

From Pills to Fentanyl: Understanding the Opioid Crisis

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Three Inextricable, Overlapping, Concurrent Crises of Opioid Consequences



Chronic pain

Patients: >90M

Cost: \$635B

(n.b.CV disease \$309

billion)



Prescription drug use/misuse

Deaths: ~15,000/yr

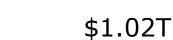
Cost: \$500B



Illicit opioid use

Death: >75,000/yr

Cost: unknown



Consequences of opioid use

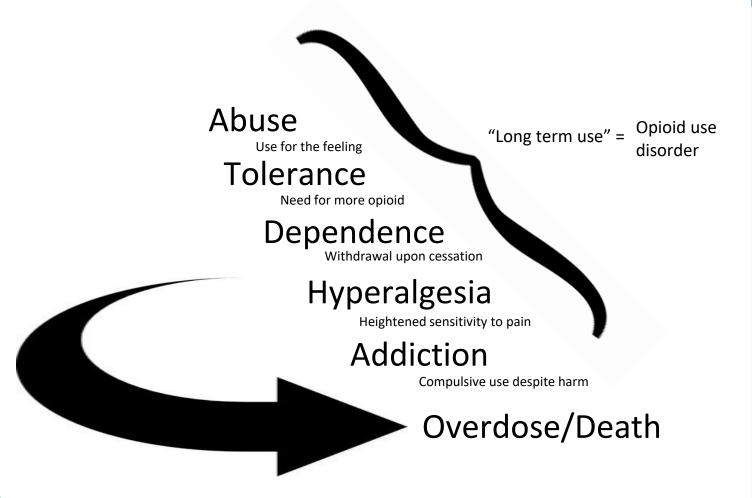


TABLE 36–3 Criteria for Opioid Use Disorder³

A problematic pattern of opioid use leading to clinically significant impairment or distress, as manifested by at least two of the following, occurring within a 12-month period:

- 1. Opioids are often taken in larger amounts or over a longer period than was intended.
- 2. There is a persistent desire or unsuccessful efforts to cut down or control opioid use.
- 3. A great deal of time is spent in activities necessary to obtain the opioid, use the opioid, or recover from its effects.
- 4. Craving, or a strong desire or urge to use opioids.
- 5. Recurrent opioid use resulting in a failure to fulfill major role obligations at work, school, or home.
- 6. Continued opioid use despite having persistent or recurrent social or interpersonal problems caused or exacerbated by the effects of opioids.
- 7. Important social, occupational, or recreational activities are given up or reduced because of opioid use.
- 8. Recurrent opioid use in situations in which it is physically hazardous.
- Continued opioid use despite knowledge of having a persistent or recurrent physical or psychological problem that is likely to have been caused or exacerbated by the substance.
- 10. Tolerance, as defined by either of the following:
 - A need for markedly increased amounts of opioids to achieve intoxication or desired effect
 - A markedly diminished effect with continued use of the same amount of an opioid

Note: This criterion is not considered to be met for those taking opioids solely under appropriate medical supervision.

- 11. Withdrawal, as manifested by either of the following:
 - The characteristic opioid withdrawal syndrome (refer to Criteria A and B of the criteria set for opioid withdrawal).
 - Opioids (or a closely related substance) are taken to relieve or avoid withdrawal symptoms.

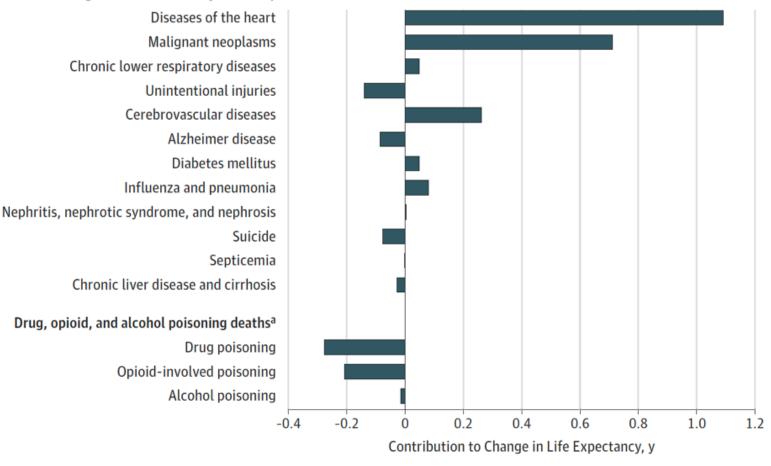


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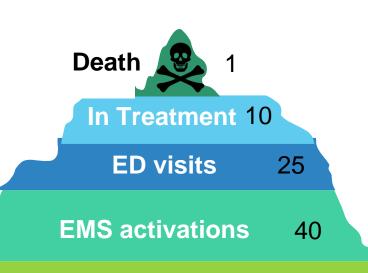
"You should just feel a tiny prick, and then a lifetime of morphine addiction."

Figure. Contributions of Selected Causes of Death to the Change in Life Expectancy in the United States, 2000-2015

12 Leading causes of death (ranked highest to lowest according to No. of deaths in year 2015)



^a In ranked cause-of-death classification, drug, opioid, and alcohol poisoning are not considered to be unique cause-of-death categories. Instead, poisoning deaths are classified as either accidental poisonings (which contribute to unintentional injuries), suicides, or homicides (ranked 16th in leading causes of death). Contributions from drug, opioid, and alcohol poisoning deaths overlap with both unintentional injury deaths and suicides and cannot be summed with these leading ranked causes of death.

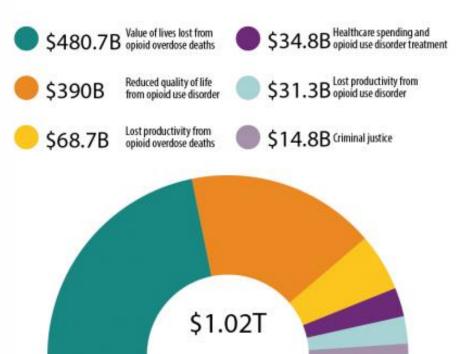


People who are opioid dependent 100

In 2017, opioid overdose deaths and opioid use disorder cost the U.S. \$1.02 trillion.

Here is what is included in the total cost.

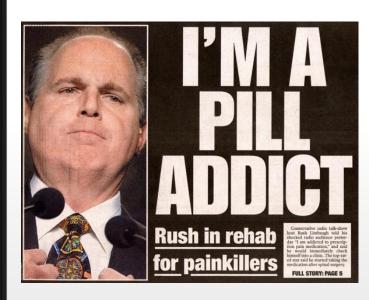
www.cdc.gov



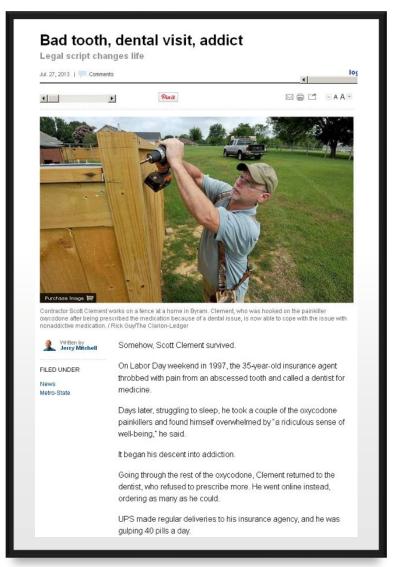
Nonmedical opioid users

Treatment admissions are for primary use of opioids from Treatment Exposure Data set
Emergency department (ED) visits are from DAWN,Drug Abuse Warning Network, https://dawninfo.samhsa.gov/default.asp
Abuse/dependence and nonmedical use in the past month are from the National Survey on Drug Use and Health

625







All opioids (e.g., oxycodone, fentanyl, hydromorphone, and morphine) bind the mu-1 and mu-2 opioid receptors

- Pain relief
- Euphoria
- Respiratory depression

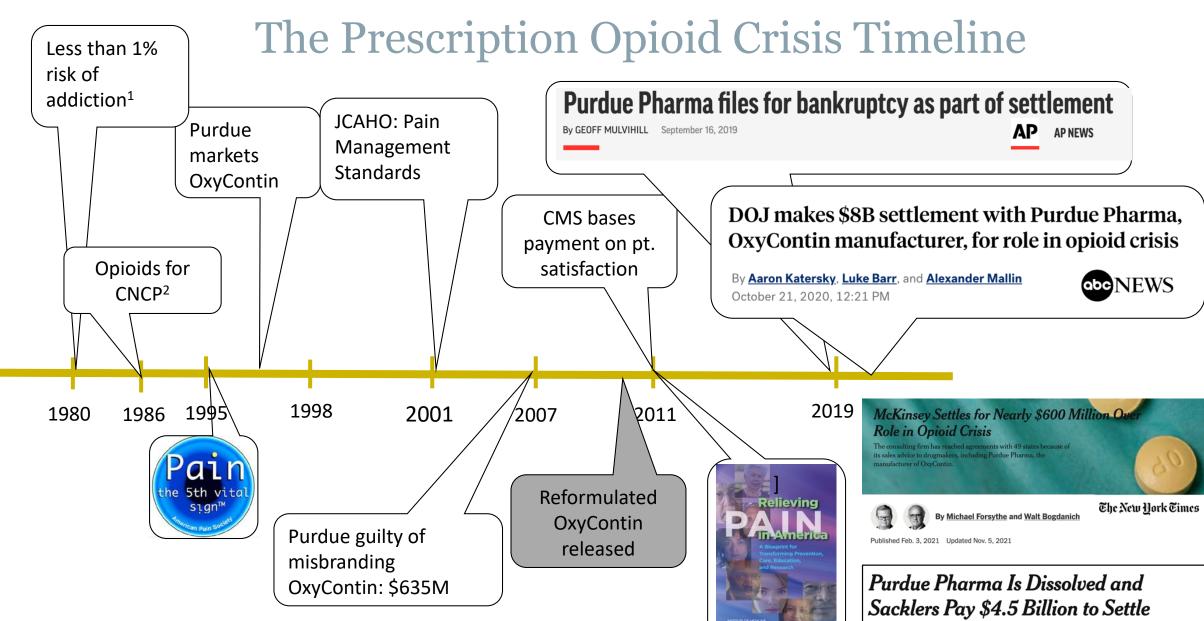
Opioid clinical effects are determined by:

- Lipid solubility
- Receptor specificity
- Binding affinity
- Dose (potency)
- Dose rate



Why isn't heroin legal?

Isn't oxy just legal heroin?



¹Porter J, Jick H. Addiction rare in patients treated with narcotics. N Engl J Med 1980;302:123.

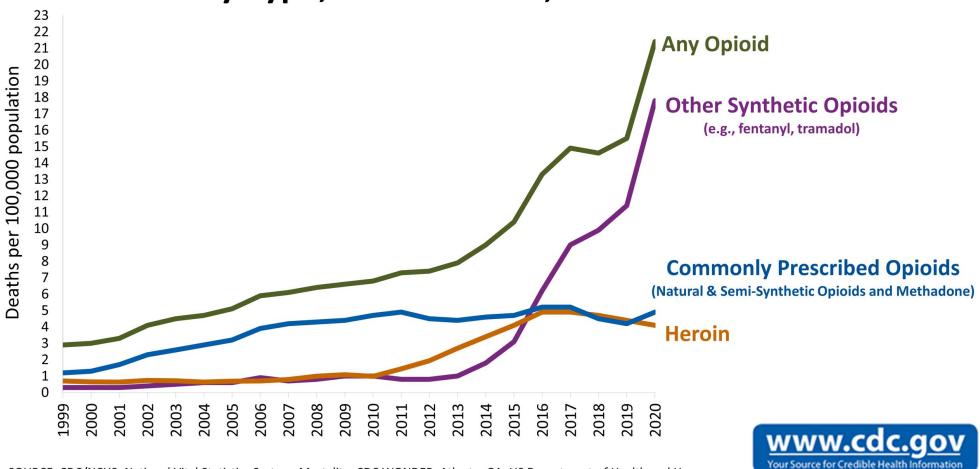
Opioid Claims By Jan Hoffman

The New Hork Times

Published Sept. 1, 2021 Updated Sept. 17, 2021

²Portenoy RK, Foley KM. Chronic use of opioid analgesics in non-malignant pain: report of 38 cases. Pain. 1986;25(2):171-86

Overdose Death Rates Involving Opioids, by Type, United States, 1999-2020

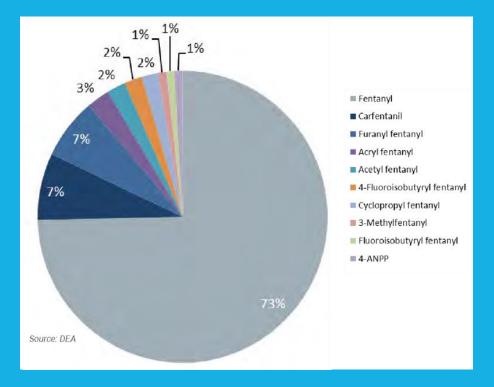


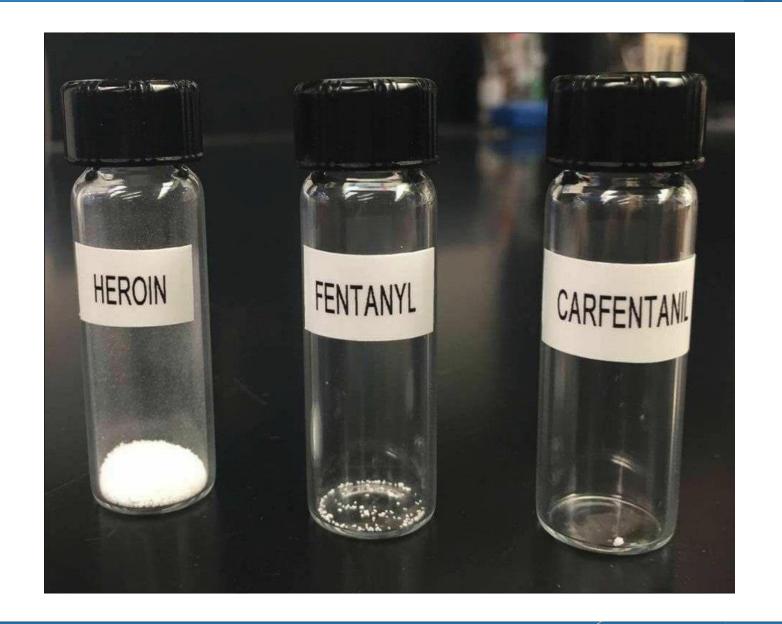
SOURCE: CDC/NCHS, National Vital Statistics System, Mortality. CDC WONDER, Atlanta, GA: US Department of Health and Human Services, CDC; 2020. https://wonder.cdc.gov/.

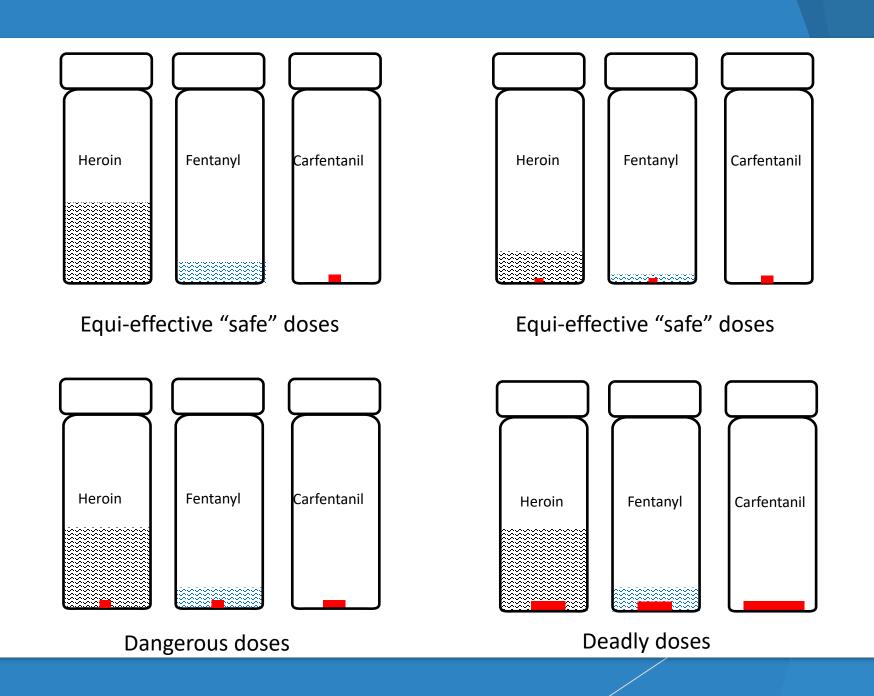
Fentanyl(s) (and Nitazenes)

- Mu opioid receptor full agonist(s)
- Highly lipophilic
 - Rapid onset when IV
 - Generally rapid offset with short term use
 - Accumulates in adipose tissue
 - ▶ Slow redistribution and long elimination time
- Apnea
 - At high doses
- Rigidity
 - ► At high infusion rates

Fentanyl/4-ANPP/Tramadol	Black	Spider & Spider (Image)	Camden Co PD	12/22/21	CAM
Fentanyl/4-ANPP/Fluorofentanyl/ Xylazine	Black	Caution (Blue Fold)	Camden Co PD	12/23/21	CAM
Fentanyl/4-ANPP/Phenethyl 4-ANPP	Red	Money Bag & Money Bag (Image)	Camden Co PD	12/31/21	CAM
Fent/Fluorofent/4-ANPP/ Phenethyl 4-ANPP	Black	Hard Knock Life (Blue Folds)	Camden Co PD	1/2/22	CAM
Fent/Fluorofent/4-ANPP/ Phenethyl 4-ANPP/	Multi	Sky Zone & Skull w/Helmet (Image)	Camden Co PD	1/2/22	CAM
Xylazine	DI I	DI 1 I	C 1 C PD	1 /4 /22	CAN
Fent/4-ANPP/Xylazine/ Phenethyl 4-ANPP	Black	Black Ice	Camden Co PD	1/4/22	CAM
Fentanyl/4-ANPP/Phenethyl 4-ANPP	Black	Hardknock Life & Illegible (Image) (Blue Folds)	NJSP AC Expwy	1/9/22	CAM
Heroin/Fentanyl/4-ANPP	Red	Power	NJSP Woodbine	12/9/21	CAP
Fentanyl/4-ANPP	Red	Mexico	NJ Transit PD	11/18/21	ESS
Fentanyl/Heroin/4-ANPP/Phenethyl 4-ANPP	Red	12:30	Irvington	12/2/21	ESS
Heroin/Fentanyl/4-ANPP/Phenethyl 4-ANPP	Multi	Pimp & Person w/Hat & Cane (Image)	Irvington	12/2/21	ESS
Fentanyl/4-ANPP/Phenethyl 4-ANPP	Blue	Goldfinger & Woman (Image)	NJSP Crime North	12/13/21	ESS
Fentanyl/4-ANPP/Phenethyl 4-ANPP	Blue	Tango & Cash & \$ (Image)	NJSP Crime North	12/13/21	ESS
Fentanyl/4-ANPP	Blue	The North Face	NJSP Crime North	12/13/21	ESS
Heroin/Fentanyl/Fluorofentanyl/4-ANPP/	Red	*Yeezy & Sneaker (Image)	Bloomfield	12/25/21	ESS
Clonidine (sedative)/Tramadol/Xylazine	Marie	Class 7 0 Classiff and Assisting Hat (Lass)	II-ll	0 /21 /21	шь
Heroin/Parafluorofentanyl/Fentanyl	Multi	Sky Zone & Skull w/Aviation Hat (Image)	Hoboken	9/21/21	HUD
Xylazine/Heroin/Parafluorofentanyl/Fentanyl	Blue	Dirty Diana & Face (Image)	Jersey City	9/21/21	HUD
Xylazine/4-ANPP/Heroin/Fentanyl	Blue	Dirty Diana & Face (Image)	Jersey City	10/4/21	HUD
4-ANPP/Heroin/Parafluorofentanyl/ Fentanyl	Orange	Givenchy & Orange (Image)	Jersey City	10/4/21	HUD
Fentanyl	Red	Section 8 & Building (Image)	Jersey City	10/22/21	HUD
Fentanyl	Black	Octopus & Octopus (Image	Jersey City	10/28/21	HUD
Fentanyl	Red	King Kong & Gorilla (Image)	Jersey City	10/30/21	HUD
Xylazine/Heroin/Fentanyl	Blue	Best Buy & BB Logo (Image)	Jersey City	11/7/21	HUD
Heroin/Fentanyl	Blue	Lemon Pepper	Jersey City	11/10/21	HUD
Heroin/Fentanyl	Red	ALPO	Jersey City	11/19/21	HUD
Xylazine/Heroin/Parafluorofentanyl	Blue	Power & Fist (Image)	Jersey City	11/30/21	HUD
Fentanyl/4-ANPP/Xylazine				10/12/21	HUN
Fentanyl/4-ANPP/Xylazine	Purple	**Keys to the Streets	NJSP Perryville	10/12/21	HUIN
Heroin/Fentanyl/4-ANPP/Phenethyl 4-ANPP/	Purple Red	**Keys to the Streets Jawbone	NJSP Perryville NJSP Crime Cent	10/12/21 10/22/21	MER
Tramadol/Xylazne	•	-	•		









What did we do about it?



- Limit opioid initiation
 - Pain management guidelines
 - Multi-modal analgesia
- Safe opioid use
 - Default prescribing EHR
 - Regulatory limits
 - Order sets
 - Nudge prescribers
 - Patient education
- Prescription Monitoring Program





University Hospital Adult Emergency Medicine Treatment of Acute Pain Guideline

- ernative therapies should be considered if there are contraindications to first line recommendations
- Consider next line therapies in a stepwise manner if pain persists 30 minutes after an IV dose OR 60 minutes after a PO dose
- Other than in the treatment of severe acute pain, the oral route is the preferred route of administration of most analgesic drugs

First Line	Second Line	Third Line	Adjunctive Therapy	Discharge
Undifferentiated abdominal pain	Undifferentiated abdominal pain	Opioid rescue*	Anti-emetics	Undifferentiated abdominal pain
Acetaminophen 975 mg PO	Ketamine 0.3 mg/kg IV over 15		Ondansetron 4 mg IV	Acetaminophen 975 mg PO q6H PRN
AND/OR	minutes		OR	AND/OR
Ibuprofen 400 - 600 mg PO			Ondansetron ODT 4 mg PO	Ibuprofen 400 mg PO q6H PRN
(If patient cannot tolerate PO,	Gastroparesis		OR	
ketorolac 15 mg IV)	Haloperidol 5 mg IV		Metoclopramide 10 mg IV	Spasmodic pain
	OR			Dicyclomine 20 mg PO q6H PRN
Spasmodic pain	Haloperidol 5 mg IM		Antacids	
Dicyclomine 20 mg PO			Mag hydroxide/aluminum	Gastroparesis
(If patient cannot tolerate PO,			hydroxide/simethicone 1200	Metoclopramide 10 mg PO q6H PRN
dicyclomine 10 mg IV)			mg/1200 mg/120 mg PO	
			AND/OR	
Gastroparesis			Famotidine 20 mg IV	
Metoclopramide 10 mg IV				
Clinical Pearls:				

- Consider underlying etiology of abdominal pain before selecting treatment option (e.g. anticholinergics and opioids counterintuitive in gastroparesis
- Ketamine: avoid use in patients with severe hypertension or history of psychosis
- NSAIDs: avoid use in third trimester of pregnancy, peptic ulcer disease, history of GI bleed, or active major bleedin
- Provide patient education regarding type of pain, medication choices, and what to expect
- Consider distractions such as music, talking to patient

Dental Pain				
First Line	Second Line	Third Line	Adjunctive Therapy	Discharge
Acetaminophen 975 mg PO	Lidocaine 2% viscous solution -	Lidocaine 1% dental block	Apply ice pack to painful area	Acetaminophen 975 mg PO q6H PRN
AND/OR	swish and spit			AND/OR
Ibuprofen 400 – 600 mg PO				Ibuprofen 400 – 600 mg PO q6H PRN
(If patient cannot tolerate PO,				AND/OR
ketorolac 15 mg IV)				Lidocaine 2% viscous solution -
				swish and spit q3 hours PRN
Officiant December				

linical Pearls:

- Provide patient education regarding type of pain, medication choices, and what to expect
- Analgesia is a temporizing measure for more definitive treatment
- NSAIDs: avoid use in third trimester of pregnancy, peptic ulcer disease, history of GI bleed, or active major bleeding

Guidelines are intended to be flexible. They serve as reference points or recommendations, not rigid oritoria. Guidelines should be followed in most cases, but there is an understanding that, depending on the patient, setting, circumstances or factors

NJPMP Home	NJ Prescription Monitoria	ng Program
Alerts	For Too Many New Jerseyans, Addiction I	-
Laws and Regulations		
Register for the NJPMP	The New Jersey Prescription Monitoring Program (NJPMP) is an important component of the New Jersey Division of Consumer Affairs' initiative to halt the abuse and diversion	Email NJ Prescription Monitoring Program
Pharmacy Reporting	of prescription drugs.	
Requirements	Established pursuant to N.J.S.A. 45:1-45 et. seq., the	Call
Frequently Asked Questions	NJPMP is a statewide database that collects prescription data on Controlled Dangerous Substances (CDS) and Human	(973) 273-8010
Suspicious Activity Report	Growth Hormone (HGH) dispensed in outpatient settings in	Inquiries about the NJPMF
and the same	New Jersey, and by out-of-State pharmacies dispensing into New Jersey.	may be forwarded to
Useful Links Law Enforcement	Pharmacies are required to report information to the NJPMP on a daily basis to the PMP Clearinghouse using	Jeffrey D. Laszczyk, Jr., PharmD NJPMP Administrator P.O. Box 47014
Treatment	the ASAP 4.2 format. Prescriptions must be reported to the database no more than one (1) business day after the	Newark, New Jersey 07101 Directions
Media	date the prescription was dispensed.	

What are doing about it?



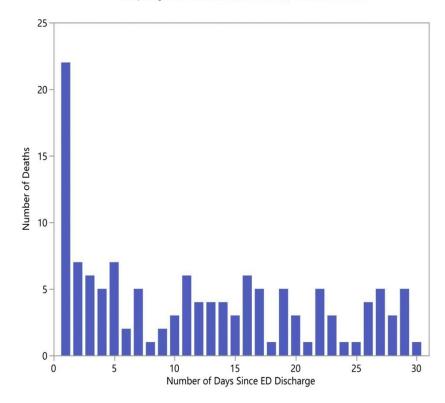
Naloxone availability without engagement into addiction treatment doesn't save lives... it only postpones death.

CARY NALOXONE

One-Year Mortality of Patients After Emergency Department Treatment for Nonfatal Opioid Overdose

Scott G. Weiner, MD, MPH*; Olesya Baker, PhD; Dana Bernson, MPH; Jeremiah D. Schuur, MD, MHS

*Corresponding Author. E-mail: sweiner@bwh.harvard.edu, Twitter: @scottweinermd.

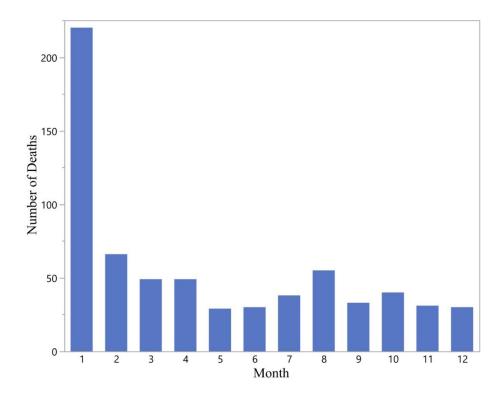


Death (rate) of 17,241 ED discharged OOD patients:

- 635 (5.5%) within 1 year
- 130 (1.1%) within 1 month
- 29 (0.36%) within 2 days
- ²/₃ died from opioid overdose

One Year Mortality of Patients Treated with Naloxone for Opioid Overdose by Emergency Medical Services

Scott G. Weiner, Olesya Baker, Dana Bernson, Jeremiah D. Schuur Subst Abus. 2022; 43(1): 99–103.



Death (rate) of 9734 EMS discharged OOD patients:

- 1465 (15.2%) 1 year
- 807 (8.3%) with 3 days
- ²/₃ died from opioid overdose

What are doing about it?







- Harm reduction
 - Naloxone distribution/prescribing
 - Fentanyl testing strips
 - Safe consumption sites

- Addiction management
 - Screening
 - Reducing barriers
 - ► Telehealth
 - Recovery coaches
 - Medication for opioid use disorder
 - Buprenorphine
 - Methadone



CRIME & DRUGS

There's no good antidote for super heroin laced with elephant tranquilizer

f 🥑 p

By Rachel Browne
October 3, 2016 | 3:10 pm

'I'm not going to let you die': Deputy collapses after handling suspected fentanyl



Handout-San Diego County Sheriff's Department.

The Sheriff's Department releases video showing the July 3 overdose. Sheriff Bill Gore calls fentanyl one of the 'greatest threats' facing region

For the record:

2:16 p.m. Aug. 10, 2021: This story misstates what was known about the medical condition of the deputy shown in the Sheriff Department's video. The conclusion that his collapse was caused by contact with fentanyl was supplied by the Sheriff's Department rather than by qualified medical experts, who have since cautioned that the risks from incidental contact are often overstated. A follow-up story can be found here.

The San Diego Union-Tribune

Officers find enough fentanyl to kill 4 million people in drug bust, officials say



The Louisville Metro Police Department reports officers found nearly 30,000 suspected fentanyl pills and at least 8.5 kilograms of fentanyl in a

storage unit. (Louisville Metro Police Department)

By Shellie Sylvestri and Jordan Gartner

v.nbc15.com/weather 18, 2022 at 4:02 PM EDT

WMTV

NATIONA

Is 'rainbow fentanyl' a threat to your kids this Halloween? Experts say no

October 11, 2022 · 5:00 AM ET



In August 2022, the Drug Enforcement Administration and law enforcement partners seized brightly colored rainbow fentanyl pills in 18 states.

Drug Enforcement Administration

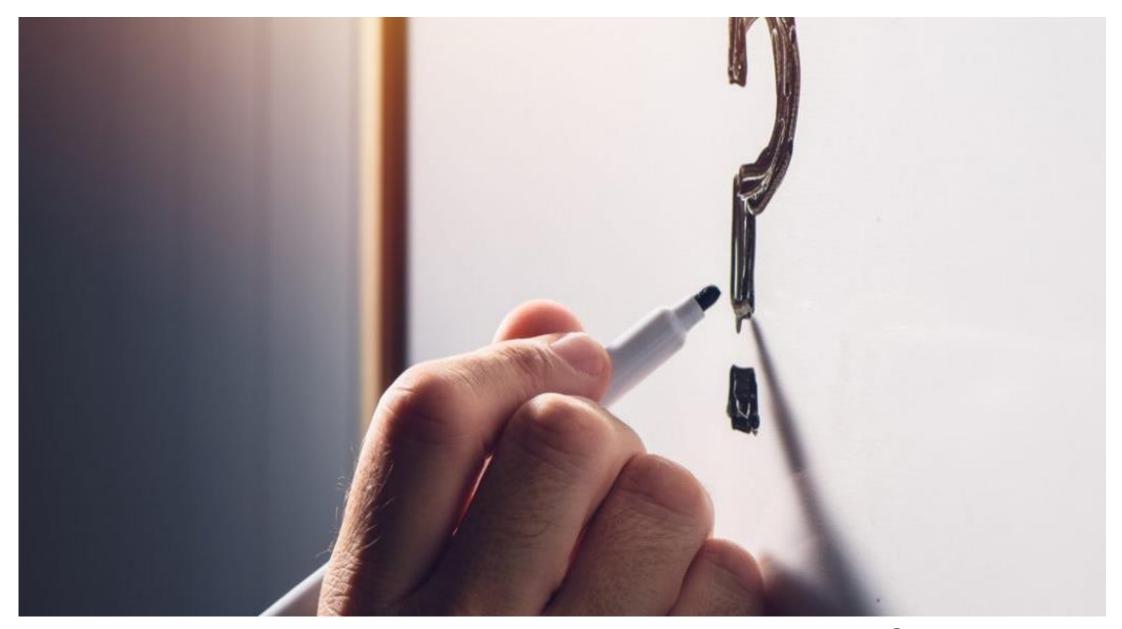
Drug Enforcement Administration





Summary

- The opioid crisis remains remains iatrogenic in part but mostly now related to fentanyl(s)
- An opioid is an opioid is an opioid (unless it is buprenorphine)
 - ► There are subtle but important pharmacological differences
- Naloxone can reverse opioid overdose (and sometimes prevent death)
 - Need to engage in treatment and recovery
- 'Be careful about reading health books. You may die of a misprint.' Mark Twain



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