GREEN DENTISTRY: What You Need to Know About Cannabis + Oral Health



Barry Taylor, DMD, FAGD June 21st, 2019



FINANCIAL DISCLOSURE I receive nothing of value from anyone

BARRY TAYLOR DMD

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OUTLINE

1. Introduction

- 2. The Plant: Cannabis
- 3. Pharmacology
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TALKING WITH PATIENTS ABOUT RISING CANNABIS TRENDS

circa 2016

"During your talk, it is recommended that you caution your patient to avoid using marijuana at least seven days before a scheduled dental appointment, especially one in which anesthesia is required and the risk of infection is high."



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TREATMENTS

Active Ingredient In Marijuana
Reduced Alzheimer's-Like Effects In
Mice

November 7, 2018 · 2:03 PM ET

But even if that's true, she said, it doesn't mean the growing number of healthy older people who smoke pot should celebrate by lighting up. "We did this same experiment in healthy mice," she said, "and they had problems learning."

PUBMED AS OF 2019



of Publications

Year

A BRIEF CANNABIS HISTORY



A BRIEF CANNABIS HISTORY

2700 BC: Pen Ts'ao Ching – Chinese Pharmacopoeia – rheumatic pain, constipation, malaria

1000 BC: India – analagesic, anticonvulsant, anti-inflammatory, anti-biotic

1880s: Irish physician Dr.

O'Shaughnessy and French Psychiatrist

Dr. Moreau bring cannabis West



Figure 1 - Age of the beginning of cannabis use as a medicine.

A BRIEF CANNABIS HISTORY

1850: Appeared in United States Phamacopeia until 1942

1937: 'Marihuana' Tax Act: **Opposed by the AMA**

1970: Federal Controlled Substance Act: **Schedule I controlled substance** No accepted medical use High potential for abuse

1970-1972: Shafer Commission under President Nixon: **Cannabis should be decriminalized**







WAPO.ST/WONKBLOG

Source: Yahoo News/Marist survey

2017 – Yahoo/Marist study 2015 – Federal study – 33 million within past year 2016 – 37.8 million smoke cigarettes every day

MEDICAL MARIJUANA VERSUS RECREATIONAL MARIJUANA

Recreational marijuana users generally seek a high THC concentration

True medicinal marijuana users seek a high CBD concentration

No prescriptions are written, a medical marijuana user has access to purchase marijuana products

NATIONAL DRUG USE 2016

Marijuana nation

% of residents age 12 or older using marijuna in the past year



WAPO.ST/WONKBLOG

Source: National Survey on Drug Use and Health

California 2016



https://www.samhsa.gov/data/sites/default/files/report_2404/ShortReport-2404.html accessed 06/10/2019

CALIFORNIA DEPARTMENT OF PUBLIC HEATLH



OLDER AMERICANS USING MARIJUANA



Source: March 2018 issue of Addiction.

KERR et al. Addiction 2018

IS MARIJUANA A GATEWAY DRUG?

<u>Yes</u>

"A large proportion of individuals who use cannabis go on to use other illegal drugs. The increased risk of progression from cannabis use to other illicit drugs use among individuals with mental disorders"

International Journal of Drug Policy

<u>No</u>

"However, the majority of people who use marijuana do not go on to use other, "harder" substances"

National Institute of Drug Abuse

MARIJUANA + OPIOIDS



AFTER LEGALIZATION OF MARIJUANA

NEGATIVE or NULL EFFECT on:

SUICIDE RATES

Anderson et al. 2014

CRIME RATES

Morris et al. 2014

OPIOID TREATMENT ADMISSIONS

PACULA et al. 2014

OPIOIDS + CANNABIS

Opioid Rx's drop 10% and 25% fewer opioid-related deaths in states with legal marijuana Less dependency No mortality

BRADFORD + BRADFORD 2016 BACHAUBER et al. 2016 OLFSON et al. 2018



After medical marijuana legalization: 23% decrease in opioid dependency 13% decrease in opioid overdoses

SHI et al. Drug and Alcohol Dependence 2017

But...

"Cannabis use appears to INCREASE rather than decrease the risk of developing opioid use disorder"

OLFSON et al.

American Journal of Psychiatry 2018

ADDICTION VS. ABUSE VS. DEPENDENCE

Abuse: Excessive use + misuse Dependence: A biologic process Addiction: Loss of control + compulsive use

DSM-V: CANNABIS USE DISORDER

- 1. Used in larger amounts or over longer period of time
- 2. Unsuccessful efforts to cut down
- 3. A great deal of time is spent obtaining cannabis
- 4. Craving
- 5. Failure to fulfill major role obligations
- 6. Continued use despite recurrent social issues

- 7. Activities are given up or reduced
- 8. Used when physically hazardous
- 9. Continued use despite persistent physical or psychologic problem
- 10.Tolerance
- 11.Withdrawal

DSM-V: CANNABIS USE DISORDER

Assigning a Diagnosis:

Mild: 2-3 symptoms

Moderate: 4-5 symptoms

Severe: 6+ of the 11 listed symptoms

Past-year prevalence of cannabis use and cannabis use disorder, Adults 18+ yo

	2001-2002	2012-2013
Any use	4.1%	9.5%
Meets criteria for CUD	1.5%	2.9%
Of those with past-year use, prevalence of CUD	35.6%	30.6%

Hasin DS, Saha TD, Kerrridge BT, et al. Prevalence of marijuana use disorders in the United States between 2001-2002 and 2012-2013. JAMA Psychiatry. 2015;72(12):1235-1242. doi:10.1001/jamapsychiatry.2015.1858

Thanks to Dr. Travis Lovejoy



Centers for Disease Control and Prevention National Center for Health Statistics

MARIJUANA ADDICTION About 1 in 10 marijuana users will become "addicted"

For people who begin using before the age of 18, that number rises to 1 in 6



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CANNABIS SATIVA



CANNABINOIDS



TRICHOMES PRODUCE

Cannabinoids: THC + CBD + many more

Terpenoids similar to pine + eucalyptus

INCREASING THC POTENCY

Average THC Percent of DEA Submitted Samples 1995 - 2013



CASCINI et al. Curr Drug Abuse Rev. 2012

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THE ENDOCANNABINOID SYSTEM




CANNABINOID RECEPTORS



CB-1 and CB-2 are G-protein Coupled Transmembrane Receptors:

- 1. Detect molecules outside the cell
- 2. Activate internal signal transduction pathways
- 3. Illicit various cellular responses

- Cannabinoid Receptor 1 (CB1) found predominately in the:
 - Central Nervous System
 - Immune System
 - Cardiovascular System
 - Respiratory System
- Cannabinoid Receptor 2 (CB2) found predominately in the:
 - Immune System
 - Musculoskeletal System

Endocannabinoid System



NEUROLOGIC EFFECTS OF CBD

CBD has low affinity for CB1 & CB2

CBD enhances the endocannabinoid system

CBD injections induce anxiolytic effects in mice

Potential effects of CBD:

Anxiolytic

Antidepressant

Neuroprotective

Anti-nausea

FOGACA et al. Neuropharmacology 2018

RECEPTOR for CBD	PHARMACOLOGIC OUTCOME
CB1	Attenuation of impaired learning, memory, and psychosis effects induced by THC
CB2	Anti-Inflammatory
GPR55	Anticancer
5ΗΤ1-α	Pain Relief (via modulation of mu and sigma opioid receptors) Anti-Anxiety
TPVR-1	Anti-Inflammatory Pain Relief
Adenosine	Anti-Inflammatory

WHITE et al. J of Clinical Psychology 2019

CANNABINOID LIGANDS



Kalapa Clinic, Spain

ENDOGENOUS CANNABINOID: ANANDAMIDE (AEA)

phospholipid precursors i.e. cell membranes



SYNTHETIC CANNABINOIDS

Nabilone (Cesmet®)

1985 – treatment of nausea and vomiting (N+V) associated with chemotherapy

Dronabinol (Marinol®)

- 1985 treatment of chemotherapy N+V
- 1992 treatment of anorexia associated with AIDS

Acomplia (Rimonabant®)

2006 – anti-obesity drug that blocked CB1 receptor

Double the risk of developing psychiatric disorders

---- withdrawn





2018 APPROVAL



ROUTES OF ADMINISTRATION

Inhalation

- Water pipe aka "bong" Marijuana cigarette aka "joint" Pipe smoking Vaporization
- Oral "Edibles"
- Sublingual spray
- Topical
 - Lotions, chapstick



KANNACORP

DECARBOXYLATION



https://ardentcannabis.com/decarboxylation-myths/

PULMONARY HAZARDS

WU et al. NEJM 1988

Increased puff volume, therefore greater inhaled volume

Increased airway retention

Increased tar inhalation, with 30% more tar retention in respiratory tract

Decreased cigarette use and frequency of smoking

Marijuana:tobacco cigarette ratio is roughly 1:5

VAPORIZABLE CANNABIS CONCENTRATES Butane Hash Oils (BHOs) used for "dabbing" "Shatter" "Oil" "Wax" "Sugar"

(Andres Rodriguez)



UNKNOWN POSSIBLE BENEFITS BUDNEY et al. 2015 Addiction

Eliminates smoke

Limits second-hand exposure

Reduces smoke-related respiratory illnesses and mortality



Acute Effects of Vaporized Cannabis SPINDLE et al. 2018 JAMA Netw OPEN

Higher peak concentrations when vaporized

Possible that some drug is destroyed when "burned"

UNKNOWN POTENTIAL HARMS **BUDNEY et al. 2015 Addiction** Health impact of long-term aerosol inhalation Increased frequency of use Decreased motivation to reduce use or quit Promotion of a pathway to use for youth Shutterstock

ONSET AND DURATION

160 (A) THC (Smoked) 140 (B) THC-COOH (Smoked) (C) THC (Oral) 120 60 40 20 0 0 1 2 з 7 4 5 6 8 Hours

Blood Levels of THC & Metabolite

INHALED:

Onset: 0-30 minutes Duration: 2 hours +/-Peak: 20-30 minutes

ORAL:

Onset: 30-120 minutes Duration: 6-8 hours Peak: 2-3 hours

HUESTIS et al. J Anal Toxicol 1992

DISTRIBUTION

Cannabinoids are highly **lipophilic** Tissue elimination in 7 days



Kreuz and Axelrod.1973. Science.

ROUTES OF ADMIN. COMPARISON

Russell et al. Int Journal of Drug Policy 2018

"Basically no experimental (e.g. randomized controlled trials) studies exist comparing the effects of different routes of administration on health outcomes, and what data exists largely focuses on associations"

METABOLISM

THC is metabolized in the **liver** by cytochrome p450 enzymes:

CYP2C9 and CYP3A4

Biotransformation of THC to active metabolite:

11-hydroxy-THC

DRUG INTERACTIONS

Inhibitors of CYP2C9:

Amiodarone — Anti-arrhythmic

Cimetidine (Tagamet[®]) — Histamine receptor antagonist

Metronidazole (Flagyl®) — Antibiotic

Fluoxetine (Prozac[®]) — Antidepressant

Fluvoxamine — Antidepressant

Fluconazole — Anti-fungal

DRUG INTERACTIONS

Inhibitors of CYP3A4:

Ketoconazole — Anti-fungal

Itraconazole — Anti-fungal

Clarithromycin — Antibiotic

Erythromycin cyclosporine — Antibiotic

Verapamil — Calcium channel blocker

DRUG INTERACTIONS

Synergism with: Opioids Benzodiazepines Sleep medications Muscle relaxants Alcohol

Oregon Poison Center at OHSU & Salem Health

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LOCAL ANESTHETIC REVIEW



Handbook of Local Anesthesia Author: Stanley F. Malamed

LOCAL ANESTHETIC REVIEW



Handbook of Local Anesthesia Author: Stanley F. Malamed



A. Evidence does show that cannabinoid agonists inhibit sodium and potassium channel conductance — therefore, cannabinoids <u>should</u> promote anesthesia

B. If cannabinoids are antagonizing local anesthesia it is likely from peripheral or central receptor desensitization

Speaking to a group of Dentists (hypothetical!)



LINARES Brazilian J of Psychiatry 2019

NITROUS OXIDE

YAJNIK et al. Drug and Alcohol Dependence 1994

Marijuana users appeared to have intensified subjective effects induced by nitrous oxide

Otherwise, no differences between marijuana users and non-users

MODERATE CONSCIOUS SEDATION

PATEL et al. Southern Medical Association 2015

Marijuana users did NOT require higher doses of sedation compared to non-users to complete a colonoscopy

Opioid users did require a significantly greater amount of fentanyl and midazolam compared to controls

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CNS: ACUTE EFFECTS

Mood: Euphoria - Relaxation - Anxiety

Psychosis: Confusion - Disorientation - Dizziness

Perception:

Intensified sensations

Time distorted

Impaired perceptual ability

Cognition:

Impaired + short-term memory loss



CNS: CHRONIC EFFECTS

Psychosis + Schizophrenia (linked with early onset use)

Depression+ Anxiety

Dependence:

Restlessness

Irritability



Insomnia
CARDIOVASCULAR SYSTEM

Elevated systolic and diastolic blood pressure

Tachycardia

Cardiovascular effects of Marijuana

↑ Sympathetic stimulation

Decreased time to angina

Increased risk of MI for 1 hour post marijuana use

American College of Cardiology 2016

RESPIRATORY SYSTEM

Bronchodilation

Stimulates ventilation

Minimal to zero respiratory depression

Chronic bronchitis

Respiratory immunosuppression



IMMUNE SYSTEM

- Immunosuppression
- Pro-inflammatory THC
- Anti-inflammatory CBD + THC
- <u>Oral</u> CBD has a beneficial action on two symptoms of established inflammation:
 - 1. Edema
 - 2. Hyperalgesia



DIGESTIVE SYSTEM

Increased appetite

Perceived improved quality of life, but no inflammation reduction for IBS, Chron's Disease and Ulcerative Colitis patients



REPRODUCTIVE SYSTEM

FEMALES: Decreased folic acid update

MALES: Decreased sperm count + decreased libido



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CANNABIS-ASSOCIATED ORAL LESIONS

'Cannabis stomatitis'

Leukoedema

Hyperkeratosis

Oral candidiasis

CANNABIS-ASSOCIATED ORAL LESIONS

+ ASSOCIATION

NO ASSOCIATION

Zhang, 1999 – H+N and oral squamous cell carcinoma
Gillison, 2008– HPV and H+N
Feng, 2009– Nasopharyngeal
Marks, 2014 – oropharyngeal

Hashibe, 2002 – H&N

Rosenblatt, 2004 – Oral Cavity

Hashibe, 2006 – H+N and esophageal

Aldington, 2008

Berthiller, 2009

Liang, 2009 – H+N and SCC

Marks, 2014 – oral and tongue

HPV + MARIJUANA + ORAL CANCER Gillison et al. J Natl Cancer Inst 2008 Gillison et al. JAMA 2012

Increasing use of MJ increases risk of HPV H&N Cancer

HPV prevalence increases with MJ use

ETIOLOGY OF ORAL CANCER

Cannabis Carcinogens:

Aromatic hydrocarbons

Benzopyrenes

Nitrosamines

ORAL CANCER SCREENING



CANNABIS STOMATITIS



ORAL CANDIDIASIS



PHARYNGEAL CANCER



HUMAN PAPILLOMA VIRUS



CANCER ANTI-CANCER POTENTIAL







"Scientists reported that THC and other cannabinoids such as CBD slow growth and/or cause death in certain types of cancer cells growing in lab dishes. Some animal studies also suggest certain cannabinoids may slow growth and reduce spread of some forms of cancer."

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Management of cannabis-induced periodontitis via resective surgical therapy

A clinical report

Fatemeh Momen-Heravi, DDS, MPH; Philip Kang, DDS



Figure 1. Clinical and radiographic presentations during initial visit and nonsurgical therapy. **A**. Initial visit, frontal view. **B**. Initial visit, facial view of mandibular anterior teeth showing papillary gingival enlargement and edematous tissue with increased probing depth (7 millimeters) at the mesiofacial aspect of teeth nos. 24 and 25. **C**. Initial visit, lingual view of anterior teeth showing nodular appearance. **D**. Radiographic bone loss seen at the mandibular anterior teeth (nos. 23-26) with more prominent bone loss at interproximal site of teeth nos. 24 and 25. **E**. Frontal view at re-evaluation visit 4 weeks after scaling and root planing.

JADA March 2017



CANNABIS USE AND DESTRUCTIVE PERIODONTAL DISEASES AMONG ADOLESCENTS

LOPEZ + BAELUM J Clin Perio 2009

Cannabis use is NOT positively associated with periodontal diseases in an adolescent population

NEW ZEALAND

DUNEDIN MULTIDISCIPLINARY HEALTH AND DEVELOPMENT STUDY

A cohort from New Zealand

Periodontal examinations at ages 26, 32 and 38 Smoking habits determined at ages 15-38 DUNEDIN MULTIDISCIPLINARY HEALTH AND DEVELOPMENT STUDY

THOMSON et al. J Clin Perio 2013

Tobacco smokers and cannabis smokers are both more likely to be in "Moderately increasing" or "Markedly increasing" trajectory groups for developing periodontitis DUNEDIN MULTIDISCIPLINARY HEALTH AND DEVELOPMENT STUDY

MEIER et al. JAMA Psychiatry 2016

"Cannabis use for up to 20 years is associated with periodontal disease but is not associated with other physical health problems in early midlife."

Study says long-term pot use causes poor gum health – but not much else

Long-term recreational cannabis users in new study had increased risk for tooth loosening and loss but not other health problems across a dozen measures



What we're seeing is that cannabis may be harmful in some respects, but possibly not in every way,' said the study's co-author. Photograph: Nelson Almeida/AFP/Getty Images

UNITED STATES OF AMERICA

NHANES 2011-2012

SHARIFF et al. J Periodontal 2015

Frequent recreational cannabis use is associated with:

Deeper probing depths More clinical attachment loss Higher odds of severe periodontitis



SAN JUAN OVERWEIGHT ADULTS LONGITUDINAL STUDY

ORTIZ et al. J Periodontal 2017

Frequent marijuana users nearly 3x more likely to have severe periodontitis

Frequent marijuana use was not associated with oral HPV infection in a Hispanic population

PERIODONTAL DISEASE PATHOGENESIS



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XEROSTOMIA

Clinical Appearance:

Oral mucosa appears dry, pale, or atrophic

Tongue may be devoid of papillae, fissured and inflamed

New and recurrent dental caries

Difficulty with chewing, swallowing, and tasting

Fungal infections are common

SALIVARY GLAND INNERVATION



CANNABIS + SALIVARY GLANDS

Agonist binding to CB-1 and CB-2 receptors causes:

- 1. Relaxation of myoepithelial cells
- 2. Inhibition of signaling within secretory cells
- 3. Modulation of neurotransmitter release from presynaptic terminals — affecting sympathetic/parasympathetics

Decreased salivation
CANNABIS + XEROSTOMIA DARLING et al. Int Dental Journal 1992

Xerostomia Experience Among: Cannabis users — 69 % Non-cannabis users — 17 %

APPETITE STIMULATION FACT OR FABLE?

FACT

Cannabis inhibits leptin — keeping you hungry

Cannabis stimulates dopamine release — making food more desirable

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SWITZERLAND

CARIES RISK IN CANNABIS + TOBACCO USERS

SCHULZ-KATTERBACH et al. 2009

21% of cannabis users regularly visited the dentist Most cannabis users brushed less frequently 63% of cannabis users consumed "sweet" foods and drinks post-cannabis use Xerostomia reported 1-6 hours after use

6X the amount of decayed surfaces

NEVADA

TOBACCO + MARIJUANA + DENTAL HEALTH

DITMYER et al. J of Adolescent Health 2013

Tobacco:

75% increase in untreated caries

Marijuana + Tobacco:

65% increase in untreated caries

CANNABIS + DENTAL IMPLANTS

Cannabis sativa smoke inhalation decreases bone filling around titanium implants: a histomorphometric study in rats



CANNABIS + DENTAL IMPLANTS **NOGUEIRA-FILHO GDA et al. J Periotondol 2010**

"Considering the limitations of the present study, the deleterious impact of cannabis sativa smoke on bone healing **may** represent a new concern for implant success/failure."

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RECOMMENDED QUESTIONS FOR MEDICAL HISTORY

Do you use marijuana?

- No Yes -If yes: How do you ingest it? Smoke – Eat – Vape – Topical Do you use a high THC or high CBD concentration? Yes — No 🕂 Don't Know How frequently do you use marijuana? Once a month or less \rightarrow Weekly — Daily

INFORMED CONSENT

Can I please just sign the form now?

American College of Dentists:

"A blanket non-treatment policy is not supported by available science or ethical analysis"

Peltier JACD 2010



RECOMMENDED RESOURCES

UW — ALCOHOL AND DRUG ABUSE INSTITUTE: <u>HTTP://LEARNABOUTMARIJUANAWA.ORG/</u> <u>FACTSHEETS.HTM</u>

PUBLIC HEALTH HOTLINE: WWW.WARECOVERYHELPLINE.ORG/

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TAKE HOME MESSAGES

1. Know your patient and her/his medical history

2. Be comfortable talking to your patients about their marijuana use

THANK YOU! TAYLORBA@OHSU.EDU

