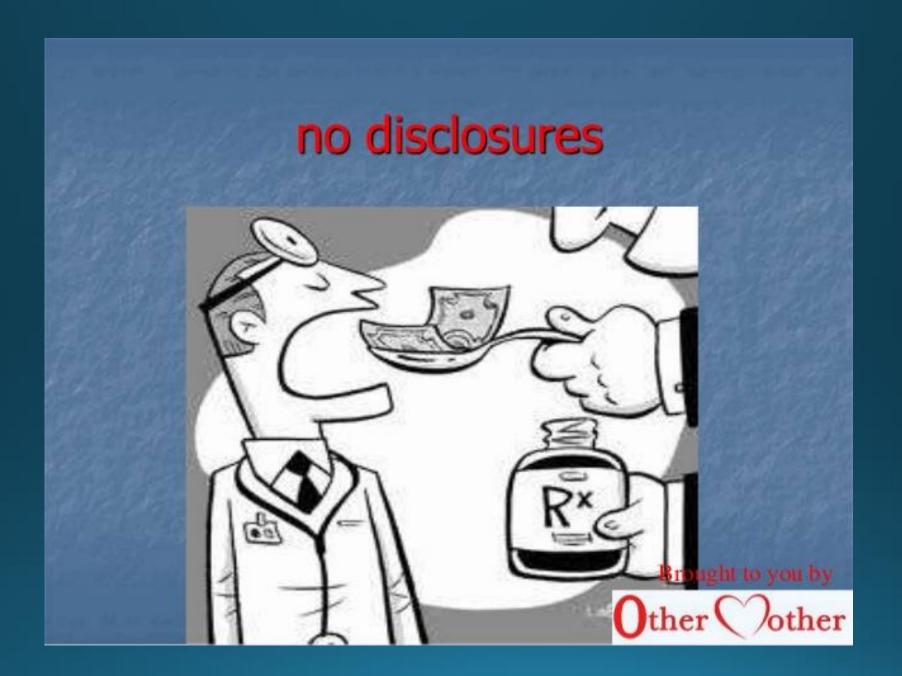
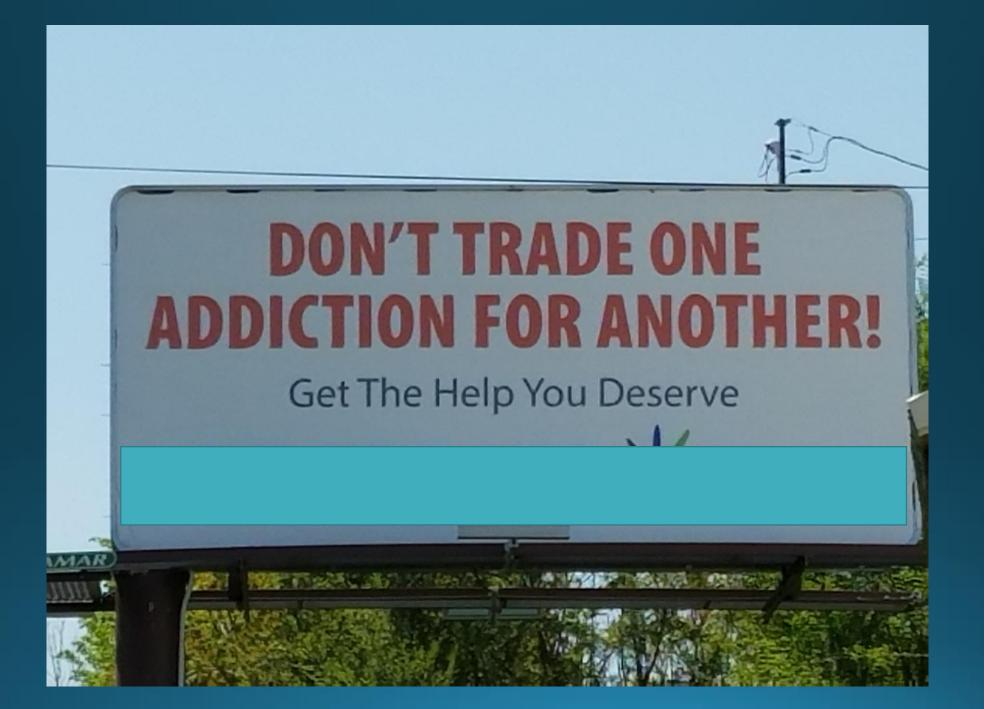
MEDICATION ASSISTED TREATMENT: An Essential Tool for Managing our **Opioid Epidemic** American College of Osteopathic Internists May 11, 2019 James H. Berry D.O. **WVUMedicine**





"The stigma surrounding the use of pharmacotherapy, in particular opioid agonist therapy, is arguably more potent and harmful than general stigma about the disease right now. Stigmatized beliefs that medication is simply a 'replacement addiction' are false and quite literally killing people."

- Dr. Richard Saitz

Chair and Professor of Community Health Sciences Boston University School of Public Health

OVERVIEW

- Epidemic
- Disease
- MAT
- Answer questions



OVERDOSE EPIDEMIC

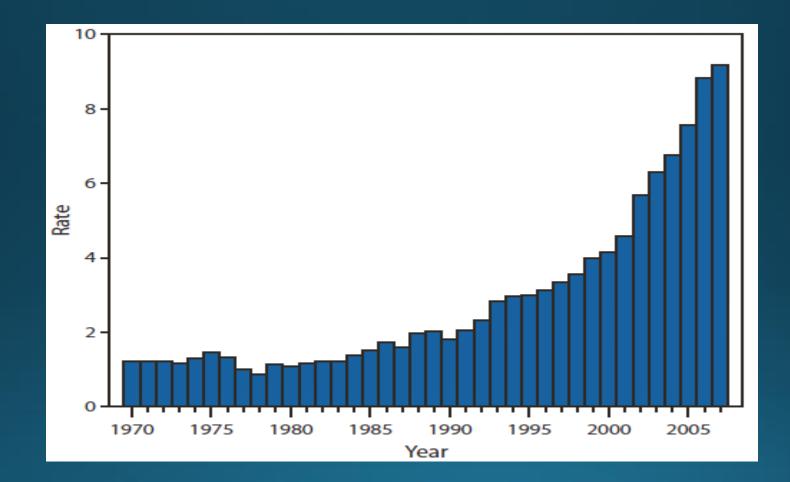
Leading cause of injury death in US

• 70, 237 deaths in 2017 • 47,600 (68%) opioids 345% 2001-2016

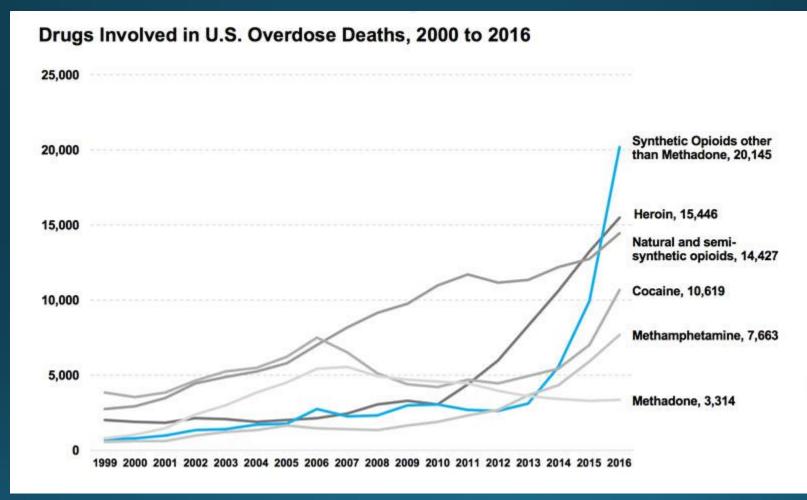
702, 568 deaths 1999 – 2017
56.8% opioids



UNINTENTIONAL OVERDOSE DEATHS



Centers for Disease Control and Prevention. CDC grand rounds: prescription drug overdoses—a US epidemic. *MMWR Morb Mortal Wkly Rep*. 2012;61(1):10-13



https://www.drugabuse.gov/related-topics/trendsstatistics/overdose-death-rates

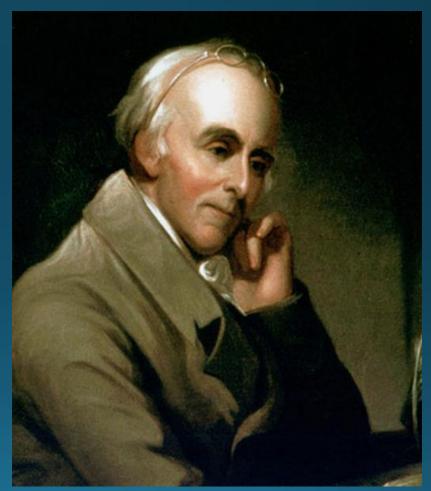
2016-2025: Projected Opioid Death Toll



Pitt AL, et al. Am J Public Health. 2018;108(10):1394-1400.

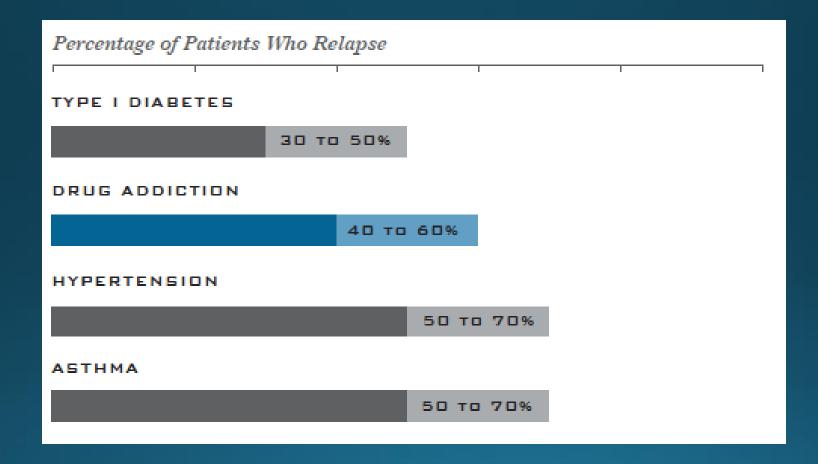
DISEASE

"My observations authorize me to say, that persons who have been addicted to them, should abstain from them suddenly and entirely. 'Taste not, handle not, touch not' should be inscribed upon every vessel that contains spirits in the house of a man, who wishes to be cured of habits of intemperance, ...habitual drunkenness should be regarded not as a bad habit but as a disease."



Dr. Benjamin Rush

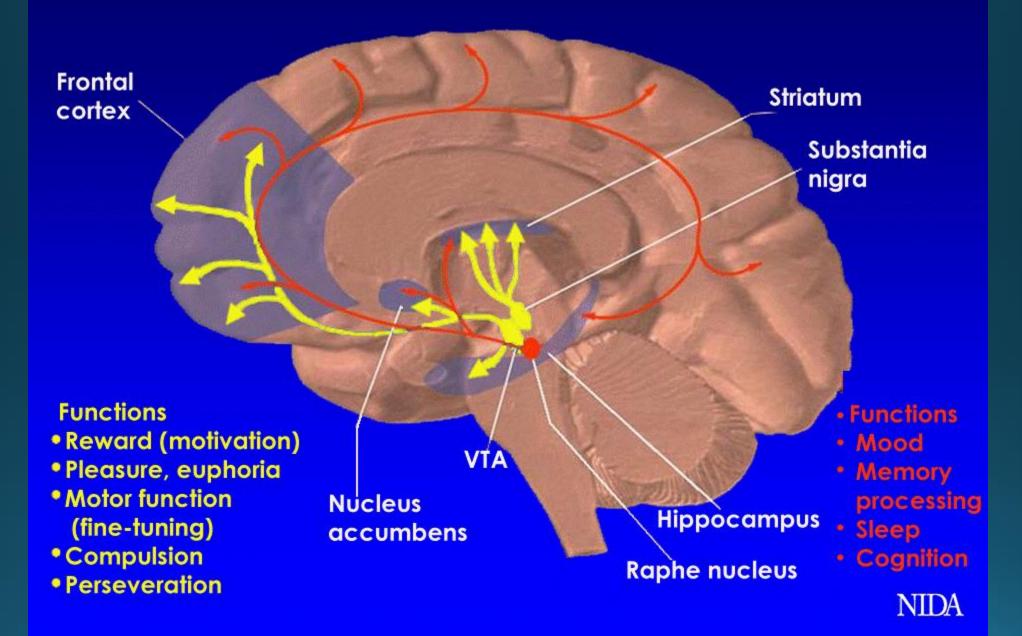
CHRONIC DISEASE





Dopamine Pathways

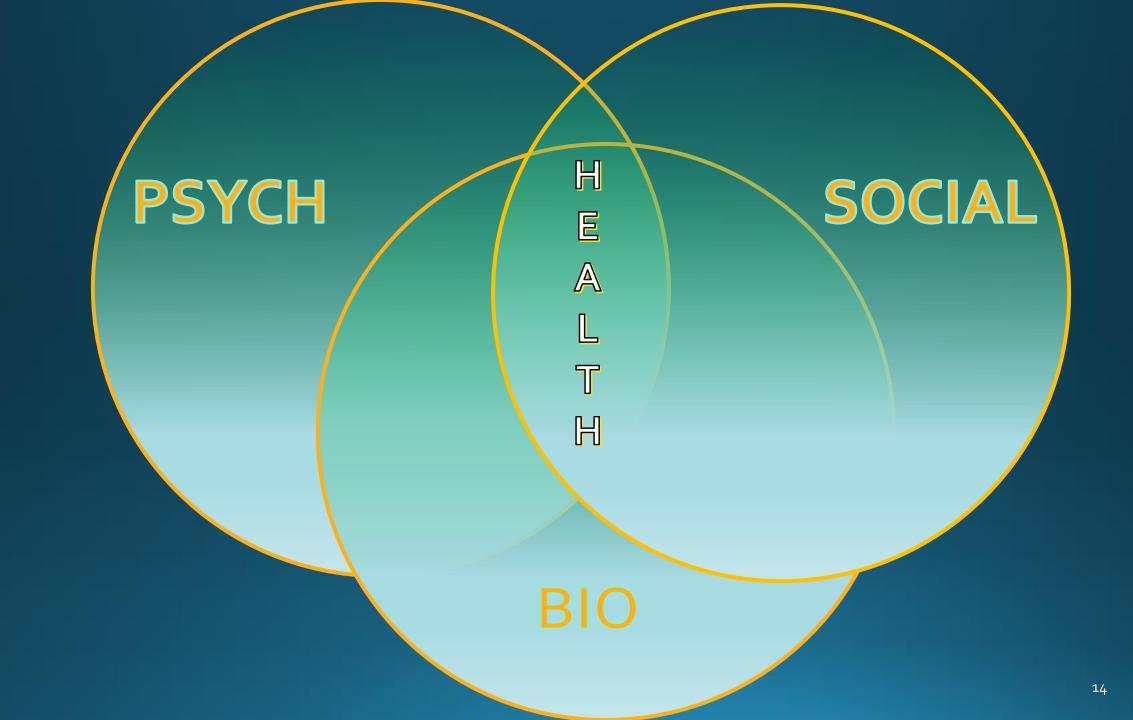
Serotonin Pathways



12

DISEASE

- <u>Addiction</u> is a primary, chronic disease of brain reward, motivation, memory and related circuitry. It is characterized by one or more of the following (ABCDE):
 - Inability to consistently abstain
 - Impairment in **behavioral** control
 - Craving
 - **Diminished** recognition of problems
 - Dysfunctional emotional response
- Cycle of relapse and remission
- This disease is often progressive and fatal
- Manage vs. Cure
- Two main goals:
 - Keep alive
 - Increase quality of life



TREATMENT

BIO

"Abstinence-Based" Vs. Medication Assisted Treatment

<u>PSYCHO</u>

Cognitive Behavioral Therapy Motivational Enhancement Therapy Contingency Management

<u>SOCIO</u>

12 Step Groups Family Therapy Spiritual Community

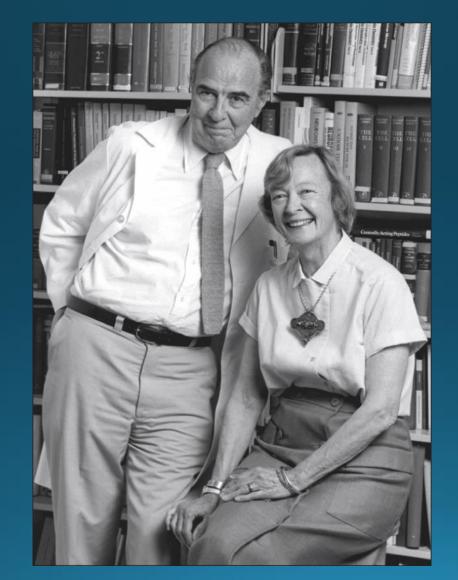
Medication Assisted Treatment (MAT) for Opioid Addiction



RECEPTORS

- Methadone full agonist
- Buprenorphine partial agonist
- Naltrexone antagonist

METHADONE



METHADONE

- Schedule II
- Use is restricted to non-residential narcotic treatment facilities:
 Opioid Treatment Programs (OTP) = Methadone Clinics
- Pain is the only legal medical indication in the office setting
- Tricky to start and stop
- Stigma

NALTREXONE





Implants (Not FDA approved)

Tablets

Injection

BUPRENORPHINE

- DATA 2000-can prescribe a schedule III-V medication
- Detoxification and maintenance
- In 2002- FDA approves Bup for office management of opioid addiction
- Docs need 8hr course & apply to SAMHSA for a waiver from the 1970's Controlled Substance Act
- 2005 lifted the 30 patient cap to 100 patients
- 2016 lifted the 100 patient cap to 275 patients
- 2018 NPs and PAs allowed to treat 30 patients

BUPRENORPHINE







BUP + NALOXONE Suboxone Zubsolv Cassipa 16mg

BUP

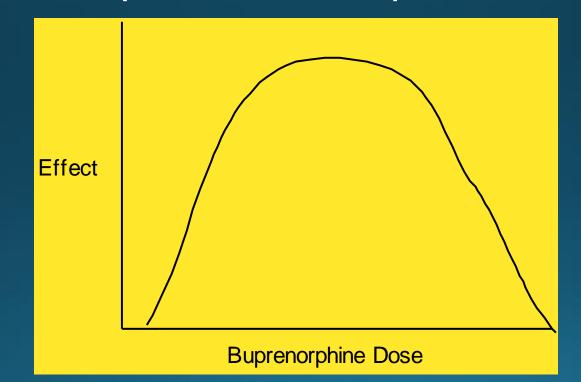
"Subutex" Probuphine (6m implant) Sublocade (1m injection)

FILM Suboxone

Bunavail

Buprenorphine Pharmacology

Bell-Shaped Dose Response Curve



Drug and Alcohol Dependence, 2003;70:S61

Buprenorphine/Naloxone



MAT

Methadone (Full Agonist)

Pros:

Stigma

Long lasting Decades of evidence Helps with pain **Cons:** Diverted Inconvenient Lethal in overdose Withdrawal Buprenorphine (Partial Agonist)

Pros:

Doctor's office Less risk of overdose Less risk of IV use Helps with pain Cons: Diverted Withdrawal Naltrexone (Antagonist)

Pros

Not diverted No risk of overdose Cons: Compliance

Pain

Medication Assisted Treatment

- 2.4 million Americans *diagnosed* with Opioid Use Disorder. ~80% do not receive treatment!2
- MAT is a safe and effective strategy for decreasing the frequency and quantity of opioid use and reducing the risk of overdose and death._{3,4}
- Although MAT has significant evidence to support it as an effective treatment, it remains highly underutilized.⁵

1.U.S. Department of Health and Human Services, Substance Abuse and Mental Health Services Administration. (2016). Key substance use and mental health indicators in the United States: Results from the 2015 National Survey on Drug Use and Health (HHS Publication No. SMA 16-4984, NSDUH Series H-51). Rockville, MD

2. U.S. Department of Health and Human Services, Substance Abuse and Mental Health Services Administration. (2016). Prescription drug use and misuse in the United States: Results from the 2015 National Survey on Drug Use and Health. Rockville, MD.

3. Kresina TF, Lubran RL. Improving public health through access to and utilization of medication assisted treatment. Int J Environ Res Public Health. 2011;8:4102-4117

4. National Institutes on Drug Abuse. Cost effectiveness of drug treatment. Retrieved from: http://www.drugabuse.gov/publications/teaching-packets/understanding-drug-abuse-addiction/sectioniv/6-costeffectiveness-drug-treatment

5. Volkow ND1, Frieden TR, Hyde PS, Cha SS. Medication-assisted therapies--tackling the opioid-overdose epidemic. N Engl J Med. 2014 May 29;370(22):2063-6. doi: 10.1056/NEJMp1402780. Epub 2014 Apr 23.







National Institute on Drug Abuse Advancing Addiction Science





FACING ADDICTION IN AMERICA

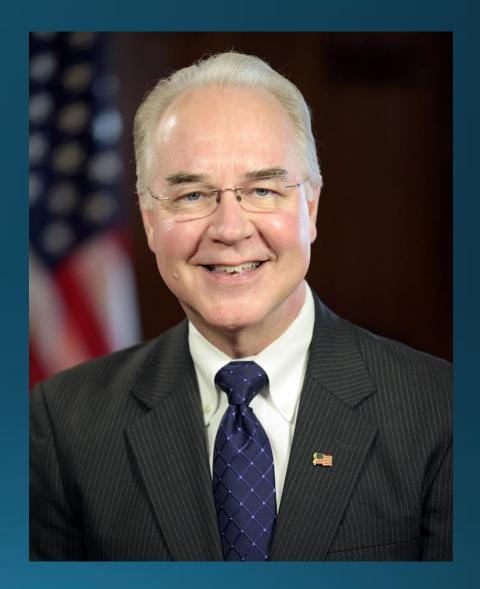
The Surgeon General's Report on Alcohol, Drugs, and Health

U.S. Department of Health & Human Services



U.S. Health and Human Services Secretary Tom Price

May 9, 2017



"No state has been hit harder by the opioid epidemic than West Virginia. More than 860 residents fatally overdosed on drugs in 2016. That's a record number...If this were any other disease, we'd have all the resources come to bear to figure out what the infection is, how did it get into our county and what are we going to do about it..." "This is a public health issue. This isn't a criminal justice issue..."

> http://www.wvgazettemail.com/news-health/20170509/trump-officialsseek-opioid-solutions-in-wv#sthash.aSj2bgfH.dpuf

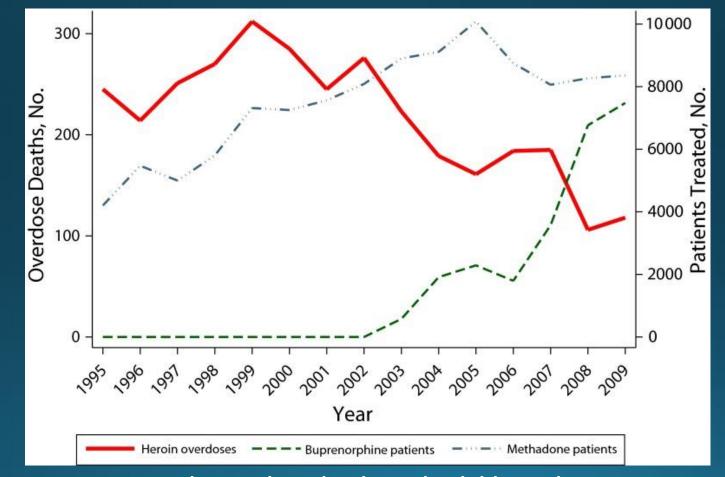
Asked about drug treatment options, Price touted faith-based programs while showing less support for medication-assisted programs in which addicts are weaned off heroin with other opioids like Suboxone and methadone.

"If we're just substituting one opioid for another, we're not moving the dial much...Folks need to be cured so they can be productive members of society and realize their dreams."

http://www.wvgazettemail.com/news-health/20170509/trump-officialsseek-opioid-solutions-in-wv#sthash.aSj2bgfH.dpuf

What Are The Benefits of MAT?

DECREASE OVERDOSE



Heroin overdose deaths and opioid agonist treatment: Baltimore, MD, 1995–2009

Schwartz, RP.Am J Public Health. 2013 May; 103(5): 917–922

Annals of Internal Medicine

ORIGINAL RESEARCH

Medication for Opioid Use Disorder After Nonfatal Opioid Overdose and Association With Mortality

A Cohort Study

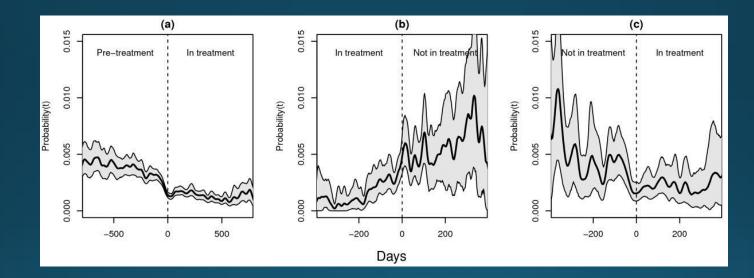
Marc R. Larochelle, MD, MPH; Dana Bernson, MPH; Thomas Land, PhD; Thomas J. Stopka, PhD, MHS; Na Wang, MA; Ziming Xuan, ScD, SM; Sarah M. Bagley, MD, MSc; Jane M. Liebschutz, MD, MPH; and Alexander Y. Walley, MD, MSc

Ann Intern Med. August 2018;169:137–145.

- 17, 568 overdose survivors over 12 months
- Methadone or Buprenorphine associated with reduced mortality
- Only 30% received MAT

DECREASE CRIMINALITY

National cohort study of all patients (n = 3221) in OMT in Norway 1997-2003



Day-to-day rates of criminal convictions. Day-to-day rates of criminal convictions (95% Cl) for all patients (a) entering OMT for the first time, (b) leaving OMT and (c) re-entering OMT. a: X-axis; days before and days after first treatment entry (n=3221). Y-axis; the day-to-day rate of criminal convictions. b: X-axis; days before and days after treatment drop out (n=1175). Y-axis; the day-to-day rate of criminal convictions. c: X-axis; days before and days before and days after treatment re-entry (n=515). Y-axis; the day-to-day rate of criminal convictions.

DECREASE HIV INFECTIONS

Study

North America

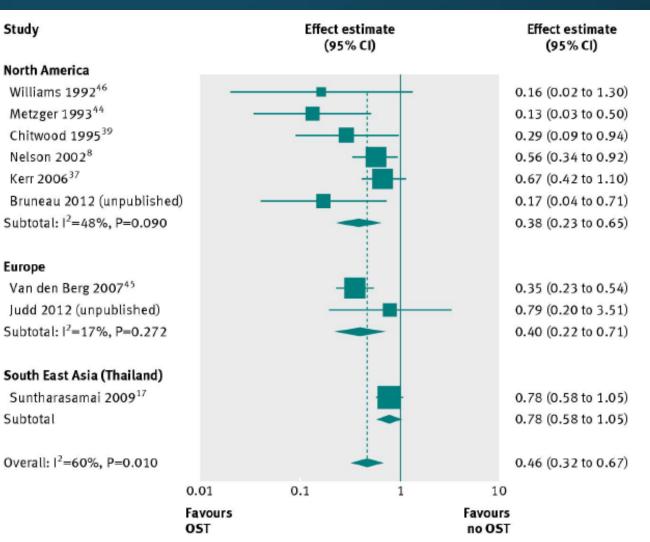
Nelson 2002⁸

Kerr 200637

Europe

Subtotal

- Meta-analysis
- 9 studies
- Overall:
 - 54% reduction
- North America:
 - 62% reduction

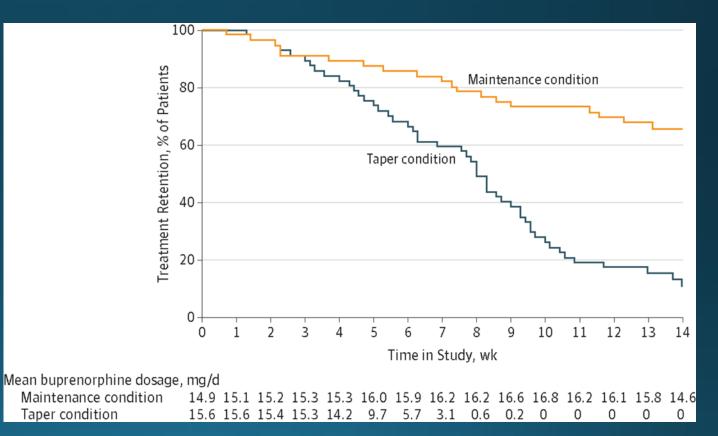


Gowing et al. BMJ 2012

INCREASE TREATMENT RETENTION

Completion of 14 week trial: 11% taper vs 66% maintenance

Mean percentage of urine negative for opioids: 35% taper vs 53% maintenance



Fiellin DA et al. JAMA Intern Med 2014

Decrease Health Care Costs

- Annual societal cost associated OUD₁:
 - \$55 Billion
 - Lost work productivity (46%)
 - Health Care (45%)
 - Criminal Justice (9%)
 - Hidden cost of comorbid conditions ~8X greater health care than non-abuser
- B-MAT more cost effective than:
 - No treatment₂
 - Detox₃
- Immediate access to OAT = \$78,257 in savings. lifetime savings = \$3.8 billion.
- B-MAT Non-Adherent Cost₄:
 - **\$27k** greater medical charges
 - **\$22k** greater overall healthcare charges



1.H.G. Birnbaum, A.G. White, M. Schiller, T. Waldman, J.M. Cleveland, C.L. Roland Societal costs of prescription opioid abuse, dependence, and misuse in the United States Pain Medicine, 12 (2011), pp. 657–667

2.B.R. Schackman, et. al. Cost-effectiveness of long-term outpatient buprenorphine-naloxone treatment for opioid dependence in primary care. Journal of General Internal Medicine, 27 (2012), pp. 669–676 3. D. Polsky, et.al.Cost-effectiveness of extended buprenorphine-naloxone treatment for opioid-dependent

youth: Data from a randomized trialAddiction, 105 (2010), pp. 1616–1624 4.Tkacz, J, et.al. Relationship between buprenorphine adherence and health service utilization and costs among opioid dependent patients.Journal of Substance Abuse Treatment (2013).

5.Krebs E, Enns B, Evans E, et al. <u>Cost-effectiveness of publicly funded treatment of aboid use</u>

disorder in California [published online November 21, 2017]. Ann Intern Med.

TOOL







BUPRENORPHINE DIVERSION

- Majority report using for therapeutic purposes
 - 97%- decrease cravings
 - 90%- prevent withdrawal
 - 29%- save \$
- Illicit use decreased with access to treatment
- 8-25% have used to get high (rates decrease over time)

Bazazi AR, Yokell M, Fu JJ, Rich JD, Zaller ND. Illicit use of buprenorphine/naloxone among injecting and noninjecting opioid users. J Addict Med. 2011;5(3):175-180.
Schuman-Olivier Z, Albanese M, Nelson SE, et al. Self-treatment: illicit buprenorphine use by opioid-dependent treatment seekers. J Subst Abuse Treat. 2010;39(1):41-50.
Cicero TJ, Surratt HL, Inciardi J. Use and misuse of buprenorphine in the management of opioid addiction. J Opioid Manag. 2007;3(6):302-308.
Cicero TJ, Ellis MS, Surratt HL, Kurtz SP. Factors contributing to the rise of buprenorphine misuse: 2008-2013. Drug Alcohol Depend. 2014;142:98-104.



SICK PEOPLE COME TO HOSPITALS

- Always open
- SUD very common and often severe
- IV use
- Early Medical Consequences
- Opioid Toxicity and Withdrawal

OPIOID TOXICITY

- Influenced by purity, loss of tolerance, alcohol/sedative mix
- clouded consciousness to coma
- Severe respiratory depression
- Constricted pupils
- Pulmonary edema
- Severe hypotension, cardiovascular collapse (hypoxia may lead to dilated pupils)
- Reversed by naloxone (narcan)



WITHDRAWAL

Typical sx's (flu-like and leaky):

- nausea/vomiting,
- cramps (abd & muscular),
- sweating,
- goose bumps/piloerection,
 rhinorrhea, diarrhea, mydriasis
- Insomnia
- fever

• Supportive care:

- Bentyl 10mg qid for diarrhea
- Immodium 4mg; 1-2 q hr prn NTE 8qd
- Motrin 600mg one q 6 prn bone pain
- Compazine 10mg TID prn nausea
- Benadryl 25mg two q 6 prn nasal cong.
- Clonidine 0.1 to 0.2mg q hour prn; up to 1.2MQ
- Buprenorphine 2mg



OPIATE WITHDRAWAL

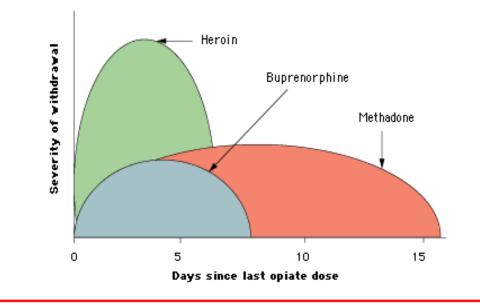
Heroin

-short 1/2 life

-4-6h after last use

Methadone

-long ½ life -24-36h after last use



Severity of opioid-withdrawal symptoms after abrupt discontinuation of equivalent doses of heroin, buprenorphine, and methadone Peak withdrawal symptoms are most severe after discontinuation of heroin. Such symptoms last longest with methadone, which has a somewhat later peak of severity. Buprenorphine has milder peak withdrawal symptoms than does methadone; the duration of symptoms is intermediate between those for methadone and those for heroin. Reproduced with permission from: Kosten, TR, O'Connor, PG. Management of drug and alcohol withdrawal. N. Engl J Med 2003; 348:1786. Copyright © 2003 Massachusetts Medical Society.

Opiate WD: Measurement

- Clinical Opiate Withdrawal Scale (COWS)
 - 11 item
 - Mild: 5-12
 - Moderate: 13-24
 - Moderately Severe: 25-36
 - Severe: >36

72-hour rule

Title 21, Code of Federal Regulations, Part 1306.07(b)

Allows to administer (but not prescribe) narcotic drugs for the purpose of relieving acute withdrawal symptoms while arranging for the patient's referral for treatment

- Not more than 1-day's medication may be administered or given to a patient at one time
- Patient must return to ED each day for no more than 72 hours
- This 72-hour period cannot be renewed or extended.



People choose to use, shouldn't we let them suffer the consequences?

Isn't MAT just a "crutch"?

Aren't you just trading one drug for another?

How long should one stay on MAT?

How about pregnancy?

SUMMARY

- The US has an addiction crisis and Appalachia is on the front lines
- Addiction is a disease that is treatable
- MAT is an evidence based tool that works
- Quality of life and extension of life are the ultimate goals
- We desperately need more clinicians willing to treat, more hospitals willing to support and fewer barriers to implementation
- Have to start where patients are

