

### New Jersey Department of Health Medicinal Marijuana Program PO 360 Trenton, NJ 08625-0360

### MEDICINAL MARIJUANA PETITION (N.J.A.C. 8:64-5.1 et seq.)

### INSTRUCTIONS

This petition form is to be used <u>only</u> for requesting approval of an additional medical condition or treatment thereof as a "debilitating medical condition" pursuant to the New Jersey Compassionate Use Medical Marijuana Act, N.J.S.A. 24:6I-3. Only one condition or treatment may be identified per petition form. For additional conditions or treatments, a separate petition form must be submitted.

NOTE: This Petition form tracks the requirements of N.J.A.C. 8:64-5.3. Note that if a petition does not contain all information required by N.J.A.C. 8:64-5.3, the Department will deny the petition and return it to petitioner without further review. For that reason the Department strongly encourages use of the Petition form.

This completed petition must be postmarked August 1 through August 31, 2016 and sent by certified mail to:

New Jersey Department of Health Office of Commissioner - Medicinal Marijuana Program Attention: Michele Stark 369 South Warren Street Trenton, NJ 08608

Please complete <u>each</u> section of this petition. If there are any supportive documents attached to this petition, you should reference those documents in the text of the petition. If you need additional space for any item, please use a separate piece of paper, number the item accordingly, and attach it to the petition.

1.	Petitioner Information
	Name:
	Street Address:
	City, State, Zip Code:
	Telephone Number:
	Email Address:
2.	Identify the medical condition or treatment thereof proposed. Please be specific. Do not submit broad categories (such as "mental illness").
	Migraine
3.	Do you wish to address the Medical Marijuana Review Panel regarding your petition?
	☐ Yes, in Person
	∑ Yes, by Telephone
	□ No
4.	Do you request that your personally identifiable information or health information remain confidential?
	⊠ Yes
	□ No
	If you answer "Yes" to Question 4, your name, address, phone number, and email, as well as any medical or health information specific to you, will be redacted from the petition before forwarding to the panel for review.



SEP 6 2016

## MEDICINAL MARIJUANA PETITION (Continued)

5.	Describe the extent to which the condition is generally accepted by the medical community and other experts as a valid,
	existing medical condition.

Migraine is widely accepted as a valid medical condition among physicians of all specialties in addition to numerous medical associations/organizations such as the American Medical Association, American Neurological Association, and the American Headache Society.

6.	If one or more treatments of the condition, rather than the condition itself, are alleged to be the cause of the patient's
	suffering, describe the extent to which the treatments causing suffering are generally accepted by the medical
	community and other experts as valid treatments for the condition.
	•

N/A

7. Describe the extent to which the condition itself and/or the treatments thereof cause severe suffering, such as severe and/or chronic pain, severe nausea and/or vomiting or otherwise severely impair the patient's ability to carry on activities of daily living.

I experience near daily headaches and frequent severe migraines. The migraine symptoms include but are not limited to, continuous sharp and/or throbbing head pain; pain in my jaw and scalp; neck pain; visual disturbances including blurry vision, tunnel vision, and aura; nausea; vomiting; diarrhea; blackouts; tinnitus; vertigo; loss of balance; motor skill impairment; aphasia; and fatigue. Most often, the migraines are debilitating to the point I must stay home and in bed and am completely unable to carry on typical activities of daily living. The migraine episodes each last up to 72 hours and are followed by 1 - 2 days of "postdrome" symptoms such as nausea, fatigue and brain fog. Every aspect of my life is negatively impacted by the migraines.

8. Describe the availability of conventional medical therapies other than those that cause suffering to alleviate suffering caused by the condition and/or the treatment thereof.

I am unable to take most of the commonly prescribed conventional medications due to negative reactions, some quite severe, or because of contraindicated health history. The medications I did not respond negatively to failed to provide reduction in symptoms.

9. Describe the extent to which evidence that is generally accepted among the medical community and other experts supports a finding that the use of marijuana alleviates suffering caused by the condition and/or the treatment thereof. [Note: You may attach articles published in peer-reviewed scientific journals reporting the results of research on the effects of marijuana on the medical condition or treatment of the condition and supporting why the medical condition should be added to the list of debilitating medical conditions.]

Articles attached

- 1: Effects of Medical Marijuana on Migraine Headache Frequency in an Adult Population, PHARMACOTHERAPY Vol 36 No. 5 2016
- 2: Comprehensive Review of Medical Marijuana Cannabinoids, and Therapeutic Implications in Migraine and Headache: What a Long Strange Trip It's Been; Baron EP, Headache 2015 Jun; 55(6): 885-916. doi: 10.1111/head.12570. Epub 2015 May 25
- 3: Cannabis for Migraine Treatment: the Once and Future Prescription? An historical and Scientific Review: Pain. 1998 May; 76(1-2):3-8.

# MEDICINAL MARIJUANA PETITION (Continued)

	Attach letters of support from physicians or other licensed health care professionals knowledgeable about the condition. List below the number of letters attached and identify the authors.
	1 - Patricia Chichon, APN
-2-1	

Signature of Patitionar

Date

8/30/2016

I certify, under penalty of perjury, that I am 18 years of age or older; that the information provided in this petition is true and accurate to the best of my knowledge; and that the attached documents are authentic.

### Effects of Medical Marijuana on Migraine Headache Frequency in an Adult Population

Danielle N. Rhyne, <sup>1</sup> Sarah L. Anderson, <sup>1</sup> Margaret Gedde, <sup>2</sup> and Laura M. Borgelt, <sup>1,3,\*</sup>

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STUDY OBJECTIVE No clinical trials are currently available that demonstrate the effects of marijuana on patients with migraine headache; however, the potential effects of cannabinoids on serotonin in the central nervous system indicate that marijuana may be a therapeutic alternative. Thus, the objective of this study was to describe the effects of medical marijuana on the monthly frequency of migraine headache.

DESIGN Retrospective chart review.

SETTING Two medical marijuana specialty clinics in Colorado.

Patients One hundred twenty-one adults with the primary diagnosis of migraine headache who were recommended migraine treatment or prophylaxis with medical marijuana by a physician, between January 2010 and September 2014, and had at least one follow-up visit.

MEASUREMENTS AND RESULTS The primary outcome was number of migraine headaches per month with medical marijuana use. Secondary outcomes were the type and dose of medical marijuana used, previous and adjunctive migraine therapies, and patient-reported effects. Migraine headache frequency decreased from 10.4 to 4.6 headaches per month (p<0.0001) with the use of medical marijuana. Most patients used more than one form of marijuana and used it daily for prevention of migraine headache. Positive effects were reported in 48 patients (39.7%), with the most common effects reported being prevention of migraine headache with decreased frequency of migraine headache (24 patients [19.8%]) and aborted migraine headache (14 patients [11.6%]). Inhaled forms of marijuana were commonly used for acute migraine treatment and were reported to abort migraine headache. Negative effects were reported in 14 patients (11.6%); the most common effects were somnolence (2 patients [1.7%]) and difficulty controlling the effects of marijuana related to timing and intensity of the dose (2 patients [1.7%]), which were experienced only in patients using edible marijuana. Edible marijuana was also reported to cause more negative effects compared with other forms.

Conclusion The frequency of migraine headache was decreased with medical marijuana use. Prospective studies should be conducted to explore a cause-and-effect relationship and the use of different strains, formulations, and doses of marijuana to better understand the effects of medical marijuana on migraine headache treatment and prophylaxis.

KEY WORDS cannabis, marijuana, migraine, headache.

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Cannabis encompasses three species: Cannabis indica, Cannabis sativa, and Cannabis ruderalis. Cannabis is composed of more than 400 compounds, with more than 60 being cannabinoids (CBs). The most common psychoactive CB is

 $\Delta^9$ -tetrahydrocannabinol (THC). Cannabidiol (CBD) is another common CB, which accounts for 40% of the plant's extract and is one of the primary constituents of medical marijuana.<sup>1</sup>

Phytocannabinoids are CBs that occur naturally in the plant (e.g., THC, CBD) and stimulate CB receptors throughout the body. The body contains endogenous CBs and receptors, which make up the endocannabinoid system. This system is responsible for maintaining homeostasis in our bodies. Research has found that the endocannabinoid system might be a target for treatment of diseases such as migraine headache (HA), fibromyalgia, neuropathic pain, and irritable bowel syndrome.3 The endocannabinoid system is common throughout the central nervous system and has presence in peripheral tissues as well. This system includes CB receptors, CB<sub>1</sub> and CB2, and ligands such as anandamide (AEA) and 2-arachidonoylglycerol (2-AG), which are located throughout the brain and influence many regulatory systems. 4 Cannabinoid, receptors are widely expressed in the central and peripheral nervous system. In the central nervous system, activation of CB<sub>1</sub> receptors leads to inhibition of the following neurotransmitters: γ-aminobutyric acid (GABA), glutamate, serotonin, dopamine, acetylcholine, norepinephrine, cholecystokinin, and D-aspartate. Cannabinoid receptors are widely expressed throughout the peripheral tissues, especially the immune system, and have antiinflammatory properties and analgesic effects. Anandamide is a partial agonist at CB receptors and binds to CB<sub>1</sub> receptors with higher affinity than CB2 receptors. Anandamide has been shown to have inhibitory effects on serotonin type 3 (5-hydroxytryptamine [HT]<sub>3</sub>) receptors, further suggesting its antiemetic and analgesic roles. It is also a 5-HT<sub>1A</sub> receptor agonist and a 5-HT<sub>2A</sub> receptor antagonist.

 $\Delta^9$ -Tetrahydrocannabinol acts as a partial agonist at CB<sub>1</sub> and CB<sub>2</sub> receptors and is structurally similar to endogenous AEA. Cannabidiol antagonizes CB<sub>1</sub> receptors at low levels in the presence of THC and acts as a potent analgesic. The mechanisms of CBs have been examined and suggest serotonergic and dopaminergic effects as well as providing antiinflammatory effects. Evidence suggests that THC affects serotonin and dopamine by inhibiting serotonin release from platelets, stimulates 5-HT synthesis, and modulates dopaminergic imbalances.<sup>3</sup> Specific conditions such as Alzheimer's disease and depression occur due to a lack of neurotransmitters.<sup>6</sup> As a result, it has been hypothesized that patients

with central nervous system disorders might have a clinical endocannabinoid deficiency. Further evidence from one study<sup>6</sup> reported reduced levels of AEA in the cerebrospinal fluid of patients with migraine HA. As a result of reduced AEA levels, the trigeminovascular system is activated, resulting in a migraine HA.

The role of serotonin in migraine HA is supported by the efficacy of serotonin agonists such as triptans for acute treatment of migraine. Other agents used for acute migraine treatment include nonsteroidal antiinflammatory drugs, acetaminophen, and antiemetic agents. In addition to acute migraine treatment, the American Academy of Neurology 2012 guidelines recommend pharmacologic agents for preventive ther-Level A treatment recommendations include certain antiepileptic drugs, β-blockers, and triptans. Current guidelines do not address the use of cannabis for the prevention or treatment of migraine HA; however, the potential effects of CBs on serotonin in the central nervous system make it possible that cannabis could be a therapeutic alternative.

Although there are no clinical trials available, to our knowledge, demonstrating the effects of marijuana on patients with migraine HA, five case reports described patients who used dronabinol with or without additional marijuana products for treatment of their vascular or migraine HAs and who experienced an overall decrease in migraine HA.<sup>8</sup> These case reports, however, lack scientific rigor and consistent reporting and do not provide detailed information about the positive or negative impact of marijuana.

Due to a lack of data on the efficacy and proposed mechanism of the pharmacologic benefit of medical marijuana in patients with migraine HA, clinical data describing the effectiveness of medical marijuana for the frequency of migraine HA are necessary. Other useful information would include the dose and type of medical marijuana being used and other clinical effects of marijuana. Thus, the primary purpose of this study was to determine the monthly frequency of migraine HA in patients diagnosed with migraine HA who used medical marijuana.

### Methods

Study Design, Setting, and Outcomes

This was a retrospective, observational chart review of patients who were seen at Gedde Whole Health, a private medical practice with offices located in Colorado Springs and Buena Vista, Colorado. The physician in these clinics specializes in applications of medical marijuana for various conditions and makes recommendations to patients for the use of medical marijuana when a patient has a qualifying medical condition based on state requirements. This study was approved by the Colorado Multiple Institutional Review Board.

The primary outcome of this study was monthly frequency of migraine HA with medical marijuana use. Secondary outcomes were type and dose of medical marijuana used, previous and adjunctive migraine therapies, and patient-reported effects.

### Patient Chart Identification and Data Collection

Charts for adult patients, aged 18-89 years old, with a primary diagnosis of migraine HA and at least one follow-up visit were included for review. Data were extracted by a single investigator for consistency. Data collection included sex, number of years with migraine HA, medical history, previous migraine therapy, adjunctive migraine therapy, number migraine HAs per month, types and doses of marijuana, frequency of marijuana use, number of migraine HAs per month at the follow-up visit, and patient-reported effects. Number of migraine HAs experienced each month and the amount of marijuana used each month were patient-reported data. Medical marijuana quantities were reported in ounces, with the exception of edible dosage forms, which were reported in milligrams. Edible doses were then converted to ounces per month based on a calculation of 100 mg/day of edible marijuana being considered equivalent to 1 oz/month of cannabis flower. This conversion was based on an approximated potency of CB in cannabis flower used by study patients of 10% (w/w), based on historical and contemporary data.9 If CB potency in cannabis flower is approximated at 10% (w/w), then 1 g of cannabis flower contains 0.1 g (or 100 mg) of CB. Given that 1 oz equals 28 g, and 1 month is approximately 30 days, then 1 oz/month is approximately 28 g/30 days, or approximately 1 g/day. Then 1 oz/month of cannabis flower roughly equals 1 g/day of cannabis flower, which converts approximately to 100 mg/day of CB. The CB conversion used is illustrated in the following equation:

No. of ounces/month≈(no. of milligrams/day)/100

When ranges of doses were reported (e.g., 1–2 oz/month), the highest dose of medical marijuana was documented.

### Statistical Analysis

Descriptive statistics were used to describe demographic and clinical data. The mean and standard deviation, median and interquartile range, and proportions were calculated for normally distributed data, nonparametric data, and nominal data, respectively. Two-tailed paired t tests were used when possible; a p value less than 0.05 was considered to indicate a statistically significant difference. All statistical tests were performed using GraphPad software (GraphPad Software, Inc., San Diego, CA).

### Results

All patient visits with dates between January 1, 2010, and September 30, 2014, were screened, and 262 patient charts with a primary diagnosis of migraine HA were identified. Of these, 121 had at least one follow-up visit recorded and were eligible for inclusion. The other 141 patient charts were excluded due to the absence of a follow-up visit.

The initial visit characteristics for the 121 included patients are shown in Table 1. Fifty-two percent of patients were female, and the average duration of migraine HA was 14 years. Eighty-two (67.8%) patients had a history of previous or current marijuana use at the initial visit. Follow-up visit characteristics are also shown in Table 1.

The primary outcome of mean number of migraine HAs per month at the initial and follow-up visits were 10.4 and 4.6 (p<0.0001), respectively. The mean time between the initial and most recent follow-up visit was 21.8 months (range 12-37 mo). A total of 103 patients (85.1%) reported a decrease in frequency of migraine HAs per month. Alternatively, 15 patients (12.4%) reported the same number of HAs per month, and 3 (2.5%) had an increase in the number of HAs per month. More than half of the patients (62 [51.2%]) reported using two or more forms of marijuana for migraine HA treatment and/or prophylaxis at the follow-up visit. The forms of medical marijuana used included vaporized (42 patients), edible (66

Table 1. Characteristics of the 121 Study Patients

Characteristic	Initial Visit	Follow-up Visit	p Value
Female	63 (52.1)	NA	NA
Mean no. of years with migraine headache	14	NA	NA
Time between initial and most recent follow-up visit (mo)	NA	21.8 [12–37]	NA
Previous marijuana use	82 (67.8)	121 (100)	< 0.0001
Mean no. of migraines/month	10.4	4.6	< 0.0001
Used migraine prescription drug therapy	59 ( <del>4</del> 8.8)	52 (43.0)	0.44
No. of migraine medications/patient	1.15 [0-2]	1.09 [0-2]	0.22
Used 1 form of medical marijuana	57/82 (69.5)	59/121 (48.8)	0.004
Used 2 forms of medical marijuana	20/82 (24.4)	51/121 ( <del>4</del> 2.1)	0.011
Used ≥ 3 forms of medical marijuana	5/82 (6.1)	11/121 (9.1)	0.597

Data are no. (%) of patients or mean [range] values unless otherwise specified. NA = not applicable.

patients), topical (15 patients), and smoked (65 patients). Follow-up visit mean monthly doses of each type of marijuana were 2.64 oz, 2.59 oz, 2.73 oz, and 1.59 oz for vaporized, edible, topical, and smoked forms, respectively. Reasons for use of medical marijuana included migraine HA prophylaxis (7 patients), acute treatment of migraine HA (4 patients), or both (110 patients). A post-hoc sample size calculation was performed by using PASS 14 (NCSS Statistical Software; NCSS, LLC, Kaysville, UT) to ensure the internal validity of these results. This analysis yielded a necessary sample size of 96 patients to achieve 80% power when the mean population difference in number of migraine headaches per month was 5.8 and the standard deviation for both groups was 10.0, which was exceeded in our study with a sample size of 121 patients.

Migraine HA prescription drug therapy was reported in 59 (48.8%) patients, with the average number of medications being 1.15 per patient at the initial visit. At the follow-up visit, 52 (42.9%) patients reported using migraine HA drug therapy in addition to medical marijuana. The average number of migraine HA medications was 1.09 per patient; however, the difference between the number of medications at the initial and follow-up visits was not statistically significant (p=0.22). There were 62 patient-reported effects, illustrated in Tables 2 and 3. Positive effects were recorded for 48 patients, with half (24 patients) of the effects being reported as prevention of migraine HA with decreased frequency of migraine HA (Table 2). These beneficial effects were reported for all forms of marijuana. In addition, migraine abortion was the second most common positive effect (14 patients). Negative patient-reported effects are shown in Table 3 (n=14). Patients who used the edible form (11 patients) were most likely to report negative effects, which included somnolence (2 patients) and difficulty controlling the effects of marijuana, including when the effects would occur and the intensity of effects (2 patients).

### Discussion

This study is one of the first to reveal that migraine HA frequency decreased in patients using medical marijuana, and the difference in frequency between the initial and follow-up visit was statistically significant (p<0.0001). Further, 90% of patients used marijuana for both treatment and prophylaxis of migraine HA. More than half of the patients at the follow-up visit reported using two or more delivery methods of marijuana for migraine HA treatment, which demonstrates that some delivery methods might be preferred for abortive treatment versus migraine HA prevention. For example, 12 patients reported migraine abortion success while using an inhaled form of marijuana. This effect was likely due to the quick onset of action with inhaled marijuana as opposed to a slower onset of action with an edible form. Although there were more overall positive effects reported, there were more negative reports for the edible form of marijuana, likely due to variability of onset of action. As previous research has shown, the pharmacokinetics of the edible forms are variable, and it could take up to 4 hours to reach peak THC concentration, with clinical effects lasting longer (e.g., up to 8 hrs). 10, 11 These pharmacokinetic factors likely led to the reported difficulty in controlling the effects of marijuana.

This study has some limitations. First, the retrospective nature of the study limits the ability to evaluate the causality of the use of medical marijuana and decrease in migraine HA frequency, and it does not allow for controlling the

Table 2. Patient-Reported Positive Effects in the 121 Patients

	No. of Patients (%)	Medical Marijuana Form (No. of Patients)				
Effect		Vaporized	Edible	Topical	Smoked	
Prevention of migraine headache with decreased frequency of migraine headache <sup>a</sup>	24 (19.8)	X	Х	Х	Х	
Aborts migraine headache	14 (11.6)	5	1	1	7	
Relieves pain	4 (3.3)		3	1		
Reduces nausea	1 (0.8)				1	
Other effects	5 (4.1)		4	1		
All positive effects	48 (39.7)					

<sup>&</sup>lt;sup>a</sup>Patients used a combination of medical marijuana forms.

Table 3. Patient-Reported Negative Effects in the 121 Patients

	No. of Patients (%)	Medical Marijuana Form (no. of patients)				
Effect		Vaporized	Edible	Topical	Smoked	
Somnolence	2 (1.7)		X			
Difficulty controlling effects of marijuana related to timing and intensity of the dose	2 (1.7)		Χ			
Increased headache and seizure	1 (0.8)	X	X			
Bad dreams	1 (0.8)				X	
Jitteriness and nausea	1 (0.8)				X	
Memory loss	1 (0.8)			X		
Other effects	6 (5.0)		X	X	X	
All negative effects	14 (11.6)					

type of dose used. This study showed a reduction in migraine HA frequency with the use of medical marijuana; however, it demonstrates the need for performing additional studies in patients with migraine HA to explore the benefits and risks of medical marijuana in a controlled environment. Second, more than half of the patients with migraine did not have a follow-up visit and were excluded from the study. The effects of marijuana are unknown for these patients, and medical follow-up was no longer required in Colorado with the legalization of marijuana in January 2014. Third, chart documentation was not consistent across every patient. For instance, documentation of clinical effects appeared for only half of the patients. Specific directions for use of medical marijuana were not recorded in the charts. In addition, most patients reported previous use of marijuana at the initial visit; however, the duration of previous use was unknown. Given that most patients had used marijuana prior to the initial visit, this study suggests that interaction with a provider may improve how prior or current marijuana use can be optimized to improve symptoms. Documentation revealed that most patients used marijuana daily; however, it is unknown if some patients used marijuana

multiple times per day. Fifty-two patients used preventive and/or abortive pharmacologic agents for migraine HA in addition to medical marijuana, but the frequency of their use was not documented. Also, information on the strains and/or amounts of CBs within medical marijuana products was not consistently documented, so this information was unable to be collected.

The ideal study design to further investigate the effects of medical marijuana on the frequency of migraine HA would be a randomized, placebo-controlled clinical trial with a marijuana washout period prior to study start. The ideal study would also provide participants with standardized quantities and potencies of medical marijuana while tracking their adherence, number of migraine HAs, and adverse effects in a systematic fashion analogous to that of a prescription drug study. Based on current federal regulations regarding research of this type and lack of consistency among cannabis and cannabis compounds, substantial changes in legislation and product manufacturing would need to occur before a study with this scientific rigor could feasibly be performed.

As health care providers enter into shared decision-making with patients experiencing

migraine HA and using marijuana, this chart review provided some insight about key messages for patients. For example, providers need to be prepared to discuss potential benefits and risks of marijuana use. In addition, given the difference in strains, doses, and formulations, it may be difficult to establish a standardized dosing schedule, and marijuana use should be accurately documented. Edible formulations have a longer onset of action and variable patient responses, so patients should be advised to start with a low dose, carefully monitor response, and titrate slowly, if needed. Use of prescription and over-the-counter medications for migraine HA should also be documented to optimize medication use.

### Conclusion

Patients using medical marijuana for migraine HA reported a statistically significant decrease in the number of migraine HAs per month. Almost all patients used marijuana daily for migraine HA prevention. Inhaled forms of marijuana were commonly used for acute migraine treatment and were reported to abort migraine HA. Overall, more positive than negative effects were reported with medical marijuana use. Edible marijuana was reported to cause more negative effects compared with other forms. Further research should be performed to determine if there is a preferred delivery method, dose, and strain of medical marijuana for migraine HA therapy as well as the potential long-term effects of medical marijuana.

### Disclosure

Danielle Rhyne and Sarah Anderson have no conflicts of interest to disclose. Margaret Gedde is the medical director and owner of Gedde Whole Health. Laura Borgelt has served as a member of the following working groups: Colorado Department of Public Health and Environment Retail Marijuana Public Health Advisory Committee (2014–present), Marijuana Pregnancy and Lactation Guidance for Colorado

Health Care Providers Committee (2014–2015), and Amendment 64 (Marijuana Legalization) Task Force Working Group: Consumer Safety and Social Issues (2013); Colorado Department of Revenue Marijuana Enforcement Division: Retail Marijuana Product Potency and Serving Size Working Group (2014); and member of the State Licensing Authority/Colorado Department of Revenue: Medical and Retail Marijuana Mandatory Testing and Random Sampling Working Group (2013) and Amendment 64 (Marijuana Legalization) and HB13-1317 stakeholder working group for rulemaking: Labeling, Product Safety and Marketing (2013). She declares no financial conflict of interest.

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Headache. 2015 Jun;55(6):885-916. doi: 10.1111/head.12570. Epub 2015 May 25.

# Comprehensive Review of Medicinal Marijuana, Cannabinoids, and Therapeutic Implications in Medicine and Headache: What a Long Strange Trip It's Been ....

Baron EP1.

### **Author information**

### Abstract

### **BACKGROUND:**

The use of cannabis, or marijuana, for medicinal purposes is deeply rooted though history, dating back to ancient times. It once held a prominent position in the history of medicine, recommended by many eminent physicians for numerous diseases, particularly headache and migraine. Through the decades, this plant has taken a fascinating journey from a legal and frequently prescribed status to illegal, driven by political and social factors rather than by science. However, with an abundance of growing support for its multitude of medicinal uses, the misguided stigma of cannabis is fading, and there has been a dramatic push for legalizing medicinal cannabis and research. Almost half of the United States has now legalized medicinal cannabis, several states have legalized recreational use, and others have legalized cannabidiol-only use, which is one of many therapeutic cannabinoids extracted from cannabis. Physicians need to be educated on the history, pharmacology, clinical indications, and proper clinical use of cannabis, as patients will inevitably inquire about it for many diseases, including chronic pain and headache disorders for which there is some intriguing supportive evidence.

### **OBJECTIVE:**

To review the history of medicinal cannabis use, discuss the pharmacology and physiology of the endocannabinoid system and cannabis-derived cannabinoids, perform a comprehensive literature review of the clinical uses of medicinal cannabis and cannabinoids with a focus on migraine and other headache disorders, and outline general clinical practice guidelines.

### **CONCLUSION:**

The literature suggests that the medicinal use of cannabis may have a therapeutic role for a multitude of diseases, particularly chronic pain disorders including headache. Supporting literature suggests a role for medicinal cannabis and cannabinoids in several types of headache disorders including migraine and cluster headache, although it is primarily limited to case based,

anecdotal, or laboratory-based scientific research. Cannabis contains an extensive number of pharmacological and biochemical compounds, of which only a minority are understood, so many potential therapeutic uses likely remain undiscovered. Cannabinoids appear to modulate and interact at many pathways inherent to migraine, triptan mechanisms ofaction, and opiate pathways, suggesting potential synergistic or similar benefits. Modulation of the endocannabinoid system through agonism or antagonism of its receptors, targeting its metabolic pathways, or combining cannabinoids with other analgesics for synergistic effects, may provide the foundation for many new classes of medications. Despite the limited evidence and research suggesting a role for cannabis and cannabinoids in some headache disorders, randomized clinical trials are lacking and necessary for confirmation and further evaluation.

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### **KEYWORDS:**

CBD; THC; cannabidiol; cannabinoids; cannabis; delta-9-tetrahydrocannabinol; headache; hemp; medical marijuana

### Comment in

• Up in Smoke: A New View on an Old Friend. [Headache. 2015]

PMID:

26015168

DOI:

10.1111/head.12570

Format: Abstract

### Send to

Pain. 1998 May;76(1-2):3-8.

# Cannabis for migraine treatment: the once and future prescription? An historical and scientific review.

Russo E<sup>1</sup>.

### **Author information**

### Abstract

Cannabis, or Marijuana, has been used for centuries for both symptomatic and prophylactic treatment of migraine. It was highly esteemed as a headache remedy by the most prominent physicians of the age between 1874 and 1942, remaining part of the Western pharmacopoeia for this indication even into the mid-twentieth century. Current ethnobotanical and anecdotal references continue to refer to its efficacy for this malady, while biochemical studies of THC and anandamide have provided a scientific basis for such treatment. The author believes that controlled clinical trials of Cannabis in acute migraine treatment are warranted.

PMID:

9696453

[PubMed - indexed for MEDLINE]

New Jersey Department of Health Medical Marijuana Program

August 18, 2016

To Whom it May Concern:

In regard to Patient:

I am a board certified Nurse Practitioner with a family practice in West Amwell, NJ and have been in practice for over 30 years. I am writing in support of my patient, and her appropriate medical need for Medical Cannabis, due to her long term history of debilitating Migraine Headaches.

has been in my practice since 2008 and we have worked to improve/resolve her migraine headaches including referrals to some of the top headache specialists on the east coast without success. She has also tried chiropractics, acupuncture, massage, herbs, and homeopathy under medical supervision without success.

She is struggling to raise a family and live a somewhat normal life, while she lives in a fog of nearly daily headaches, with migraines at least daily for half of the month with chronic fatigue and memory loss.

We have talked about the use of medical cannabis and she and her husband have reviewed the literature and feel she absolutely should try this valid option.

Having reviewed the medical literature and discussed the side effects and contraindications, I feel it is a very valid, potentially very beneficial option for and she is an appropriate candidate for its legal use in the state of New Jersey.

Please consider my petition at your earliest convenience.

Thank you for your consideration of this matter and do not hesitate to contact me if I can be of any future assistance.

Sincerely,

Talua & Chickon APN
Patricia G. Chichon APN

609-397-1466(f-1013)