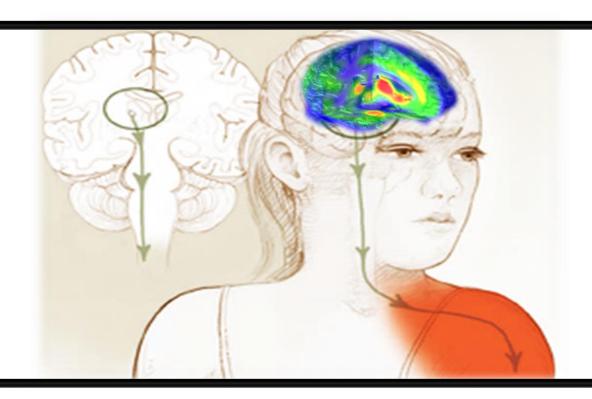
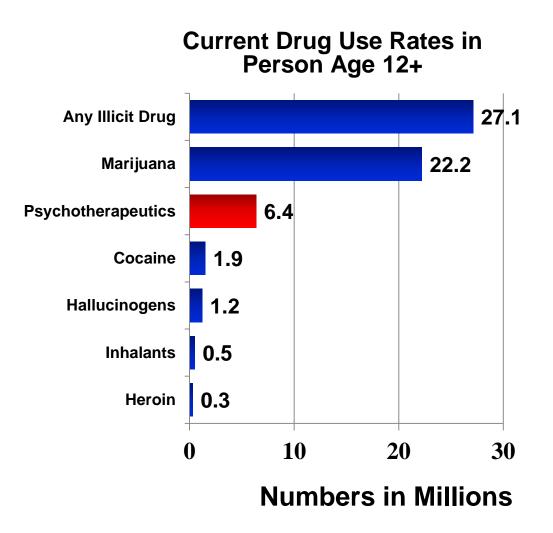
Chronic Pain in Women and its Relationship to Opioid Addiction

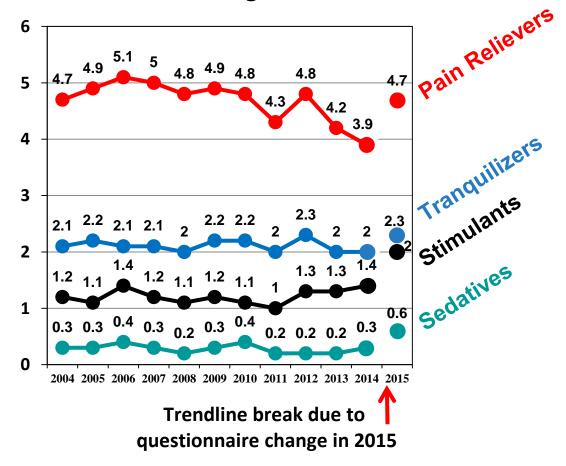




Prescription Drug Misuse/Abuse is a Major Problem in the US

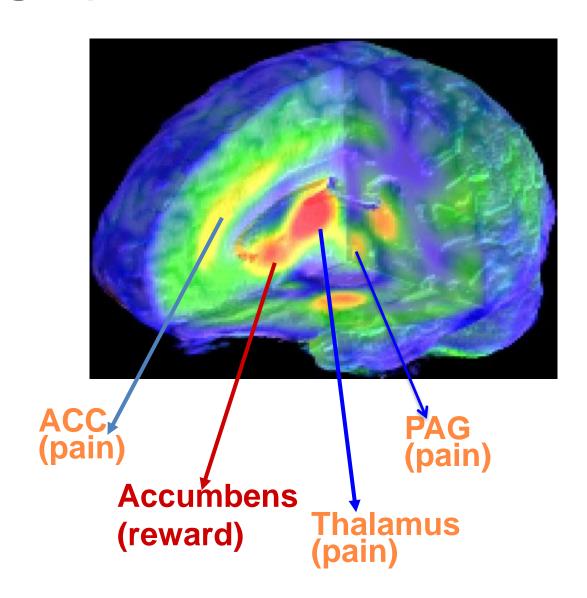


Past Year Nonmedical Use of Psychotherapeutic Drugs Persons Ages 12+

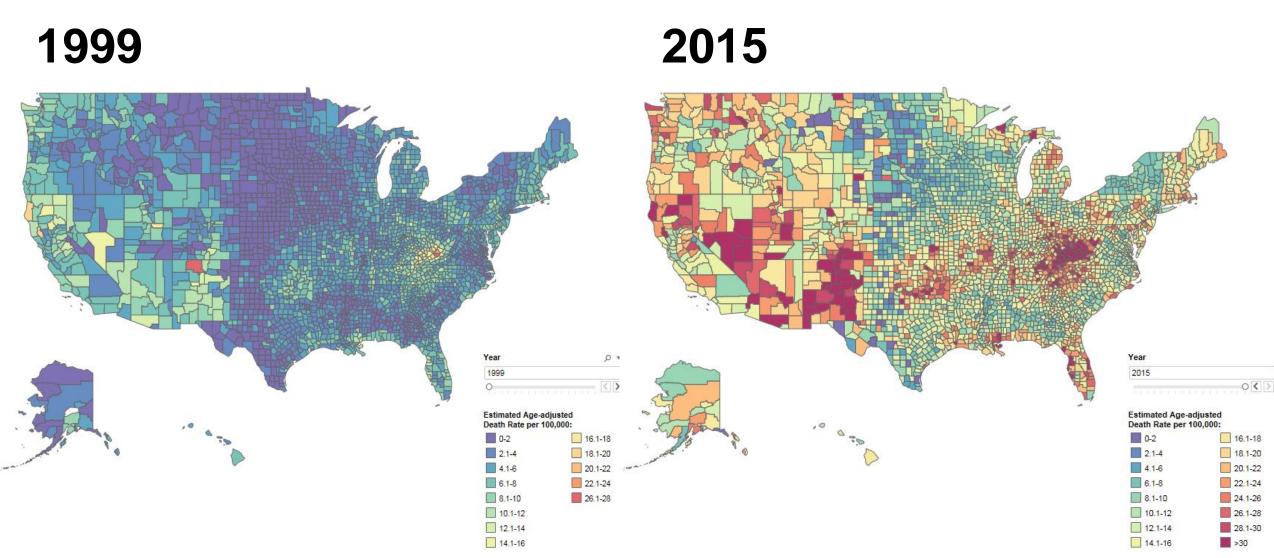


Source: SAMHSA, 2015 National Survey on Drug Use and Health, 2016.

Analgesic Mechanisms of Mu Opiate Drugs (Heroin, Vicodin, Morphine)



Overdose Death Rates

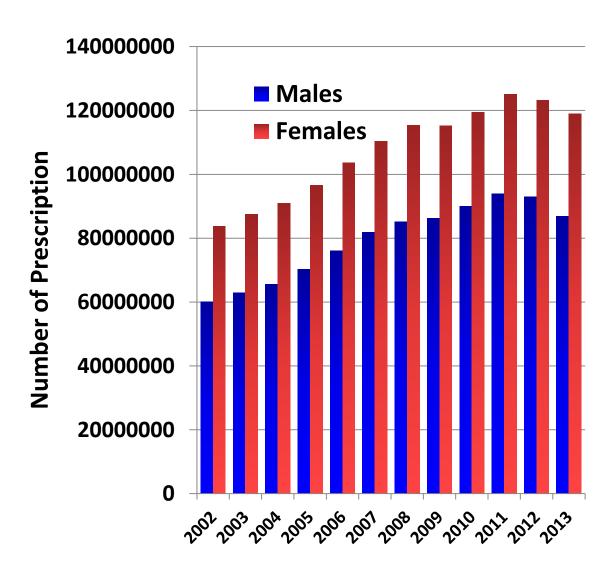


Opioid Prescriptions 1991-2011

Opioids 250 Hydrocodone Oxycodone 200 Prescriptions (millions) 150 76 78 80 86 91 96 100 50 91 07 09 11

IMS's Source Prescription Audit (SPA) & Vector One®: National (VONA)

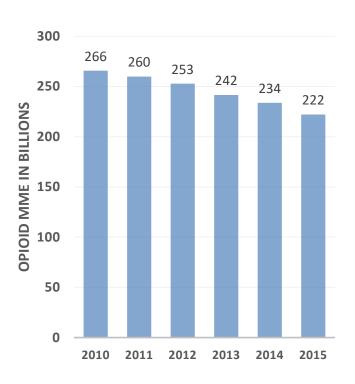
Number of Opioid Prescriptions U.S. Retail Pharmacies, 2002-2013



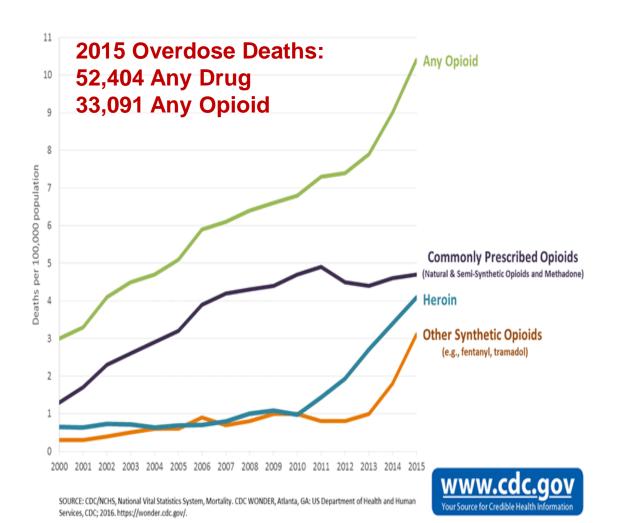
Source: IMS Health, National Prescription Audit

Opioid Prescriptions have started to Decrease but Opioids Fatalities are still Increasing

Opioid morphine milligram equivalents (MME) dispensed fell by over 15% from 2010-2015



Opioid OD Deaths US, 2000-2015



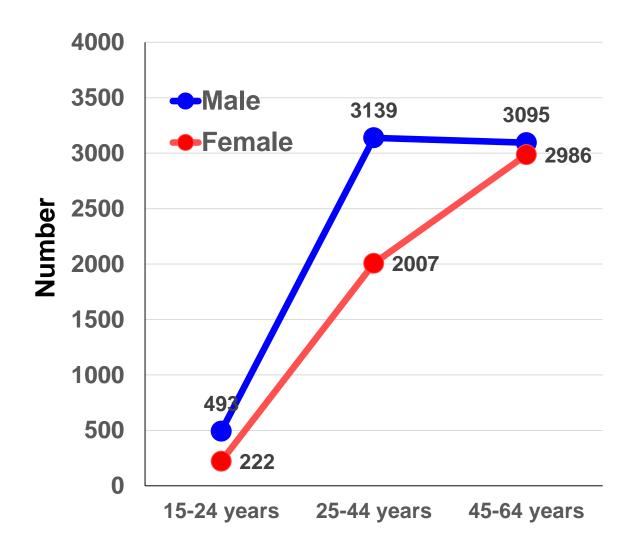
Source: IMS Health, U.S. Outpatient Retail Setting

The Changing Face of Heroin Use in the US

90 Men 80 Women 70 Total Sample, % 60 40 30 20 10 1960s 1970s 1980s 1990s 2000s 2010s Decade of First Regular Opioid Use

Cicero TJ et al., JAMA Psychiatry 2014.

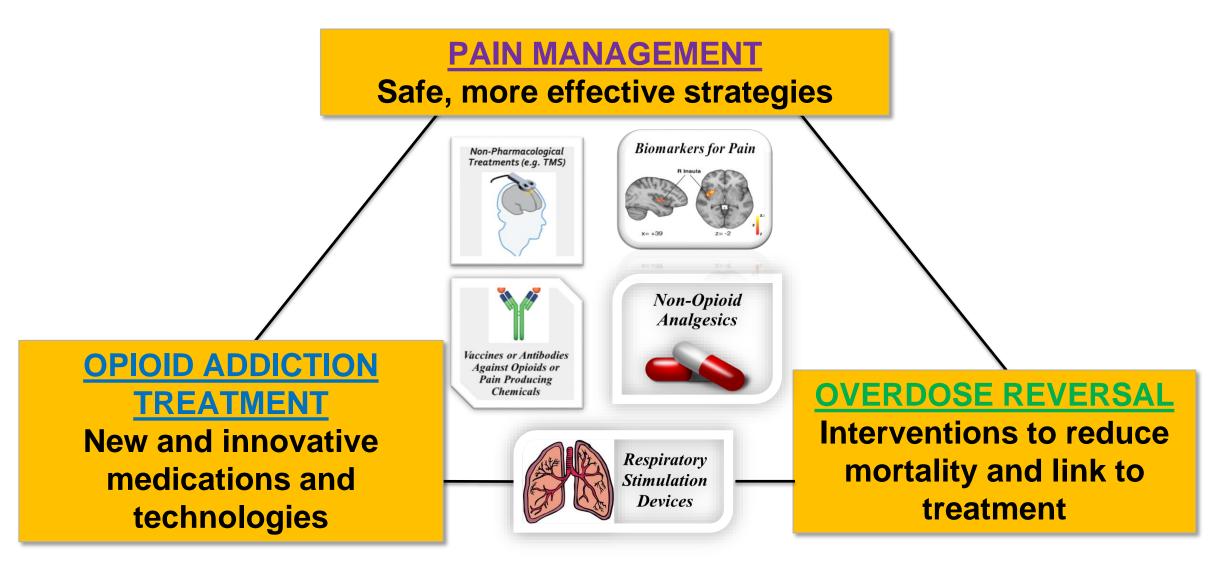
Deaths Involving Natural and Semi-Synthetic Opioids

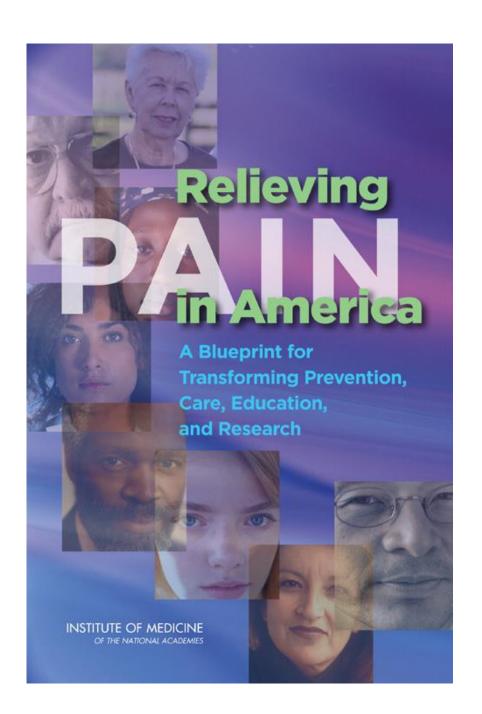


Rudd RA et al., MMWR Morb Mortal Wkly Rep 2016

NIH OPIOID RESEARCH INITIATIVE

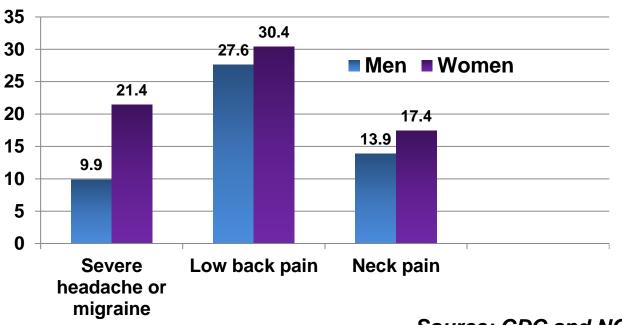
Using Research to End the Opioid Crisis





Women suffer more pain in many categories

Rates of U.S. Adults > 18 and Older Reporting Pain, 2015

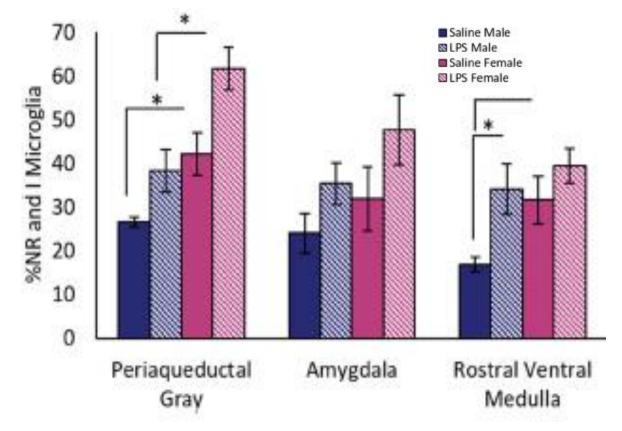


Source: CDC and NCHS, 2015

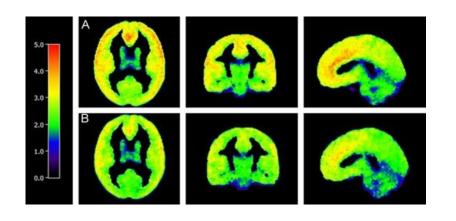
 Women's reports of pain more likely to be dismissed

Sex Differences in Microglia Activity in Periaqueductal Gray in Response to Inflammatory Pain

Treatment with TLR4 Agonist LSP Increased Microglial Activation in Females

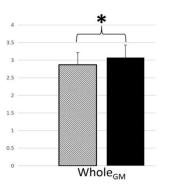


Gender Differences in Kappa Opioid Receptor Availability



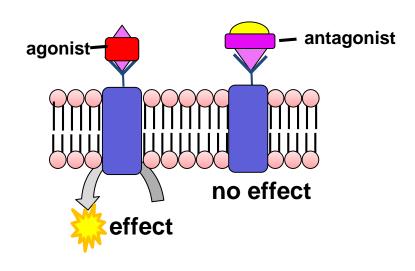
Males

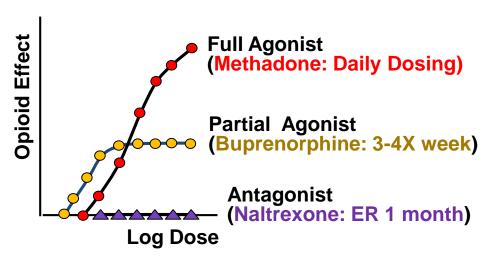
Females



- Males had higher K receptor availability than females presumably from increased dynorphin, which is aversive.
- This might help explain gender differences in pain catastrophizing

Medications Assisted Therapies for OUD





DECREASES:

- Opioid use
- Opioid-related OD deaths
- Criminal activity
- Infectious disease transmission

And INCREASES

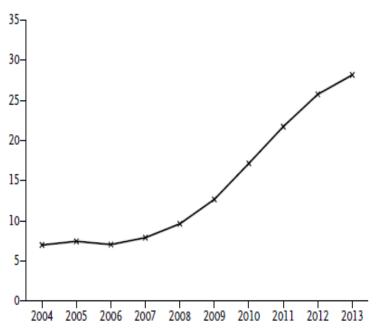
- Social functioning
- Retention in treatment
- Outcomes in NAS

MAT is <u>highly underutilized</u>

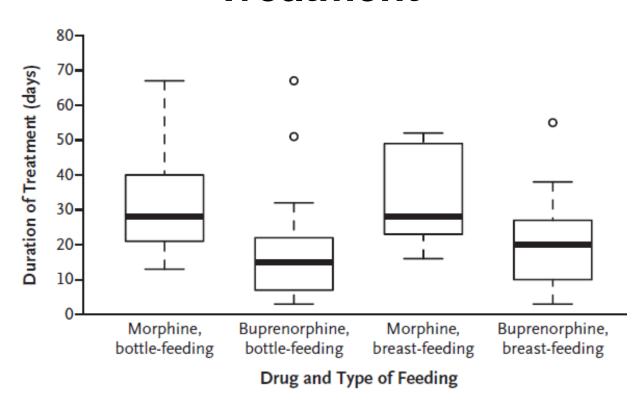
Opioid Use and Misuse During Pregnancy

17% of pregnant women are prescribed an opioid during pregnancy

NICU Admissions for Neonatal Abstinence Syndrome NAS (Number per 1000 Admissions)



Buprenorphine for NAS Treatment

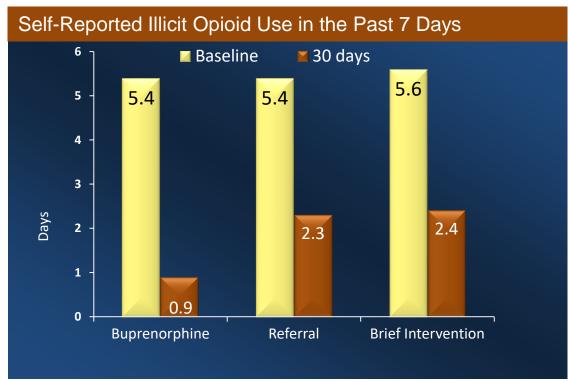


Buprenorphine resulted in a shorter duration of NAS treatment and length of hospital stay than treatment with morphine

Source: Tolia VN, Patrick SW, et al. NEJM 2015;372:2118-2126.

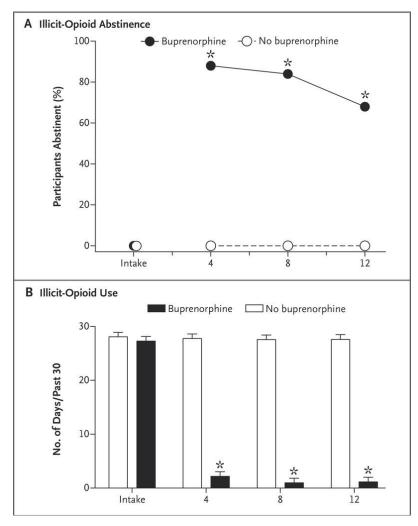
Emergency Department-Initiated Buprenorphine

- Reduced self-reported, illicit opioid use
- Increased engagement in addiction treatment
- Decreased use of inpatient addiction treatment services



D'Onofrio G et al., JAMA April 28, 2015.

Abstinence from Illicit Opioids over 12 Weeks with Interim Buprenorphine



Sigmon SC et al. N Engl J Med 2016.

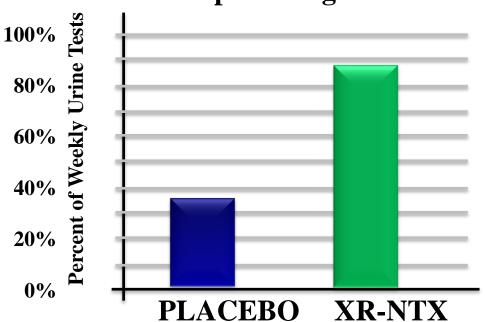
Extended Release Formulations

Vivitrol®



IM Injection q 4 weeks for 24 weeks

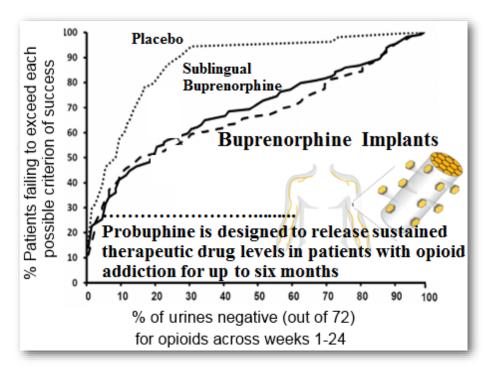
Median % Opioid-Negative Urines



Placebo: N=124 XR-NTX: N=126

PROBUPHINE®





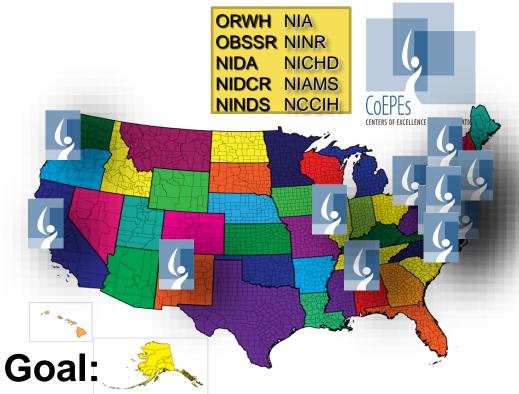
Rosenthal et al., Addiction 2013;105.

FDA approval – May 26, 2016

Education

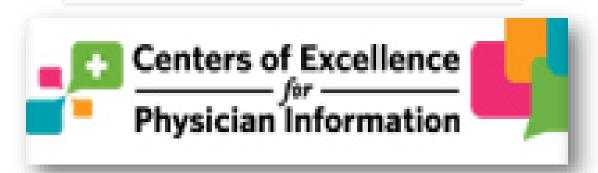
PAIN

NIH Pain Consortium Centers of Excellence in Pain Education



Improve pain treatment through education





Goal:

Prevent SUD and improve outcomes in addiction through education of health care providers

















