discharge to community	62 (67.4)	24 (48.0)	0.024
Conditions in medical history			
Chronic pain	38 (41.3)	32 (64.0)	0.010
Depression	20 (21.7)	21 (42.0)	0.011
Cancer	36 (39.1)	27 (54.0)	0.09
Metastatic	14 (15.2)	18 (36.0)	0.005
Admission characteristics			
Hospital length of stay	6.6 ± 8.2	6.6 ± 6.4	0.98
Admission pain score	6.3 ± 1.8	6.9 ± 2.0	0.07
Discharge pain score	1.9 ± 2.5	3.4 ± 2.7	0.002
Measures of morbidity			
Cumulative illness burden (CIRS-G total score)	17.2 ± 5.3	19.6 ± 5.0	0.010
Multimorbidity severity (number of organ system	ns with CIRS-G score of ≥ 2) ^a		
Three or more systems	89 (96.7)	49 (98.0)	1.00
Four or more systems	80 (87.0)	45 (90.0)	0.59
Five or more systems	59 (64.1)	40 (80.0)	0.049

 TABLE 3. CHARACTERISTICS OF PATIENTS ADMITTED TO MEDICINE SERVICE WITH MODERATE-TO-SEVERE

 PAIN BY OPIOID DOSE PRESCRIBED DURING 24 HOURS BEFORE DISCHARGE

Each organ system scored 0-4 for illness severity: 0=none, 1=mild, 2=moderate, 3=severe, 4=extremely severe.

^aAll patients in both groups had CIRS-G score of ≥ 2 in two or more organ systems.

MEDD, morphine equivalent daily dose in milligrams per day.

These findings have a number of key implications. First, older patients are frequently prescribed opioids. Our findings align with several other studies examining opioid use in older adults. Higher rates of opioid prescriptions to older patients with musculoskeletal morbidity, chronic pain, and cancer in our study mirror a large-scale national study that found adults

 TABLE 4.
 MULTIVARIABLE ANALYSIS OF FACTORS

 Associated with Opioid Prescription at Discharge

	Discharge opioid prescription	d
Independent variable	Odds ratio (95% confidence interval)	р
Age (continuous) Female Race (White vs. non-White) Chronic pain Cumulative illness burden (CIRS-G total score)	0.98 (0.95-1.01) 1.28 (0.73-2.23) 1.25 (0.71-2.18) 2.59 (1.45-4.61) 1.08 (1.03-1.14)	$\begin{array}{c} 0.23 \\ 0.39 \\ 0.44 \\ 0.001 \\ 0.003 \end{array}$

Adjusted for age, gender, and race.

with primary discharge diagnoses of musculoskeletal injury, pain, and cancer were significantly more likely to receive opioids during their hospitalization.³¹ Recent studies of opioid prescribing in hospitalized older adults¹⁶ and Medicare beneficiaries³² also found associations between opioid prescribing and musculoskeletal morbidity. Second, our findings demonstrated overall high illness burden in these hospitalized older adults. Third, the presence of multimorbidity increased the risk of opioid prescribing. Given that multimorbidity also increases risk for opioid-related adverse events,14-16 higher illness burden may inform the relationship between opioids and adverse outcomes during and posthospitalization. Maiti et al.¹⁶ also found that older adults exposed to opioids before and during hospitalization had higher rates of poor outcomes (e.g., longer hospital stay, discharge to skilled nursing facility, and 30-day readmission) than nonexposed older adults. Maiti and coauthors¹⁶ did not find a difference in multimorbidity (assessed through the Charlson Comorbidity Index [CCI]³³) between older medical inpatients exposed and not exposed to opioids. This discrepancy could be explained by differences in study population, differences in the time points of our opioid measures, and by data element differences between the CIRS-G and CCI. However, the discrepancy may

reflect the importance of comorbidity severity in this relationship, which the CIRS-G captures more fully than the CCL.^{25,34}

The association between opioid prescriptions and higher illness burden has several potential explanations. Prescribing physicians may be concerned about the safety of nonopioid analgesics in older patients, particularly those with certain comorbidities such as renal and cardiovascular impairment or those at increased risk for gastrointestinal bleeding.^{12,35} For example, some adjuvant analgesics have anticholinergic effects, which can lead to falls or other adverse cognitive and physical outcomes in elderly.¹² Opioids may be perceived as safer for those with liver disease (in the case of acetaminophen), renal disease, or a history of gastrointestinal bleeding (in the case of nonsteroidal anti-inflammatory agents). We found that patients who received opioid prescriptions, however, were also more likely to be prescribed an array of analgesics orders for acetaminophen and adjuvant analgesics.

Little evidence exists to guide clinical decision making about opioid prescribing to older adults with pain. Systematic reviews of opioid efficacy and safety have found little data³⁶ and low-quality evidence³⁷ on outcomes in older adults, and have rarely compared opioids with other analgesics. Studies about the role of opioids with falls and fractures in older adults have reported conflicting results.^{38–42} Furthermore, underuse of opioids may carry risk for older adults with acute pain. In a study of delirium predictors in older adults after hip fracture, delirium incidence increased as opioid dose decreased (from >30 to 10–30 to <10 mg MEDD).²⁹

Evidence for optimal analgesic therapies for older adults is even more limited in the context of multimorbidity. A 2010 meta-analysis of opioid treatment outcomes in older adults found only six studies reporting results on participants aged \geq 65 years, and these excluded patients with "significant comorbidity."⁴³ Furthermore, studies of opioid therapy in older adults with specific comorbidities (e.g., dementia) have focused on postoperative pain, rather than acute or chronic pain.^{29,41,44}

The common co-occurrence of pain and multimorbidity in our cohort and other inpatient and community settings^{2,13,45} indicates that physicians regularly face the question of how to safely and effectively treat pain in older adults with multimorbidity. More evidence is needed to guide practice. Clinical research examining opioid efficacy and safety in such a complex population is likely to be challenging, in part, because of growing restrictions on opioid use. Care models such as the Program of All-Inclusive Care for the Elderly⁴⁶ and homebased primary care⁴⁷ that provide close follow-up of older adults with multimorbidity may be ideal settings for studying analgesic-related health outcomes. Such research will be increasingly important as U.S. population ages and as state and federal governments seek to limit opioid misuse and health care expenditure.

Limitations

This study has several limitations. First, we present data from a single urban academic medical center. However, given the extensive EMR review required to assess comorbidity severity, a large multicenter study on our topic is unlikely. In addition, morbidity measure accuracy could be affected by underdiagnosis of conditions such as cognitive impairment, or by retention of resolved illnesses in the EMR. We addressed the latter by thoroughly reviewing admission and discharge notes and medications to determine the presence and severity of ongoing illnesses.

Conclusion

Multimorbidity and high illness burden are common among older medical inpatients with pain. Higher comorbidity severity and illness burden is associated with opioid prescribing, and may contribute to posthospitalization adverse events in older adults given opioids. As the population of older adults with multimorbidity grows, additional studies will be critical to identify safer and more effective pain management strategies.

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Author Disclosure Statement

The authors have no conflicts of interest.

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