INTRODUCTION
The increase in the consumption of opioids has contributed to a subsequent increase in opioid use disorder and the complications thereof, both with prescribed medical use as well as illicit use. Therefore, adequately managing postoperative pain with the judicious use of opioids has become an area of active research. Numerous studies have assessed the prevalence and risk factors of opioid use for extended periods of time (often termed “prolonged opioid use”).

Opioid prescriptions around surgical intervention present an opportunity for opioid dependency to develop or worsen. For example, patients who already filled opioid prescriptions before surgical intervention are at increased risk for additional prescription requests after surgery. Similarly, Katzman et al reported that patients with intermittent opioid use before surgery were more likely to have persistent opioid use postoperatively. These findings are concerning considering the clear association between longer-term opioid use and the development of tolerance and addiction.

Currently, there is no clear consensus in the literature as to what constitutes prolonged opioid use in surgical

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patients, and current attempts at defining prolonged opioid use are flawed. The Center for Disease Control and Prevention (CDC) defines long-term opioid therapy as “use of opioids on most days for >3 months.” This definition does not account for patient populations naïve to opioid therapy, or those undergoing minor procedures with minimal postoperative pain.

Although hand surgeons often perform minor surgical procedures, prolonged opioid use and over-prescription of opioids continue to be a problem. For example, one recent study found that over 75% of patients undergoing carpal or cubital tunnel release fill a perioperative opioid prescription, and 13% of these patients continued to fill prescriptions 90–180 days after surgery. Despite the over-use and over-prescription of opioids in hand surgery, only one study has evaluated the rates and risk factors of prolonged use in postoperative hand surgical patients, and no study has attempted to define and standardize appropriate timeframes for prolonged postoperative opioid use by procedure.

To ensure that definitions of prolonged opioid use are practically useful in a hand surgical practice, this study has three aims: (1) to analyze and compare the definitions of prolonged opioid use among surgical specialties; (2) to investigate the relationship between the definition and prevalence of prolonged opioid use; and (3) to address shortcomings of current definitions by providing new criteria for prolonged opioid use following hand surgical procedures.

**METHODS**

This systematic review was conducted in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines. A literature search was performed for all published clinical studies on opioid use after surgery within the Embase, Web of Science, and PubMed databases on June 10, 2020. Our search strategy included all English articles published in the years 1990 through 2020 (see table, Supplemental Digital Content 1, which details search strategy in PubMed, Web of Science, and Embase. http://links.lww.com/PRSGO/B984). Since this systematic review does not involve interactions with human subjects, it is exempt from IRB review.

**Eligibility Criteria**

The inclusion criteria required studies to have at least one explicit definition of prolonged opioid use, to include patients undergoing surgery, and to include patients receiving opioids postoperatively. We included all studies using terms related to prolonged opioid use including “chronic,” “long-term,” “sustained,” “persistent,” and others. We excluded all animal studies, studies that only evaluated preoperative opioids, and studies with secondary surgery within the opioid use evaluation time if the second procedure was unaccounted for.

**Study Selection and Data Collection**

The review process was conducted independently by two authors (S.N. and S.M.), whereas a third author (Y.H.) mediated discrepancies. The initial search returned 5085 clinical studies on postoperative opioid use. Eight hundred duplicates, 16 non-English articles, and 61 articles outside the date range were removed. All remaining 4208 titles and abstracts were manually screened to determine eligibility. After the initial screening, 339 articles met the preliminary eligibility criteria. Of these, 209 articles were excluded during full-text analysis, leaving a total of 130 eligible for data collection. The data was manually analyzed and extracted in a standardized fashion (Fig. 1). The references of the included studies were cross-checked for additional studies but none met the inclusion criteria. No formal assessment of the potential bias among the included studies was performed.

**Outcome Measures**

The primary outcome measure was the definition of prolonged opioid use. These definitions were categorized based on the timeframe used in each study to distinguish between “regular” and “prolonged” (or any equivalent term) durations of opioid use; no other factors were used in categorization to avoid overspecification. Other collected variables included the prevalence of prolonged opioid use (as a percentage of the population at-risk) and mean or median patient age (depending on what was used by the study). Although multiple tools for synthesis of means and medians into an overall mean have been published, all of these have their limitations and, therefore, we decided not to use them. We treated age as a descriptive rather than an explanatory variable and categorized the means and medians into decades ranging from “less than 20 years” to “80 years or more.” We collected risk factors for prolonged opioid use and determined whether or not oral morphine equivalents (OMEs) were reported.

The publications were also grouped by “surgical specialty,” placing studies into categories based on the specialty that predominantly performs the procedures studied and the scope of the journal in which the publication was published. For example, a study of patients that underwent total knee arthroplasty would be categorized as “orthopedic.” Included surgical specialties were orthopedic, spine, general, surgical oncology, urologic, gynecologic, cardiac, plastic, otolaryngology, pediatric, and mixed for studies with procedures spanning multiple surgical specialties. The publications were also grouped by “surgical specialty,” placing studies into categories based on the specialty that predominantly performs the procedures studied and the scope of the journal in which the publication was published. For example, a study of patients that underwent total knee arthroplasty would be categorized as “orthopedic.” Included surgical specialties were orthopedic, spine, general, surgical oncology, urologic, gynecologic, cardiac, plastic, otolaryngology, pediatric, and mixed for studies with procedures spanning multiple surgical specialties.

**Takeaways**

**Question:** Which definitions are used to define prolonged postoperative opioid use in surgical research?

**Findings:** While many surgical specialties maintain similar definitions of prolonged opioid use in research studies, these do not seem to take into account the intensity and duration of pain after various surgical procedures. Therefore, these definitions may not correspond to what is considered prolonged use in clinical care. We propose definitions to be used during the postoperative use of opioids following hand surgery.

**Meaning:** Consensus on what constitutes prolonged postoperative opioid use may help physicians and researchers to determine its prevalence among patients that undergo hand surgery.
specialties. Spine surgery was considered separately from other orthopedic surgical subspecialties due to the fundamental clinical differences in surgical procedures and subsequent rehabilitation.

Patient cohorts were also categorized by their preoperative opioid use, the definition of which may differ between individual studies. Cohorts could be categorized as naïve (if patients used no opioids in the preoperative period), nonnaive (if patients used opioids preoperatively), combined (when both naïve and nonnaive patients were included in a single study), or unknown (if a study did not specify preoperative opioid use).

The prevalence of prolonged opioid use was separately presented as means and SDs for different definitions and...
different surgical specialties. Furthermore, the mean number of months used to define prolonged opioid use was presented for each surgical specialty; all definitions that were not time-based were excluded from this analysis. Last, we performed a Spearman’s correlation analysis to evaluate the relationship between definition timeframes and prevalence of prolonged opioid use.

RESULTS

General Characteristics

A total of 130 studies were included for data extraction (see table, Supplemental Digital Content 2, which details overview of collected data, http://links.lww.com/PRSGO/B985; see table, Supplemental Digital Content 3, which lists the studies that were included as part of the systematic review, http://links.lww.com/PRSGO/B986). Among the 83 studies that reported a mean or median age, a cohort age between 50 and 59.9 years was the most common and found in 28 studies (33.7%). The range of age groups included patients from less than 20 (n = 2) to “80 or more” (n = 1) years old. Of the 130 included studies, 47 did not report a mean or median age (Fig. 2). Among the included studies, 56.9% reported OMEs and 43.1% did not.

Preoperative opioid use varied, but combined cohorts with both naive and nonnaive patients represented 60.0% of the included studies. An opioid-naive cohort was present in 37.7% of the included studies, 1.5% of studies did not clearly state the preoperative opioid status of their patients and only 0.8% of studies specifically assessed cohorts that were nonnaive to opioid use.

Definitions

The definitions of prolonged opioid use were categorized into different timeframes, ranging from 1 to 24 months postoperatively, based on the first day the authors of a given publication considered opioid use to be prolonged. The 130 studies provided a total of 150 definitions due to 14 studies that used two or more definitions. Of these 150 definitions, 22 did not define prolonged opioid use by means of a postoperative timepoint and were therefore excluded from quantitative analysis, leaving a total of 128 eligible definitions.

The most common timepoint after which opioid use was considered prolonged was 3 months, which accounted for 67.2% of eligible definitions. Following prolonged use after 3 months, the most frequent postoperative time points were 6 months in 8.59%, 12 months in 7.81%, and 1 month in 5.47% of eligible definitions (Fig. 3).

Postoperative Timeframe and Prevalence of Prolonged Opioid Use

For each of the timeframes mentioned in Figure 3, the mean prevalence of prolonged opioid use was calculated. The highest mean prevalence of prolonged opioid use was found in studies that used a cutoff of 1.5 months with 36.0% (SD 17.5) of patients, 24 months (34.2%, SD 6.83), and 12 months (25.7%, SD 19.9), respectively. The lowest mean prevalence, at 1.60%, was reported by the single study that utilized a cutoff of 9 months (Fig. 4). No significant association was found between the definition timeframes and the prevalence of prolonged opioid use (Spearman’s correlation, P = 0.56).

Relationship between Surgical Specialty and Prolonged Opioid Use

The included studies were categorized according to the procedures being performed and the scope of the
**Fig. 3.** Definitions of prolonged opioid use. *N = 150, of which 22 were ineligible for time-based analysis.

**Fig. 4.** Prolonged opioid use by definition. *N = 122, due to eight studies not reporting a percentage of prolonged opioid use.
The most represented specialties in our analysis were orthopedic (n = 43), spine (n = 32), and mixed (n = 21), whereas the least represented specialty was pediatric surgery (n = 1) as shown in the footnote of Figure 5.

The mean prevalence of prolonged opioid use was calculated for each surgical specialty. The specialties with the highest prevalence of prolonged opioid use were otolaryngology (30.8%, SD 11.7) and spine (25.8%, SD 20.2), whereas gynecologic (3.80%, SD 3.30) and pediatric (4.80%, n = 1) were the lowest (Fig. 5). There were eight studies that defined prolonged opioid use but unfortunately did not report the percentage of patients who met those criteria.

The mean time after which opioid use was considered prolonged was calculated for each specialty. As shown in Figure 6, the mean of 6.90 months in spine surgery is an outlier compared to the mean timeframes of the other specialties.

Risk Factors for Prolonged Opioid Use

A wide variety of risk factors were reported to be significantly associated with prolonged opioid use. Preoperative opioid use was the most common risk factor (48.5% of studies), followed by mood disorders in 26.9% of studies. Other notable risk factors included pain or chronic pain (21.5%), tobacco use (21.5%), anxiety disorders (19.2%), and history of substance use (18.5%), as shown in Figure 7. Furthermore, for patients from the United States, being treated outside of the Northeastern United States was reported as a risk factor in six (4.62%) studies.

DISCUSSION

This systematic review highlights similarities in defining prolonged opioid use within existing literature, which was relatively consistent across various surgical specialties but without overall consensus. A timeframe of 3 months after surgery was found in 67.2% (n = 86) of the 128 time-based definitions, with 11 of the 12 categories using a timeframe between 2.50 and 4.17 months. Only spine surgery used a considerably different timeframe, at a mean of 6.90 months. Despite the seeming similarity of prolonged opioid use for varying specialties, there are no procedure-specific definitions for prolonged opioid use, particularly in the field of hand surgery. These would be useful for the practicing surgeon, as postoperative requirements generally vary for different operations and prolonged opioid use rates for elective hand surgical procedures are high.12

Orthopedic (n = 43) and spine (n = 32) surgeries were the most represented categories in the literature while other categories, such as gynecologic (n = 4) surgery, were less represented. There was only one publication in hand surgery that reported rates of prolonged opioid use.12 Similarly, on average 25.8% of spine surgery patients and 18.9% of orthopedic surgery patients were identified as prolonged opioid users, compared to only 3.80% of patients in gynecologic surgery. One possible explanation for the variability in specialty representation is that the perceived extent of prolonged opioid use within each field guides the number of publications. Most notable is that the average timeframe used to define prolonged opioid use in spine surgery is rather long (6.90 months). If Spine surgery used a timeframe to define prolonged use that was in agreement with other fields (eg, 3 months), the
proportion of patients with prolonged opioid use would likely increase. This emphasizes the need to reach a consensus within each surgical field on the definitions of prolonged opioid use, so that the results from research efforts correspond well with clinical experiences.

Although not all studies assessed risk factors for the development of prolonged opioid use, a selection of risk factors were commonly found. These included preoperative opioid use, a history of mood disorders, chronic pain, anxiety disorders, tobacco use, and a history of substance

Fig. 6. Mean timeframe to define prolonged opioid use by specialty. *N=128, due to 22 definitions not being time-based.

Fig. 7. Risk factors for prolonged opioid use.
use disorder. Specific to the United States, six studies reported being treated outside of the Northeastern United States as a risk factor for prolonged opioid use. A possible explanation for the regional differences could be increasing efforts in the Northeastern United States to reduce opioid use after an initial surge, such as the implementation of statutory prescription limits.\textsuperscript{16,17}

Consensus on definitions of prolonged opioid use will provide a benchmark for future research and a timeframe for clinicians to mitigate prolonged opioid use. Complete uniformity in the definition of prolonged opioid use may not contribute to achieving these aims. For example, both the studies by Steen et al\textsuperscript{18} and Zaveri et al\textsuperscript{19} used the timepoint of 3 months after surgery to define prolonged opioid use after major limb amputation and minor outpatient surgery, respectively. Clinically, using 3 months to define prolonged opioid use for both major limb amputations and less invasive procedures such as a carpal tunnel release fails to account for the expected differences in both the intensity and duration of pain.

In the absence of a procedure and specialty-specific consensus for postoperative prolonged opioid use within hand surgery, we propose criteria for four different categories of procedures within hand surgery along with example procedures for each of these categories (Table 1). These criteria represent a synthesis of our clinical expertise and aim to provide a boundary after which opioid use can be considered prolonged following various hand surgical procedures, which can be applied in research settings. The aim of the study was not to dictate an optimal prescription regimen for individual patients, which is at their treating physicians’ discretion. However, although clinician judgment will remain fundamental to the tailoring of opioid prescriptions based on patients’ risk factors, behavior, and complications, the purpose of these criteria is to lay the foundation upon which future clinical research can define procedure-specific prolonged opioid use for clinical application. Future research efforts should be undertaken to reach a procedure-specific consensus on how to define prolonged opioid use and these studies should involve the public, perhaps by means of a Delphi process.

In addition to the proposed criteria, the reported risk factors should be considered to counsel at-risk patients. Additionally, opioids should be given in conjunction with acetaminophen or non-steroidal anti-inflammatory drugs since this has been demonstrated to reduce postoperative opioid dependence.\textsuperscript{20} Furthermore, the authors suggest that the evaluation of OMEs may be helpful in critically reviewing the amount of opioids prescribed to individual patients, and should be standard practice in research on this topic. Last, dose tapering strategies have been demonstrated to be effective in weaning patients off their opioid therapies.\textsuperscript{10}

The results of this review should be interpreted in the light of its limitations. While the populations studied were cohorts of patients with postoperative opioid use, we were unable to analyze surgical outcomes and their effect on prolonged opioid use as this information was not included in the studies we analyzed. During full-text review the authors found that a vast majority of the included studies evaluated representative populations for their respective surgical procedures, to avoid introducing bias by specifically selecting patients with better or worse surgical outcomes. A second limitation is the limited number of studies for some of the surgical specialties. Still, this also suggests that some surgical fields may benefit from further research on prolonged opioid use to evaluate the extent of this problem in that field. Last, even though gray literature was not analyzed, our data provide a focused report on what the primary literature considers prolonged opioid use.

Several findings of this review may inspire future research efforts. The first of these findings is that only four papers studied a patient population with a mean or median age below 30 years, indicating that younger patients may be underrepresented in research on this topic.\textsuperscript{21–24} Therefore, it is not well understood whether the extent of this problem is different in patients of this age, and whether risk factors for prolonged use may be different. Since only one paper (0.8%) focused solely on a non-naive cohort, the authors hypothesize that research efforts

| Table 1. Proposed Criteria for Defining Prolonged Opioid Use in Hand Surgery Based on Clinical Expertise of Authors |
|---------------|---------------|---------------|
| **Type of Surgery** | **Example Procedures** | **Proposed Criteria** |
| **Minor soft-tissue procedures** | · Carpal tunnel release | Opioid use beyond 2 wks (14 d) should be considered abnormal |
| | · Trigger finger release | |
| | · Dupuytren fasciectomy | |
| | · First compartment release for de Quervain tenosynovitis | |
| | · Flexor tendon repair or tenolysis | |
| | · Extensor tendon repair or tenolysis | |
| | · Minor excisions (eg, mucoid cyst) | |
| **Major soft-tissue procedures** | · Extensive debridement and soft-tissue coverage with local tissue rearrangement or advancement | Opioid use beyond 1 mo (30 d) should be considered abnormal |
| | · Flap coverage for soft-tissue defect | |
| | · Skin grafting for significant burn injuries or soft-tissue defect | |
| **Minor bone procedures** | · ORIF or closed reduction and pinning of fractures from the metacarpal level and distal CMC | Opioid use beyond 1 mo (30 d) should be considered abnormal |
| | · Amputation of digit | |
| | · CMC arthroplasty | |
| | · PIP joint arthroplasty | |
| **Major bone procedures** | · ORIF of radius, ulna, or humerus | Opioid use beyond 6 wks (42 d) should be considered abnormal |
| | · Total shoulder arthroplasty | |
| | · Major limb amputation | |
| | · Corrective osteotomy | |

CMC, carpometacarpal; ORIF, open reduction internal fixation; PIP, proximal interphalangeal.
specifically in nonnaive patients may aid in identifying additional risk factors in this already at-risk population. Furthermore, the authors suggest that future research efforts may benefit from standardization of terminology and methods to report the amount of opioid use. Last, although the proposed categorizations of procedures within hand surgery are based on both literature review and clinical experience, a prospective evaluation is warranted to determine their validity. The authors encourage academic discussion of the criteria to identify potential shortcomings and to expand the list of procedures.

CONCLUSIONS

Although there are similarities in the definitions of postoperative prolonged opioid use, there is poor consensus about procedure-specific definitions of prolonged opioid use after surgery. The authors therefore emphasize the need for consensus building about the definition of prolonged opioid use following hand surgery, so that future research efforts may accurately estimate the extent of this problem and that clinicians may specifically counsel at-risk patients.

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