

Treatments for Adolescents/Young Adults with Opioid Use Disorder

Webinar
May 28th, 2009

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Opioid Pharmaceuticals - Then

THE NEW YORK MEDICAL JOURNAL. 39

BAYER PHARMACEUTICAL PRODUCTS

Send for samples and Literature to

ASPIRIN
The substitute for the salicylates

HEROIN
The sedative for coughs

LYCETOL
The uric acid solvent

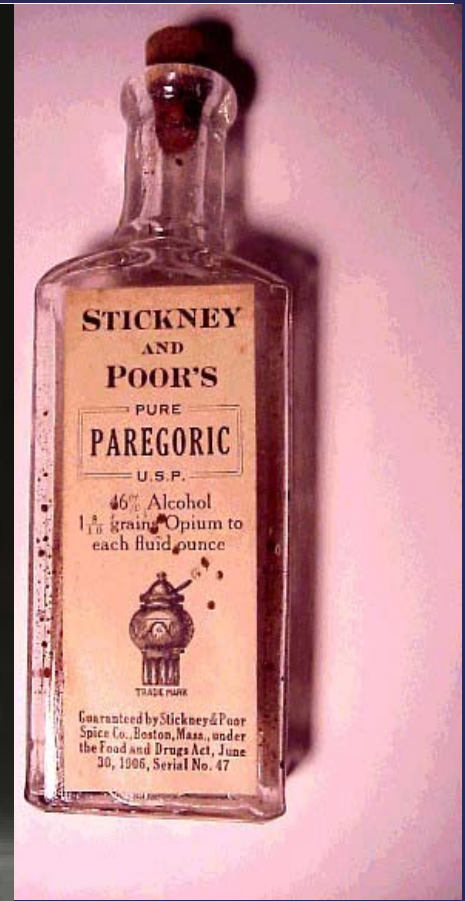
EUOPHEN
The sedative for coughs

HEROIN-HYDROCHL.
The sedative for coughs

SALOPHEN
The antirheumatic and antineuralgic

FARBENFABRIKEN OF ELBERFELD CO.

40 STONE STREET, NEW YORK.



Opioid Pharmaceuticals – and Now



C-III	ABBOTT LABORATORIES	C-II	PURDUE PHARMA L.P.	C-II	ENDO PHARMACEUTICALS
 <p>10 mg/660 mg</p> <p>Vicodin HP® (hydrocodone bitartrate/ acetaminophen tablets, USP)</p>		 <p>10 mg</p> <p>20 mg</p> <p>40 mg</p> <p>80 mg</p> <p>160 mg</p> <p>Controlled-Release Tablets</p> <p>OxyContin® (oxycodone HCl)</p>		 <p>2.5 mg/325 mg</p> <p>5 mg/325 mg</p> <p>7.5 mg/325 mg</p> <p>7.5 mg/500 mg</p> <p>10 mg/325 mg</p> <p>10 mg/650 mg</p> <p>Percocet® (oxycodone HCl/acetaminophen, USP)</p>	

Opioids: Heroin

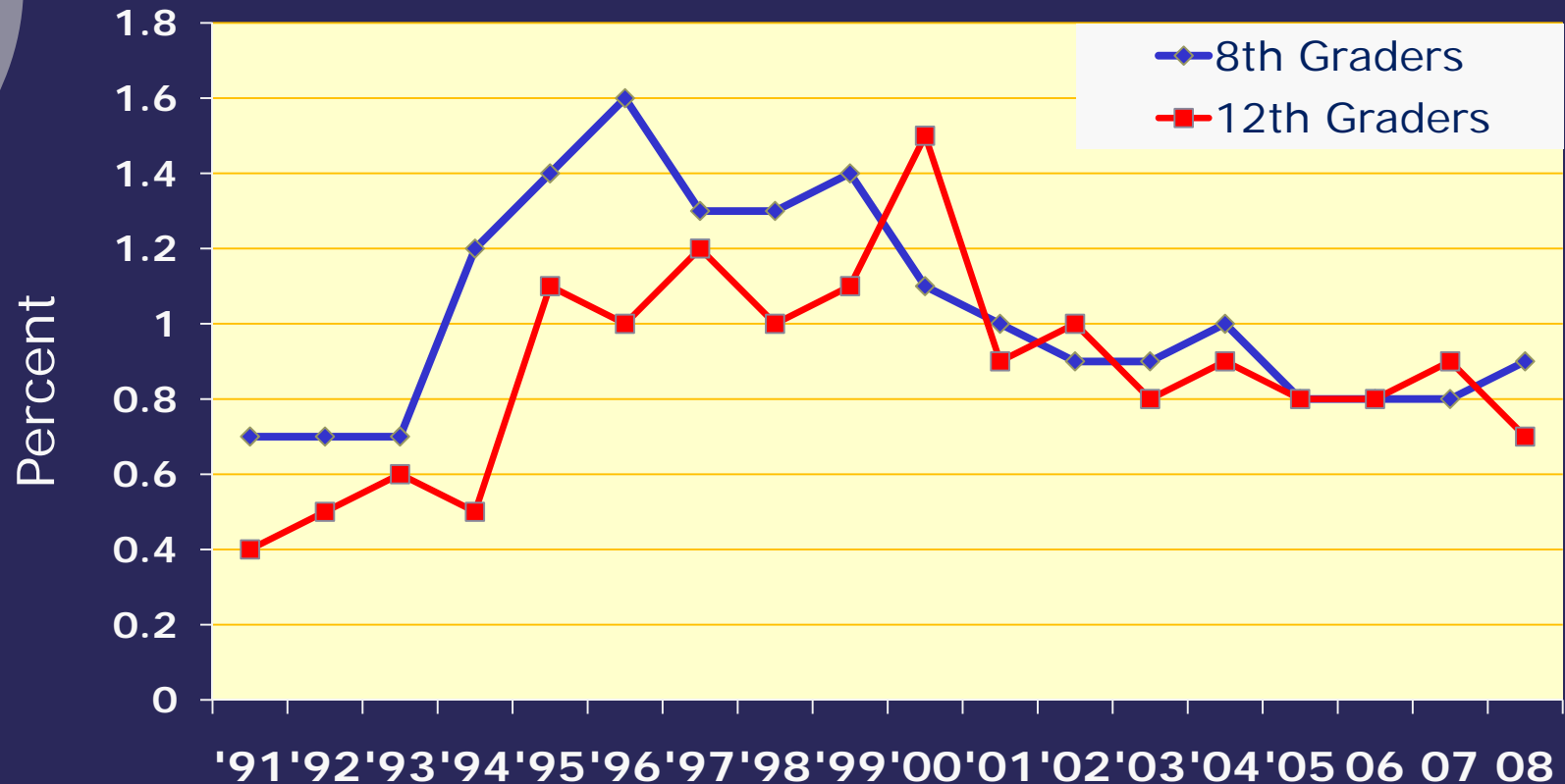




Prevalence

Heroin Use From 1991-2007 (MTF)

Annual Use Prevalence: 8th and 12th Graders

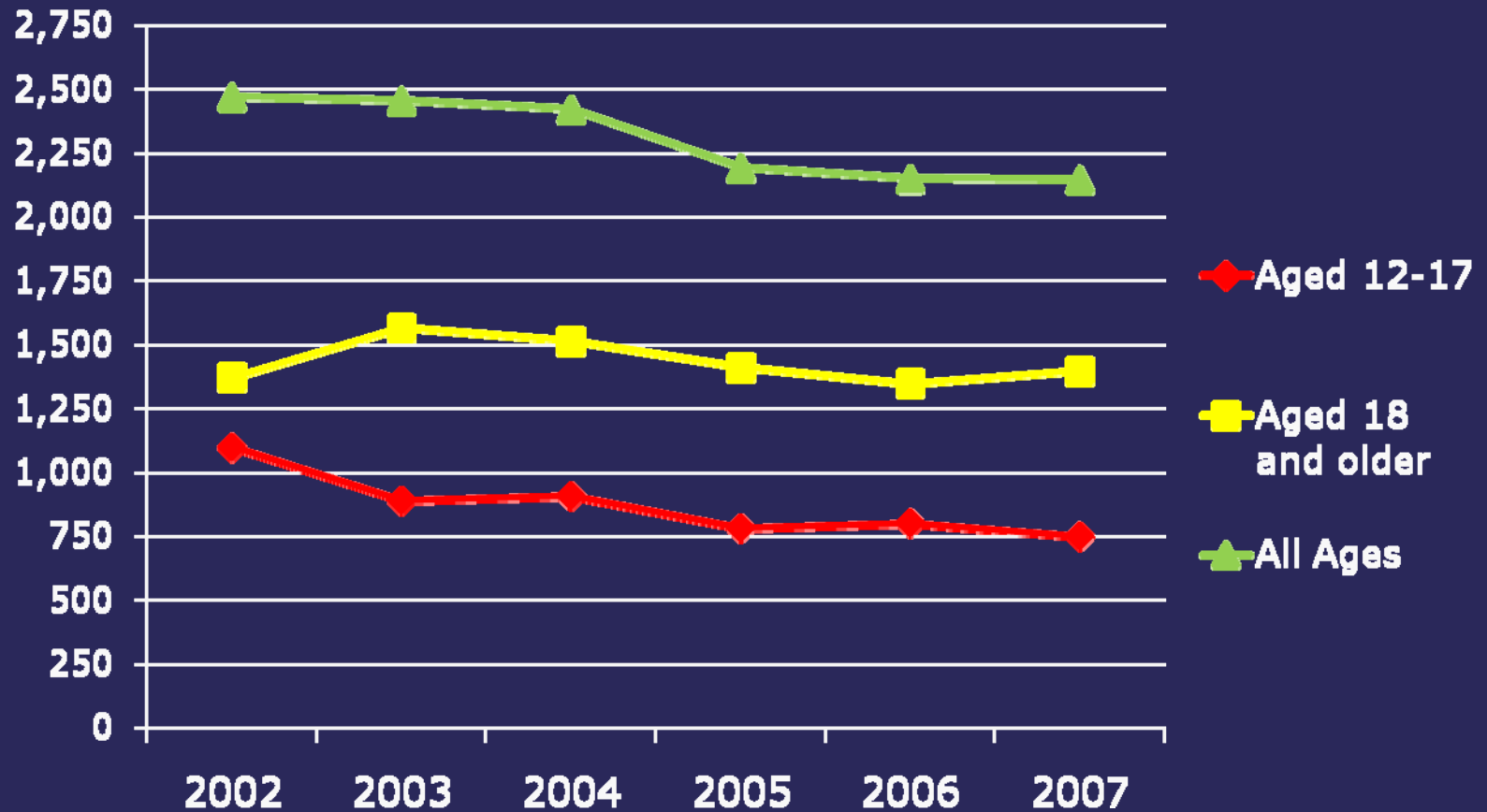


Non-Medical Prescription Opioid Use

MTF: Annual Use Prevalence 12th Graders

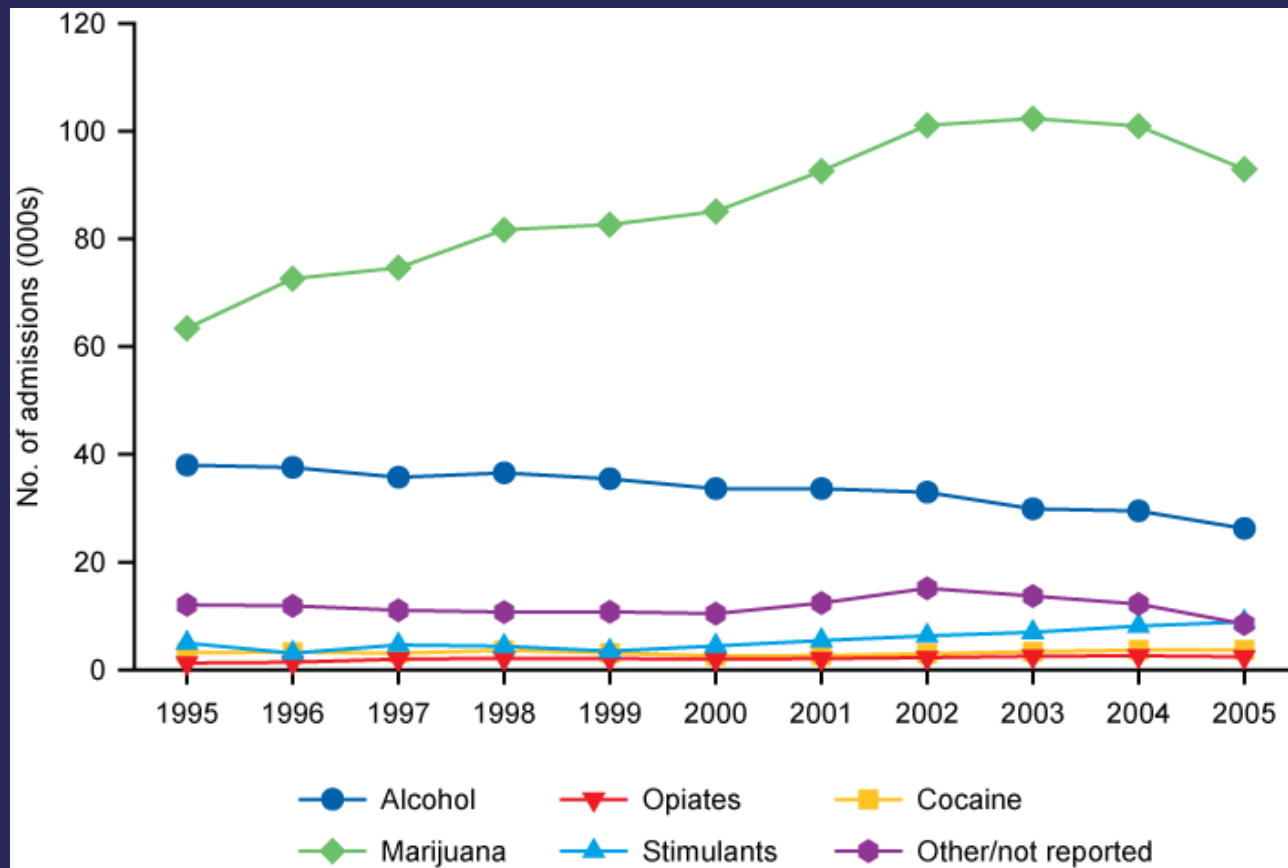


New Non-Medical Users of Prescription Opioids, By Age Group: 2002-2007



Source: NSDUH 2003-2007

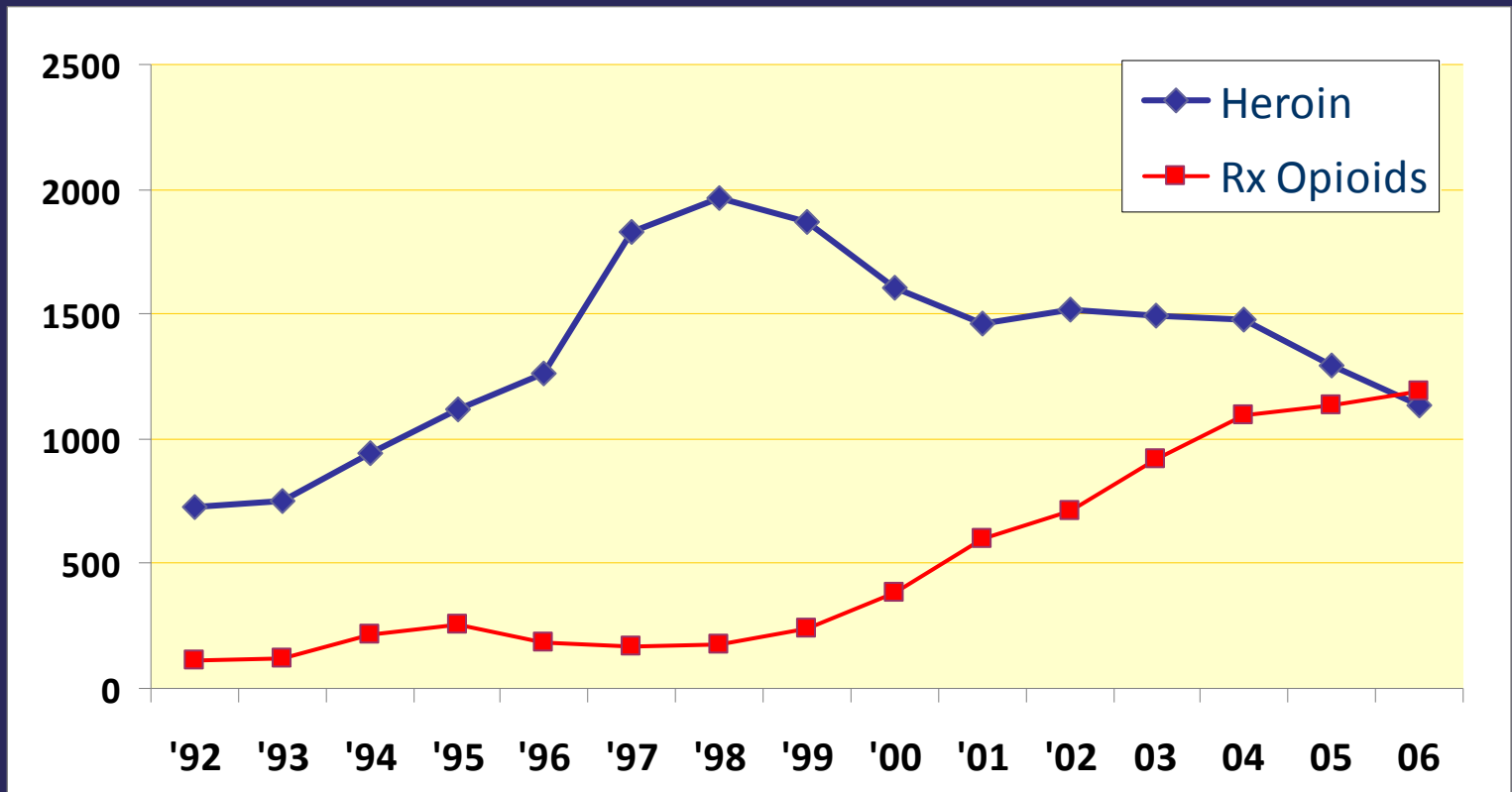
Adolescent Admissions by Primary Substance: TEDS 1995-2005



Opioid Admissions aged 12-17 yrs

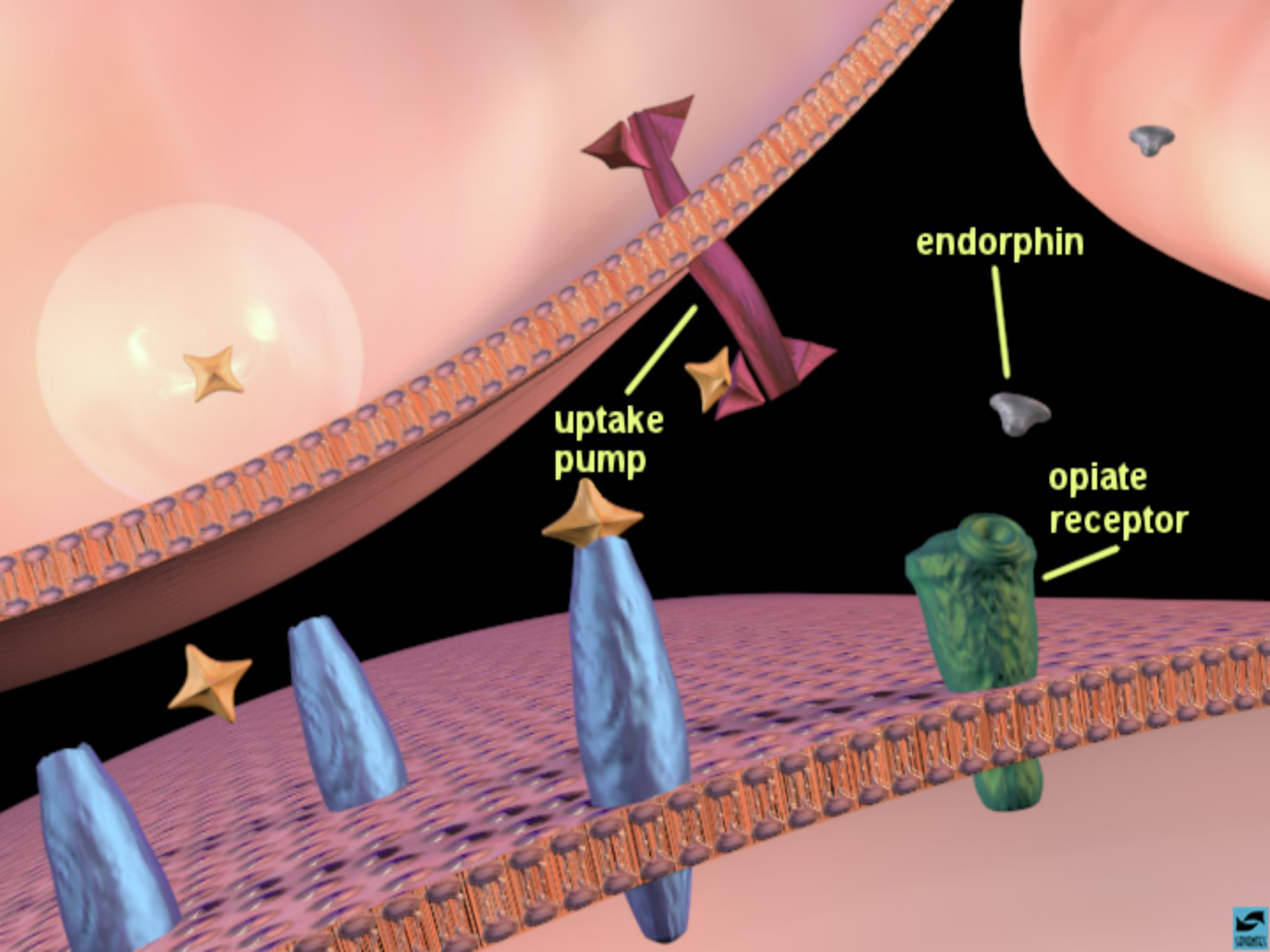
Treatment Episode Dataset (TEDS): 1996-2006

Number of Admissions





Opioids and Brain Mechanisms



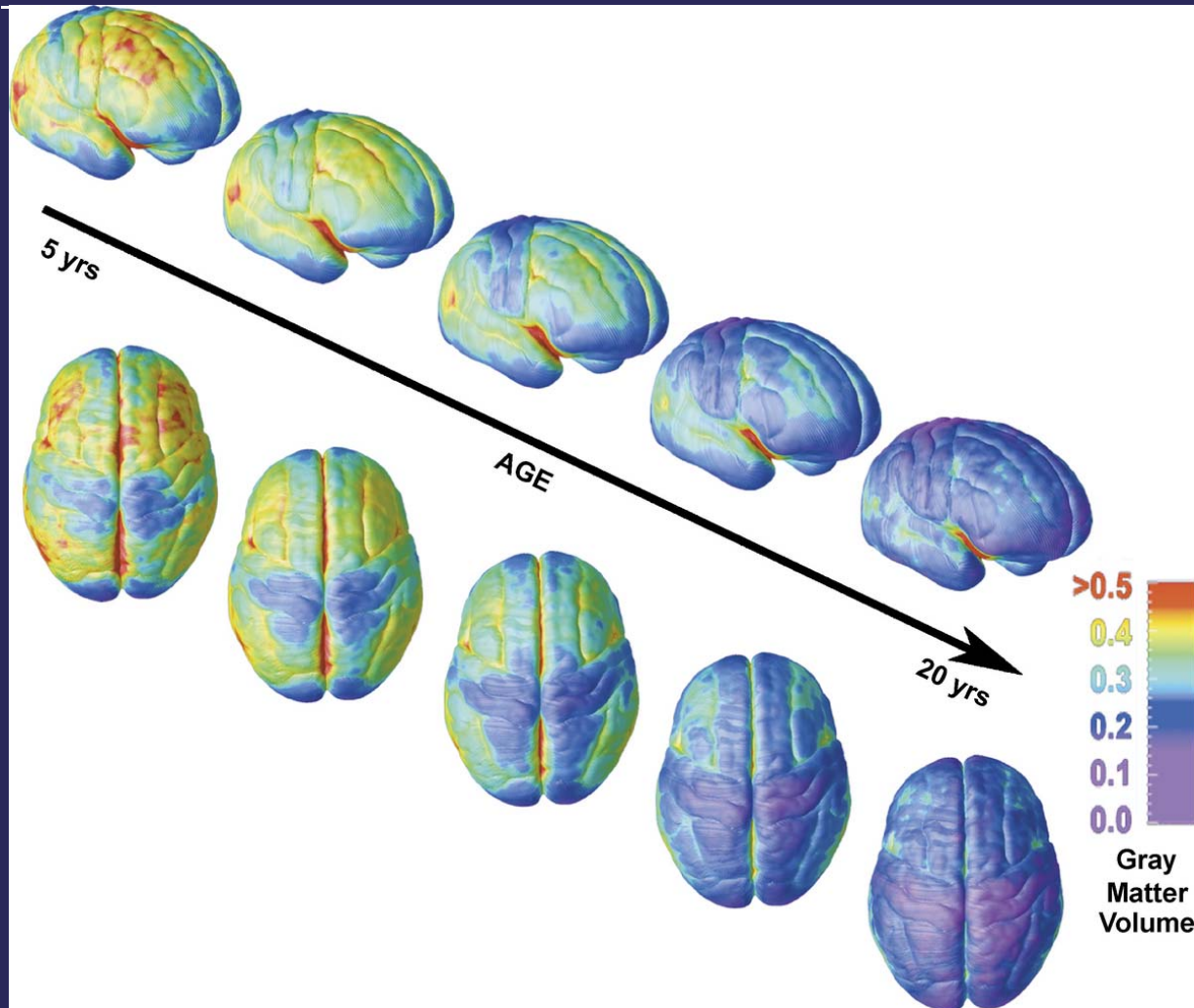
Opioid Addiction

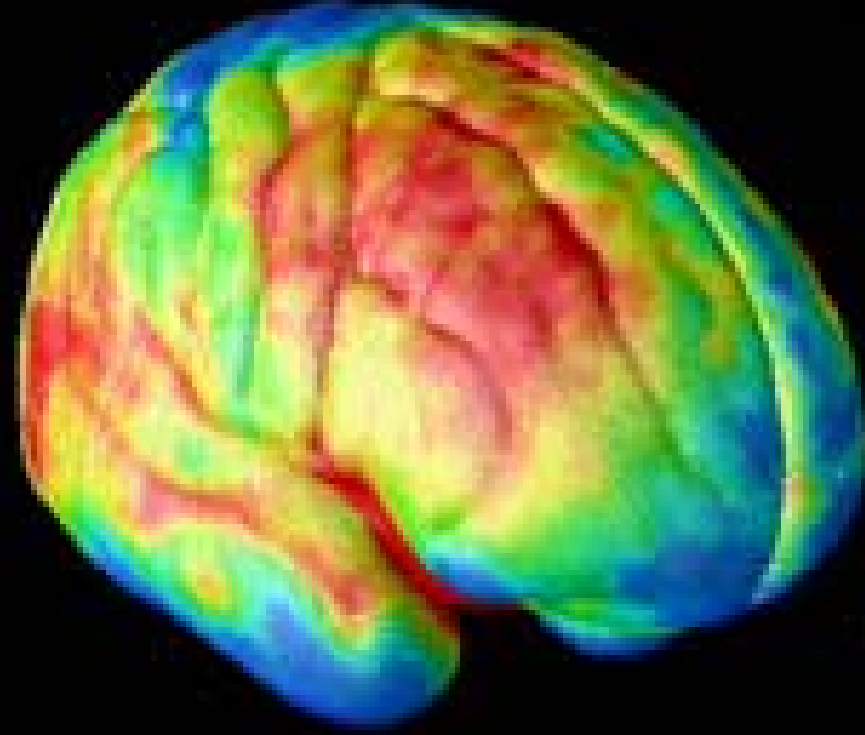
- Opioids attach to specific receptors - **mu** receptors.
- Activation of these receptors - a **pleasure response**.
- Repeated stimulation of these receptors leads to **tolerance** – requiring more drug for same effect.



Adolescent Brain

Dynamic sequence of gray matter maturation over the cortical surface





Gray Matter Amount

1.0
0.9
0.8
0.7
0.6
0.5
0.4
0.3
0.2
0.1
0.0




In a developing nervous system

- Difficulty in decision making
- Difficulty understanding consequences of behavior
- More vulnerable to memory and attention problems

In the Context of Drug Abuse:

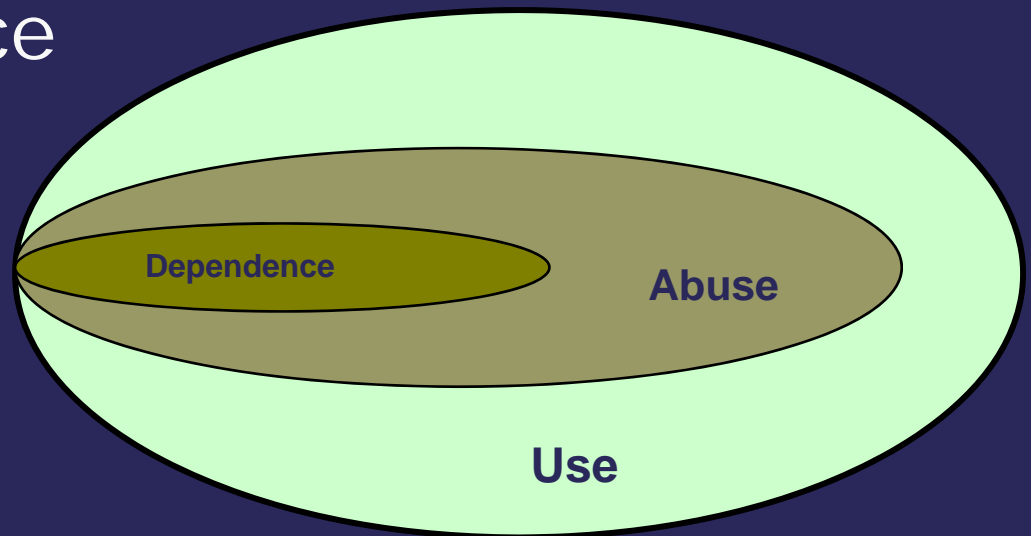
- **Less susceptible to intoxication**
- Animal studies show greater self-administration, and therefore **higher rates of drug dependence**



Clinical Characteristics of Adolescents with OUD

Terminology

- Use
- Abuse
- Dependence
- Substance Use Disorder = Abuse or Dependence



An Overview of Published Studies

Samples	Treatment	Community
Type of opioid	Heroin	Rx Opioid
Age	16-22yrs	>16years
Gender	15-48% Female	More male
Race	Mostly White	Mostly White
School	Poor attendance	Lower performance
Psychiatric	41% clinical Dx	-
Polysubstance	Common	Common
Legal Problems	Common	30-87% sold drugs
% IDU	45-75	5-6

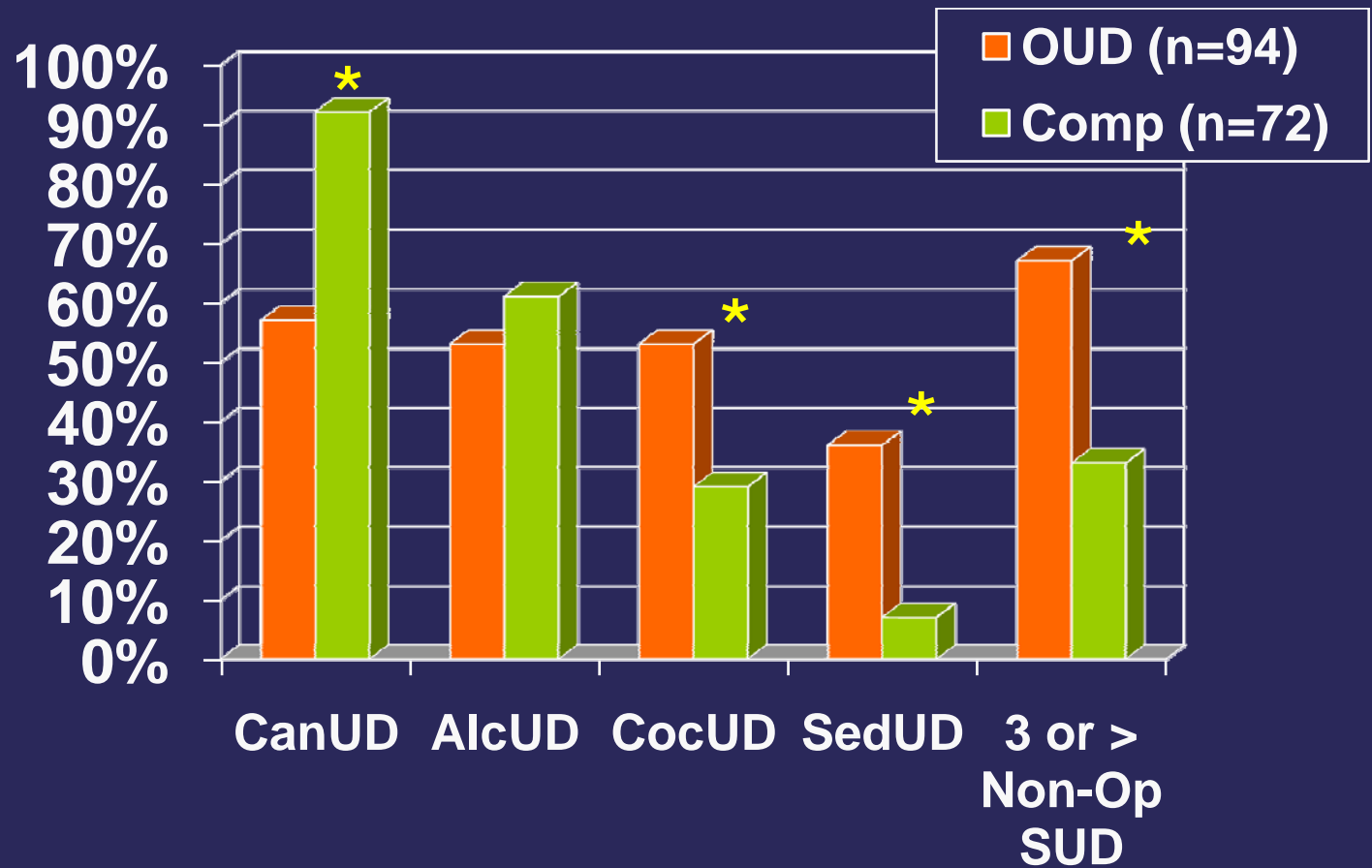
[Hopfer et al., 2000; Clemmey et al., 2004; Gordon et al., 2004; Marsch et al., 2005; Pugatch et al., 2001; McCabe et al., 2005; Sung et al., 2005]

Demographics & Social Characteristics

	<u>OOD</u> (94)	<u>Comparison</u> (72)
<u>Matching Criteria:</u>		
% 18 year-olds	33	33
% Female	45	35
% Residential setting	69	76
% Past 30 D Cocaine use*	46	22
(50 Cocaine Matched cases)		
<u>Other:</u>		
% Caucasian Race*	89	51
% Baltimore City Residence*	30	50
% Still in School*	38	60
% Court Ordered	18	27
% on Probation	35	44
% Guardian both parents	19	23

* Denotes statistical significance

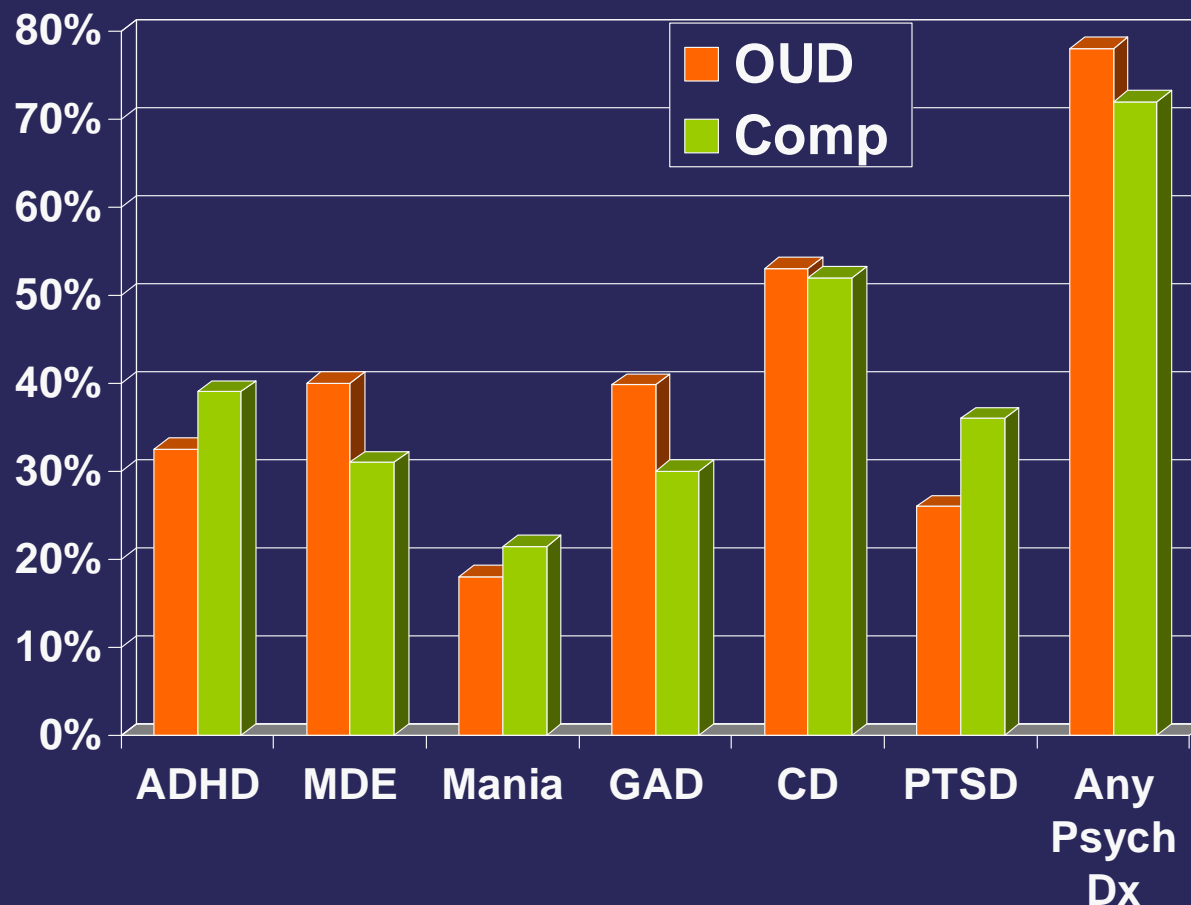
Meeting DSM-IV Criteria Past Year (Non-nicotine/non-opioid) Substance Use Disorders



* Denotes statistical significance

Results based on The Composite International Diagnostic Interview (CIDI)

Meeting DSM-IV Criteria for Current Psychiatric Disorders



Results based on The Diagnostic interview for children and adolescents (**DICA**)

Among OUD Youth: Age of Onset Issues

- Cann/Alc use d/o = **14** y (onset regular use = 12.7-13.5y)
- OUD = **15.3** y (onset reg. use = 15.1 y)
- Coc use d/o = **15.7** y* (onset reg use = 15.3 y)

- ODD/CD/hypomania/mania = **10.3-11.1**y
- GAD = **11.8** y
- MDE = **13.1** y
- PTSD = **13.9** y

HIV Risk Behaviors (Past 30 Days)

	<u>OUD</u>	<u>Comparison</u>
% Any Injection Drug Use*	41	01 (n=1)
% Sharing Needles *	51	-
# Times sharing needles *	25	-
% Sexually active	76	81
% Always Unprotect. Sex	41	29
% 2 or > sex partners	38	35

* Denotes statistical significant differences



Study Summary

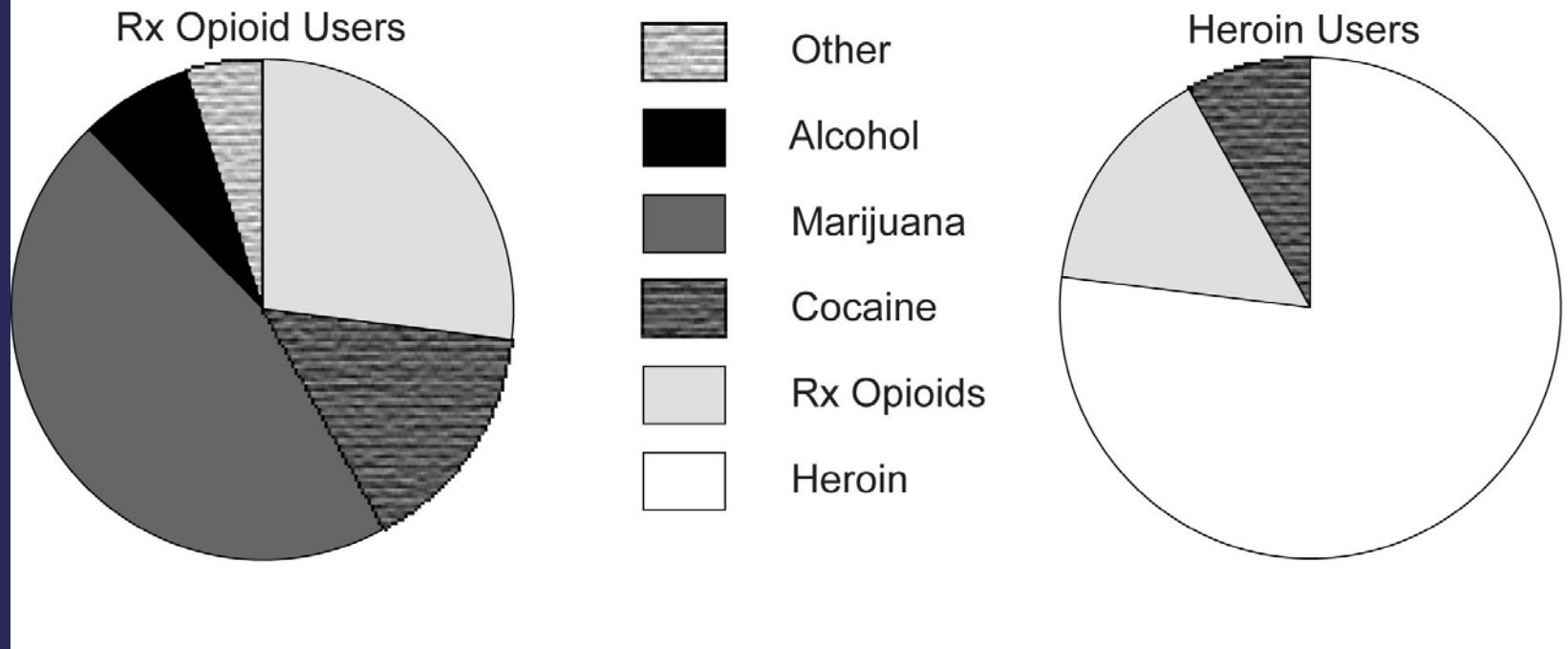
OOD youth more likely to be

- Caucasian, older teens
- Live outside Baltimore city
- Drop out of school
- Concurrent other SUD and 3 or > SUD
- IDU-related HIV/Hepatitis-C risk
- Rapid progression to OUD

But similar to Cann/Alc use d/o youth

- high rates of psychiatric disorders
- high rates of sexual-risk behaviors
- high rates of illegal behaviors

Self-reported “Drug of First Choice”





Treatment Options

Treatment Options for Adolescent OUD

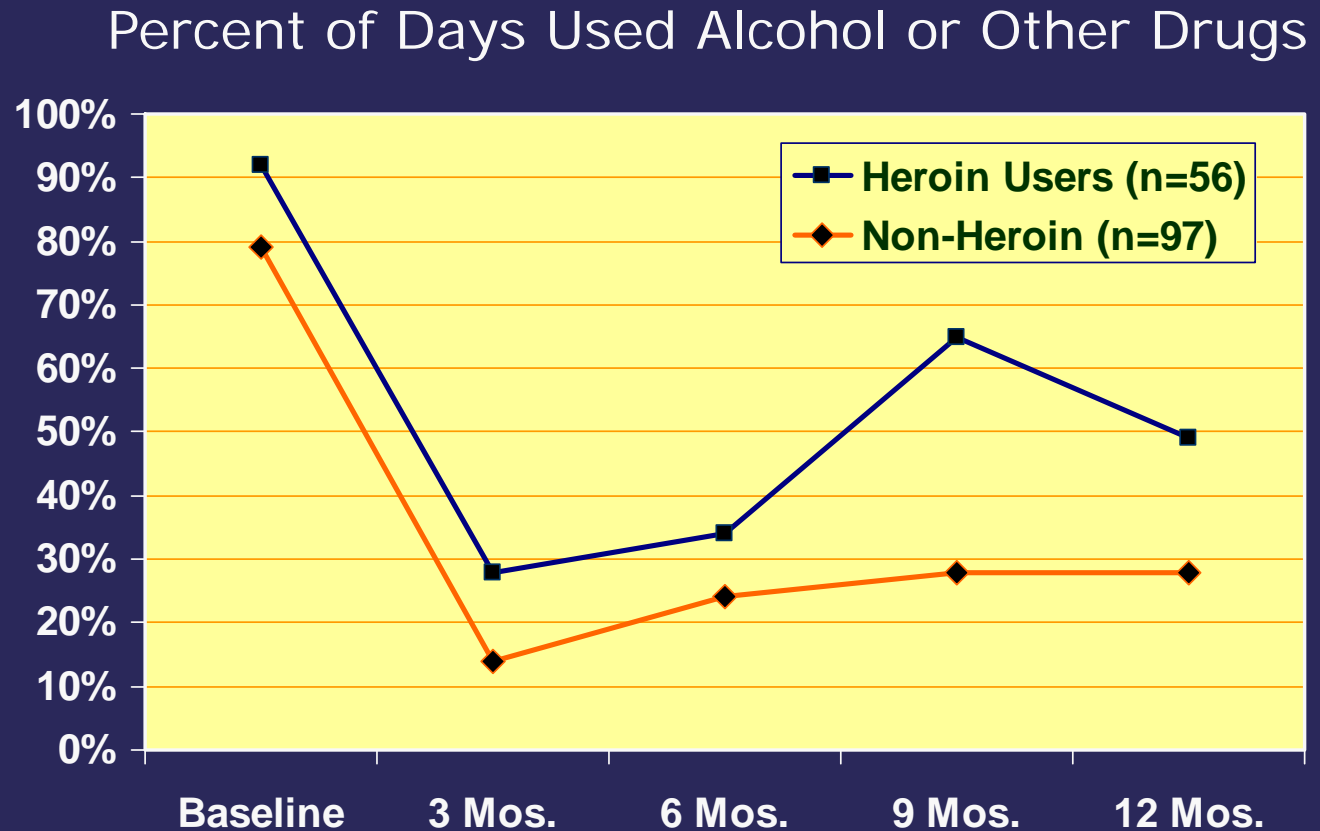
I. Pharmacological:

- Buprenorphine
- Methadone
- Naltrexone

II. Psychosocial:

- Short-term residential treatment (ASAM Level III, non-specific SUD treatment)
- Therapeutic community
- Contingency management
- Individual and group counseling

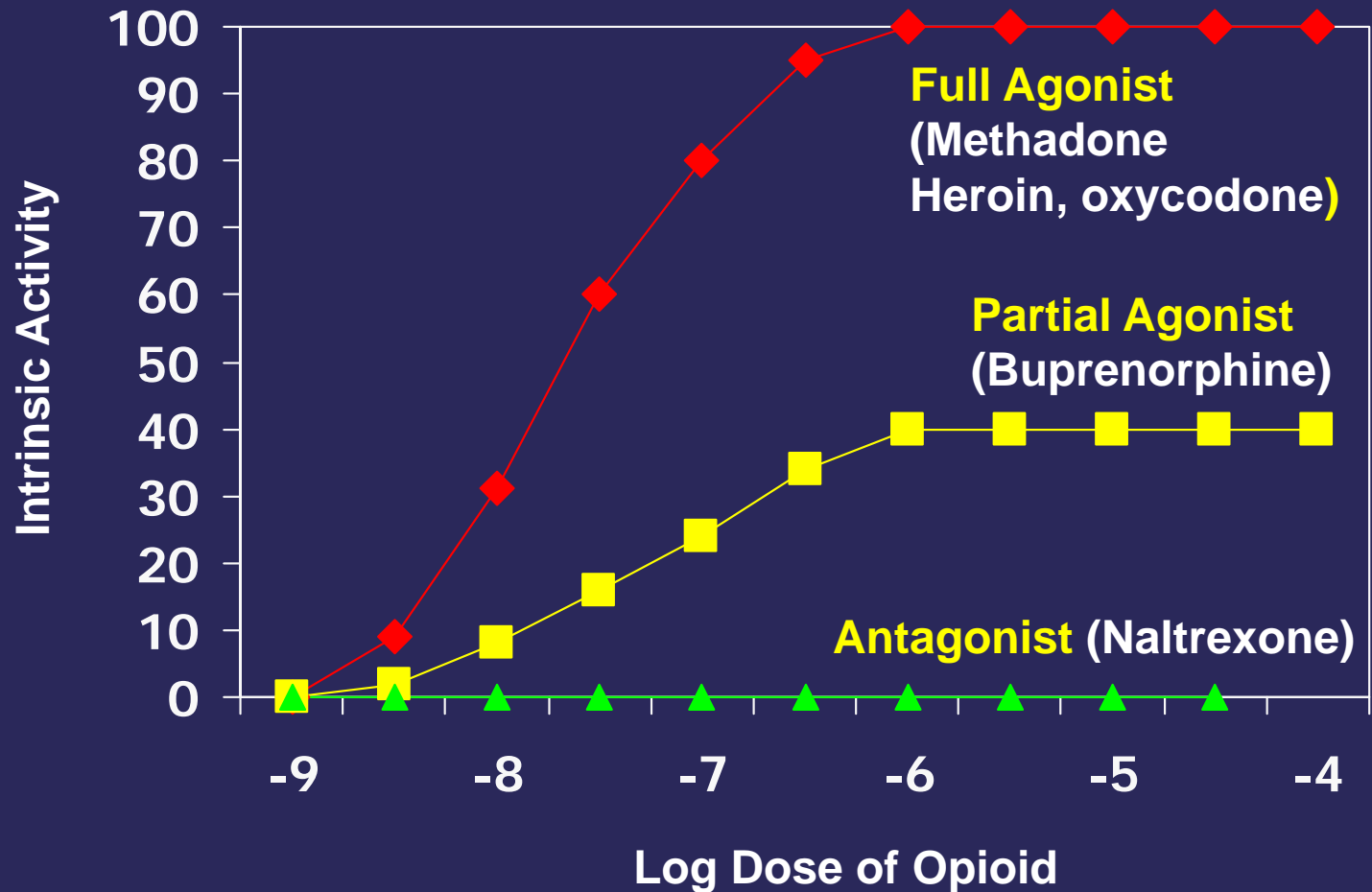
Short-Term Residential Treatment Outcome (Baltimore Site)





Medication-assisted Treatments

Full Agonist, Partial Agonist, and Antagonist of Opioids



Buprenorphine: Preparations and Characteristics

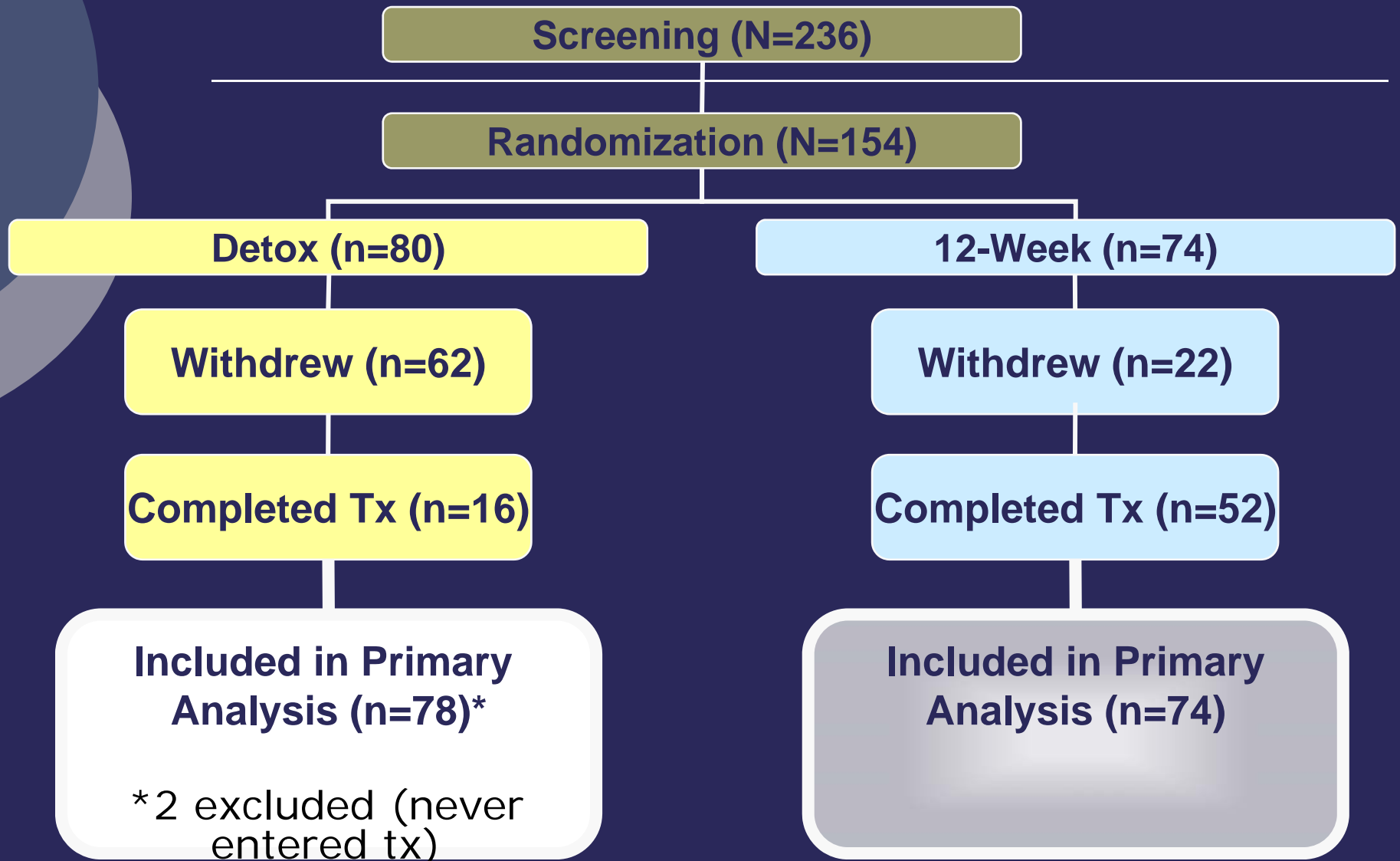
- Buprenorphine combined with naloxone, marketed as **Suboxone®**,
- **Naloxone** is not absorbed sublingually/orally but would precipitate withdrawal **if injected**
- **Subutex®** - stand alone preparation
- **Suboxone®**, available in **2mg** and **8mg** tablets.
- **Suboxone** is to be placed under the tongue and must be allowed to dissolve sublingually

First Controlled Trial using Buprenorphine for Adolescent OUD

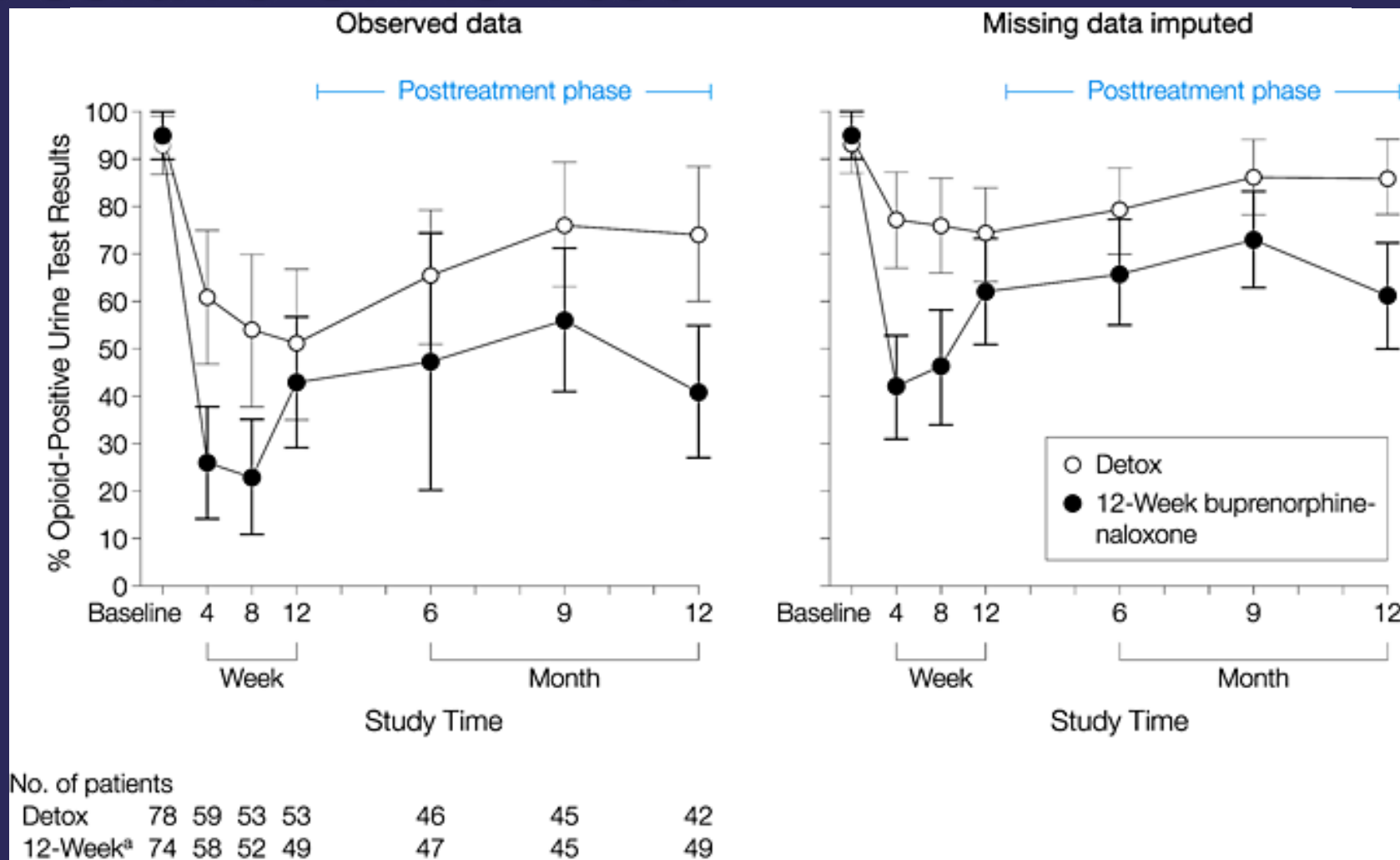
Marsch et al, 2005

- Sample: 38, 13-17 y o.
- Duration: 28-day detox
- Treatment Groups: 6-8mg buprenorphine SL vs. Clonidine 0.1-0.3mg P.O.
- Results:
 - Greater Treatment retention (Bup vs. clon) (72% vs. 39%)
 - > percent of opioid negative urines (64% vs. 32%)

NIDA CTN –Multisite Study Design



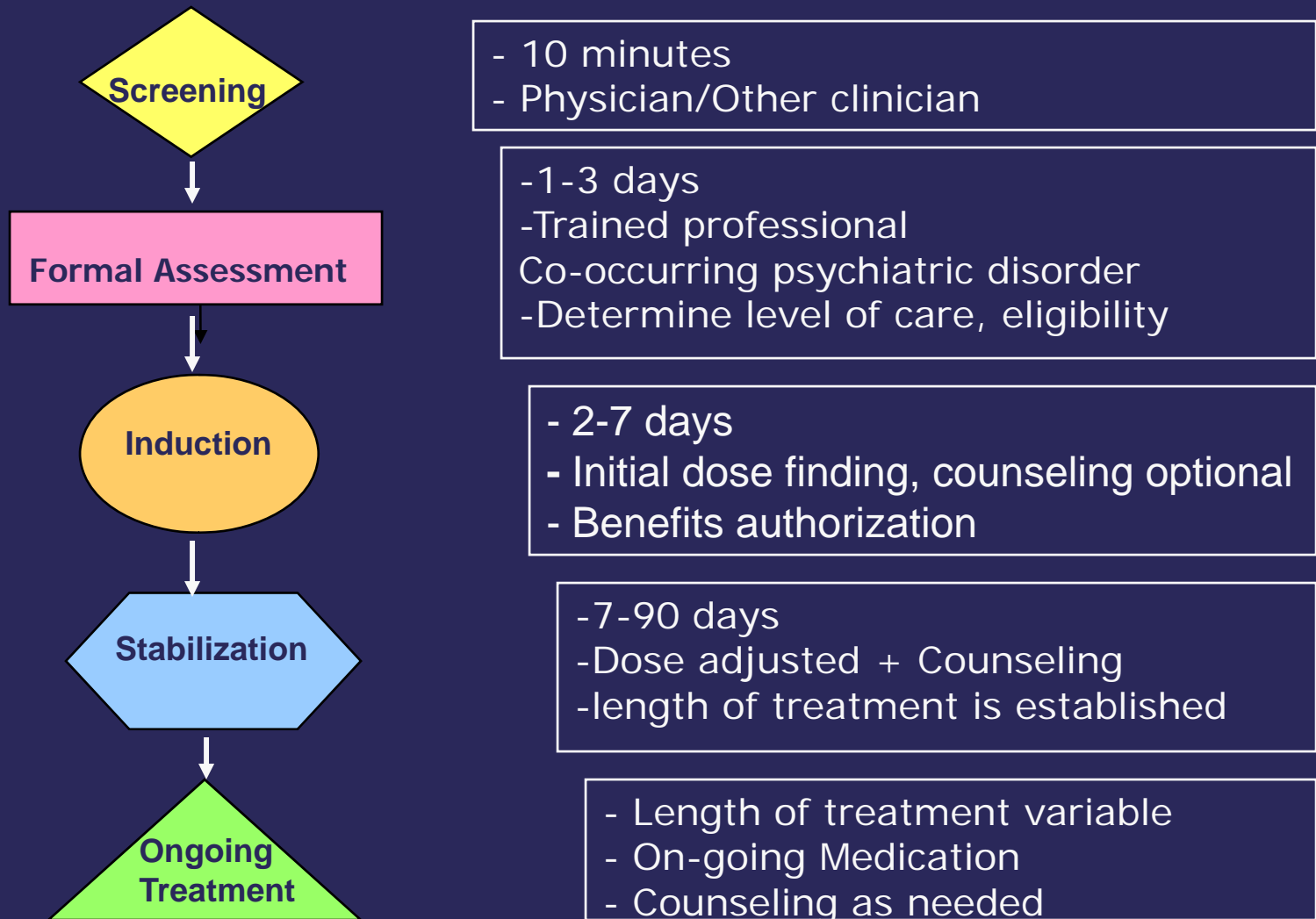
Primary Outcome: Percent of Opioid-Positive Urine Test



Lessons Learned

- Longer treatment seems to be better
- Compared to 2-week detox group, the 12-week group showed:
 - Fewer opioid positive urines
 - Greater retention in active treatment phase
 - Lowered use of marijuana and cocaine use and injection drug use
 - Effect only during active treatment with buprenorphine

Buprenorphine Tx – A Flow Diagram



Treatment Phases

○ Detoxification

- Not effective as stand alone (Mattick, 1996)
- High rates of relapse (Broers 2000, Vaillant, 1988)

○ Maintenance

○ Medication-free

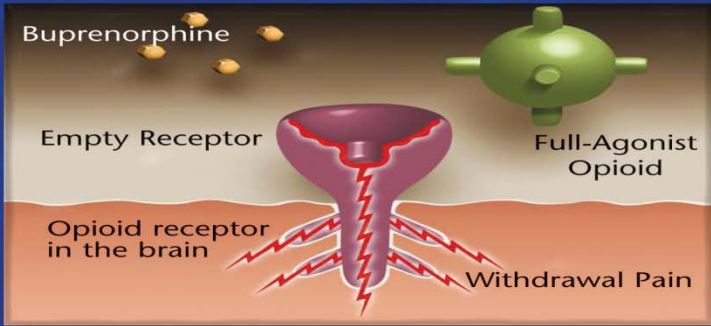
Future Directions

- Are there differences in Tx outcomes between Rx opioid and heroin users?
- Additional questions regarding buprenorphine Tx - duration, dosing, medication compliance issues, etc.
- Is naltrexone effective for this age group? Will it be better accepted?
- Explore other psychosocial treatments as platform treatments – e.g. CM
- Examine integration of psychiatric and HIV-risk reduction treatments

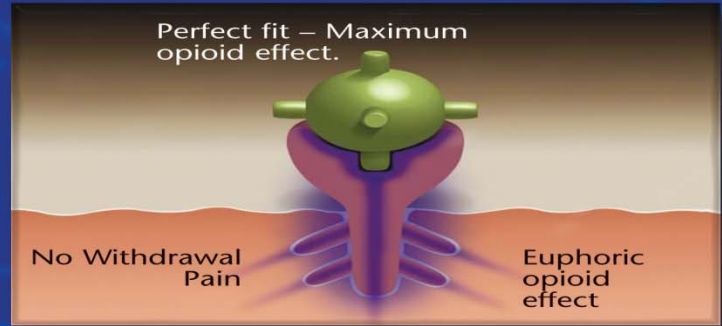


Online Resources

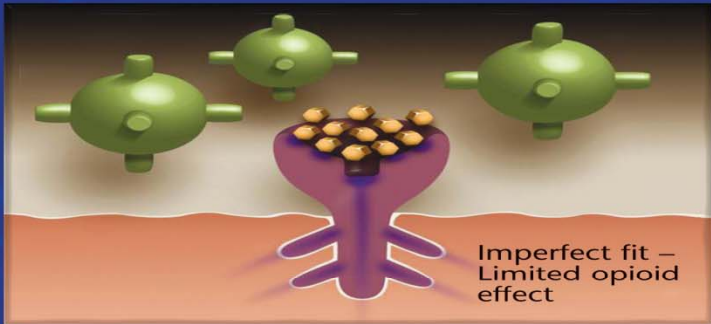
How Buprenorphine Works



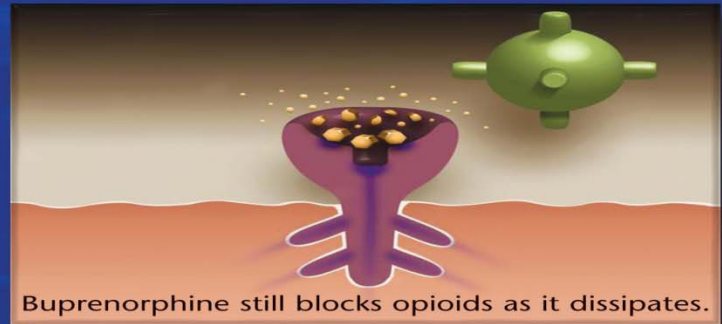
Opioid receptor is empty. As someone becomes *tolerant* to opioids, they become less sensitive and require more opioids to produce the same effect. Whenever there is an insufficient amount of opioid receptors activated, the patient feels discomfort. This happens in withdrawal.



Opioid receptor filled with a full-agonist. The strong opioid effect of heroin and painkillers can cause euphoria and stop the withdrawal for a period of time (4-24 hours). The brain begins to crave opioids, sometimes to the point of an uncontrollable compulsion (addiction), and the cycle repeats and escalates.



Opioids replaced and blocked by buprenorphine. Buprenorphine competes with the full agonist opioids for the receptor. Since buprenorphine has a higher affinity (stronger binding ability) it expels existing opioids and blocks others from attaching. As a partial agonist, the buprenorphine has a limited opioid effect, enough to stop withdrawal but not enough to cause intense euphoria.



Over time (24-72 hours) buprenorphine dissipates, but still creates a limited opioid effect (enough to prevent withdrawal) and continues to block other opioids from attaching to the opioid receptors.


The above illustrations are for educational purposes and do not accurately represent the true appearance.



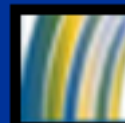
The National Alliance of Advocates
for Buprenorphine Treatment
naabt.org

10M 6/07
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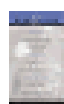


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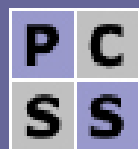
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PCSSmentor.org

Physician Clinical Support System

An Educational Resource for Those Treating Patients with Opioid Dependence

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What is the Physician Clinical Support System? (PCSS)

The SAMHSA-funded PCSS is designed to assist practicing physicians, in accordance with the Drug Addiction Treatment Act of 2000, in incorporating into their practices the treatment of prescription opioid and heroin dependent patients using buprenorphine.

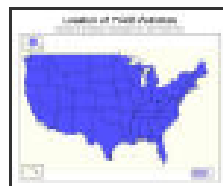
The PCSS service is available, at no cost, to interested physicians and staff, to assist in implementing office-based treatment of opioid dependence with buprenorphine. The essential elements of the PCSS are a national network of trained physician mentors with expertise in buprenorphine treatment and skilled in clinical education, who will be supported by [NATIONAL EXPERTS](#) in the use of buprenorphine and a [MEDICAL DIRECTOR](#).

The [PCSS MENTORS](#) are members of medical specialty societies and provide mentoring support and educational services based on evidence-based practice guidelines. The efforts of PCSS are coordinated by a [STEERING COMMITTEE](#) composed of representatives from the Federal government, the leading addiction medicine societies, along with primary care and psychiatric organizations that represent the target physician populations.

It is estimated that in its first year of operation the PCSS will provide clinical support services to primary care physicians, pain specialists, psychiatrists, and other non-addiction medical practitioners in an effort to increase access to this form of treatment. The PCSS serves to significantly increase access to buprenorphine treatment among the millions of untreated opioid dependent patients.

The PCSS is designed to offer support to clinicians on a number of [TOPICS](#).

The PCSS is active in 48 states, Washington DC, and Puerto Rico. Click [here](#) or on the image below to see the [PCSS ACTIVITIES MAP](#).



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