

The Other Opioid Crisis: Heroin and Fentanyl

Kevin L. Zacharoff, MD, FACIP, FACPE, FAAP

Disclosures

Nothing to disclose



Learning Objectives

- Describe the opioid overdose crisis in the United States today
- Discuss common beliefs and inconsistencies about the role of prescription opioids in the opioid crisis
- Identify the intersection of illicit drug use and clinical pain practice in today's "opioid epidemic"
- Provide clinically relevant recommendations for navigating the current landscape without depriving pain care to patients in need





Is There More than one Opioid Crisis?

The Facts: No Lack of Media Attention Painweek The Facts: No Lack of Media Attention The Facts: No Lack of Media Attention Painweek The Facts: No Lack of Media Attention The Facts: No Lack of Media Attention Painweek The Facts: No Lack of Media Attention The Facts: No Lack of Media Att

- People are dying…
- People are angry and reacting
- People are scared
- Overdoses from drugs (<u>ALL</u> drugs, legally prescribed or not) including opioids, hallucinogens, cocaine, etc
 - -~ **64,000 deaths** nationwide in 2016
 - •**121%** from 2015
 - More than 42,000 (66%) involved some type of opioid

There Is More Than One Opioid Crisis. To better tackle the epidemic, Kentucky needed to know which drugs were causing lethal overdoses. By Kathryn Casteel Published Jan. 17, 2018. https://fivethirtyeight.com/features/there-is-more-than-one-opioid-crisis/. Accessed January 18, 2018.

The Facts

- Drug(s) responsible may be unknown
 - Overdose is usually about RESPIRATORY DEPRESSION
- Sometimes causes are not reported
 - · May depend on the state
 - · Trends can be difficult to identify
- This may lead to a lack of focus

There Is More Than One Opioid Crisis. To better tackle the epidemic, Kentucky needed to know which drugs were causing lethal overdoses. By Kathryn Casteel Published Jan. 17, 2018. https://fivethirtyeight.com/features/there-is-more-than-one-opioid-crisis/. Accessed January 18, 2018.

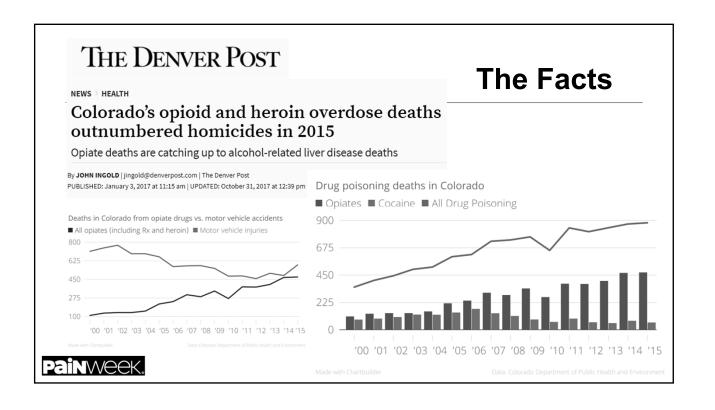
Painweek.

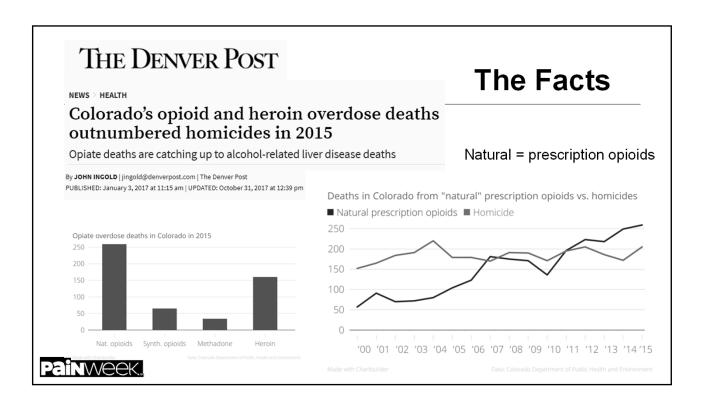
Identifying drugs in overdose deaths

In each state, 2016

	_	CASES IN WHICH NO DRUG WAS SPECIFIED		
STATE	ALL DEATHS	TOTAL	SHARE	
Louisiana	996	473	47.5%	
Pennsylvania	4,627	2,075	44.8	
Alabama	756	308	40.7	
Montana	119	46	38.7	
Indiana	1,526	547	35.8	
Delaware	282	99	35.1	
Nebraska	120	37	30.8	
Arkansas	401	115	28.7	
Florida	4,728	1,144	24.2	
Idaho	243	55	22.6	
New Jersey	2,056	461	22.4	
Mississippi	352	78	22.2	
Wyoming	99	21	21.2	
California	4,654	930	20.0	
Kansas	313	62	19.8	
Colorado	942	172	18.3	
Kentucky	1,419	253	17.8	
Missouri	1,371	199	14.5	
North Dakota	77	11	14.3	
Arizona	1,382	196	14.2	

The Facts: Kentucky as an Example The most common drugs found in Kentucky's overdose victims Based on an analysis of 1,471 drug overdose deaths in 2016 Than One Opioid Crisis. To better tackle the epidemic, Two-drug combinations ndividual drugs Three-drug combinations Kentucky needed to know which drugs DRIIG SHARE DRIIGS SHARE DRIIGS SHARE were causing Morphine 45.4% Heroin, morphine 24.3% Heroin, morphine, codeine 16.6% lethal overdoses. By Kathryn Casteel Fentanyl Fentanyl, morphine 23.7 Heroin, morphine, fentanyl Published Jan. Morphine, codeine, fentanyl 17, 2018. https://fivethirty Alprazolam Heroin, codeine 16.6 Heroin, morphine, THC-COOH eight.com/featu тнс-соон Gabapentin, morphine 14.2 Heroin, codeine, fentanyl 7.6 res/there-is-Heroin 24.7 Morphine, THC-COOH 12.8 Fentanyl, morphine, THC-COOH 7.3 opioid-crisis/ Accessed Codeine 20.7 Alprazolam, morphine 12.6 Alprazolam, heroin, morphine 7.3 January 18, 2018. Ethanol 18.4 Methamphetamine, amphetamine 12.0 Gabapentin, heroin, morphine 7.1 Oxycodone 17.2 Heroin, fentanyl 11.7 Gabapentin, fentanyl, morphine 6.8 Methamphetamine 17.1 Oxycodone, oxymorphone 11.5 Alprazolam, codeine, morphine In cases in which multiple drugs are detected, every individual drug — as well as every two- and three-drug combination — is counted separately.





THE DENVER POST

POLITICS > COLORADO POLÍTICS

Here's how Colorado is combating the prescription opioid and heroin epidemic

Colorado is exploring opioid prescription limits, prescriber report cards and new treatment options.

By **JOHN FRANK** | Jfrank@denverpost.com | The Denver Post PUBLISHED: November 5, 2017 at 12:01 am | UPDATED: November 6, 2017 at 11:02 am

The nation's drug epidemic kills someone in Colorado about every 9 hours and 36 minutes, a fact that rings like a siren for state leaders who are combating the leading driver: prescription and illicit opioids.

The most alarming trend is the rate of Seroin deaths. The number of overdoses caused by heroin, an opioid, continues to skyrocket, contributing to 228 deaths last year, compared with just 79 five years earlier, according to state

The Facts





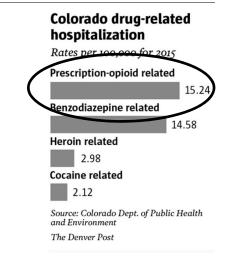
THE DENVER POST

- "The state tracks opioid prescriptions, but doctors aren't required to check the database"
- "The vast majority of states require clinicians to check the Prescription Drug Monitoring Program (PDMP) database before writing most opioid prescriptions"

By **JOHN FRANK** | Jfrank@denverpost.com | The Denver Post PUBLISHED: November 5, 2017 at 12:01 am | UPDATED: November 6, 2017 at 11:02 am

Painweek.

The Facts



THE DENVER POST

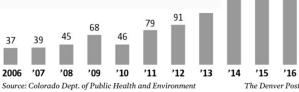
The Facts

228

151

A crackdown on overprescribing isn't enough — drug treatment options remain a challenge.

According to policymakers, Colorado's initial efforts to restrict opioid prescriptions may have an unintended effect: pushing addicts to opioids such as heroin or fentanyl. The trend is spotlighting the limited treatment options for drug users who want help.



By **JOHN FRANK** | jfrank@denverpost.com | The Denver Post PUBLISHED: November 5, 2017 at 12:01 am | UPDATED: November 6, 2017 at 11:02 am



THE DENVER POST The Facts Colorado opioid-related deaths Doctors will soon get opioid prescription "report cards." Rates per 100,000 for 2016 Male ■ "The idea is that Female To educate doctors about the dangers of prescribers might be 6.3 overprescribing opioids, Colorado is testing a prescribing more than relatively new idea that is showing early 20 to 24 years average for their results - prescriber report cards. 25 to 34 years particular specialty" 35 to 44 years The reports, also known as scorecards, will ■ "They'll say 'Oh, I better 45 to 64 years give clinicians a summary of their prescriber look at things more history and how they rank with their peers in closely" the same specialty in terms of dosage, White, non-Hispani duration and type of drug. ■ "What we are hoping to White, Hispanic Black/African-American see is a change in Asian/Pacific Islander prescriber behavior" American Indian By JOHN FRANK | jfrank@denverpost.com | The Denver Post PUBLISHED: November 5, 2017 at 12:01 am | UPDATED: November 6, 2017 at 11:02 am Source: Colorado Dept. of Public Health and Environ Painweek

IMPORTANT SOCIETAL QUESTIONS



Is the Current Climate Living up to Promises Made in the Past?

Does the chronic pain patient still have rights?

Is this a problem about chronic pain patients abusing prescription pain medications?

Bill of Rights for People with Chronic Pain

The right to have your report of pain taken seriously and to be treated with dignity and respect by doctors, nurses, pharmacists and other healthcare professionals.

The right to have your pain thoroughly assessed and promptly treated.

The right to be informed by your doctor about what may be causing your pain, possible treatments, and the benefits, risks and costs of each.

The right to participate actively in decisions about how to manage your pain.

The right to have your pain re-assessed regularly and your treatment adjusted if your pain has not been eased.

The right to be referred to a pain specialist or other healthcare provider if your pain persists.

The right to get clear and prompt answers to your questions, take time to make decisions, and refuse a particular type of treatment if you choose.

The Clinical

Implications

THE DENVER POST

NEWS > HEALTH

Chronic pain patients say they are hurt by Colorado's opioid prescription guidelines

Some Colorado doctors are refusing to prescribe opioids to chronic pain sufferers



David and Marjorie Orthman discuss David's painful form of cystic fibrosis Dec. 6, 2016 at their home.

By JOHN INCOLD | Jingold@denverpost.com | The Denver Post
PUBLISHED: December 6, 2016 at 9:27 nm | UPDATED: December 7, 2016 at 10:31 am

The Denver Post C

Now, across Colorado and the rest of the nation, these policies intended to address opioid abuse have unexpectedly harmed patients who depend on the drugs to treat chronic conditions, pain specialists and patient advocates say. The policies are supposed to offer guidance — helpful

prescribing more than a certain amount of opioids to any one patient.

advice to doctors to be cautious in

Officials at the Colorado Department of Regulatory Agencies are looking at revising the guidelines. The Colorado Consortium for Prescription Drug Abuse Prevention, a group of medical and drug experts, is also working on ideas to protect pain patients while cracking down on opioid abuse.







Drug overdose deaths in the United States continue to increase in 2015

- The majority of drug overdose deaths involve an opioid¹
- Since 1999 the # of overdose deaths quadrupled²
 - -Prescription opioids
 - -Heroin
- ■91 Americans die every day from an opioid overdose



Rudd RA, Seth P, David F, Scholl L. Increases in Drug and Opioid-Involved Overdose Deaths — United States, 2010–2015 MMWR Morb Mortal Wkly Rep. ePub: 16 December 2016. DOI: http://dx.doi.org/10.15585/mmwr.mm655051e1 CDC. Wide-ranging online data for epidemiologic research (WONDER). Atlanta, GA: CDC, National Center for Health Statistics; 2016. Available at http://wonder.cdc.gov.



Understanding the Epidemic

The Facts



Heroin use is trending up

- From 2002-2013, past month/year heroin use and addiction have all increased among 18-25 year olds¹
- Among new heroin users, 75% report abusing prescription opioids prior to using heroin²
- Heroin-related deaths more than tripled from 2010-2015
 - -The largest increase was for those heroin-related deaths involving synthetic opioids - FENTANYL



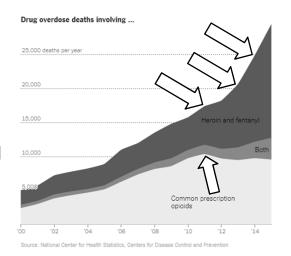
Centers for Disease Control and Prevention. Demographic and Substance Use Trends Among Heroin Users — United States, 2002–2013. MMWR 2015;

The Facts: Times are Changing

The New York Times

Short Answers to Hard Questions About the Opioid Crisis

- Characteristics of opioid drug overdoses:
 - -Fast
 - -Deadly
 - -Scary
 - -Socioeconomic status neutral
 - -Abuse-history neutral
 - -Increasing exponentially







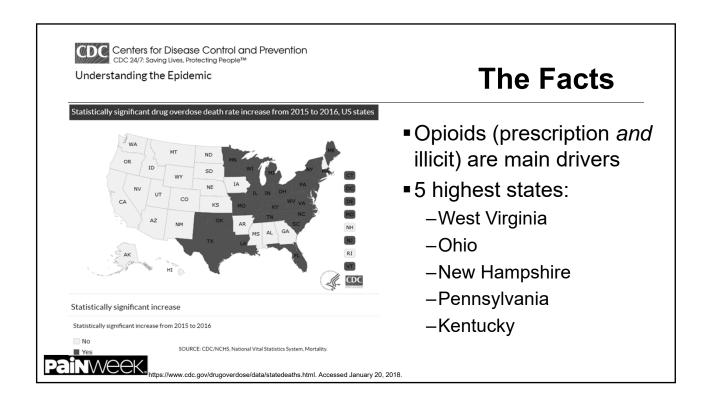
Understanding the Epidemic

The Clinical Implications



- Improved opioid prescribing
- Expanded access to substance abuse treatment
- Naloxone
- Prescription Drug Monitoring Programs (PDMPs)
- State-level strategies to prevent high risk prescribing and opioid overdoses
- Improved detection of illegal opioid use by law enforcement







The "Other" Epidemic

Heroin

- Highly addictive
- It is an opioid
- Made from morphine
 - ~3 times more potent
- A natural substance
 - -Extracted from the opium poppy plant
 - Asia
 - Mexico
 - Colombia

The Facts





PainWeek.

https://www.drugabuse.gov/publications/drugfacts/heroin. Accessed January 22, 2018.



Understanding the Epidemic

- Heroin use has been increasing¹
 - -Men
 - -Women
 - -Most age groups
 - -All income levels
- Past misuse/abuse of prescription opioids is the strongest risk factor for starting heroin²

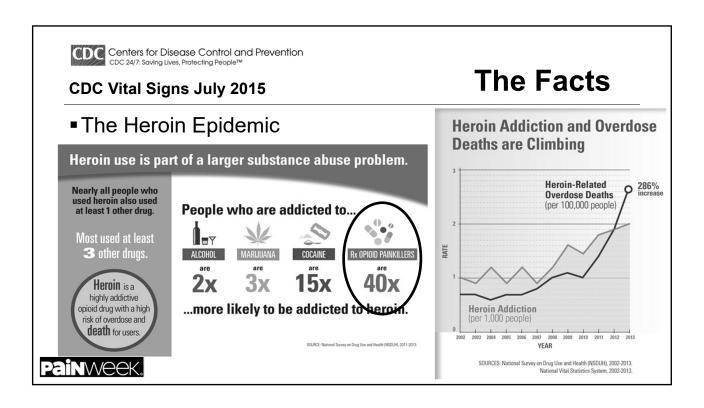
The Facts

	raphic G		
SEX	2002-2004*	2011-2013*	% CHANGE
Male	2.4	3.6	50%
Female	0.8	1.6	100%
AGE, YEARS			
12-17	1.8	1.6	
18-25	3.5	7.3	109%
26 or older		1.9	58%
RACE/ETHNICITY			
Non-Hispanic white		3 1.7	114%
Other		1.7	
ANNUAL HOUSEHOLD	INCOME		
Less than \$20,000	3.4	5.5	
\$20,000-\$49,999		2.3	77%
\$50,000 or more		1.6	
HEALTH INSURANCE C	OVERAGE		
None	4.2	6.7	60%
Medicaid	4.3	4.7	-
Private or other	0.8	1.3	63%

*Annual average rate of heroin use (per 1,000 people in each group)



. Centers for Disease Control and Prevention. Vital Signs: Today's Heroin Epidemic – More People at Risk, Multiple Drugs Abused. MMWR 2015.
Compton WM, Jones CM, and Baldwin GT. Understanding the Relationship between Prescription Opioid and Heroin Abuse. NEJM.





"The relationship between prescription opioid abuse and increases in heroin use in the U.S. is under scrutiny. These substances are all part of the same opioid drug category and overlap in important ways"





https://www.drugabuse.gov/publications/research-reports/relationship-between-prescription-drug-abuse-heroin-use/introduction. Accessed January 30, 2018
 https://d14rmgtrwzf5a.cloudfront.net/sites/default/files/rx_and_heroin_rrs_layout_final.pdf. Accessed January 30, 2018.



Research Report Series

The Facts

- "Current" ~ 2013-2015 research regarding prescription opioids and heroin shows:
 - -Prescription opioid abuse is a risk factor for heroin use
 - Heroin use is *rare* in prescription drug abusers (<4% start within 5 years)
 - -Although similar effects, risk factors are different
 - A subset of people who abuse prescription opioids may progress to heroin use
 - -Availability of drug(s) is associated with increased use and overdose
 - -Heroin use is driven by cost advantage and availability
 - -Emphasis is needed on prevention and treatment



https://www.drugabuse.gov/publications/research-reports/relationship-between-prescription-drug-abuse-heroin-use/introduction. Accessed January 30, 2018
 https://d14rmgtrwz/5a.cloudfront.net/sites/default/files/rx_and_heroin_rrs_layout_final.pdf. Accessed January 30, 2018.



Research Report Series

The Facts

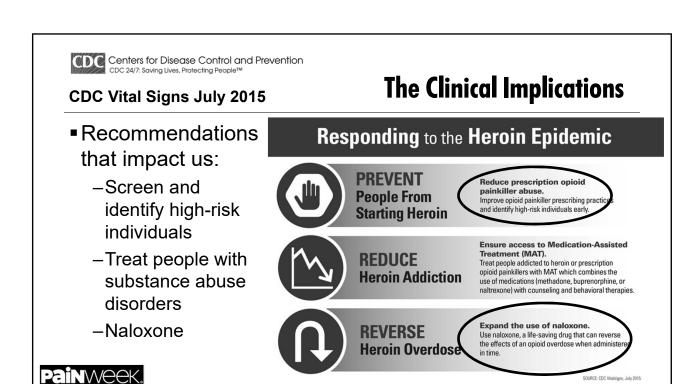
- And...
 - -"Analyses suggest that those who transition to heroin use tend to be frequent users of multiple substances (polydrug users)"
 - Jones CM. Heroin use and heroin use risk behaviors among nonmedical users of prescription opioid pain relievers United States, 2002-2004 and 2008-2010. Drug Alcohol Depend. 2013;132(1-2):95-100.

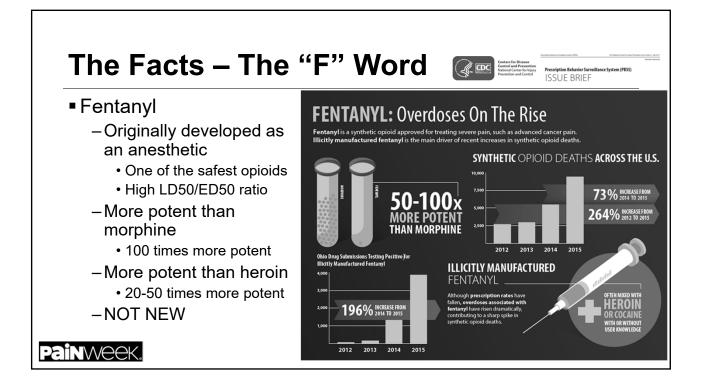
 $A + B \neq C$





. https://www.drugabuse.gov/publications/research-reports/relationship-between-prescription-drug-abuse-heroin-use/introduction. Accessed January 30, 2018. https://d14rmgtrwzf5a.cloudfront.net/sites/default/files/rx_and_heroin_rrs_layout_final.pdf. Accessed January 30, 2018.





Case 1

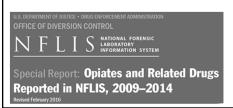
- 42-year-old Hispanic male
 - -Addicted to heroin
 - -Comes to the same street corner every day to buy heroin
 - First dose usually free
 - -He's actually buying heroin laced with fentanyl
 - -"It's a new epidemic" he says
 - -"If you catch a bag of pure fentanyl, that Narcan ain't bringing you back"
 - -"I just watched my friend die from fentanyl"

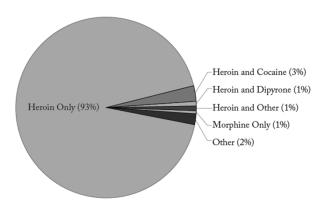




Heroin and Fentanyl: A "Perfect" Pair

- The overwhelming majority of the time fentanyl is paired with heroin
- To get a better "high"
- "Better" economic profile





Note: Percentages may not sum to total because of rounding.

Fentanyl - Drug seizures involving fentanyl are going up dramatically Drug seizures containing fentanyl 30.000 Sentanyl are going up dramatically Drug seizures containing fentanyl 30.000 Sentanyl reports doubled in 2016 Sentanyl reports

The Facts **Fentanyl** Defined geographic Figure 2 NFLIS regional trends in fentanyl reported per 100,000 persons aged 15 or older, January 2001-December 20151 distribution Number of Fentanyl Reports (per 100,000) 2 9 6 51 ■ West ▲ Midwest 12 Northeast South 2001 2002 2003 2004 2005 2006 2007 2008 2009 2010 2011 2012 2013 2014 2015 Note: U.S. census 2015 population data by age were not available for this publication. Population data for 2015 were imputed. NFLIS Brief: Fentanyl, 2001-2015 ¹A dashed trend line indicates that estimates did not meet the criteria for precision or reliability, including the 2002 estimate in the West and the 2001 estimate in the Northeast. Painweek.

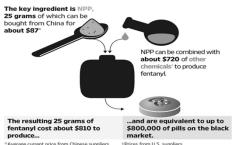
The Chinese Connection Fueling America's Fentanyl Crisis

A vast network beginning in China feeds fentanyl, a deadly synthetic opioid, to the U.S., Mexico and Canada

By JEANNE WHALEN and BRIAN SPEGELE Updated June 23, 2016 1:44 a.m. ET

Painweek.

Traffickers manufacturing fentanyl often purchase the key ingredient from China, which doesn't regulate its sale. Here's how the chemical building blocks become a highly profitable street drug.



Criminal Chemistry



- Fentanyl can be manufactured anywhere
 - -Synthetic
- N-Phenethyl-4piperidinone is NPP
 - Intermediate precursor to fentanyl
- It is cheap
- It is not going anywhere

Terminology

The Facts

- A very small amount of fentanyl is potentially fatal
- This applies to licit or illicit fentanyl
 - Illicitly manufactured fentanyl is likely no more dangerous than legally manufactured fentanyl

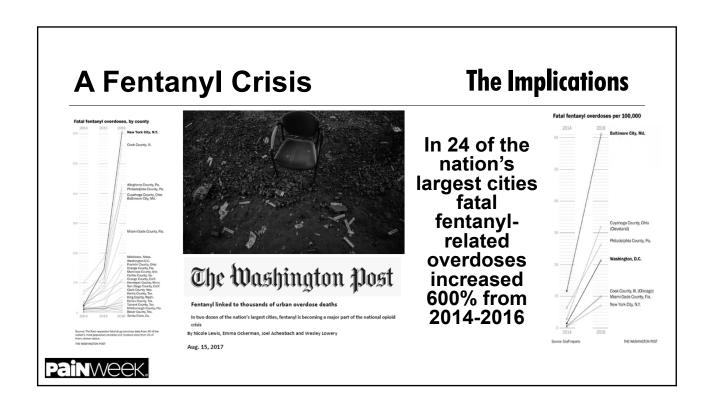


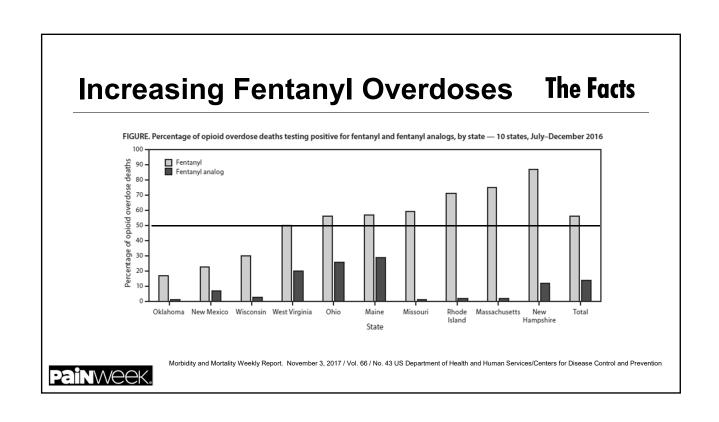
- Fentanyl label:
 - FENTANYL CITRATE SHOULD BE ADMINISTERED ONLY BY PERSONS SPECIFICALLY TRAINED IN THE USE OF INTRAVENOUS ANESTHETICS AND MANAGEMENT OF THE RESPIRATORY EFFECTS OF POTENT OPIOIDS

THE WALL STREET JOURNAL

- AN OPIOID ANTAGONIST, RESUSCITATIVE AND INTUBATION EQUIPMENT AND OXYGEN SHOULD BE READILY AVAILABLE
- Fentanyl may cause muscle rigidity, particularly involving the muscles of respiration
- DIMINISHED SENSITIVITY TO CO2 STIMULATION MAY PERSIST LONGER THAN DEPRESSION OF RESPIRATORY RATE







Different Types of Fentanyl and Presentations

- Different formulations and varying potencies:
 - Fentanyl
 - Analogs
 - · Acetyl fentanyl
 - Ocfentanil
 - Carfentanyl
 - Remifentanyl
 - Alfentanyl
 - Sufentanyl
 - Furanylfentanyl
- Presentations
 - Powder
 - Counterfeit pills
 - -Etc





Carfentanyl

- Synthetic
- Large animal anesthetic
- 100 times more potent than fentanyl
- 10,000 times more potent than morphine
- Airborne/skin exposure often fatal
 - –2012 Moscow Theater Hostage Crisis

The Facts



By Lynh Bui and Peter Hermann April 26, 2017
Elephant tranquilizer is the latest lethal



Members of the Royal Canadian Mounted Police go through a decontamination procedure in Vancouver, British Columbia, in June 2016 at intercepting a package containing approximately one kilogram (2.2 pounds) of the opioid carfentanil imported from China, (Royal Canadia

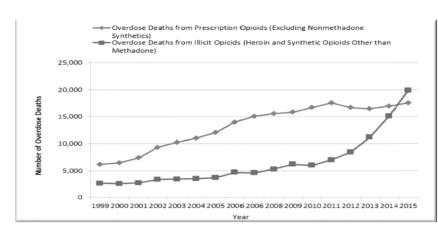


- Fentanyl in the wrong hands is deadly
- Potentially contributing significantly to overdose/fatal overdose risk
- Analogs not often tested for in routine toxicology testing
- Naloxone is a bridge to survival but not a final solution
 - -Re-narcotization from fentanyl is common
 - -CO₂ will persist
 - -Diminished hypoxic drive may persist



SUMMARY

The Co-existing Opioid Crises The Facts

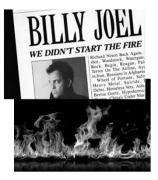






We Didn't Start the Fire...Or did We?

- Significant controversy exists to the relationship between prescription opioids and the heroin/fentanyl epidemic
 - -Tracking back to prescribers
 - The 1990s when opioid prescribing increased
 - Pain being designated as the "fifth vital sign" in 2000
 - Pain Bill of Rights
 - Evolution of "pill mills"
 - -Related to cost and availability
 - · Heroin and fentanyl are cheaper and stronger





Case 2

- ■26-year-old white female
 - -Has 2 children ages 4 & 7 who live with their godmother
 - Became addicted to opioids after being prescribed oxycodone post C-section
 - -Addiction to oxycodone transitioned to heroin
 - -Addiction to heroin transitioned to heroin/fentanyl
 - -"If there's no fentanyl in it, I don't want it at all"
 - -Tried 14-day rehab without success
 - "This is all I know anymore... It's all I know"







Who and What is Our Responsibility?

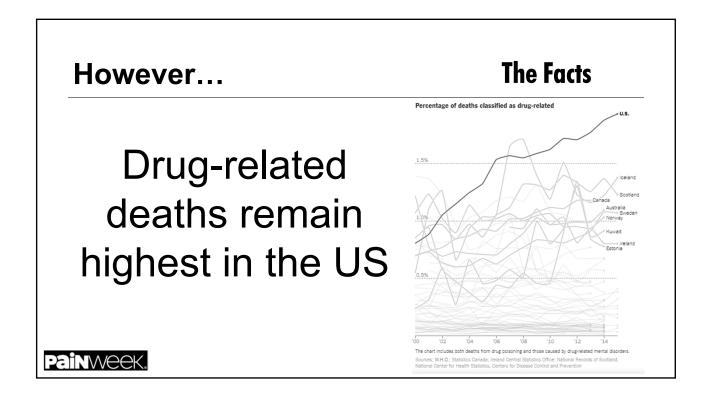
We Are Involved Like it or Not

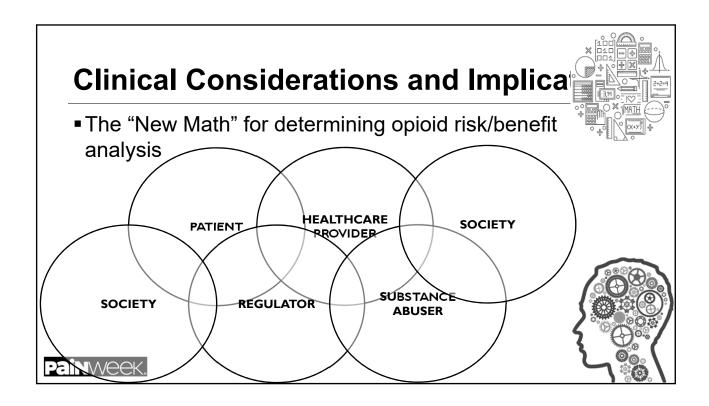
Does the United States Own the Problem??

- We are not alone
- This is a global issue
- In 2015, approximately ¼ billion people abused/misused drugs
 - ~29.5 million (0.6% of global adult population) engaged in problematic use/suffered from substance use disorder
- 70% of the global burden of disease caused by drug use disorders attributable to opioid use









Final Thoughts

- We must consider the parallel "opioid" crises that exist today Our role in these problems
- Heroin and fentanyl are not going anywhere
- Remember the "New Math"
- Naloxone is likely something to keep top of mind
- Avoidance of "risky" situations is critical
- Strongly consider and document appropriate candidates for fentanyl-based therapies
- Everyone is looking at us as an integral part of solving this "other" opioid crisis
- Everyone is looking to us to be a part of the solution





Final Thoughts

But... Let's not make patients "pay" for these crises



"Cure sometimes, treat often, comfort always."

— Hippocrates



