Treatment of Heroin or Other Opioid Addiction in Adolescents

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Recreational Heroin & Other Opioid Use Among Youth

 Recreational use and dependence on heroin and other opioids among adolescents is a significant and, in some countries, a growing public health concern.

In the U.S., the prevalence of heroin use among 8th, 10th, & 12th graders increased from 0.4-0.6% a decade ago to 1.0-1.6% in recent years (Monitoring the Future, 2006)

About 13% of 8th graders, 17% of 10th graders & 27% of 12th graders say heroin is "fairly or very easy to get" (MTF, 2006)

Recreational Heroin & Other Opioid Use Among Youth

 The increased number of young heroin users has been largely attributed to the decreased price and increased purity of heroin, which allows for intranasal use.

Purity of heroin in U.S. increased from an average of approx.
7% a couple of decades ago to approx. 69% (DEA, 2003)

 Many adolescents initiate heroin use by snorting it; however, they are at great risk of becoming injection drug users.

 Heroin-using adolescents have the highest rate of injection drug use compared with youth using other substances.

Recreational Heroin & Other Opioid Use Among Youth

- 2.6%, 3.8% & 4.3% of 8th, 10th & 12th graders, respectively used OxyContin, and 3%, 7% & 9.7% used Vicodin in last year (Monitoring the Future, 2006)
- About 13% of 8th graders, 22% of 10th graders & 40% of 12th graders say narcotics are "fairly or very easy to get" (MTF, 2006)
- Opiates are currently the second most commonly used illicit drugs among youth in the U.S.
- These trends of increased recreational opioid use have paralleled a sizeable and continuing growth of the availability of new prescription analgesics available in the pharmaceutical market.

Research on Treatment for Opioid-Dependent Adolescents

- A few treatment studies were conducted in the 1960s and 1970s with opioid-dependent youth
- These studies typically did not have control groups or use random assignment, most did not specifically focus on youth under age 18, & may not reflect characteristics of the current cohort of opioid-abusing youth.
 - We launched a line of clinical research to identify effective treatments for this understudied population of youth.

Overview of First, Randomized, Controlled Trial

(Marsch et al., 2005, Archives of General Psychiatry)

 First study in science-based, data-driven effort to produce information needed to effect large-scale change

 Double-blind, double-dummy study designed to compare the relative efficacy of buprenorphine and clonidine in the detoxification of opioid-dependent youth (28-day detoxification; ages 13-18 eligible)

 Informed by the scientific literatures on both effective treatment for opioid-dependent adults & effective treatment of adolescent substance abusers in general

Behavioral Interventions

- All adolescents were also provided with a multicomponent, behaviorally-based treatment program:
 - Individual behavior therapy, including family therapy (based on efficacious Community Reinforcement Approach)
 - Voucher-based Contingency Management (incentives for drug abstinence as measured via thrice weekly urinalysis and clinic attendance)
 - Outreach component to engage adolescents in recreational and other activities to increase non-drug sources of reinforcement

Post-Detoxification Interventions

- All adolescents were provided with 2 months of aftercare, including:
 - Individual Counseling
 - Urinalysis (semi-quantitative)
 - Naltrexone A pure opioid antagonist; prevents receptor activation by other opioid compounds and blocks effects of other opioid drugs
 - Referral to a community-based treatment facility

Participant Characteristics

| CHARACTERISTIC (% or M + SD) | BUPRENORPHINE | |
|---|----------------------|------------|
| Age | 17.3 (0.7) | 17.4 (0.7) |
| Age of First Opiate Use | 15.0 (1.6) | 14.7 (1.7) |
| Gender (% Male) | 50% | 28% |
| Race (% Caucasian) | 100% | 94% |
| Route of Opiate Use (% Injecting) | 33% | 39% |
| Primary Opiate Used (% Using Heroin) | 55% | 50% |
| # of Days Used Opiates in Last 30 Days | 27.7 (3.0) | 27.7 (4.8) |
| # of Prior Outpatient S. Abuse Treatmen | ot 0.9 (1.05) | 1.1 (1.13) |
| # of Prior Inpatient S. Abuse Treatment | 0.8 (1.06) | 0.4 (0.85) |

Other Participant Characteristics/ Life History Variables

 Participants reported significant exposure to risk and high levels of risk behavior. For example,

93% had a family member who drank/used drugs regularly; 45% had a family member with significant mental health problems

44% had experienced a significant family crisis and 41% had someone close to them reject them

36% had witnessed severe violence or abuse, and 31% percent of female participants reported having been raped

Over half (53%) had a family member who engaged in illegal activity, and over 70% had witnessed the arrest of a friend, relative, or neighbor

Other Participant Characteristics/ Life History Variables

Participants criminal activity included:

91% had committed a crime, most commonly shoplifting (73%) & drug dealing (57%)

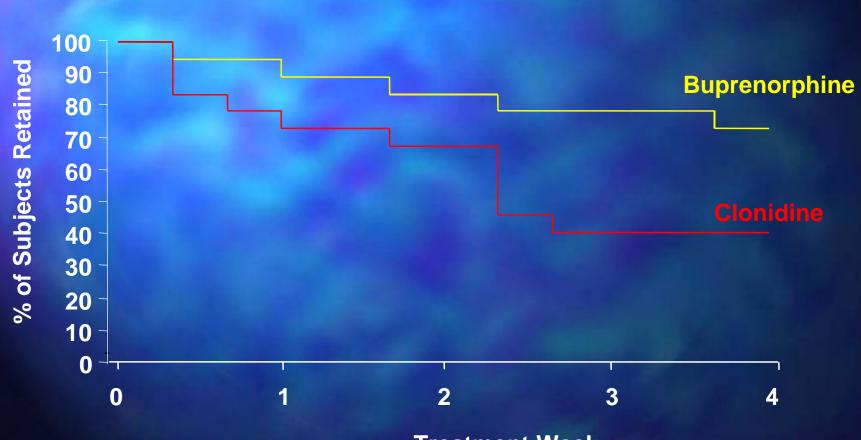
Age of first occurrence of criminal activity was 14 years on average

54% had been picked up by police

42% had been on probation

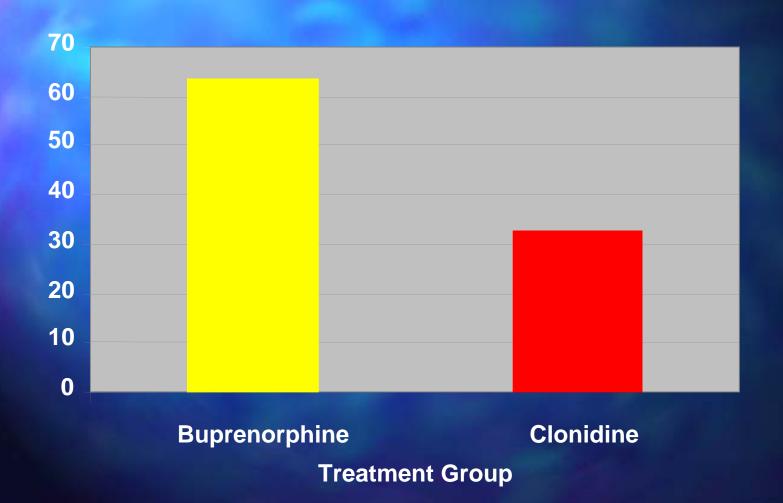
24% spent time in juvenile detention or jail

Treatment Retention

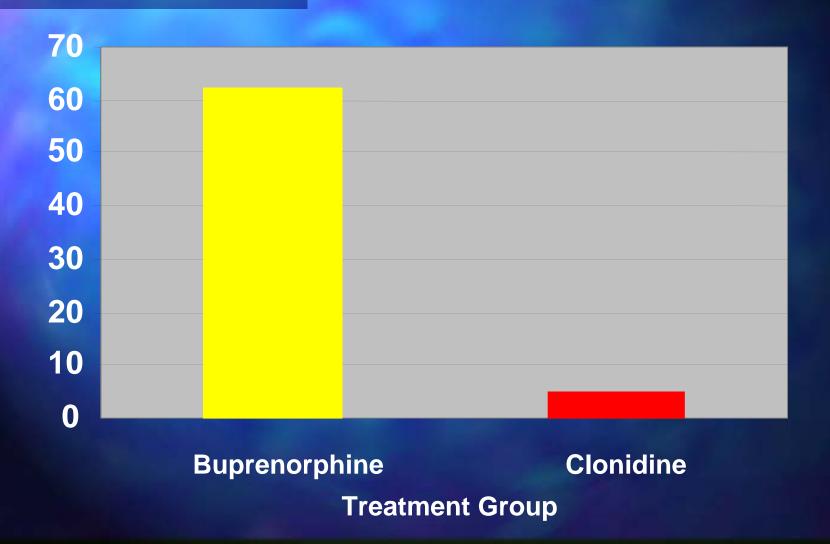


Treatment Week

Mean Percent Opiate Abstinence

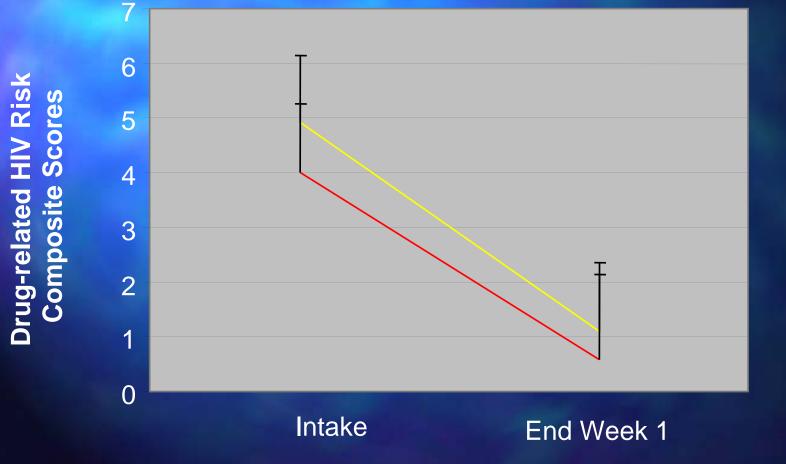


Percent of Participants Initiating Natrexone Post-Detoxification



HIV Risk Behavior (HRBS) (time effect: <u>p</u> = .0005)

BuprenorphineClonidine



Sub-group Analyses: Outcome by Gender

Baseline characteristics of males & females were similar.

 Both males & females had significantly better outcomes from buprenorphine & behavioral treatment compared to clonidine & behavioral treatment.

 However, females achieved greater opioid abstinence and reductions in HIV risk behavior relative to males during buprenorphine/behavioral treatment.

Current Clinical Research in New York City

 Can treatment outcomes be improved if duration of medication taper is increased?

<u>Phase 1:</u> Random Assignment to 28 or 63-day buprenorphine taper

 Can incentives contingent on naltrexone consumption increase compliance with naltrexone and reduce relapse?

<u>Phase 2:</u> Random Assignment to receive/not receive voucher incentives contingent on naltrexone

 Do various sub-populations of opioid-dependent youth have differential treatment outcomes (e.g., based on demographics, other drug use, psychological variables)?

Summary of Research Results to date

- Expanded science-based prevention & treatment interventions are needed for the emerging cohort of opioid-dependent adolescents
- Although both clonidine and buprenorphine were shown to be quite safe, results indicate that combined behavioral & buprenorphine treatment is more efficacious than combined behavioral & clonidine treatment
- Due to the nature and pharmacology of opioid drugs, pharmacotherapy appears to be a critical component of successful treatment of opioid dependence (to stabilize brain neurochemistry).

Summary of Research Results to date

- Naltrexone appears to have considerable utility in preventing relapse to opioid use after an adolescent is no longer dependent.
- Treatment outcomes appear optimal when medication is provided along with intensive behavioral therapy (to promote alternative rewarding behaviors & strengthen inhibitory control).
- Psychotherapy should address adolescent-specific issues (e.g., school re-entry, securing a degree, self-control training)
- Psychosocial treatment should address high rates of psychiatric comorbidity to be optimally effective

Summary of Research Results to date

- Early intervention is key (to prevent transition from abuse to dependence or from intranasal to injection opioid use)
- Extended therapeutic treatment may be important for relapse prevention.
- Given this group's extensive involvement in the criminal justice system, there may be many opportunities for offering effective treatment to youth within this system
- Providing science-based treatment to this young population greatly reduces their likelihood of continued and escalating substance involvement and may prevent a substance-abusing life trajectory.