

AGA Clinical Practice Update on Diagnosis and Management of Cannabinoid Hyperemesis Syndrome

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Background

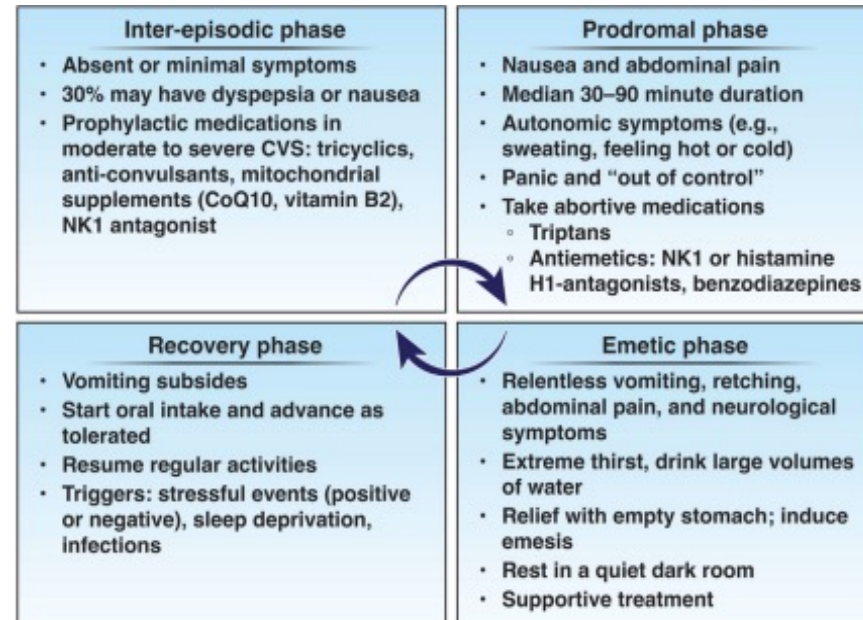
- Cannabinoid Hyperemesis Syndrome (CHS) is a subtype of cyclical vomiting syndrome (CVS)
- Cannabis plant has 400+ chemicals -- most importantly tetrahydrocannabinol (THC) and cannabidiol (CBD)
 - THC → psychoactive ingredient (not CBD).
- Marijuana is widely used-- 48.2 million people used in 2019
 - 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^a

- Stereotypical episodes of vomiting regarding onset (acute) and duration (<1 week)
- 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

^aCriteria fulfilled for the last 3 months with symptom onset at least 6 months before diagnosis. Adapted from Stanghellini et al.¹



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



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Pathophysiology

- 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

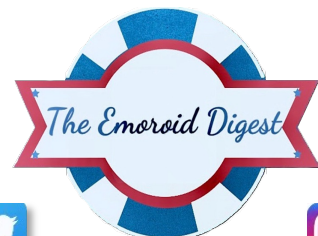
Diagnostic Criteria for CHS

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

Management

- 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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