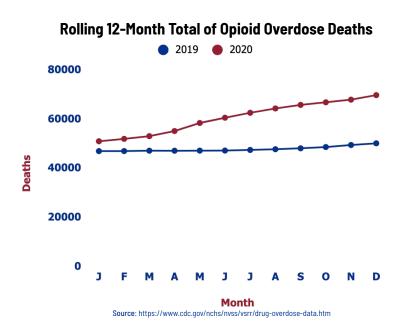
Opioids: The Epidemic Within the Pandemic



Opioid Epidemic The increased prescription of opioids for pain management in the 1990s combined with the rise in distribution and access to heroin have accelerated opioid use in the United States¹. In the last decade, widespread use of synthetic opioids has caused a spike in opioid-related deaths — over **57,000** deaths by overdose in 2020 were due to synthetic opioids^{2,3}. Even with a **44**% decrease in opioid prescriptions in the past decade, drug overdose deaths continue to rise⁴.

Pandemic Impact In the 12-month period beginning in March 2020 — when pandemic shutdowns began in the U.S. — drug overdose deaths from opioids rose 38%². The COVID-19 pandemic has escalated the challenge of combating the already tragic opioid epidemic.

In an analysis of electronic medical records, hospital patients who received a diagnosis of OUD were more likely to have COVID-19⁵. The negative impacts on heart and lung health from OUD increase susceptibility of those with OUD for severe COVID-19.



How does opioid use lead to opioid use disorder?

In addition to alleviating pain, opioids induce feelings of pleasure by hijacking the body's natural reward system. Tolerance to opioids can build after repeated use, leading to increased amounts being required to reach the same pleasurable feelings or to sustain pain relief. **Opioid use disorder (OUD)** — commonly referred to as opioid addiction — is a disease that consists of chronic opioid use despite negative consequences. Opioid withdrawal, which occurs when usage declines or stops, can be difficult to tolerate.

- Of those using opioids for chronic pain, 8%-12% develop an OUD⁶.
- Among heroin users, **80**% first misused prescription opioids.
- OUD is pervasive in the U.S., affecting over **2.1 million** Americans⁷.
- More than 73,000 Americans died of an opioid overdose in 2020².
- According to an estimate from the CDC, OUD costs in the U.S. total \$78.5 billion each year⁸.

What are opioids? Opioids are substances used to manage pain. They are produced naturally in the body, are found in nature, and can be chemically synthesized.

Why is OUD difficult to overcome?

- Long-term opioid use alters brain function.
- Access to care is a major challenge compromising efforts to provide treatment and care^{9,10,11,12}.
- People with OUD often face other health conditions: alcoholism, chronic pain, and Hepatitis C¹³. In 2016,
 62% of adults with OUD had a co-occurring mental illness (anxiety, depression, PTSD, ADHD, etc.)¹⁴.
 These comorbidities can further complicate treatment and recovery.

How are overdoses treated?

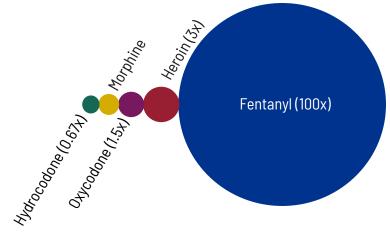
Naloxone is a life-saving medication used to reverse opioid overdose¹⁵. It is available as both a nasal spray (brand name Narcan) and an injectable (brand name Evzio). These forms can be administered without medical training.

Naloxone is available in all 50 states and can be obtained without a prescription in most states. While it cannot reverse overdose of non-opioid drugs, it will not cause harm if administered for overdose of other drug classes¹⁵.

COVID-19's impact on overdose therapy

During the pandemic, there was a 26% reduction⁴ in the number of naloxone prescriptions filled in retail pharmacies, reducing the prevalence of one of the most powerful tools for reversing opioid overdose.

Opioid Strength (Relative to Morphine)



Sources: https://jpet.aspetjournals.org/content/136/1/43 https://www.ncbi.nlm.nih.gov/books/NBK537482/table/appannex6.tab2/

The increase in overdose deaths in 2020 was caused, in part, by the rise in the prevalence of fentanyl³. Fentanyl is a synthetic opioid 100 times more potent than morphine, and is added to other drugs (often without the knowledge of the user) to increase potency.

Therapies for Opioid Use Disorder

While multiple therapies are available to treat OUD, there is no cure.

Pharmaceutical Intervention (Medication-Assisted Treatment)

Buprenorphine: An opioid medication used to treat OUD — it has reduced side effects compared to methadone in the supernorphine and naloxone: Combining naloxone with buprenorphine helps patients manage with drawal symptoms and reduces the misuse potential of buprenorphine 17 .

Methadone: An opioid medication used to manage opioid withdrawal symptoms — it can be used as a maintenance therapy 18 .

Naltrexone: A treatment that blocks the effects of opioids and reduces cravings^{i,19}.

ⁱThe injectable form is long-lasting, which can help with medication adherence.

Behavioral Intervention

Contingency management: Rewards patients with gifts or incentives for drug abstinence²⁰.

Community reinforcement: Weekly therapy sessions help patients overcome their addiction²¹.

12-step therapy program: Structured self-help groups work through a 12-step process to achieve abstinence²².

COVID-19 Disrupted Access to Therapy

Research suggests that the COVID-19 pandemic exacerbated the existing shortage of methadone clinics for those pursuing treatment²³. Social distancing and other COVID-19 restrictions prevented in-person 12-step meetings, which are a critical facet of recovery for many people with OUD²⁴. Though some services were made available via telehealth, internet access is a factor limiting who can benefit from telehealth programs²⁵.

Research at Work

Medical University of South Carolina (MUSC)

The Center for Opioid and Cocaine Addiction (COCA) seeks to understand the neurological mechanisms that contribute to relapse and design treatments that prevent relapse.

Icahn School of Medicine at Mount Sinai

Researchers are investigating the genetic basis and neural circuits involved in drug addiction. Funding from NIH's Helping to End Addiction Long-term (HEAL) initiative allows researchers to discover new treatments for drug addiction and accelerate early-phase clinical trials for non-addictive pain therapies.

Virginia Tech

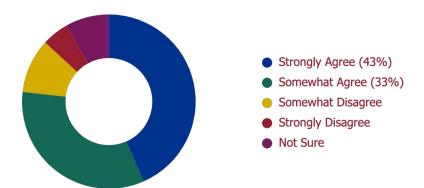
A grant from the National Institute on Drug Abuse (NIDA) at NIH allows Virginia Tech researchers to work on the development of a vaccine to counteract opioid addiction.

Yale University School of Medicine

A grant from NIH's HEAL initiative will enable researchers at Yale to launch the IMPOWR-YOU Research Center, which will harness an integrative approach to conducting opioid research.

Please indicate the extent to which you agree that opioid abuse and addiction is a major problem in your community.

Source: A Research! America poll of U.S. adults conducted in partnership with Zogby Analytics in January 2021.



The Future of Opioid Research

Elucidating the biological basis of opioid addiction, developing better opioid blockers that are more effective against stronger, synthetic opioids, and investigating new methods of treating chronic pain are a few of the ways research will save countless lives in confronting the opioid epidemic.

Emerging investigations looking at the intersection of the opioid epidemic and COVID-19 pandemic range from assessing the effects of the pandemic on harm reduction to the association of chronic pain and COVID-19 diagnosis or complications.

Resolving complex regulatory issues that bear on opioid research can lead to faster progress. There is interest in Congress on this issue, including hearings, such as https://example.com/TheOverdose Crisis: Interagency Proposal to Combat Illicit Fentanyl-Related
Substances and legislation, including the Substances Act of 2021.

Sources:

- 1. Volkow et. al. "The Changing Opioid Crisis: development, challenges and opportunities." 2021 2021, 26(1): 218–233.
- ${\it 2.~"Vital~Statistics~Rapid~Release."~National~Center~for~Health~Statistics.~2021}$
- 3. "Overdose Deaths Reached Record High as the Pandemic Spread." New York Times. 2021.
- 4. "2021 Overdose Epidemic Report." AMA. 2021.
- 5. "COVID-19 and Substance Use." National Institute on Drug Abuse. 2021.
- 6. "Opioid Overdose Crisis." National Institute on Drug Abuse. 2021.
- 7. Dydyk AM, Jain NK, Gupta M. "Opioid Use Disorder." StatPearls. 2021.
- 8. Florence CS, Zhou C, Luo F, Xu L. "The Economic Burden of Prescription Opioid Overdose, Abuse,
- and Dependence in the United States, 2013". Med Care. 2016: 54(10): 901-6.
 9. "Vital Statistics Rapid Release." National Center for Health Statistics. 2021.
- Mallow et al. "Geographic variation in hospital costs, payments, and length of stay for opioid-related hospital visits in the USA." J Pain Res. 2018:11:3079–3088.
- 10. "How much does opioid treatment cost?" NIDA. 2018.
- 11. "Guidelines for the Psychosocially Assisted Pharmacological Treatment of Opioid Dependence." WHO. 2009.
- 12. Hoffman et al. "Opioid use disorder and treatment: challenges and opportunities." BMC Health Serv Res. 2019:19:884.

- 13. "Treating patients with opioid disorders is not just about treating addiction. Here's why." STAT News. 2017.
- 14. "When Addiction and Mental Illness Collide." NIH HEAL.
- 15. "Life-Saving Naloxone from Pharmacies." CDC. 2019.
- 16. "U.S. overdose deaths topped 100,000 in one year, officials say." STAT News. 2021.
- 17. "Buprenorphine." SAMHSA. 2021.
- 18. "Methadone." SAMHSA. 2021.
- 19. "Naltrexone." SAMHSA. 2021.
- 20. "Contingency Management Interventions/Motivational Incentives (Alcohol, Stimulants, Opioids, Marijuana, Nicotine)" NIDA. 2020.
- 21. "Community Reinforcement Approach Plus Vouchers (Alcohol, Cocaine, Opioids)." NIDA. 2020.
- 22. "12-Step Facilitation Therapy (Alcohol, Stimulants, Opiates)." NIDA. 2020.
- 23. Joudrey et al. "Methadone Access for Opioid Use Disorder During the COVID-19 Pandemic Within the United States and Canada." JAMA Netw Open. 2021:4(7).
- 24. "Recovery Resources Adapt To COVID-19 Social Distancing, 'Stay-At-Home' Orders." WFYI. 2020.
- 25. "How the COVID Pandemic Made the Opioid Epidemic Worse, Even as Telehealth Helped." U.S. News and World Report. 2021.

