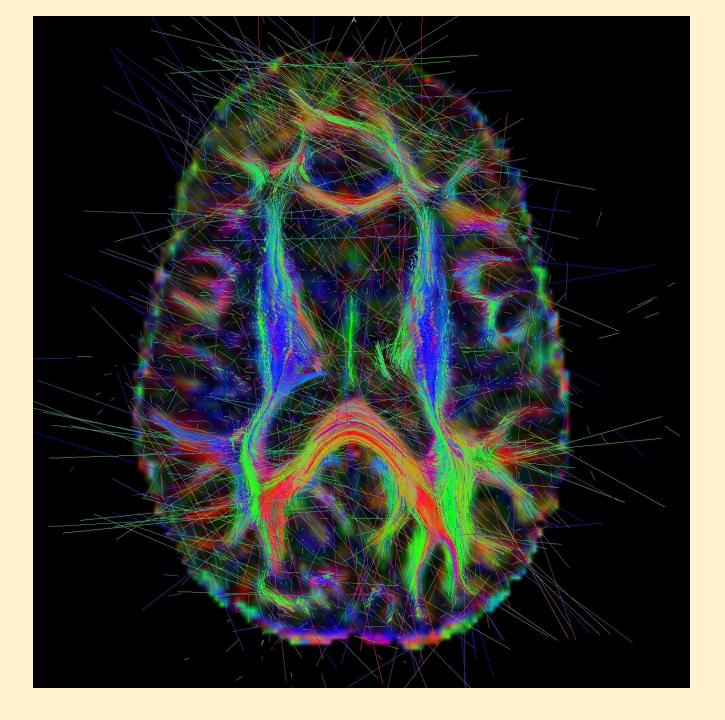
# THE ADOLESCENT BRAIN

& The Physiology of Addiction

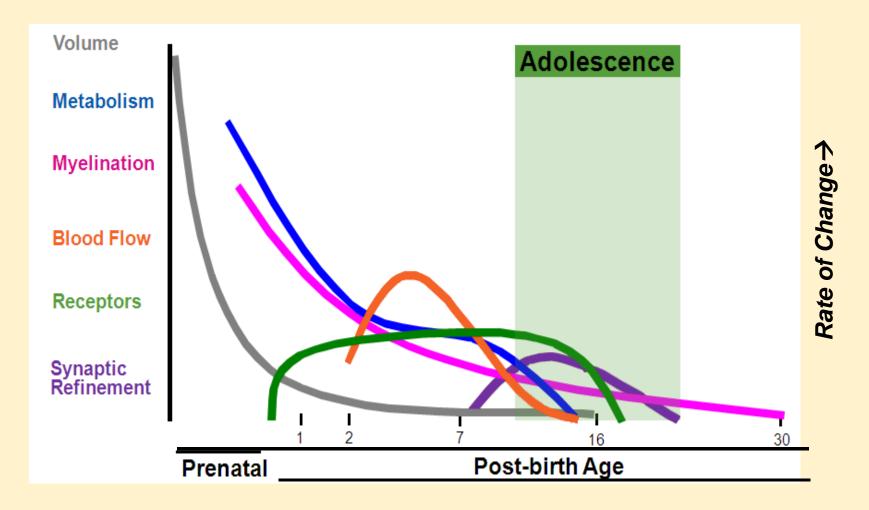
## Ruth A. Potee, MD

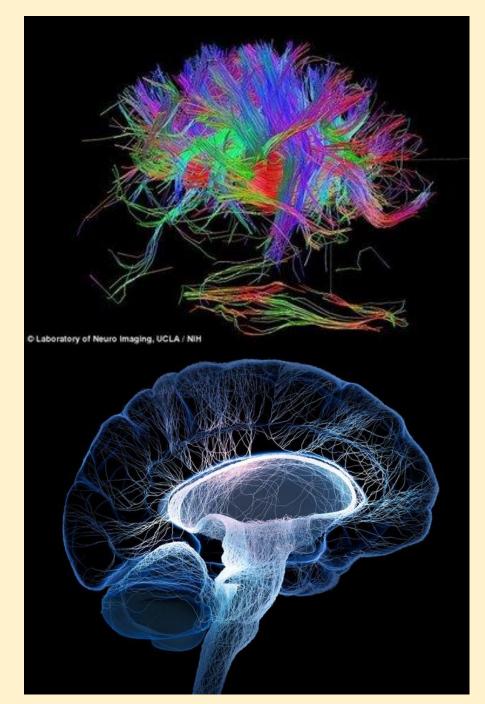
Family Physician & Addictionologist, Valley Medical Group Medical Director, Franklin County House of Corrections Medical Director, Franklin Recovery Center Medical Director, Pioneer Valley Regional School District Chair, Department of Medicine, Baystate – Franklin Medical Center Co-Chair – Healthcare Solutions of the Opioid Taskforce

> Thanks to Tess Jurgensen and the OTF of Franklin County and North Quabbin for organizing these forums



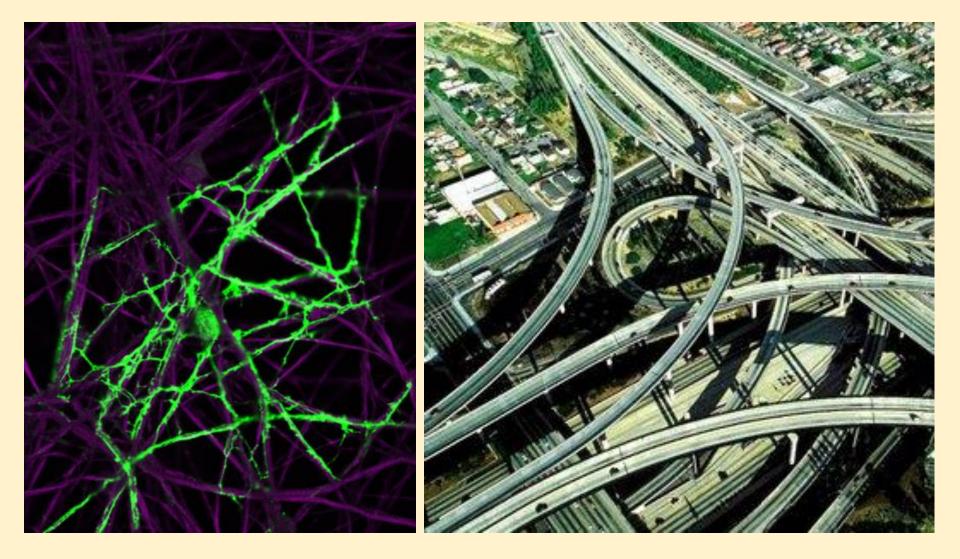
## **Brain Development**







## Synaptic Refinement



# Myelination

Hypersensitive to Social Exclusion

More Risky

Impulsive

Behavior

### This is Normal Development

Preference for Physical Activity and Sensation Seeking



Emotions Felt Very Intensely

Less than Optimal Planning Less Consideration of Negative Consequences Strongly Influenced by Friends and Peers "I'M NOT WHO YOU THINK I AM; I'M NOT WHO I THINK I AM. I AM WHO I THINK I AM. YOU THINK I AM.

The Looking Glass Self Adolescents are more likely to compare themselves to others, feel that they are being compared or judged and place higher value on that judgement

### **HIJACKING THE BRAIN**

New research suggests that the brain's reward system has different mechanisms for craving and pleasure. Craving is driven by the neurotransmitter dopamine. Pleasure is stimulated by other neurotransmitters in "hedonic hot spots." When the craving circuitry overwhelms the pleasure hot spots, addiction occurs, leading people to pursue a behavior or drug despite the consequences.

#### PATHWAYS TO CRAVING

Desire is triggered when dopamine, which originates near the top of the brain stem, travels through neural pathways to act on the brain. Drugs increase the flow of dopamine.

#### Ventral tegmental area (VTA)

Dopamine is produced here and flows outward along neurons distributed throughout the brain's reward system.

#### September 2017

NATIONAL GEOGRAPHIC

ADDICTION

Basic visceral sensations and reactions to pleasure, such as smiling, originate from this hot spot.

**Brain stem** 

#### Animal experiments show that damaging this hot spot can turn something that once gave pleasure into a source of disgust.

#### **Dorsal striatum**

Neurons here help form habits by identifying enjoyable patterns, such as the anticipation of buying drugs.

#### **Prefrontal cortex**

The amino acid glutamate, produced here, interacts with dopamine to spark visualizations that cue cravings.

#### Amygdal

Neurons here are stimulated by learned emotional responses, such as memories of cravings and pleasure.

#### Orbitofrontal cortex

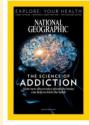
This hot spot gives a sense of gratification but is also the first to shut down if a person has indulged too much.

#### Nucleus accumbens

A hot spot within this key part of the craving circuitry amplifies the response to pleasure.

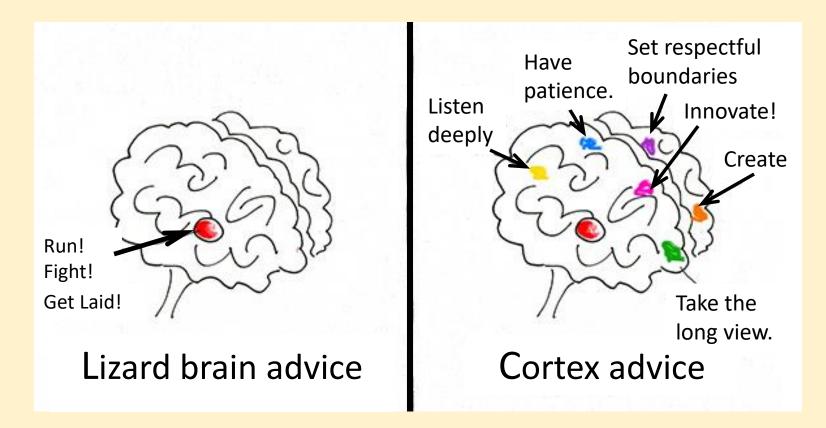
#### PLEASURE HOT SPOTS

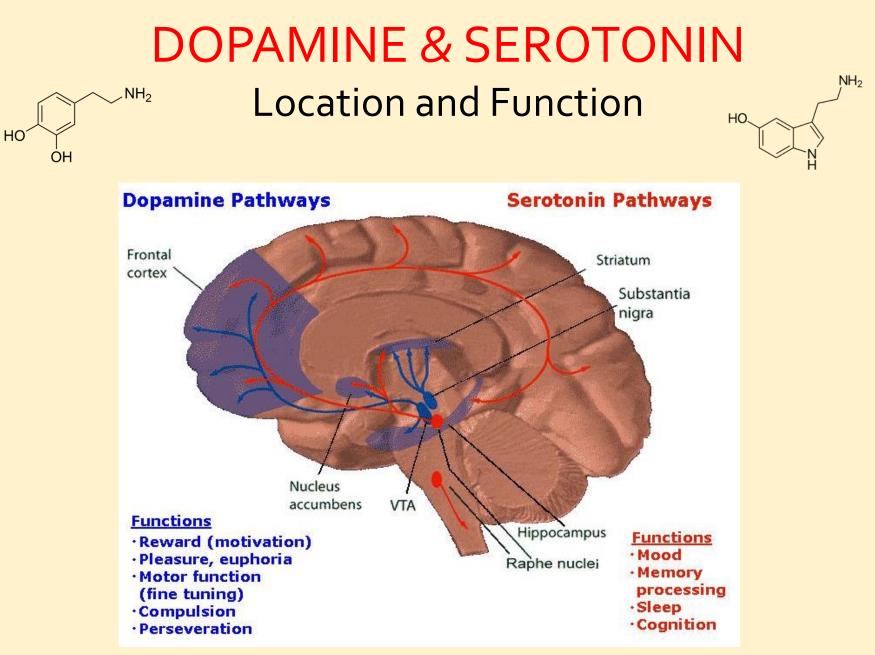
A system of small hedonic hot spots, unrelated to dopamine, provides temporary sensations of pleasure and forms a feedback loop with the reward system that controls desire.



September 2017

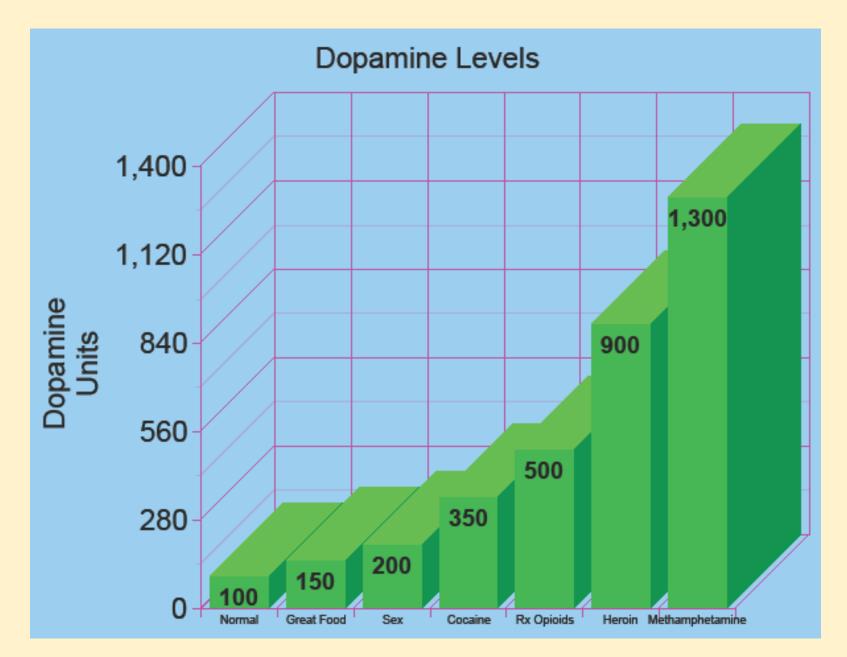
## EAT, DRINK, HAVE SEX (AND USE DRUGS)



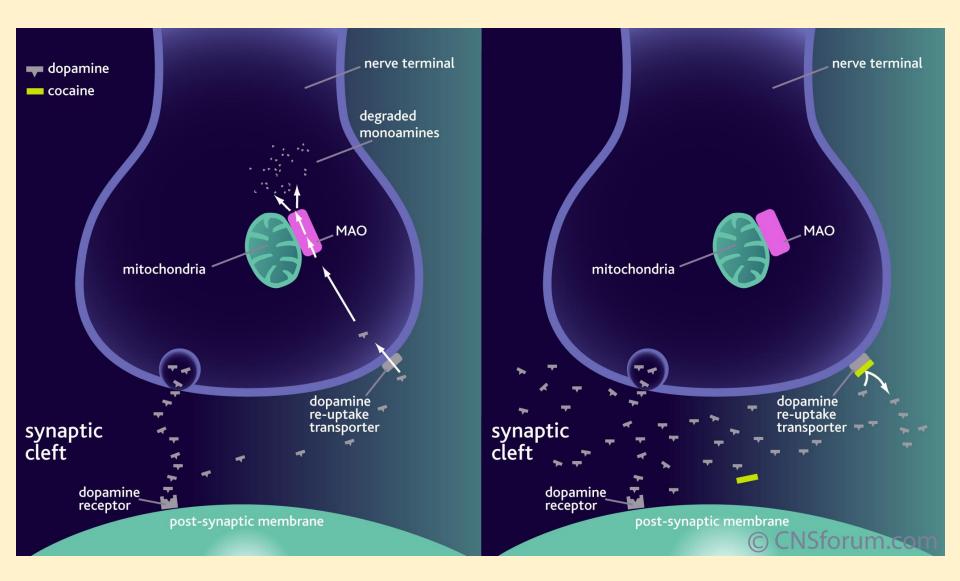




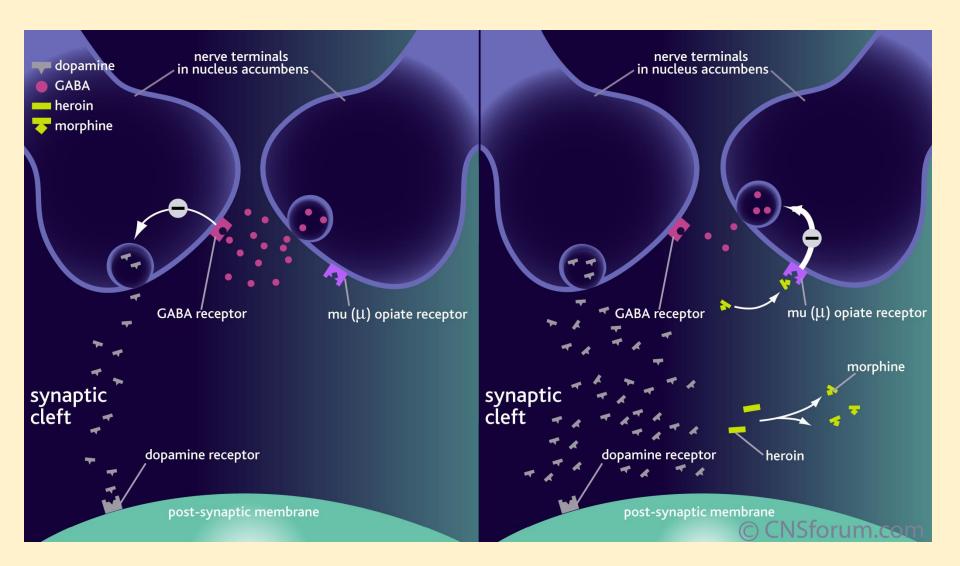


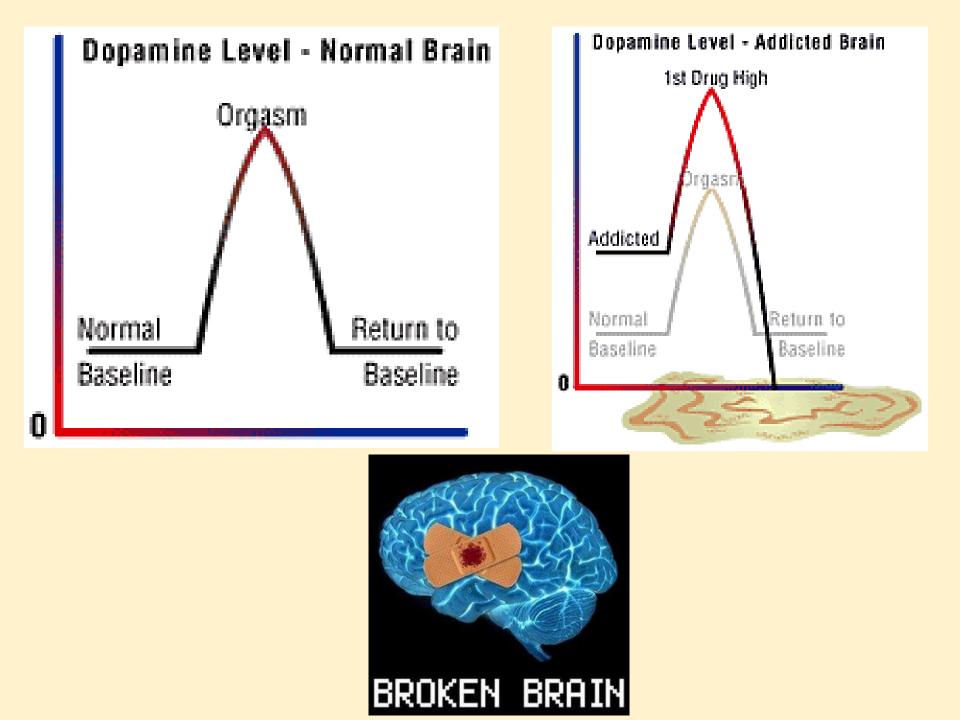


# How Cocaine Works



# How Heroin Works





## Functionally...

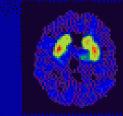
### **Dopamine D2 Receptors are Decreased by Addiction**

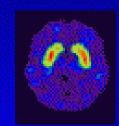


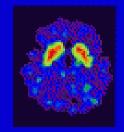




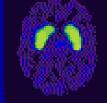


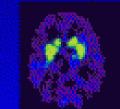


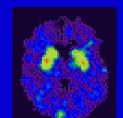






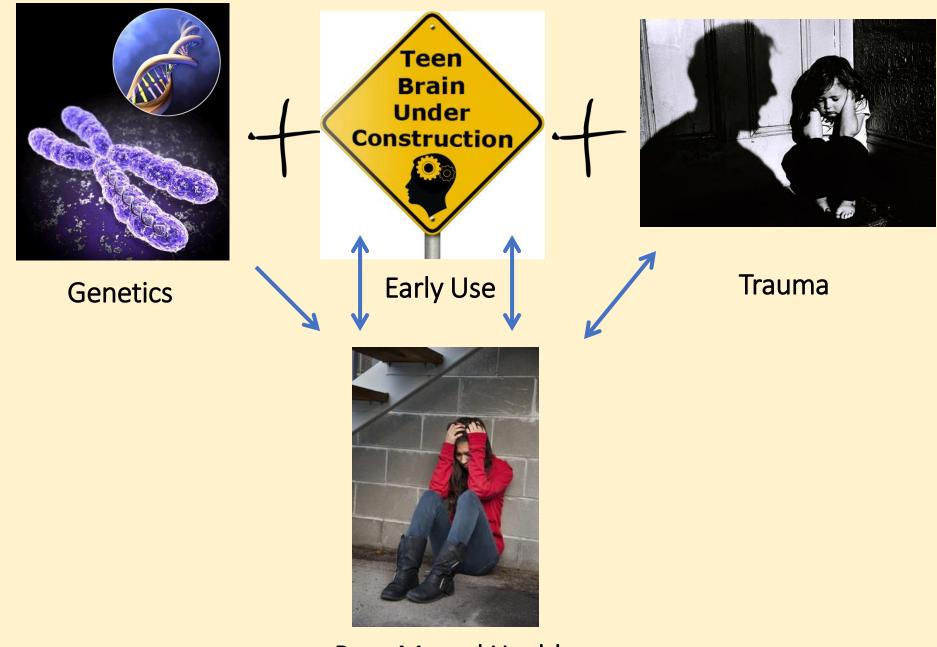






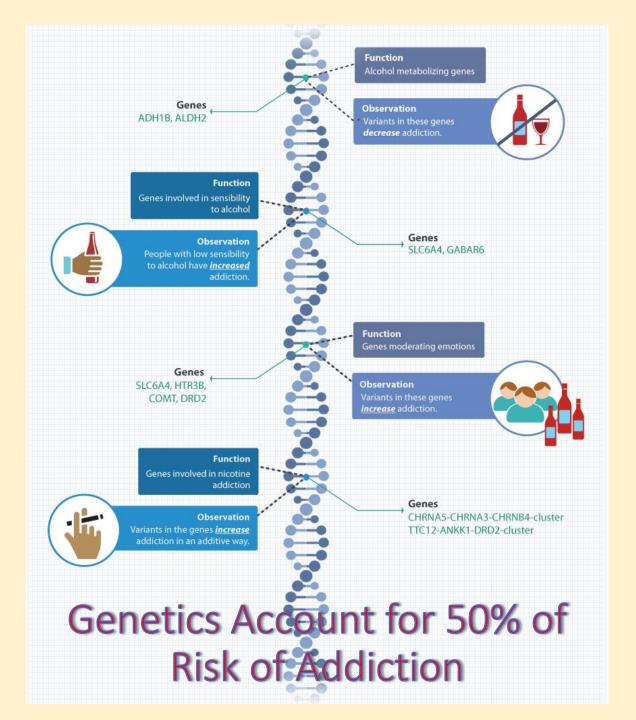






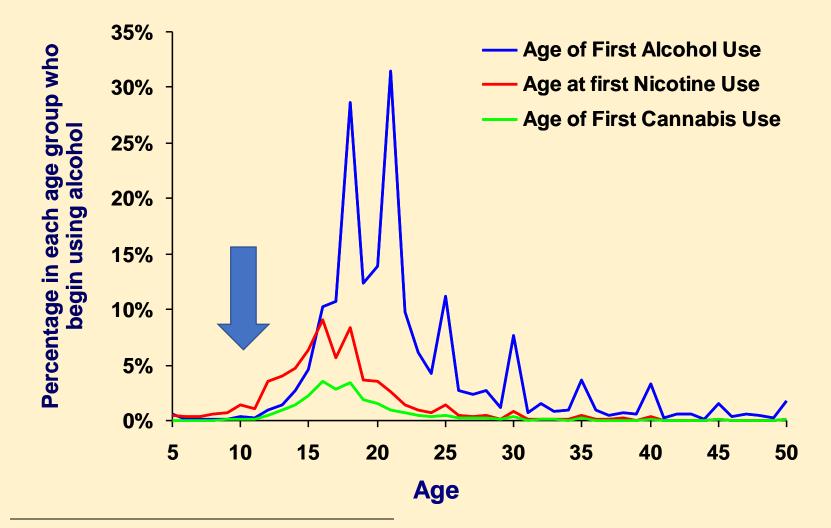
**Poor Mental Health** 





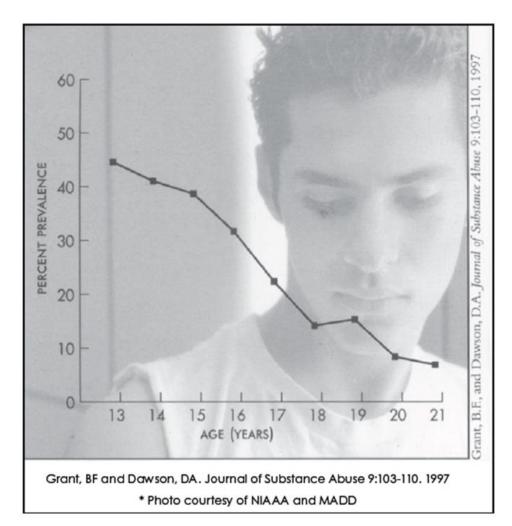


## Addiction is a Developmental Pediatric Disease



Source: NIAAA National Epidemiologic Survey on Alcohol and Related Conditions, 2003

# **Teen Alcohol Use Wires The Brain For Addiction**

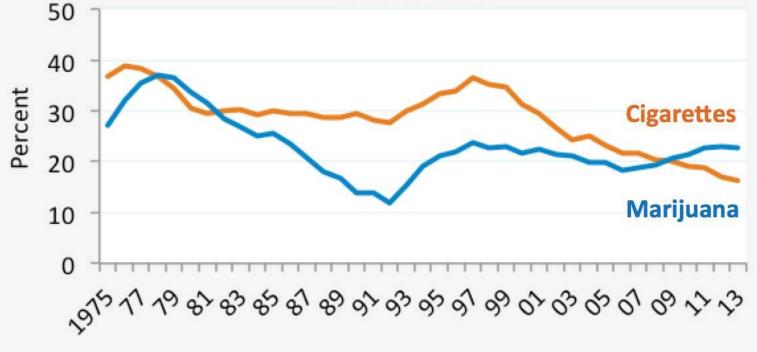


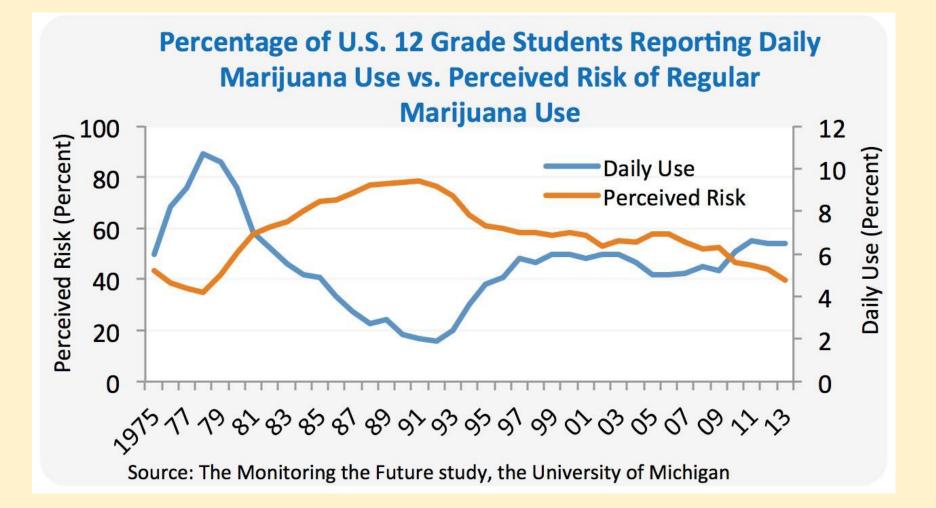
40% of kids who begin drinking at age 15 will become alcoholics.

Only 7% of those who begin drinking at age 21 become alcoholics.

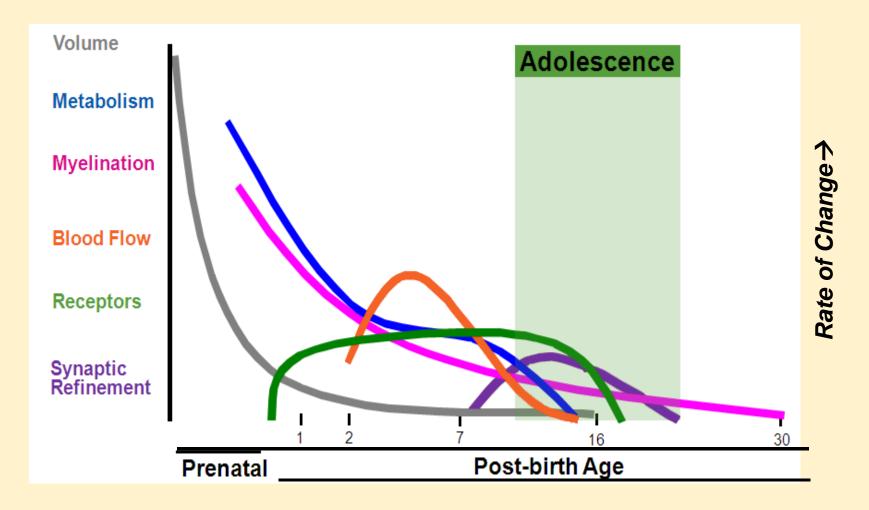


### Percentage of U.S. 12th Grade Students Reporting Past Month Use of Cigarettes and Marijuana, 1975 to 2013

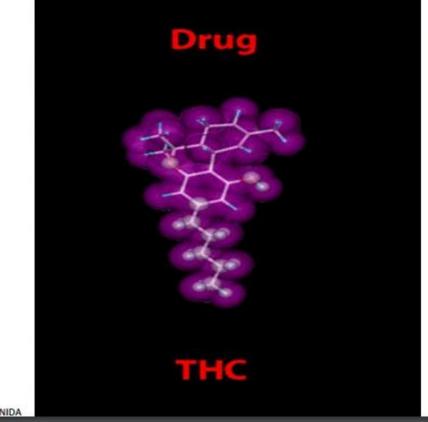




## **Brain Development**

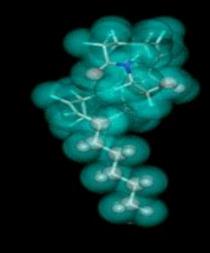


## Marijuana (Tetrahydrocannabinol



## Endo-cannabinoid (Anandamide)

### **Brain's Chemical**



### Anandamide

Source: NIDA





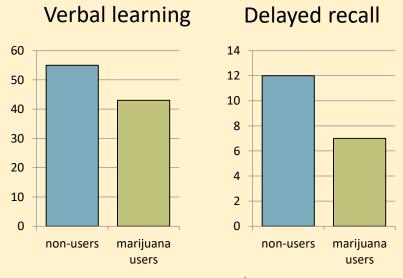


## Deficits in Cognitive Functioning Among Active Marijuana Users

Many studies show that adolescents who use marijuana heavily tend to score worse than non-users on tests of:

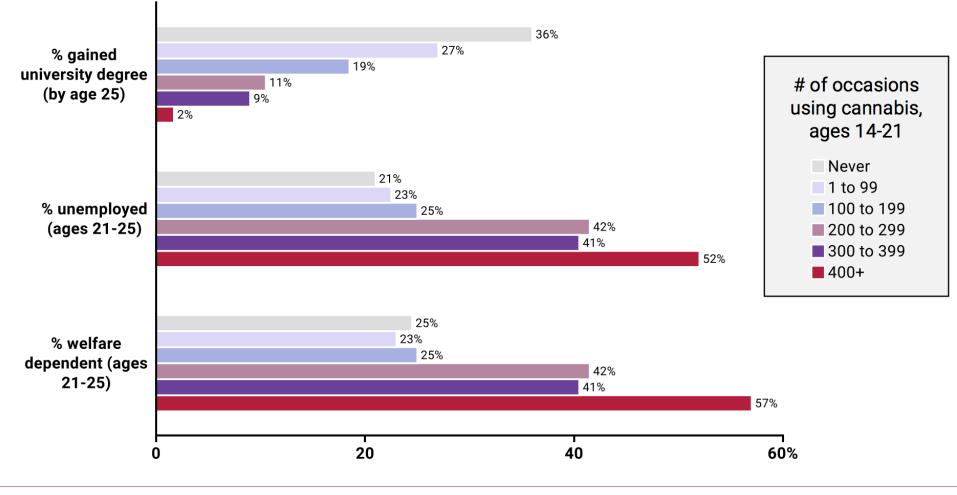
- attention
- verbal learning
- memory
- processing speed

... even when they are not high.



Messinis, et al 2006

### New Zealand study showing relationship between cannabis use and social outcomes





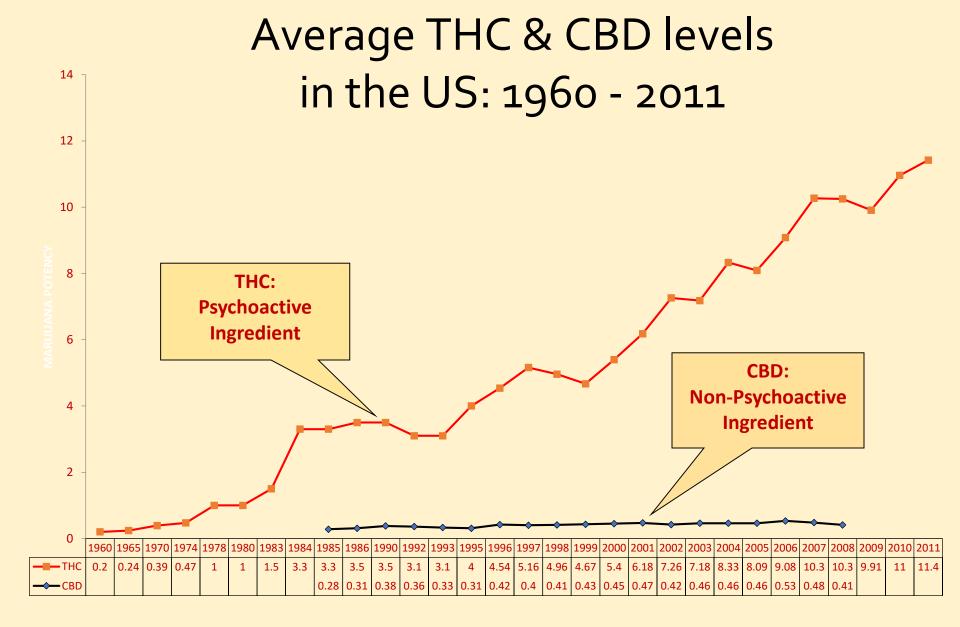
Source: Fergusson and Boden. Addiction, 103, pp. 969-976, 2008 [New Zealand study]

Loss of Adult IQ with Marijuana Dependence in Adolescence

Findings:

- Those who developed marijuana dependence before age 18 showed IQ decline in adulthood.
- The longer their dependence persisted, the greater the decline, with a decline of 8 IQ points for the most persistent users.
- Those who began using in adulthood did not show IQ decline.
- Quitting in adulthood did not restore functioning in those who began in adolescence.

Persistent cannabis users show neuropsychological decline from childhood to midlife Madeline H. Meier, Avshalom Caspi, Antony Ambler, HonaLee Harrington, Renate Houts, Richard S. E. Keefe, Kay McDonald, Aimee Ward, Richie Poulton, and Terrie E. Moffitt PNAS October 2, 2012. 109 (40) E2657-E2664



Data from the NIDA-sponsored Potency Monitoring program at the University of Mississippi, showing average THC and CBD levels in samples of marijuana seized by federal, state and local governments in each year shown.

## **THC Concentrates**



"Green Crack" wax



"Ear Wax"



### Butane Hash Oil (BHO)



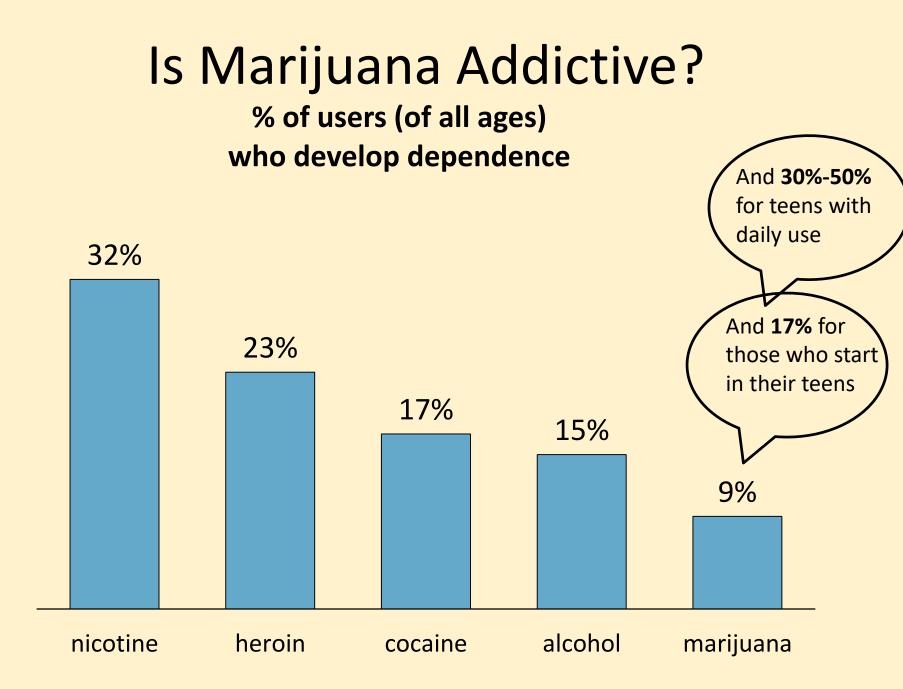
Hash Oil Capsules











#### TEENS MORE LIKELY TO USE MARIJUANA THAN CIGARETTES

### Daily use among 12th graders



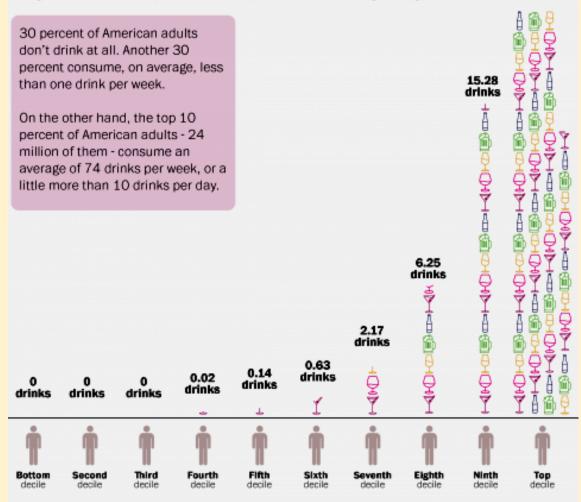


DRUGABUSE.GOV

## Think you drink a lot?

#### Time for a stiff drink

Average number of drinks per capita consumed in the past week, by decile, among adults aged 18 and over.



WASHINGTONPOST.COM/WONKBLOG

Source: "Paying the Tab," by Philip J. Cook

73.85

drinks

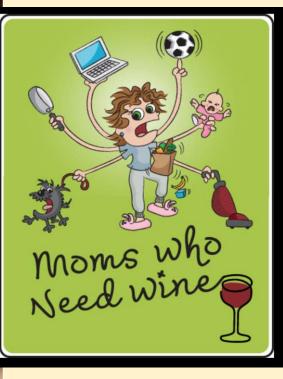














# Association of parental supply of alcohol with adolescent drinking, alcohol-related harms, and alcohol use disorder symptoms: a prospective cohort study

Richard P Mattick, Philip J Clare, Alexandra Aiken, Monika Wadolowski, Delyse Hutchinson, Jackob Najman, Tim Slade, Raimondo Bruno, Nyanda McBride, Kypros Kypri, Laura Vogl, Louisa Degenhardt

Interpretation Providing alcohol to children is associated with alcohol-related harms. There is no evidence to support the view that parental supply protects from adverse drinking outcomes by providing alcohol to their child. Parents should be advised that this practice is associated with risk, both directly and indirectly through increased access to alcohol from other sources.

### Countries by their Alcohol Consumption

3

Rank	Country	Consumption Per Capita in Liters	
1	Belarus	17.5	
2	Moldova	16.8	
3	Lithuania	15.4	
4	Russian Federation	15.1	
5	Romania	14.4	
6	Ukraine	13.9	
7	Andorra	13.8	
8	Hungary	13.3	
9	Czech Republic	13	
10	Slovakia	13	
11	Portugal	12.9	
12	Serbia	12.8	
13	South Korea	12.3	
14	Finland	12.3	
15	Latvia	12.3	
16	Croatia	12.2	
17	France	12.2	
18	Australia	12.2	
19	Luxembourg	11.9	
20	Ireland	11.9	
21	Germany	11.8	
22	Denmark	11.4	
23	Bulgaria	11.4	
24	Belgium	11	
25	South Africa	11	



## Adverse Childhood Experiences – ACEs

Emotional Neglect Abuse Household Dysfunction

### Relationship of Childhood Abuse and Household Dysfunction to Many of the Leading Causes of Death in Adults

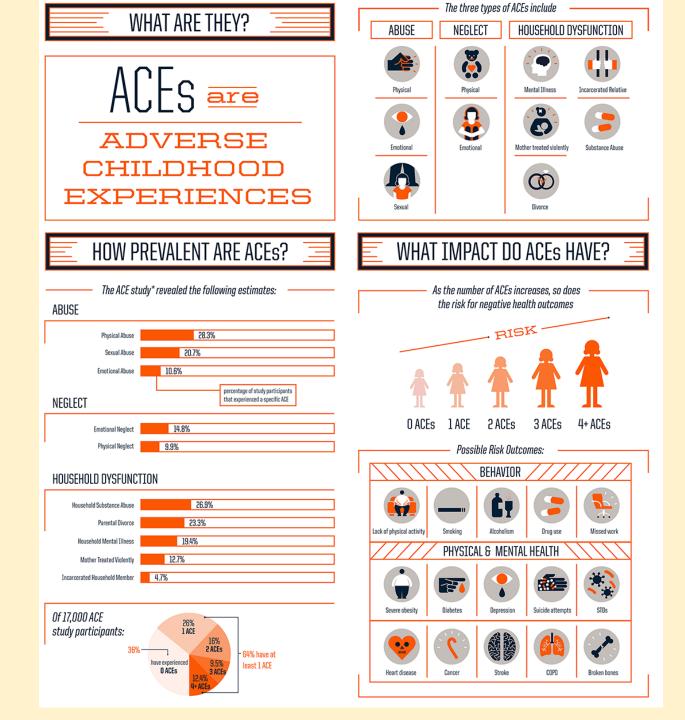
#### The Adverse Childhood Experiences (ACE) Study

Vincent J. Felitti, MD, FACP, Robert F. Anda, MD, MS, Dale Nordenberg, MD, David F. Williamson, MS, PhD, Alison M. Spitz, MS, MPH, Valerie Edwards, BA, Mary P. Koss, PhD, James S. Marks, MD, MPH

- **Background:** The relationship of health risk behavior and disease in adulthood to the breadth of exposure to childhood emotional, physical, or sexual abuse, and household dysfunction during childhood has not previously been described.
- Methods: A questionnaire about adverse childhood experiences was mailed to 13,494 adults who had completed a standardized medical evaluation at a large HMO; 9,508 (70.5%) responded. Seven categories of adverse childhood experiences were studied: psychological, physical, or sexual abuse; violence against mother; or living with household members who were substance abusers, mentally ill or suicidal, or ever imprisoned. The number of categories of these adverse childhood experiences was then compared to measures of adult risk behavior, health status, and disease. Logistic regression was used to adjust for effects of demographic factors on the association between the cumulative number of categories of childhood exposures (range: 0–7) and risk factors for the leading causes of death in adult life.
- **Results:** More than half of respondents reported at least one, and one-fourth reported  $\geq 2$  categories of childhood exposures. We found a graded relationship between the number of categories of childhood exposure and each of the adult health risk behaviors and diseases that were studied (P < .001). Persons who had experienced four or more categories of childhood exposure, compared to those who had experienced none, had 4-to 12-fold increased health risks for alcoholism, drug abuse, depression, and suicide attempt; a 2-to 4-fold increase in smoking, poor self-rated health,  $\geq 50$  sexual intercourse partners, and sexually transmitted disease; and a 1.4-to 16-fold increase in physical inactivity and severe obesity. The number of categories of adverse childhood exposures showed a graded relationship to the presence of adult diseases including ischemic heart disease, cancer, chronic lung disease, skeletal fractures, and liver disease. The seven categories of adverse childhood exposure were likely to have multiple health risk factors later in life.
- **Conclusions:** We found a strong graded relationship between the breadth of exposure to abuse or household dysfunction during childhood and multiple risk factors for several of the leading causes of death in adults.

Medical Subject Headings (MeSH): child abuse, sexual, domestic violence, spouse abuse, children of impaired parents, substance abuse, alcoholism, smoking, obesity, physical activity, depression, suicide, sexual behavior, sexually transmitted diseases, chronic obstructive pulmonary disease, ischemic heart disease. (Am J Prev Med 1998;14:245–258) © 1998 American Journal of Preventive Medicine

	Findin	g Your ACE Score	
While you were grow	ving up, during your fir	st 18 years of life:	
	er adult in the household insult you, put you down		
	at made you afraid that y Yes No	you might be physically hurt? If yes enter 1	
Push, grab, sla	er adult in the household ap, or throw something a	often or very often t you?	
Ever hit you so	o hard that you had mark Yes No	s or were injured? If yes enter 1	
Touch or fondle	on at least 5 years older e you or have you touch or	than you <b>ever</b> their body in a sexual way?	
Attempt or actu	**	vaginal intercourse with you? If yes enter 1	
4. Did you often or ve No one in your		ight you were important or special?	
	In't look out for each othe Yes No	er, feel close to each other, or support each other? If yes enter 1	
5. Did you <b>often or ve</b> You didn't have	ery often feel that e enough to eat, had to v or	wear dirty clothes, and had no one to protect you?	
Your parents w it?	vere too drunk or high to	take care of you or take you to the doctor if you need	led
	Yes No	If yes enter 1	
	ever separated or divord Yes No	ed? If yes enter 1	
		slapped, or had something thrown at her?	
Sometimes, o	often, or very often kicke	ed, bitten, hit with a fist, or hit with something hard?	
	ly hit at least a few minu Yes No	tes or threatened with a gun or knife? If yes enter 1	
	yone who was a problen Yes No	n drinker or alcoholic or who used street drugs? If yes enter 1	
	nember depressed or me Yes No	entally ill, or did a household member attempt suicide? If yes enter 1	?
10. Did a household m	nember go to prison? Yes No	If yes enter 1	
Now add up y	your "Yes" answers:	This is your ACE Score.	



# Talk Early, Talk Often

# "Delay, Delay, Delay or Avoid Altogether"



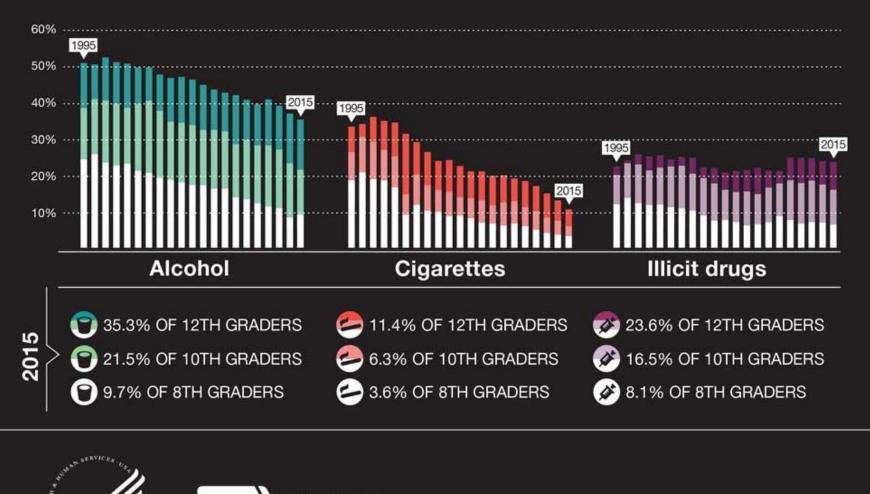
Genetics

Trauma

Early Use

### LAST TWO DECADES OF ALCOHOL, CIGARETTE, AND ILLICIT DRUG USE\*

\*Past-month use



National Institute

on Drug Abuse



## WWW.DRUGABUSE.GOV

# TOBACCO PRODUCT USE BY HIGH SCHOOL STUDENTS (2011-2016)

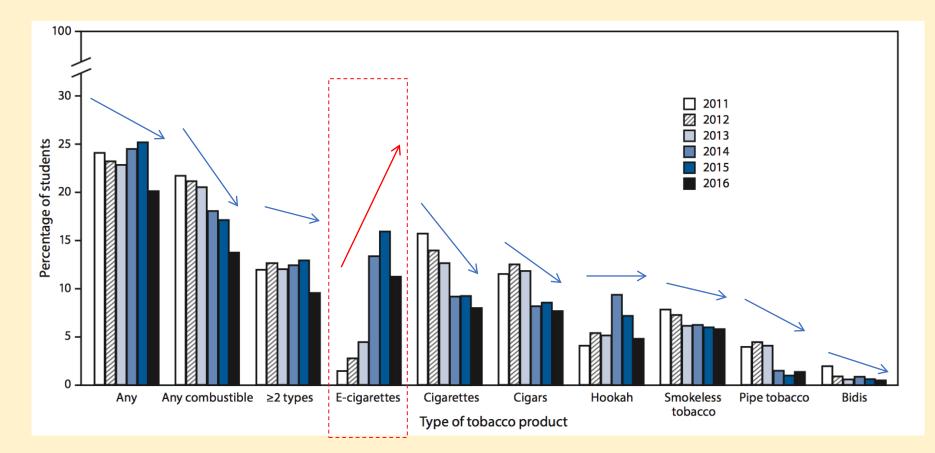
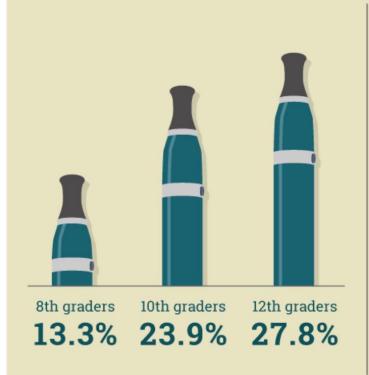
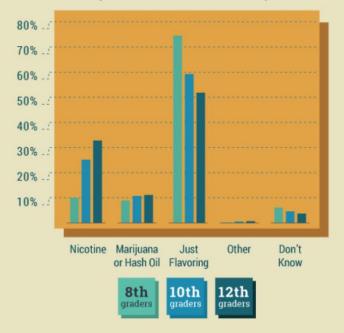


Figure 1. Estimated percentage of high school students who currently use any tobacco products, any combustible tobacco products, ≥2 tobacco products, and selected tobacco products – National Youth Tobacco Survey, United States. 2011-2016.

### PAST-YEAR E-VAPORIZER USE AND WHAT TEENS ARE INHALING



When asked what they thought was in the e-vaporizer mist students inhaled the last time they smoked, these were their responses:



NEARLY 1 IN 3 STUDENTS IN 12TH GRADE REPORT PAST-YEAR USE OF E-VAPORIZERS, RAISING CONCERNS ABOUT THE IMPACT ON THEIR LONG-TERM HEALTH.





DRUGABUSE.GOV

The Nicotine Content of a Sample of E-cigarette Liquid Manufactured in the United States				
Raymond, Barrett H. BSN, RN, FNP-S, NREMT-P; Collette-Merrill, Katreena PhD, RN; Harrison, Roger G. PhD; Jarvis, Sabrina DNP, FNP-BC, ACNP-BC, FAANP; Rasmussen, Ryan Jay MS, RN, FNP				
Journal of Addiction Medicine: December 26, 2017 - Volume Publish Ahead of Print - Issue - p doi: 10.1097/ADM.000000000000376 Original Research: PDF Only				
BUY PAP				
Abstract Author Information				

SD = 3.3) nicotine. The labeled 18 mg/mL samples measured as little as 35% less nicotine and as much as 52% greater nicotine. In the 35 samples labeled 0 mg/mL, nicotine was detected (>0.01 mg/mL) in 91.4% of the samples (range 0–23.9 mg/mL; M = 2.9, SD = 7.2). Six samples from 2 manufacturers labeled as 0 mg/mL were found to contain nicotine in amounts ranging from 5.7 to 23.9 mg/mL.

popular flavors from each manufacturer were purchased in nicotine concentrations of 0 and 18 mg/mL. Of the samples purchased (n = 70), all were labeled as produced in the United States of America. The researchers anonymized the samples before sending them to an independent university laboratory for testing.

**Results:** The 35 e-liquid samples labeled 18 mg/mL nicotine measured between 11.6 and 27.4 mg/mL (M = 18.7, SD = 3.3) nicotine. The labeled 18 mg/mL samples measured as little as 35% less nicotine and as much as 52% greater nicotine. In the 35 samples labeled 0 mg/mL, nicotine was detected (>0.01 mg/mL) in 91.4% of the samples (range 0–23.9 mg/mL; M = 2.9, SD = 7.2). Six samples from 2 manufacturers labeled as 0 mg/mL were found to contain nicotine in amounts ranging from 5.7 to 23.9 mg/mL.

**Conclusion:** This study demonstrates the nicotine labeling inaccuracies present in current e-liquid solutions produced in the United States. Incorrect labeling poses a significant risk to consumers and supports the recent regulation changes enacted by the US Food and Drug Administration. Additional routine testing of nicotine concentrations should be conducted to evaluate the effectiveness of the regulations on future e-liquid production.

### One Mad Hit Juice Box

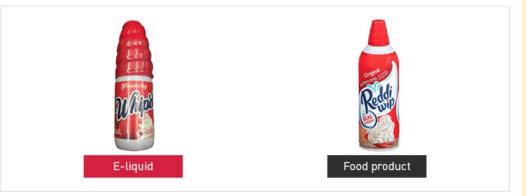
• NEwhere Inc. d/b/a Mad Hatter Juice





### Whip'd Strawberry

- Cosmic Fog Vapors d/b/a Next Day Vapor Products
- Elite Vaporworks



### V'Nilla Cookies & Milk

- Tinted Brew, Inc.
- Fog It Up





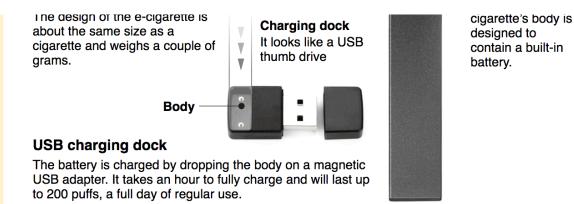
# JUUL e-cigarettes

Some e-cigarette critics are calling the JUUL e-cigarette the "Apple of vaping" or the "iPhone of vaping." With its relatively low entry price, sleek portable design, ease of use, and nicotine head rush it generates for users, JUULS are catching on with younger people.

JUUL flavor pods



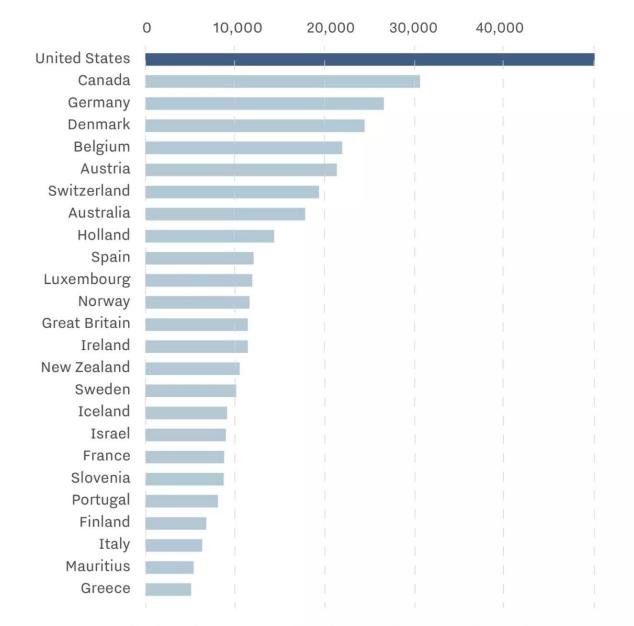
\*Contains one JUULpod each of Virginia Tobacco, Cool Mint, Fruit Medley and Creme Brulee. Each JUULpod contains 0.7mL with 5% nicotine by weight, approximately equivalent to 1 pack of cigarettes or 200 puffs.



UNE 15, 201

They're the most powerful painkillers ever invented. And they're creating the worst addiction crisis America has ever seen.

### Standard daily opioid dose for every 1 million people



Source: United Nations International Narcotics Control Board Credit: Sarah Frostenson

Vex

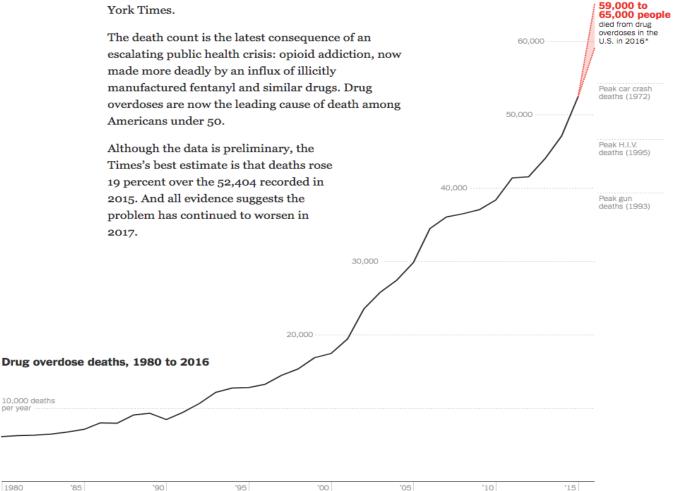


# Drug Deaths in America Are **Rising Faster Than Ever**

#### By JOSH KATZ JUNE 5, 2017

New data compiled from hundreds of health agencies reveals the extent of the drug overdose epidemic last year.

AKRON, Ohio - Drug overdose deaths in 2016 most likely exceeded 59,000, the largest annual jump ever recorded in the United States, according to preliminary data compiled by The New York Times.

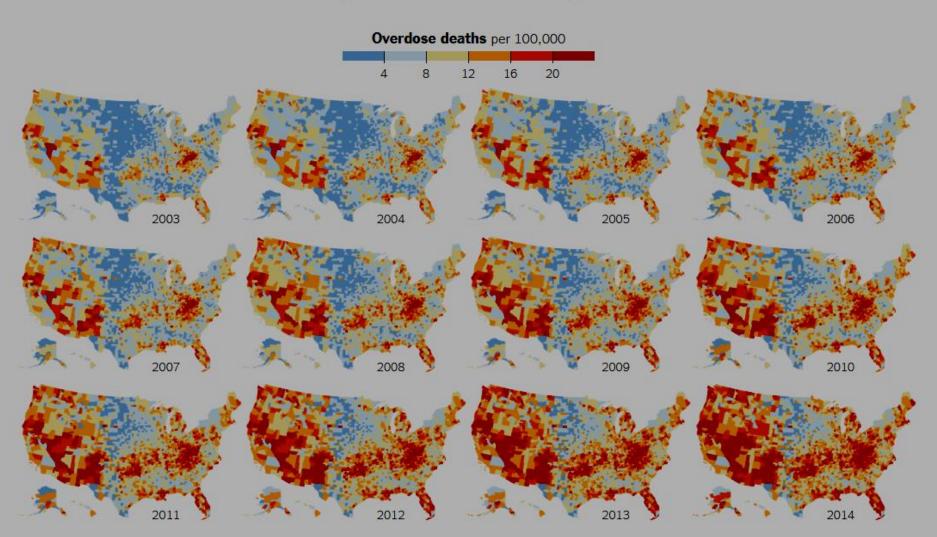


\*Estimate based on preliminary data

1980

# **Deaths Ripples Across America**

By HAEYOUN PARK and MATTHEW BLOCH JAN. 19, 2016



NYT January 19, 2016







### Florida Shutting 'Pill Mill' Clinics



At an Aug. 15 news conference, Florida officials displayed prescription drugs that were surrendered to the state by pain clinics.

By LIZETTE ALVAREZ Published: August 31, 2011

WEST PALM BEACH, Fla. - Florida has long been the nation's center of the illegal sale of prescription drugs: Doctors here bought 89 percent of all the Oxycodone sold in the country last year. At its peak, so many out-of-staters flocked to Florida to buy drugs at more than 1,000 pain clinics that the state earned the nickname "Oxy Express."

#### Related

Times Topics: OxyContin (Drug) Prescription Drug Abuse

R Enlarge This Image



Florida Gov. Rick Scott attends a

But with the help of tougher laws, officials have moved aggressively this year to shut down so-called pill mills and disrupt the pipeline that moves the drugs north. In the past year, more than 400 clinics were either shut down or closed their doors.

Prosecutors have indicted dozens of pill mill operators, and nearly 80 doctors have seen their licenses suspended for prescribing mass quantities of pills without clear medical need.

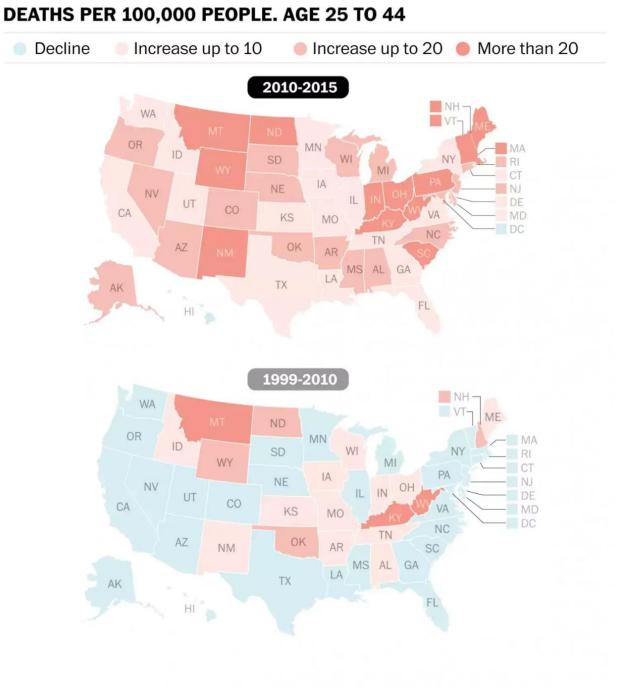
RECOMMEND TWITTER INKEDIN COMMENTS (58) E-MAIL B PRINT SINGLE PAGE REPRINTS + SHARE ACADEMY AWARD® WINNER ANTHONY HOPKINS

70



A cap, containing what looks like a dusting of heroin, sells for \$10.



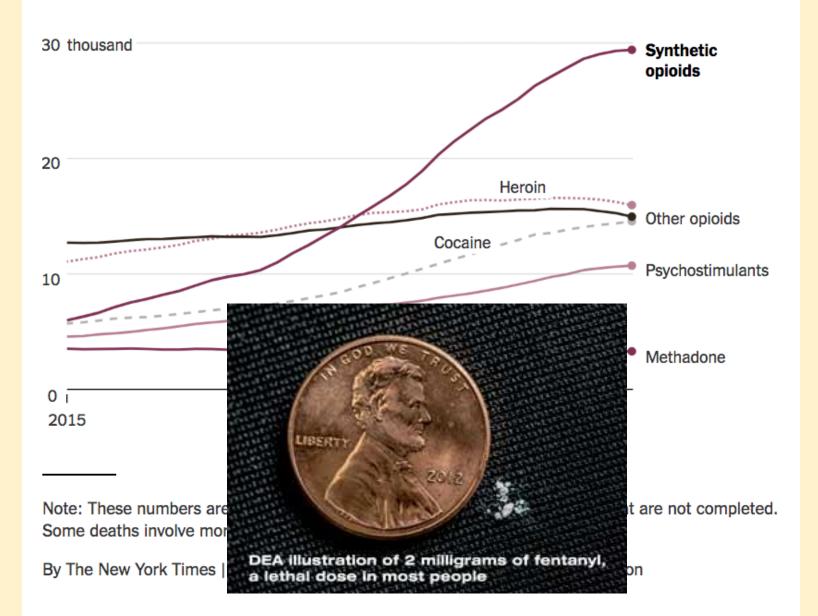


Source: Centers for Disease Control and Prevention

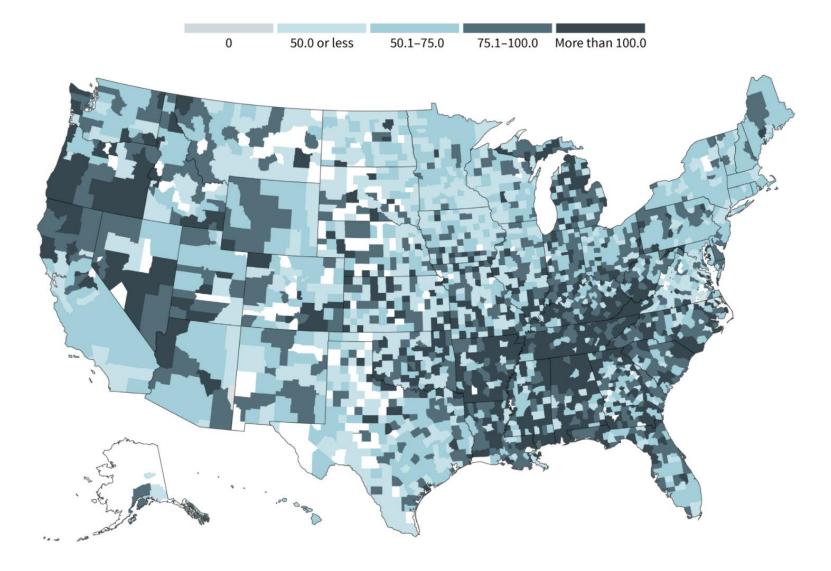
THE WASHINGTON POST

## Synthetic Opioids Are Driving Up the Overdose Rate

Overdose deaths in thousands in preceding 12 months



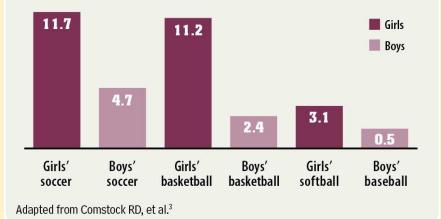
The map below shows the amount of retail opioid prescriptions dispensed per 100 people in 2016.





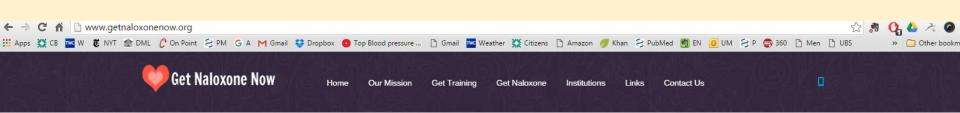
## **FIGURE 2** High school ACL injury rates

High school anterior cruciate ligament (ACL) injury rates per 100,000 athlete exposures (AEs) by sport and sex. (An athlete exposure is 1 athlete participating in 1 practice or competition.)











#### INDIVIDUALS-GET TRAINED!

In as little as 20 minutes, you can learn techniques to save the life of a loved-one, friend, co-worker, neighbor ... anyone ... who is experiencing an overdose caused by prescription narcotics or heroin. You can also find out where you can obtain the lifesaving, easy-to-use antidate palaxone as well as access relevant

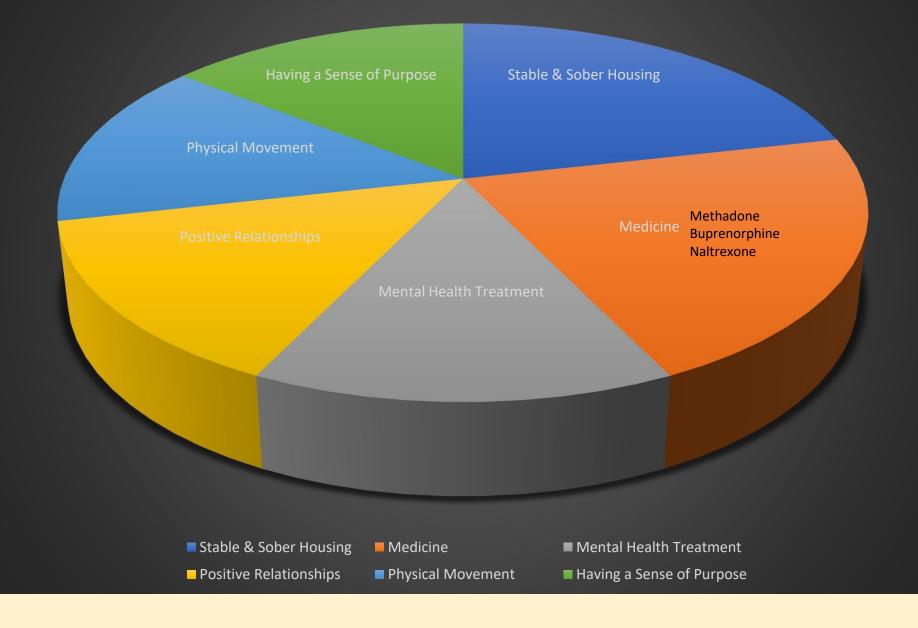
#### INSTITUTIONAL PROGRAM

Central to our mission to save lives is our institutional program. Targeted online training is available to professional first responders (police officers, firefighters, EMTs) through use of our first responder computerbased module.

#### HOW CAN YOU HELP

Our computer-based training modules can help you give someone a second chance, and spare those who love them the pain of a preventable loss of life or brain injury. The online training is provided free of charge. If you wish to obtain CE or CASAC credits (pending accreditation) after completing the module, a nominal

## What It Takes To Get Better



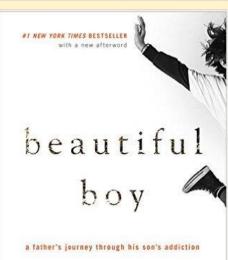
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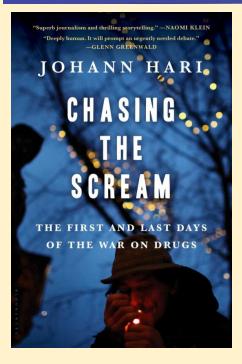
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