Predictors of Opioid Prescription Among a Sample of Patients with Acute Musculoskeletal Pain at a Tertiary Care Hospital in Saudi Arabia

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Background: Musculoskeletal pain is one of the most complex and debilitating types of pain. Although different pharmacologic treatments are available, very few studies have explored the predictors for opioid analgesics prescription to manage this type of pain.

Objective: The aim of this study was to explore the predictors for opioid prescription in patients with acute musculoskeletal pain in Saudi Arabia.

Methods: This was a single-center, retrospective chart review of adult patients (≥18 yrs.) with an acute nociceptive musculoskeletal pain at a university-affiliated medical center in Riyadh, Saudi Arabia. Cancer patients and those with chronic neuropathic pain were excluded. Patients' age, gender, number of comorbidities, duration of pain management, number of clinic visits for pain, and Numeric Pain Rating Scale (NPRS) scores at rest and with normal activities were collected. Multiple logistic regression was conducted to examine the relationship between the type of musculoskeletal pain and the prescription of opioid analgesics controlling for NPRS score on activity, age, gender, number of comorbidities, duration of pain treatment, and number of clinic visits for pain.

Results: The mean age of the 227 patients, who met the inclusion criteria, was 39 years and 68% of them were male. Sixty-three percent of the patients were prescribed opioid analgesics, and 61% of them had shoulder pain, 29% had back pain, and 10% had lower extremity pain (eg, hip, thigh, lower leg, knee, ankle, and foot pain). Tramadol was the most commonly prescribed opioid analgesic (82%), followed by codeine (13%). Ninety-seven percent of patients who were prescribed non-opioid analgesics had shoulder pain. Patients with shoulder pain had lower odds of receiving opioid analgesics (OR=0.019, P<0.0001, 95% CI=0.004-0.081) in comparison to their counterparts who had lower extremity or back pains. Moreover, the higher the pain score on activity was, the higher odds of receiving opioid analgesics (OR=1.317, P<0.0001, 95% CI=1.029-1.685).

Conclusion: Future studies should explore the impact of different opioid prescribing policies to improve the quality of patient care and reduce the unnecessary prescribing of opioids for patients with non-cancer musculoskeletal pain.

Keywords: analgesics, opioid, musculoskeletal pain, shoulder pain, Saudi Arabia

Introduction

Musculoskeletal disorders include more than 150 diagnoses affecting muscles, bones, joints and associated tissues, such as tendons and ligaments, as stated in the International Classification of Diseases. These conditions can occur anytime during people's lifetime resulting in persistent pain that negatively impact their quality of life. Musculoskeletal pain is one of the most commonly encountered pain

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in healthcare settings, and is considered the largest cause of disability in many countries, such as the United Kingdom.²⁻⁴ It was found to be one of the three major causes of the global burden of non-communicable diseases in 2016 representing about 16% of all disability adjusted life-years (DALYs).5 Moreover, it was reported that musculoskeletal pain and disorders account for 22.6% of the total costs of illnesses in Sweden. In the United Kingdom. the cost of opioid prescriptions for musculoskeletal pain management varied from £3 to £4,844 per patient per year, and the cost of opioid overprescription could exceed £100 million per year assuming that 40% of patients with musculoskeletal pain are overprescribed.² The cost of managing musculoskeletal pain is not limited to the cost of prescriptions or clinic visits; the productivity loss and absenteeism due to musculoskeletal pain is also significant. In Saudi Arabia, few research studies have explored the prevalence of musculoskeletal pain among the public. In a questionnaire-based study that explored the prevalence of musculoskeletal pain among 165 construction workers in the cities of Riyadh and Dammam, 48.5% of the respondents reported having musculoskeletal pain, such as low back and knee pain.8 Musculoskeletal pain was found to be also prevalent among medical professionals. In a study that explored the prevalence of musculoskeletal pain among medical and surgical residents at a tertiary care center in Riyadh, the majority of the respondents (82.5%) reported suffering from musculoskeletal pain at least once during their medical training, and the low back pain was the most prevalent (53%). Low back pain is also prevalent among dental practitioners according to multiple studies that explored the prevalence of musculoskeletal pain among dentists the Kingdom. 10-12

Although multiple non-pharmacologic treatments, such as physiotherapy and acupuncture, are used in the management of musculoskeletal pain, pharmacotherapy (eg, nonsteroidal anti-inflammatory drugs (NSAIDs), selective COX-2 inhibitors, oral and injectable corticosteroids, and opioids) remains the mainstay of musculoskeletal pain treatment. 13 Opioids are the most commonly prescribed analgesics used in the management of acute musculoskeletal pain. Nearly 90% of patients who visit pain clinics in the United Kingdom are prescribed some forms of opioid analgesics.² Although opioids are deemed to be the most effective analgesics in the management of acute musculoskeletal pain, their use is associated with higher risk of abuse, misuse, and serious adverse events. 14 Therefore, the lowest effective dose of opioids should be used for the shortest appropriate period of time whenever needed in the management of musculoskeletal pain. 15 Additionally, it was found that the early use of opioids in the management of both acute and chronic musculoskeletal pain has been associated poorer clinical outcomes especially in the case of low back pain according to a recently published systematic review of clinical practice guidelines. 16 However. another systematic review and meta-analysis has found that the use of opioids in the management of low back pain is associated with greater pain relief in the short term, but this effect was not observed in the long term. 17

There has been an upward trend in the opioids prescribing rate for both acute and chronic musculoskeletal pain globally over the last two decades particularly in the United States. 18,19 It is believed the rate of opioids prescription for patients with musculoskeletal pain has seen a 50% increase between the years of 2001 and 2010. 18 This was accompanied with a significant increase in the rates of opioid-related death. 20 Unfortunately, research studies aimed at exploring the rates of opioids consumption are scarce in the Middle East.²¹ Although the rates of opioids utilization in the management of pain states have significantly increased in the world over the last two decades, some Middle Eastern countries have seen either a modest increase or a decrease in the rates of opioids utilization. ^{21,22} This could be attributed to several reasons, such as poor palliative care services, lack of essential opioid stock, and strict regulations.²² Although no study has explored the pain management adequacy among patients with musculoskeletal disorders in Saudi Arabia, many believe that there is an underutilization of analgesics in general, and opioids in particular, in the management of musculoskeletal pain.²² This assumption is mainly based on the findings of previously published studies that assessed the pain management adequacy among patients with more severe disease states, such as cancer and cardiac surgery.^{23,24} Moreover, very few tertiary care hospitals in Saudi Arabia have pain management clinics, and those with such clinics require special forms to be filled for any opioid prescriptions by a medical specialist or consultant only. These regulations besides the local drug enforcement laws that require a narcotic pharmacist to be in charge of auditing all narcotic prescriptions in each public or private hospital as well as the fact that those who engage in any prescription drug diversion can be sentenced to death could explain the reason behind the underutilization of opioid analgesics in Saudi Arabia. 22,25

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Understanding the predictors for opioids prescription should help in formulating a safe and effective opioid prescribing protocol for patients with musculoskeletal pain.

Therefore, the aim of this study was to explore the predictors for opioids prescription among a sample of patients with acute musculoskeletal pain in Saudi Arabia.

Methods

Study Design, Setting, and Subjects

This was a single-center, retrospective medical chart review study that was conducted at King Khalid University hospital, which is a university-affiliated hospital in Riyadh, Saudi Arabia. The hospital's bed capacity is 1,200 beds, and it provides primary, secondary, and tertiary health services to university faculty and their families as well as to Saudi citizens on a referral basis. Although the hospital has recently established pain clinics for patients with chronic pain, patients with acute pain are treated with opioid or non-opioid analgesics based on physicians' discretion. Furthermore, there are no published pain management guidelines for physicians to follow or adhere to. However, only licensed medical specialists or consultants can prescribe opioids for legitimate medical indications.

Adult patients (≥18 years old) with an acute nociceptive musculoskeletal pain who have been seen in the orthopedic outpatient clinics were identified via the hospital electronic health record (EHR) system (eg, Cerner Millennium). Patients with cancer, neuropathic pain, prior use of opioids or substance abuse history, osteoarthritis, and those with other pain disorders that necessitate long-term pain management (eg, inflammatory bowel disease, fibromyalgia, post-trauma or post-surgical pain) were excluded. Systematic random sampling was used by choosing every 100th patient health record of all patients who were prescribed opioid and/or non-opioid analgesics for acute musculoskeletal pain between May 2015 and December 2017. Patients' age, gender, body mass index (BMI), number of clinic visits, pain location, number of comorbidities, number of prescription medications, duration of pain treatment which was written in the prescriptions for the different dispensed analgesics, and the Numeric Pain Rating Scale (NPRS) scores at rest and with normal activities were collected for both cases (patients on opioid analgesics) and controls (patients on non-opioid analgesics). The NPRS is an 11-point verbal

scale which has been validated to assess the pain intensity across multiple patient populations and in different health-care settings. ²⁶ The study was approved by the institutional review board of King Saud University College of Medicine (Research Project No. E-17-2587). No patient consent form was required per the institutional policy since the study was a retrospective chart review, and no personal identifiers were collected. Furthermore, the study adhered to the ethical principles of the declaration Helsinki with regard to data storage and handling. ²⁷

Statistical Analysis

Descriptive statistics using means, standard deviations, frequencies, and percentages were used to describe the patient characteristics. The baseline characteristics for the cases (patients on opioid analgesics) and controls (patients on non-opioid analgesics) were compared using Chisquare test, Fisher's exact test, and Student's t-test as appropriate. The pain location was dichotomized into shoulder pain versus the other types of pain. Multiple logistic regression analysis to identify the predictors for opioids prescription was conducted and included the following variables: age, gender, NPRS score on activity, number of comorbidities, number of clinic visits, duration of pain treatment, and pain location. 28 The minimum sample size for multiple logistic regression that included seven predictor variables for opioids prescription with at least 10 events (cases on opioids) per variable matched to controls (patients with musculoskeletal pain on non-opioid analgesics) based on a 1:1 ratio was estimated to be 140 patients.²⁹ All statistical analyses were conducted using SAS® version 9.4 (SAS institute, Carv. NC, USA).

Results

Out of 1,651 randomly selected patients who received prescription analgesics between May 2015 and December 2017, 227 patients met the inclusion criteria and were included in the study. Approximately 64% of the patients with acute musculoskeletal pain were prescribed opioid analgesics. The mean age of the patients was 39 years with no significant difference between the cases (patients on opioid analgesics) and controls (patients on non-opioid analgesics). Most of the patients were male (68.28%) with no significant difference in the percentages of males or females between the two groups. Most of the sampled patients had shoulder pain (60.79%), followed by back pain (29.07%), and lower extremity pain (eg, hip, thigh, lower leg, knee, ankle, and foot pain) (10.13%). The

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majority of the patients with shoulder pain (97.56%) were prescribed non-opioid analgesics (eg, nonsteroidal antiinflammatory drugs, celecoxib, and acetaminophen), whereas only two patients with back pain and none of the patients with lower extremity pain were prescribed non-opioid analgesics (P<0.0001). Approximately 21% of the patients who were prescribed opioid analgesics had more than one comorbidity, and 17.24% were taking more than two prescription medications in comparison to 3.66% and 4.88%, respectively, among their counterparts who were prescribed non-opioid analgesics (P < 0.05). The mean number of clinic visits for the patients was 1.72 visits with no significant difference between the two groups. Pain treatment mean duration was significantly higher among patients treated with opioid analgesics in comparison to their counterparts on non-opioid analgesics (19.14 days versus 10.44 days, P<0.0001). The mean NPRS scores with activity for patients who were prescribed opioid analgesics was slightly but significantly higher than their counterparts who were prescribed nonopioid analgesics (3.73 versus 3.06, P=0.0012). On the other hand, the mean NPRS scores at rest were not significantly different between the two groups. NO significant difference between the NPRS scores with activity and rest for the different pain locations (eg, lower limb, shoulder, and back pain) was found. The baseline characteristics of the patients are shown in Table 1. There were 158 opioid prescriptions dispensed for 145 patients who received

Table I Patients' Baseline Characteristics

Characteristics	Use of Opioid Analgesic		P-value	Total (n=227)
	No (n=82)	Yes (n=145)		
Age, years; mean ± SD	36.35±16.87	41.03±18.95	0.065	39.33±18.32
BMI, mean ± SD	27.83±5.44	28.39±6.59	0.510	28.19±6.19
Number of clinic visits, mean ± SD	1.43±2.59	1.89±3.10	0.246	1.72±2.93
Gender, n(%)				
Female	23(28.05)	49(33.79)	0.371	72(31.72)
Male	59(71.95)	96(66.21)		155(68.28)
Pain location, n(%)				
Lower extremity	0(0.00)	23(15.86)	<0.0001*	23(10.13)
Shoulder	80(97.56)	58(40.00)	<0.0001*	138(60.79)
Back pain	2(2.44)	64(44.14)	<0.0001*	66(29.07)
Pain severity on activity, mean ± SD				
Lower extremity	_	3.57±1.78	0.071	3.56±1.78
Shoulder	3.01±1.23	3.74±1.61		3.32±1.44
Back pain	5.00±0.0	3.78±2.02		3.82±1.99
Overall	3.06±1.25	3.73±1.81	0.0012*	3.48±1.66
Pain severity at rest, mean ± SD				
Lower extremity	_	2.217±1.57	0.359	2.22±1.57
Shoulder	2.31±1.24	2.71±1.60		2.48±1.41
Back pain	4.50±0.71	2.79±1.86		2.85±1.86
Overall	2.36±1.27	2.66±1.71	0.1313	2.56±1.57
Duration of pain treatment, mean ± SD	10.44±10.45	19.14±22.19	<0.0001*	15.99±19.25
Number of prescription medications, n(%)				
0–2	78(95.12)	120(82.76)	0.0069*	198(87.22)
>2	4(4.88)	25(17.24)		29(12.78)
Comorbidities, n(%)				
0–1	79(96.34)	114(78.62)	0.0002*	193(85.02)
>	3(3.66)	31(21.38)		34(14.98)

Notes: The percentages in each column were calculated based on the total number of patients for each arm of treatment (non-opioid vs opioid analgesics) and total sample size: The P-values were calculated using Chi-square, Fisher's exact test, and Student's t-test as appropriate: *P<0.05. Abbreviation: BMI, body mass index.

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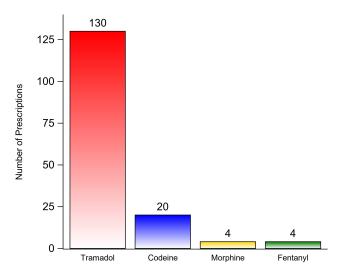


Figure 1 The type and number of opioid analgesic prescriptions for 145 patients.

opioid analgesics, and tramadol was the most commonly prescribed opioid (82.27%), followed by codeine (12.66%), morphine (2.53%), and fentanyl (2.53%) as shown in Figure 1. Patients with shoulder pain had 98.1% lower odds of receiving opioid analgesics in comparison to their counterparts with back or lower extremity pain (OR=0.019, 95% CI= 0.004-0.081, P<0.0001) controlling for age, NPRS score with activity, gender, number of comorbidities, duration of pain treatment, and number of clinic visits. On the other hand, patients with lower limb or back pain had more than 50 times higher odds of receiving opioid analgesics in comparison to their counterparts with shoulder pain (OR=53.59, 95% CI= 12.34--232.74, P<0.0001) controlling for age, NPRS score with activity, gender, number of comorbidities, duration of pain treatment, and number of clinic visits. Moreover, every additional point in the NPRS score with activity was associated with 31.7% increase in the odds of receiving opioid analgesics (OR=1.317, 95% CI= 1.029-1.685, P=0.0285) controlling for pain location, age, gender, number of comorbidities, duration of pain treatment, and number of clinic visits. The results of the multiple logistic regression analysis are shown in Table 2.

Discussion

The utilization of opioids in the management of chronic and acute pain has significantly increased worldwide especially in the western countries, such as the United States, where it largely contributed to the current opioid epidemic. However, there is a lack of studies that explored the predictors for opioids prescription in the

Table 2 The Odds of Opioids Prescription Among Patients with Musculoskeletal Pain

Variables	Odds Ratio (OR)	95% Confidence Interval (CI)	P-value
Shoulder pain	0.019	0.004-0.081	<0.0001*
Lower extremity or back pain	53.59	12.34-232.74	<0.0001*
NPRS score on activity	1.317	1.029-1.685	0.0285*
Age	0.995	0.972-1.019	0.6842
Female	0.745	0.325-1.708	0.4871
Number of comorbidities	1.345	0.769–2.354	0.2991
Duration of pain treatment	1.03	1.00-1.061	0.0534
Number of clinic visits	1.064	0.944–1.199	0.3076

Note: *P<0.05.

management of musculoskeletal pain in the Middle East where the medical practice and the regulations that govern the prescription of opioids are starkly different. 21,22 Thus, the findings of this study shed light on the prescribing pattern of opioid analgesics in the management of acute musculoskeletal pain, which is largely unknown in Saudi Arabia. Contrary to what was expected, opioids are not infrequently prescribed in the management of acute pain among the patients in the study. The vast majority of patients seen in the outpatient orthopedic clinics for acute back and lower extremity pain were prescribed opioid analgesics. Patients with lower extremity and back pain have more than 50 times higher odds of receiving opioid analgesics in comparison to their counterparts with shoulder pain despite controlling for a multitude of factors that are known to influence opioids prescription.²⁸ The high utilization rates of opioids in the management of lower extremity and back pain observed in this study are consistent with the reported high opioids utilization rates in the literature. 18-20,30 The use of opioids in the management of acute back and lower extremity pain for short term (eg, <90 days) result in greater pain relief according to a moderate level of evidence. 17 However, the results of recently published systematic reviews are against the use of opioids for acute and chronic musculoskeletal pain conditions. 16 Moreover, the use of opioids in the management of acute and chronic low back pain have not shown to be superior when compared to other non-opioid analgesics.³¹

On the other hand, shoulder pain was associated with significantly lower odds of opioid prescription. The reasons behind that can be attributable to the increasing evidence that discourage the use of opioids in the

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management of shoulder pain. 32,33 In a retrospective cohort study that examined the impact of opioids use on clinical outcomes among patients who underwent arthroscopic rotator cuff repair, patients who were treated with opioids pre-operatively did not reach the same level of functionality and required higher dosages of opioids postoperatively.³³ Additionally, patients with upper extremity pain are believed to require shorter period of opioids treatment in comparison to patients with other types of musculoskeletal pain.³⁴ However, the use of opioids in the management of other types of musculoskeletal pain, such as back pain, is not encouraged either.³¹ Therefore, it is unclear why patients with back or lower extremity pain were more likely to be treated with opioid analgesics in comparison to their counterparts with shoulder pain despite the fact that the pain severity, as measured by the NPRS, was controlled for. This can be partly explained by the fact that physicians have different beliefs about the consequences of their prescribing decisions, emotionally charged physician-patient interactions, and patients' and physicians' characteristics, and preferences. 35,36 The severity of pain, as measured by the NPRS, was positively associated with opioids prescribing, which is consistent with the preponderance of evidence.²⁸ However, female gender, age, duration of pain treatment, and number of clinic visits were not associated with opioid prescribing among the sample of patients in this study as shown in the adjusted odds ratios. On another note, tramadol was the most commonly prescribed opioid analgesics. This can be due to the beliefs among some physicians that tramadol is a safe and "opioid-lite". 37 However, the use of tramadol for acute pain is associated with high risk of misuse, adverse effects, and chronic use after an acute pain episode.38

Although this is the first study to the best of our knowledge that explored the predictors for opioids prescribing in Saudi Arabia, it has multiple limitations that must be acknowledged. First, this was a retrospective chart review study that is fraught with information bias, such as documentation and recall bias.³⁹ In addition, this was a single-center study with limited generalizability.⁴⁰ Furthermore, important variables, such as patient education level, prescribers' characteristics (eg, education, years of experience, previous history of prescribing opioids), patient satisfaction with care, and patients' perceptions about the adequacy of pain management, have not been collected and were not controlled for.²⁸ Moreover, not all patients with acute musculoskeletal

pain were included in this study although systematic random sampling was used. Despite all of the abovementioned limitations, this study highlights the gaps in knowledge about opioids prescribing for acute musculoskeletal pain in Saudi Arabia. Although it is largely believed that pain management is inadequate for some patient populations, such as cancer, this might not be the case for patients with acute musculoskeletal pain.²⁴ Future studies should examine the appropriateness and predictors of opioids prescribing in Saudi Arabia for patients with acute and chronic pain at a national level in order to formulate a national policy for opioids prescribing that ensures safe and effective management of pain.

Abbreviations

NPRS, Numeric Pain Rating Scale; DALYs, disability adjusted life-years; EHR, electronic health record.

Data Sharing Statement

Study data are available from the corresponding author (Yazed AlRuthia) upon request.

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Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

Disclosure

The authors report no conflicts of interest in this work.

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