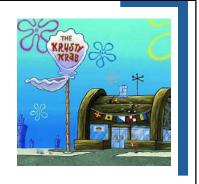
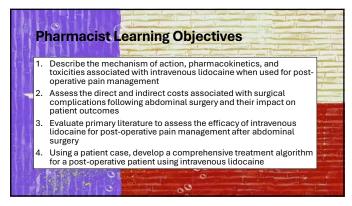
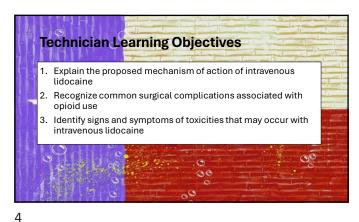
Can You Feel It Now Mr. Krabs? Evaluating IV Lidocaine for Post-Operative Pain Management after Abdominal Surgery Nicholas Martin, PharmD PGY-1 Pharmacotherapy Resident UIW Feik School of Pharmacy

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3

Abbreviations					
Abbreviation	Meaning	Abbreviation	Meaning		
APS	American Pain Society	POD	Post-operative day		
ORADEs	Opioid-related adverse drug events	MSOFA	Modified sequential organ failure assessment		
POI	Post-operative ileus	MDE	Morphine dose equivalents		
VAS	Visual analog scale	IBD	Inflammatory bowel disease		
IQR	Interquartile range	ITT	Intention-to-treat		
ADR	Adverse drug reaction	PCA	Patient controlled analgesia		



5

# **Knowledge Check**

7

Which of the following best describes the principle of multimodal analgesia in postoperative pain management?

- A. Using a single analgesic medication at high doses to control pain
- B. Combining multiple medications and techniques that target different pain pathways to enhance pain relief and reduce side effects
- C. Focusing on opioid utilization for pain relief
- D. Utilizing non-pharmacological methods, such as physical therapy, to manage pain

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- D. Utilizing non-pharmacological methods, such as physical therapy, to manage pain



## **Post-Operative Pain**

- More than 80% of patients who undergo surgical procedures experience acute postoperative pain
- Less than half of patients who undergo surgery report adequate pain relief
- 2016 APS guidelines for management of postoperative pain recommends the use of multimodal analgesia

J Pain, 2018, 17(2): 131-157

10

8

#### **Providing Pain Control**

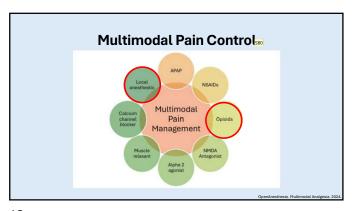
- The Joint Commission emphasizes a patient's right to pain relief but emphasizes:
  - $\circ\,\text{Safe}$  use of opioids
  - Patient education on pain management
  - Multimodal approach to pain control

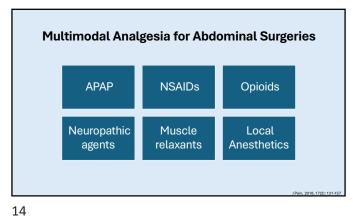


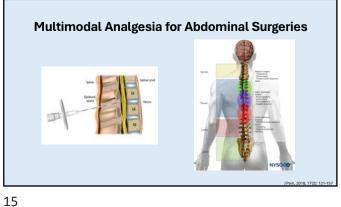
TIC, 2017, Hospital and Hospital Clinics Manual: Leaders



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## **Epidural Contraindications**

Active infection

16

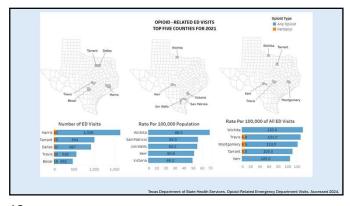
- Allergy to anesthetic agent
- Use of anticoagulation, clotting disorders
- Spinal deformities
- Patient refusal or inability to cooperate

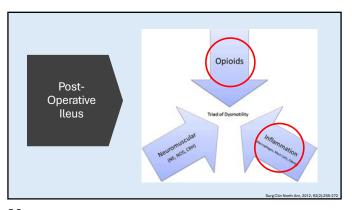




17 18

Make sure to explain not all these are used routinely. I like that this SB0 image shows the options in pretty much our order of prevalence/usage starting with APAP and going clockwise. Berman, Dr. Sarah E., 2024-12-16T21:33:41.226





## **Knowledge Check**

Which of the following is NOT a contributing factor to the etiology of postoperative ileus?

- A. Opioids
- B. Inflammation
- C. Neuromuscular factors
- D. Increased physical activity after surgery

## **Knowledge Check**

Which of the following is NOT a contributing factor to the etiology of postoperative ileus?

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21 22

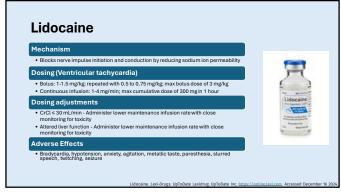
## **Post-Operative Ileus (POI)**

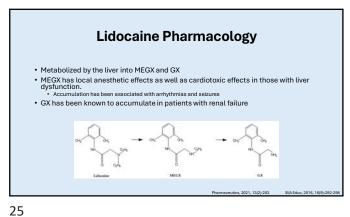
- Delays recovery time and prolongs hospitalization
- Up to 25% of colectomy patients experience POI, which doubles their cost of care
- Estimated to cost the healthcare system an additional \$750 million USD annually from prolonged initial hospitalization alone



Sung Clin North Am. 2012. 92(2):2

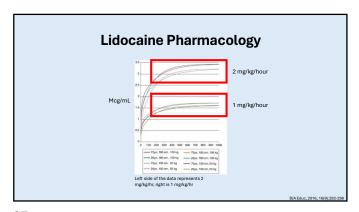
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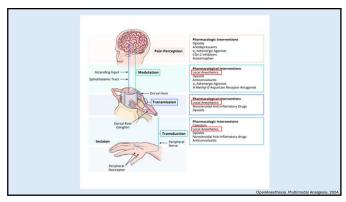
Lidocaine Pharmacology: Analgesic • Effects attributed to NMDA and voltage gated calcium receptors Pain intensity Injury Animal studies with IV allodynia lidocaine showed reduced hyperalgesia and modulation of inflammation Stimulus intensity

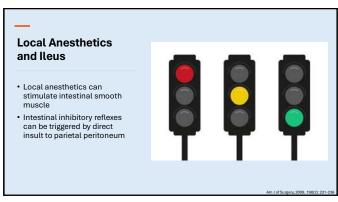
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**Toxicity** Relationship of Signs and Symptoms of Narrow therapeutic Lidocaine Toxicity to Serum Concentration range (1.5-5 µg/mL) • Lidocaine levels recommended in: o Durations >24 hours Heart failure, liver and renal dysfunction

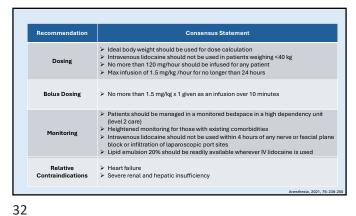
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29 30









Objective • Evaluate the use of intravenous lidocaine in ICU patients as an adjunct analgesic agent for pain **Study Design** • Multicenter (2 institutions), retrospective, prepost intervention, observational chart review

• ≥ 18 years old ICU stay ≥ 24 hours Systemic lidocaine infusion for any duration as an adjunct therapy for pain management during ICU stay **Exclusion** · Received intravenous lidocaine within 24 hours of admission · Patients who received ketamine infusion in conjunction with intravenous lidocaine or within 24 hours of discontinuation · Started on intravenous lidocaine as part of comfort care due to terminal

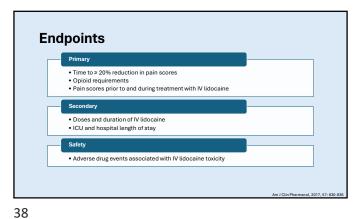
35 36

Intervention

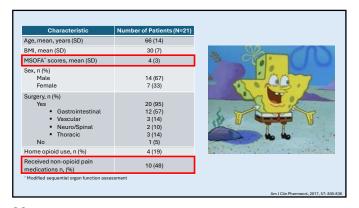
Initiation, dosing, and duration determined at provider discretion

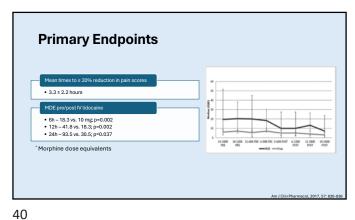
Continuous infusion at an average rate of 0.93 mg/min for a mean duration of 48 hours

Pain scores recorded for 24 hours prior to IV lidocaine initiation



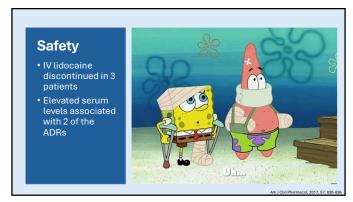
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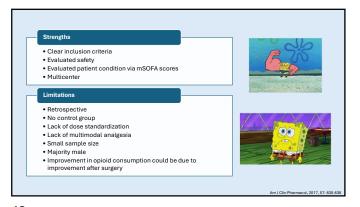


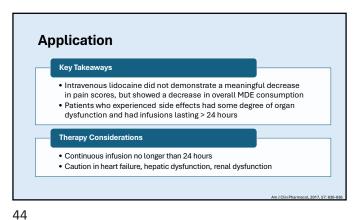
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Patient	Patient Factors	Noted Reaction	Dose and Serum Level	Timing
Patient #1	History of diastolic heart failure and atrial fibrillation     SCr of 1.56 mg/dL (baseline 1)     BUN of 40 mg/dL (baseline 18)     Decreased urine output (0.1-0.2 mL/[kg*hr])	Less responsive on Riker Sedation- Agitation scale (from 5 to 2)	Dose – 1 mg/min Serum level – 8.4 μg/mL	26 hours after IVL initiation
Patient #2	History of congestive heart failure     SCr of 1.51 mg/dL (baseline 0.86)     Elevated AST/ALT of 339 and 523, respectively (baseline WNL)	Episodes of dizziness when attempting to stand up	Dose - 0.5 mg/min Serum level: #1 – 3.2 µg/mL #2 – 11.5 µg/mL	#1 – 19 hours afte IVLI initiation #2 – 63 hours afte IVLI initiation
Patient #3	History of aortic valve replacement     SCr of 2.58 mg/dL (baseline 1.06)	Disorientation and confusion	Dose – 2.1 mg/min Serum level not collected	19 hours after IVL



41 42







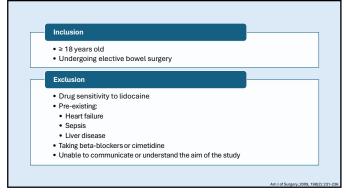
Objective

• Evaluate postsurgical pain, return of bowel function, amount of morphine used, and length of stay in patients undergoing elective bowel surgery who receive IV lidocaine compared to placebo

Study Design

• Randomized, double-blinded, placebo-controlled trial

5 46



Intervention

Intravenous lidocaine at 1 mg/min^ for 24 hours with a morphine PCA

Control

Normal saline for 24 hours with a morphine PCA

\*Corresponds to a dose range of 0.63-0.95 mg/kg/hour based on mean patient weights

Am J of Surgery, 2009, 198(2): 231-236.

47 48

Endpoints

VAS scores at 6,
18, and 24 hours
after surgery

Morphine use at 6,
18, and 24 hours
after surgery

Time to first flatus
(hours)

Hospital length of
stay (days)

**Baseline Characteristics** Characteristic Lidocaine (N=11) Placebo (N=11) 60 ± 5.7  $65 \pm 3.5$ Age, years 55% Sex, male (%) 55% Weight (kg) 83 ± 11 74 ± 11  $2.18 \pm 0.18$  $2.0 \pm 0.19$ ASA score Data represented as mean ± SEM unless otherwise noted

50

52

49

Pain Scores and Morphine Use					
VAS Score	Lidocaine Group	Placebo Group	P-value		
VAS at 6 hours	44.6 ± 5.59	54.0 ± 5.36	0.241		
VAS at 18 hours	38.8 ± 8.8	49.3 ± 6.2	0.2606		
VAS at 24 hours	26.1 ± 8.2	45.4 ± 6.4	0.08		
Morphine Use	Lidocaine Group	Placebo Group	P-value		
Morphine use at 6 hrs	17.5 ± 3.2	17.3 ± 3.4	0.97		
Morphine use at 18 hrs	19.0 ± 3.8	14 ± 2.5	0.2987		
Morphine use at 24 hrs	11.63 ± 3.8	8.36 ± 1.8	0.4496		
Total morphine	47.2 ± 8.2	39.7 ± 5.3	0.4525		

Endpoint	Lidocaine	Placebo	P-value
Flatus (h)	68.2 ± 9.7	86.9 ± 13.6	0.2802
Bowel movement (h)	88.3 ± 6.08	116.2 ± 10.1	0.0286
Days in the hospital	3.76 ± 0.24	4.93 ± 0.42	0.0277

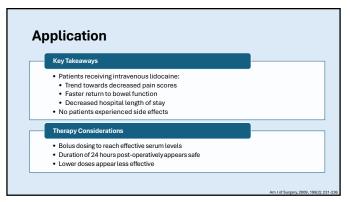
51

Strengths

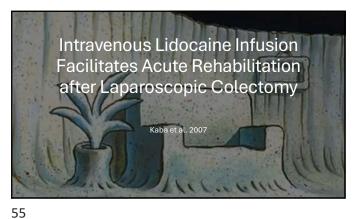
• Randomized
• Effective double-blinding
• Standardization of pain management protocol with morphine PCA
• Evaluated patient safety

Limitations

• Small sample size
• Single, community hospital
• Low doses of lidocaine relative to patient weight
• No primary outcome or study power defined



53 54



Objective • Evaluate the effects of intravenous lidocaine on facilitation of acute rehabilitation after laparoscopic colectomy **Study Design** • Randomized, double-blinded, placebocontrolled trial in Belgium

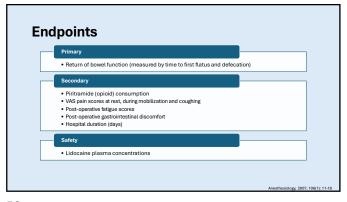
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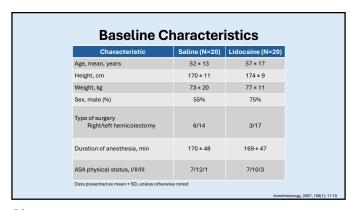
 Elective laparoscopic colectomy for nonmalignant disease ASA status I-III Exclusion >70 years old • History of gastroduodenal peptic ulcers Renal or hepatic failure Psychiatric disorders · Steroid treatment • Chronic treatment with opioids

Intervention • IV lidocaine bolus of 1.5 mg/kg, followed by mg/kg/hour during surgery, followed by mg/kg/hour for 24 hours post-operatively Control • Equal volumes of normal saline \* All patients received acetaminophen 2 g IV 30 minutes before the end of surgery and then every 6 hours and ketorolac 30 mg IV every 8 h

58

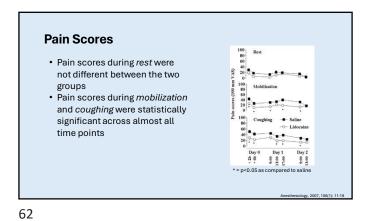
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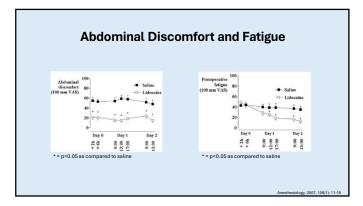




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Endpoint	Saline	Lidocaine	P-value
First flatus (hours)	28 [25-33]	17 [11-24]	<0.001
Defecation (hours)	51 [41-70]	28 [24-37]	0.001
Hospital stay (days)	3 [3-4]	2 [2-3]	0.001





Hours after surgery	Saline	Lidocaine	P-value
0-2	8 [4-11]	2 [0-5]	0.002
2-6	3 [0-9]	2 [1-3]	0.46
6-20	7 [4-16]	3 [2-9]	0.06
20-24	6 [3-7]	1 [0-1]	<0.001
0-24	22 [14-36]	8 [5-18]	0.005
Reported as median [IQR]			

63 64

Time	Plasma Concentration, µg/mL, mean ± SD	Highest Plasma Concentration, µg/mL
5 minutes*	1.6 ± 0.9	3.5
15 minutes*	1.3 ± 0.4	2.1
60 minutes*	1.8 ± 0.5	2.6
End of surgery	2.4 ± 0.6	4.0
End of 24-hour infusion	2.7 ± 1.1	4.6

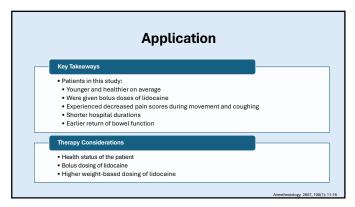
Strengths

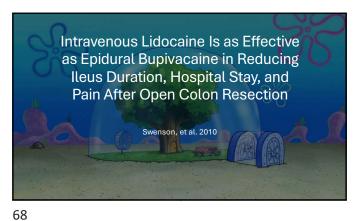
• Randomized
• Balanced groups
• Standardization of surgical procedures and pain management
• Bolus and weight-based dosing utilized

Limitations

• Small sample size
• Single center
• Primary outcome was return of bowel function

65 66





Objective

• To compare perioperative administration of IV vs. epidural local anesthetic in combination with epidural hydromorphone in patients undergoing open colon surgery

Study Design

• Randomized control trial, single center in Virginia

Inclusion

18-75 years old
Scheduled for elective colon resection
ASA status I-III

Exclusion

Allergy to local anesthetics
Myocardial infarction within 6 months before surgery
Liver disease or renal impairment (CrCl < 60 mL/min)
Systemic corticosteroid use
Chronic use of opiates
Unwillingness or contraindication to epidural analgesia
Pregnancy or active breast feeding

69 70

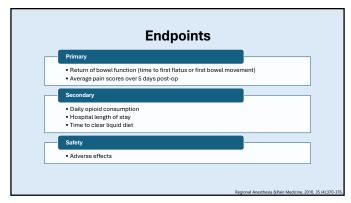
Intervention

• IV lidocaine bolus of 1.5 mg/kg at induction, followed by a maintenance dose of 1 mg/min in patients <70 kg or 2 mg/min for patients ≥70 kg
• Continued for 5 days or until first flatus

Control

• T8-T12 epidural with bupivacaine 0.125% and hydromorphone 6 μg/mL

@ 10 mL/hour within 1 hour of the end of surgery
• Continued until first flatus, but could be continued at discretion of team

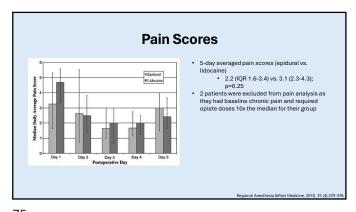


71 72

**Baseline Characteristics** 49 (36-54) 52 (40-62) 0.23 Female sex (%) 4 (20%) 12 (55%) 0.021 BMI, kg/m² 28 (22-31) 25 (19-29) 0.20 ASA Score (%) 1 (5%) 18 (95%) 0 (0%) 1 (5%) 14 (64%) 7 (32%) 0.014 69.9 hours (±28.23 91.6 hours (±41.05 hours) Duration of infusion, hours N/a orted as median [IQR] unless other

Endpoint	Epidural	IV Lidocaine	P-value
First flatus	1.6 (1.2-3.4)	2.7 (1.9-3.5)	0.17
First bowel movement	3.0 (1.7-4.5)	2.9 (2.3-3.6)	0.99
Time of advancement to clear liquid diet	3.6 (2.6-4.8)	2.9 (2.7-3.7)	0.47
Hospital length of stay	5.3 (4.7-7.9)	5.1 (4.8-5.9)	0.80

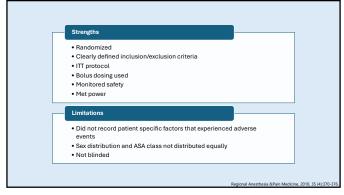
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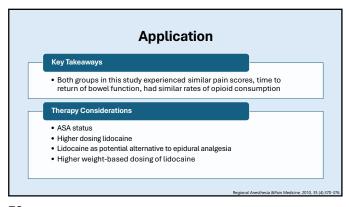
	Op. Day	POD 1	POD 2	POD 3	POD 4
Epidural group, mg	25 (11-50)	57 (27-100)	40 (7-74)	29 (12-89)	30 (18-87)
Lidocaine group, mg	17 (8-56)	48 (30-83)	23 (17-76)	20 (14-64)	7 (4-59)
P-value	0.884	0.961	0.883	0.657	0.111

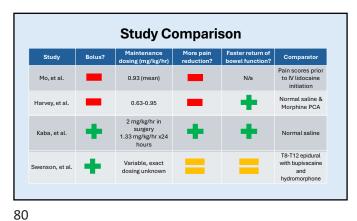
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Patient	ADR	Dose and serum level	Timing of level
Patient #1 (lidocaine group)	Facial paresthesia, perioral numbness, shortness of breath, and palpitations	Dose – Not noted  Serum level – <5 µg/mL	Not listed
Patient #2 (lidocaine group)	Hospital day 4 – disoriented with visual hallucinations Hospital day 7 – ventricular tachycardia that required cardioversion and placement of implantable cardioverter defibrillator	Dose - Not noted  Serum level – 6.5 µg/mL	Hospital da
Patient #3 (epidural group)	Atrial fibrillation	Dose - Not noted  Serum level – not collected	N/a

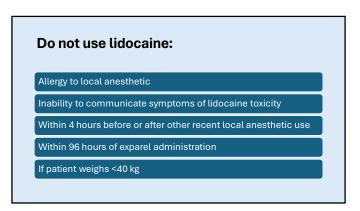


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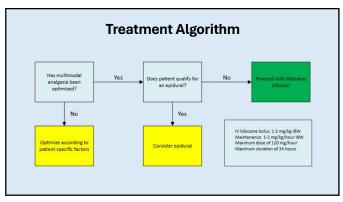




What to do with this information?

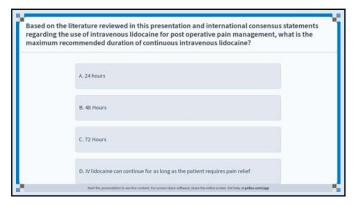


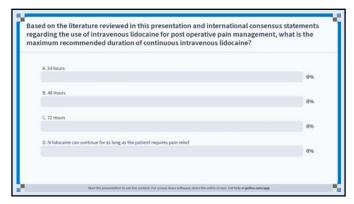
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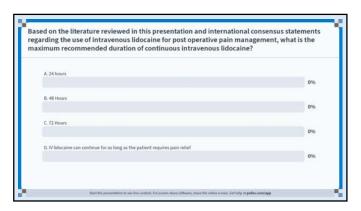


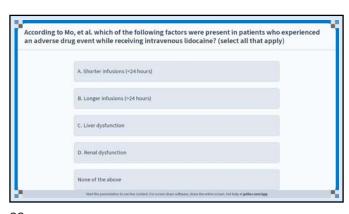
Post-Presentation Questions

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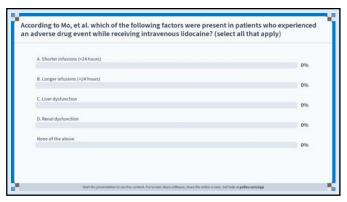


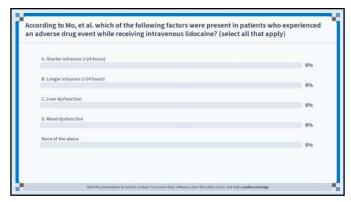






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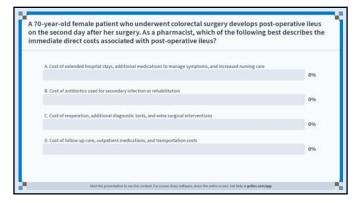
A 70-year-old female patient who underwent colorectal surgery develops post-operative ileus on the second day after her surgery. As a pharmacist, which of the following best describes the immediate direct costs associated with post-operative ileus?

A. Cost of extended hospital stays, additional medications to manage symptoms, ...

B. Cost of antibiotics used for secondary infection or rehabilitation

C. Cost of reoperation, additional diagnostic tests, and extra surgical interventions

D. Cost of follow-up care, outpatient medications, and transportation costs



91 92

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A Cost of extended hospital stays, additional medications to manage symptoms, and increased nursing care

O%

B. Cost of antibiotics used for secondary infection or rehabilitation

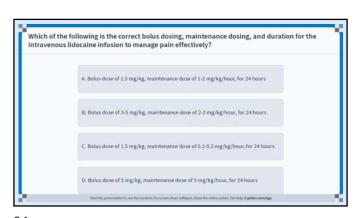
O%

C. Cost of reoperation, additional diagnostic tests, and extra surgical interventions

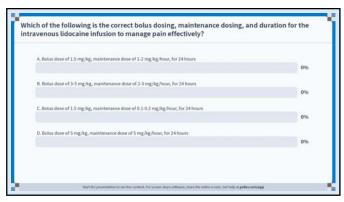
O%

D. Cost of follow-up care, outpatient medications, and transportation costs

O%



93 94





95 96

#### **Resources for Pharmacists**

- Chou R et al. Management of Postoperative Pain: A Clinical Practice Guideline From the American Pain Society, the American Society of Regional Anesthesia and Pain Medicine, and the American Society of Anesthesiologists' Committee on Regional Anesthesia, Executive Committee, and Administrative Council. J Pain. 2016 Feb;17(2):131-57. doi: 10.1016/j.jpain.2015.12.008. Erratum in: J Pain. 2016 Apr;17(4):508-10. doi: 10.1016/j.jpain.2016.02.002. Dosage error in article text. PMID: 26827847.

   Fool et al. The upper first transport of the state of the state
- Foo I, et al. The use of intravenous lidocaine for postoperative pain and recovery: international consensus statement on efficacy and safety. Anaesthesia. 2021;76(2):238-250. doi:10.1111/anae.15270

## **Special Thanks**

- Dr. Sarah Berman, PharmD, BCCCP o Faculty Mentor
- Dr. Kyllie Ryan-Hummel, PharmD, BCCCP
   o Critique

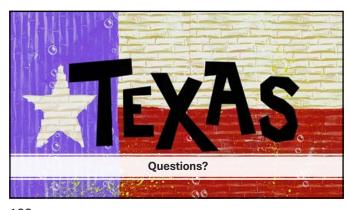


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# Co-Curricular Credit





99 100