

Objectives

- Discuss common classes of drugs of abuse, their clinical effects, and management.
- Recognize over-the-counter (OTC)
 medications commonly abused and their
 clinical effects in the setting of overdose.
- Discuss contaminants and adulterants, such as xylazine, identified in drugs of abuse.
- Evaluate the utility of various drug screens in testing for commonly abused substances including synthetics.

Opioids

- Include both prescription and illicit formulations
 - Multiple routes of exposure including ingestion, inhalation, parenteral
- Classic triad of symptoms includes CNS depression, respiratory depression, and pinpoint pupils
 - Other common effects include hypotension and bradycardia
 - These symptoms can also occur with clonidine, tizanidine, and imidazoline decongestants



Morphine and codeine are naturally occurring alkaloids from the opium poppy seed

- T_{1/2} varies widely from relatively short (morphine) to long (methadone)
 - Antidote is naloxone
 - Start low, particularly in patients who are habituated, and be ready to titrate upwards quickly
 - Duration of naloxone is 30-90 minutes
 - If patient experiences recurrent symptoms, consider continuous infusion
 - Synthetic opioids may require higher dosing

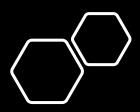
Common Opioids

Drug	Type of Activity	Usual Adult Dose*(mg)	Elimination Half-Life(h)	Duration of Analgesia (h)
butorphanol (Stadol®)	Mixed	2	5-6	3-4
codeine	Agonist	60	2-4	4-6
fentanyl (Duragesic®, Sublimaze®)	Agonist	0.2	1-5	0.5-2
heroin	Agonist	4	(b)	3-4
hydrocodone (Lortab®, Vicodin®)	Agonist	5	3-4	4-8
hydromorphone (Dilaudid®)	Agonist	1.5	1-4	4-5
meperidine (Demerol®)	Agonist	100	2-5	2-4
methadone (Dolophine*, Methadose*)	Agonist	10	20-30	4-8 ^d
morphine	Agonist	10	2-4	3-6
nalbuphine (Nubain®)	Mixed	10	5	3-6
oxycodone (Percocet®, OxyContin®)	Agonist	4.5	2-5	4-6
pentazocine (Talwin®)	Mixed	50	2-3	2-3
propoxyphene (Darvon®, Darvocet®)	Agonist	100	6-12	4-6
tramadol (Ultram®)	Agonist€	50-100	6-7.5	4-6

Usual dose: Dose equivalent to 10 mg of morphine
 Rapidly hydrolyzed to morphine
 Also inhibits serotonin reuptake
 Sedation and coma may last 2-3 days

TABLE 4. Classification of Opioids

Derivation	Opioid			
From opium	Opium, morphine, codeine, thebaine			
Semisynthetic	Heroin, hydrocodone, hydromorphone, oxycodone			
Synthetic	Methadone, propoxyphene, meperidine, fentanyl			



- We all know about the opioid epidemic...
- What happens with increased demand for prescription opioids and decreased supply?



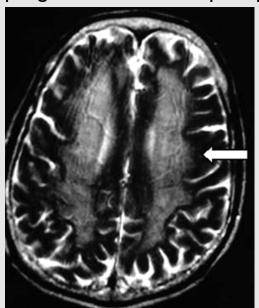


Heroin use rises dramatically

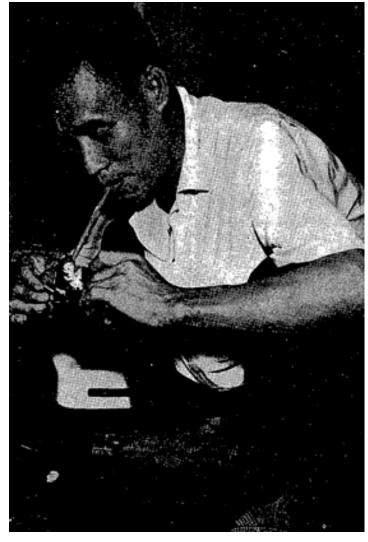


"Chasing the Dragon"

- Users inhale the white pyrolysate [vapor] that is generated by heating heroin on aluminum foil
- Pharmacokinetics similar to IV heroin
- Spongeioform leukoencephalopathy







Special considerations

- **Body stuffer** an individual who hastily ingests illegal drug packets to evade law enforcement officials
- Ingestion often involves more than one type/class of drugs. Leakage from these poorly wrapped packets can produce mild to severe toxicity
- Body stuffers usually deny the ingestion, and are often not discovered until symptoms develop

Special considerations

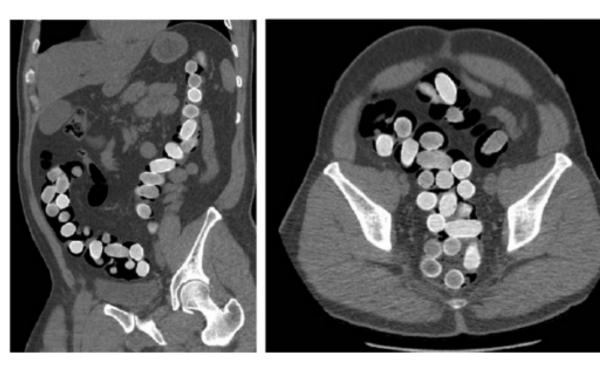


- Body packer —individual who ingests or inserts into body orifices, then transports wrapped packages of an illegal substance in an attempt to evade detection by law enforcement. Also known as a "mule." Usually seen in those transporting cocaine or heroin.
- Generally asymptomatic; however, severe acute drug toxicity can develop rapidly from leakage of contents or rupture of the drug packets
- Bowel obstruction is another potential complication, especially in opioid body packers





Multiple drug-filled packets distributed through colon.



CT showing multiple packets in the colon again, with difference in density suggesting difference in purity of content. Note lack of oral or intravenous contrast.



Decontamination

• Body packers and stuffers – whole bowel irrigation (WBI)

• PEG solution (Golytely®) - continue until rectal effluent is clear and no packets are detected on a contrast study of the bowel

• Enemas or endoscopy should generally be avoided due to the risk of package rupture

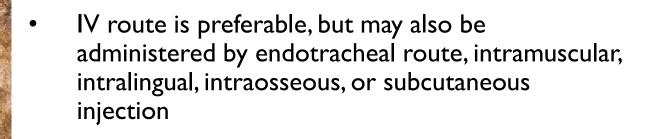
Antidote

- naloxone (Narcan[®])
- naltrexone (Naltrex[®], Vivitrol[®]) an opioid antagonist used for the treatment of opioid dependence. NOT for use in the overdose setting

Naloxone

- Initial dose is 0.4 2 mg IV push, may be repeated in 2 mg increments
- Large (>10 mg) doses may be required for treatment of buprenorphine or fentanyl (or fentanyl derivative) overdose
- In the habituated patient, smaller doses (0.1 or 0.2 mg increments) may be used intravenously and titrated to desired effect to minimize the precipitation of a withdrawal syndrome

Naloxone (Narcan®)



- If a reversal response occurs, patients should be observed for 4 hours after the final dose for any recurrent sedation/respiratory depression
 - Recommend 8-hour observation time in the setting of long-acting opioids such as fentanyl, methadone and buprenorphine
 - Naloxone may not completely reverse the respiratory depression seen with buprenorphine



Intranasal naloxone has seen a tremendous increase in use in emergency situations. Easy for minimally-trained personnel to administer.

Continuous-infusion (CI) naloxone

- May be used in patients with persistent or recurrent effects
- An initial dose of IV naloxone is given to achieve reversal of opioid effects, then a continuous infusion is administered to maintain this state of antagonism
- DOSE Use 2/3 of the initial effective naloxone bolus on an hourly basis

- Add ten times this dose to each liter of D5W and infuse at a rate of 100 mL/hour
- Titrate dose to response
- Mixtures should be used within 24 hours
- Observe for recurrence of toxic symptoms for at least 2 hours after infusion is discontinued

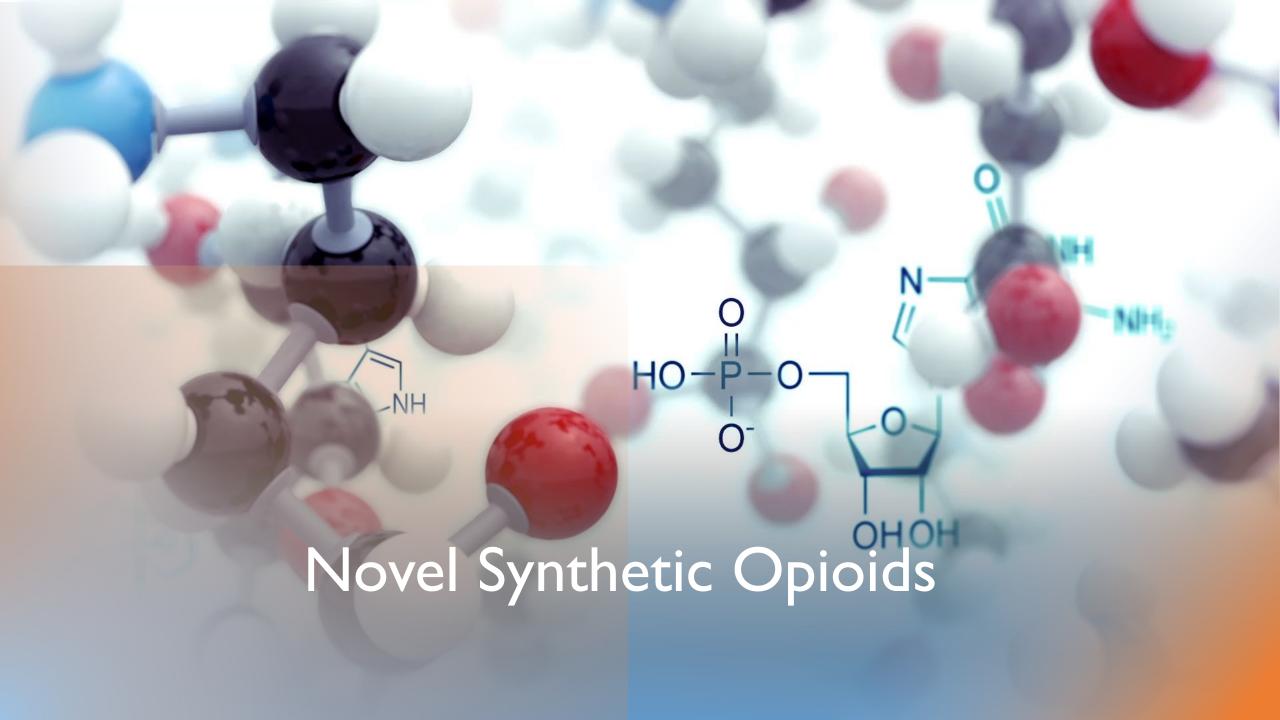
Continuous-infusion (CI) naloxone

Example

- I.5 mg of naloxone required to achieve satisfactory response
- Starting dose for CI naloxone will be I mg (2/3 of 1.5 mg) per hour
- Mix 10 mg of naloxone in 1000 mL of D5W
- Infuse at 100 mL/hr; titrate to response

Adverse Effects of Naloxone

- Adverse effects reported with naloxone include hypertension, laryngospasm, and pulmonary edema
- Precipitation of withdrawal may occur in the habituated patient
 - Although the symptoms are subjectively severe (e.g., n/v, diarrhea, diaphoresis, agitation), they are generally not life-threatening



Classes of Novel Synthetic Opioids (NSOs)

Fentanyl and it's derivatives

- Ones you may have heard of "historically":
 - α-methylfentanyl/3-methylfentanyl (China white)
 - Acetylfentanyl
 - Carfentanil (Wildnil)
 - Fluorofentanyls (i.e. para-fluorofentanyl) (briefly in 1980s then again in 2016)
 - Still seeing many cases of this (330 cases in Q1 of 2024)
- New analogues:
 - Ortho-methylfentanyl (first detected in Dec 2023)

Other non-fentanyl synthetic opioids

- Ones you may have heard of "historically":
 - AH-7921 (Doxylam) (a benzoamide)
 - U-47700 (closely related to AH-7921)
 - MT-45 (a piperazine)
- Nitazene analogues
 - N-pyrrolidino etonitazene (first detected in May 2021)
 - Protonitazene (first detected in May 2021)
 - Metonitazene (first detected in July 2020)

Classes of Novel Synthetic Opioids (NSOs)

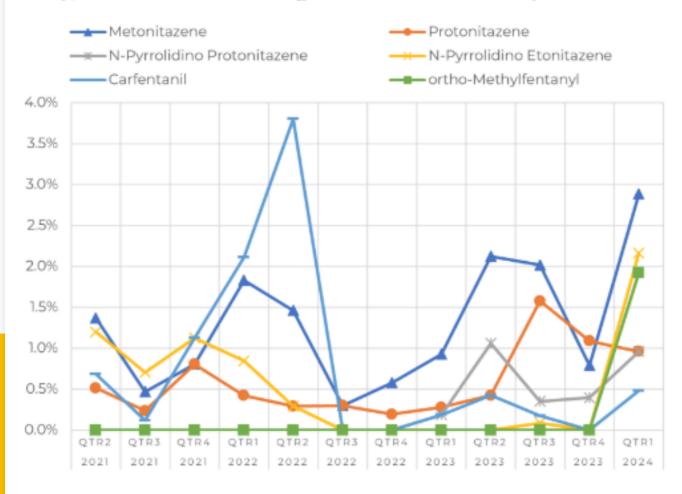
Current 5 Most Commonly Detected NSOs:

(according to April 2024 Quarterly Report from The Center for Forsenic Science Research and Education (CFSRE)):

- I. Ortho-methylfentanyl
- 2. N-pyrrolidino etonitazene
- 3. Metonitazene
- 4. Protonitazene
- 5. Carfentanil

SELECT POSITIVITY: Q2 2021 TO Q1 2024

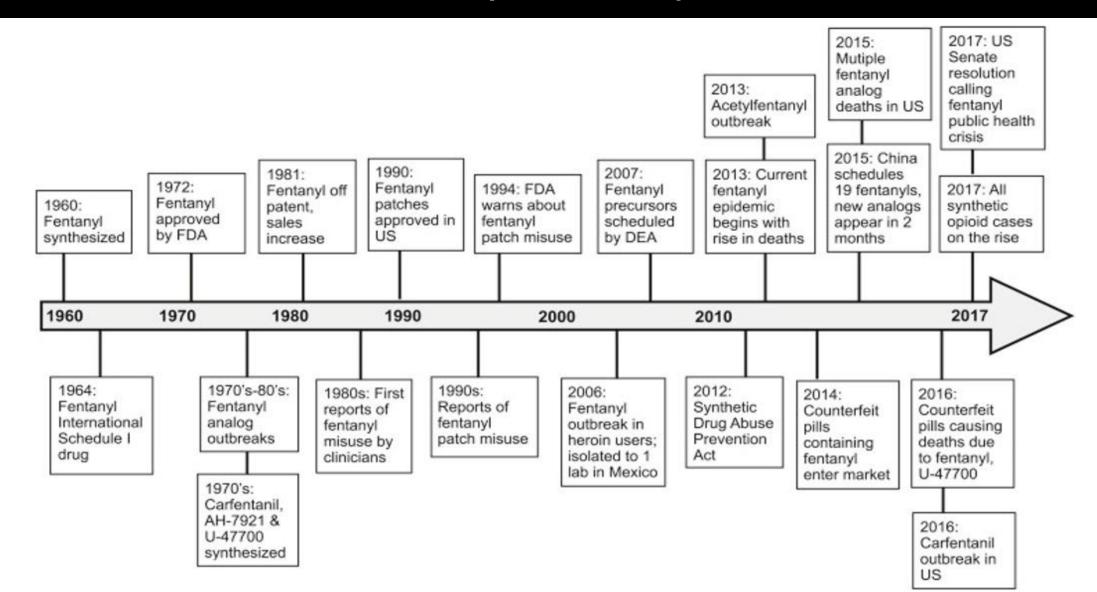
Positivity plots are derived from a select toxicology data source that has been consistently monitored since 2018.



Where do NSOs Come From?

- Online (including the dark web) labs in China and overseas
 - Shipped to US
 - For example fluorofentanyls being manufactured in Mexico by cartels as a result of the scheduling of Fentanyl precursors
- Head shops
- Drug dealers
- Often sold as heroin or mixed with other drug supplies to unsuspecting users
- May be ingested, insufflated, smoked, or injected

Timeline of Synthetic Opioid Events

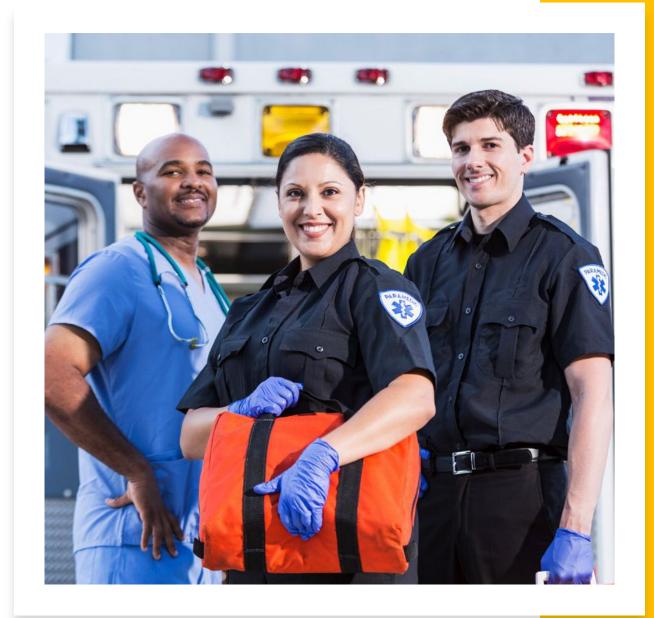


Demographics

- The most US deaths from NSOs have occurred in the Northeast and South
- Primarily affects the same demographic group that uses heroin: non-Hispanic white males aged 25-44 years, lower SES
- Other potential risks: chronic pain, obesity, uneducated, mental health issues

What About First Responders?

- Risk of exposure to first responders is very low
- For routine handling of drug, nitrile gloves provide sufficient dermal protection
- In rare cases where there are aerosols, an N95 respirator provides sufficient protection
- No persistent effects from NSO exposure are expected in the absence of prolonged hypoxia

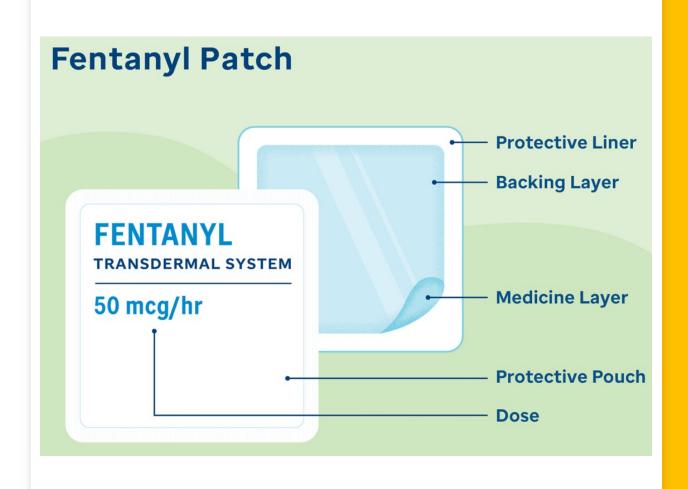


Fentanyl

- Available in many formulations
 - Sublingual Spray (Subsys) 100 mcg, 200mcg, 400 mcg, 600 mcg, 800 mcg, 1200 mcg, 1600 mcg/spray
 - Patches (Duragesic) 12.5 mcg/hr, 25 mcg/hr, 37.5 mcg/hr, 50 mcg/hr, 62.5 mcg/hr, 75 mcg/hr, 87.5 mcg/hr, 100 mcg/hr,
 - Lozenge on a Handle (Actiq) 200 mcg, 400 mcg, 600 mcg, 800 mcg, 1200 mcg, 1600 mcg
 - Buccal Tablets (Fentora, Abstral) 100 mcg, 200 mcg, 400 mcg, 600 mcg, 800 mcg
- Highly lipid soluble and quickly crosses BBB
- Short duration of action

Abuse of Fentanyl Patches

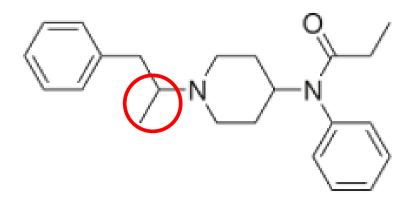
- Inhalation:
 - Place patch in a glass container, heat, and smoke
 - Scratch and smoke the patches
- Ingestion
 - Remove gel contents and ingest
 - Cut frozen patches into pieces and chew or suck on them ("Chiclets")
- Injection:
 - Remove gel contents and inject
 - Simmer patches in water and inject the liquid extract IV



Residual Amount of Fentanyl in Patch

Patch Strength	Dose in Patch (mg)	Dose in Patch (mcg)	mcg used in 72 hours	Residual amount of drug in patch (mcg)
25 mcg/hour	2.75	2750	1800	950
50 mcg/hour	5.5	5500	3600	1900
75 mcg/hour	8.25	8250	5400	2850
100 mcg/hour	П	11000	7200	3800

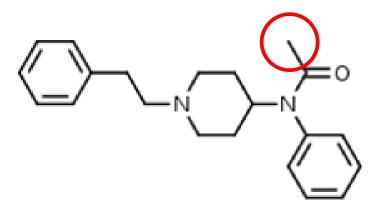
α-Methylfentanyl (China White)



- First designer fentanyl (California, 1970s)
- Used as a heroin substitute or laced in heroin

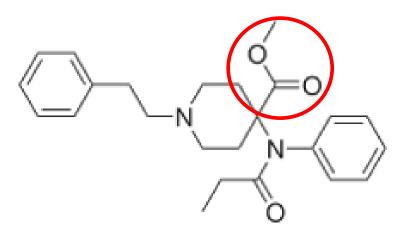


Acetylfentanyl



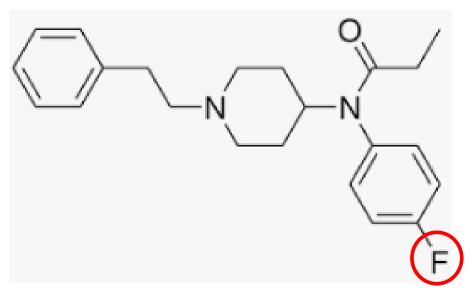
- Created by removing one methyl group from fentanyl to form desmethylfentanyl
- Somewhat weaker than fentanyl
- Public health alert first issued in early 2013

Carfentanil (Wildnil®)



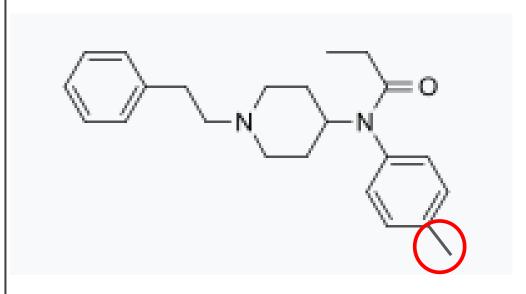
- Most potent commercially used opioid
- Marketed as a large animal sedative (elephants and rhinos) – not for human use!
 - First introduced as vet medicine in 1986
- May appear in heroin
- Dramatic increase in shipments from Chinese chemical companies into the US in 2016
- Current "outbreak" is different from historical outbreaks where found drug material that was solely carfentanil and multiple deaths in a centralized area
 - Now seeing it across the country and almost always mixed with fentanyl (not high purity)

Para-fluorofentanyl



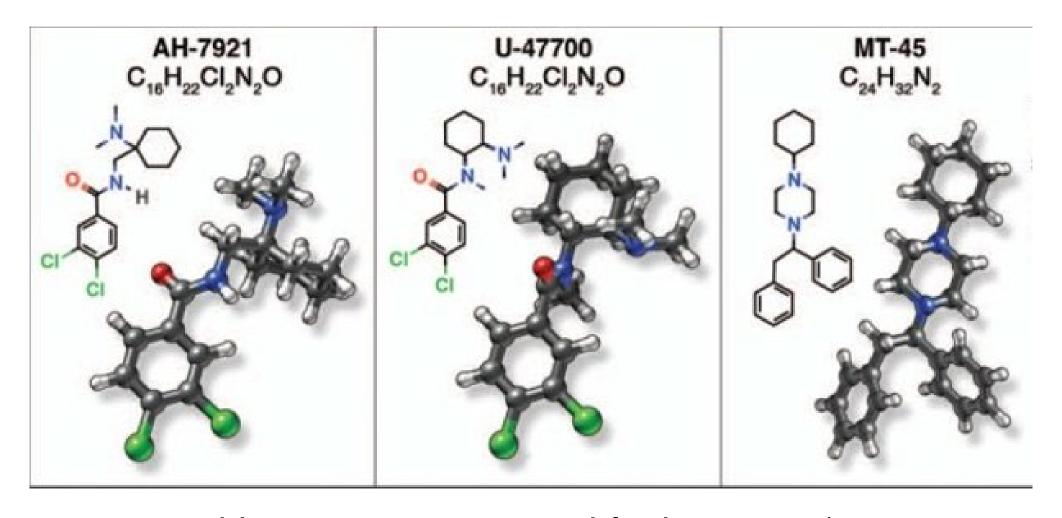
- Three times less potent than fentanyl
 - However, concentrations found in drug deaths suggest that deaths can occur with concentrations similar to those involving fentanyl
- Almost always found together with fentanyl (93% of the time)
- 1980s → 2016 → dropped off in 2018 after core-structure scheduling → reemerged in Q4 of 2020 (per CFSRE)

Ortho-methylfentanyl



- Relatively new found in four blood toxicology specimens in British Columbia, Canada
- Not yet identified in any drug materials (as of Dec 2023)

Other Novel Synthetic Opioids



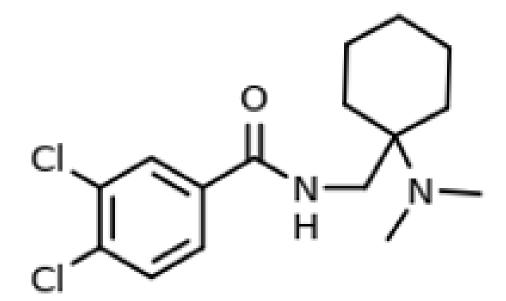
Note: none are approved for human use!

AH-7921 (Doxylam)

BEIMONT PARK
SPESSORS FALSE SELECTION
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Two horses that raced at Belmont Park have tested positive for the synthetic opioid known as AH-7921

- Unique effects in overdose: tachycardia, HTN, seizures
- Not the same as doxylamine!
- Scheduled by the US in 2016
- Less commonly seen



U-47700 (U4 or pink heroin or pinky)

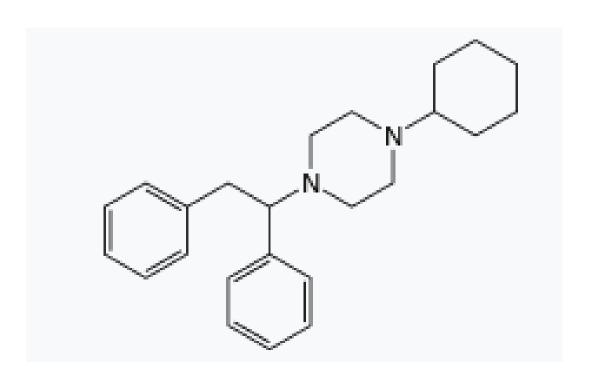
- Structural isomer of AH-7921
- According to user reports, has a longer-lasting effect than AH-7921
- Induces significant euphoria which is short-lived and causes an urge to keep redosing



$$CI \underbrace{\hspace{1cm} 0 \hspace{1cm} N^{N}}_{N} \underbrace{\hspace{1cm} N^{N}}_{N}$$

MT-45 (IC-6)

- Unusual side effects: hearing loss, hair depigmentation, alopecia, cataracts, Mees lines on nails
- About 80% of potency of morphine



Core Structure Scheduling – Fentanyl analogues

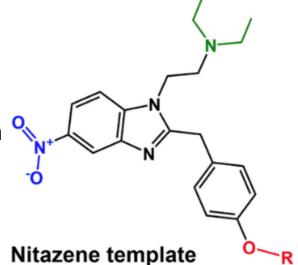


The Scheduling Cycle – Nitazene analogues

- DEA moved to control fentanyl analogues (core structure scheduling) in spring of 2018
 - Had dramatic effect in eradicating fentanyl analogues in the supply in the United States
 - Exceptions: carfentanil and now orthomethylfentanyl
 - As a result opened the door to the proliferation of nitazene analogues
 - Starting with etonitazene
 - Early on would see a nitazene analogue emerge – DEA would schedule it – would see that nitazene analogue go away
 - More recently seeing some scheduled nitazene analogues persisting and proliferating in supply
 - Metonitazene cyclical patterns of increase/decrease in the supply (not clear why)

Nitazene Analogues

- First developed in the 1950s-1960s as opioid analgesics but never approved to market
 - Largely investigational/chemistry research
- Structurally unrelated to fentanyl
- Hundreds-thousands fold more potent than morphine and 10-40x more potent than fentanyl
 - Different from fentanyl analogues of which majority are less potent than fentanyl
- Often mixed with fentanyl or other agents



R = methyl. metonitazene

R = ethyl, etonitazene

R = isopropyl, isotonitazene

R = propyl, protonitazene

R = butyl, butonitazene

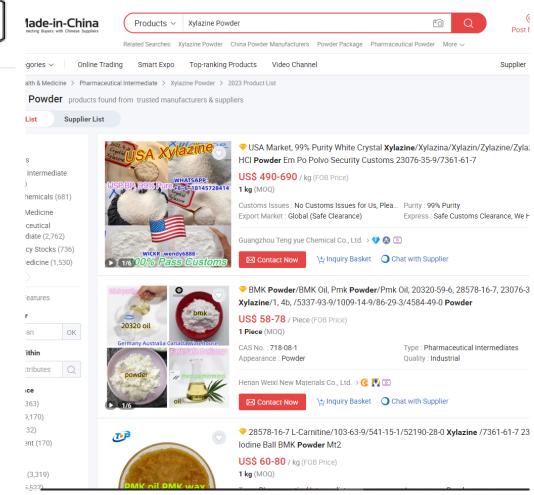
Table of Comparative Opioid Potencies

Opioid	Potency Compared to Morphine	Potency Compared to Heroin
Morphine	1	~1/3
Heroin	2-5x	1
AH-7921	1.7x	
MT-45	3.5 x	
Butyrylfentanyl	7x	
U-47700	7.5x	
Acetylfentanyl	16-80x	5-15x
Fentanyl	50-100x	
3-methylfentanyl	1,000-7,000x	
Carfentanil	10,000x	4,000x

Xylazine ("Tranq")



- Commonly mixed with fentanyl
 - Fentanyl results in more euphoria but is shorter-acting than heroin
- Powerful sedative FDA approved for veterinary use
- Alpha-2 agonist (like clonidine and guanfacine)
- DEA has seized xylazine/fentanyl mixtures in 48/50 states
- Since 2021: >90% of "dope" samples tested from Philadelphia have contained xylazine and fentanyl with xylazine purity ranging from 5% to > 70%
- New Jersey: about 60% of samples contained xylazine
- In 2022, approximately 23% of fentanyl powder and 7% of fentanyl pills seized by the DEA contained xylazine.
- From June 2022 to June 2023, the Oklahoma ME office saw 9 cases of fentanyl and xylazine.
- Xylazine powder can be purchased online from Chinese suppliers.



Trigger Warning: Graphic image of necrotic wound on next slide

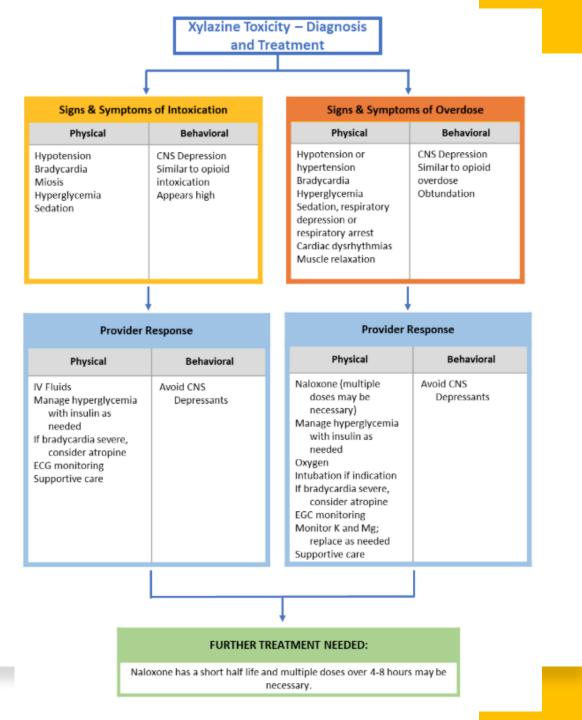
Xylazine ("Tranq")

- Also, can develop severe wounds, including necrosis that may lead to amputation (seeing it even in some patients who deny injecting – rather use via insufflation)
- Potential pathophysiology of xylazine-associated soft tissue infections is possibly multifactorial:
 - Prolongs effects of fentanyl = patients have longer downtime = possible pressure necrosis
 - Peripheral vasoconstrictor = poorer blood flow resulting in an ischemic-like necrosis
 - Possible association with skin-picking behavior which can initiate wounds
 - May be compounded by poor nutritional status in users during chaotic use
 - Some animal evidence that xylazine is directly cytotoxic
 - Repeated trauma due to injection at site where developing wound (some patients state easier to inject directly into a vessel)



Xylazine ("Tranq")

- Mixed reports about whether or not xylazine responds to naloxone
- Potential for physiologic dependence and withdrawal with chronic use/cessation of use has been reported
 - Patients have reported anecdotally that xylazine withdrawal feels different from opioid withdrawal
 - Manifested by significant anxiety



EMNote.org

Narcotic (Opioid) Toxidrome

Mnemonic: "CPR-3H"

C: Coma

P: Pinpoint pupils

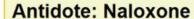
R: Respiratory depression

H: Hypotension

H: Hypothermia

H: Hyporeflexia

NOTE: Meperidine (Demerol) will not cause miosis



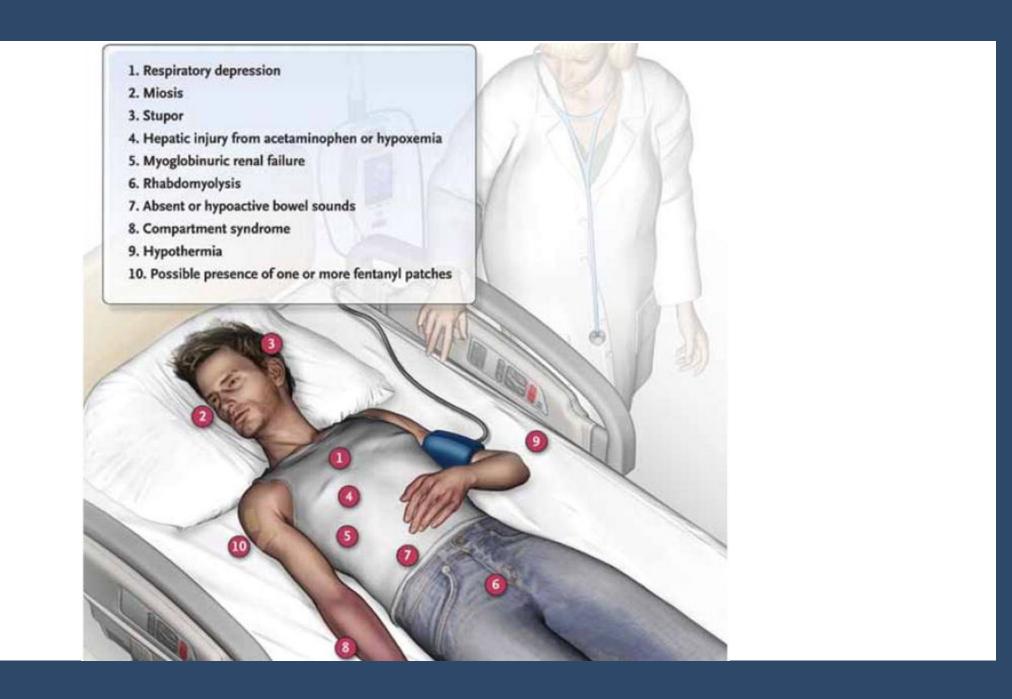
Start with 0.04 mg and titrate up q 2-3 min as need for ventilation to 0.5 mg, 2 mg, 5 mg, up to max 10-15 mg





@ jackcfchong

Review:
Opioid
Toxidrome



Workup

- UDS generally NOT helpful
- Immunoassay that tests for opiates (morphine, heroin, and codeine)
- Semisynthetic opioids like hydrocodone may show up when taken in high doses
- Synthetic opioids, including fentanyl and other NSOs, will NEVER show up
- Must use GC/MS or LC/MS to detect NSOs

Stimulants

- Cocaine/methamphetamine
- Exposure by many routes swallowed, snorted, smoked, injected
- Primary effect is sympathetic stimulation
- Risks include tachydysrhythmias, seizures, rhabdomyolysis

Stimulants

- Duration of action of cocaine significantly less than for meth
- Treatment primarily sedation, possible cooling methods
- Cocaine combined with ethanol results in the formation of cocaethylene, which has a significantly longer half-life
- Cocaine only detectable for < I day in standard UDS --- benzoylecgonine metabolite generally detectable for 3-5 days





Hallucinogens

- Includes LSD, MDMA (Ecstasy), ayahuasca, psilocybin, etc.
 - Most frequent routes of exposure are ingestion, buccal absorption
- Patients experience sensations/images that are perceived as real
- Synesthesia may occur
- Dissociative effects occur with PCP



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Psilocybe genera spores, provided hydrated in aqueous solution. Spores are supplied in 10cc B-D syringes with removable sterile tip cap attached and a sterility packaged 1.5 inch 16 gauge needle. Each Microscopy Kit will include one glass microscope slide, cover slip, cardboard protective case, and instructions on how to properly view your spores under a microscope. Every request containing spores for microscopy will receive one microscopy slide kit per order. Quantity and wholesale discounts available, click on each item for more information and detailed pricing.

California, Idaho, and Georgia residents: Orders requesting Psilocybe Genera Spores shipped to California, Idaho, and Georgia will be refused, voided, or refunded. Possession of these mushroom spores may be illegal in CA, ID, and GA without the proper permissions.

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Psilocybe cubensis: 4 Pack Customer's Choice Grab Bag Special



Our price: \$55.55

Psilocybe cubensis: Albino A+ Spore Syringe Microscopy Kit



Our price: \$18.50

Quantity 1 ~

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Infuse hallucinogenic mushrooms into:

- Gummies,
- Chocolate bars,
- Potions/shots









Polkadot Classic Belgian Milk Chocolate Bar \$25.00

ADD TO CART

Polkadot Cookie Crisp Chocolate \$30.00

ADD TO CART

Polkadot Cookies & Cacao Chocolate \$30.00 \$25.00

ADD TO CART

Polkadot Cookies and Cream Chocolate \$25.00

ADD TO CART









Polkadot Dark Chocolate Bar

\$25.00

ADD TO CART

Polkadot Dark Matcha Chocolate

\$30.00

ADD TO CART

Polkadot Ferrero Rocher Chocolate Bar

\$25.00

ADD TO CART

Polkadot Frosted Flakes Chocolate Bar

\$25.00

ADD TO CART



2nd annual

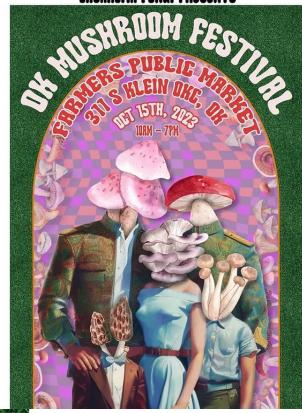
festival will be in

October 2024

WHY OKLAHOMA NEEDS MORE OKLAHOMA'S NATIVE MOREL OKLAHOMA'S POISONOUS MUSHROOMS BY JAMES FROM @MYCTYSONMUSHROOMS THE IMPORTANCE OF MUSHROOM TESTING BY CHRIS PRULI FROM TRYPTOMICS HB2107 - OKLAHOMA'S PSILOGYBIN



OKLAHOMA FUNGI PRESENTS



Stimulants & Hallucinogens

Current 5 Most Commonly Detected Novel Stimulants and Hallucinogens:

(according to April 2024 Quarterly Report from The Center for Forsenic Science Research and Education (CFSRE)):

- I. N,N-dimethylpentylone
- 2. Pentylone
- 3. 2F-2oxo-PCE
- 4. Methylmethcathinone
- 5. Chloromethcathinone

Synthetic Cathinones

- Methylone (2009)
- Butylone (2013)
- n-ethylpentylone (2014)
- Eutylone (2014, surge in 2017-2021)
- N,N-dimethylpentylone (late 2021, surge in 2022)
 - WHO put on the list for international scheduling in March 2024
- All –ylone's (cross hybrid of MDMA) -- different from hexedrones (-drone's) which don't have the methylenedioxy group

Cathinone Mephedrone Methylone NHCH₃ NHCH₃ **Amphetamine**

Synthetic Cathinones

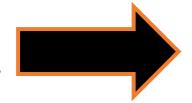
AKA "Bath Salts"



- Drugs designed for a specific pharmacologic effect similar to meth or MDMA.
 - The predominant effects seen to date have been severe agitation, paranoia, hallucinations, and violent behavior.



These Bath Salts



Brandon DeLeon, High On Cloud 9 Bath Salts And Four Loko, Tries To Bite Police: Report (VIDEO)

Also see:

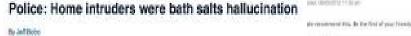
- Elevated temperature/fever
- Rapid heart rate,
- High blood pressure
- Seizures
- Possible muscle breakdown leading to kidney failure
- Extraordinary compulsion to repeat use of the drug

Dangerous Bath Salts Drugs Linked To Nearly 23,000 Hospital ER Visits In 2011



FOLLOW: Bath Salts Emergency Department, Bath Salts Hospital, Bath Salts Hospital Er, Bath Salts Hospital Er Visits, Healthy Living News

Dangerous "bath salts" drugs were linked to tens of thousands of visits to hospital emergency departments in 2011, according to a new government report.



CHURCH HEL. — Police said a Hawkins County man who called \$11 Friday morning complaining of intruders in his home committing thefts was actually alone and hallucinating on beth sate.

Published Rebrusny 27th, 2012 8:44 pm.

About 9 a.m. Friday, Hawkins County Sheriffs Office Deputy Josh Byot responded to the home of Ronald Wayne Sellers, 37, 366 Tipton Road near Church Hill after Sellers called 911 and stated someone had been inside his residence removing forms.

After Byrd and other deputies arrived at the residence, they were told by Selfers that the intruders were in the woods behind his house.

Byid added, "Lasked Mr. Sellers if the subjects were still standing in the location, and he stated. They are still standing there. Don't you see them?" Officers could not see the subjects."





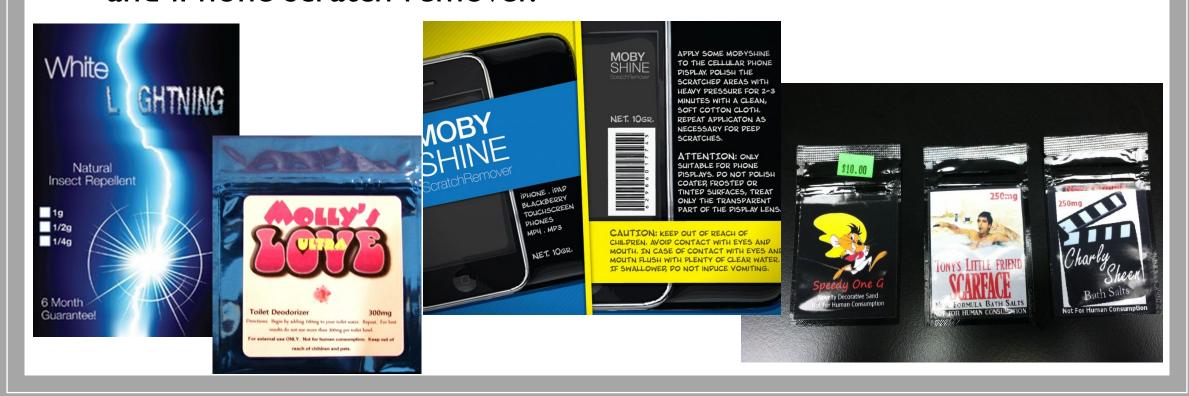


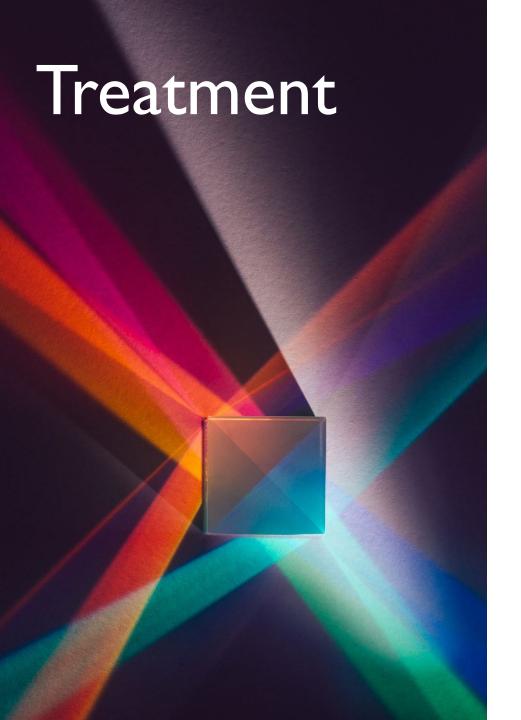
Cross-Dressing Man High on Bath Salts Kills Neighbor's

B 100 1 100 100

Ritan Maylan — Folice in Charleston, West Virginia, found Mark Thompson wandering around the woods in a bre and parties and covered in the blood of his neighbor's pygray guat, which he had killed in his bedroom. He had been high on "bath salus" for three days.

- As with the synthetic cannabinoids, these products are represented to be a legal product.
- Most common is "bath salts," though more recently they've been marketed as insect repellents, plant food, toilet bowl deodorizer, and iPhone scratch remover.









- For the patient experiencing hallucinations, minimize external stimuli (e.g. place in darkened room, no loud noises).
- Liberal use of benzos are the mainstay of therapy
- Be aware that effects of some of these compounds have been reported to last up to 48 hours



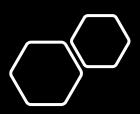
Synthetic Cannabinoids

Emerged on the recreational market in 2010



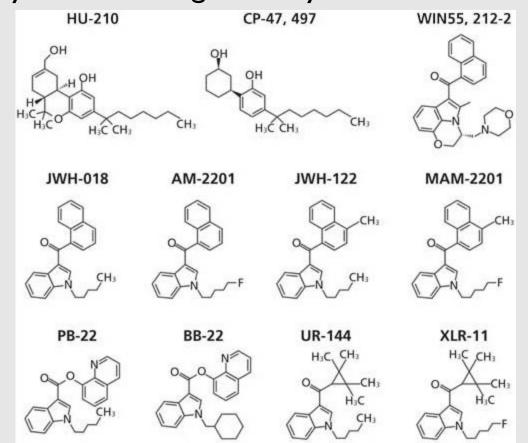
Brand names were big – remember, the people selling these products are in it to make money

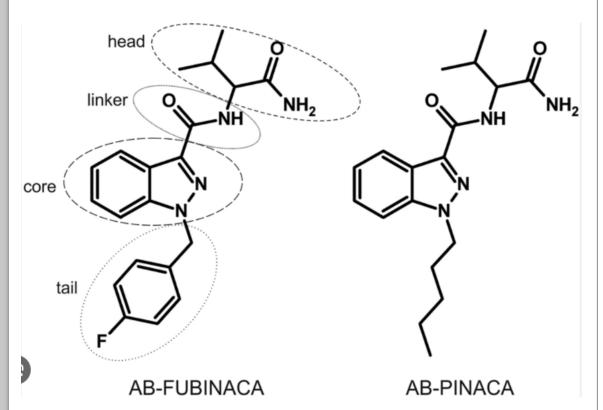




Synthetic Cannabinoids: Challenging to Study

- From chemistry perspective
 - Unlike benzos, opioids, stimulants, and hallucinogens, they are difficult to detect in labs when mixed with other drugs
 - Need own detection methods
- Many labs unwilling to study because of difficulty





Synthetic Cannabinoids Making a Resurgence

- July 2021- China announced plan to ban synthetic cannabinoids based on structural class
 - Similar to what US did with fentanyl analogues
 - Labs in China ceased in production of synthetic cannabinoids
- 2022 into 2023: emergence of inactive drugs & unusual structures with weird names
- End of 2023 into 2024: market responded to ban – China ban did not include the precursor to synthetic cannabinoids so labs producing precursors (synthetic cannabinoids without the tail) then ship them somewhere + add on the tail

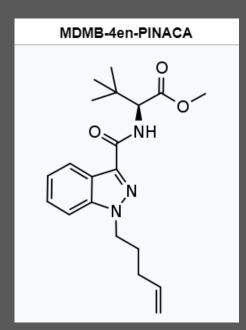
Synthetic Cannabinoids

Current 5 Most Commonly Detected Novel Synthetic Cannabinoids:

(according to April 2024 Quarterly Report from The Center for Forsenic Science Research and Education

(CFSRE)):

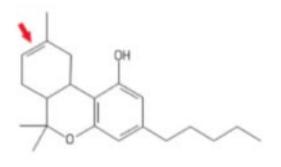
- I. MDMB-4en-PINACA
- 2. 5F-MDMB-PINACA (5F-ADB)
- 3. MDMB-INACA
- 4. ADB-4en-PINACA
- 5. 4CN-CUMYL BINACA



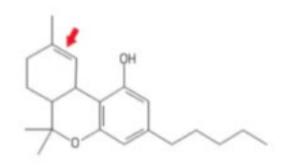




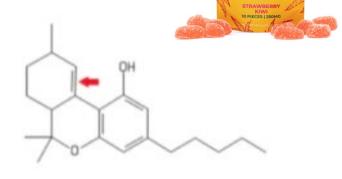
Head shops still exist but now selling semi-synthetic cannabinoids (derived from CBD): delta-8, delta-10, HHC, THC-P







delta-9-THC



delta-10-THC



Galaxytreats.com

Home / Cosmic Krispies / Cosmic Krispies (100MG Delta 8 THC)

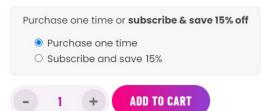


COSMIC KRISPIES (100MG DELTA 8 THC)

\$11.99 - OR SUBSCRIBE AND SAVE 15%

★ ★ ★ ★ (87 customer reviews)

Get ready to launch with our 100mg Delta 8 THC Fruit Cereal Cosmic Krispies. Give it a try and explore the galaxy of uplifting and motivating feeling that Delta 8 THC has to offer!













Utility of UDS



- Though these drugs act on the cannabinoid receptors they are not structurally similar to THC.
- Therefore, they do not show up on a typical urine drug screen.
- In order to be detected, more specific (and expensive) tests such as GC-MS LC-MS/MS must be employed.

Effects & Treatment

- Mild to moderate toxicity: similar to THC
 - Alterations in mood/perception, red eyes, increase in heart rate
 - Potential for hyperemesis with excessive use
 - Other effects: increased blood pressure, agitation, tremors, paranoia, and hallucinations
- Severe toxicity:
 - Severe agitation, hyperthermia, recurrent seizures, psychosis
 - Possible muscle breakdown
- Treatment:
 - Symptomatic and supportive
 - Most do not require specific treatment but some have been intubated for days

Chicago Tribune

APRIL 6, 2018, 4:05 PM

Rat poison likely to blame for deadly side effects of synthetic pot in Illinois

Photo Illustration of K2. (Christopher Smith / Chicago Tribune)

Contaminants?

Contaminants?

Tainted synthetic marijuana linked to severe bleeding cases in Florida

By Nicoletta Lanese published 2 days ago

Florida's Poison Control Centers urge people to go to the emergency room if they experience bleeding after using synthetic marijuana, or "spice."









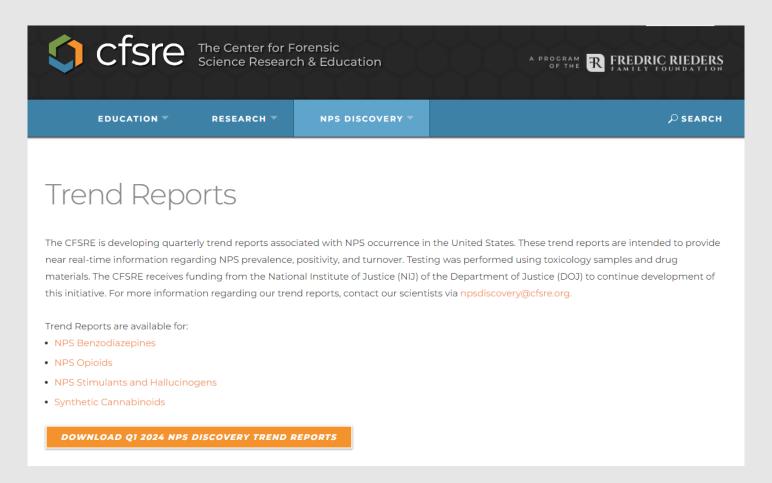






The Center for Forensic Science Research & Education

https://www.cfsre.org/nps-discovery/trend-reports



Kratom

- Derived from the plant Mitragyna speciosa, which is native to southeast Asia
- Typically, the leaves of the plant are ingested (often in capsule form), but leaves can also be chewed; occasionally smoked
- Unregulated at the federal or state (Oklahoma) level
- In low doses, has stimulant and antidepressant effects
- In higher doses, opioid effects begin to be observed
- Frequently sold in "smoke shops" and convenience stores



ONE GRAM

Tianeptine ("Gas Station Heroin")

 Not approved for use in the US but is marketed in Europe, Asia, and Latin America as a prescription antidepressant

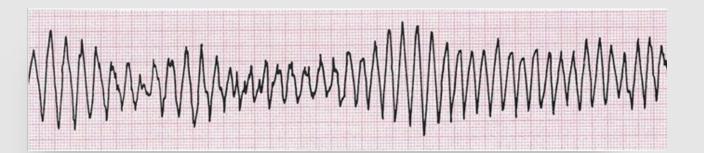


- Relatively easy to obtain at convenience stores, gas stations, and on the internet
- Potential for physical dependence and withdrawal with chronic use/cessation of use









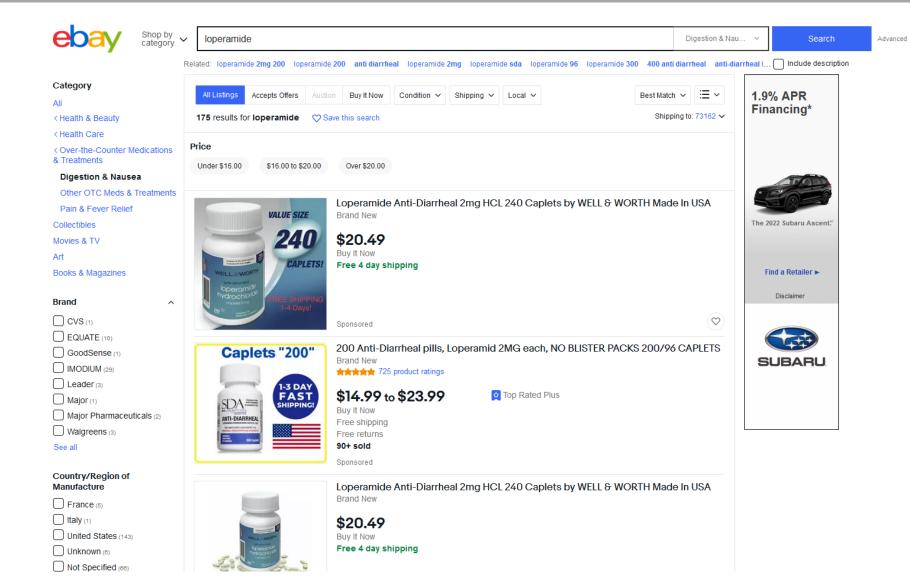
Loperamide

- Sold OTC for treatment of diarrhea
- In addition to opioid effects, can cause cardiac conduction disturbances (QRS/QTc widening), torsades de pointes





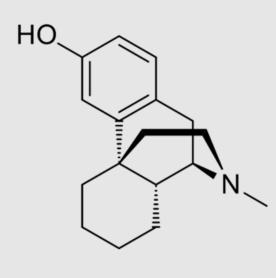
Often taken with a p-glycoprotein inhibitor (omeprazole, lansoprazole) to increase CNS penetration





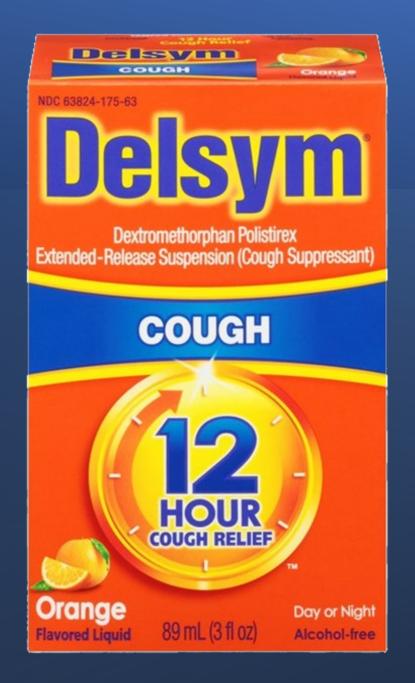
$$H_3C$$

dextromethorphan



levorphanol

- A dextrorotary analog of levorphanol, a powerful narcotic
- Causes dissociative effects, opioid-like effects at high doses

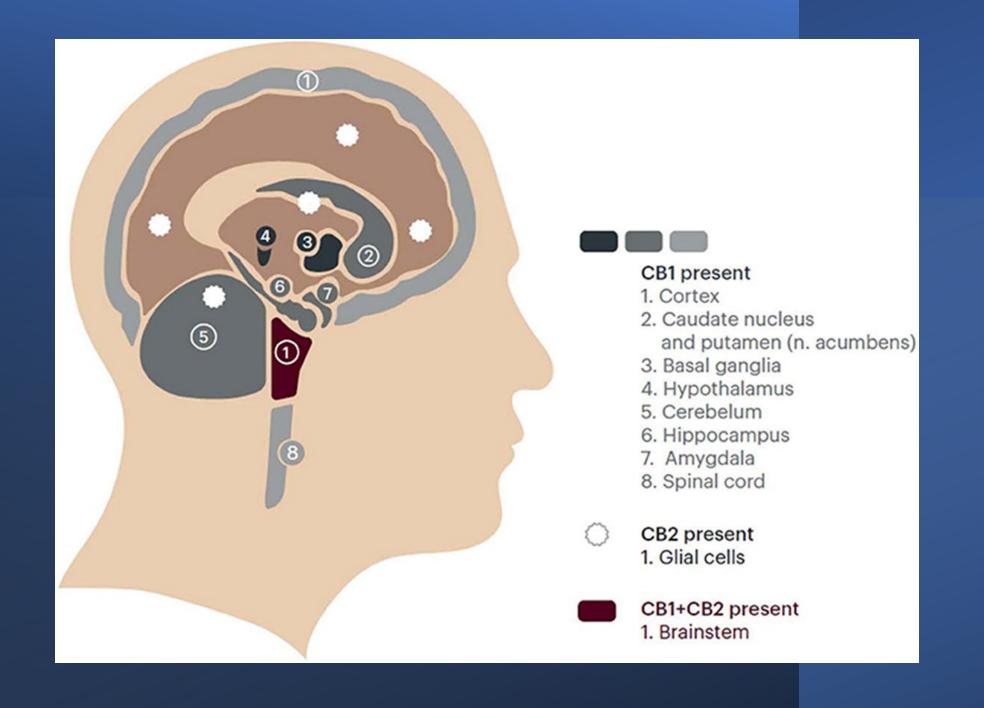




Marijuana



- The human body produces endocannabinoids, i.e., endogenous compounds (anandamide, 2-arachidonoylglycerol) that act at CBI and CB2 receptors
- Cannabinoids derived from plants are called phytocannabinoids: THC, cannabidiol, and cannabinol



Brain

- · Learning, memory, cognition
- Motor control
- Anxiety and depression
- Appetite and food intake
- Reward and addiction
- Neuroprotection
- Neural development
- Sleep

Liver

- Ascites formation
- Lipogenesis
- Fibrosis
- Insulin resistance

Reproductive system

- Fertility regulation
- Embryo implantation
- Embryonic development

Cardiovascular system

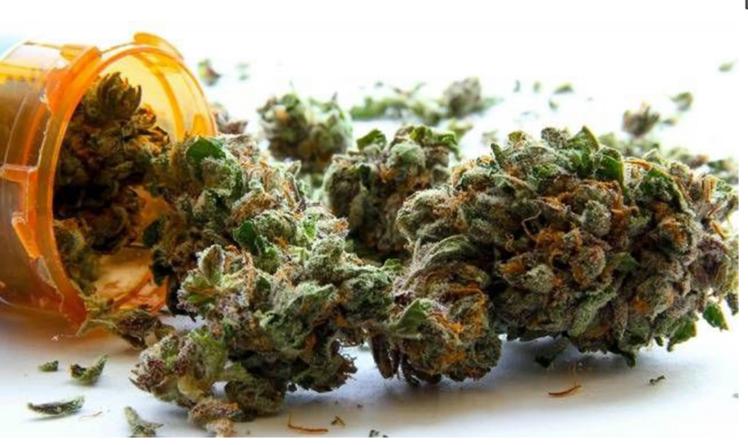
- Negative inotropy
- Vasodilation
- Cardiac function

Skeletal muscles

- Energy metabolism
- · Muscle fiber formation

GI tract

- GI motility
- Enteroendocrine function
- Intestinal barrier function
- Energy balance
- Nociception
- Immune modulation
- Nausea and emesis
- · Intraocular pressure
- · Bone remodeling
- Bronchodilation
- Cancer
-



Effects vary by species / strain

Sativa: More euphoria, stress relief

Indica: Relaxation, physical

(especially pain) relief

Sativa and Indica are often combined



THCV CBG CBC CBL CBT CBE

Sabinene Delta-3 Carene Alpha-Phellandr Beta-Mrycene Alpha Terpinene monene

CANNABINOIDS

TERPENOIDS

FLAVANOIDS



Genesis Testing Labs 1620 South Main St., Unit A Grove, OK 74344 (918)791-3460 OMMA # LAAA-EKZG-UNEB

Certificate of Analysis

Client Name Full Tilt Cannabis
Address 4017 27th Ave. No

4017 27th Ave. NE Miami, OK 74354

Sample Type

Phone

 Sample ID
 SA-071521-10329

 Strain
 Platinum OG 3141

Sample Batch # Date Received

License Number

PAAA-CLAZ-QRID 253-331-0941

Concentrate

POG-062421 07/15/2021

Sum	mary of F
file	
Result mg/g	
1.91571	
6.22605	7 I
0.91954	
	7 [
34.8276	
	7 [
996.964	
1.9636	7 [
90.9165%	7 [
909.165	
	Result mg/g - 1.91571 - 6.22605 0.91954 - 34.8276 - 996.964 1.9636 90.9165%

Microbiologi	cal Profile	
All Results	Pass	

Mycotoxin Screening	
All Results	Pass

Residual Solvents	
All Results	Pass

Heavy Metals Screening	
All Results	Pass

Pesticide Screening	
All Results	Pass

Water Activi	Water Activity	
All Results	NT	

Terpene Profile		
Terpene	Result %	
Camphene	0.0137931	
Myrcene	0.136398	
β-pinene	0.0477011	
3-Carene		
Delta-limonene		
Cymene		
β-Ocimene		
Eucalyptol		
y-Terpinene		
a-Terpinene		
Terpinolene	0.0145594	
Linalool	0.0559387	
Isopulegol	0.0810345	
Geraniol		
β-Caryophyllene	1.14215	
Humulene	0.356897	
Nerolidol		
Guaiol		
Caryophillene oxide		
α-Bisabolol	0.0204981	
Total Terpene %	1.86897	

...

OMMA Patient Advisory:

Our QA lab has confirmed the presence of THC-O-acetate (a manufactured THC compound) in a concentrate by the name of Platinum OG (BA#-POG-062421). This manufactured THC compound has been shown to be harmful when added to other THC-containing products.



PATIENT ADVISORY

Our QA lab, Metis, has confirmed the presence of THC-O-acetate in a concentrate by the name of Platinum OG (BA#-POG-062421).

THC-O-acetate is NOT a medical marijuana product and has been shown to be very harmful when added to other THC-containing products, causing side effects such as seizures, difficulty speaking, and vomiting.

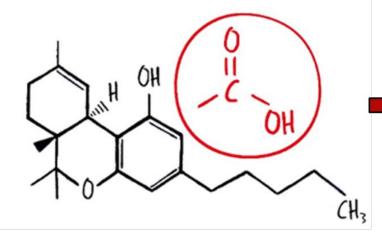
If you've had negative health outcomes after using a medical marijuana product purchased from a licensed dispensary, please immediately contact a medical provider or the Oklahoma Center for Poison and Drug Control (800-222-1222).



Oct. 15, 2021

THCA

Can't bind to CB receptors





THC

Bioavailable and ready to bind with CB receptors















Other Resources

- erowid.org
 - A rich source of information, including "trip reports," pharmacology information, legal status, etc.
 - Information is usually reliable
- bluelight.ru
- drugs-forum.com
- tripsit.me



OKLAHOMA PQSON CENTER 1-800-222-1222

Drugs of Abuse

Eszter Moore, Pharm.D., CSPI Oklahoma Poison Center