

Effect of Opioid Use Disorder on Brain Chemistry

David Butterfield, PharmD, BCPS, BCPP Clinical Pharmacy Specialist-Psychiatry



- Understand the major ways that opioids impact the brain
- Explain the neurobiological mechanisms behind how opioids affect the brain
- Understand how medication assisted treatment (MAT) reverses some of the affects opioids have on the brain
- Understand how stigma can negatively impact treatment for opioid use disorder





Effect of Opioids on Pain Relief Over Time



Accessed from: https://www.oregonlive.com/health/2017/12/opioids_rewire_the_brain_ohsu.html

PO

0

Ø





6

Ø

Neurobiology of Dependence and Withdrawal

- Locus Coeruleus is an area of the brain that is critically involved in dependence and withdrawal with opioids
- Repeated exposure to escalating dosing of opioids alters the brain so it functions "normally" when opioids are present and "abnormally" when opioids are absent
- Clinical important alterations:
 - Tolerance- need to take higher dosage of drugs to achieve the same effect
 - Dependence- susceptibility to withdrawal symptoms





A 0 0 Ø













Discontinuation of Opioids



A

0

Ø





 $(\bigcirc$ 0 6 Ø





(O)

6

Ø

0

60

Volkow, et al. New England Journal of Medicine 2016; 363-371.

Withdrawal/Negative Affect



PO

0

Ø

0





PO

0

Ø

0



Stage of Addiction	Shifting Drivers Resulting from Neuroadaptations					
Binge and intoxication	Feeling euphoric	1	Feeling good	1	Escaping dysphoria	
Withdrawal and negative affect	Feeling reduced energy	5	Feeling reduced excitement	5	Feeling depressed, anxious, restless	
Preoccupation and anticipation	Looking forward	E>	Desiring drug	1	Obsessing and planning to get drug	

5

Ø



- Two factors in dopamine release in Nucleus Accumbens
 - Glutamate that drives dopamine release
 - Auto-receptor ("brakes") that shut down further release when dopamine concentrations become excessive
- Opioid use initially bypasses these "brakes" leading to large amount of dopamine release
 - With repeated use brain increases the number and strength of these "brakes"
 - Reduces resting dopamine levels- leading to dysphoria



- Memories or environmental cues that lead to drug wanting or craving
 - Craving can increase glutamate activity which drives the resting dopamine levels higher
 - Glutamate can also drive resting noradrenaline levels higher potentially worsening withdrawal symptoms → increased craving



Biologic and Social Factors

- Family history
- Early exposure to drug use (particularly adolescence)
- High risk environments (e.g. easy access to drugs, normative attitudes towards drug taking, high stress environment, lack of familial support)
- Mental illness (e.g. mood disorder, ADHD, anxiety)



- Help prevent relapse while the brain is healing and emotional and decision making capacities are being restored
- Medication assisted treatment play an essential role in reducing the three main stages of addiction
 - Dosed appropriately to not provide euphoria
 - Prevent withdrawals
 - Prevent cravings
- Usually medication assisted treatment is given in conjunction with psychotherapy



Ø

- Brief motivational models
- Contingency management models
- Cognitive behavioral therapy



Important to Remember



- Substance use disorder is a chronic brain disorder from which people can and do recover.
- Addictive substances can lead to dramatic changes in brain function and reduce a person's ability to control his or her substance use.



Dehumanization of the individual based on their social identity or participation in a negative or an undesirable social category.

Four Identified Types of Stigma

- Public
 - Endorsement by the public of negative attitudes against a specific stigmatized group, which manifests in discrimination towards individuals belonging to that group.
- Perceived
 - Stigmatized individuals think that most people believe common negative stereotypes about individuals belonging to the same stigmatized category.
- Enacted
 - Direct experience of discrimination and rejection from members of the larger society.
- Self
 - Negative thoughts, feelings, and diminished self-image resulting from identification with the stigmatized group and anticipation of rejection from the larger society.

Consequences of Stigma

2

- People with substance use disorders are viewed more negatively than people with physical or psychiatric disabilities.
- The terminology often used can suggest that substance use disorders are the result of a personal failing/choice.
- The term "abuse" is highly associated with negative judgments and punishment.
- Even trained clinicians are likely to assign blame when someone is called a "substance abuser" rather than a "person with a substance use disorder."
- Negative attitudes among health professionals have been found to adversely affect quality of care and subsequent treatment outcomes.

Language Matters

- American Society of Addiction Medicine and others have recommended the adoption of clinical, non-stigmatizing language for substance use.
- "Person-first language" has been widely adopted by professional associations to replace negative terms that have been used to label people with other health conditions and disabilities.
- "Person with a mental health condition" or "person with a disability" carry neutral rather than pejorative connotations, and distinguish the person from his/her diagnosis.

Person with a Substance Use Disorder

2

- Person-first language is the accepted standard for discussing people with disabilities and/or chronic medical conditions.
- Use of the terms "abuse" and "abuser" negatively affects perceptions and judgments about people with substance use disorders.
- Terms such as "addict" and "alcoholic" can have similar effects.

Person in Recovery

- Various terms are used colloquially to label the people with SUD, including the terms "clean" and "dirty."
- Instead of "clean,"
 - "negative" (for a toxicology screen)
 - "not currently using substances" (for a person)
- Instead of "dirty," the term
 - "positive" (for a toxicology screen)
 - "currently using substances" (for a person)
- The term "person in recovery" refers to an individual who is stopping or at least reducing substance use to a safer level, and reflects a process of change.

Medication-Assisted Treatment

- Terms "replacement" and "substitution" have been used to imply that medications merely "substitute" one drug or "one addiction" for another. This is a misconception.
- The dosage of medication used in treatment for opioid use disorder does not result in a "high," rather it helps to reduce opioid cravings and withdrawal.
- "Medication-assisted treatment" (MAT) is used to refer to the use of any medication approved to treat substance use disorders and may be combined with psychosocial support services.

Summary

- Opioids impact the brain in many different ways that can lead to addiction
- Opioid tolerance develops very quickly requiring higher doses of opioids for the same effect
- Opioid use can quickly lead to dependence and withdrawal
- Three main stages of addiction:
 - Binge/intoxication
 - Withdrawal/negative affect
 - Pre-occupation/craving
- Treatment can give the brain time to heal and restore neural circuitry

Know The Facts









2		

2

Effect of Opioid Use Disorder on Brain Chemistry

David Butterfield, PharmD, BCPS, BCPP Clinical Pharmacy Specialist-Psychiatry