

## OBJECTIVES

Review of chronic, relapsing model of addiction

Comparison with other chronic diseases

Lessons from patients

### Addiction

- 24.6 million adults age 12+ live with a Substance Use disorder
- Only 10%, or 1/10 individuals, sought or received treatment for their addiction
- Opioid overdose is now the #1 cause of preventable death
  - higher than car accidents
  - higher than gun violence.
- We are 3 times more likely to die of an opioid overdose than a car accident and most car accidents are substance related.

NSUDH-2013 (national survey on drug use and health)



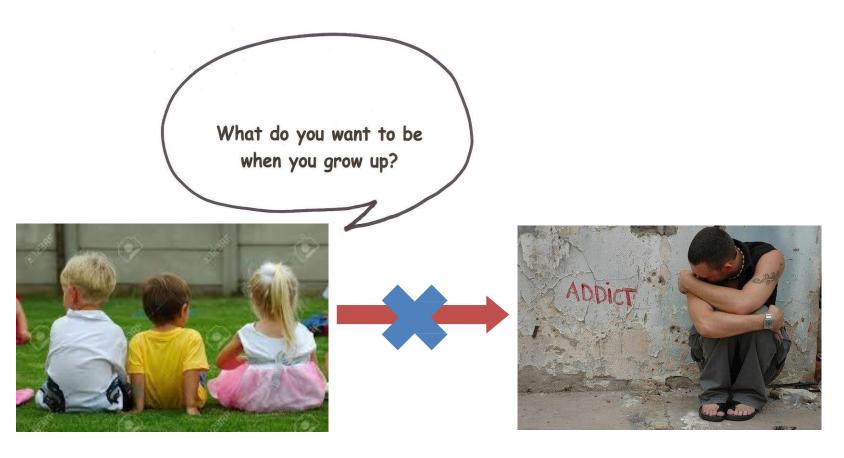
#### "Addiction Is Irrational"

 Primary, chronic brain disease is characterized by compulsive drug seeking and use despite harmful consequences

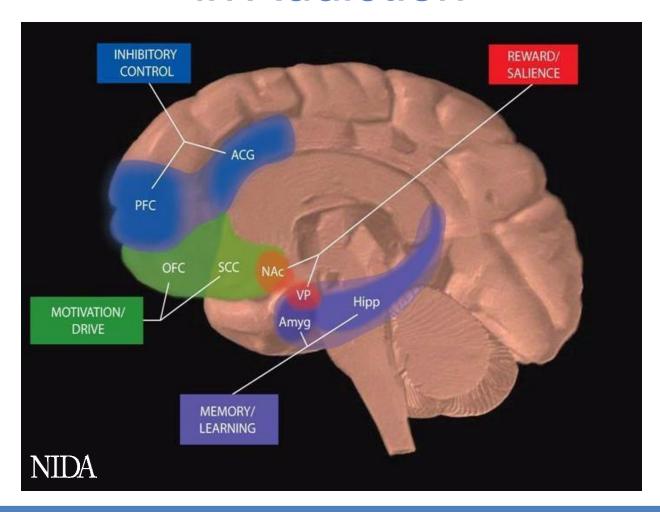
- Involves cycles of relapse and remission
- 40-60% genetic
- Without treatment addiction is progressive and can result in disability or premature death

American Society of Addiction Medicine. April 12, 2011. <a href="www.asam.org">www.asam.org</a> NIDA. August, 2010 <a href="http://www.drugabuse.gov/publications/science-addiction">http://www.drugabuse.gov/publications/science-addiction</a>

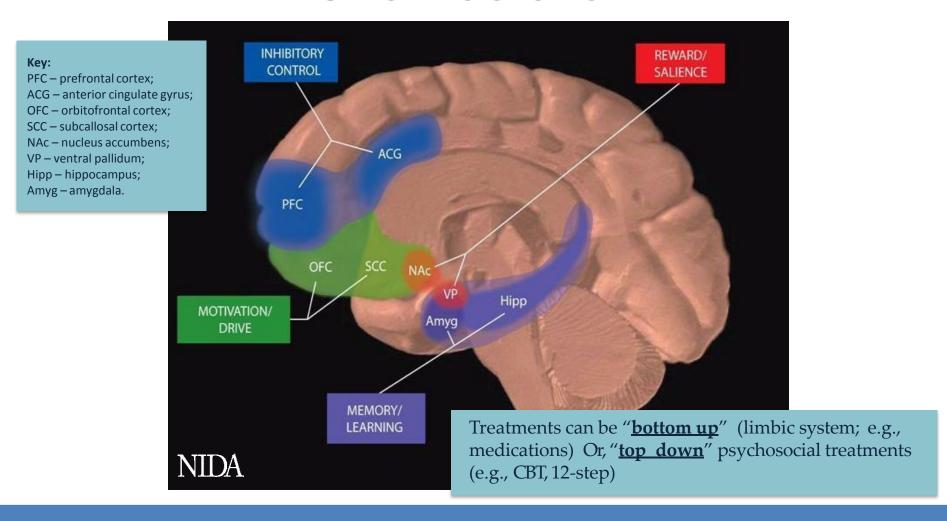
# Childhood Dreams and Aspirations



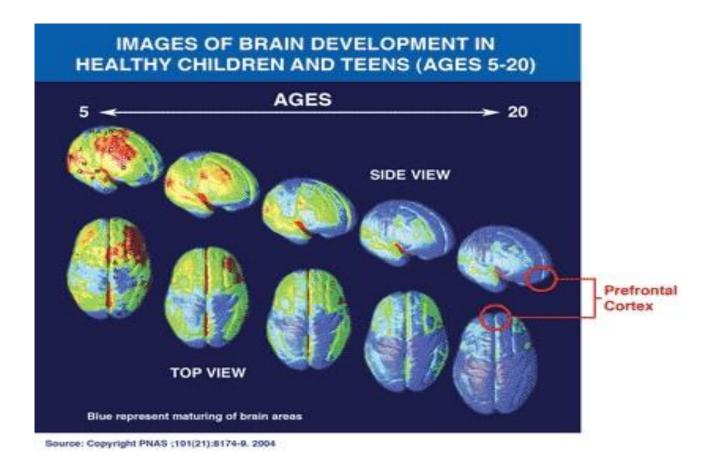
# Circuits Involved in Addiction



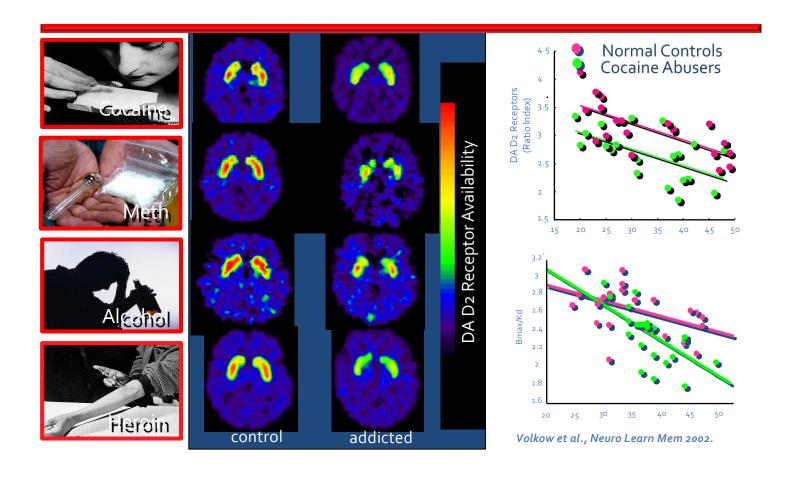
## Circuits Involved in Drug Abuse and Addiction



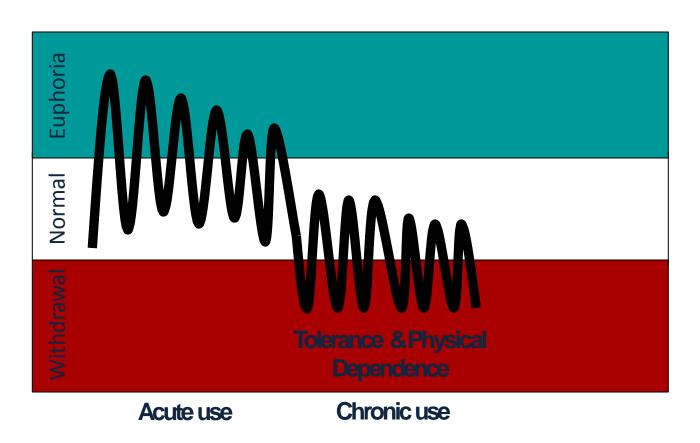
# Addiction is a Developmental Disease



## Dopamine D2 Receptors are Lower in Addiction



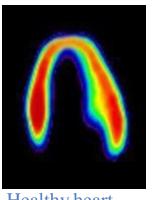
# Natural History of Opioid Use Disorder



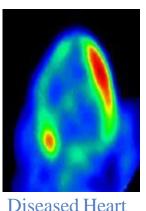
Slide courtesy of Dan Alford, 2012

### Addiction is Similar to **Heart Disease**

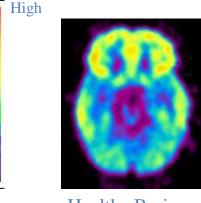
Decreased Heart Metabolism in **Coronary Artery Disease** 



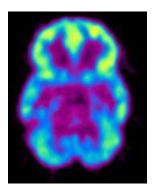
Healthy heart



Decreased Brain Metabolism in Addiction

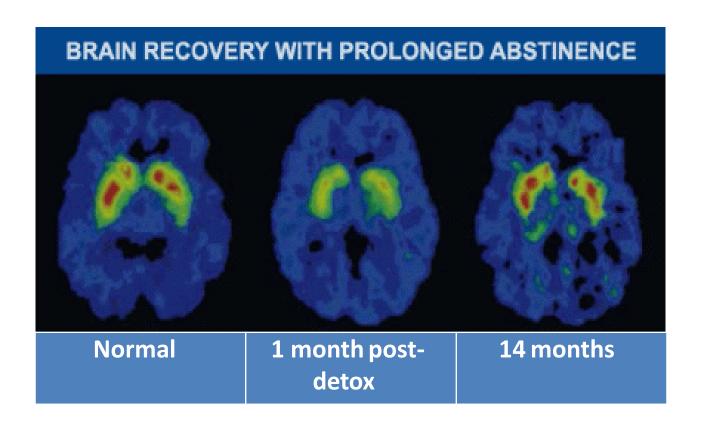


Healthy Brain



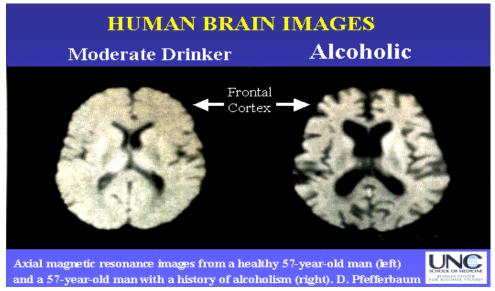
**Diseased Brain** 

### Functional Recovery Takes Time

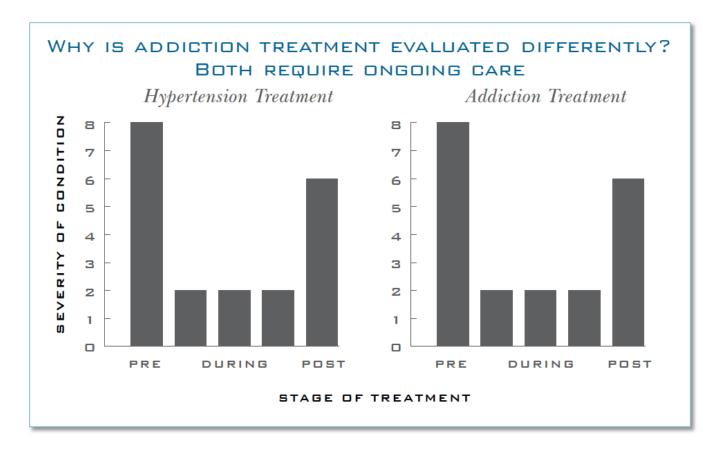


Source: Volkow, ND et al., Journal of Neuroscience 21, 9414-9418, 2001



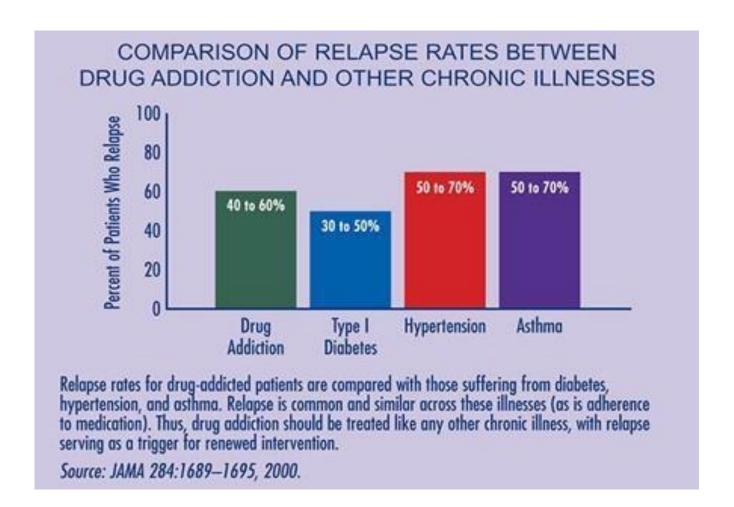


## Addiction is a Treatable Disorder



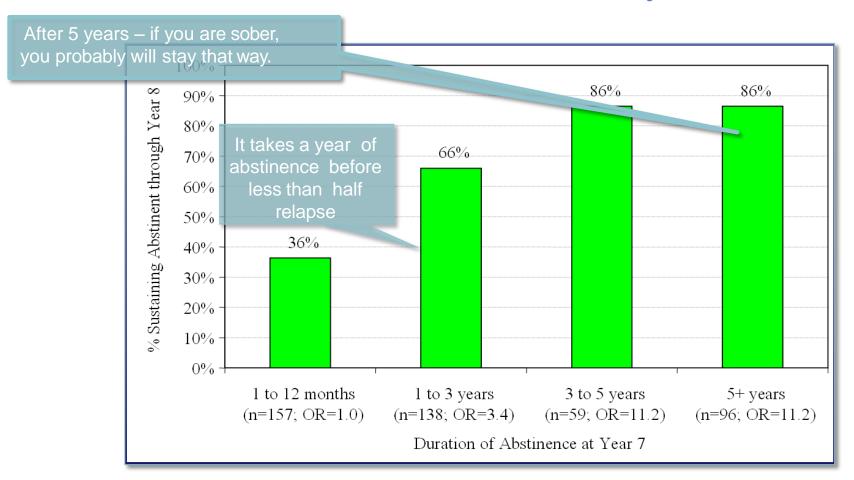
NIDA. Principles of Drug Addiction Treatment. 2012. McLellan et al., JAMA, 284:1689-1695, 2000.

### Comparable Relapse Rates



Slide courtesy of NIDA, Drugs, Brain Behavior: the Science of Addiction

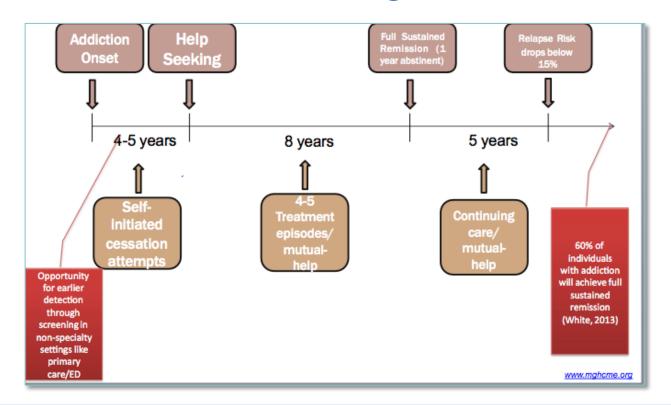
## Extended Abstinence is Predictive of Sustained Recovery



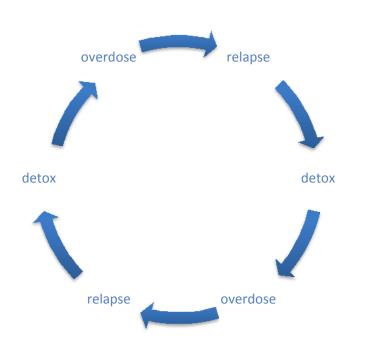
Slide courtesy of NIDA, Drugs, Brain Behavior: the Science of Addiction

# For More Severely Addicted Individuals ...

course of SUD and achievement of stable recovery can take a long time ...



## Acute Care Model as We Know It





"Treatment"

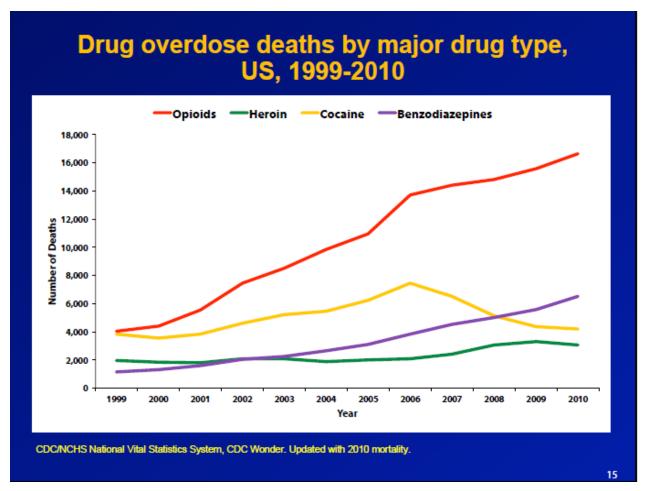
## Relapse Requires Increased Support

- We label patients as "not ready" or "non compliant"
- We ask them to seek a higher level of care on their own, when most ill
- We refer them for "higher level of care" yet many of those programs are not evidence based, and are essentially lower level of care
- What would we do if a cancer survivor had a lymphoma recurrence after years of remission?

## Who is at Risk for an Overdose?

- Patients receiving opioids from multiple prescribers / pharmacies
- Patients taking high daily doses of opioids
- Men ages 35-55 years old
- Whites/ American Indians
- Medicaid
- Rural
- Patients taking Opioids and Benzodiazepines

# OD on Benzodiazepines is Increasing



### Benzodiazepines 1996-2013

Fatal overdoses have nearly quadrupled

- Combination of Alcohol and Benzodiazepines
- Combination of Opioids and Benzodiazepines
- The number of adults filling a RX increased 67% from 8.1 million to 13.5 million

#### **Alcohol Withdrawal**

- Begins 6- 8 hours after last drink
- Peaks 24 hours after last drink
- Typically resolves in 7 days

#### **Alcohol Withdrawal**

- 5 % of patients with AUD will develop more than mild to moderate symptoms
- Chronic medical conditions, nutritional deficiencies and electrolyte abnormalities can contribute and complicate course

#### Alcohol Withdrawal Seizures

- Estimated to occur in 5-15 % of patients
- DT's estimated to occur in 5% of patients admitted for Alcohol Withdrawal
- Mortality rate is 20% for patients who have DT's

Source: APA Textbook of Substance Abuse Treatment, 4th edition

### Us Opioid Related Deaths

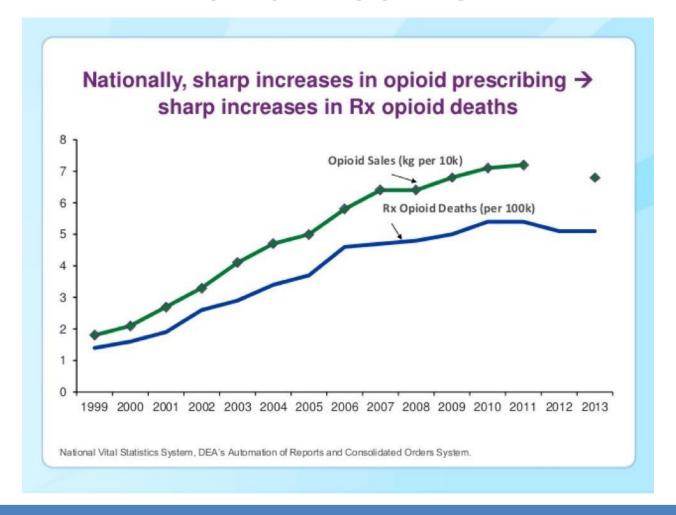
Opioid analgesics 5 Deaths per 100,000 population 3 Heroin 2006 2000 2002 2004 2008 2010 2012 2013 Year

Figure 1. Age-adjusted rates for drug-poisoning deaths, by type of drug: United States, 2000–2013

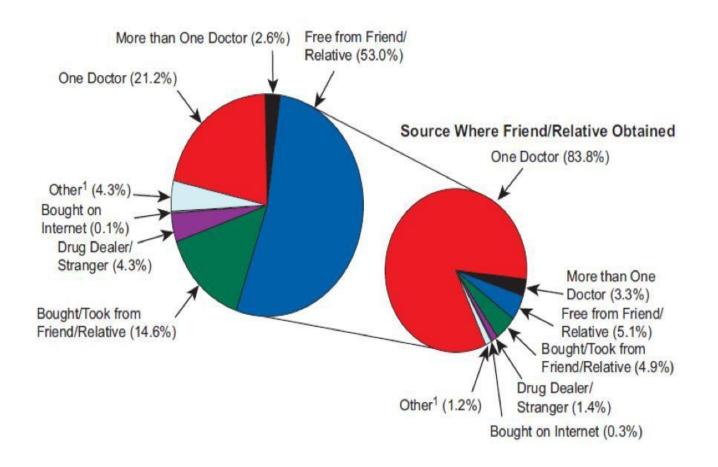
NOTES: The number of drug-poisoning deaths in 2013 was 43,982, the number of drug-poisoning deaths involving opioid analgesics was 16,235, and the number of drug-poisoning deaths involving heroin was 8,257. A small subset of 1,342 deaths involved both opioid analgesics and heroin. Deaths involving both opioid analgesics and heroin are included in both the rate of deaths involving opioid analgesics and the rate of deaths involving heroin. Access data table for Figure 1 at: http://www.cdc.gov/nchs/data/databriefs/db190\_table.pdf#1.

SOURCE: CDC/NCHS, National Vital Statistics System, Mortality.

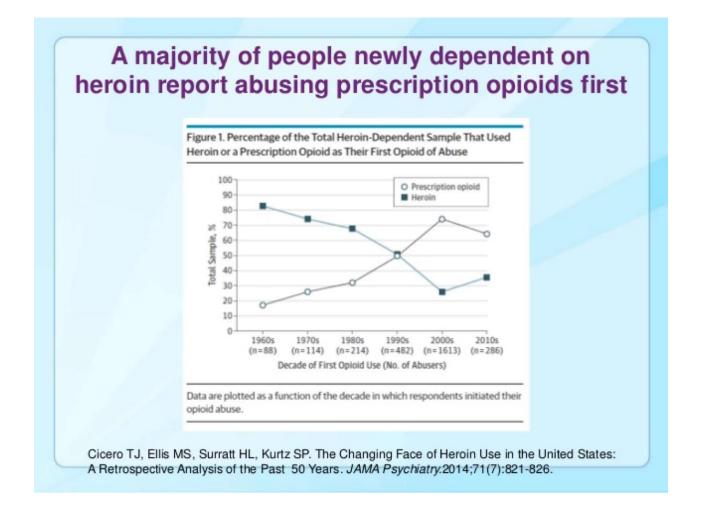
## Prescribing Patterns and Deaths



### Sources of Pain Meds



### Rx Opioids As Gateway To Heroin



### Heroin Use is Part of a Larger Substance Use Problem

Nearly all people who used heroin also used at least 1 other drug.

Most used at least 3 other drugs.

Heroin is a highly addictive opioid drug with a high risk of overdose and death for users.

People who are addicted to...









2x

3x

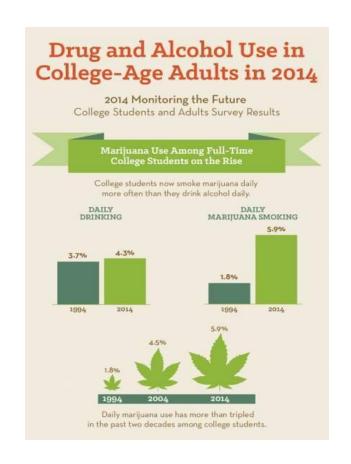
15x

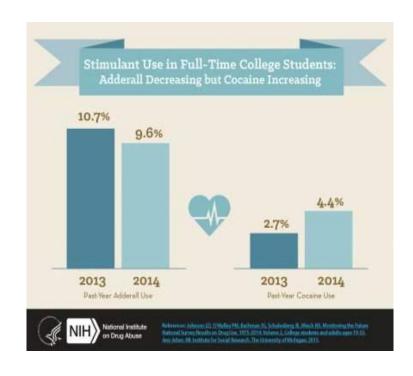
40x

...more likely to be addicted to heroin.

SOURCE: National Survey on Drug Use and Health (NSDUH), 2011-2013.

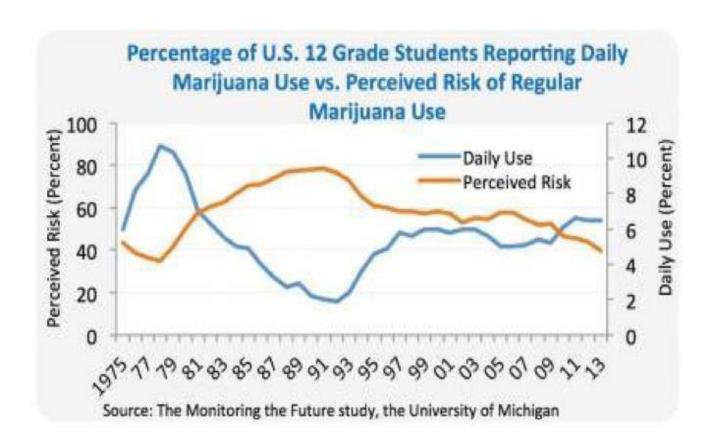
# Monitoring the Future 2014 College and Adult





Source: National Institute on Drug Abuse; National Institutes of Health; U.S. Department of Health and Human Services

### Perceived Harm and Drug Use



## The Changing Face of Heroin Addiction

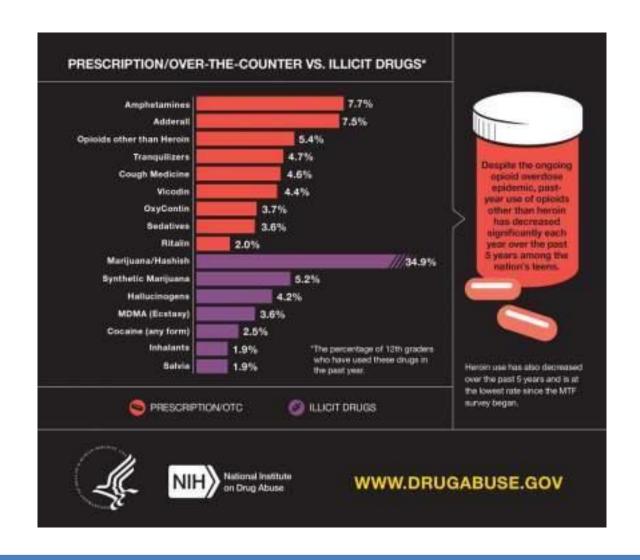
### Addiction



Your medicine cabinet could be a gateway to heroin.

CDC Vital Signs: Demographic and Substance Use Trends Among Heroin Users — United States, 20,02,013,000

### Prescription/OTC vs. Illicits



#### Medication RX Numbers -2013

- PERCOCET (Hydrocodone / Acetaminophen)
   is the #1 most prescribed medication in the USA
- XANAX is the #1 most prescribed Psychiatric medication in the USA

### Assessment of Need for Treatment

- ✓ Confirm OUD
- ✓ Establish current use, when, what, how much
- ✓ Assess social supports
- ✓ Evaluate degree of motivation
- ✓ Identify co-morbid medical and psych history
- ✓ PE: skin for inj sites, older tracking, body systems, cardiac murmurs, rhythm abnx

- ✓ Lab testing- blood and urine tox, liver enz, bili, Hep B, C, HIV
- ✓ Medications

**Be sure to CHECK THE PMP!** 

# Access to Treatment – Gap

- 2.5 million Americans 12 and over have opioid use disorders
- 120 people a day die of substance related overdoses
- Fewer than 1 million received treatment
- We let people "hit rock bottom"



# Treating a Bio-behavioral Disorder Must Go Beyond Just Fixing the Chemistry

We need to treat the whole person

Pharmacological Treatments (Medications)

**Medical Services** 



Behavioral Therapies

Social Services

# Your Parents Were Right

At least about one thing...



Beliefs
Perception
Respect
Empathy
Stigma

# Stigma

- Set of negative and often unfair beliefs that a society or group of people have about something
- A mark of disgrace or dishonor

#### October 2015: Charleston, West Virginia



"We can't fight this epidemic without removing stigma."

President Obama, 10/21/2015



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### Words to Avoid

- Addict, Abuser, Junkie
- Abuse
- Clean or dirty
- Habit
- Replacement or Substitution therapy
- (MAT)

#### Words to Use

Person first language focuses on the person, not the disorder

- Addiction
- Misuse
- Substance Use Disorder
- Medication Treatment (caution "MAT")
- Person with Patient

# Does AA "cause" better outcomes or is AA participation an outcome of better

#### The Bradford Hill Criteria

- 1: Strength of Association. The stronger the r the dependent variable, the less likely it is that
- 2: Temporality. It is logically necessary for a
- **3:** Consistency. Multiple observations, of an a circumstances and with different measurement
- **4: Theoretical Plausibility.** It is easier to account theoretical basis for such a conclusion.
- **5:** Coherence. A cause-and-effect interpretatic conflict with what is known about the variables competing theories or rival hypotheses. In othe other knowledge.
- **6: Specificity in the causes.** In the ideal situat showing that an outcome is best predicted by o claim.
- 7: Dose Response Relationship. There should (i.e., the independent variable) and people's stavariable).
- **8: Experimental Evidence.** Any related resear inference more plausible.
- **9: Analogy.** Sometimes a commonly accepted area.

- Using accepted scientific standards (Bradford Hill criteria) and the most rigorous scientific methods (i.e., RCTs, instrumental variables analysis, PS matching), evidence indicates causal therapeutic benefit of AA
- The one exception is "specificity" (e.g., other interventions could also cause these benefits)
- But given AA is available free of charge in practically every US community and that an intervention's "Impact" is a product of = reach x effectiveness (Glasgow et al, 2003), AA canbe considered a clinical and public health ally in ameliorating the prodigious burden of disease attributable to alcohol addiction

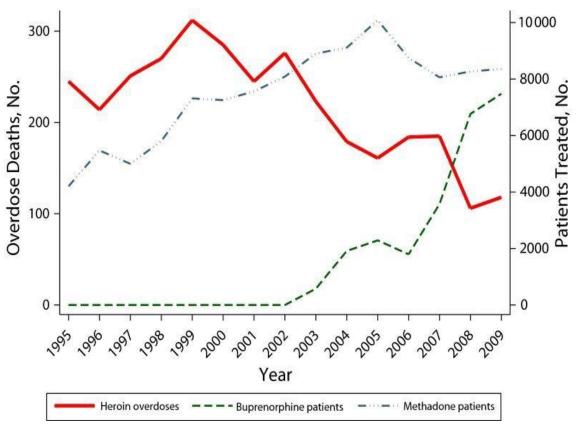
### Medication Treatment

- "MAT" = Medication PLUS counseling and behavioral therapies
- "Opioid Agonist Therapy," Medication, or Treatment preferred
  - Reduces drug use
  - Reduces the risk of infectious disease transmission
  - Reduces criminal activity
  - Reduces the risk of overdose
  - Reduces death
  - Increases treatment retention
  - Improves social functioning
  - Cost-effective
  - Safe

# Overdoses Symptomatic of Untreated Disease "A key driver of the overdose epidemic is underlying substance-use disorder. Consequently, expanding access to addiction-treatment services is an essential component of a comprehensive response.

- 50% of Addiction treatment centers offer medication
- <38% of eligible patients are offered medications</li>
- <5% of physicians are waivered to prescribe buprenorphine

#### Methadone and Suboxone save lives



Schwartz, RP, Gryczynski J, O'Grady, Ke et al. Opioid agonist treatments and heroin overdose deaths in Baltimore, MD, 1995-2009. Am J Public Health 2013;103:917-22

# Medication Therapy

- Opioid Agonists
  - Full: Methadone (Methadose or Dolophine)
  - Partial: Buprenorphine/Naloxone, Buprenorphine (Suboxone or Subutex)
- Opioid antagonist
  - Naltrexone (Revia or Vivitrol)

## Methadone

opiate receptor opiate

- Long acting, full opioid agonist
- Binds to and occupies mu

  opioid receptors
- Prevents euphoria from other mu agonists
- Alleviates withdrawal symptoms
- Administered in licensed OTP
- Federal law: initial dose 10-30 mg, not to exceed 40 mg in day 1
- Suppresses cravings (60-120mg+)
- Can prolong QTc with risk of Torsades de Pointes
- Respiratory depression can be a side effect at any dose
- Increases overdose risk significantly if mixed with sedative hypnotics and ETOH

# Methadone Myths

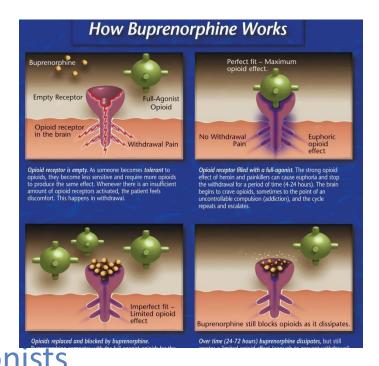
- "Liquid handcuffs"
- Substitutes one addiction for another
- Prevents true recovery
- Should not be used long term
- Rots teeth
- Damages bones
- Turns people into "zombies"
- Causes overdoses



# Buprenorphine

#### Major Paradigm Shift: DATA 2000

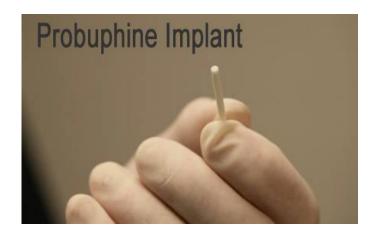
- Partial agonist antagonist
- Sublingual
- Higher affinity for the mu opioid receptor compared to full agonists
- Slow to dissociate
- Will displace full agonists from the receptor with decreased opioid effect → precipitated withdrawal
- Relieves cravings without producing euphoria or dangerous side effects of other opioids
- Naloxone to deter IV use, not active sublingually



## Naltrexone

- Full mu opioid antagonist
- Blocks the euphoric effect of mu opioid agonists
- No dependence, no need to wean
- Not scheduled no special training or license needed
- Reduces relapse rates
- Will precipitate withdrawal if agonists (full or partial) are occupying mu receptors
- Must be 7-10 days opioid free
- Increased risk of overdose if try to overcome blockade
- Increased risk of overdose end of month or missed dose because of loss of tolerance
- Monthly IM dosing improves adherence, low adherence with oral dosing
- Substantially less stigma

# On the Horizon... Probuphine



Investigational subdermal implant delivers buprenorphine continuously for six months following a single treatment.

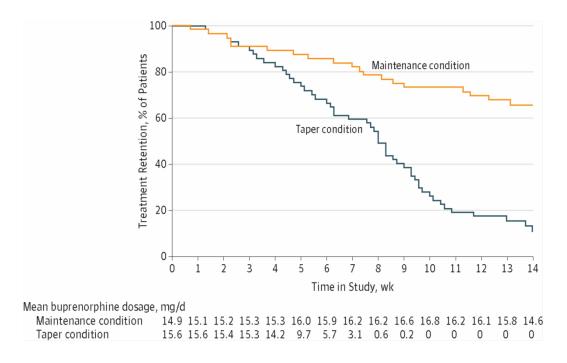
Currently under FDA review.

# To taper or to maintain?

- No question, actually.....
- Longer treatment, better outcomes
- Consistent with chronic disease model
- Think DM, CAD, COPD
- As with any medication no set limit
- Minimum of 12 months, but better outcomes with longer durations
- Continually reassessed and individualized

#### From: Primary Care-Based Buprenorphine Taper vs Maintenance Therapy for Prescription Opioid Dependence: A Randomized Clinical Trial

JAMA Intern Med. 2014;174(12):1947-1954. doi:10.1001/jamainternmed.2014.5302



Results: Completion of 14 week trial: taper 11% vs maintenance 66% Mean percentage of urine negative for opioids: taper 35% vs maintenance 53%

#### Figure Legend:

Treatment Retention and Mean Buprenorphine Dosage for Patients With Prescription Opioid Dependence. Patients were assigned to the taper or the maintenance condition. Buprenorphine treatment was administered as a tablet formulation of buprenorphine hydrochloride and naloxone hydrochloride in a 4:1 ratio.

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# Imagine Sobriety...

- After multiple detoxes, long term programs, losses, overdoses....
- You achieve sobriety
- You are engaged in counseling
- You are engaged in a treatment community
- You are exercising and eating healthfully
- You are in college or have a job
- You have your family back
- You feel "normal

# BUT....

#### You are on agonist therapy/medication

- You are told by your support network that you are not sober
- You are "trading one addiction for another," using a "crutch"
- You are told you cannot engage in peer support groups that bolster your sobriety
- You are badgered by your insurance company for repeated authorizations as to why you need it
- You are asked by your family and doctors when you are going to get off the of the medication

# "People Don't Fail Treatment Treatment Fails People"

- Deconstruct the relapse with your patient
- Good people make bad decisions when SUD active
- Change takes time, patience and trust
- When diseases flare, we increase care or enlist the care of other team members. This is no different
- Trust is an important tool
- Positive reinforcement (contingency management)
- Competing priorities
- Communicate with others
  - "No one size fits all" just like other diseases
  - Diet controlled pre-diabetes, oral agents, insulin for DM
  - Diet, exercise, statin, beta-blocker, ASA, ACE inhibitors for heart disease

#### Lessons Learned

- Listen to your patients
- It's hard to have an addiction
- Diversion happens
- Most have used Suboxone in the past and can do home inductions
- Don't get caught up in the dose-splitting hairs
- Don't forget about "pseudo-addiction"
- Take sleep disturbance seriously advance Suboxone
- Take report of cravings extremely seriously treat

### **Addiction Treatment**

For each treatment plan, try to consider the level of care –

What does THE PATIENT want to craft as a realistic and concrete plan focused on engagement, patient preference and safety

Meeting patients where they are . . but never leaving them where we found them.

### What Next?



