



Drivers of in-hospital opioid consumption in patients undergoing lumbar fusion surgery

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Background: With the current opioid crisis, as many as 38% of patients are still on opioids one year after elective spine surgery. Identifying drivers of in-hospital opioid consumption may decrease subsequent opioid dependence. We aimed to identify the drivers of in-hospital opioid consumption in patients undergoing 1–2-level instrumented lumbar fusions.

Methods: This is a retrospective cohort study. Electronic medical record analysts identified consecutive patients undergoing 1–2 level instrumented lumbar fusions for degenerative lumbar conditions from 2016 to 2018 from a single-center hospital administrative database. Oral, intravenous, and transdermal opioid dose administrations were converted to morphine milligram equivalents (MME). Linear regression analysis was used to determine associations between postoperative day (POD) 4 cumulative in-hospital MMEs and the patients' baseline characteristics including body mass index (BMI), race, American Society of Anesthesiologists (ASA) grade, smoking status, marital status, insurance type, zip code, number of fused levels, approach and preoperative opioid use.

Results: A total of 1,502 patients were included. The mean cumulative MMEs at POD 4 was 251.5. Linear regression analysis yielded four drivers including younger age, preoperative opioid use, current smokers and more levels fused. There were no associations with surgical approach, zip code, ASA grade, marital status, BMI, race or insurance type.

Conclusions: Use of preoperative opioids and smoking are modifiable risk factors for higher in-hospital opioid consumption and can be targets for intervention prior to surgery in order to decrease in-hospital opioid use.

Keywords: In-hospital opioid consumption; driver; lumbar fusion surgery; degenerative lumbar spine disease; opioid crisis; opioid dependence; chronic opioid use

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Introduction

It is obvious that the United States is amidst an opioid crisis. This epidemic of opioid use, misuse, and abuse had become a critical public health issue. Americans, comprising only 4.6% of the world's population, consume 80% of the global opioid supply (1). More than 2 million people are addicted to prescription opioids (2). Drug overdose is the leading cause of accidental death in the US, with more than 20,000 overdose death related to prescription pain relievers in 2015 (3). To help address this desperate public health issue, surgeons must make their best effort to decrease their role in this epidemic, as surgery is associated with an increased risk of chronic postoperative opioid use (4-7).

Prevalence of opioid dependence after spine surgery is high. As much as 38% of patients undergoing major spine surgery are still on opioids one year after surgery (8). A higher rate of preoperative opioid use in patients with spinal diseases may contribute to postoperative opioid dependence (8-10). However, even patients without preoperative opioid use have an increased risk of subsequent chronic opioid dependence in the postoperative period (4,5). The later statistic is what should drive us as surgeons to be educated and thoughtful in how we administer opioids to our patients.

Identifying the drivers of in-hospital opioid consumption can be useful to potentially decrease the incidence of postoperative opioid dependence as in-hospital opioid administration can be a modifiable factor. The aim of this study was to identify the drivers of in-hospital opioid consumption in patients undergoing 1–2-level instrumented lumbar fusions.

We present the following article in accordance with the STROBE reporting checklist (available at <http://dx.doi.org/10.21037/jss-20-626>).

Methods

This is a retrospective cohort study at a single institution. Electronic Medical Record analysis identified consecutive patients who underwent a one- or two-level instrumented lumbar fusions for degenerative lumbar conditions from 2016 to 2018 using the hospital administrative database. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by institutional ethics board of University of Louisville/Norton Healthcare (# 18.1197) and individual consent for this retrospective analysis was waived.

Postoperative opioids were administered by nurses based on doctors' PRN order and pain severity. In-hospital, total daily opioid consumption, including oral, intravenous, or transdermal administration was calculated and converted to the morphine milligram equivalents (MMEs) using MME conversion factors (*Table 1*). In brief, MMEs of each opioid was calculated based on the following formula.

$$\text{MMEs} = \text{total dose (mg)} \times \text{MME conversion factor} \quad [1]$$

Then, MMEs of used opioids were added.

Linear regression analysis was used to determine associations between postoperative day (POD) cumulative in-hospital MMEs and the patients' baseline characteristics including body mass index (BMI), race, American Society of Anesthesiologists (ASA) grade, smoking status, marital status, preoperative daily opioid use, insurance type, zip code, number of fused levels, approach and preoperative opioid use. Preoperative opioid use including type of opioid, dose and frequency, was asked before surgery. Zip codes were classified into medically "underserved" or not based on government designation; areas with a high prevalence of health conditions in combination with a higher than average poverty rate and less access to healthcare, were designated as "underserved". Cumulative MME's up to POD 4 were used as the majority patients were discharged by POD 5.

Statistical analysis

A multivariate linear regression analysis was performed to identify risk factors for in-hospital opioid consumption. All statistical analyses were performed using SPSS Statistics 25 (IBM Corp., Armonk, NY). A statistical significance was defined as P value <0.05.

Results

A total of 1,502 patients, 601 (40%) male, mean age of 57.5 years, were included. Patients' demographic data are shown in *Table 2*. The mean BMI was 31.6, and patients were predominantly Caucasian. 39.3% were current smokers, while former smokers comprised 44.2%; 26.5% lived in underserved zip codes. The majority of patients (77.6%) underwent single level fusion. Anterior lumbar interbody fusion (ALIF) was done in 13.1% while combined antero-posterior procedure was done in 12.5%. Only 163 (11%) reported active daily opioid use prior to surgery. Total cumulative MMEs are shown in *Table 3*. Cumulative MMEs reached plateau at POD 4 being 251.5 ± 203.7 (*Figure 1*).

Table 1 MME conversion factors of commonly used opioids

Factors	Numbers
Tramadol	0.1
Morphine	1
Hydrocodone	1
Oxycodone	1.5
Hydromorphone (Dilaudid)	4

MME, morphine milligram equivalent.

Table 2 Demographic data of the 1,502 patients in the study

Variables	Numbers
Mean age (years), mean \pm SD	57.5 \pm 12.9
Male (%)	40.0
BMI, mean \pm SD	31.6 \pm 6.6
Race (%)	
Caucasian	87.0
Asian	0.1
Hispanic	0.6
African American	9.8
Missing	2.5
ASA (%)	
1	0.6
2	24.1
3	65.3
4	2.0
5	0.1
Missing	7.9
Smoking status (%)	
Non-smoker	16.4
Former smoker	44.2
Current smoker	39.3
Marital status (%)	
Married	61.2
Single	18.0
Divorced	5.5
Widowed	12.8

Table 2 (continued)**Table 2** (continued)

Variables	Numbers
Insurance type (%)	
Government	63.8
Private	36.2
Underserved ZIP code (%)	26.5
Number of fused levels (%)	
1	77.6
2	22.4
Surgical approach (%)	
PLF	33.8
ALIF	13.1
TLIF/PLIF/XLIF/DLIF	40.6
Combined antero-posterior	12.5
Length of stay (days), mean \pm SD	3.5 \pm 2.7
Preop daily opioid use (%)	10.9

BMI, body mass index; ASA, American Society of Anesthesiologists score; PLF, posterior lumbar fusion; ALIF, anterior lumbar interbody fusion; TLIF, transforaminal lumbar interbody fusion; PLIF, posterior lumbar interbody fusion; XLIF, extreme lateral interbody fusion; DLIF, direct lateral interbody fusion.

Younger age, preoperative opioid use, current smokers and more levels fused were associated with greater cumulative in-hospital MMEs with the R square of 0.079 (Table 4). There were no associations with surgical approach, zip code, ASA grade, marital status, BMI, race or insurance type.

Discussion

Amid the current opioid epidemic, it is imperative for surgeons to identify high risk patients preoperatively, as part of an effort to minimize chronic opioid use after surgery. Additionally, it would be wise from our specialty perspective, to take ownership of, or at minimum to reduce surgeon contribution to this atrocious public health problem. A number of risk factors for postoperative chronic opioid use have been reported, most commonly and obvious is preoperative opioid use, followed by younger age, depression, substance use disorder, preoperative pain

Table 3 In-hospital opioid consumption

Variables	Numbers (mean \pm SD)
MME on the day of surgery	60.7 \pm 43.5
Cumulative POD 1 MME	140.0 \pm 96.0
Cumulative POD 2 MME	199.6 \pm 144.6
Cumulative POD 3 MME	233.9 \pm 179.2
Cumulative POD 4 MME	251.5 \pm 203.7
Cumulative POD 5 MME	260.2 \pm 221.3
Cumulative POD 6 MME	265.2 \pm 234.8
Cumulative POD 7 MME	268.3 \pm 249.4

POD, postoperative day; MME, morphine milligram equivalent.

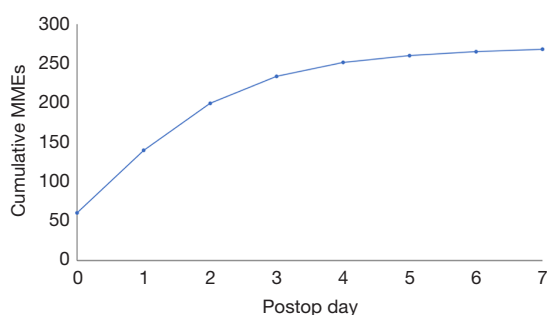


Figure 1 Postoperative in-hospital opioid consumption. Cumulative morphine milligram equivalents (MMEs) were increased postoperatively and reached plateau on the postop day 4.

Table 4 Multivariate linear regression analysis of the variables for postoperative cumulative MMEs

Variables	Standardized coefficients	P value
Age	-0.245	<0.001
Preoperative opioid use	0.102	<0.001
Smoking status	-0.065	0.015
Number of fused levels	0.065	0.016

MME, morphine milligram equivalent.

conditions, and smoking (8-13). These studies did not focus on in-hospital opioid administration even though it may well be critical, in particular for opioid naïve patients (14,15). Ge *et al.* showed that the amount of in-hospital opioid consumption following transforaminal lumbar interbody fusion (TLIF) is associated with postoperative

chronic opioid dependence in patients with and without preoperative opioid use. Compared with patients receiving 250–500 in-hospital MMEs, those receiving <250 in-hospital MMEs had a 3.73 times lower probability of requiring opioids at 6 months follow-up whereas those receiving >500 in-hospital MMEs had a 4.84 times greater probability of requiring opioids at 6 months (16). This study also demonstrated more than 500 MMEs during admission is a risk factor for postoperative chronic opioid use (16).

Of the four identified factors associated with in-hospital opioid consumption in this study, preoperative opioid use and smoking are potentially modifiable. There are numerous reports examining the association between preoperative opioid use and postoperative chronic opioid dependence (6,8,10-12). A recent systematic review revealed the incidence of persistent postoperative opioid use, whose definition was variable depending on studies (90-day to 1-year postop), ranges from 35% to 77% for patients with preoperative opioid use and from 0.6% to 26% for opioid-naïve patients in major surgical procedures (11). In addition, duration of preoperative opioid use was the most important predictor of continued use following lumbar spine surgery (13). Considering these findings, it is optimal to minimize preoperative opioid exposure, and to spend time educating patients about the potential downstream undesired affects. In our cohort, only 11% of patients were using daily opioids preoperatively. However, some studies have reported that up to 72.1% of patients presenting for major spine surgery were chronically using opioids before surgery (8). Preoperative opioid use was still risk factor for postoperative chronic opioid use in this study (8) despite this big discrepancy in the rate of preoperative opioid use might affect the results of statistical analysis. Thus, weaning patients off opioids prior to surgery may decrease in-hospital consumption and subsequently decrease chronic opioid use.

We also identified smoking as another modifiable factor. Multiple studies have showed the association between smoking and postoperative opioid dependence (6,11,17). Smoking has negative impact on all aspects of surgical treatment in spine surgery, including patient-reported outcomes, infection rate, pseudarthrosis, reoperation rate and complications (18-21). Literature supports smoking cessation as an effective tool in mitigating negative outcomes in spine surgery (21). The current study provides another reason for advocating smoking cessation prior to lumbar fusion surgery.

We also identified two risk factors, younger age and more levels fused, that are more difficult to be modified. Debate exists in terms of the impact of age on postoperative opioid dependence. Sharma *et al.* investigated risk factors associated with opioid dependence in 10,708 patients undergoing surgery for degenerative spondylolisthesis, in which younger age was an independent predictor of opioid dependence following surgery (12). On the other hand, Sun *et al.* examined risk factors associated with opioid dependence in patients undergoing major surgery, and identified age older than 50 years was associated with chronic opioid use with odds ratio of 1.74 (5). Interestingly, Kalakoti *et al.* reported that younger patients had a lower likelihood of opioid use following PLIF or TLIF whereas younger age was associated with higher likelihood of opioid use following posterior lumbar fusion (PLF) (9). However, the majority of previous studies did not find an association between age and chronic opioid dependence (8,10). We found only one study focusing on in-hospital opioid consumption (16). This study did not show a difference between in-hospital opioid consumption among patients of different age, but showed patients taking preoperative opioids were significantly younger than opioid naïve patients. Further study is necessary to conclude for the impact of age.

With regards to more levels requiring fusion or surgical invasiveness, a more invasive procedure intuitively may require more opioids. Modern technology allows for more complex spine procedures and spine surgery has become more invasive. Opioids are indispensable in perioperative pain management in spine surgery. Opioids are a necessary component of postoperative pain control, but efforts should be made to limit the amount. To decrease perioperative opioid consumption, multi-modal pain control (MMPC) approach has been developed (22). MMPC uses multiple agents that target several different pathways and mediators involved in nociception to improve analgesic effect (22,23). MMPC has been reported to be associated with less postoperative pain and opioid consumption (24-29). MMPC may be a good option to minimize in-hospital opioid consumption and subsequent chronic opioid dependence. Opioid free anesthesia (OFA) techniques have been developed which even further reduces in hospital exposure to opioids (30). Whether it is exposure, dosage, or duration of opioids, in theory they all can contribute to a long-term dependence and if possible, we should try to minimize our opioid consumption.

There are several limitations in this study. First, this is a

retrospective study in a single institution, making external validity unclear. Second, we did not evaluate chronic opioid dependence after discharge. Partly because of the inherent flaws in studying out of hospital opioid consumption. Typically, this is self-reported and the validity of actual usage versus other alternatives cannot be verified. Also, preoperative opioid use was based on patients' report and might have recall bias. In turn, in-hospital consumption is monitored strictly. Further studies are warranted to see the impact of in-hospital opioid consumption on the transition to chronic opioid dependence (14,15).

In conclusion, use of opioids prior to surgery and smoking are modifiable risk factors for higher in-hospital opioid consumption and can be targets for intervention prior to surgery in order to decrease in-hospital opioid use. Additionally, although levels to be fused or surgical invasiveness and age are not necessarily modifiable, they are still identifiable risk factors. When counselling patients on appropriate expectations prior to undergoing lumbar fusion surgery, it potentially beneficial to identify high-risk patients and counsel them on opioid consumption.

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Footnote

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by institutional

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