

Buprenorphine in Palliative Care

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Conflicts of Interest

- None of the presenters have a conflict of interest to disclose



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Objectives

- Discuss ways to incorporate buprenorphine into the treatment of cancer related pain or pain secondary to serious illness in patients at risk for or with substance use disorder
- Develop methods to educate clinicians on the appropriate use of buprenorphine to treat both pain and substance use disorder
- Propose ideas to decrease stigma related to the buprenorphine in those with serious illness



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Key Abbreviations

- BUP: buprenorphine (for the purposes of this presentation, this includes formulations, with or without naloxone, FDA approved for the treatment of Opioid Use Disorder)
- MEDD: Morphine Equivalent Daily Dose
- MOUD: Medication for Opioid Use Disorder
- OUD: Opioid Use Disorder
- SUD: Substance Use Disorder
- SL: sublingual
- HPM: hospice and palliative medicine
- IDT: interdisciplinary team



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Case

- 58 y.o. woman admitted to the hospital with respiratory distress, was intubated and during hospitalization diagnosed with small cell lung cancer with extensive osseous metastases
- PMH: Intravenous drug use (IVDU) – heroin; lumbar abscess s/p laminectomy; chronic low back pain
- SOC: lives in boarding house; TOB use, IVDU heroin; twin daughters with active IVDU, one hospitalized at same time with endocarditis



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Case (continued)

- While hospitalized on tapering methadone for treatment of substance use disorder and pain
- Due to perceived intolerance to methadone, hospitalist rotated to Oxycodone ER and IR
- Discharged with #60 Oxycodone ER 30 mg and #60 oxycodone/APAP 5/325 mg
- Follow-up scheduled for palliative medicine to manage pain



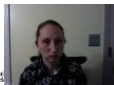
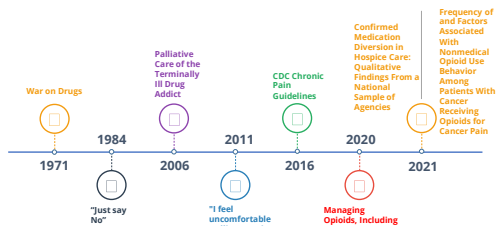
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Case (continued)

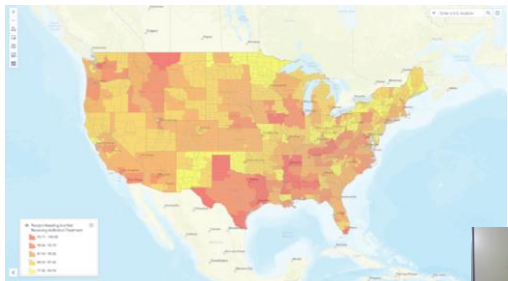
- Initial palliative visit– offered Suboxone (buprenorphine) – refused
- Harm reduction approach
 - Prescribed only Oxycodone ER
 - No breakthrough opioids
 - Weekly prescriptions
- Continued to use heroin/fentanyl
- WHAT WOULD YOU DO??

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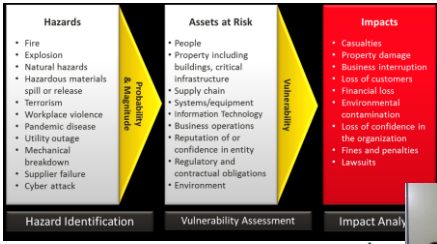


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Risk Assessment



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Risk Mitigation and Harm Reduction

- Mitigation strategies aim to reduce possibility of damage from hazards
- Harm reduction strategies aim to reduce the extent of damage from the hazards



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Risk Assessment for SUDs

RELATIONSHIPS

Risk factors

Family members who use drugs or alcohol

Family history of mental illness

Adverse childhood experiences

Protective factors

Family involvement

COMMUNITIES

Risk factors

Neighborhood poverty

Neighborhood violence

Protective factors

Availability of faith-based resources

Community activities

SOCIETY

Risk factors

Norms and Laws favorable to substance use

Racism

Lack of economic opportunity

Protective factors

Hate crime laws

Policies limiting availability of



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Screening Tools for Current SUDs

- ADULT SUBSTANCE USE SURVEY (ASUS)
- ALCOHOL USE DISORDERS OBSERVATION TEST (AUDIT)
- CASE, CASE-AD MODIFICATION
- CRAFT
- DRUG ABUSE SCREENING TEST (DAST)
- DRUG USE SCREENING INVENTORY-REVISED (DUSI-R)
- MICHIGAN ALCOHOL SCREENING TEST (MAST)
- MASSACHUSETTS YOUTH SCREENING INSTRUMENT-VERSION 2 (MAYSI-2)
- PERSONAL EXPERIENCE SCREENING QUESTIONNAIRE (PESQ)
- RAPID ALCOHOL PROBLEMS SCREEN (BAPAS)
- SUBSTANCE ABUSE SURTLE SCREENING INVENTORY, 3RD EDITION (SASSI-3)
- SUBSTANCE ABUSE SURTLE SCREENING INVENTORY-ADOLESCENT (SASSI-A2)
- TRIAGE ASSESSMENT FOR ADDICTIVE DISORDERS (TAAAD)
- TUD-100 (TUD-100) SCREENING TEST (TUD)
- TREAT
- COMPOSITE INTERNATIONAL DIAGNOSTIC INTERVIEW VERSION 3.1 (CIDI-V3.1)
- UNCOPE
- COMPREHENSIVE ADDICTION AND PSYCHOLOGICAL EVALUATION (CAAPE)
- GLOBAL APPRAISAL OF INDIVIDUAL NEEDS-INITIAL (GAINI)
- STRUCTURED CLINICAL INTERVIEW FOR DSM (SCID)
- LEVEL OF CARE INDEX-2 REVISED (LCOI-2R) FOR ADULTS
- ADDICTION SEVERITY INDEX (ASI) AND TREATMENT SERVICE REVIEW (TSR)
- FAMILY ASSESSMENT FORM (FAF)
- GLOBAL APPRAISAL OF INDIVIDUAL NEEDS-MRO (GAINI-MRO)
- LEVEL OF CARE INDEX-2 REVISED (LCOI-2R)
- RECOVERY ATTITUDE AND TREATMENT EVALUATION (RAATE)
- RISK INVENTORY FOR SUBSTANCE-AFFECTED FAMILIES
- STRUCTURED DECISION-MAKING (SDM) FAMILY AND CHILDREN ASSESSMENT



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Tools for Risk of Problematic Opioid Use

- ORT, ORT-R = for future development of OUD on opioids
- SOAPP, SOAPP-R, SOAPP-SF
- COMM

Review Article

Assessment tools for problematic opioid use in palliative care: A scoping review

Jenny Lau^{1,2,3,4}, Paolo Mazzotta^{5,6}, Rouhi Fazelzad⁷, Suzanne Ryan^{8,9}, Alissa Tedesco⁸, Andrew J. Smith⁸, Abhimanyu Sud⁸, Andrea D. Farlan^{10,11} and Camilla Zimmermann^{12,13}

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 https://doi.org/10.1177/08980101211041289

SAGE



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Risk Mitigation for SUD

- | | | |
|--|---|---|
| <p>Universal preventive interventions</p> <p>Designed to reach entire groups or populations.</p> <p><i>Examples:</i> might target schools, whole communities, or workplaces</p> | <p>Selective interventions</p> <p>Target biological, psychological, or social risk factors that are more prominent among high-risk groups</p> <p><i>Examples:</i> include prevention education or peer support groups for adults</p> | <p>Indicated preventive interventions</p> <p>Target individuals who show signs of being at risk for a substance use disorder.</p> <p><i>Examples:</i> support services for patients who violate drug policies or screening and consultation for patients with alcohol-relate</p> |
|--|---|---|



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Harm Reduction in SUD

- OEND = Overdose Education and Naloxone Distribution
- Fentanyl test strips
- Never Use Alone
- National Suicide Prevention Hotline
- Safer injection practices
- Safe Syringe Programs
- PrEP = pre-exposure prophylaxis anti-retroviral medication



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Harm Reduction in Pain Management

- Co-prescribing laxatives
- Ambulatory devices
- Continuous Positive Airway Pressure
- Opioid Overdose Education and Naloxone Distribution (OEND)
- National Suicide Hotline
- **BUPRENORPHINE!!!**



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Systemic Bias versus Implicit Bias

- Systemic Biases are institutional.
- Implicit Biases are individual.



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Bup Systemic Bias

- Insurance Companies
- Lack of education regarding substance use disorders
- Lack of education regarding buprenorphine
- Media coverage
- Regulations
- Political climate = dichotomous thinking



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Bup Implicit Bias

- Patient education
- Provider education
- Population knowledge/experience



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Stigma and Bias

- Known factor in healthcare disparities
- Words matter!

Avoid	Try Instead
Abuser, addict	Person with an opioid use disorder
Abuse, misuse	Substance use, Non-medical use
Lapse, relapse, slip	Recurrence of symptoms
Clean (regarding person)	In remission or recovery
Clean & Dirty (regarding urine drug test results)	Negative or positive (test results)
Medication Assisted Treatment (MAT), Opioid Replacement Therapy (ORT), Opioid maintenance therapy, Opioid substitution therapy	Medication for addiction treatment, Medication for opioid use disorder



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Addictionary®, Recovery Research Institute.
 Ashford RB et al. Drug Alcohol Depend. 2016; 169:131-8.
 Kelly JL et al. JAMA. 2016; 315(12):1202-7.
 FitzGerald C et al. BMC Med Ethics. 2017; 18(1):19.

Language Matters in Pain Management

legitimate pain	pain, discomfort, suffering
drug seeking	symptoms of OUD
be liberal	titrate dose
be lenient	working with patient
patient reports	patient describes, patient experiences



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Reframing the Pain & Palliative Conversation

You all never had pain like mine.	You don't feel heard.
Are you calling me a liar?	I will help you with what you tell me.
Don't call me an addict.	You're concerned about becoming dependent on these medications.
Just make me comfortable.	Let's focus on you as a person focusing on the disease and treatments that are not as helpful anymore.



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Pharmacology & Physiology



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Buprenorphine Preparations*

FDA Indication	Preparation	Method of Administration	Patient Population (Opioid naïve or tolerant)
Pain	Belbuca®	Buccal	Naïve
	Butrans®	Transdermal Patch	
	Buprenex®	Intravenous	Naïve (tolerant in specific circumstances)
Medication for Opioid Use Disorder (MOUD) Buprenorphine/ Naloxone	Suboxone®	Sublingual	Tolerant
	Zubsolv®		
	Bunavail®	Buccal	

*Excludes all long-acting injections/implants for MOUD

Information based on associated package inserts



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The Conundrum

- 1) BUP for MOUD
 - Most widely utilized of MOUD medications
 - Complex pharmacology
- 2) Difficulty of acute pain management in opioid tolerant patients



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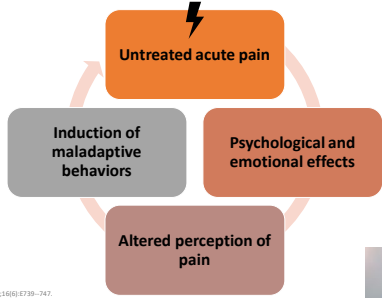
Physiologic Changes from Chronic Opioid Exposure

- Hyperalgesia
 - ↑ sensitivity to pain
 - ↓ pain tolerance
- Opioid Induced Hyperalgesia (OIH)
- Neuroplasticity effects on long-term tolerance in OUD
- Endocrinopathies
 - Dysregulation of stress response
 - Sex hormone deficiency

De Andrade, L, et al. Pain Manag. 2015;5(3):167-173.
Makris, PC. Drugs. 2016;80(2):19-25.
Zhou Y, et al. J Pain Behav. 2016;23:237-251.



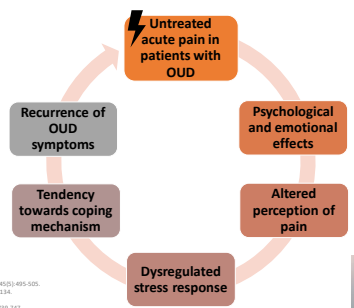
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Bounev V, et al. Pain Physician. 2013;16(6):739-747.



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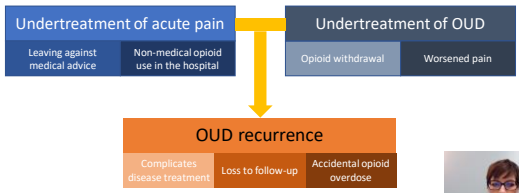


Washford A, et al. J Drug Alcohol Abuse. 2019;45(1):45-50.
 Alford DP, et al. Am J Addict Med. 2008;14(2):117-134.
 of Pain M. Psychosom Med. 2018;30(2):1-9.
 Bounev V, et al. Pain Physician. 2013;16(6):739-747.



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Risks of Undertreatment



Herscher M, et al. Med Clin North Am. 2020;104(4):695-708



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Common Misconceptions

- MOUD provides analgesia
- Use of opioids for analgesia may result in addiction recurrence
- Full agonist opioids used “on top” of BUP won’t provide analgesia
- Reporting pain → manipulation to obtain opioids, drug seeking, manifestation of worsening addiction
- Naloxone in Suboxone® blunts effects of opioids for analgesia

Alford DP. *Ann Intern Med*. 2006;144(2):127-134.
 Haber LA et al. *J Hosp Med*. 2019;14:e23-9.
 Hencher M et al. *Med Clin North Am*. 2020;104(4):695-708.
 Lembke A et al. *Pharm Med*. 2019;20(1):42-8.



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Buprenorphine Pharmacology & Clinical Implications

- Naloxone absorption
- Partial agonist
 - ↓ peak effect
 - ↓ adverse effects
- Affinity for mu opioid receptor
 - Precipitated withdrawal not relevant when continuing therapy

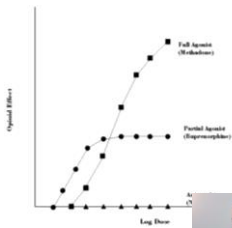


Figure 2-1, TIPS 40, SAMHSA 2004.

Haber LA. *J Hosp Med*. 2019;14:e33-5.
 van Nieuwenhuijzen IC. *Drug Rev (Stuttg)*. 2016;66(11):562-570.



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Buprenorphine Pharmacology & Clinical Implications

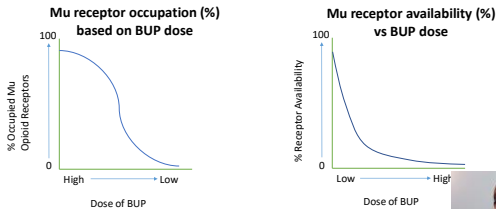


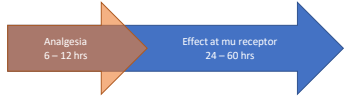
Figure (left) adapted from Figure 1A. *Drug Alcohol Depend*. 2014;144:3-11. Figure (right) adapted from Figure 1. Warner NS. *Mayo Clin Proc*. 2003 Jun;78(6):658-663.



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Buprenorphine Pharmacology & Clinical Implications

- Potency
 - 16 mg SL BUP ~480 MEDD
- SL elimination half-life
 - Duration of analgesia vs OUD treatment



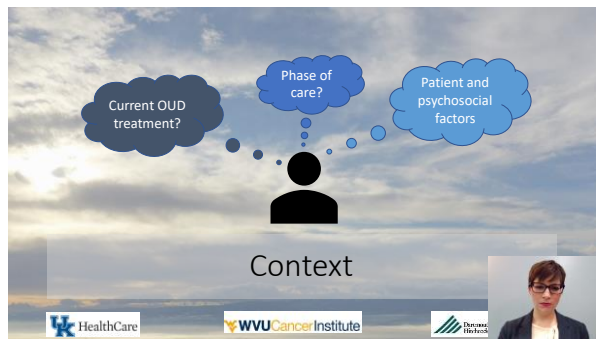
Hercher M. Med Clin North Am. 2020;104(4):695-708
Children WJ. Stat Sci and Comput. #211. 2020.



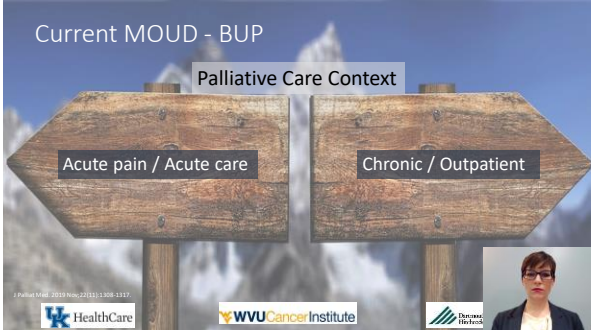
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Advantages of Continuing BUP

- Don't have to cover large MEDD deficit
- Evidence to support BUP + IR opioids achieving analgesia
- Avoid risk of DIScontinuity in care
- Avoid re-induction complexities
- Decrease risk of OUD relapse and accidental overdose



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Advantages of Continuing BUP

- Reduced incidence of hyperalgesia
- Schedule 3 controlled substance
 - Fewer prescribing restrictions
- Safety in renal failure/dialysis?
 - Renal elimination of active metabolites
- Neuropathic pain?



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Scenario - Stopping BUP for MOUD

- Underlying concept - improved OPIOID analgesia
 - Other meds/modalities can and should be used to treat pain in an opioid tolerant patient
- Baseline MEDD must be covered
 - 16 mg SL BUP ~480 MEDD
 - Concerns for withdrawal & craving
 - Analgesia not possible until baseline requirements met
- Prescribed full agonist opioids WITHOUT BUP exposes patient to trigger in a tenuous and vulnerable state

Waltcher M. Add Clin North Am. 2020;104(4):695-708



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Active OUD – Not on MOUD

- Evaluate Total Pain and optimize non-opioid options
 - Opioid-sparing modalities
 - Address psychological dimensions
- Define expectations, goals of therapy, and frequent monitoring
- Consider long-acting therapies and abuse-deterrent formulations
- Collaborate with addiction specialist, when possible
 - Consider methadone or **BUP/NAL** in divided dose

J Palliat Med. 2019 Nov;23(11):1308-1317. Am Soc Clin Oncol Educ Book. 2019 Jan;39:24-35.



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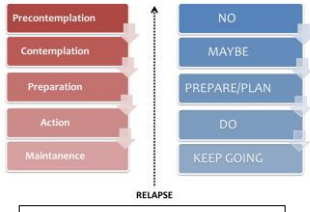
An Intervention: Palliative Care Buprenorphine Education

- Background:
- 13% of palliative providers reported having DATA waiver to prescribe buprenorphine as treatment for opioid use disorder.
 - 21% of HPM Fellows reported they were satisfied with how they treat symptoms in patients with opioid misuse and/or substance use disorder.
 - 41% of HPM Fellows reported feeling trained to manage opioid misuse.
 - Palliative providers reported least comfort in managing patients with non-opioid substance use disorder



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Transtheoretical Model Stages of change



https://commons.wikimedia.org/wiki/File:Transtheoretical_Model_-_Stages_of_change.jpg

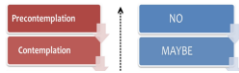


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Buprenorphine in Palliative Care

"I don't DO addiction! I don't DO chronic pain!"
BUT.....

- 7% hospice discharge
 - FOR CAUSE
 - STABILITY
- Upstream palliative care
- Diversion and OUD
- OUD/SUD prevalence in patient population



MAYBE.....

- I might need to diagnose OUD
- I might need more tools in the toolbox



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Buprenorphine in Palliative Care

Where do I start?

EDUCATE MYSELF on long term opioids, chronic pain management, opioid use disorder

- Opioid Misuse = Nonmedical opioid use = Unhealthy Opioid use OR OUD
- Multimodal Pain Management
- Interdisciplinary Addiction Care
- Neurobiologic similarities between complex opioid dependence and mild OUD
- **BUPRENORPHINE???**

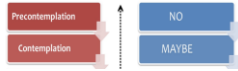


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Buprenorphine in Palliative Care

EDUCATE OTHERS

- 1. engage in conversations
- 2. dispel myths
- 3. decrease moral distress
- 4. garner support



- interdisciplinary team
- referring teams
- patients and caregivers



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HPM Bup Education

Intervention:

- Regular meetings of HPM, pain management, and addiction medicine clinicians to discuss challenging palliative care cases
- Two DATA waiver training live sessions for providers in central KY

Aims:

- Increase HPM provider awareness of opioid use disorder, chronic pain, and interventional pain treatment options
- Provide DATA waiver training on buprenorphine for opioid use disorder to HPM providers.



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Buprenorphine in Palliative Care

Plan the Logistics:

- Diagnoses Inclusion
- Prescribing Team
- Prior Authorizations & Administrative Team
- Peer Network for entire IDT (psychosocial and behavioral included!!!)



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HPM Bup Education

Results from survey study of HPM clinicians:

- Increased comfort and feeling supported in managing patients with substance use
- Increase comfort in talking about goals of care and substance use with patients
- Increased likelihood of referring to interventional pain and nonpharmacologic pain modalities
- Increase in considering buprenorphine as an option

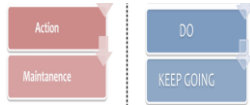


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Buprenorphine in Palliative Care

Implementation & Maintenance:

- Revisit policy and procedures as necessary
- Share experiences
- Peer Network

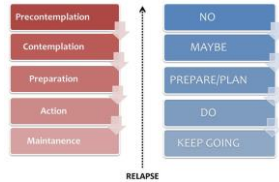


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Buprenorphine in Palliative Care

Next steps:

- Share more cases!!!
- Expand the hospice & palliative literature on buprenorphine
- Decrease systemic barriers and bias
- Encourage DATA waiver in HPM training
- Expand the palliative buprenorphine peer network



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Case (continued)

- Admitted for 3 days of chemotherapy due need for IV access
- Went through withdrawal due to concurrent use of heroin/fentanyl in addition to Oxycodone ER
- Started on buprenorphine/naloxone
- Remained on buprenorphine with good pain control
- Occasionally used methamphetamine and cocaine but there was no evidence of recurrence of opioid use



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Case (conclusion)

- Developed brain metastases, neurologic decline and after brief ICU stay, transitioned to comfort focused care
- Daughters both entered treatment for substance use disorder and were able to spend quality time with her
- Transferred to hospice center for end of life care and was surrounded by family and friends



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Take Home Messages

- Buprenorphine can be a useful medication for use in those with serious illness for both pain and substance use disorder.
- Education and collaboration with colleagues can both promote the better treatment of pain and substance use.



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Questions?

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- Monika Holbein - monika.holbein@hsc.wvu.edu
- Katrina Nickels - katrina.nickels@uky.edu



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