Medical Cannabis for the Treatment of Fibromyalgia

George Habib, MD, MPH†‡ and Suheil Artul, MD§∥

Background: Fibromyalgia is a chronic pain syndrome, characterized by chronic musculoskeletal pain, fatigue, and mood disturbances. There are nearly no data on the effect of medical cannabis (MC) treatment on patients with fibromyalgia.

Methods: Data were obtained from the registries of 2 hospitals in Israel (Laniado Hospital and Nazareth Hospital) on patients with a diagnosis of fibromyalgia who were treated with MC. After obtaining patient consent, demographic, clinical, and laboratory parameters were documented. All the patients also completed the Revised Fibromyalgia Impact Questionnaire regarding the period before and after MC treatment.

Results: Thirty patients were identified, and 26 patients were included in the study. There were 19 female patients (73%), and the mean age of the study group was 37.8 ± 7.6 years. The mean dosage of MC was 26 ± 8.3 g per month, and the mean duration of MC use was 10.4 ± 11.3 months. After commencing MC treatment, all the patients reported a significant improvement in every parameter on the questionnaire, and 13 patients (50%) stopped taking any other medications for fibromyalgia. Eight patients (30%) experienced very mild adverse effects.

Conclusions: Medical cannabis treatment had a significant favorable effect on patients with fibromyalgia, with few adverse effects.

Key Words: fibromyalgia, medical cannabis, treatment

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Fibromyalgia is one of the most common chronic pain syndromes.1 It is characterized by diffuse musculoskeletal pain, in addition to extreme fatigue and mood and sleep disturbances.1 The pathogenesis of fibromyalgia is not clear. It usually affects women more than men and has a genetic preponderance.2 Its prevalence in the general population is estimated to be approximately 7%, and it is more common among women than men.3

Fibromyalgia can have tremendous physical, as well as psychological, impacts on patients.4 For example, many patients may be unable to accomplish various tasks at work and home, resulting in physical disability, which can be accompanied by anxiety and depression. Unfortunately, in most patients, fibromyalgia is chronic, and the main treatment is pain control medications. These medications include simple analgesics, pregabalin, and opioids.5,6 Patients with fibromyalgia may also benefit from tricyclic antidepressants, benzodiazepines, and other types of antidepressants.7 However, many of these medications are associated with adverse effects, which affect compliance. As a result, many patients with fibromyalgia experience continuous pain.

Cannabis is derived from the cannabis plant and is considered an illicit drug in most countries, including Israel. However, it is widely used illegally or legally in some countries where cannabis use is not outlawed.8 The 2 main cannabis plant species are Cannabis sativa and Cannabis indica.9 Most species today are a hybrid of the two, with cannabis derived from C. sativa designed mainly for morning or daytime use because it induces calmness and good sleep.10 The flowers of the cannabis plant contain more than 100 types of phyto cannabinoids. Most research has focused on Δ-9-tetrahydrocannabinol and cannabidiol, both of which have the highest concentrations of phyto cannabinoids.11

In recent years, cannabinoid receptors have been discovered that supply the MC. The patient is advised to consume the same amount of cannabis used. In general, the patients in these studies reported favorable effects of cannabis use. A systematic review of the use of synthetic cannabinoids in fibromyalgia (nabilone, 2 studies) found evidence (very low quality) of a greater reduction in pain and limitations in health-related quality of life in the synthetic cannabinoid group as compared with a placebo group in 1 study and better effects of synthetic cannabinoids on sleep than amitriptyline in another study.18 The aim of the present study was to examine the effects of licensed MC on patients with fibromyalgia in an Israeli population.

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PATIENTS AND METHODS

This was a retrospective review of patients with fibromyalgia who were treated with MC. Data were obtained on patients with fibromyalgia from the registries of Laniado Hospital, Netanya, and Nazareth Hospital, Nazareth, Israel. All the patients met the diagnostic criteria for fibromyalgia. The patients were contacted and asked to participate in the study.

The inclusion criteria were patients older than 18 years and being able to sign a consent form. The exclusion criteria were patients with malignancy-associated or other rheumatic disease–associated fibromyalgia.

All the patients signed a consent form, and the study was approved by the local ethics committee of Laniado Hospital.

After obtaining consent, demographic, clinical, and laboratory parameters were documented. Demographic data included age and sex. Clinical data included the duration of symptoms of fibromyalgia, time of disease diagnosis, medical treatment before and after MC treatment, dose of MC, mode of consumption, adverse effects of MC, employment status, impact of MC treatment on return to work (part-time or full-time work), and number of hours of work. Laboratory parameters included serological markers and vitamin B12 status.

In addition, all the patients completed the Revised Fibromyalgia Impact Questionnaire (FIQR) on the period before and after MC treatment. The patients were asked to document any medical treatment they had received for fibromyalgia in the 2 months prior to starting MC treatment and during the 2 months while receiving MC treatment. Simple analgesics were considered paracetamol, dipyrone, or orphenadrine citrate combined with paracetamol. Mild opiates were considered tablets containing equal to or less than 30 mg of codeine or equal to or less than 100 mg tramadol per day. Strong opiates were considered any treatment containing oxycodone, patches containing fentanyl or buprenorphine, and tablets containing more than 200 mg tramadol per day. The participants were also asked about adverse effects associated with the use of MC. In the questionnaire, all the participants were asked to describe their experience of MC treatment in their own words.

For statistical analysis, Wilcoxon signed rank test was conducted to compare the results of questionnaire data before MC treatment with those after MC treatment. A $\chi^2$ test was performed to compare the number of patients receiving different types of medications prior to MC treatment with the number of patients receiving the same types of medications while receiving MC treatment.

RESULTS

Thirty patients were identified. One patient could not be contacted. Two patients had cancer, and 1 patient had inflammatory joint disease. Thus, 26 patients were included in the study. Nineteen patients (~73%) were females, and the mean age of the study group was 37.8 ± 7.6 years. The mean duration of fibromyalgia diagnosis was 4.3 ± 2.64 years. The mean dose of MC was 26 ± 8.3 g per month, and the mean duration of treatment was 10.4 ± 11.3 months, with a median duration of 3 months. No patient ceased MC treatment.

Table 1 summarizes the demographic and clinical parameters of the patients. All the patients smoked or inhaled MC. One of the 26 patients used a combination of smoking and oral oil drops. Table 2 summarizes the mean score for each item in the FIQR before and after MC treatment. Table 3 summarizes the various medications patients took in the 2 months prior to MC treatment and in the 2 months under MC treatment. In the study group, 13 patients (50%) ceased taking any medication other than MC. Twelve patients (~46%) reduced the dose/number of medications by at least 50% as compared with the dose/number of medications prior to MC treatment. Table 4 summarizes the adverse effects of MC treatment reported by the patients.

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Results (%)</th>
</tr>
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<tbody>
<tr>
<td>Sex, female: male</td>
<td>19:7</td>
</tr>
<tr>
<td>Age, y</td>
<td>37.8 ± 7.6, 27–52</td>
</tr>
<tr>
<td>Duration of fibromyalgia symptoms, y</td>
<td>7.6 ± 6.3, 1–26</td>
</tr>
<tr>
<td>Duration of fibromyalgia since diagnosis</td>
<td>4.21 ± 2.59, 0.5–10</td>
</tr>
<tr>
<td>Mean dose of MC, g/mo</td>
<td>26 ± 8.3, 20–50</td>
</tr>
<tr>
<td>Duration of MC treatment, mo</td>
<td>10.4 ± 11.3, 1–42</td>
</tr>
<tr>
<td>No. tender points</td>
<td>15.7 ± 2.2, 12–18</td>
</tr>
<tr>
<td>No. patients with headache</td>
<td>25 (~96)</td>
</tr>
<tr>
<td>No. patients with irritable bowel/bladder syndrome</td>
<td>9 (~35)</td>
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</tbody>
</table>

DISCUSSION

The main finding of the present study was that MC treatment was associated with significant favorable outcomes in every item evaluated in the FIQR. In some cases, the improvement was so marked that the patients completely ceased treatments they had taken previously. In other cases, the patients significantly reduced the dose or type of medication they had taken prior to MC treatment.

The patients expressed the effects of the treatment in their own words, and their responses were dramatic. Very rarely as physicians have we encountered such responses in real-life medicine, except possibly among patients treated with steroids for inflammatory conditions, such as polymyalgia rheumatica or rheumatoid arthritis.

Examples of the patients’ responses were as follows: “I wish I had received this treatment when I was first diagnosed with fibromyalgia,” “I returned to be the same person as before,” “I regained my health,” and “This is a miraculous treatment.”

For some items of the FIQR, all the patients reported a favorable outcome. These included the effect of MC on pain and energy levels. For other items, the impact was less prominent, yet significant. These included the impact of MC on memory problems and daily activities of living, such as household activities (e.g., cleaning the house and changing bed sheets) and shopping (e.g., carrying grocery bags).

Another major benefit of MC treatment was a lack of serious adverse effects. The patients reported a few mild adverse effects, including dry mouth, redness of the eyes, and feeling hungry. These adverse effects appeared from the start of the treatment. The first 2 adverse effects were usually transient, lasting only a few weeks, and were mainly encountered in cases where the mode of MC was smoking. Many patients adapted to feeling hungry by eating prior to the use of MC.

The mean dose of MC consumed in the present study was relatively low (26 ± 8.2 g per month) as compared with that
Please rate your level of tenderness to touch. 8.74 ± 1.61, 5
Please rate your level of anxiety. 6.84 ± 3.68, 0
Please rate your level of memory problems. 6.96 ± 2.73, 0
Please rate your level of depression. 8.40 ± 1.38, 5–10
Please rate the level of your sleep. 9.23 ± 1.59, 4–10
Please rate your level of energy. 9.37 ± 0.79, 8
Please rate your level of pain. 9.21 ± 0.95, 7
Fibromyalgia prevented me from accomplishing goals for the week. 9.17 ± 1.06, 7
Go shopping for groceries. 8.67 ± 1.42, 5
Sit in a chair for 45 min. 8.89 ± 1.36, 6
Change bed sheets. 8.71 ± 2.18, 3
Climb 1 flight of stairs. 7.88 ± 1.83, 4
Lift and carry a bag full of groceries. 8.52 ± 1.68, 5
Vacuum, scrub, or sweep floors 9.51 ± 1.21, 7
Prepare a homemade meal. 8.38 ± 1.62, 5
Walk continuously for 20 min. 8.35 ± 2.13, 3

TABLE 3. Meds Consumed Prior to and Under MC Treatment

<table>
<thead>
<tr>
<th>Medication</th>
<th>Prior to MC Treatmenta</th>
<th>Under MC Treatmenta</th>
<th>P</th>
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</thead>
<tbody>
<tr>
<td>Simple analgesics</td>
<td>12 (~46)</td>
<td>3 (~15)</td>
<td>0.000</td>
</tr>
<tr>
<td>NSAIDs</td>
<td>19 (~73)</td>
<td>2 (~8)</td>
<td>0.000</td>
</tr>
<tr>
<td>Simple opiates</td>
<td>4 (~15)</td>
<td>0 (0)</td>
<td>0.055</td>
</tr>
<tr>
<td>Pregabalin</td>
<td>7 (~27)</td>
<td>0 (0)</td>
<td>0.005</td>
</tr>
<tr>
<td>Strong opiates</td>
<td>20 (~77)</td>
<td>5 (~19)</td>
<td>0.000</td>
</tr>
<tr>
<td>Benzodiazepines</td>
<td>7 (~27)</td>
<td>1 (~5)</td>
<td>0.027</td>
</tr>
<tr>
<td>Tricyclics</td>
<td>4 (~15)</td>
<td>0 (0)</td>
<td>0.055</td>
</tr>
<tr>
<td>Other antidepressants</td>
<td>8 (~31)</td>
<td>3 (~12)</td>
<td>0.107</td>
</tr>
</tbody>
</table>

NSAIDs indicates nonsteroidal anti-inflammatory drugs.

consumed by patients who receive MC for other indications, such as cancer pain. In the latter case, the dosage exceeded 60 g per month (personal communication, Dr. Salem Billan, oncologist, Rambam Medical Center, Haifa, oral communication, on July 1, 2017). In the present study, many patients continued to take 30 or even 20 g per month, which were the lowest starting doses, suggesting that consumption of approximately 1 g or less a day could be sufficient to control most symptoms of fibromyalgia. The findings of the present study should reassure health policy makers and health care providers that most fibromyalgia patients will remain on a relatively low dose.

There are no studies on tolerance among MC users. However, a previous study of other recreational cannabis users found no tolerance to subjective effects of cannabis.\(^2^1\)

In the current study, 12 patients (46%) reported either an improvement in their capacity to work or return to full-time work (data not shown). The aforementioned finding has implications for the patient, the patient's family, and society. A literature search revealed no studies on this issue of return to work, among patients treated with MC for different indications. However, a large study of the impact of illicit use of cannabis reported detrimental effects on employment and labor force.\(^2^2\)

Although previous research proposed a role for endocannabinoid deficiency in fibromyalgia,\(^2^3\) the potential role of endogenous cannabinoids in the pathogenesis of fibromyalgia remains unclear. More studies are needed to clarify their role. The distribution of cannabinoid receptors in the body may favor the proposed theory of central sensitization in the pathogenesis of fibromyalgia.

The long-term effects of MC treatment remain unclear. All MC request forms submitted to the Israeli Medical Cannabis Agency of the Ministry of Health that are signed by all patients by the Israeli Agency of the Ministry of Health that are signed by all patients (data not shown). The aforementioned finding has implications for the patient, the patient's family, and society. A literature search revealed no studies on this issue of return to work, among patients treated with MC for different indications. However, a large study of the impact of illicit use of cannabis reported detrimental effects on employment and labor force.\(^2^2\)

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TABLE 4. Adverse Effects of MC Treatment Reported by the Patients

<table>
<thead>
<tr>
<th>Adverse Effect</th>
<th>No. Patients (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Dry mouth</td>
<td>7 (~27)</td>
</tr>
<tr>
<td>Red eyes</td>
<td>7 (~27)</td>
</tr>
<tr>
<td>Hunger feeling</td>
<td>4 (~15)</td>
</tr>
</tbody>
</table>
the consistent findings of the impact of MC on many of the FIQR items, especially on pain, allude to the validity of the results.

The main drawback of the present study was its retrospective nature, where patients were asked to answer questions regarding the period prior to their use of MC. However, most patients (~54%) answered the questionnaire a relatively short period after starting MC treatment (i.e., ~3 months). Second, as mentioned earlier, fibromyalgia is not among the indications for MC treatment. Based on personal experience, fewer than 5% of requests for MC treatment for fibromyalgia are approved. Thus, it seemed unpractical to administer the questionnaire a priori to all patients with fibromyalgia whose doctor submitted a form for MC licensing.

REFERENCES


