# AGA Clinical Practice Update on Diagnosis and Management of Cannabinoid Hyperemesis Syndrome By: Andres Rodriguez, DO, MBA

## Background

- Cannabinoid Hyperemesis Syndrome (CHS) is a subtype of cyclical vomiting syndrome (CVS)
- Cannabis plant has 400+ chemicals -most importantly tetrahydrocannabinol (THC) and cannabidiol (CBD)
  - THC → psychoactive ingredient (not CBD).
- Marijuana is widely used-- 48.2 million people used in 2019
  - Medical marijuana legalized in 37 states, recreational in 27 states
  - Higher THC concentration in cannabis products being sold in dispensaries
- Unique adverse effects → nausea, vomiting, abdominal pain
  - Cases of up to 30 episodes of vomiting daily
- Paradoxical effects patients may note relief of symptoms such as emesis with cannabis use

# **Cyclical Vomiting Syndrome**

- Table 1. Rome IV Criteria for Cyclical Vomiting Syndrome in Adults<sup>®</sup>
- Stereotypical episodes of vomiting regarding onset (acute) and duration (<1 week)</li>
- At least 3 discrete episodes in the previous year and 2 episodes in the past 6 months, occurring at least 1 week apart
- Absence of nausea and vomiting between episodes, but other milder symptoms can be present between episodes
- Supportive remarks: history or family history of migraine headaches

<sup>a</sup>Criteria fulfilled for the last 3 months with symptom onset at least 6 months before diagnosis. Adapted from Stanghellini et al.<sup>1</sup>

#### Inter-episodic phase

- Absent or minimal symptoms
- 30% may have dyspepsia or nausea
- Prophylactic medications in moderate to severe CVS: tricyclics, anti-convulsants, mitochondrial supplements (CoQ10, vitamin B2), NK1 antagonist

**Recovery phase** 

Start oral intake and advance as

Triggers: stressful events (positive

or negative), sleep deprivation,

Resume regular activities

Vomiting subsides

tolerated

infections

Triptans
 Antiemetics: NK1 or histamine
 H1-antagonists, benzodiazepines

#### / Emetic phase

- Relentless vomiting, retching, abdominal pain, and neurological symptoms
- Extreme thirst, drink large volumes of water

**Prodromal phase** 

Nausea and abdominal pain

Autonomic symptoms (e.g.,

Panic and "out of control"

Take abortive medications

Median 30-90 minute duration

sweating, feeling hot or cold)

- Relief with empty stomach; induce emesis
- Rest in a quiet dark room
- · Supportive treatment

# Epidemiology

- Prevalence in emergency departments (ED), primary care, and gastroenterology clinics
  - Prevalence of CHS in EDs doubled between 2017-2021 in North America
  - Highest prevalence: 16-34 age range
- Males > females
- Cannabinoid hyperemesis syndrome (CHS)
  - associated with chronic (typically years) and heavy (usually daily / near-daily) cannabis use



Citations: Rubio-Tapia, et al. (2024). AGA Clinical Practice Update on Diagnosis and Management of Cannabinoid Hyperemesis Syndrome: Commentary. Gastroenterology 2024; 166: 930-934; UpToDate - Cannabinoid Hyperemesis Syndrome

#### Pathophysiology

### **Diagnostic Criteria for CHS**

### Management

- THC activates these receptors:
  - CB1 → brain and gut
    - effects on anxiety, depression, GI secretions, emesis and appetite control
  - CB2 → inflammatory (immunocyte/macrophage) and epithelial cells/neurons (sensory neurons)
    - effects on inflammation and nociception
  - Transient receptors of vanilloid
    type 1 channel
    - affects vagus nerve and gut function

# The Emoroid Digest

## 1. Clinical features

- episodic vomiting episodes,
  3+ episodes annually
- 2. Cannabis use patterns
  - duration of cannabis use > 1 year before symptom onset
  - frequency more than 4 times per week, on average
- 3. Cannabis cessation
  - resolution of symptoms after a period of abstinence from cannabis use for at least 6 months or at least equal to the total duration of 3 typical vomiting cycles in that patient

\* a negative result on a urine THC metabolite immunoassay (i.e. drug screen) likely excludes CHS

- Mainstay of treatment: TCA
  - Amitriptyline with minimal effective dose being 75-100 mg at bedtime
    - start with 25 mg → up-titrate with weekly increments to reach minimal effective dose with close monitoring of adverse effects
- Topical capsaicin (0.1%) cream applied to upper abdomen
  - may improve symptoms by activation of transient receptor potential vanilloid type 1 receptors
- Avoid X opioids
  - Can worsen nausea
  - addiction risk, tolerance, dependence
- Anti-emetics as needed
- Counseling to achieve marijuana cessation is necessary for successful treatment
  - Stopping use immediately may incite withdrawal symptoms and high frequency of recidivism
- Co-management with psychologist/psychiatrist may be helpful
- Ongoing clinical trials regarding topical capsaicin, benzodiazepines, haloperidol, promethazine, olanzaprine, ondansetron

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