



Hi my name is Dr. Brenda Nordstrom, and this presentation is on the relationship be tween adverse childhood experiences other traumas in pain management.

I have a bit of an eclectic nursing background and I am currently teaching nursing w hile also working as a relief nurse in home hospice. However, what peaked my inter ested in connecting trauma with pain management was my work with human traffic king survivors. I am a strong advocate for trafficked persons and volunteer for a loc al service provider and I am a healthcare trainer for my regional human trafficking t ask force. The more I interacted with not only human trafficking survivors but other s who have experienced adverse childhood experiences and other trauma, I realized that nurses needed to learn about and understand the link between the mental and physical aspects of the pain experience.

When I first ventured into providing education about trauma and pain it was specifically for hospice nurses. However, I think the information is very applicable for all nursing specialties.



# Early History Native to Central Asia since prehistoric times Fiber, food, medicine, rituals Found with ancient artifacts 2,500-year-old "Siberian Ice Princess" Chinese cannabis medicine began 1,800 years ago

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The cannabis plant is native to Central Asia since prehistoric times, and it quickly became will know as it traveled west. It originated in the Kush Himalayan foothills then migrated across the globe, primarily due to the herb's role as a fiber and psychoactive flower. Neolithic people had many uses for the cannabis plat during ancient times. It provided fiber for textiles, and it was a food source. It also quickly became a part of medicinal concoctions as was part of ritual preparations. It is hypothesized that people began smoking cannabis around the 5<sup>th</sup> century B.C. as part of burial ceremonies.

In 1993, a very fascinating find was made in the mountains of Russia— a well pre served mummy who was known as the "Siberian Ice Princess". She was a 2,500-y ear-old mummy who was buried with cannabis. Scientists' examination revealed that she suffered from metastatic breast cancer and other ailments. They believ ed she was medicating with cannabis and therefore it was buried with her. As China was the epicenter of hemp cultivation — both textile and medicinal vari eties — it is no wonder that written evidence demonstrates that the Chinese peo ple used cannabis for medicinal purposes more than 1,800 years ago. They eve n developed a unique specialty called Achene (a-ken) that utilized the seeds of the cannabis flower. Cannabis seeds were also consumed as food during ancient Chinese times.

## **Early History**

- Shen-Nung (2,700 BC) emperor and medicine man
  - More than 100 ailments
  - Pain, malaria, mental illness, cough, seizures, insomnia
- Hua T'o (approx. 117-207 AD) founder of Chinese surgery
  - "Ma-yo" = boiled hemp compound, mixed with wine
  - Abdominal & other surgeries





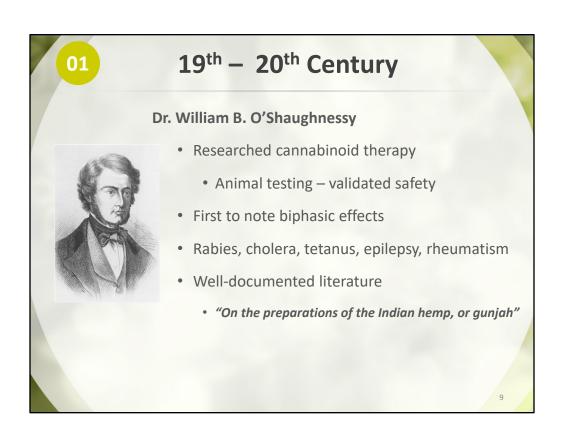
Eventually, the cannabis plant migrated around the globe. Starting with India, th en to the Middle East, Central Asia, Africa, Europe, & then ultimately finding it's way to the Americas

## Early History --> 1800s

- Colonial-era settlers required to grow hemp
- 1600s Virginia Assembly law
- Hemp & cannabis used for sailing
  - Rigging, sails, rope, & textiles



Medicinal cannabis developed reputation as curative



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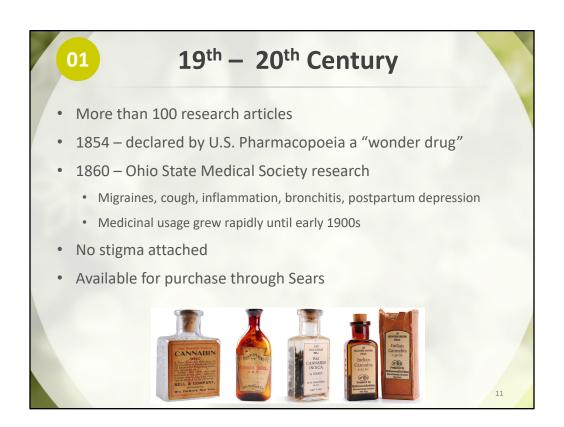


## 19<sup>th</sup> – 20<sup>th</sup> Century

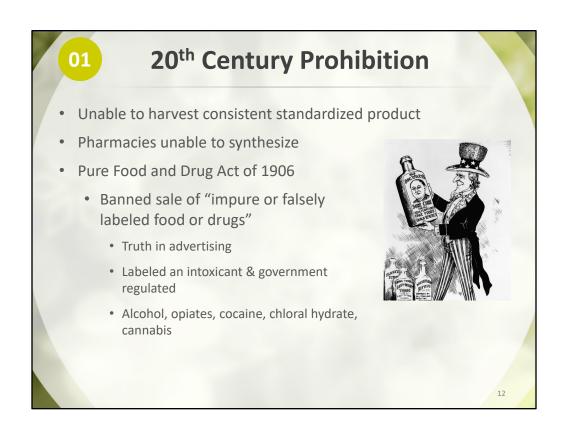
## Dr. J. R. Reynolds

- Physician to Queen Victoria of England
- Cannabis for menstrual cramps & sleep
- Published in *The Lancet* 
  - "Therapeutic uses and toxic effects of Cannabis Indica"
- Prescribed cannabis >30 years
  - Insomnia, dysmenorrhea, neuralgia, tics, & spasms
- Advised "start low, go slow"

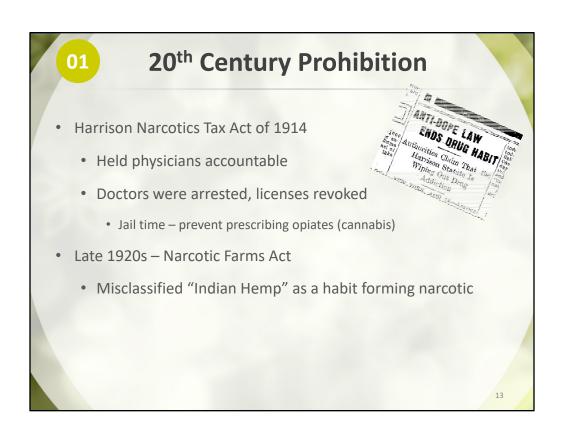




Pg 11-12



Pg 11-12 Required listing all ingredients on the label – cocaine in coca cola



Pg 14-15

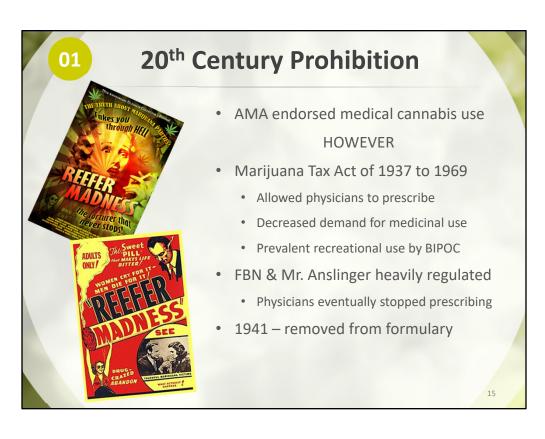
# **20<sup>th</sup> Century Prohibition**

- 1930s Federal Bureau of Narcotics (FBN)
  - Harry J. Anslinger director, alcohol prohibitionist enforcer
  - Late 1930s cannabis prohibition replaced alcohol

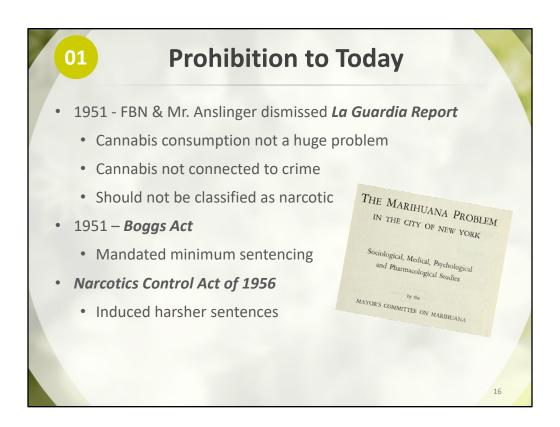


MARIJUANA
IS THE MOST
VIOLENCECAUSING DRUG
IN THE HISTORY
OF MANKIND

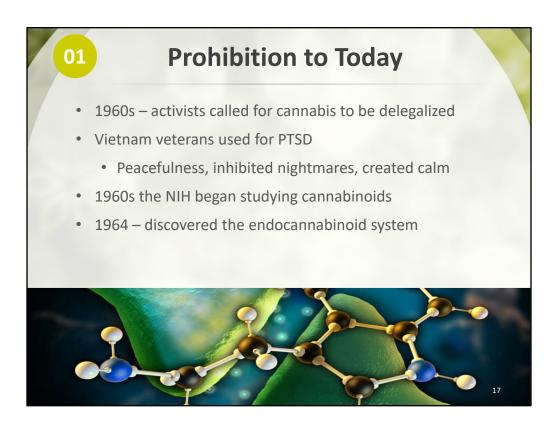
- HARRY J. ANSLINGER -



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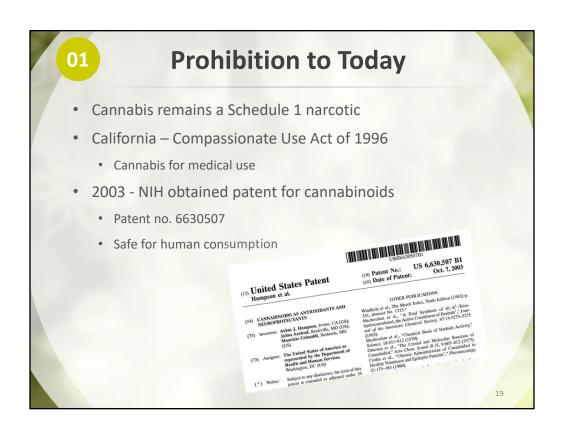


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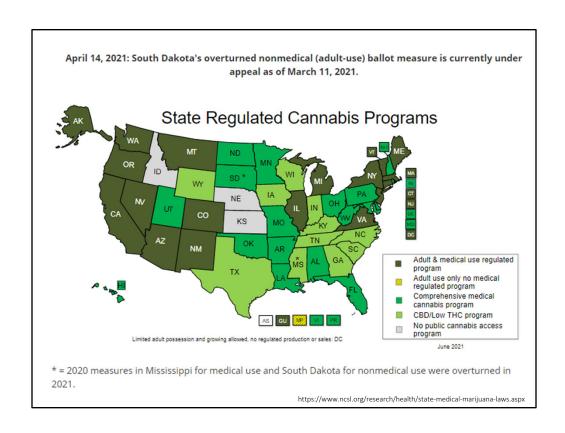


Pg 19-21 Dr. Machoulam

# Prohibition to Today 1970 – Controlled Substances Act Established five drug schedules Placed cannabis in Schedule I Past research deemed "folk medicine"



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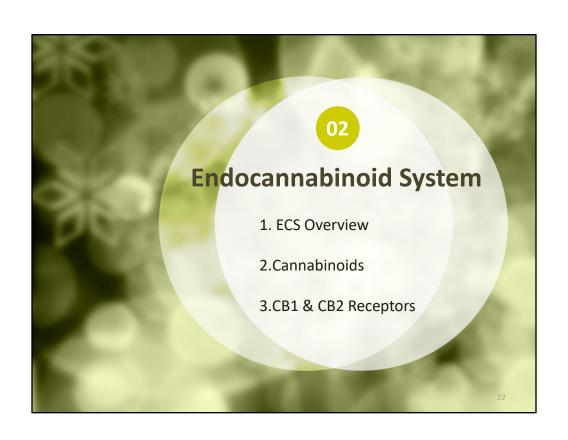


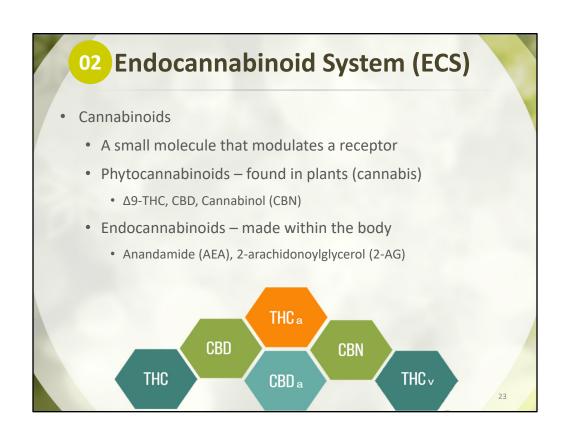


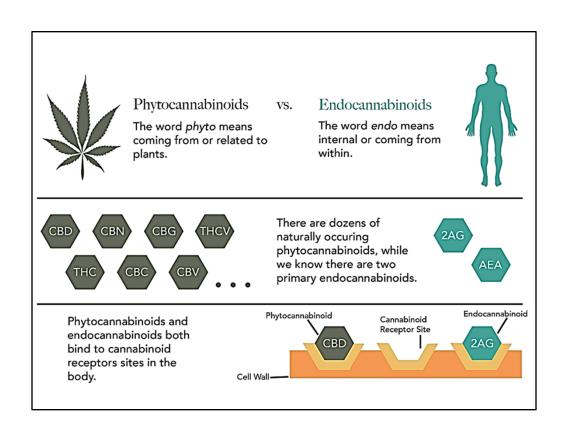
MICHIGAN REGULATION AND TAXATION OF MARIHUANA ACT - Initiated Law 1 of 2018

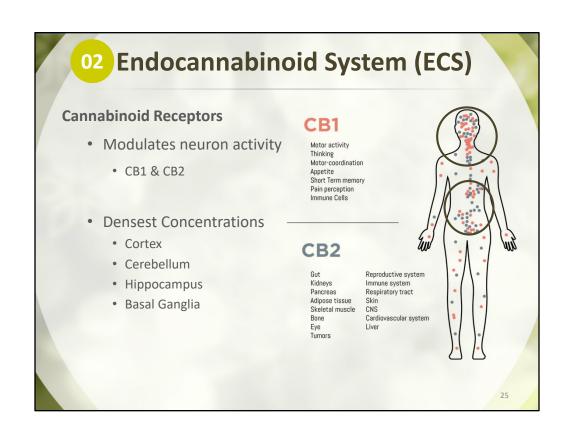
An initiation of legislation to allow under state law the personal possession and use of marihuana by persons 21 years of age or older; to provide for the lawful cultivation and sale of marihuana and industrial hemp by persons 21 years of age or older; to permit the taxation of revenue derived from commercial marihuan a facilities; to permit the promulgation of administrative rules; and to prescribe c ertain penalties for violations of this act. If not enacted by the Michigan State Le gislature in accordance with the Michigan Constitution of 1963, the proposed le gislation is to be voted on at the General Election, November 6, 2018. https://www.legislature.mi.gov/(S(s4fcgz5wtdxhdedbievvhv1d))/mileg.aspx?page=shortlinkdisplay&docname=mcl-Initiated-Law-1-of-2018

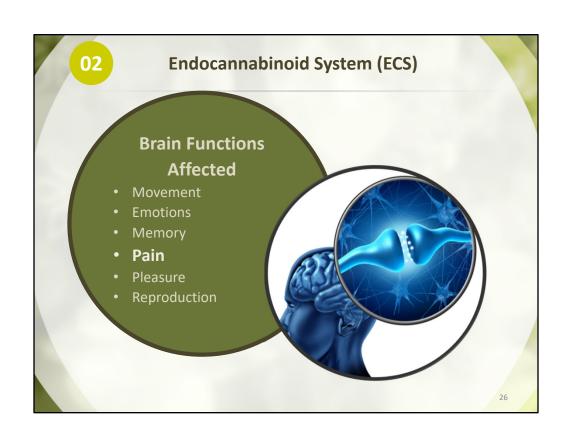
local options to ban the sale of recreational marijuana

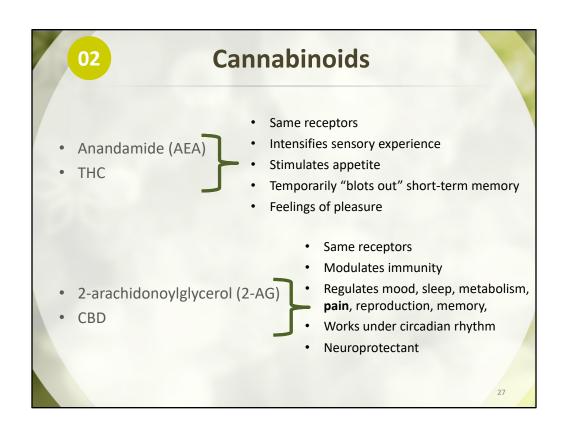












Pain receptors Perception of pain Sleep Anxiety Quality of life





For example, people who use marijuana recreationally often smoke it to achieve a high, rather than to ease chronic pain or other conditions.

THC is the mind-altering chemical in marijuana primarily responsible for produci ng euphoria. THC-based medications are often used to increase appetite and re duce nausea. The chemical is also used to decrease pain, inflammation and mus cle control problems. That being said, not all chemicals in marijuana produce a e uphoric effect. Many medical cannabis patients use the drug only for its therape utic properties, aiming to minimize marijuana's side effects, including the high. This is where cannabidiol, or CBD, is significant.

Cannabidiol is non-psychoactive: it does not have mind-altering effects. It also minimizes the euphoric effects of marijuana caused by THC. CBD can be useful in reducing pain and inflammation, as well as controlling epileptic seizures. It may also be effective in treating mental illness or addiction, according to the National Institute on Drug Abuse.

Recreational marijuana generally contains high levels of THC and lower levels of CBD. Medicinal marijuana is often rich in CBD, with less or no THC. Therefore, m any medical marijuana users will feel the therapeutic effects without a high.

## **Medical Marijuana**

## **Medical Marijuana – Most common qualifying conditions:**

- ALS
- Cachexia
- Chronic pain
- Cancer pain
- Crohn disease/irritable bowel syndrome
- Epilepsy/seizures
- HIV/AIDS
- Multiple sclerosis
- Persistent muscle spasms
- PTSD



The National Academies of Sciences, Engineering, & Medicine (2017) The Health Effects of Cannabis and Cannabinoids: The Current State of Evidence and Recommendations for Research

## 03 Cannabis for Pain Management

### **Current State of Evidence – Israel**

- Chronic Pain
- Spasticity multiple sclerosis or paraplegia
- Sleep disorders
- Psychosis
- Anxiety disorders
- Crohn's disease



44% decrease in opioid consumption



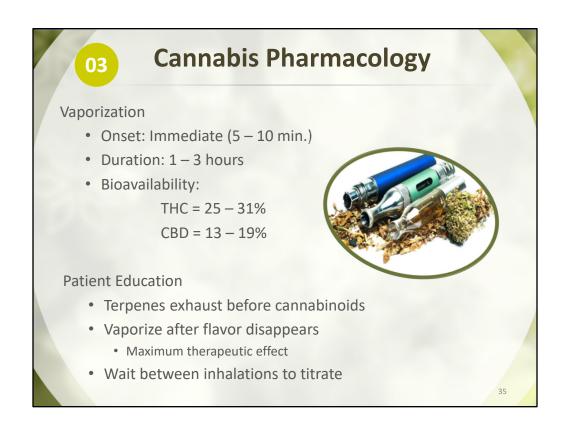
### Fast facts

Flower may not contain high enough concentration of cannabinoids needed for some conditions. This is why gas or liquid chromatography lab testing is so important

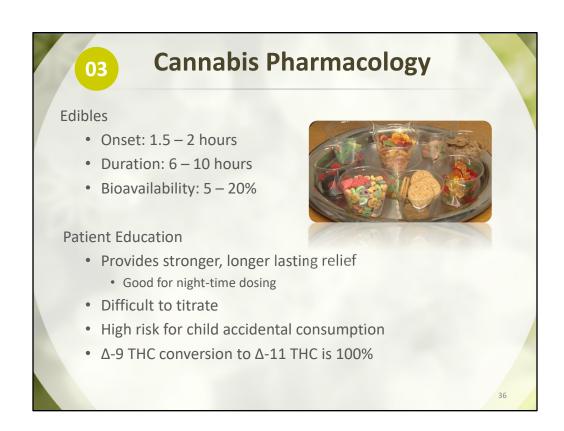
A powerful strain of cannabis contains 20% THC where as a dab can reach 70-8 0% concentration of THC

# Cannabis Pharmacology Smoking Onset: Immediate (5 – 10 min.) Duration: 2 – 4 hours Bioavailability: THC = 25 – 31% CBD = 13 – 19% Patient Education Wait 15 min between inhalations Titrate for desired effect Do not "hold" smoke Water pipe & bongs – larger doses

Waiting 15 minutes between inhalation assists with the titration to desired effect – which is to relieve the targeted symptom, not the psychoactive effect. Water pipes and bongs deliver larger doses of cannabinoids per inhale than trad itional smoking. Water can remove plant matter and tar. However, water that coo Is the smoke can trap therapeutic cannabinoids.



Vaporizers heat the oils at a lower temperature and does not burn plant matter. It is a very efficient method. Vaporizing is less likely to cause irritating respirator y side effects. Types of vaporizers include conduction (heats a metal plate), con vection (hot air circulates around the plant), and oil cartridge vapor pens. The v apor pens are the least expensive option, it is discreet and easy to use. Patients can also purchase a variety of prefilled cartridges for different dosing options.



Does not provide immediate relief

Smoking results in only 20% of delta-9 THC metabolizing into delta-11 THC, ing estion results in 100%

Overconsuming edibles can lead to accidental cannabis poisoning Candies, gummy bears, chocolates, cookies, pills, and lozenges



# **Cannabis Pharmacology**

Oral Mucosal – tinctures, sprays, sublingual drops

- Onset: 1 15 min (1.5 hours if swallowed)
- Duration: 2 8 hours
- Bioavailability: 1 12%



## **Patient Education**

- Variety of CBD:THC ratios
- Best under the tongue or between gums & cheek
- Avoid eating or drinking for 10 min



# **Cannabis Pharmacology**

Topicals – creams, salves, roll-ons, pain patches

- Onset: Immediate
- Duration: 30 min 3 hours
- Bioavailability: NA



### **Patient Education**

- Non-psychoactive unless VERY high THC component
- Good for localized pain
  - Arthritis, inflammation, eczema/psoriasis, chronic pain

# OB Side Effects of Cannabis Use

### **Short-term side effects**

- Altered senses
- · Altered sense of time
- · Changes in mood (suicide risk)
- Impaired body movement (fall risk)
- Impaired cognition & memory
- With high doses
  - Hallucinations
  - Delusions
  - Paranoia & psychosis

Anyone with severe diseases of the liver or kidneys should also take special prec aution that the metabolic breakdown of cannabinoids does not worsen their con ditions

Individuals with a history of suicide attempt or who are at risk for suicide and th ose with schizophrenia, bipolar disorder, or other psychotic condition should be informed about the risks of cannabis use and be advised to not use cannabis. The risk of suicide and cannabis use is a contentious area of study. Current findings are contradictory, and more research is needed to confirm any association be tween cannabis use and suicide risk while controlling for numerous confounding variables. Individuals with a greater risk of psychological disturbances and suicidal ideation should take precautions when using cannabis as a therapeutic. Substance-induced psychosis is most frequently the result of large doses of THC. These events are typically acute and resolve quickly.

Consuming more than the recommended dose of an edible product is the leading cause of experiencing paranoia and psychosis

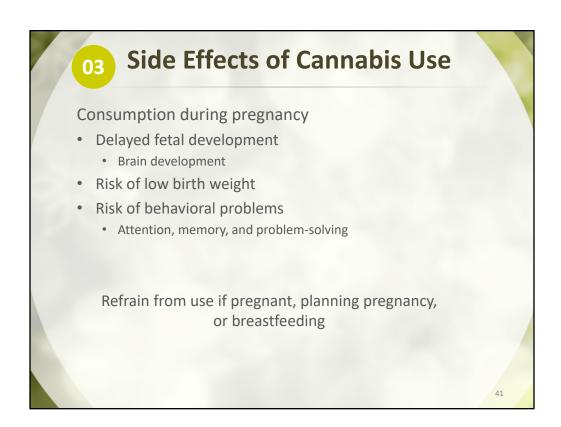
# Side Effects of Cannabis Use Long-term side effects Clinical studies are VERY limited. Physical effects Breathing problems Increased heart rate Nausea and vomiting Cannabinoid hyperemesis syndrome (CHS) Risk of stroke

Breathing problems - Smoking marijuana may irritate the lungs. If smoked frequently, the effects are similar to tobacco smoke – daily cough & phlegm with frequent respiratory illness. Those who smoke marijuana joints tend to inhale more deeply and hold the smoke in their lungs longer. Once study documented that smoking cannabis in this way can result in 4x the exposure to carbon monoxide and 3 – 5x more tar than smoking a single cigarette. However, researchers have not identified a higher risk of lung cancer. This may be due to the interaction of cannabinoids with the endocannabinoid system.

Smoking cannabis can raise the resting heart rate for up to 3 hours, dilate blook vessels , and make the heart work harder. Smoking cannabis also decreases exercise test durati on on maximal exercise tests and increases the heart rate at submaximal levels of exerci se. This could be a concern for patients with a cardiac condition.

Nausea & vomiting – regular, long-term marijuana use increases the risk of developing cannabinoid hyperemesis syndrome. CHS only occurs in daily long-term users (often ex ceeding 3 to 5 times daily for several years). Using cannabis also affects the GI tract due to the large number of CB1 receptors in this area – potentially inhibiting gastric emptying, tightening of the pyloric sphincter, and altered motility. CHS is believed to have 3 phases – pre-emetic, hyper-emetic, and recovery. It is during the hyper-emetic phase that medical treatment is often obtained in the ED. Due to these phases, CHS is often misdiagnosed as cyclic vomiting syndrome. Of interesting note – many patients with CHS report obtaining some relief of symptoms with frequent hot showers. Discontinuing cannabis use is the only treatment of CHS.

There is limited evidence of statistical association between cannabis use and the risk of having a stroke. However, the cardiovascular effects have been proposed as a possible mechanism.



Based on study of mothers who had used cannabis at least once per week befor e and throughout pregnancy versus nonusers. However, when adjusted for othe r drug use such as cocaine or opiates, there was no significant association betw een cannabis use and lowered birth rate. All studies of cannabis use and pregnancy involved women who were polysubstance drug users.



# O3 Cannabis – Drug Interactions

# Hepatic/intestinal enzyme CYP3A4 metabolism

## Increased level of drug

- Amiodarone
- **Statins**
- Pimozide
- Erythromycin
- Ketoconazole
- Verapamil
- Ritonavir
- Buprenorphine (sedation)
- Narcotics & benzos (sedation)
- Phenobarbital (sedation)

## Increased effect of cannabis

- Diltiazem
- Ketoconazole
- Ritonavir
- Buprenorphine (sedation)
- · Narcotics & benzos (sedation)

https://reference.medscape.com/ drug-interactionchecker

# Cannabis Use Disorder DSM-V diagnosis Approximately 9% adults will develop CUD Adolescent users – increases to 17% Risk factors may include Family or personal history of SUD Low socioeconomic status Traumatic childhood Sexual abuse Other mental health disorders

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Long term cannabis use can potentially lead to cannabis use disorder (CUD), esp ecially in individuals predisposed to addiction.

Cannabis dependence and withdrawal is poorly understood but seems to occur with high doses taken multiple times a day.

Withdrawal symptoms include irritability, nervousness, sleeping difficulties, dysph oria, decreased appetite, restlessness, strange or vivid dreams, and anxiety.



# **Cannabidiol - CBD**

- Synergistic relationship with THC
- Types of CBD products:
  - Full-spectrum
    - Utilizes total hemp flower with all cannabinoids & terpenes
    - THC is less than 0.3%
    - Potential for positive drug test
  - Isolate
    - Pure CBD 0% THC
    - Other cannabinoids & terpenes removed
  - Topical
    - Good for local pain targets (arthritis)

Quality varies which affects effectiveness!

03

# **Cannabidiol - CBD**

- Over 100 clinical trials as of 2019 including
  - Autism spectrum disorder
  - PTSD
  - Anxiety
  - Substance use disorder
  - Chronic pain/pain
  - Epilepsy
  - Prevention of GVHD
  - Diabetes

# **Cannabidiol - CBD** 03 Documented therapeutic effects: Anti-inflammatory Analgesic Antianxiety Antitumor Antioxidant Neuroprotectant Antidepressant Antiemetic antipsychotic • Inhibition of TNF-α (arthritis) Immunomodulatory properties Attenuation of fibrosis & myocardial dysfunction (diabetes) • Inhibition of macrophage recruitment (Crohn's disease)

TNF- $\alpha$  = tumor necrosis factor alpha (a cytokine, a small protein used by t he immune system for cell signaling; triggers a series of various inflammat ory molecules)

# Generally considered safe but may cause: Drowsiness Lightheadedness Nausea Diarrhea Dry mouth Liver damage (rare)

CBD is metabolized, among others, via the CYP3A4 enzyme. Various drugs such as ketoconazol and clarithromycin inhibit this enzyme. This leads to slower CBD degradation and can consequently lead to higher CBD doses that are longer ph armaceutically active.

The only observed side effects were reduced sucrose preference, reduced food c onsumption and body weight in some animals treated with CBD (50 mg/kg). CBD, which is nonhedonic, can reduce heroin-seeking behavior after, for example, cueinduced reinstatement.

03

# **CBD** – **Drug Interactions**

# May enhance sleepiness with:

- Opioids
- Benzos
- Antipsychotics
- Antidepressants
- Antihistamines
- Alcohol

## Increased risk of diarrhea:

- Metformin or other diabetes medications
- Prilosec or other GERD medications

**CBD** – **Drug Interactions** 03 Metabolized by cytochrome P450-complex enzymes Approximately 60% of prescribed drugs Either increases or decreases drug effect Amiodarone Fentanyl Amitriptyline levothyroxine Clindamycin Nortriptyline Clonidine Quinidine Cyclosporine Theophylline Digitoxin Warfarin \* Monitor when used with beta blockers, warfarin, statins, benzos, cardiovascular & epilepsy meds

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# **Prescriptions & Synthetics**

## **Epidiolex**

- Only FDA-approved prescription CBD
- Treatment for seizures

### Nabiximols (Sativex)

- 1:1 mix of THC and CBD
- Oromucosal spray
- Licensed in UK in 2010 (now licensed in several countries)
  - MS symptoms bladder problems, pain, muscles spasms
  - · Rheumatoid arthritis
- Investigational in the U.S.
  - · Spasticity due to MS only

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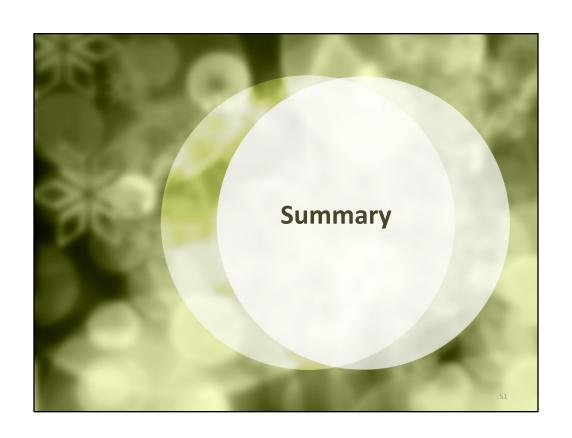
EPIDIOLEX is the first and only FDA-approved prescription CBD used to treat sei zures associated with Lennox-Gastaut syndrome (LGS), Dravet syndrome, or tube rous sclerosis complex (TSC) in patients 1 year of age and older.

### Nabilone

•Synthetic cannabinoid similar to THC

## Dronabinol

- Synthetic THC
- Marinol



# **Summary**

- Patients fear stigma with use
- Evidence of effectiveness for pain management
- "start low go slow"
- True medical use = THC + CBD
  - No psychoactive effects
- Inpatient pain management
  - Cannabis use discontinued
  - May require increased opioid dosing
- Out patient pain management
  - Cannabis use decrease opioid dose
- Does have drug-drug interactions

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