OTHER DRUGS OF ABUSE

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Name	Commercial Interests	Relevant Financial Relationships: What Was Received	Relevant Financial Relationships: For What Role	No Relevant Financial Relationships with Any Commercial Interests
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Hallucinogens



Definition

- Produce alterations in thought, mood and perception
- Produce minimal autonomic side effects or craving
- Fail to produce excessive stupor or central stimulation



"Illusionogen"

- Illusions = alteration or enhancement of existing sensory perception
- May be more accurate term
 - Reality testing is generally intact
 - Effect varies greatly with expectations and environment



Question #1

Classical Hallucinogens increase the activity of which of the following neurotransmitter systems?

- A. Dopamine
- B. Serotonin
- C. Acetylcholine
- D. Gamma-amino-butyric acid (GABA)
- E. Glutamate



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Classical Hallucinogens

- 5HT_{2A} agonists or partial agonists → Serotonergic Hallucinogens
- 2 subclasses of arylalkylamines
 - Indolealkylamines
 - Bind at multiple receptors (5HT_{2A}, 5HT_{2B}, 5HT_{2C}, 5HT_{1A})
 - Phenylalkylamines
 - Fairly selective for 5HT_{2A}
 - Not all arylalkylamines are hallucinogenic
 - Stimulants
 - Empathogens



Effects of Hallucinogens



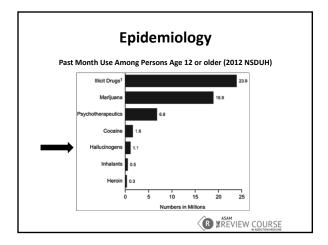
- Somatic
 - Dizziness
 - Weakness
 - Tremors
 - Nausea
 - Drowsiness
 - Paresthesias
 - Blurred Vision
- Perceptual
 - Altered shapes and colors
 - Difficulty focusing on objects
 - Heightened sense of hearing
 - Synesthesias
 - Stimulation of one sensory pathway leads to experiences in a second pathway (tasting colors, seeing music)

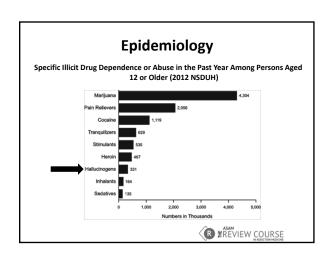


Effects of Hallucinogens

- Alterations in mood
- · Tension and anxiety
- Distorted sense of time
- Difficulty expressing thoughts
- Depersonalization
- Dreamlike feeling
- Visual hallucinations or illusions

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Idolealkylamines

- DMT (N,N-Dimethyltryptamine)
 - Prototype of this subclass
 - Naturally occurring (plants, toad)
 - Rapid onset (<5 min), short duration of action (30 min)
 - Characteristically unpleasant odor
 - Routes of administration include inhalation (smoking) and injection (rare)
 - Can be taken orally (component of ayahuasca)



Ayahuasca

- Brew containing DMT, MAOis, and other hallucinogens
- Used ceremonially in parts of the Amazon and in some Native American religions
 - Legalized for religious use among Native Americans in the US



Idolealkylamines

- Psilocybin, Psilocin
 - Found as naturally occurring tryptamine in certain varieties of mushrooms (shooms, shrooming)
 - Detachment from reality
 - Inability to discern fantasy from reality
 - Can lead to panic attacks, psychosis
 - Rapid tolerance to effects
 - Cross tolerance with LSD



Idolealkylamines

- · Psilocybin, Psilocin
 - Reported mystical-like experiences
 - Inner peace, patience, optimism, self-confidence
 - Adverse Effects
 - Nausea, vomiting, anxiety
 - May interact with MAOi
 - Duration: 4-6 hours



Idolealkylamines

- LSD (lysergic acid diethylamide) aka acid
 - Highly potent
 - Not naturally occurring
 - Schedule I
 - Water soluble, clear white odorless crystals
 - Distributed on sheets of thin blotter paper with dried solution of LSD (most common)
 - Breath mints, sugar cubes (dropping acid)
 - Pressed into pills (microdots) or thin gelatin squares (window panes)



LSD

- Onset: 30-60 min, Peak: 2-4 hours, Duration: 8-12 hours
- Effects
 - Perceptual
 - Altered shapes and colors, heightened sense of hearing
 - Somati
 - Nausea, blurred vision, dizziness
 - Psychic
 - Depersonalization, visual hallucinations, alterations in mood



Idolealkylamines

- Beta-carbolines
 - Harmaline, Harmine
 - Not technically classical hallucinogens
 - Lack 5-HT_{2A} agonist effects although they do bind at receptor
 - Act as MAOi
 - Poorly studied
 - Naturally occurring (South American plants)
 - Can be component of ayahuasca



Phenylalkylamines

- Largest group of classical hallucinogens
 - Mescaline, Peyote
 - DOM, DOB, DMA, MDA, Nexus (2-CB)
 - Modifications of mescaline-like substances



Mescaline/Peyote

- Buttons from top (crown) of peyote cactus
 - Buttons are removed & dried, then chewed or soaked in water
 - 6-10 buttons for intoxication
 - Use legalized within Native American Church
- Slow onset (30-60 min)
 - First hour
 - Minor perceptual changes, increased resp rate, nausea
 - Next several hours (5-10)
 - Visual illusions/hallucinations
 - Synesthesias



DOM

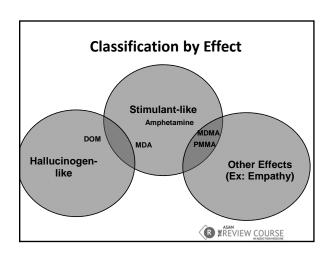
- Results from structural modification of mescaline-like substances
- · Extremely potent
- Used as model hallucinogen in drug discrimination studies



MDA

- Produces stimulant and hallucinogenic effects
 - Similar to combined effects of cocaine and LSD
- Can be modified to MDMA (ecstasy)
 - Stimulant effects
 - Empathogenic
- Has been represented and sold as MDMA





Salvia



- Herb found in southern Mexico and South America used in healing rituals
 - Traditionally ingested by chewing fresh leaves or by drinking extracted juices
 - Sometimes smoked when used as drug of abuse
- Active ingredient in Salvia is salvinorin A, a kappa opioid agonist



Salvia

- Not currently regulated by Controlled Substances Act or scheduled by the DEA
 - DEA has listed Salvia as a drug of concern
 - Several states and countries have passed legislation to regulate its use
 - May be classified as Schedule I



Salvia Effects

- · Intense and short-lived
 - Onset < 1 minute
 - Duration < 30 minutes
- Changes in visual perception
- Increased sense of well-being
- Feelings of detachment
- Modified perception of external reality and the self
 → decreased ability to interact with surroundings



Hallucinogen Intoxication

- Clear Sensorium
- Intact Memory
- Hyperalert
- EEG = arousal
- Intact reality testing
 - Can sometimes be reasoned with or calmed by talking
- Visual Hallucinations >> Auditory



Hallucinogen Persisting Perception Disorder (HPPD)

- Re-experiencing of perceptual symptoms experienced while intoxicated following cessation of use = flashbacks
- Unrelated to dose or number of exposures
- Usually resolves within 1-2 years of last use
- Can be triggered by other substance use



Dissociatives

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Definition

- NMDA receptor antagonists
 - Glutamate activates NMDA receptors to filter sensory stimuli
 - Dissociatives noncompetitively block NMDA receptors → sensory overflow



Members of the Class

- Arlcyclohexylamines
 - PCP
 - Ketamine
- Dextromethorphan
- Nitrous Oxide

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Effects

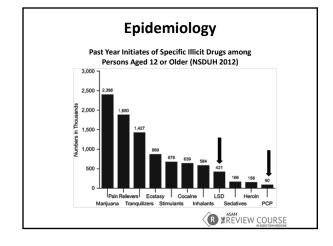
- Dissociation
- Sensory isolation
- Mental Distortions
- Increased HR, BP, Temp

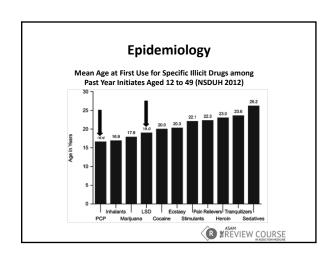
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Epidemiology

- Men > Women
- More common in large urban areas
- Often used in combination with alcohol or other illicit substances

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Phenylciclidine (PCP, angel dust)

- Developed as IV anesthetic
 - No longer FDA approved
 - Associated with prolonged delirium
 - Schedule I
- Risk of seizures or death
- Available as powder, tablets, liquid, and sprayed onto plant leaves and then smoked



PCP

- Psychotomimetic = Models psychosis
 - Positive Symptoms
 - Delusions, hallucinations
 - Negative Symptoms
 - · Blunted affect, asociality
 - Individuals with schizophrenia are more susceptible to prolonged psychosis after use



PCP Intoxication

- Nystagmus (rotary, vertical, horizontal)
- Hyperreflexia
- HTN
- · Feelings of invulnerability
 - Agitation, violence



Management of PCP Intoxication

- Low stimulus environment
- Check serial CPKs
- Risk of rhabdomyolysis
- Can progress to renal failure
- · Acidification of urine
 - Increases renal clearance, but not advised bc inc. urinary myoglobin
- Benzos, Antipsychotics as needed



PCP Effects

- · Vary widely with dose
 - Confusion, delirium, psychosis



- Semi-coma and coma (less common)



Coma with seizures (rare)



Ketamine (K, Special K)

- FDA approved for general anesthesia in animals and humans
- Schedule III
- Administered as IV or IM in medical settings
- Abused by inhalation, smoking, or oral administration
- Less potent, shorter acting than PCP



Ketamine

- Clinical Effects
 - Analgesia
 - Dissociation
 - Amnesia
 - Delirium (higher doses)
 - Long term
 - Dysphoria, memory impairment, apathy, irritability



Intoxication and Overdose

- Nystagmus (vertical and/or horizontal)
- Cardiovascular and renal complications
- Numbness of entire body
- Spacey feeling
- · Rare visual hallucinations



Dextromethorphan (DXM)

- · Widely available in over-the-counter cough medicines
 - Capsules, tablets, lozenges, syrup
 - Anti-tussive dose <120mg Daily
 - 300-1800mg produce PCP-like effects



Dextrorphan (DXO)

- DXM is metabolized to dextrorphan (DXO) , an NMDA antagonist
 - Weaker sigma opioid agonist and stronger NMDA antagonist than DXM
 - Relatively inactive at mu, kappa and delta opioid receptors
 - Lacks conventional opioid properties
 - Respiratory depression at massive doses



Effects of Dextromethorphan

- Drowsiness, Dizziness, Blurred Vision, Slurred Speech
- Euphoria and Hallucinosis
- N/V, Hypertension, Diaphoresis
- Significant Serotonergic Properties
 - Increase synthesis & release
- Inhibit reuptake
- Risk of serotonin syndrome



Dextromethorphan (DXM)

- Significant serotonergic properties
 - Increase synthesis and release of serotonin
 - Inhibit reuptake
 - Risk of serotonin syndrome



Dextromethorphan (DXM)

- Popular with children and adolescents
- Perceived as "SMART" choice
 - Stigma
 - <u>M</u>oney
- Access
- **R**isks
- <u>T</u>esting



DXM Overdose

- Deaths have been reported with large doses (reports of 200x recommended dose)
 - CNS & Respiratory Depression, Seizure, Arrhythmias
- Additional ingredients in cough medicine increase hazards
 - Pseudoephedrine
 - Antihistamines
 - Acetaminophen



Dissociative Intoxication & Overdose

- PCP included on most screening panels, but need special testing (GCMS) for ketamine, DXM
- Increased serum CPK & urine myoglobin
- Rarely see dilated pupils
 - Different from stimulants, hallucinogens, opioid withdrawal
- Visual hallucinations relatively rare



Inhalants



Question #2

All of the following are true regarding inhalant use except:

- A. Many inhalants are legally available.
- B. Inhalants cause short and long term damage to multiple organ systems.
- C. Inhalant use increases with age.
- D. Onset of action is rapid and duration is short.
- E. Sniffing, huffing and bagging are methods of abusing inhalants.



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Inhalants

- Breathable chemicals that can be self-administered
- Deliberately inhaled to cause a high
- Also known as
 - Whippets
 - Poppers
 - Huff
 - Bang
 - Kick
 - Sniff



Terminology

- Sniffing = inhaling from an open container
- Huffing = holding fabric soaked in substance to the nose or mouth and inhaling
- Bagging = concentrating vapors in a bag and inhaling



Types of Inhalants

Volatile Solvents

- Adhesives
- Aerosols
- Solvents
- Cleaning agents
- Aromatic Hydrocarbons (Toluene, Xylene)
 - Highly abused due to intense euphoric effects



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Types of Inhalants

Anesthetics

- Ethyl ether
- Chloroform
- Methyl chloride
- Trichloroethylene
- Chloral hydrate



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Types of Inhalants

Nitrous oxide

- Dissociative Anesthetic Gas
- Widely available
- Directly depresses ventilation and myocardial function
 - Use of nitrous can result in:
 - Hypoxia (displacement of O₂)
 - Hypotension & arrhythmias
 - Depression, psychosis, memory loss
 - Peripheral neuropathy
 - Bone marrow suppression

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Types of Inhalants

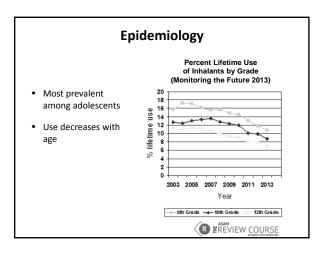


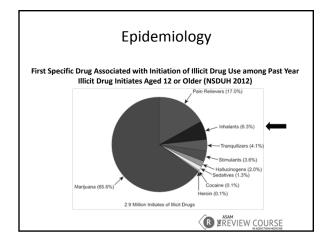
Amyl-, Butyl-, Cyclohexyl- and Isobutyl- Nitrites

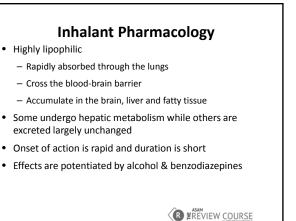
- Amyl nitrite was originally used as a vasodilator to relieve angina
- Sold in ampules (bulbs) that can be broken to release the vapors
 the sound that resulted caused them to be referred as poppers
- Reputed to act as a sexual enhancer (causes smooth muscle relaxation)



Sources of Inhalants POSSIBLE CONTENTS PRODUCT Amyl, butyl, cyclohexyl nitrite; butane Butane Lighter fluid n-Hexane, tetrachloroethylene, xylene Benzene, toluene, xylene, (lead) Household cleaners Gasoline Butane, propane Hair spray Mothballs Naphthalene, paradichlorobenzene Acetone, toluene Toluene, trichloroethylene, xylene Nail polish remover Paint thinner Xvlene Markers Refrigerant Acetone, benzene, n-Hexane, toluene Rubber cement Butane, propane, toluene Amyl, butyl, cyclohexyl nitrite Spray paint Video head cleaner R EREVIEW COURSE







Abuse Liability

- Quick acting
- Short duration
- Free or low cost
- Easily available
- Generally not prosecuted
- Difficult to test for
- Not perceived as dangerous



Inhalant Presentation

- · Prototypical patient
 - Adolescent male
 - Odor of paint or solvents
 - "Glue Sniffer's Rash" around nose & mouth
 - Conjunctival irritation
 - Cough, nasal discharge, dyspnea, rales, rhonchi



Acute Effects of Inhalants

- Euphoria
- Disinhibition
- Dizziness / lightheadedness
- · Slurred speech
- Ataxia
- Drowsiness
- Increased incidence of accidents and injuries



Toxic Effects & Overdose

- · Respiratory depression
- Arrhythmias
- Chest pain
- Vague muscle & joint pains
- · Asphyxia, cardiac arrest and death can occur



Sudden Sniffing Death

- Acute cardiotoxicity in otherwise healthy users
- Occurs when intoxicated individual engages in strenuous activity
- · Increased sensitivity of myocardium to norepinephrine
- · Most common with butane or CFCs



Chronic Effects of Inhalants

- Cardiac
 - arrhythmia
 - cardiomyopathy heart block
- Pulmonary
 - pneumonitis
 - emphysema
- hypoxia
- aspiration pneumonia Dermatological
- - perioral infection - rash

- Gastrointestinal
 - hepatitishepatorenal failure
- Genitourinary
 glomerulonephritis
 - Goodpasture's syndrome hypokalemia
- Hematopoietic
 - aplastic anemia
 - leukemia
- bone marrow suppressionMusculoskeletal



Neurological Effects of Inhalants

- Peripheral neuropathy
- Trigeminal neuralgia Optic neuritis
- Delirium
- Dementia
 - Decreased attention
 - Impaired learning and
 - information processing
 - Apathy
- Poor memory
- Impairment in working memory and executive functioning
- Cerebellar atrophy
 - Tremors
 - Ataxia
 - Gait abnormalities
- Irreversible white matter changes
 - Most common with toluene (aromatic hydrocarbon)
 - Correlate with lower
 - performance IQ scores



Fetal Effects of Inhalants

- Inhalants cross the placenta
- · Low birth weight
- Facial and limb abnormalities
- Microcephaly
- Developmental delays

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Treatment Considerations

- User may experience prolonged residual effects because chemicals are stored in fatty tissue
- Neurological impairment is often present and cognition should be continually assessed
- Talk therapy may not be appropriate for patients with neurological or cognitive dysfunction
- Short attention span, poor impulse control, and poor social skills may not be appropriate for group therapy



Steroids



Question #3

- Which of the following is incorrect regarding steroid use:
 - The term anabolic refers to the smooth muscle building properties of steroids.
 - B. The term androgenic refers to the masculinizing properties of
 - C. Anabolic-androgenic steroids can be legally prescribed.
 - Cycling, stacking and pyramiding are terms used to describe illicit steroid use.
 - E. Liver damage is a known complication of steroid use.



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Anabolic - Androgenic Steroids

- Anabolic = skeletal muscle-building
- Androgenic = masculinizing
- Includes testosterone and >100 related synthetic substances

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History

- 1935: researchers discovered that testosterone increased muscle mass
- 1940s 1950s: used to improve performance in sports
- 1950s: brought to US for use in weightlifting
- 1975: banned in Olympics
- 1990: U. S. Federal Law reclassified all anabolic steroids as controlled substances, schedule III



Intended Use

- Can be legally prescribed
 - Steroid hormone deficiency
 - · Delayed puberty
 - Loss of lean muscle mass
 - Cancer
 - HIV
 - Can lead to generalized muscle hypertrophy
 - Especially in arms, shoulders, neck and chest



Misuse

- AAS are abused in an attempt to enhance performance and/or improve physical appearance
 - May be taken at 10-100 times the intended dose
- · Routes of Administration
 - Ora
 - Anadrol (oxymetholone), Oxandrin (oxandrolone), Dianabol (methandrostenolone), Winstrol (stanozolol)
 - _ IM
 - Deca-Durabolin (nandrolone decanoate), Durabolin (nandrolone phenpropionate), Depo-Testosterone (testosterone cypionate), Equipoise (boldenone undecylenate)









Terminology

- Cycling
 - Steroids taken for weeks or months alternating with rest periods
- Stacking
 - Combination of several different types of steroids and/or routes of administration
- Pyramiding
 - Slow escalation of steroid use (increasing the number, dose and/or frequency), with peak amount at mid-cycle and tapering toward the end of the cycle



Sources of Illicit Steroids

- Often sold at gyms, competitions, and through mail order operations
 - Most illegal steroids are smuggled from countries that do not require a prescription for the purchase of steroids
 - These "steroids" may actually contain inactive products that do not have anabolic properties
- Illegally diverted from U.S. pharmacies or synthesized in amateur laboratories
- Obtained by Rx under false pretenses

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Epidemiology

- 3 most common populations
 - Athletes
 - Used for performance enhancement
 - Aesthetes
 - Used to improve physical appearance
 - Most common among adolescents
 - Fighting Elite
 - Used to increase aggression and/or job performance (security, law enforcement)



Medical Side Effects

- Acne
 - Especially on face, shoulders and back
- Liver damage
 - Jaundice
 - Hepatomegaly
 - Liver cysts
- Fluid Retention

- Joint Pain
- Increased LDL, decreased HDL
- Injectable
 - Risk of infection, abscess, HIV, Hep C, endocarditis



Gender-Specific Side Effects

Women

Men

- · Deepening of voice
- · Testicular atrophy
- · Facial hair
- Prostatic Hypertrophy
- Menstrual changes
- Gynecomastia
- Male-pattern baldness
- Baldness
- · Genital hypertrophy
- Infertility



Psychiatric Side Effects

- Aggressive / violent behavior
 - "Roid Rage"
- Hypomania or Mania (high doses)
- Paranoia
- · Extreme irritability
- Impaired judgment
- Delusions

- Treatment
 - Remove AAS
 - Use mood stabilizers or anti-psychotics as needed
 - Generally resolves within 1-2 weeks after cessation of AAS use in psychiatrically healthy individuals



Withdrawal Syndrome

- · Steroid Withdrawal Associated Depression
 - Depressed mood
 - Fatigue
 - Restlessness
 - Anorexia
 - Insomnia
 - Decreased libido
 - Rarely, suicidal ideation
- Can be responsive to SSRIs



Associated Syndromes

- Comorbid substance abuse, especially opioid abuse/dependence
 - Opioids used to counteract insomnia and irritability resulting from the steroids and pain from excessive exercise
- Body Dysmorphic Disorder / Muscle Dysmorphia
 - Perception of self as small and frail
 - Compulsive weight-lifting / body building
 - Avoidance of beach, locker room, showers



Treatment

- Rarely seek treatment
 - Steroids are not euphorigenic
 - No immediate high
 - Goal is long-term reward associated with physical changes
 - May be seen as socially acceptable or positive
 - Help to achieve idealized male form
 - Some have little respect for doctors
 - Feel they have no understanding of bodybuilding



Summary

- Hallucinogens
 - DMT, Ayahuasca, Psilocybin, LSD, Harmaline, Mescaline, DOM, MDA, Salvia
- Dissociatives
 - PCP, Ketamine, Dextromethorphan, Nitrous Oxide
- Inhalants
 - Volatile Solvents, Anesthetics, Nitrous Oxide, Nitrites
- Steroids



References

- Principles of Addiction Medicine: The Essentials
- Monitoring the Future
 http://www.monitoringthefuture.org
- National Survey of Drug Use and Health <u>https://nsduhweb.rti.org/</u>

