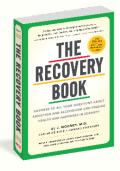
From Science to Serenity: An experienced addictionist discusses recent developments in addiction research

April 9, 2019

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 - Member: NC Substance Abuse Professional Practice Board
 - Medical Director, Healing Transitions; Raleigh, NC
 - Adjunct Associate Professor of Family Medicine; UNC Chapel Hill



FROM SCIENCE TO SERENITY:

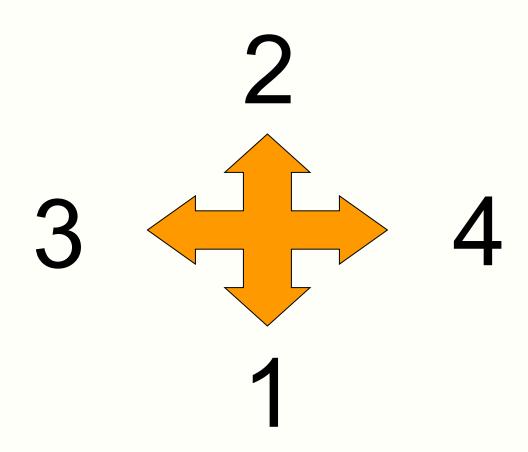
An experienced addictionist reflects on alcoholism and recovery

- 1. Review critical addiction knowledge
- 2. Highlight 5 most important recent addiction research directions
- 3. Challenges to finding solutions to the addiction epidemic

Three most important enduring principles of addiction knowledge

- 1. Drugs of addiction
- 2. Tolerance & dependence
- 3. Commercialization of science

Addiction 101: Drugs of addiction



Addiction 101: Drugs of addiction



Drugs of addiction

Kratom

Ultram

Methadone

Buprenorphine

Heroin

Codiene

Morphine

Dextromethorphan

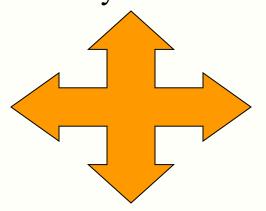
"Krokodil"

Suboxone

Carfentanil

Cocaine Bath Salts
Ritalin "Molly"
Crack Flakka
Adderal

Crystal Meth



"Special K"
Marijuana
LSD
Peyote
PCP
Spice

Benadryl
Zoloft
Prozac
Paxil
K-2

Alcohol

Xanax Ativan Restoril Valium Klonopin Ambien Butalbital

Soma Methaqualone

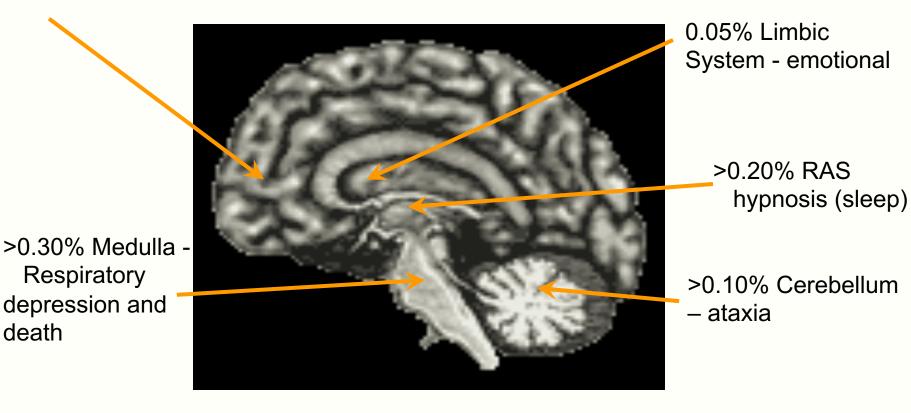
Neurontin Sonata

Alcohol effect on brain

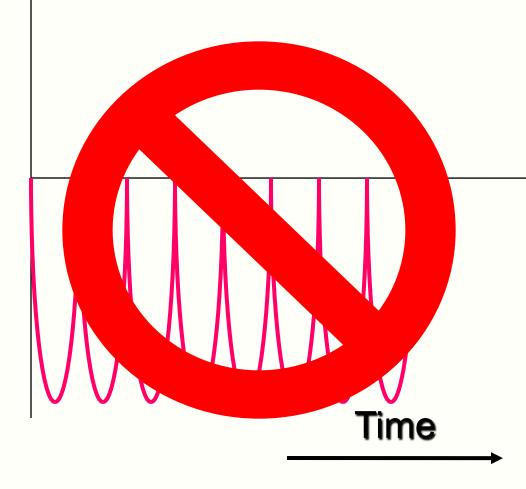
< 0.05% Prefrontal Cortex - disinhibition

Respiratory

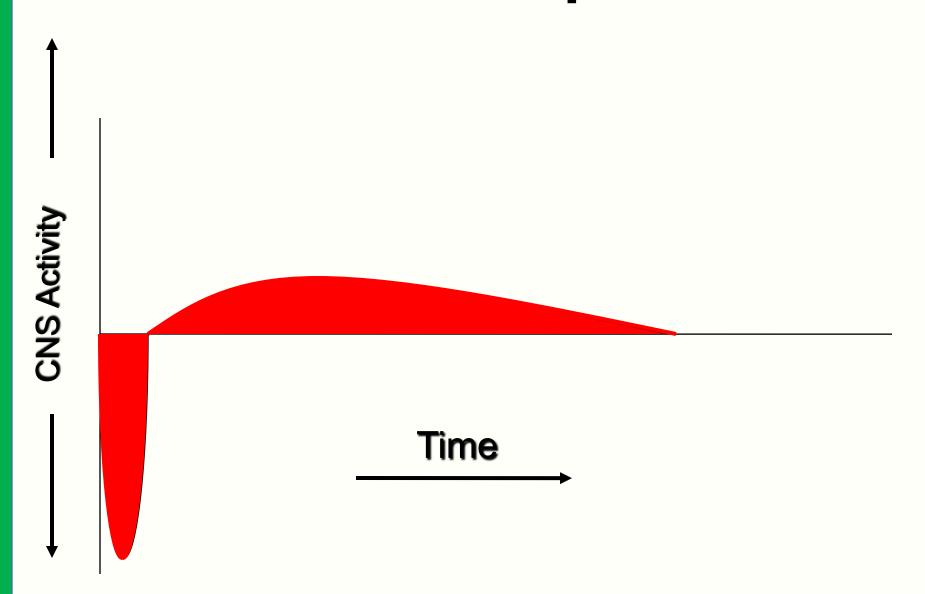
death

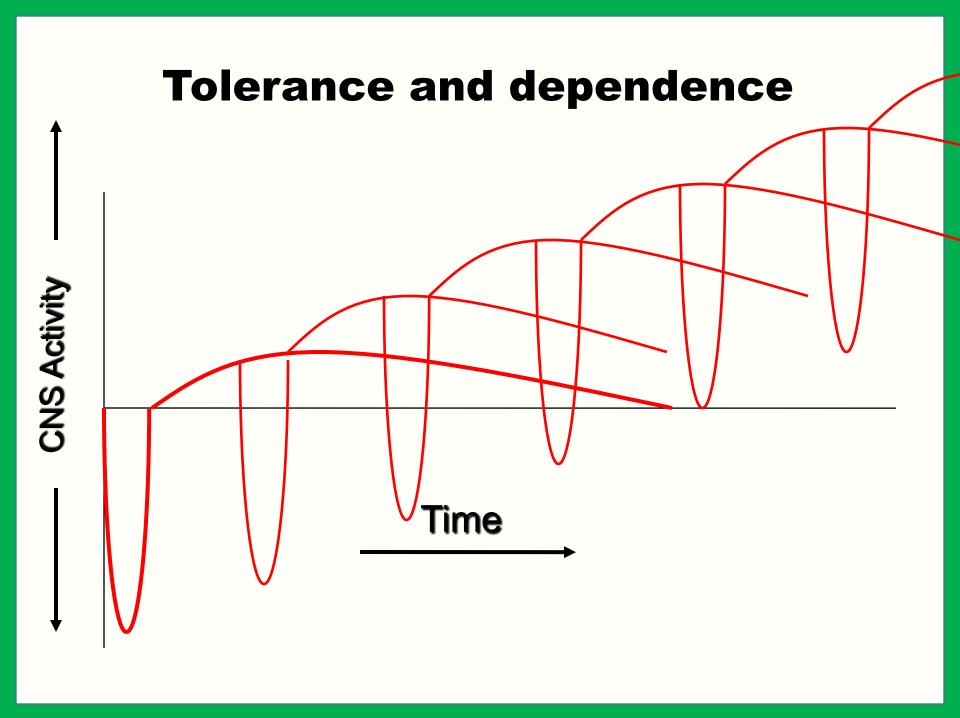


Addiction 101: Tolerance and dependence

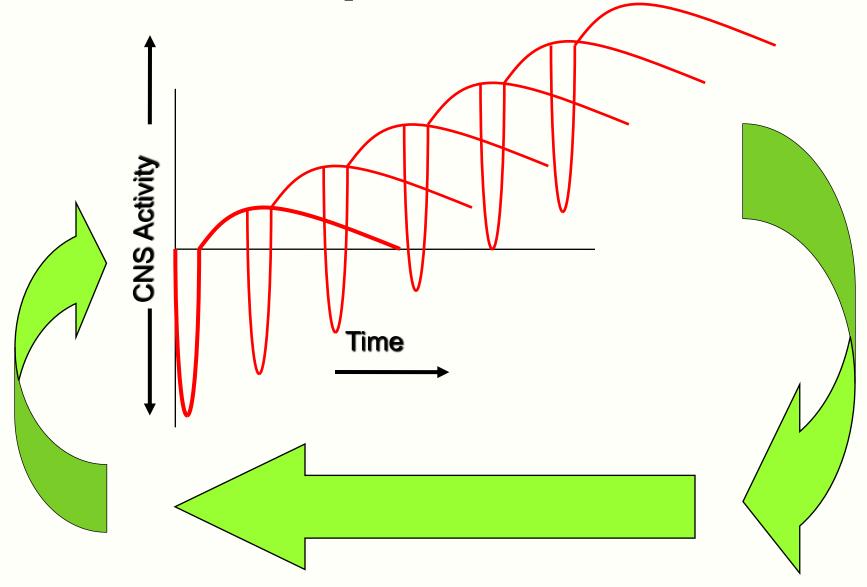


Tolerance and dependence





Tolerance and dependence vs. addiction



Consequences of commercializing science

- Flaws from overreliance on an "evidence based" approach
 - Always retrospective, so evidence serves best as the foundation of good practice
 - Evidence can support practices based on untruths (antibacterials, allergies)
 - Inaccurate observations may reflect the truth

Consequences of commercializing science

- Flaws from overreliance on an "evidence based" approach
 - Always retrospective, so evidence serves best as the foundation of good practice
 - Evidence can support practices based on untruths (antibacterials, allergens)
 - Inaccurate observations may reflect the truth
 - Under control of those with money and power
- Market model of recurring revenues (vaccines vs life style Rxs)
- Historical cycle of drug solutions to drug problem

Willingway tranquillizer rule

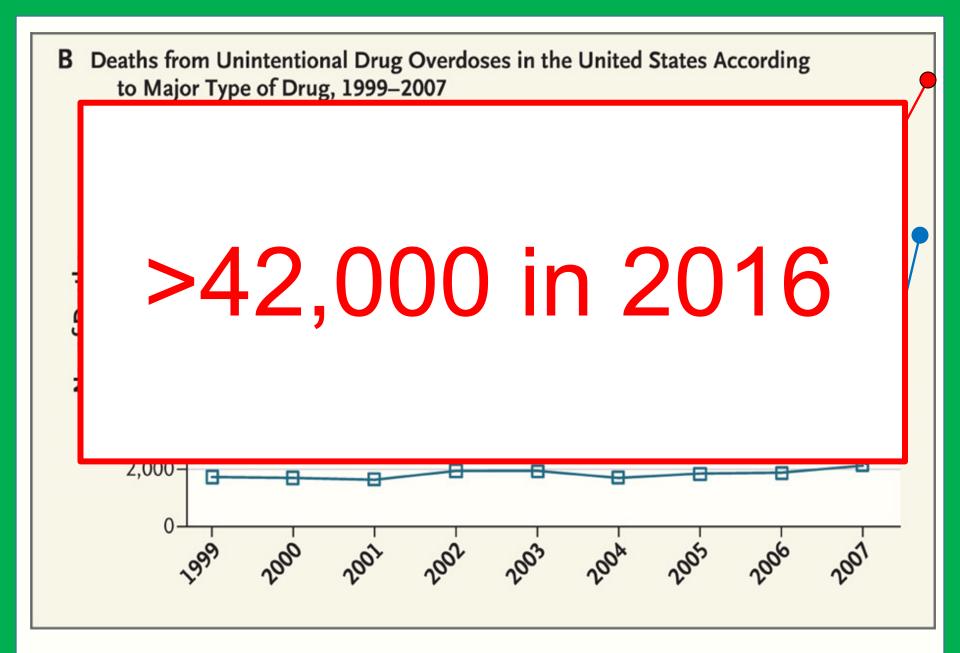
- There are old drugs
- And there are new drugs
- The old drugs are addicting
- And the new drugs are non-addicting
- When the new drugs become old drugs
- They become addicting and are replaced by new non-addicting drugs

Disease facts – #1 health problem

- Fatal illness
 - One alcohol related death every 10 seconds (globally)
 - 100,600 deeths annually
 - 170 deaths/day from overdose alone (USA)
 - YPLL (2.3 iviillion) Cancer, Heart Disease
 - >80% unanticipated deaths in community mental health
- Most important Health problem
 - \$466 billion annual cost of NOT treating
 - No claims data to use for policy formulation
- 7% U.S. population afflicted
- ¼ of inspital admissions (#1 diagnosis of military admissions)
- Family morbidity 43% (76,000,000)
- 100% increase in healthcare costs
- Primary cause of preventable birth defects
- Treatment yield of \$7 for \$1 expense
- \$39 Yield for each \$1 spent in medical settings
- Treatment wasses >00% (EAA LIIMS)
- Prevention reduces risk by 75%

Disease facts – #1 health problem

- Fatal illness
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- Most important Health problem
 - \$466 billion annual cost of NOT treating
 - No claims data to use for policy formulation
- 7% U.S. population afflicted
- 1/4 of hospital admissions (#1 diagnosis of military admissions)
- 4% of NICU admission are for neonatal abstinence syndrome
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- 100% increase in healthcare costs
- Primary cause of preventable birth defects
- Treatment yield of \$7 for \$1 expense
- \$39 Yield for each \$1 spent in medical settings
- Treatment success >90% (FAA HIMS)
- Prevention reduces risk by 75%



Global annual reported polio cases, 1974-2002

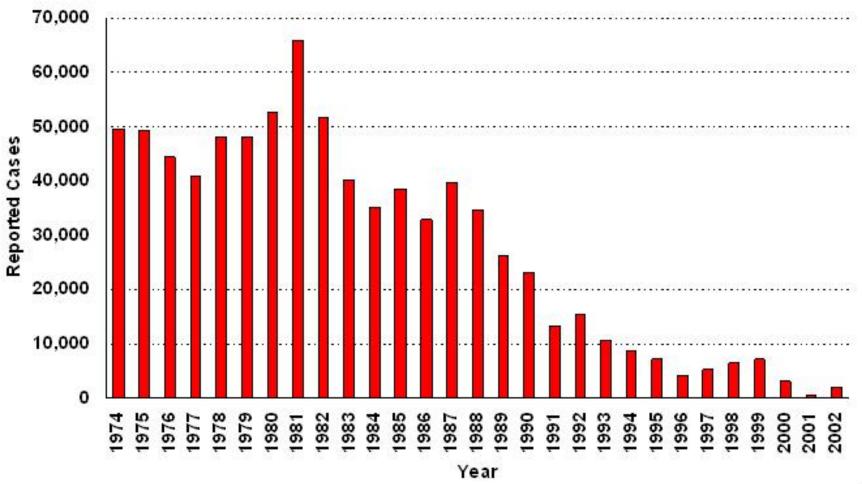
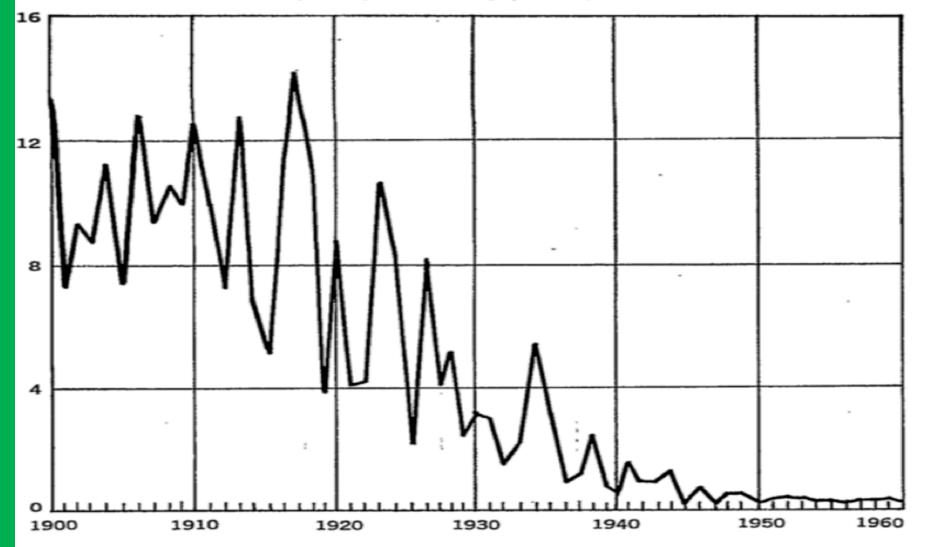
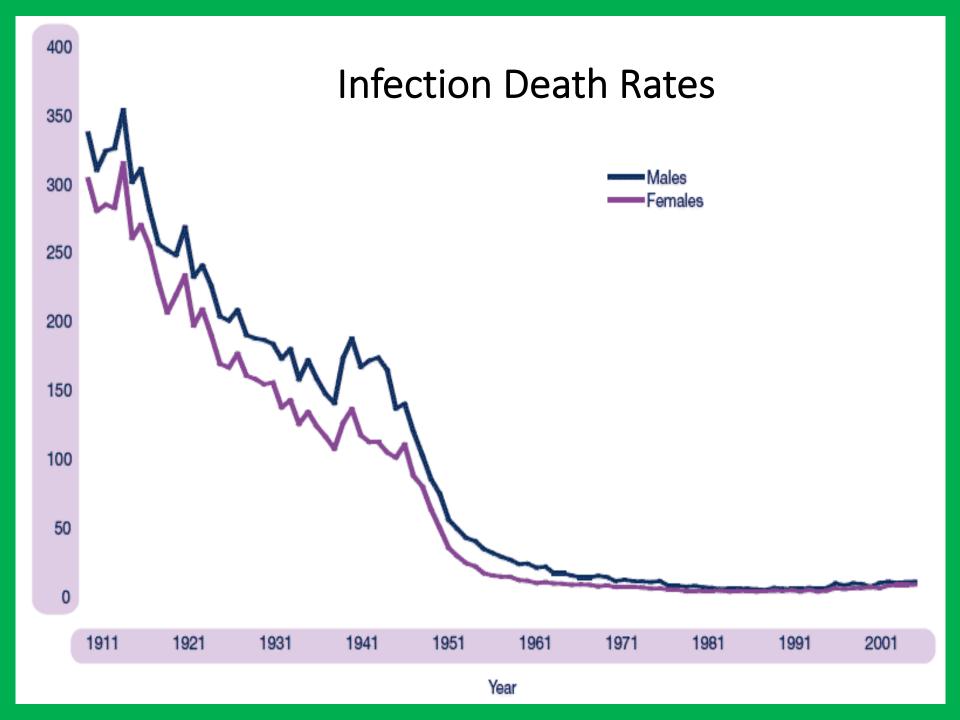




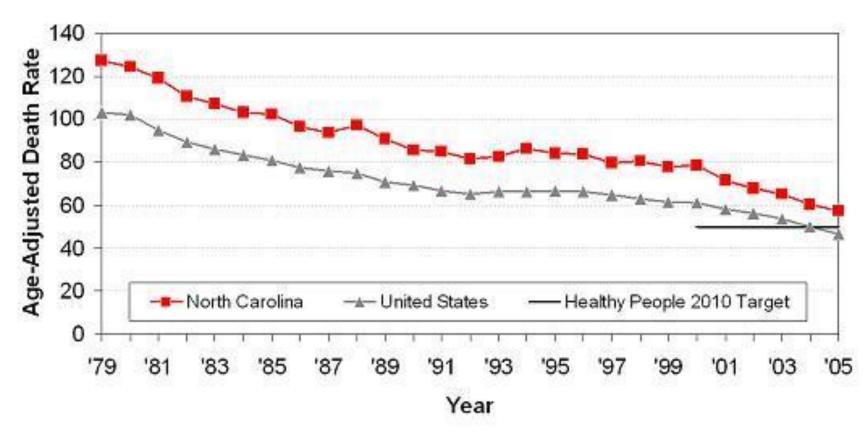
Figure 19.—Death Rates for Measles: Death-registration States, 1900–32, and United States, 1933–60

(Rates per 100,000 population).

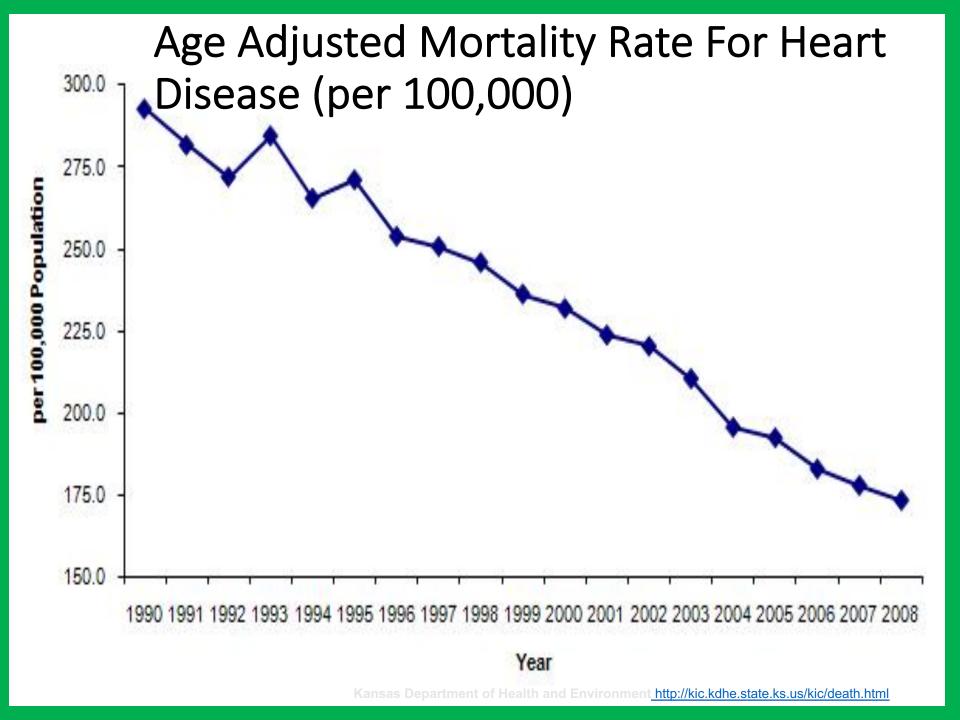


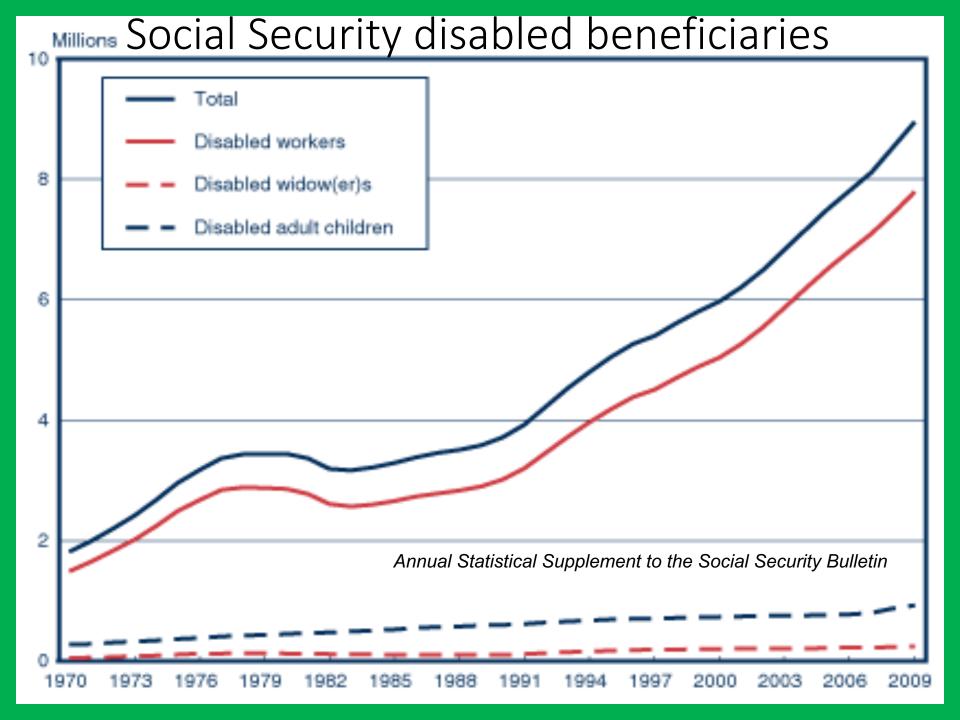


Stroke Death Rates, 1979-2005



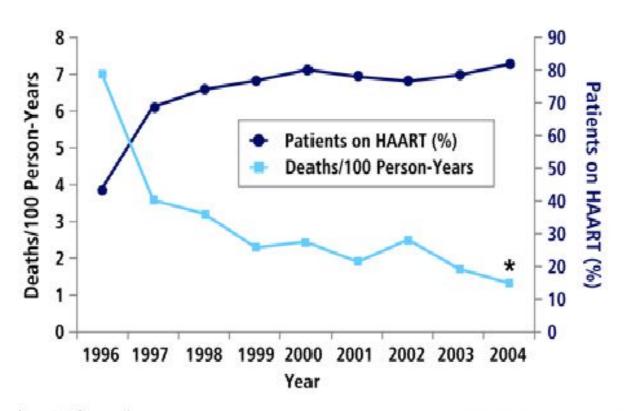
1999-2005; ICD-10 codes I60-I69; 1979-1998; ICD-9 codes 430-434,436-438 multiplied by comparability ratio of 1.0588. Rates per 100,000 population, age-adjusted to the 2000 U.S. standard population. Data Source: Compressed Mortality File, CDC Wonder.





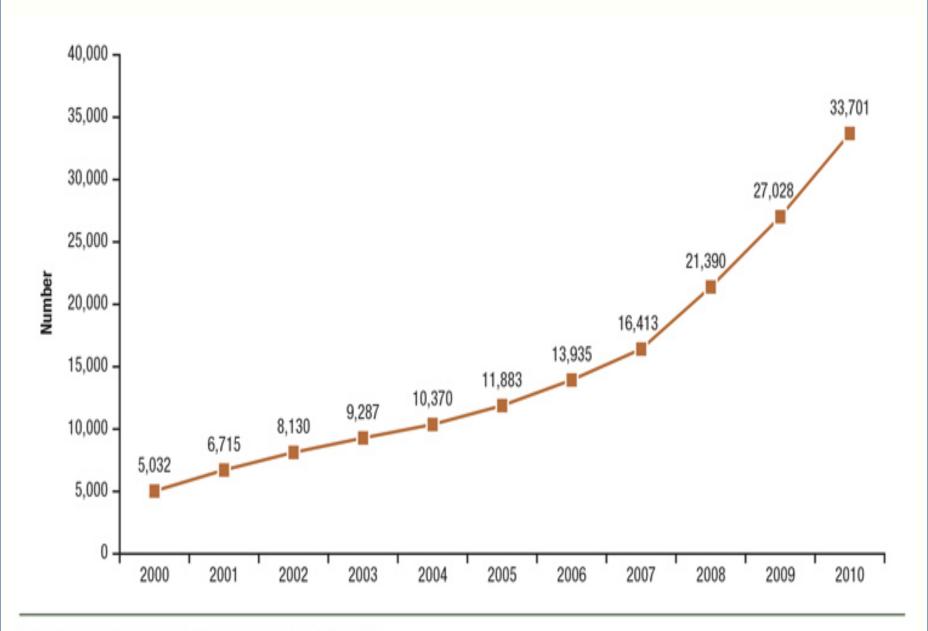
HIV Mortality with more access to treatment

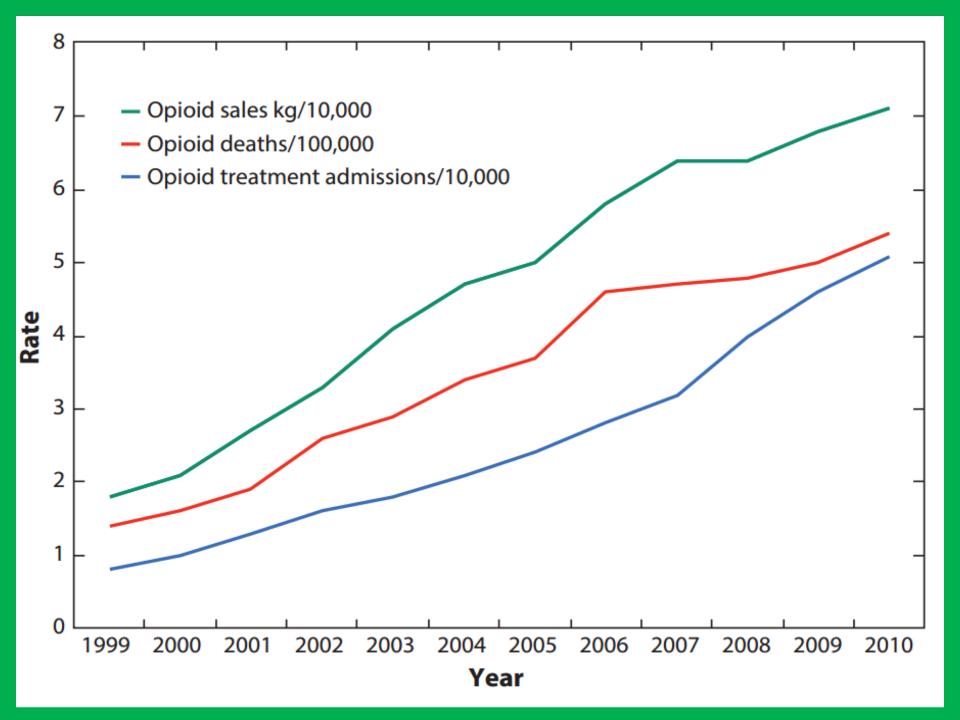
Mortality and HAART Utilization Over Time

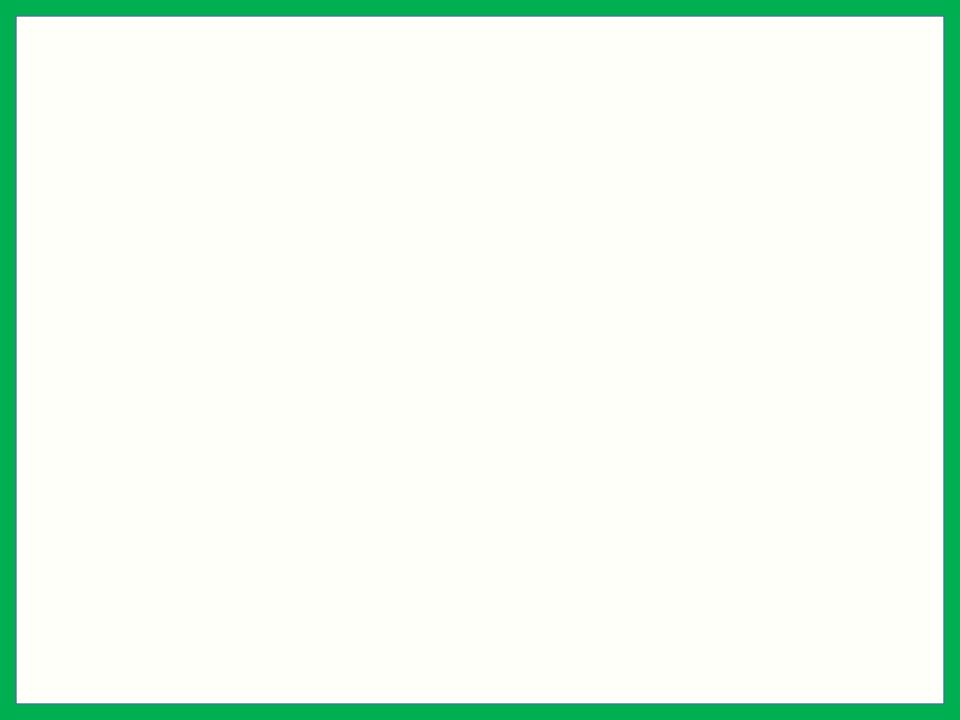


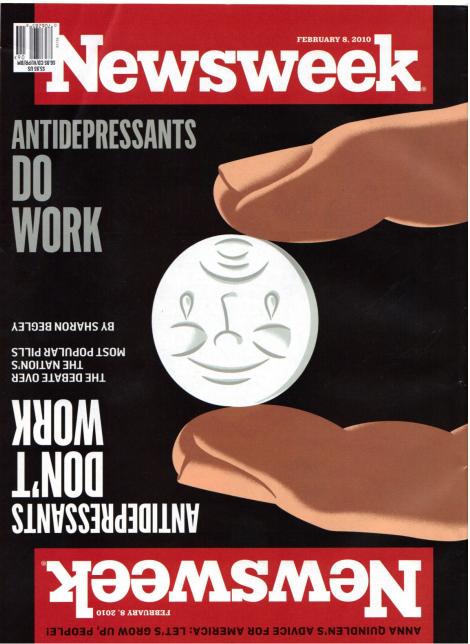
*P=.008 for trend
Palella FJ, et al. J Acquir Immune Defic Syndr. 2006;43:27-34.











Andrews PW, Thomson JA Jr, Amstadter A and Neale MC (4/24/2012) Primum non nocere: an evolutionary analysis of whether antidepressants do more harm than good. Front. Psychology 3:117.

How to be "blinded by the evidence" in a "post-truth" era • Scientific methods can be used in ways that corrupt measurement of

- Scientific methods can be used in ways that corrupt measurement of meaningful outcomes
 - 1. Length of study misrepresents outcomes of practice usage (Xanax, FDA)
 - 2. Measure effects to justify use of the medication (Ambien, methadone)
 - 3. Ignore illness criteria which don't support intended study outcomes (DSM V exclusion for SUD)
 - 4. Avoid measurement of side effects (atypicals, antidepressants)
 - 5. Emphasize statistical endpoints irrelevant to efficacy (acamprosate) (Drug Court)
 - 6. Pay researchers to publish positive outcomes (Bankole Johnson)
 - 7. Don't report negative results (Irving Kirsh FOIA discoveries)
 - 8. Overemphasize positive results (Acamprosate NNT)
 - 9. Publish research ghostwritten by marketing staff under physician bylines
 - 10. Avoid comparison with effective alternatives (Phenob, Suboxone, abstinence)
 - 11. Generalize data to populations excluded from study (alcoholics)
 - 12. Infer efficacy by ignoring oscillations in illness severity
 - 13. Only count results that look good (look for "response rate") (Naloxone)
 - 14. Define a disease by an anticipated drug effect ("Serotonin deficiency", SWSD)

Fines and Settlements for illegal or unethical drug marketing practices

- Astra Zeneca \$520,000,000 Seroquel
- Pfizer \$2,300,000,000 Geodon, Lyrica ...
- Purdue Pharma\$634,500,000 Oxycontin
- •Lilly \$1.400,000,000 Zyprexa
- •Glaxo-Smith-Kline \$3,000,000,000 Paxil, Welbutrin, ...
- •Abbott \$1,600,000,000 Depakote
- Johnson and Johnson \$2,200,000,000 Risperdal
- Bristol-Myers Squibb \$515,000,000 Abilify

Most important recent addiction research directions

- 1. What causes it? Allergy: "New" theory explains addiction insanity
- 2. What keeps it going? Denial: Neuroscience of cognitive dissonance
- 3. Why can't we stop it? Motivation: The science of readiness to change
- **4. Why is healing from it so slow?** Lifting the Fog: Neuroplasticity
- **5. What keeps it away forever?** Lifetime Healing: Recovery Zone System

1. Allergy & Immunity

Insanity

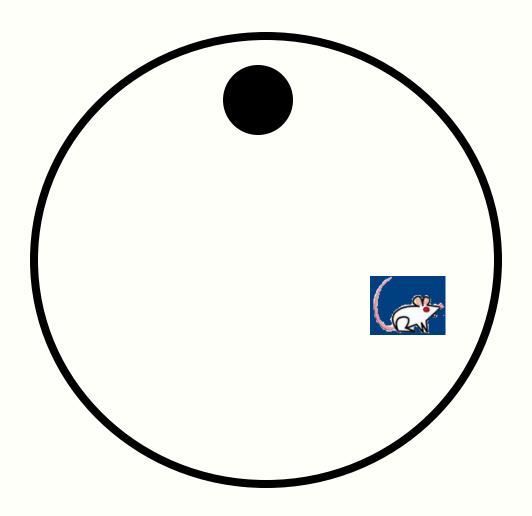
•Doing the same thing over and over again expecting a different result.

Addiction Insanity

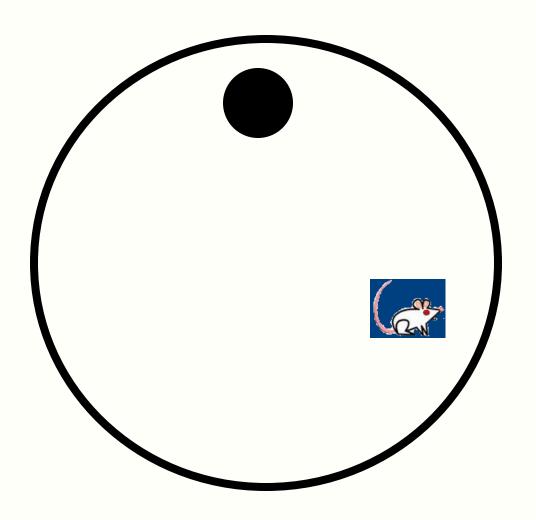
 Continuing to drink or use drugs in spite of awareness that one should not. "The doctor's theory of an allergy interests us"

 Alcoholics Anonymous – The Doctor's Opinion (1939)

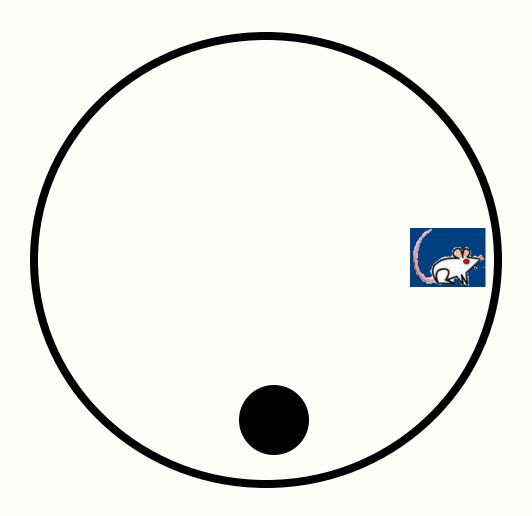
Non-alcoholic rat



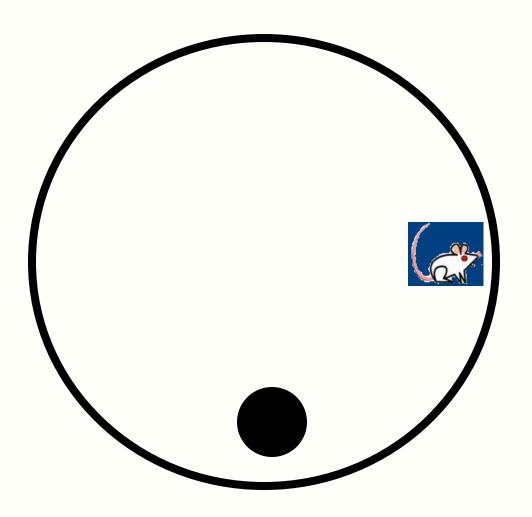
Alcoholic rat



Non-alcoholic rat

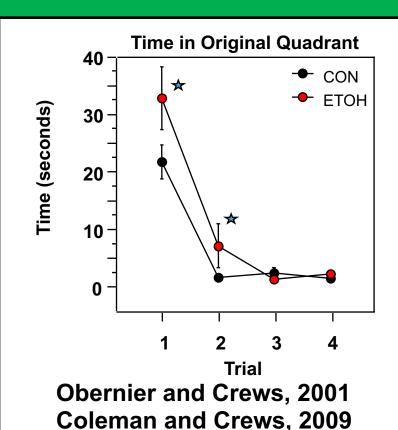


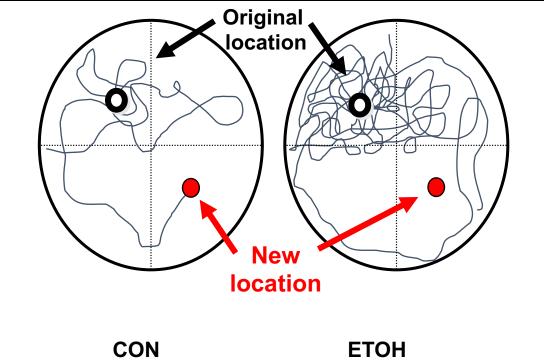
Alcoholic rat



Binge Ethanol treated animals have persistent reversal learning deficits.

Ethanol binge treatment of adolescent mice or rats results in persistent reductions In reversal learning, an indicator of disrupted frontal cortical and learning networks. Adolescent ethanol results in adult relearning deficits in both Morris water maze and Barnes spacial learning maze.





Search path of rats. Open circle original platform-red circle new location. Binge ethanol treated rats perseverate on old location.

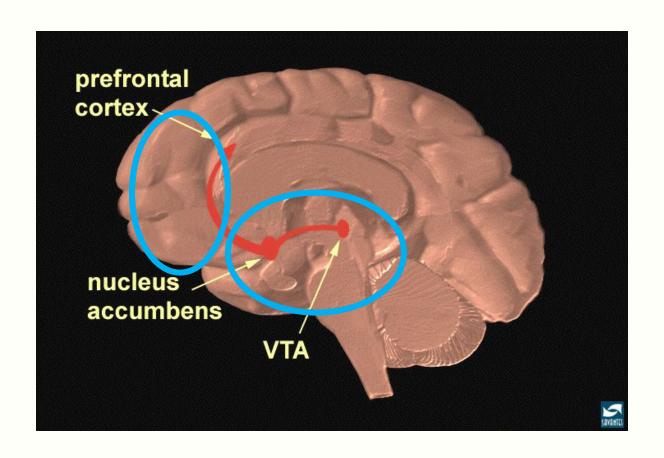
Immune theory of addiction: Induction of innate immunity

Genetics (DNA) Allergen **Transcription** Innate immune proteins Innate immune reaction Impaired function: prefrontal cortex Alcoholic insanity

2. Neuroscience of cognitive dissonance

Formula of mature human behavior

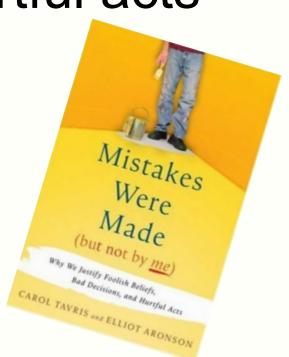
F (Feelings) +T (Thoughts) =A (Actions)



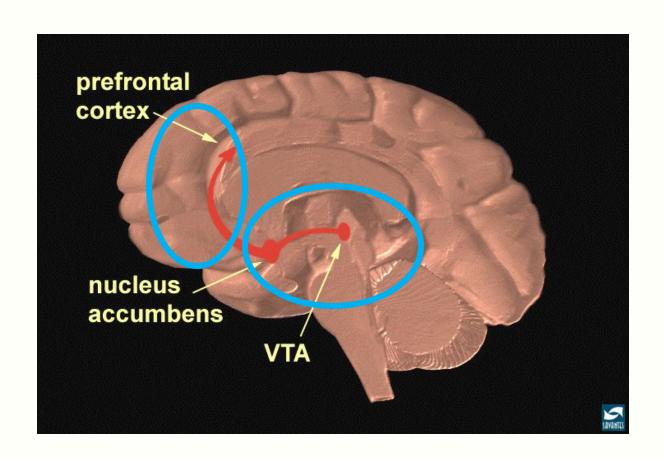
Denial & cognitive dissonance

"Mistakes were made, but not by me: Why we justify foolish beliefs, bad decisions, and hurtful acts"

Carol Tavris and Elliot Aronson

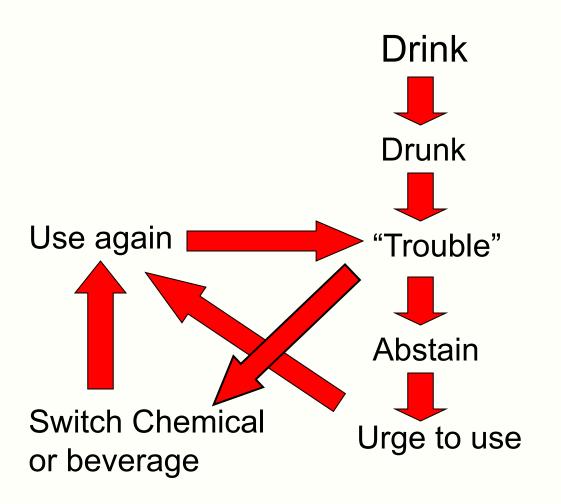


Hierarchy of human behavior



Alcoholic denial resolves conflict caused by drinking too much

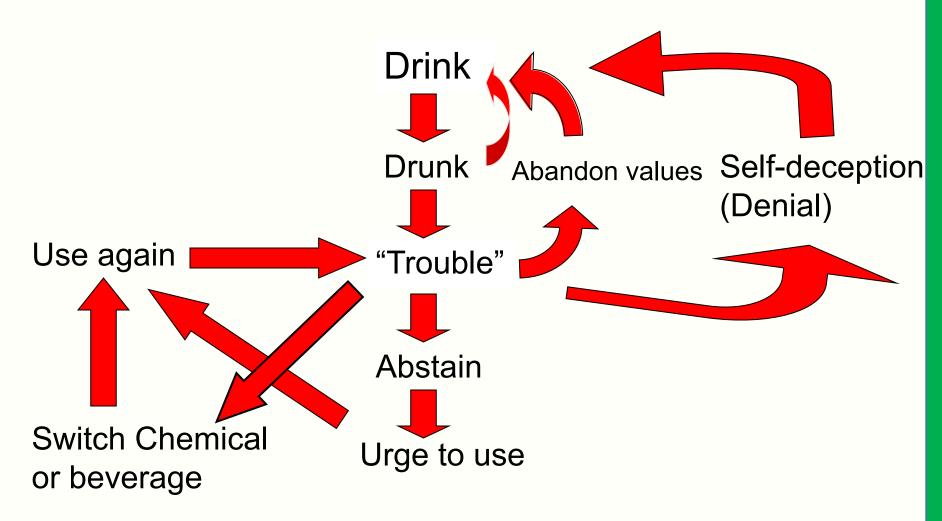
Addiction Cascade



Addiction Cascade

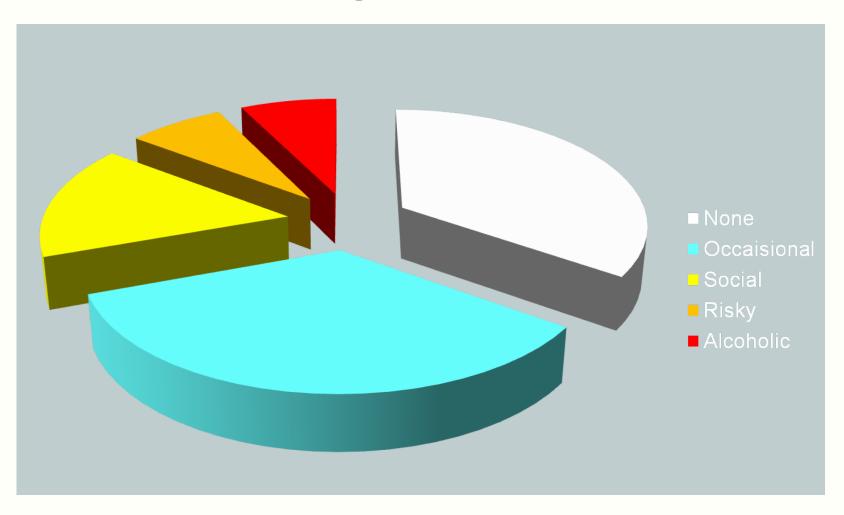
- ➤ Conflict between values and behavior
 - Drinking and drug use is causing harm
 - BUT
 - Drinking (or using drugs) is a need that must be satisfied
- >=
- ➤ Cognitive dissonance
 - an uncomfortable feeling or stress caused by holding two contradictory ideas simultaneously

Addiction Cascade



3. Science of readiness

Identifying the problem: Drinking in America



Motivational Readiness

Maintenance (Steps 10-12)

85%

Action (Steps 4-9)

Preparation (Steps 1,2 &3)

Contemplation

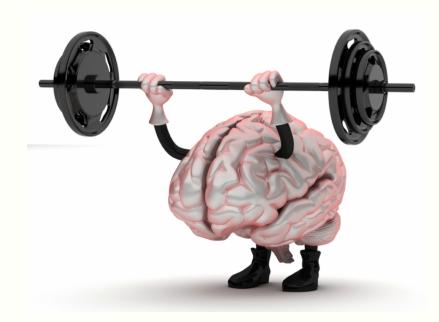
Precontemplation

4. Neuroplasticity

Neuroplasticity:

 The capacity of neuronal substance to change in response to environment and experience

"Psychic change"



Types of neuroplasticity

- Negative (brain impairment)
- Positive (brain healing)

- Adding connections
- Removing connections (pruning)
- Sensitizing connections
- Desensitizing connections
- Adding neurons (neurogenesis)

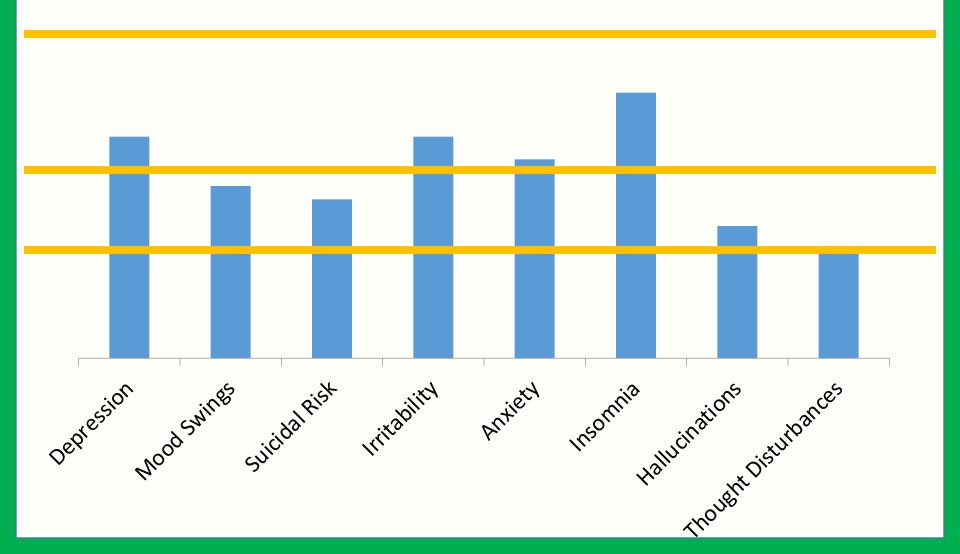
Heal Your Brain

- Thoughts & actions remold brain:
 - Spur new neurons
 - Create new connections; prune old
 - Restore neurotransmitter function
 - Restore frontal lobe (exec brain) relationship with limbic brain (primitive) -> better decisions, behavior

Drain the Swamp and see the stumps: Clinical correlation of recent addiction neuroscience and psychiatric co-morbidity

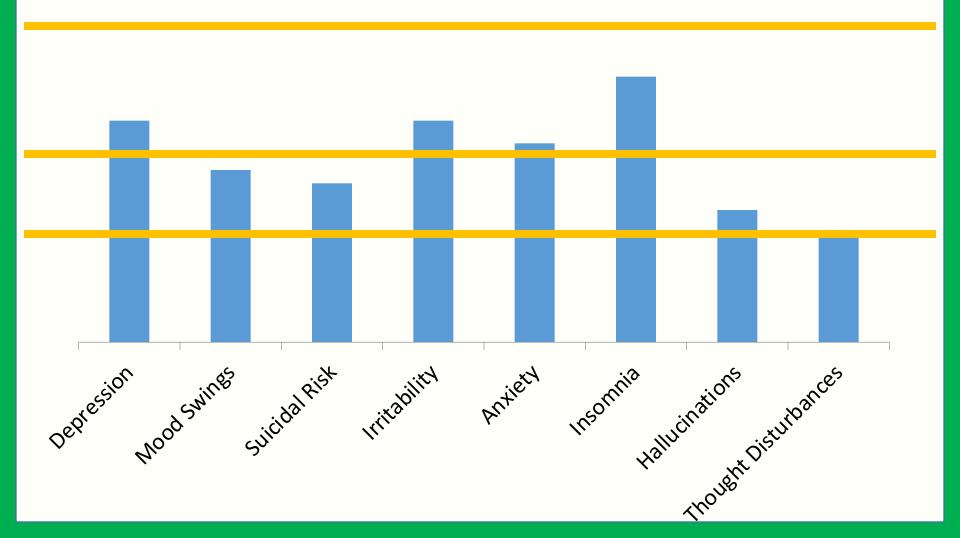


Stress, fatigue, sleep and sensory deprivation etc. lower threshold for latent emotional symptoms



Impact of threshold changes of addiction on mental co-morbidity

(binge drinking and induction of innate immune genes create co-morbidity) (and neuroplasticity of recovery helps resolve it)



DIAGNOSTIC AND STATISTICAL MANUAL OF MENTAL DISORDERS

FIFTH EDITION

DSM-5

AMERICAN PSYCHIATRIC ASSOCIATION

DSM5 exclusionary criteria

> Major Depressive Disorder:

C. The episode is not attributable to the physiological effects of a substance or to another medical condition.

Bipolar I Disorder:

D. The episode is not attributable to the physiological effects of a substance (e.g., a drug of abuse, a medication, other treatment) or to another medical condition.

Note: A full manic episode that emerges during antidepressant treatment (e.g., medication, electroconvulsive therapy) but persists at a fully syndromal level beyond the physiological effect of that treatment is sufficient evidence for a manic episode and therefore, a bipolar I diagnosis.

Bipolar II Disorder:

F. The episode is not attributable to the physiological effects of a substance (e.g., a drug of abuse, a medication or other treatment).

DSM5 exclusionary criteria

- Obsessive Compulsive Disorder:
 - C. The obsessive-compulsive symptoms are not attributable to the physiological effect of a substance (e.g., a drug of abuse, a medication) or another medical condition.
- Anxiety Disorder:
 - E. The disturbance is not attributable to the physiological effects of a substance (e.g., a drug of abuse, a medication) or another medical condition (e.g., hyperthyroidism).
- Schizophrenia
 - E. The disturbance is not attributable to the physiological effects of a substance (e.g., a drug of abuse, a medication) or another medical condition.
- Personality Disorder:
 - F. The enduring pattern is not attributable to the physiological effects of a substance (e.g., a drug of abuse, a medication) or another medical condition (e.g., head trauma).

5. Recovery Zone System

http://bit.ly/recoveryzones



The Recovery Zone System

- Three Recovery Zones; blueprint for life
 - 1) Red Zone: Stop.
 Activate your recovery.
 - 2) Yellow Zone: Proceed with caution.
 Build or rebuild your life.
 - 3) Green Zone: Go.

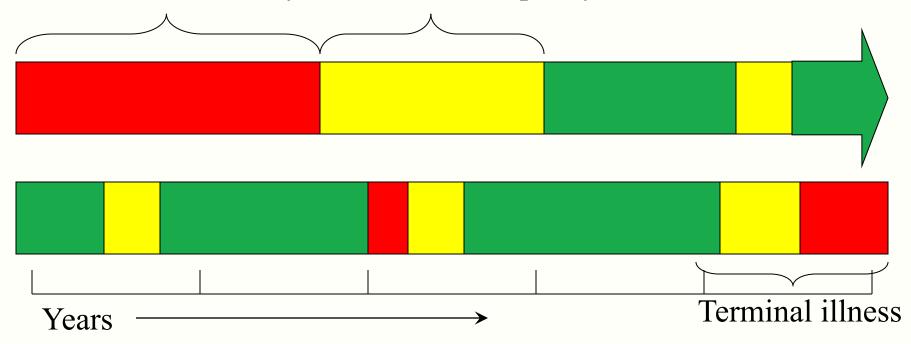
 Celebrate your life.

Recovery Zone System

- The Recovery Zones
 - Red Activate
 - Yellow Build
 - Green Celebrate
- Recovery Zone ReCheck
- TAMERS

Zones in the recovery timeline

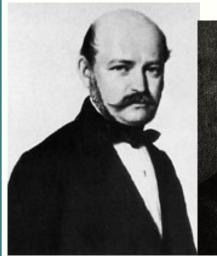
Activate recovery Build life to capacity

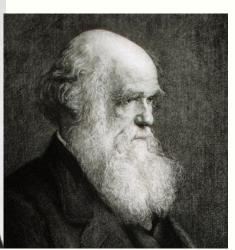


TAMERS

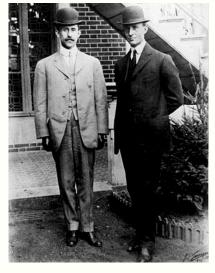
- Think about recovery, Talk about recovery
- Act on recovery, connect with others
- Meditate and Minimize stress
- Exercise and Eat well
- •Relax
- •Sleep

Overcoming Barriers to Scientific Progress













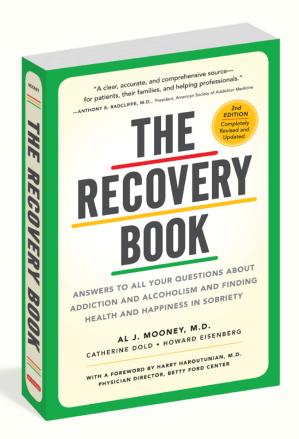




THE RECOVERY BOOK

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http://bit.ly/CornerstoneHIMS2019