

# Treatment of Opioid Use

Jason Beaman, D.O., M.S., M.P.H., FAPA  
Chair, Psychiatry & Behavioral Sciences  
Assistant Clinical Professor

Board Certified:

Forensic Psychiatry  
Psychiatry  
Family Medicine  
Addiction Medicine



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# Disclosures

None



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# Objectives

- Understand different opioid conditions
- Understand the treatment of the above
- Understand new emerging trends in treatment



- Overdose
- Withdrawal
- Opioid Use Disorder



# Opioid Overdose Triad

- Triad of symptoms defines the common signs/symptoms of opioid overdose syndrome:
  - Altered mental status
  - Depressed respirations
  - Miotic pupils
- This triad has a sensitivity of 92% and a specificity of 76%
- Differential:
  - Hypoglycemia
  - Acidemia
  - Complications from end-stage liver disease



# Naloxone

- Displaces other opioids bound to mu opioid receptors allowing respiration to take place
- Binds to mu opioid receptors but does not activate the receptors
- Begins working in 1-3 minutes
- Effects last 30-90 minutes in duration
- Not addictive and will not cause overdose
- Can cause withdrawal in some individuals: nausea, vomiting, chills, muscle discomfort, confusion



# Naloxone for Opioid Overdose

- Naloxone blockade is short-lived versus opioid agonist systemic duration
- Monitor short-acting opioid overdose patients for at least 12 hours
- Monitor methadone overdose patients for 24-48 hours
- May not work as well for buprenorphine or fentanyl overdose



# Opioid Withdrawal

- Results from immediate or rapid cessation of opioids when an individual is physiologically dependent
- Unlikely to produce severe morbidity and mortality
- CNS depression that occurred during use is replaced with CNS noradrenergic neuron hyperactivity in the locus coeruleus
  - Effects can be reversed with clonidine, lofexidine, and opioids





# Opioid Withdrawal

- Increase in Gamma-Aminobutyric Acid (GABA) effects and decrease in dopamine release from the nucleus accumbens cause dysphoria, depression, and cravings
  - Effects can be reversed only with opioids
- Onset of withdrawal symptoms depends on half-life of used drug
  - Heroin, Morphine: 4-6 hours
  - Hydrocodone, Oxycodone, Hydromorphone, Oxymorphone: 8-12 hours
  - Methadone, Buprenorphine: 24-36 hours



# Spontaneous Opioid Withdrawal Timeframes

Drug	Begin Withdrawal	Peak	Lasts
Morphine Heroin Oxycodone Hydrocodone	6-12 Hours After Last Use	36-72 Hours	Approximately 5 Days
Methadone	36-72 Hours After Last Use  *May be as short as 20 hours for rapid metabolizers	96-144 Hours	Several Weeks in Duration

# Signs/Symptoms of Opioid Withdrawal

- Tachycardia
- Hypertension
- Restlessness
- Irritability
- Insomnia
- Opioid craving
- Pupillary dilation
- Lacrimation
- Rhinorrhea
- Piloerection
- Yawning
- Sneezing
- Mild anorexia
- Nausea
- Vomiting
- Diarrhea
- Muscle cramping/pain
- Anxiety



# Clinical Opiate Withdrawal Scale (COWS)

- Clinical staff administered tool that evaluates 11 symptoms/signs of opioid withdrawal based on intensity
  - Pulse rate, sweating, restlessness, pupil size, bone/joint aches, runny nose or tearing, GI upset, tremor yawning, anxiety/irritability, gooseflesh
- Evaluates severity withdrawal based on those 11 domains
  - Score 5-12: mild withdrawal
  - Score 13-24: moderate withdrawal
  - Score 25-36 moderately severe withdrawal
  - Scores greater than 36: severe withdrawal



# Dangers of Opioid Withdrawal



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- Behavioral alone
- Medication Assisted Treatment alone
- Combination of the above



# Behavioral Treatment

- Motivational Interviewing
- Individual psychotherapy
- Relapse prevention
- 12 Step groups



# Motivational Interviewing

- Engages patient in a non-confrontational explorative conversation
- Evaluates pros and cons
- Primary modality for exploring change





# Individual Psychotherapy

- May help to explore underlying personality traits that can contribute to use
- Explore relationship problems to help increase insight and support



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# Relapse Prevention

- Recognition of people, places and things
- Relapse is a natural process of recovery
- If relapse occurs:
  - Increase treatment
  - Minimize extent and length of use



# 12 Step Groups

- Varying evidence of efficacy
- Some individuals find it helpful
- Requires some basic religious acceptance
- Located around the world



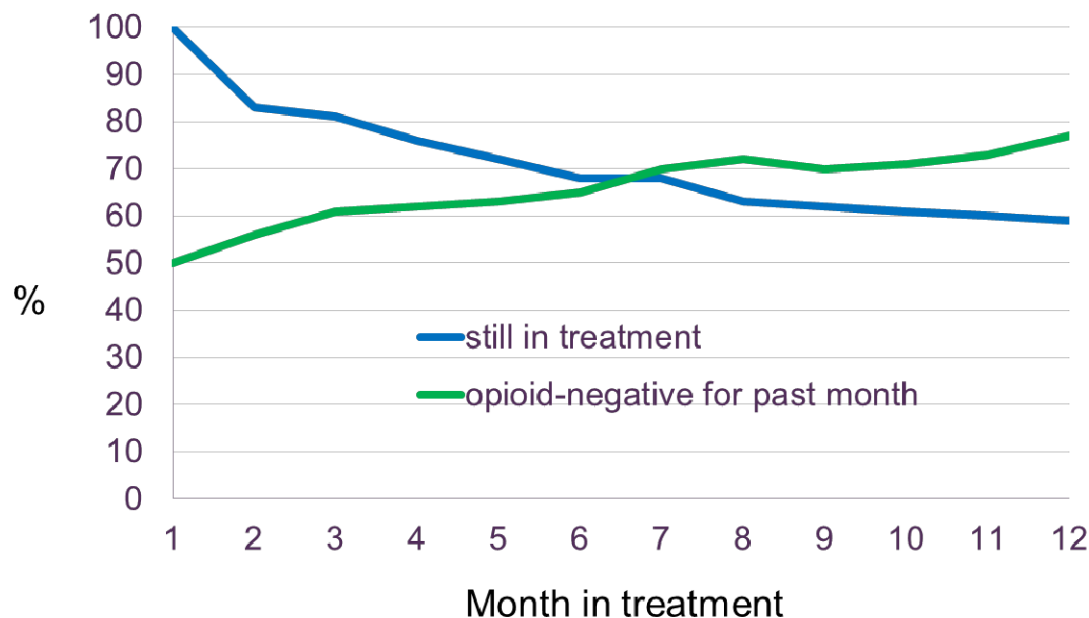
# Medication Assisted Treatment

- Methadone
- Buprenorphine
- Naltrexone



# Treatment Retention and Decreased Illicit Opioid Use on MAT

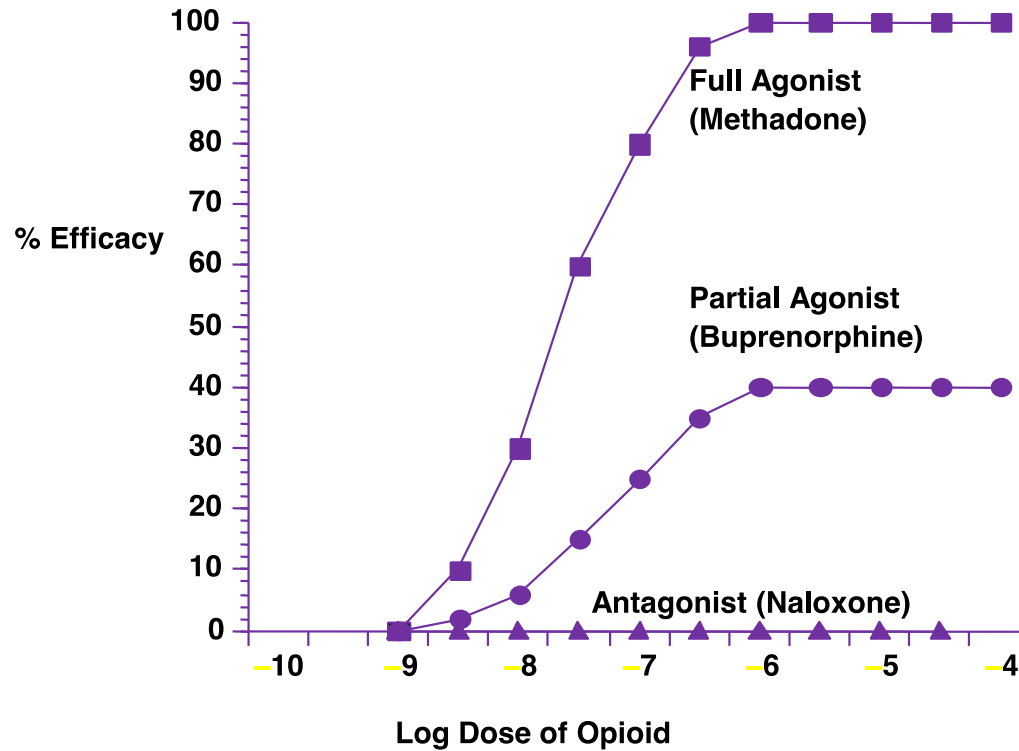
- Buprenorphine promotes retention, and those who remain in treatment become more likely over time to abstain from other opioids



Kakko et al, 2003  
Soeffing et al., 2009



# Opioid Ligand Pharmacology



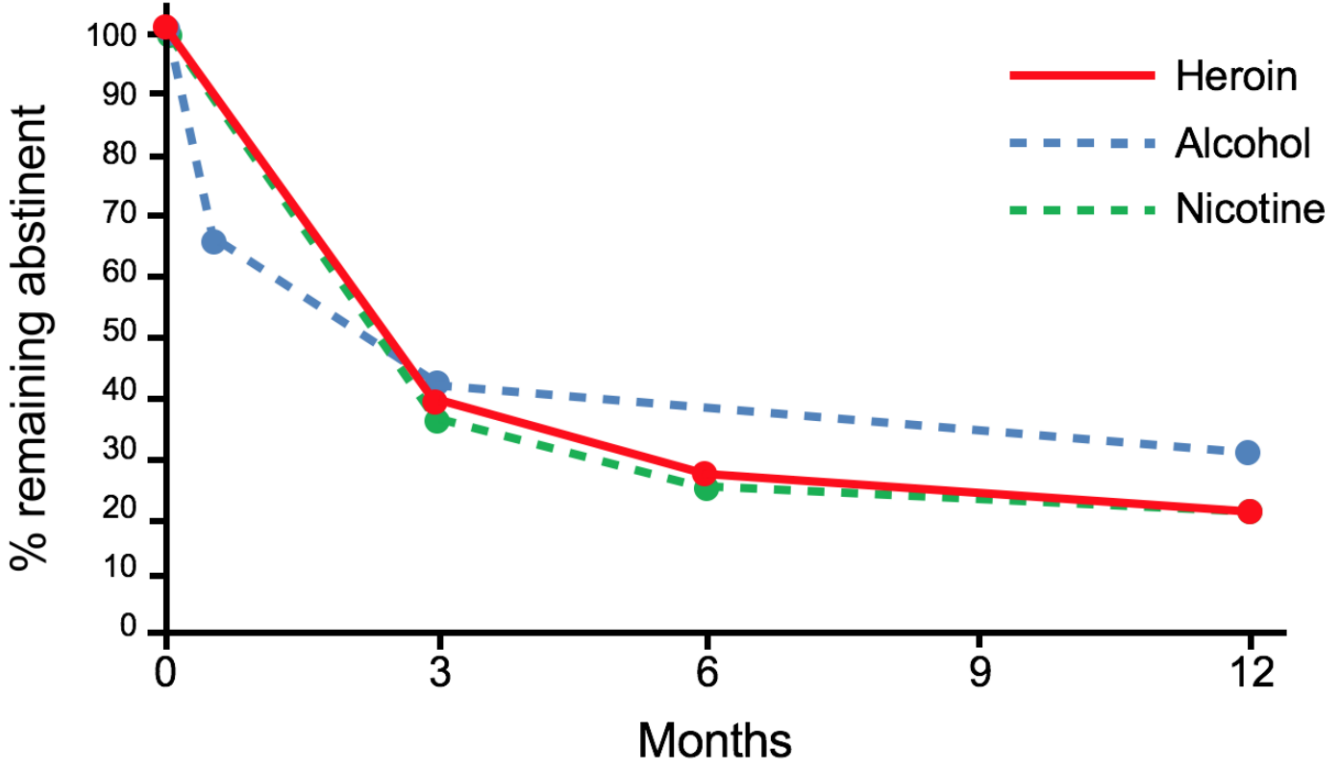
SAMHSA, 2018  
Orman & Keating, 2009



Providers  
Clinical Support  
System



# Abstinence Without MAT

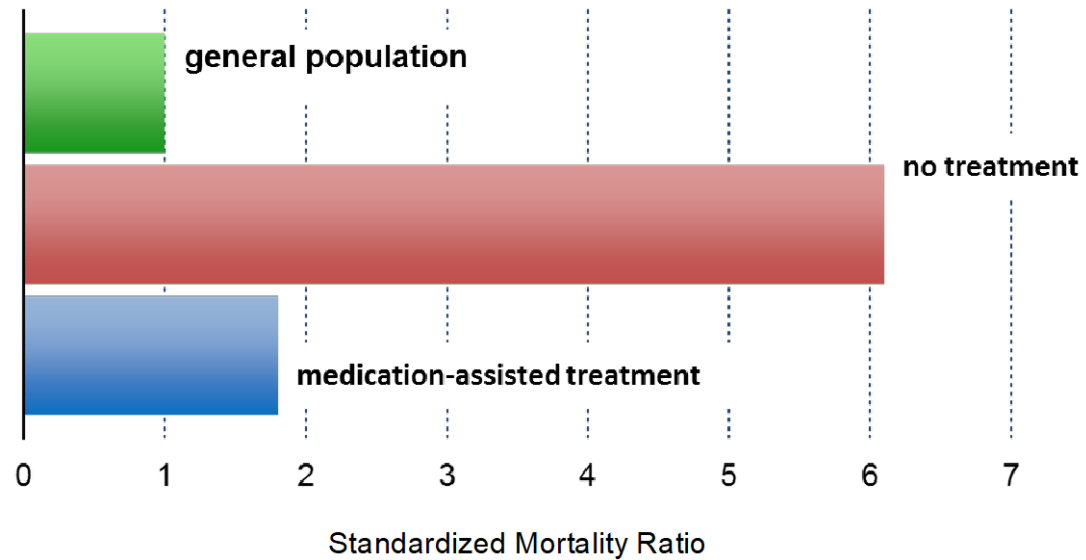


Hunt et al., 1971



# Benefits of MAT: Decreased Mortality

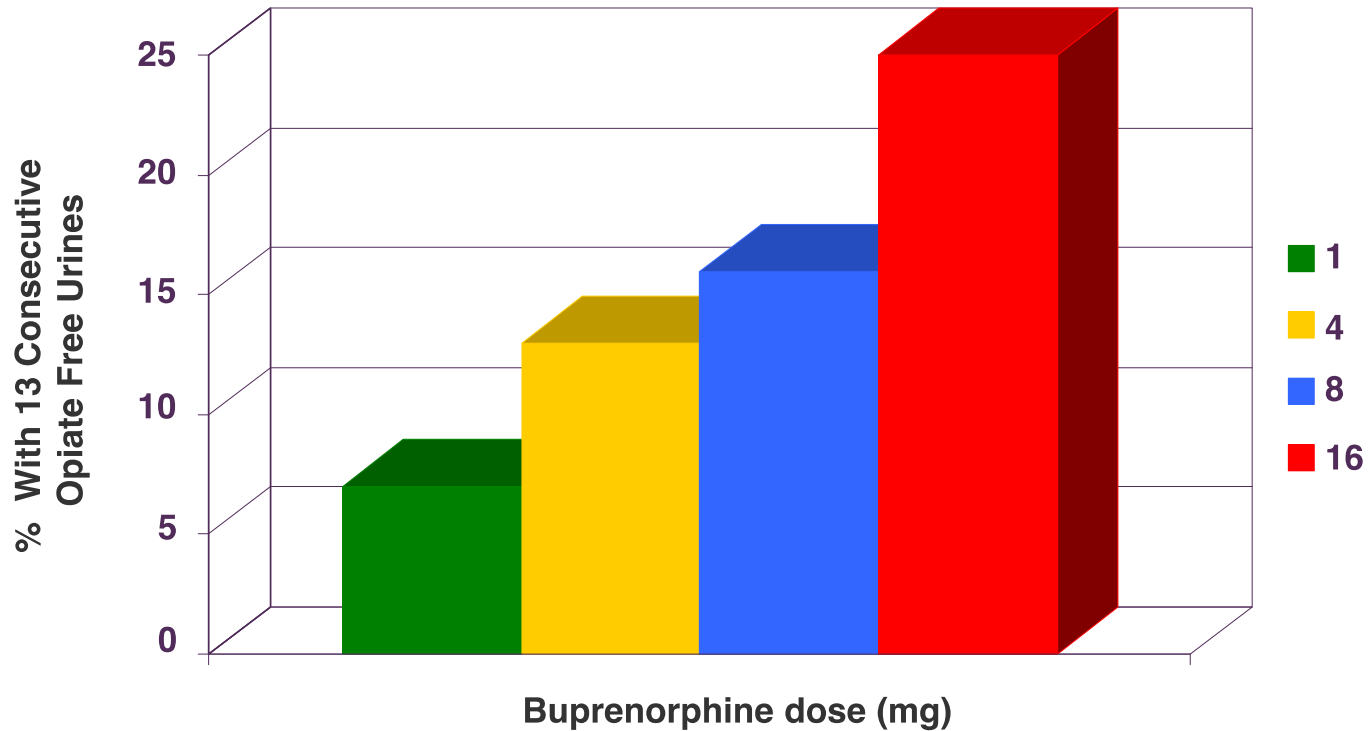
## Death rates:



Dupouy et al., 2017  
Evans et al., 2015  
Sordo et al., 2017

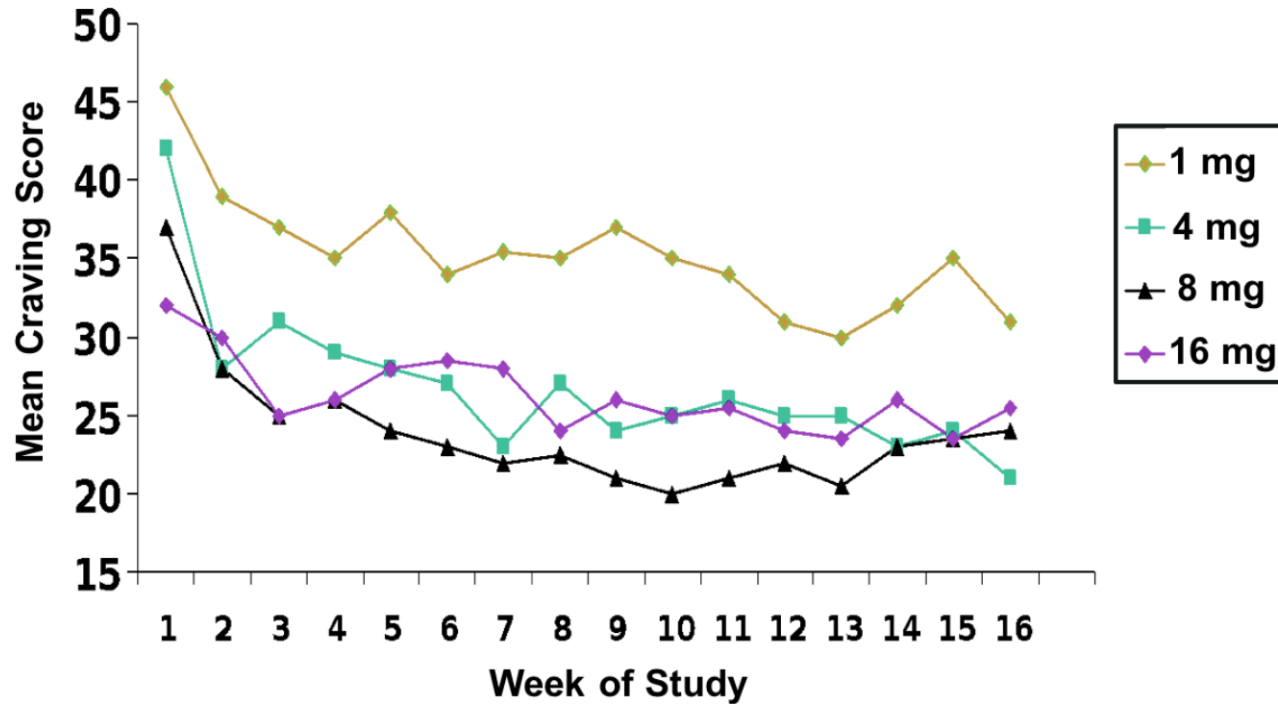


# Buprenorphine Dosing: Efficacy



Ling et al., 1998

# Mean Heroin Craving: 16 Week Completers: Reduced Craving with Therapeutic Buprenorphine Doses

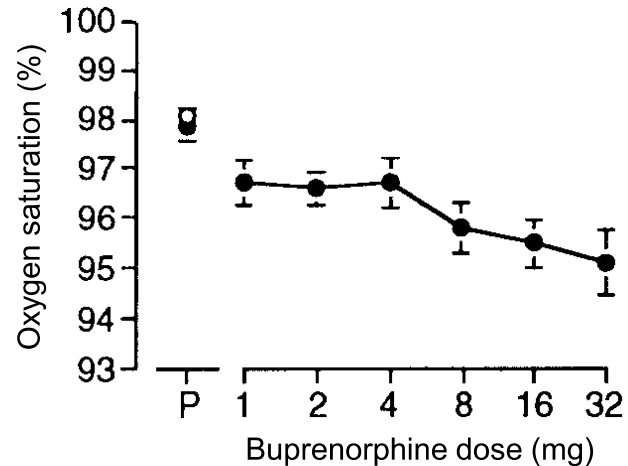
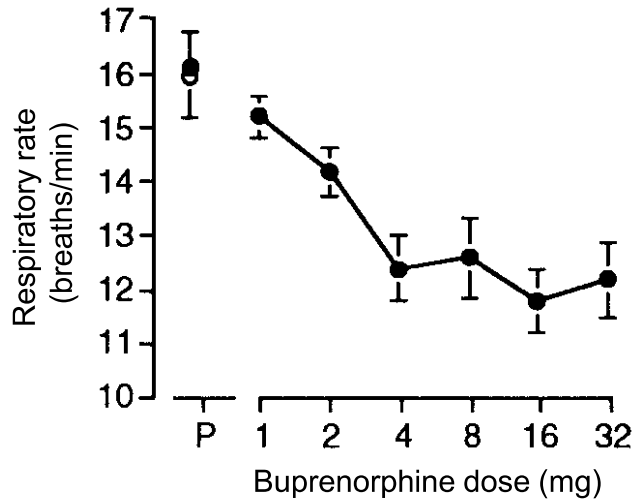


Ling et al., 1998



# Buprenorphine Dosing: Safety

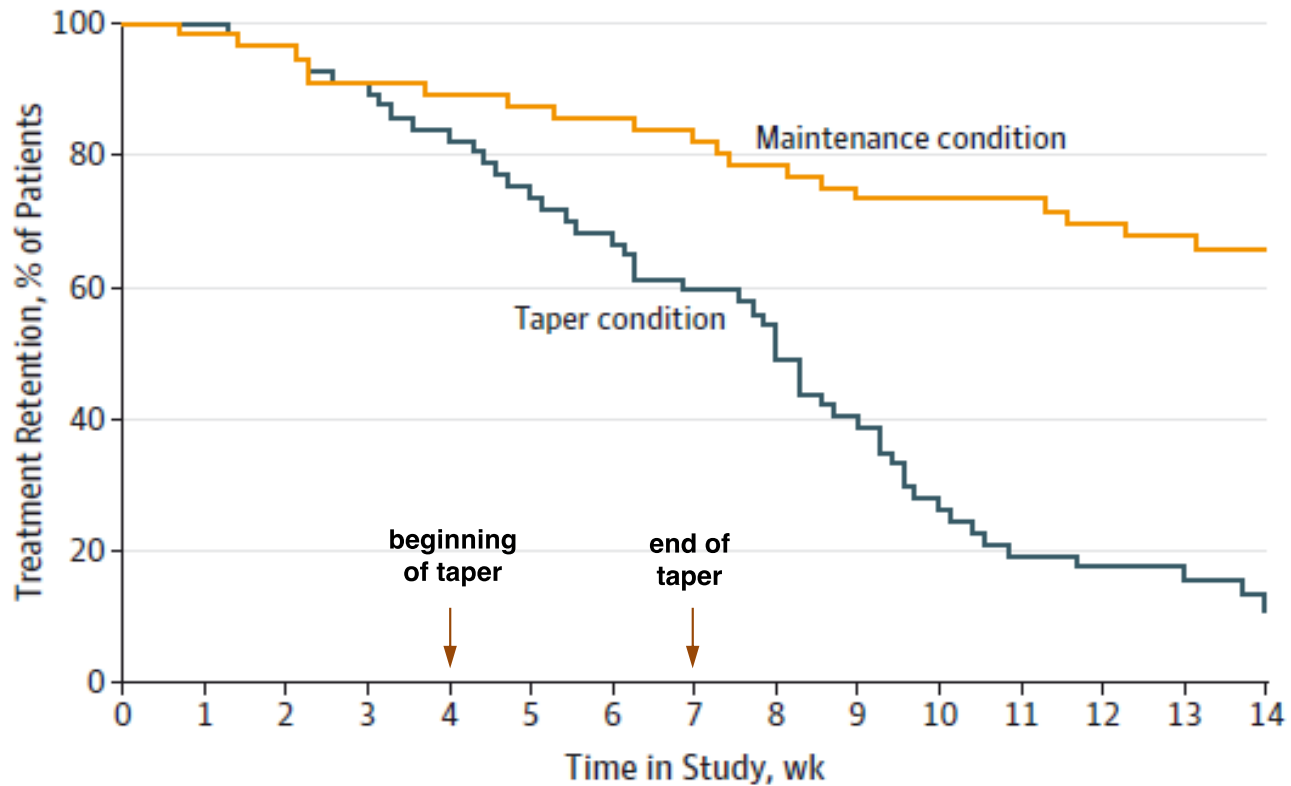
- Cognitive and psychomotor effects appear to be negligible



- Nearly all fatal poisonings involve multiple substances

Hakkinen et al., 2012  
Walsh et al., 1994

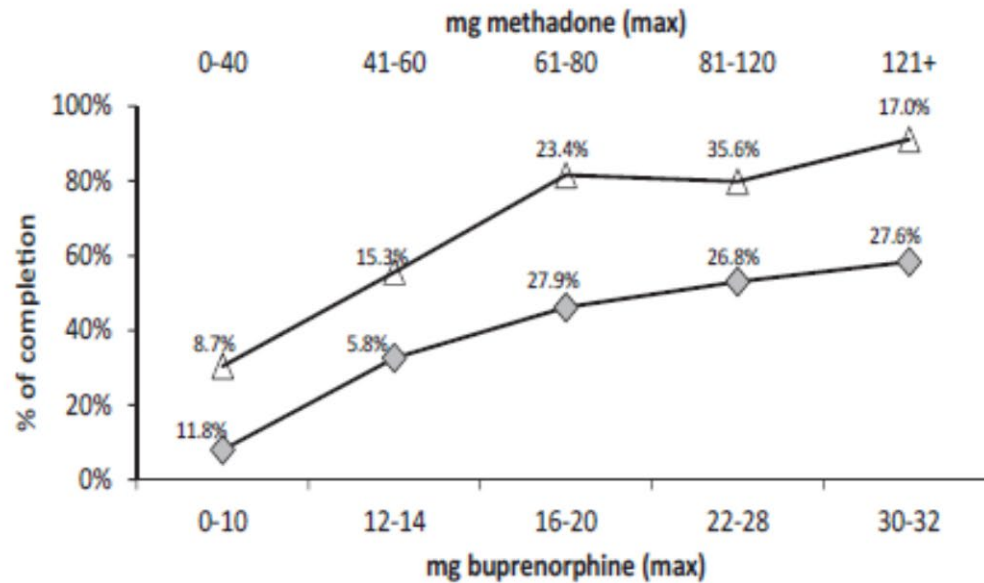
# Buprenorphine: Maintenance vs. Taper



Fiellin et al., 2014



# Benefits of Methadone: Treatment Retention



◆ Buprenorphine (% = % of buprenorphine participants prescribed in that dose range)

△ Methadone (% = % of methadone participants prescribed in that dose range)

Hser et al., 2014



Providers  
Clinical Support  
System

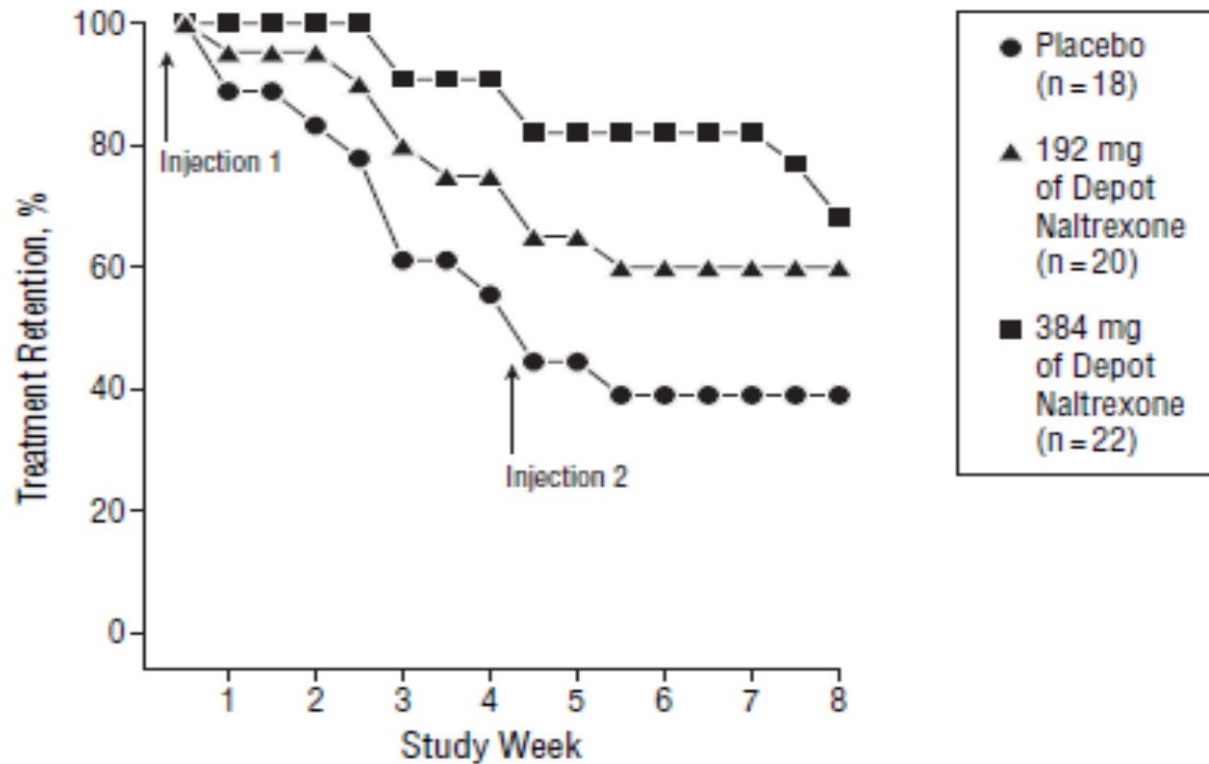
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# Naltrexone: Dose Response



Comer et al., 2011



## Conclusion

- Opioid use has various manifestations
- Physicians should be aware of the condition their patient is experiencing
- An evidenced based individualized treatment plan can help a person enter recovery



# Questions

**Questions?**

[Jason.beaman@okstate.edu](mailto:Jason.beaman@okstate.edu)



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