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The Rise of Fentanyl in the Midst of an Already Troubling Opioid Epidemic

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Each day we hear more and more stories of addiction, overdose, and deaths with heartbreaking stories of the individuals and the families who have been impacted by the opioid epidemic. In an attempt to address this crisis, more aggressive clinical intervention and opioid management programs are being adopted, and opioid use has steadily decreased for workers' compensation patients; however, according to an [NSC survey](#), 75% of employers are being directly impacted by employee opioid misuse. In another [study](#) of workers' compensation claimants who were initially treated with an opioid prescription, approximately 30% continued to fill opioid prescriptions beyond 90 days from injury, and unfortunately, overdose deaths attributed to synthetic opioids and heroin are increasing and are more than [making up for](#) reductions in prescription opioid deaths.

Due to its addictive properties, approaches that include opioid prescriptions can be followed by physical dependency, addiction, personal struggles, job losses, financial ruin, the need for substance use disorder treatment, and in some cases death by overdose or suicide.

Complex sequences of events have propelled the overdose crisis forward including the dramatic rise in the supply of opioids coupled with inadequate access to treatment programs to address their use. During the COVID-19 pandemic, these already concerning circumstances seemingly pushed even more people to turn to drugs at a time when treatment facilities and other services were shutting down. As it stands today, [drug overdose deaths](#) in the U.S. have topped 100,000 annually, and deaths from synthetic opioids (primarily fentanyl) and psychostimulants such as methamphetamines are on the rise. [Experts](#) sum all this up this way: "If it's easier to get high than to get treatment, people who are addicted will get high. The U.S. has effectively made it easy to get high and hard to get help."

As synthetic opioids began to appear more and more in the illegal drug supply, increasingly so did [fentanyl](#), which was diverted for abuse due to its powerful potency and euphoric properties, added to heroin to increase its

effects, or disguised as more intoxicating heroin. Sadly, often drug users believe they are purchasing heroin or other substances of abuse such as cocaine and actually don't know that they are purchasing fentanyl or products laced with fentanyl. To address misconceptions, it's critical that we understand the connection between fentanyl's appropriate uses and the devastating effects it can have when no guards are in place to prevent abuse.

Fentanyl is a powerful synthetic opioid and a comparatively recent addition to the list of all opioid medications used in clinical practice. First created by physician researcher Paul Janssen in 1960 and introduced in medicine as a general anesthetic in 1968, it continues to occupy a critical role in the treatment of a set of clearly-specified uses. It differs from other opioids (morphine, hydrocodone, oxycodone) in its high affinity for mu receptors, where it is 50 to 100 times more potent than morphine, contributing both to a benefit in effectiveness for treating severe pain and caution regarding its overdose potential.

In medical practice, fentanyl is often used in hospitals, intravenously, to enhance anesthesia, and for pain management in obstetric and surgical settings. In the COVID-19 era, fentanyl is used in endotracheal intubation procedures that support emergency ventilation in the sickest patients. In outpatient settings, a variety of dosage forms (transdermal patch, buccal films, lozenge, lollipop) are available by prescription for use in opioid-tolerant* patients for severe chronic pain, cancer pain, and breakthrough pain episodes, which are often short-lived but severe in intensity. Fentanyl also plays an important treatment role in comfort care for terminally ill patients and in battlefield settings for pain from acute trauma and wounds.

The synthetic analogues of fentanyl (acetylfentanyl, carfentanil, etc.), with even greater potency, are the subject of significant recent concern for illicit use and a growing contribution to drug overdoses. The illicit manufacture of fentanyl and fentanyl analogues incorporated into other substances of abuse are driving an overdose death trend in recent times for synthetic opioids other than methadone. These overdoses now eclipse those related to heroin or prescription opioids. The Centers for Disease Control and Prevention (CDC) [reported](#) that over half of all opioid overdose deaths involved synthetic opioids other than methadone, with illicit versions of fentanyl being the most common. The U.S. Drug Enforcement Administration (DEA) has issued a [temporary order](#) placing fentanyl-related substances (excluding fentanyl itself) in Schedule 1 of the Controlled Substances Act.

A number of concerned clinician authors have recently called for a [naming distinction](#) between fentanyl used in medical settings for critical applications and fentalogues used in illicit channels that are responsible for a growing number of overdose deaths. They note that "linchpins of the opioid epidemic, such as overdose, misuse, and abuse occur largely outside of their use as legitimate pain treatments," [citing](#) the rate of opioid use disorder among chronic pain patients as roughly 2 – 8%. While recognizing that a naming difference is largely symbolic, they also feel it's important to establish a distinction in the mind of the public, government officials, and the media between a medicine of clinical importance and the illicit forms of fentanyl that are driving a nation-wide epidemic of overdose deaths.

Despite continuous visibility in the [media](#) and public [education campaigns](#) conveying the dangers of fentanyl, access and exposure don't seem to be waning as fentanyl continues to enter the U.S. in large part illegally from [Mexico](#). With the U.S. claiming an overdose death every five minutes, the DEA has declared there are direct [links](#) between fentanyl-related overdose deaths and criminal drug networks. To help illustrate the severity of this problem, over a two-month period the Phoenix Arizona DEA Field Division launched a public safety surge targeting criminal drug networks utilizing social media apps to distribute deadly drugs. During this time over three million fentanyl pills and 45 kilograms of fentanyl powder were [seized](#). Obviously, more must be done to address the deadly cycle of drug misuse, abuse, and overdoses and the synthetic opioids made available through illegal drug supplies.

**The clinical literature generally defines patients who are considered opioid-tolerant as those who are taking at least 60 mg of oral morphine per day (or its equivalent dose with a different opioid) around-the-clock for one*

week or longer.



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