### The New Neurobiology of Addiction

Petros Levounis, MD, MA

Professor and Chair, Department of Psychiatry Rutgers New Jersey Medical School

Community Opioid Symposium

Woman's Foundation & Acadiana Area Human Services District

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#### **Disclosures**

I have no competing interests in this presentation.

#### **Learning Objectives**

- 1. Identify the origins and current state of the opioid crisis in America.
- 2. Describe the basic elements of drive and cognition in the neurobiology of addiction.
- 3. Discuss the clinical use of pharmacotherapy and psychosocial interventions in the treatment of substance use disorders.

#### **Outline**

- 1. The Basic Model
- 2. Neurobiology
- 3. New Neurobiological Concepts
- 4. Treatments
- 5. New Directions
- 6. Conclusions

# The Basic Model

# 

#### A Biopsychosocial Illness



Addiction

Brain Switch

#### The Root Cause of the Disaster

#### ADDICTION RARE IN PATIENTS TREATED WITH NARCOTICS

To the Editor: Recently, we examined our current files to determine the incidence of narcotic addiction in 39,946 hospitalized medical patients¹ who were monitored consecutively. Although there were 11,882 patients who received at least one narcotic preparation, there were only four cases of reasonably well documented addiction in patients who had no history of addiction. The addiction was considered major in only one instance. The drugs implicated were meperidine in two patients,² Percodan in one, and hydromorphone in one. We conclude that despite widespread use of narcotic drugs in hospitals, the development of addiction is rare in medical patients with no history of addiction.

JANE PORTER
HERSHEL JICK, M.D.
Boston Collaborative Drug
Surveillance Program
Boston University Medical Center

Waltham, MA 02154

- 1. Jick H, Miettinen OS, Shapiro S, Lewis GP, Siskind Y, Slone D. Comprehensive drug surveillance. JAMA. 1970; 213:1455-60.
- 2. Miller RR, Jick H. Clinical effects of meperidine in hospitalized medical patients. J Clin Pharmacol. 1978; 18:180-8.

#### **The False Promise**



Extra strength pain relief free of extra prescribing restrictions.

Telephone prescribing in most statesUp to five refills in 6 months

No triplicate Rx required

#### Excellent patient acceptance.

In 12 years of clinical experience, nausea, sedation and constipation have rarely been reported.1

COMPARATIVE PHARMACOLOGY OF TWO ANALGESICS					
	Constipation	Respiratory Depression	Sedation	Emesis	Physical Dependence
HYDROCODONE		X			X
OXYCODONE	XX	XX	XX	XX	XX

1991 and Catalano RB. The medical approach to management of pain caused by cancer. Semin. Oncol. 1975; 2: 379-92 and Router JB, et. al. The chronic pain syndrome: misconceptions and management. Ann. Intern. Med. 1990 589-96.

#### The heritage of VICODIN," over a billion doses prescribed.2

- VICODIN ES provides greater central and peripheral action than other hydrocodone/acetaminophen combinations.
- Four to six hours of extra strength pain relief from a single dose
- The 14th most frequently prescribed medication in America<sup>2</sup>

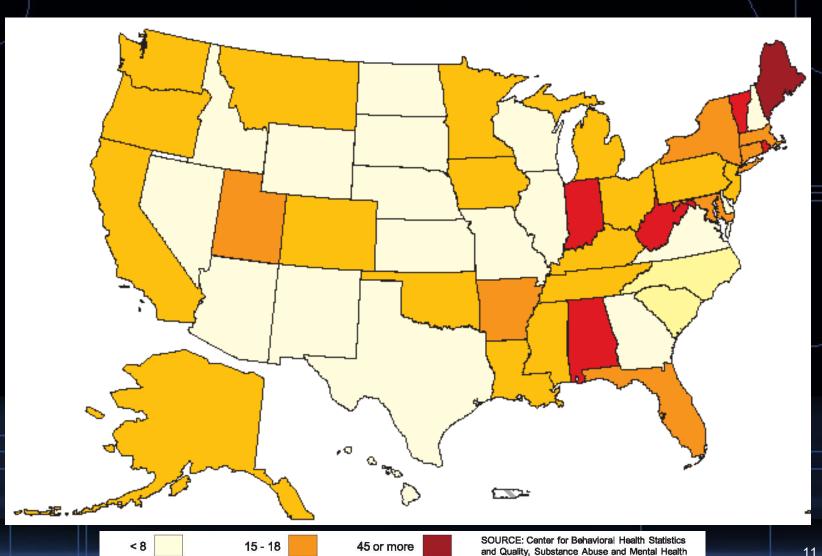


Tablet for tablet, the most potent analgesic you can phone in.

#### Money and Influence



Primary non-heroin opioid admission rates (per 100,000)

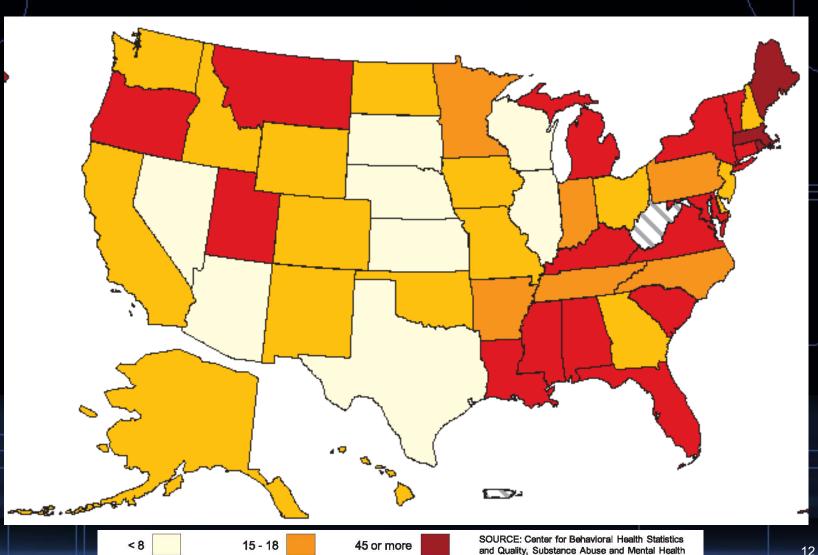


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Services Administration, Treatment Episode Data

Primary non-heroin opioid admission rates (per 100,000)

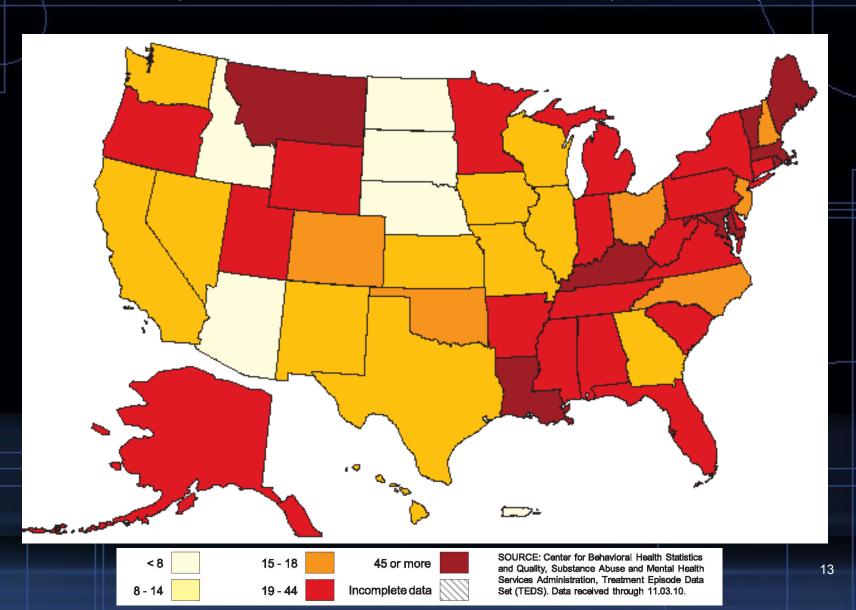


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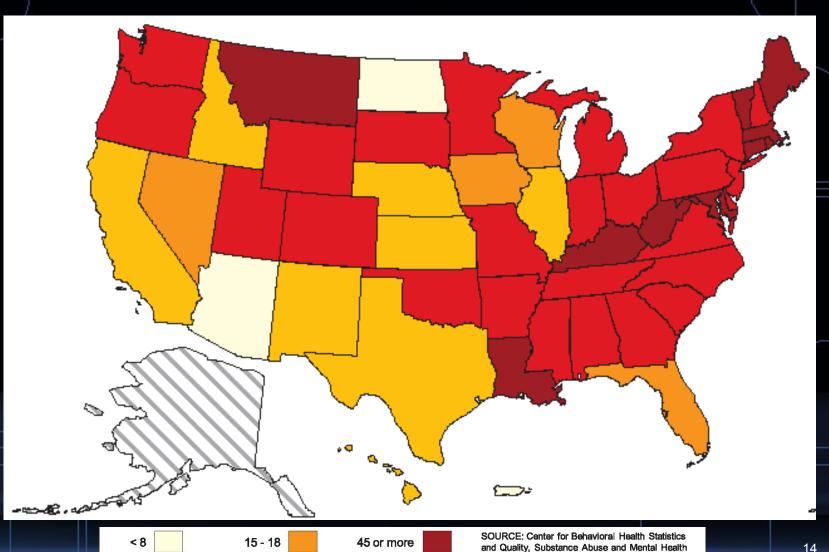
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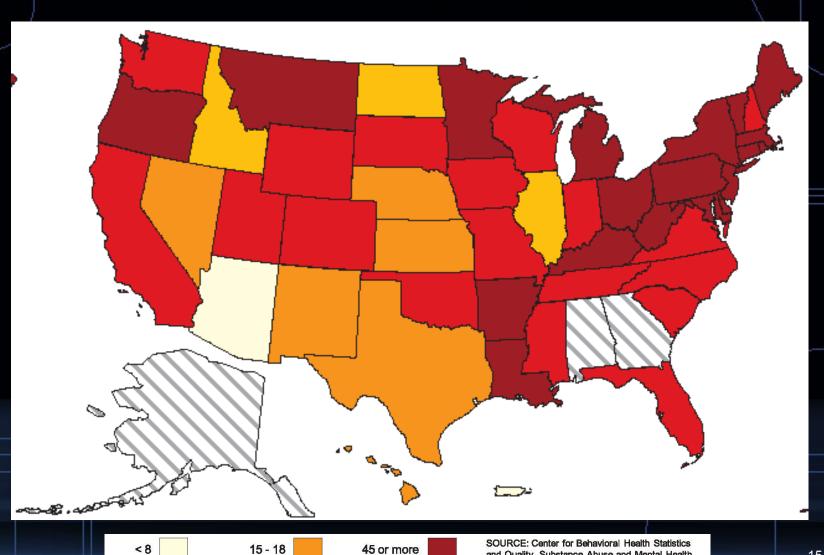


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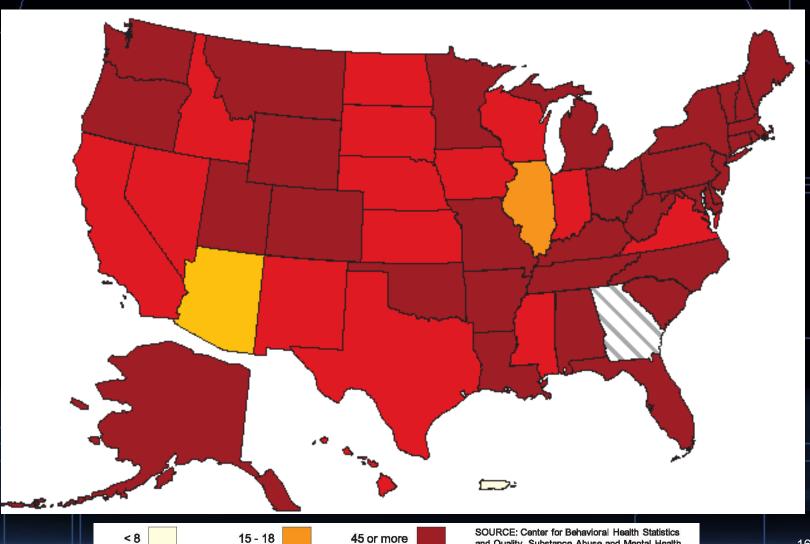


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and Quality, Substance Abuse and Mental Health

Services Administration, Treatment Episode Data

Primary non-heroin opioid admission rates (per 100,000)



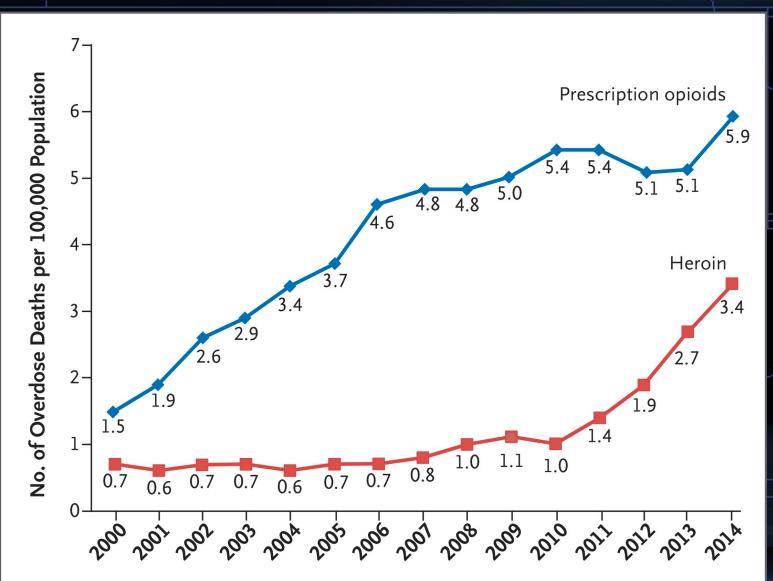
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and Quality, Substance Abuse and Mental Health

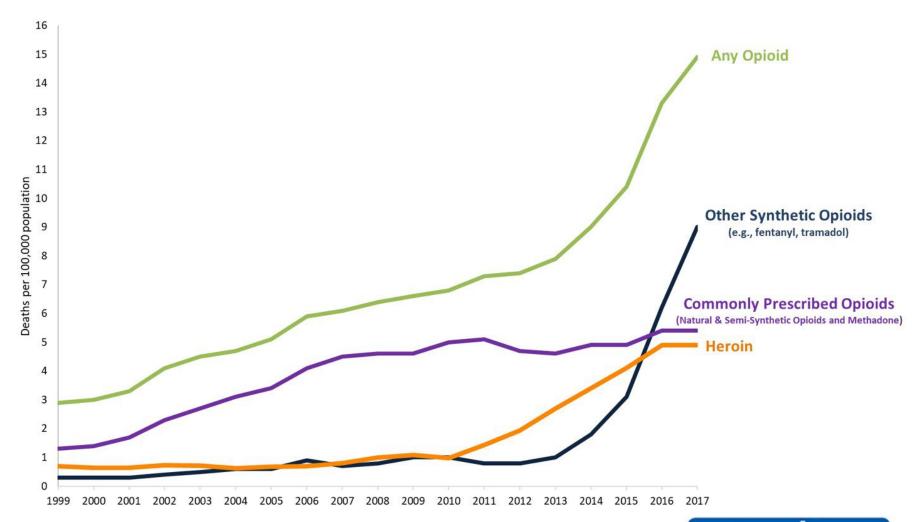
Services Administration, Treatment Episode Data

#### From Pills to Heroin



#### From Heroin to Fentanyl

Overdose Death Rates Involving Opioids, by Type, United States, 2000-2017



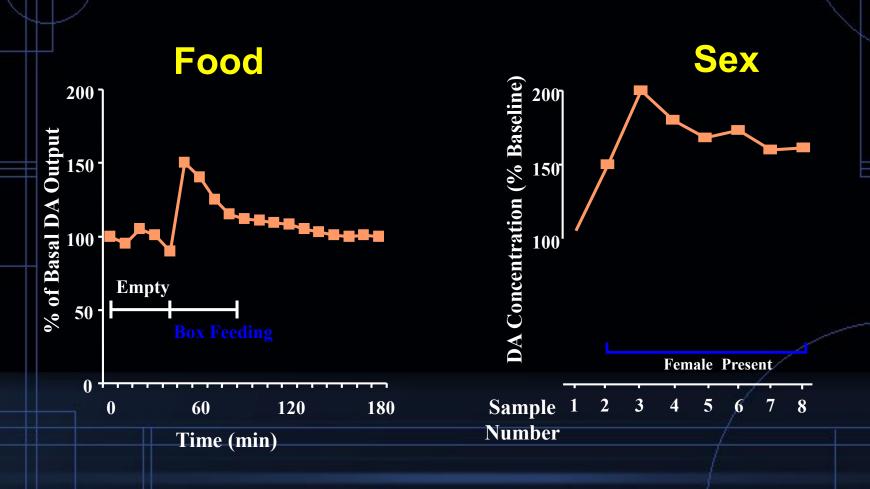


#### From Fentanyl to Carfentanil



# Neurobiology 20

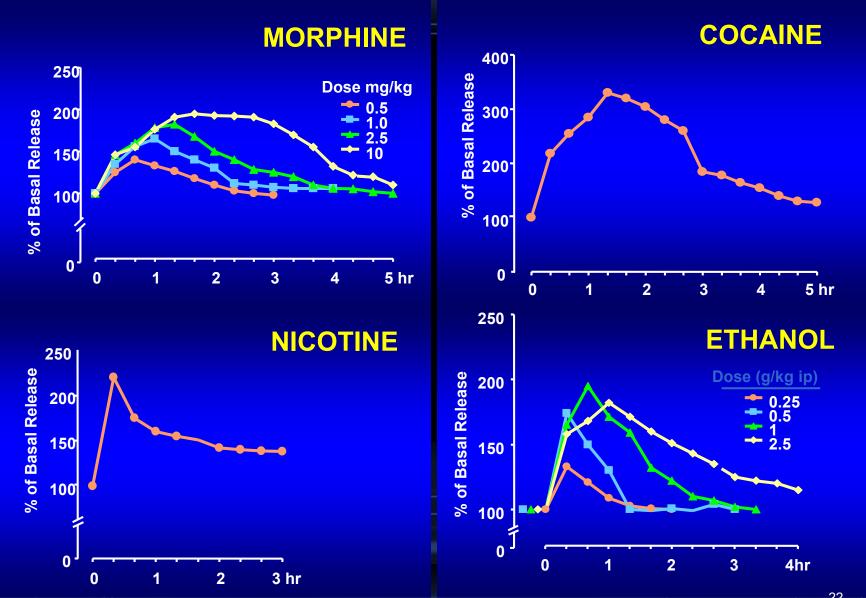
#### **Natural Rewards**



Adapted from: Di Chiara et al, Neuroscience, 1999

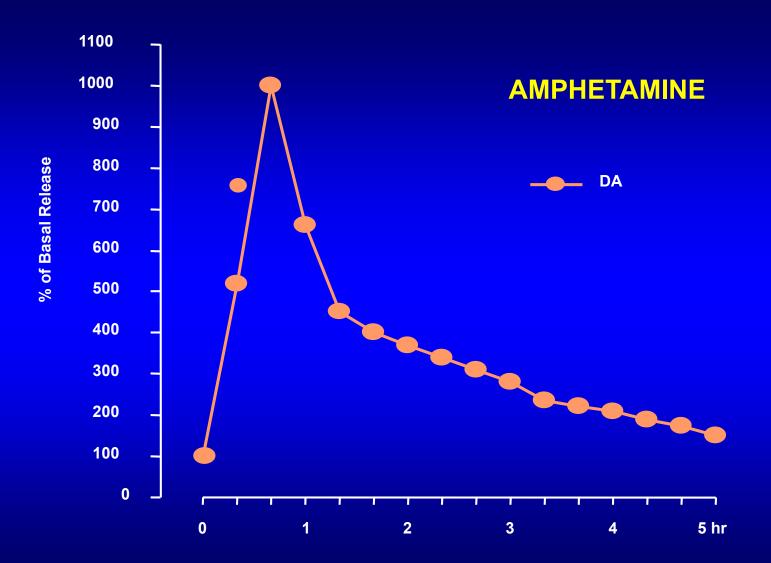
Adapted from: Fiorino and Phillips, J Neuroscience, 1997

#### **Effects of Drugs on Dopamine Levels**



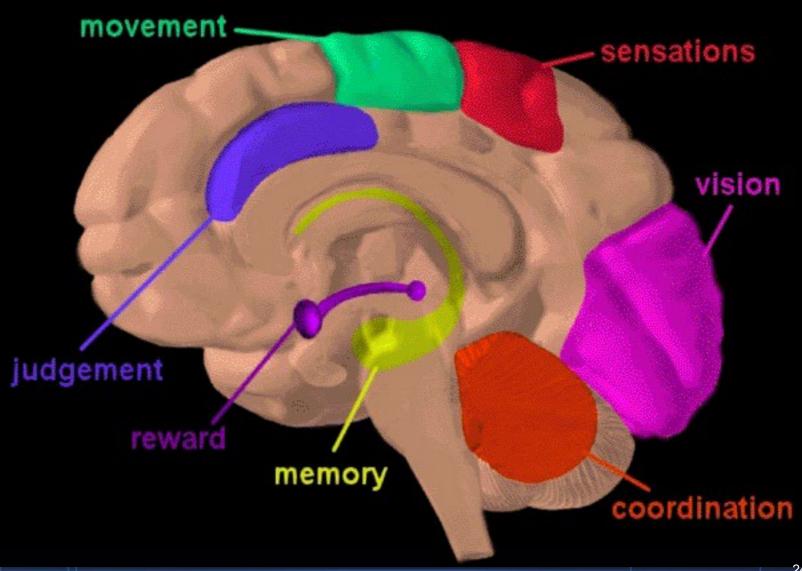
Adapted from: Di Chiara and Imperato, *Proceedings of the National Academy of Sciences USA*, 1988; courtesy of Nora D Volkow, MD

#### **Effects of Amphetamines on Dopamine Levels**



Adapted from: Di Chiara and Imperato, *Proceedings of the National Academy of Sciences USA*, 1988; courtesy of Nora D Volkow, MD.

#### **Pleasure-Reward Pathways**



### 3

## New Neurobiological Concepts



#### **Three Novel Areas**

- Motivational Circuitry
- Antireward Pathways
- Interoception

#### Motivation: The Stinking Thinking Part



#### **Reward Systems**

#### **GAME 1**

A. A sure gain of \$250.

B. 25% chance to gain \$1,000, 75% chance to gain nothing.

#### **Reward Systems**

#### **GAME 1**

- A. A sure gain of \$250.
- B. 25% chance to gain \$1,000, 75% chance to gain nothing.

84%

16%

#### **Antireward Systems**

#### GAME 2

A. A sure loss of \$750.

B. 25% chance to lose nothing, 75% chance to lose \$1,000.

#### **Antireward Systems**

#### GAME 2

- A. A sure loss of \$750.
- B. 25% chance to lose nothing, 75% chance to lose \$1,000.

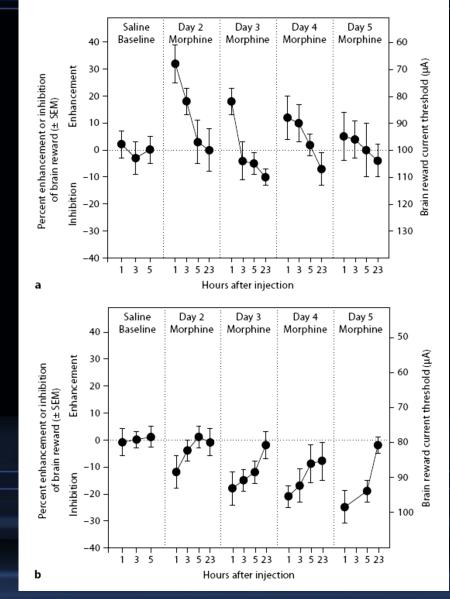
13%

87%

#### **Human Nature**

- People avoid risks to ensure gains (even small gains).
- People take risks (even big risks) to avoid definite losses.
- Psychology trumps probability.

#### Reward and Antireward Systems



#### sensual touch

thirst

temperature

INTEROCEPTION

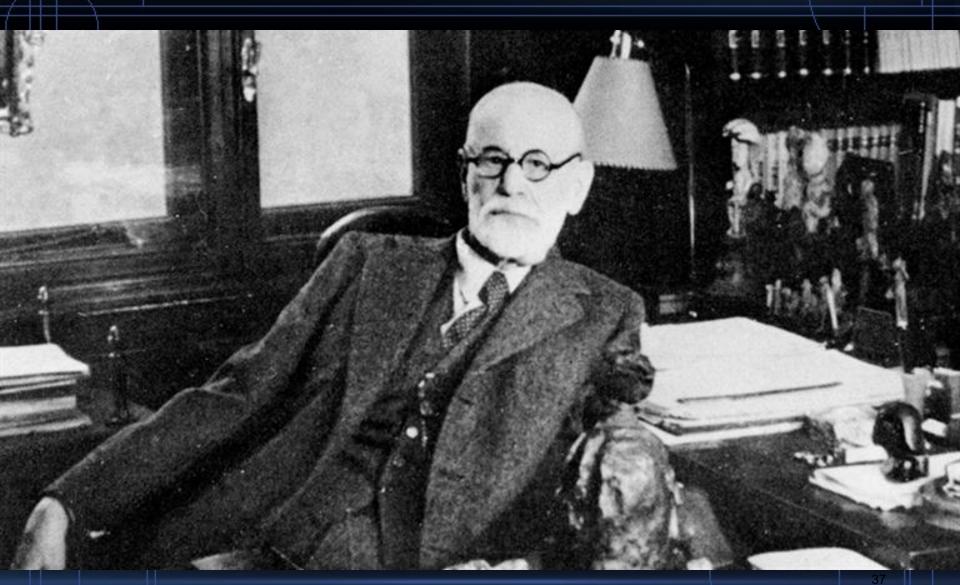
PAIN

hunger

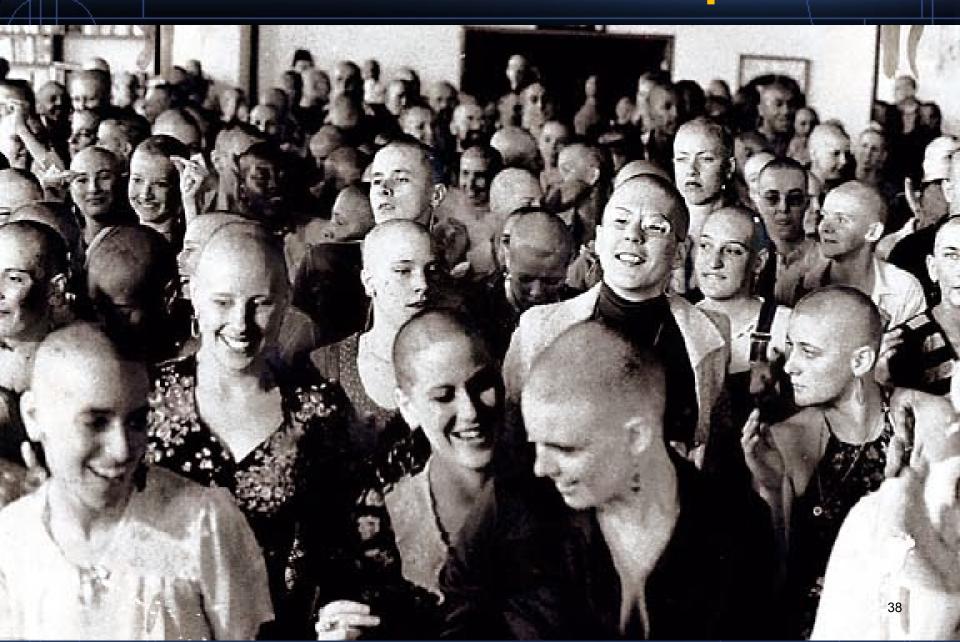
breathlessness



## 1st Wave: Psychoanalysis



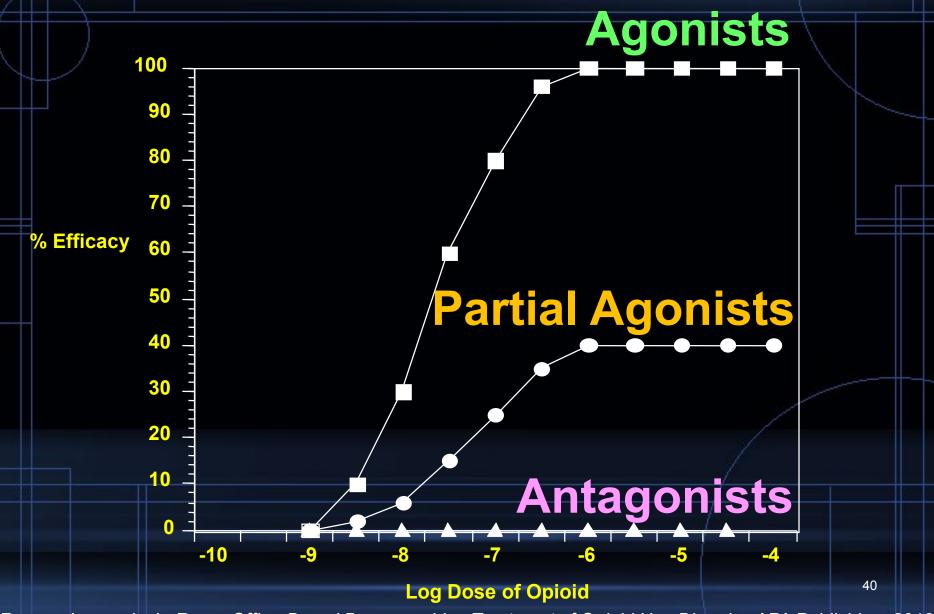
## 2<sup>nd</sup> Wave: Boot Camps



### 3rd Wave: The Current Approach

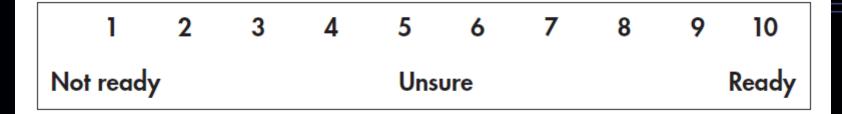
- Medications
- Counseling & Psychotherapy
- Mutual Help





Renner, Levounis, LaRose, Office-Based Buprenorphine Treatment of Opioid Use Disorder, APA Publishing, 2018.

### **Motivational Interviewing**



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### **Attitudes Towards AA**

### MEDICAL STAFF PATIENTS

- 1. Housing
- Government 2. God
- 4. Outpatient Treat. 4. AA
- 5. Job
- 6. Community
- Inner Peace
- 9. God
- 10. Spirituality
- 11. AA

- Inner peace

  - 5. Housing
  - 6. Spirituality
- 7. Trusting People 7. Outpatient Treat. 7. Inner Peace
  - 8. Community
  - 9. Government
  - 10. Trusting People 10. Spirituality
  - 11. Job

### What Med Staff Think **Patients Think**

- 1. Housing
- 2. Outpatient Treatment
- 3. Medical Services 3. Medical Services
  - 4. Job
  - 5. Trusting People
  - 6. AA

  - 8. Community
  - 9. Government

  - **11.** God



# New Directions

### 4th Wave: Mindfullness

"Between stimulus and response there is a space. In that space is our power to choose our response. In our response lie our growth and our freedom."

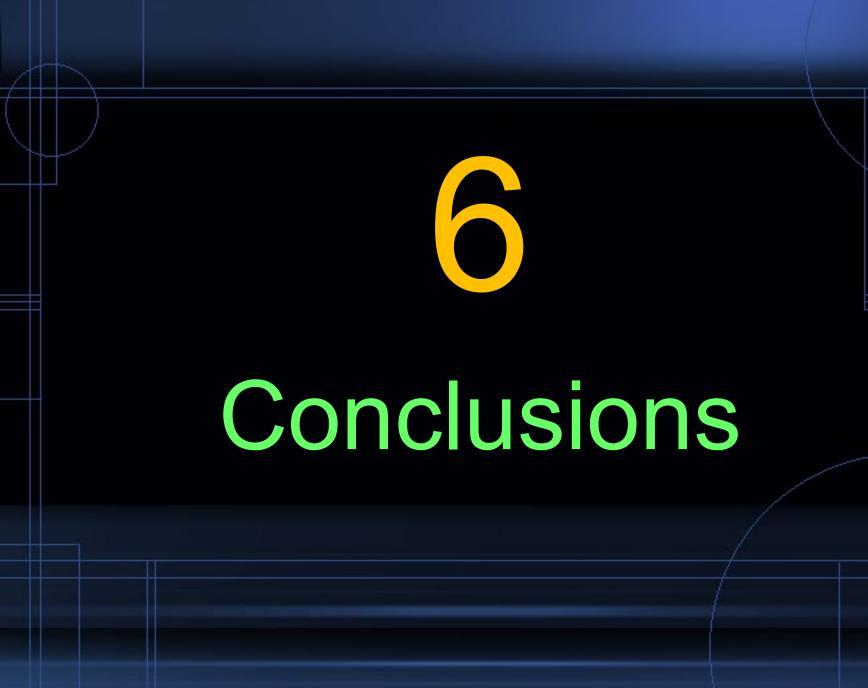
Viktor E. Frankl

### And Back to Psychodynamics...



\*p<0.05, \*\*\*p<.001 based on logistic regression analysis adjusted for race, age, educational level, personal income, employment status, relationship status, health insurance status, geographic location, MSA, age at alcohol onset, and family history of AOD problems. Reference group was "heterosexual" group.

McCabe SE, *Addiction*, 2009; courtesy of Sean McCabe, PhD. Levounis, Drescher, and Barber, *The LGBT Casebook*, 2012.



- 1. A medical mistake, a greedy industry, and a tricky brain are primarily responsible for the opioid epidemic.
- 2. Addiction is the war between the hijacked pleasure/reward pathways of the brain and the frontal lobes.
- 3. Motivational circuitry, the anti-reward pathways, and interoception complete the STOP-GO model of addiction.
- 4. Partial agonists, Motivational Interviewing, and 12-step work are the first line interventions in 2019.
- 5. Mindfulness and a renewed consideration for psychodynamic psychotherapy are next frontiers in the psychosocial treatment of addiction.

# Thank you

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