Neurobiology of Addiction

LOWELL J ROBERTSON MD

INTERNAL/ADDICTION MEDICINE DIRECTOR OF DETOXIFICATION AND INPATIENT SERVICES GRAND ARC, TULSA

Objectives

Understand	Understand addiction as an illness
Understand	Understand the biologic basis of addiction
Understand	Understand how current treatment trends work towards the above





What is Addiction?

Compulsive behavior despite harmful consequences









Addiction

A brain disease, not a moral failure





What is Addiction?

Cravings

Loss of Control

Physiologic Dependence

Addiction is a Disease

4. ADDICTION CAN BE TREATED

Partial Recovery of Brain Dopamine Transporters in Methamphetamine (METH) Abuser After Protracted Abstinence

Normal Control

METH Abuser (1 month detox) METH Abuser (24 months detox) ml/gm

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Source: Volkow, ND et al., Journal of Neuroscience 21, 9414-9418, 2001.

What Does it Take to be Addicted?

• Bad childhood?

• Traumatic event?

• Bad friends?

What Does it Take to be Addicted?

Exposure to addictive substance

Brain's Response to Exposure

1. Activation of reward circuit

2. Change in receptor density

3. Alteration in neurotransmitter levels



Activation of Reward Circuit

- An area of the brain that is involved in a feedback loop of reinforcement:
 - Nucleus Accumbens
 - Ventral Tegmental Area







www.drugabuse.gov

The Reward Circuit



Reward Circuit

• Tells the individual what to repeat in order to obtain reward

• Builds pathways and connections that remain



Reward Circuit



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Change in Receptor Density

• When more of a substance is available, the body makes more receptors

• This requires more of the substance to feel the same effects



Some drugs target the brain's pleasure center

Brain reward (dopamine pathways)



These brain circuits are important for natural rewards such as food, music, and sex.





While eating food

Dopamine Transporter

Dopamine

While using cocaine

Typically, dopamine increases in response to natural rewards such as food. When cocaine is taken, dopamine increases are exaggerated, and communication is denied.

Alteration in Neurotransmitter Level Drugs Food







Other Considerations

- Childhood is important
- Genetics is the most important factor
- Do not fall for the fallacy that some patients can be safely exposed to an addictive substance based on their background



Adverse Childhood Experiences



Growing up in a household where:



There are adults with alcohol and drug problems



There are adults with mental health problems



There is domestic violence



There are adults who have spent time in prison



Parents have separated

ACE's and Addiction -The Original Ace's Study



Felitti, Vincent, Anada, Robert F. et al. Relationship of Childhood Abuse and Household Dysfunction to Many of the Leading Causes of Death in Adults.

ACE's and Addiction -The Original Ace's Study

Ever Used illicit drugs



Felitti, Vincent, Anada, Robert F. et al. Relationship of Childhood Abuse and Household Dysfunction to Many of the Leading Causes of Death in Adults.

ACE's and Addiction -The Original Ace's Study



Felitti, Vincent, Anada, Robert F. et al. Relationship of Childhood Abuse and Household Dysfunction to Many of the Leading Causes of Death in Adults.

Other Studies

- Early initiation of drug use
 - Each ACE increases risk 2-4 fold
- Drug use problems, addiction, and parenteral drug use
 - 5 or more ACE's increase risk 7-10 fold

Dube S, Felitti V, Dong M, Chapman D, Giles W, Anda R. Childhood Abuse, Neglect, and household dysfunction and the risk of illicit drug use: The Adverse Childhood Experiment Study; Pediatrics 2003 (11)3:564-572

Overall Addiction Risk Attributed to ACE's

•67%

ACE's in Individuals Entering Treatment

A 1-point increase in the ACE is associated with a 1.1. factor increase in reporting a lifetime overdose

Higher ACEs were associated with lower age at initiation

Stein, M., Conti, M. Adverse childhood experience effects on opioid use initiation, injection drug use, and overdose among persons with opioid use disorder. Drug and Alcohol Dependence (179). August 2017. 325-329.

Allostatic load and disrupted neurological development

Complex trauma/ACE

Race/social conditions/local context

Generational embodiment/historical trauma

GENETICS

Research suggests that:

- •Genetics account for 40-60% of a person's risk of developing a substance use problem.
- •Alcohol addiction is about 50% heritable, while addiction to other drugs is as much as 70% heritable.
- •Shared genetic risk variants across different substance use disorders provide insight into the mechanisms that underlie these disorders.

Genetics of Addiction

Alcoholism and genetics

Substance Use Disorder As Chronic Illness

McLellan 2000

Epigenetics

• Epigenetics is the regulation of gene expression. Epigenetic mechanisms include: DNA modifications, DNA methylation, Histone modifications.

• Epigenetic adaptations can affect:

- Susceptibility to addiction
- Response to drugs of abuse
- Response to pharmacotherapy for addictions
- Health
- Expression of traits passed to children

CONCLUSIONS

- Addiction is a disease caused by several factors
- Addiction is inevitable in certain situations
- Addiction is created by the science of the substance (medication)
- All individuals (patients) are at risk
- Childhood is important
- Mother Teresa would get addicted if treated long enough

LOWELLMD@GMAIL.COM LROBERTSON@GRANDMH.COM