



Hello, my name is Eric Chen from Boston University and today we'll be talking about safe postoperative prescribing of opioids.



Opioid analgesia has long been used for pain control after surgery but the first question is – is surgery a risk factor for chronic use?

Here we show data from large insurance database studies looking at both elderly Canadians as well as privately insured American adults. And they showed that anywhere from one to six percent of patients used opioids for over three months after their surgery. These studies looked at a variety of surgical procedures and found evidence of chronic use after even minor surgery such as carpal tunnel releases and major, more invasive surgeries such as total knee arthroplasties.

	% of Patients	
	Estimated Incidence of Chronic Pain	Estimated Chronic Severe (Disabling) Pain (>5 out of 10
Amputation	30–50	5–10
Breast surgery (lumpectomy and mastectomy)	20–30	5–10
Thoracotomy	30-40	10
Inguinal hernia repair	10	2-4
Coronary artery bypass	30-50	5–10
Caesarean section	10	4

Related to chronic opioid use and somewhat going hand in hand with it, is also the incidence of chronic postsurgical pain. This study by Sinatra found high rates of chronic pain after surgeries such as amputations and thoracotomies but also present after more elective or less invasive cases such as inguinal hernia repairs and Caesarian sections. And so all surgery due to its inherently invasive nature is a source of acute pain.



But how exactly does the experience of acute pain become chronic? Here this figure sort of conceptually shows this transition starting from surgery causing inflammation, which causes initially a transient activation of nociceptive fibers, which then become sensitized and then activated in a more sustained manner. This ultimately results in a structural remodeling of the central nervous system, which results in hyperactivity and chronic pain. And evidence suggests that these changes are mediated by alterations in expressions of neurotransmitters, receptors, ion channels, as well as structure, connectivity and survival of neurons.



Can we predict which patients are at higher risk of developing chronic postsurgical pain? Indeed some risk factors have been identified. Much of the experience of postoperative pain can be attributed to psychosocial factors such as anxiety, depression as well as catastrophization or essentially a patient's ability to cope with their individual surgical experience. However, beyond patient related factors such as younger age or pain syndromes, there are some operative variables that can play a role in increasing the risk for chronic postsurgical pain, such as the surgical procedure or technique or presence of any sort of ischemia or nerve injury during or after surgery.



Here we give the example of the StarT back streaming tool, which is a nine question survey that can help stratify patients into being low, medium or high risk of chronic back pain disability. As you can see, this survey includes questions regarding physical symptoms such as radiating pain down the legs, shoulder and neck pain, as well as more psychosocial questions such as worrying thoughts or not enjoying things that the patient used to enjoy.

Over-prescription After Surgery			
Reference	Surgery	Over-prescription	
Kim et al. JSBS. 2016	Upper extremity procedures	84% did not complete meds	
Rodgers et al. J. Hand Surg. 2012	Upper extremity procedures	77% reported taking half or less	
Chapman et al. Hand. 2017	Carpal tunnel release	4.3 pills consumed on average	
Bartels et al. Plos One. 2016	Cesarean section	83% reported taking half or less	
Bartels et al. Plos One. 2016	Thoracic surgery	71% reported taking half or less	
Hill et al. Annals Surgery. 2016	General surgery procedures	71.3% of pills not consumed	
Bates et al. J. Urology. 2011	Urologic procedures	42% of meds not consumed	
Harris et al. JAMA Derm. 2013	Dermatologic procedures	86% had leftover pills	

Now in the effort to sort of prevent patients from having too much pain, which may lead to chronic pain, have surgeons sort of taken the pendulum and swung it too far? And are they overprescribing opioids after surgery? Several studies out of the past several years have shown there is evidence for over prescription after surgery. Here are just some examples of some of these studies from the upper extremity literature, general surgery literature, as well as urologic and dermatologic literature.



And so the bulk of this presentation is going to center around strategies to reduce opioid prescribing. These strategies can be generally divided into three sections, which we'll talk about individually, namely provider based strategies as well as procedure and patient based strategies.





We'll start by talking about provider based strategies. These strategies primarily center around guidelines for appropriate opioid prescription as well as the use of non-opioid analgesia in a multi-modal fashion.



Here is a clinical practice guideline on the management of postoperative pain, released in 2016, from the American Pain Society, the American Society of Regional Anesthesia and Pain Medicine, and the American Society of Anesthesiologists.



Shown here is a summary of their strong recommendations, namely conducting a preoperative evaluation, providing patient centered education including the tapering of analgesics after discharge. Monitoring patients in terms of their analgesic regimens as well as for adverse reactions or signs of dependence and abuse. As well as the encouragement of PO opioids over parental routes. They do recommend that PCAs be used but caution against the routine use of basal infusions.

Highlighted in green is their recommendation to offer multi-modal analgesia such as the use of acetaminophen or nonsteroidal anti-inflammatories or other non-opioid medication such as gabapentin or region nerve blocks. Furthermore, they also recommend establishing or advocating for organizational structures to be put in place for safe effective postop analgesia such as the use of an acute pain service or sort of spinal or regional anesthesia when appropriate.



Here is another practice guideline from 2012, from the American Society of Anesthesiologists...





...highlighting many of the same concepts, schedules, multi-modal analgesia, which they found had been associated with superior pain relief, decreased opioid use and decreased opioid related adverse events.

Oral Analgesics for Acute Postoperative Pair				
Oral analgesics for postop pain ~50,000 participants in ~460 high-quality studies				
Analgesic(s)	Dose (mg)	NNT vs Placebo for at least 50% maximum pain relief over 4-6 hours		
SINGLE AGENTS:				
Ibuprofen	600	2.7		
Naproxen	500	2.7		
Celecoxib	400	2.6		
Acetaminophen (APAP)	1000	3.6		
Oxycodone	15	4.6		
Codeine	60	12.0		
Gabapentin 250 11.0		11.0		
COMBINATIONS:				
Ibuprofen + APAP	400+1000	1.5		
Ibuprofen + oxycodone	400+5	2.3		
APAP + oxycodone	325+5	5.4		
APAP + codeine	300+30	6.9		

Now what about some evidence for multi-modal analgesia? I'm sure that we've all had patients who have reported that medications such as ibuprofen or Tylenol do not work for them and that opioids are the only effective pain medications for them. However, there is evidence such as this Cochrane Review that suggests that non-opioid medications can work just as good if not better than opioid medications such as Oxycodone or codeine and that combinations of these medications work even better.

Slide 16

(Oral Opioid and Nonopioid Equivalents for Acute Pain					
	RCT comparing 4 oral analgesics for mode in 416 patients in 2 urban EDs (baseline m	erate to severe acute extremity pain nean pain score was 8.7)				
	Analgesic Groups Decreases in Pain Scores					
	ibuprofen 400 mg + APAP 1000 mg	4.3 (95% CI 3.6 to 4.9)				
	oxycodone 5 mg + APAP 325 mg 4.4 (95% Cl 3.7 to 5.0)					
	hydrocodone 5 mg + APAP 300 mg 3.5 (95% Cl 2.9 to 4.2)					
	codeine 30 mg + APAP 300 mg 3.9 (95% Cl 3.2 to 4.5)					
	 Adverse events were not assessed. 					
 CONCLUSIONS: For patients presenting to the ED with acute extremity pain, there were no statistically significant or clinically important differences in pain reduction at 2 hours among single-dose treatment with nonopioid and opioid analgesics 						
с	hang AK, et al. JAMA. 2017 Nov 7;318(17):1661-1667.	16				

Here is another study looking at patients who present to the emergency room with acute extremity pain and they found that there was no statistically significant or clinically important difference in pain reduction over two hours between single doses of non-opioid and opioid analgesics. Though NSAIDs are effective as a non-opioid analgesic and does not have the risk for dependence or abuse that opioids do, they do have their own sort of side effects, namely effects possibly on bleeding as well as GI problems and also on fracture healing.

studies in hum	ans on the	effect of NSAI	Ds on fracture healing
References	Design	NSAID	Outcome
Vo effect Wurning et al.	Prospective	Indomethacin	Hip replacement: no effect on periprosthetic
Davis and Ackroyd	RCT	Fluriprofen	Colles fracture
Adolphson et al.	RCT	Piroxicam	Colles fracture
Sculean et al.	RCT	Rofecoxib	Periodontal defects after periodontal surgery with enam matrix proteins
Meunier et al.	RCT	Celecoxib	Prosthesis migration, pain scores, range of motion, and subjective outcomes after total knee replacement
Vitale et al.	Retrospective	Ketorolac	Reoperation after scoliosis surgery
Pradhan et al.	Retrospective	Ketorolac (48 h)	Spinal fusion rate
Sucato et al.	Retrospective	Ketorolac	Pseudoarthrosis after spinal fusion
Horn et al	Retrospective	Ketorolac	Nonunion after spinal fusion

While the primary evidence for inhibition of fracture healing by NSAIDs primarily has been shown in animal data as well as with more potent NSAIDs such as indomethacin or ketorolac, many studies in humans have been variable and inconclusive in their findings. So in cases where bony healing is required after surgery, NSAIDs should be used in more of a case to case basis based on the risk of delayed bony healing.



I do believe that a multi-modal pain strategy is one of the most effective ways to decrease opioid use after surgery. And there are sort of many opportunities in a patient's perioperative course where a comprehensive multi-modal strategy can be used. For example, before surgery, patients can work on decreasing their pre-op opioid utilization, which has shown to be a risk factor for chronic postoperative use. Physicians and providers can set expectations for the postoperative pain experience. Pain is normal after surgery. It is expected. But it will resolve. Patients can also work on some self-management strategies such as meditation and music, as well as sort of trying to develop their own personal coping strategies for the postoperative pain.

Intraoperatively the use of field blocks or regional blocks can be highly effective in providing sort of short term analgesia and decreasing the total opioids used during surgery.

Postoperatively in addition to different types of medications that work on different areas of the pain pathway, cryotherapy with ice or some type of cooler can be very effective as well as cognitive and psychological therapies, as well as a specific plan for the patient regarding how to use opioid medications and how to eventually wean themselves off of it as they recover from surgery.



Short-acting opioids should really be used. There are different kinds of nonsteroidals such as selective COX-2 inhibitors as well as nonselective ones that can be used. Gabapentin has also been shown to be effective in decreasing opioid use and obviously Tylenol as well. To the patients who feel that these non-opioid medications are not effective, I think that's important to emphasize to them that they can be and even if they don't notice an effect, to still try to take them and encourage them to use them as first-line therapy as it may help decrease their opioid use even if they don't realize it.





The next section is more procedure based, which is essentially evidence based prescribing based on the type of surgery being performed.

How much opioid analgesia is needed after surgery?

And the big question here is how much opioid analgesia is needed after surgery?

Pain Medication After Surgery and Injury			
Differences Operative T	in Prescription of reatment of Hip ar States and th	Narcotic Pain I nd Ankle Fractu ne Netherlands	Medication After tres in the Unite
Mark Vrahas, MD, David Ring, MD, and Peter Kloen, MD, PhD Discharged on Opioids Amsterdam MGH			
Discharged o	n Opioids	Amsterdam	MGH
Discharged of Hip Fractures	n Opioids	Amsterdam 0%	MGH 77%
Discharged of Hip Fractures Ankle Fractur	n Opioids es	Amsterdam 0% 6%	MGH 77% 82%
Discharged of Hip Fractures Ankle Fracture Femur Fracture	n Opioids es s ²	Amsterdam 0% 6%	MGH 77% 82%
Discharged o Hip Fractures Ankle Fractur Femur Fracture IM Nail Femur	n Opioids es s ² Opioid Requirement (2 weeks post-op)	Amsterdam 0% 6% Adequate Pain Control	MGH 77% 82% Accurate Pre- conception of Pair
Discharged of Hip Fractures Ankle Fracture Femur Fracture IM Nail Femur Vietnam	n Opioids es S ² Opioid Requirement (2 weeks post-op) 0.9 MME/kg/day	Amsterdam 0% 6% Adequate Pain Control 92%	MGH 77% 82% Accurate Pre- conception of Pair 76%

And the answer is it varies. So here are two studies showing the use of opioid medications after surgery in American patients and in those in the Netherlands or in Vietnam. So here on the top this was a study that looked at opioid use after hip and ankle fracture surgery, comparing the use in the Netherlands as well as here at Mass General in Boston. They showed that patients in the United States, nearly all of them got some sort of opioid medication at discharge, whereas in the Netherlands only a small minority of patients did.

On the bottom is a study comparing opioid utilization after femur fracture surgery, comparing the use between American patients and Vietnamese patients. And they showed that American patients used nearly 30 times more opioids than the Vietnamese ones in the two weeks postoperatively. Surprisingly 92 percent of Vietnamese patients reported adequate pain control whereas only 20 percent of the American patients did. And they sort of attributed this difference into the pre-conception of pain. Most American patients reported that their perioperative experience was way more painful than they had originally imagined, whereas 76 percent of the Vietnamese patients stated that the surgery hurt as much as they thought it was going to.

I think these studies are interesting and they really show that there is somewhat of a cultural or psychological factor in the experience of pain after surgery.



And so what is the ideal length of an opioid prescription? So this study by Scully, et al, looked at the need for receiving more than one opioid refill as a function of the initial length of prescription. And they sort of found that the length of prescription, if it were too short, would place patients at higher risk of needing multiple refills as well as if the prescription was too long, then they would also be at high risk of requiring a refill. Amongst different surgeries, they recommended that for patients undergoing musculoskeletal procedures that the prescription length be from between six to 15 days; for women's health surgeries four to 13 days; and for general surgery procedures four to nine days.



Here is a tool that was developed for opiate prescribing after hand surgery. It was sort of named the Pink Card and it was given to all the providers and provides sort of guidelines for postoperative analgesia after hand and upper extremity surgery. You can see on one side they have sort of best practices in terms of sort of multi-modal analgesia. Sort of on the second side they have opioid guidelines where they list the recommended amount of opioids prescribed based on the type of procedure. With smaller surgeries such as carpal tunnel releases, trigger finger releases, requiring no opioids, up to 20 or 40 tabs for more invasive and painful surgeries.



They found that by using this Pink Card they were able to produce a specifically significant reduction in opioid prescribing in their departments.



Here is another study presented last year at the Hand Society meeting, highlighting surgeon education and opioid guidelines for carpal tunnel and distal radius fracture surgeries, where they had different recommendations for the amount to prescribe based on whether the patient was older or younger than 65. And they found that this recommendation was able to decrease the amount prescribed after both surgeries.

Slide 27	
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Procedure-based Reduction					
 General Surgery Procedures Surgeon Education Opioid Guidelines 					
	# Pills	# Pil	lls		
Procedure	Recommendation	Before	After	Change	
Partial mastectomy	5	20	5	-74%	
Partial mastectomy sentinel lymph node biopsy	10	24	10	-59%	
Laparoscopic cholecystectomy	15	35	19	-45%	
Laparoscopic inguinal hernia repair	15	34	19	-43%	
Open inguinal hernia 15 33 18 -45%					

This is the hand literature, the study by Hill, et al, used a similar surgeon education opioid guideline protocol where they first surveyed a large number of their patients and asked them to log their opioid use and then created recommendations based on the patient's reported use. And they came up with these recommendations.

They then applied these sort of recommendations and best practices to all their other patients and they were able to show quite a significant reduction in opioid prescription after a variety of general surgery procedures.

In addition to lowering the amount of opioids prescribed, they also found that the amount of refills that patients required was not significantly affected, suggesting that they were not sort of under prescribing but merely reducing the rate of over prescription.





Finally the last section or strategy to reduce opioid prescribing centers around the patient, which essentially tries to create a plan that tailors opioid prescribing to an individual patient's specific utilization.

Patient-based Redu	uction
 Inpatient Ortho Procedures Surgeon Education Opioid taper 	Opioid Pain Control Plan: This extended taper is recommended for treatment of your acute post-operative pain. Take your medications as needed. DO NOT exceed the daily maximum amount specified below. Track the amount you are taking daily and bring this log to your follow-up appointments for review. Dispose of excess medications responsibly (www.mass.gov).
	Day - Daily Maximum Amount Taken
	1 - 11 tablets
	2 - 9 tablets
	3 - 8 tablets
	F 6 tablets
	6 - 5 tablets
	7 - 4 tablets
	8 - 3 tablets
	9 - 3 tablets
	10 - 2 tablets
	11 - 2 tablets
	12 - 2 tablets
	13 - 2 tablets
Chen EY, et al (unpublished). Boston Medical Center.	14 - 1 tablets

So this is an initiative that we have implemented here at Boston University, which centers around this opioid taper, which is based on a patient's 24 hour opioid utilization prior to discharge. And essentially provides a 14 day taper, which tells patients the daily maximum of tablets that they can take and includes a space where they can log their use for review with their physicians.



This taper was implemented around May 2017, and we found that it has been able to produce a significant reduction in opioid prescription at discharge after inpatient orthopedic surgeries.



So finally, how do we implement these changes? I think the first step would be to gather data on the opioid prescription patterns in your own practice. I mean, once you have an assessment of the opioid prescribing patterns at your own institutions you can then sort of compare them to existing evidence and consensus for sort of reasonable opioid requirements.

I also do feel that it's important to survey patients regarding their postoperative opioid use to sort of get a better sense as to whether or not they're getting too much or too little opioids.



Oral Oral

IV / Parenteral

IV / Parenteral

0

32

Select Other Select Other

Select Other

Chen EY, et al (unpublished). Boston Medical Center.

Select Other

Just to recap, there are several decision aids that have been reported to be effective, such as the Pink Card. Guidelines have been released to help guide clinical practice and those can be consulted as well.



In terms of institutional support with the advent and proliferation of electronic medical records, I think that it's important to have good default settings for opioid medications and that they're not ordered to be prescribed too frequently or having default values that make it easier for the prescriber to prescribe a reasonable amount rather than sort of having to make up a number on the fly.

I think policy changes can also be effective, telling patients that you don't refill prescriptions overnight or weekends can prevent s ort of unwanted phone calls and also give patients the expectation to either try to taper their own medications if a weekend is coming up. Or to try to learn how to wean or go longer and longer without opioids.

Obviously everyone would love additional staffing and resources but if they are available I think they can be immensely useful in sort of following up and getting data from patients as well as training providers on responsible opioid prescribing as well.



All medications have side effects. So patients should be counseled on how to recognize these side effects and to report them. And I will just add that though we encourage multi-modal therapies, the combination of benzodiazepines and opioids together does increase the risk of an overdose.



I also think printed handouts and educational materials can be extremely useful. That can detail multi-modal therapy, the ability to log their own opioid use as well as information for contacting their providers should they run into any problems or concerns.

Summary
Post-op pain can be effectively managed with fewer opioids than currently prescribed
Various strategies exist for standardizing post-op opioid prescribing
Education can help improve patient's post-operative experience

So in summary, postop pain can be effectively managed with fewer opioids than are currently being prescribed. I do think that the pendulum has swung a bit too far in the emphasis on eliminating postoperative pain and that many studies have shown over the past couple of years that patients can have effective analgesia with less opioid medications being prescribed.

There are various strategies that exist for standardizing postop opioid prescribing. I think that standardization is important because having a standardized protocol in place allows for better setting of expectations and a more reproducible outcome after surgery.

And finally, education can help improve a patient's postoperative experience. And the most effective time to discuss these things is before the patient undergoes surgery.

I thank you for your time and attention. I hope that some of this information can be used to help your own patients with their pain after surgery. Thank you.