

60th Annual Meeting

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Reframing Opioid Therapy: Buprenorphine's Role in Modern Pain Management

Joseph Cammilleri, Pharm.D., BCACP

UF Health Jacksonville
Ambulatory Care Clinical Pharmacist, Pain Management
PGY2 Pain and Palliative Care Program Director
Clinical Assistant Professor, University of Florida College of Pharmacy

1

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Disclosure

I do not have a vested interest in or affiliation with any corporate organization offering financial support or grant monies for this continuing education activity, or any affiliation with an organization whose philosophy could potentially bias my presentation (within the last 12 months).

2

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Presentation Objectives

Pharmacist:

- Outline the key features that make buprenorphine's pharmacology unique.
- List evidence-based therapeutic uses of buprenorphine.
- Evaluate the role of buprenorphine as a first-line opioid for chronic pain management.
- Formulate an acute pain management plan for patients maintained on buprenorphine.

Pharmacy Technician:

- Understand features that make buprenorphine's pharmacology unique.
- List evidence-based therapeutic uses of buprenorphine.
- Describe the role of buprenorphine as a first-line opioid for chronic pain management.
- Identify an acute pain management plan for patients maintained on buprenorphine.

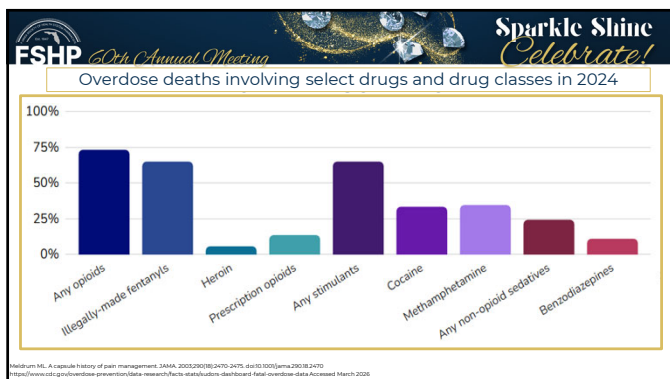
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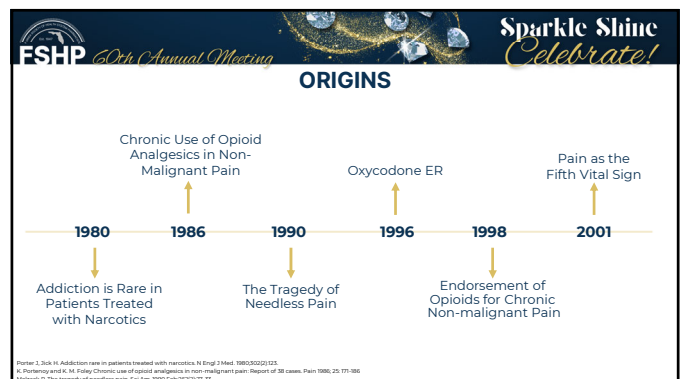
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THE OPIOID CRISIS

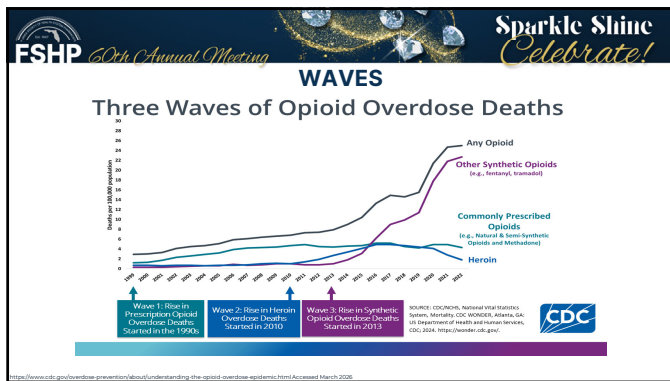
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6



7



8

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WHY IT MATTERS TO PHARMACISTS & TECHNICIANS

- Buprenorphine's unique pharmacological profile
- Rising prevalence of opioid use disorder (OUD)
- Pharmacists' expanding role in medications for opioid use disorder (MOUD)
- Acute pain management challenges
- Stigma reduction and patient advocacy

Image Generated with Gorki Imagine - March 30, 2020

9

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MECHANISM OF ACTION

Mu Receptor:
Partial Agonist

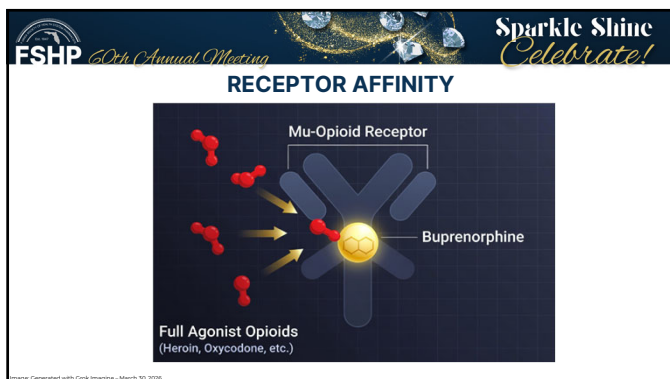
Kappa Receptor:
Full Antagonist

Delta Receptor:
Full Antagonist

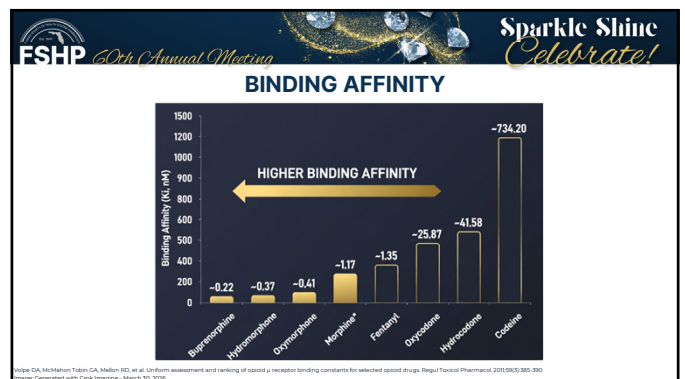
ORL-1 Receptor:
Partial Agonist

Codin L et al (2020). A Unique Opioid for the Treatment of Chronic Pain. Pain Ther. 2020(5):e1-e4.

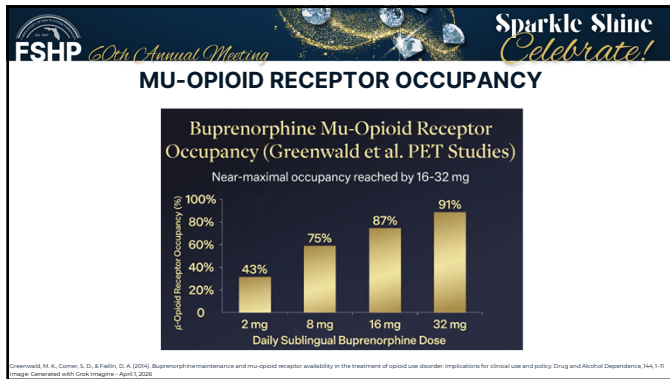
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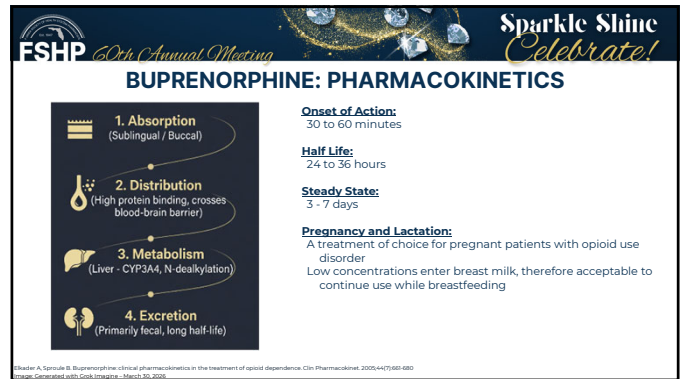
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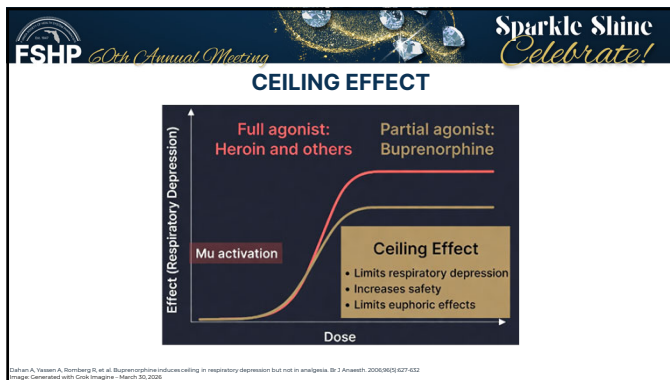
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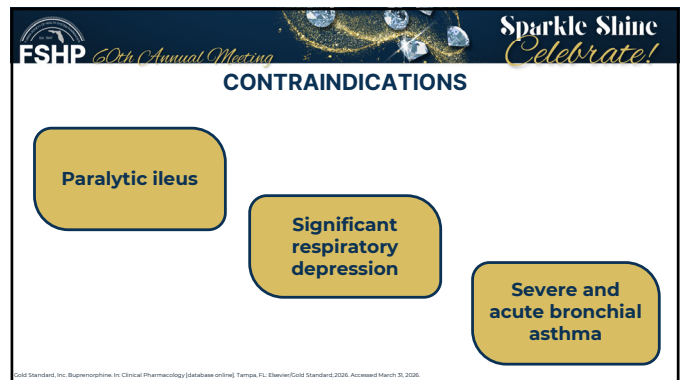
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14



15



16

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ADVERSE DRUG REACTIONS

Adverse Reaction	Frequency
Nausea	Very common
Headache	Very common
Dizziness	Common
Somnolence/Drowsiness	Common
Vomiting	Common
Constipation	Common
Rash (Transdermal patch)	Common
Dry mouth	Uncommon
Respiratory Depression	Uncommon
QT prolongation	Not significant
Dental problems	Rare to uncommon

Gold Standard, Inc. Buprenorphine. In: Clinical Pharmacology [database online]. Tampa, FL: Elsevier/Gold Standard 2020. Accessed March 30, 2020. Image Generated with Gink Imagine - April 3, 2020

17



18

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OTHER USES

Chronic pain in patients with OUD

- Lower risk of misuse compared to full opioid agonists
- Manage pain while treating OUD

Acute pain management

- Sublingual formulation
- Increase frequency
- In combination with multimodal therapy

Opioid withdrawal management

Connor R, Voonanath O, Sandakul A. Buprenorphine in StarParks [Internet]. Treasure Island (FL): StarParks Publishing; 2020. Item 5 [updated 2020; cited March 9, 2020].

19

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BUPRENORPHINE: PRODUCTS

Brand Name	Generic Name	Route	Indication	Formulation	Frequency
Butrans	Buprenorphine	Transdermal	Analgesia	Topical patch	Weekly
Belbuca	Buprenorphine	Transmucosal	Analgesia	Buccal film	Daily to Q12h
Buprenex	Buprenorphine	IV/IM	Analgesia	Solution for injection	Q6h prn
Suboxone	Buprenorphine/Naloxone	Sublingual	OUD	Film, Tablet	Daily to Q8h
Subutex	Buprenorphine	Sublingual	OUD	Tablet	Daily to Q8h
Zubsolv	Buprenorphine/Naloxone	Sublingual	OUD	Tablet	Daily to Q8h
Sublocade	Buprenorphine	Subcutaneous	OUD	ER solution for injection	Monthly
Brixadi	Buprenorphine	Subcutaneous	OUD	ER solution for injection	Weekly, Monthly

Gold Standard Inc. Buprenorphine. In: Clinical Pharmacology [database online]. Tampa, FL: Elsevier/Elsevier Standard; 2020. Accessed March 9, 2020.

20

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BUPRENORPHINE/NALOXONE

- Naloxone added to deter parenteral misuse
- Buprenorphine has a 10-fold greater binding affinity for the mu receptor
- TI/2 for naloxone in 30-40min vs Buprenorphine 24-60hrs
- Buprenorphine/Naloxone is misused
 - Injection
 - Snorting
- Combination may not be safer after discontinuation
- Naloxone may cause nausea, HA, anxiety

Blakes CK, Morrow JD. Reconsidering the Usefulness of Adding Naloxone to Buprenorphine. Front Psychiatry. 2020 Sep 15;11:549272.
 Wang E, Cummings C, Trevisan L, Finkel G. Buprenorphine alone or with naloxone (which is safer)? Psychopharmacol. 2020; 235:244-42.
 Buprenorphine/Naloxone. In: Clinical Pharmacology [database online]. Tampa, FL: Elsevier/Elsevier Standard; 2020. Accessed March 9, 2020.

21

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BUPRENORPHINE FOR PAIN MANAGEMENT

22

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BUPRENORPHINE TRANSDERMAL & TRANSMUCOSAL

PRODUCT	DOSING
Buprenorphine transdermal patch (Butrans®)	OME < 30: 5 mcg/hr patch OME 30-80: 10 mcg/hr patch OME > 80: use not recommended Max Dose: 20 mcg/hr patch
Buprenorphine buccal film (Belbuca®)	OME < 30: 75 mcg transmucosally once or twice daily OME 30 - 89: 150 mcg transmucosally twice daily OME 90 - 160: 300 mcg transmucosally twice daily OME > 160: use not recommended Max Dose: 900 mcg twice daily

OME=Oral morphine equivalent

Purdue Pharma, P. (2020). Butrans® (Buprenorphine) transdermal system, CII [Package insert]. U.S. Food and Drug Administration.

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BUPRENORPHINE TRANSDERMAL APPLICATION

- Apply to clean, dry, intact, relatively hairless skin
- Rotate sites with each new patch. Do not use the same site for at least 21 days
- Press firmly with the palm of your hand for 15 seconds. Do not rub
- Each patch is worn for 7 full days (change on the same day and time each week)
- Do not cut the patch

Purdue Pharma, P. (2020). Butrans® (Buprenorphine) transdermal system, CII [Package insert]. U.S. Food and Drug Administration.

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BUPRENORPHINE TRANSDERMAL COUNSELING



- Avoid heat exposure (heating pads, hot baths, saunas, electric blankets, sunbathing, fever)
- Do not apply to broken, irritated, or recently shaved skin
- Avoid lotions, oils, or soap on the site before applying
- May take up to 72 hours to feel the full pain-relief effect
- If the patch falls off, apply a new one and note the new change day
- Report severe application site reactions (redness, swelling, rash, blisters).
- Avoid driving or operating (may cause drowsiness or dizziness)
- Store used patches safely and dispose of them properly (fold sticky sides together and flush or use disposal program)

Purdue Pharma L.P. (2020). Butrans® (buprenorphine) transdermal system, CII [Package insert]. U.S. Food and Drug Administration. Image Generated with Ceph Imagina - April 1, 2020.

25

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BUPRENORPHINE TRANSDERMAL PEARLS



- Transition from short acting opioid to the patch
- OME 30-60 : Start at 10mcg patch and allow the same short acting opioid use for the first 3 days
- Nasal fluticasone sprayed under the patch may help with irritation/rash
- Works best in opioid naive patients
- OME >60 Patch doesn't seem to work well
- Short acting opioids for breakthrough pain effective

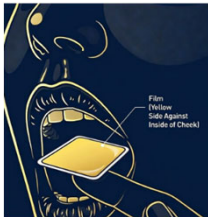
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26

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BUPRENORPHINE BUCCAL APPLICATION

- Wet the inside of your cheek with your tongue or rinse your mouth with water.
- Remove the film and place it on your fingertip with the yellow side facing up.
- Press the yellow side firmly against the inside of your moistened cheek (left or right).
- Hold it in place with your finger for 5 seconds.
- You can use a second finger on the outside of your cheek for better positioning.
- Leave the film in place until it completely dissolves (usually within 30 minutes).
- Do not touch, move, chew, or swallow the film.
- You may talk and swallow saliva, but do not eat or drink anything until it is fully dissolved.



Collegium Pharmaceuticals, Inc. (2020). BELBUCA® (buprenorphine) buccal film, CII [Package insert]. U.S. Food and Drug Administration. Image Generated with Ceph Imagina - April 1, 2020.

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BUPRENORPHINE BUCCAL COUNSELING



- Apply every 12 hours (twice daily), at the same times each day
- After the film dissolves: Rinse your mouth with water and swallow
- Wait at least 1 hour before brushing your teeth
- Do not apply to open sores, lesions, or irritated areas in the mouth
- If two films are used, apply the second film to the opposite cheek after the first one has been placed
- Avoid Eating or drinking until the film is completely dissolved

Collegium Pharmaceuticals, Inc. (2020). BELBUCA® (buprenorphine) buccal film, CII [Package insert]. U.S. Food and Drug Administration. Image Generated with Ceph Imagina - April 1, 2020.

28

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BUPRENORPHINE BUCCAL PEARLS



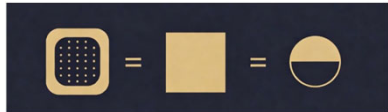
- Gets to higher doses than the transdermal patch
- Copay assistance available
- Short acting opioids for breakthrough pain effective
- If not "working" ensure proper use

Collegium Pharmaceuticals, Inc. (2020). BELBUCA® (buprenorphine) buccal film, CII [Package insert]. U.S. Food and Drug Administration. Image Generated with Ceph Imagina - April 1, 2020.

29

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BUPRENORPHINE DOSING EQUIVALENTS



- A 20 mcg/hour buprenorphine transdermal patch delivers approximately 0.48 mg of buprenorphine per day and is roughly equivalent to 1.5 – 2 mg per day of sublingual buprenorphine
- A 20 mcg/hour buprenorphine transdermal patch is roughly equivalent to 150 mcg of buprenorphine buccal film every 12 hours (total 300 mcg per day)

Collegium Pharmaceuticals, Inc. (2020). BELBUCA® (buprenorphine) buccal film, CII [Package insert]. U.S. Food and Drug Administration. Purdue Pharma L.P. (2020). Butrans® (buprenorphine) transdermal system, CII [Package insert]. U.S. Food and Drug Administration. Image Generated with Ceph Imagina - April 1, 2020.

30

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SPLITTING BUPRENORPHINE

- Package insert explicitly states that the film/tablets should not be cut.
 - Concern that the active ingredients may not be uniformly distributed throughout the film.
- Cutting films or tablets is common practice
- Reindel KL, et al. An Exploratory Study of Suboxone (Buprenorphine/Naloxone) Film Splitting: Cutting Methods, Content Uniformity, and Stability. J Am Pharm Assoc. 2019;59(4):548-554.
 - Content uniformity remained acceptable = 1/2 film had 50% of drug



Reindel KL, et al. An Exploratory Study of Suboxone (Buprenorphine/Naloxone) Film Splitting: Cutting Methods, Content Uniformity, and Stability. J Am Pharm Assoc. 2019;59(4):548-554. Content Uniformity and Stability. April 3, 2019.

31

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BUPRENORPHINE/NALOXONE FOR CHRONIC PAIN MANAGEMENT

32

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ACUTE PAIN IN PATIENTS USING SUBLINGUAL BUPRENORPHINE



Mild to moderate pain

- Continue usual buprenorphine dose
- May split it into q6-8h dosing for better analgesic coverage
- Non-pharmacologic: ice, elevation, positioning, relaxation techniques, splinting
- Optimize non-opioids
 - NSAIDs, gabapentinoids, APAP
- Consider temporarily increasing total daily buprenorphine
 - 24-32 mg/day in divided doses

Shen L, et al. Buprenorphine management in the perioperative period: educational review and recommendations from a multisociety expert panel. Reg Anesth Pain Med. 2021;46(2):840-859. Content Uniformity and Stability. April 3, 2019.

33

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ACUTE PAIN IN PATIENTS USING SUBLINGUAL BUPRENORPHINE

Moderate to severe pain

- Continue buprenorphine at the home dose
- May split it into q6-8h dosing for better analgesic coverage
- For high baseline doses (>16 mg/day)
 - Consider a temporary reduction to 8-16 mg/day
- Optimize non-opioids and multimodal
- Use full mu-opioids
 - High-affinity (hydromorphone, fentanyl)
 - Likely need higher doses



Shen L, et al. Buprenorphine management in the perioperative period: educational review and recommendations from a multisociety expert panel. Reg Anesth Pain Med. 2021;46(2):840-859. Content Uniformity and Stability. April 3, 2019.

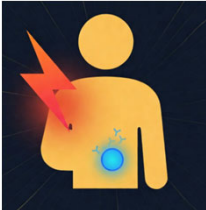
34

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ACUTE PAIN IN PATIENTS USING LONG-ACTING BUPRENORPHINE

Moderate to severe pain

- Continue buprenorphine
- Schedule elective surgery for end of dose or change to SL prior to surgery
- Optimize non-opioids and multimodal
- Use full mu-opioids
 - High-affinity (hydromorphone, fentanyl)
 - Likely need higher doses



Shen L, et al. Buprenorphine management in the perioperative period: educational review and recommendations from a multisociety expert panel. Reg Anesth Pain Med. 2021;46(2):840-859. Content Uniformity and Stability. April 3, 2019.

35

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STIGMA REDUCTION

- Buprenorphine is legitimate medicine for a chronic condition
- People on stable on buprenorphine can work, parent, and live fulling lives
- Common myth: Buprenorphine just replaces one addiction with another.
 - Buprenorphine helps stabilize brain chemistry safely
- People often develop OUD after an injury that led to prescribed opioids



Wendler, Jennifer B. "Suboxone National Survey: Misconceptions." The Opioid Journal vol. 16 no. 1, Spring 2018, pp. 23-28. PINE.

36

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KEY TAKE HOME POINTS

Buprenorphine is safer than full opioid agonist

Buprenorphine is indicated for opioid naïve patients and is considered a first line option in patients who require opioid therapy

For acute pain in patients taking buprenorphine: continue buprenorphine, optimize non-opioid, multi modal therapy, and use high affinity full opioids if needed

37

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Questions?

38

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Reframing Opioid Therapy: Buprenorphine's Role in Modern Pain Management

Joseph Cammilleri, Pharm.D., BCACP
joseph.cammilleri@ufhealth.org

Thank you!

39