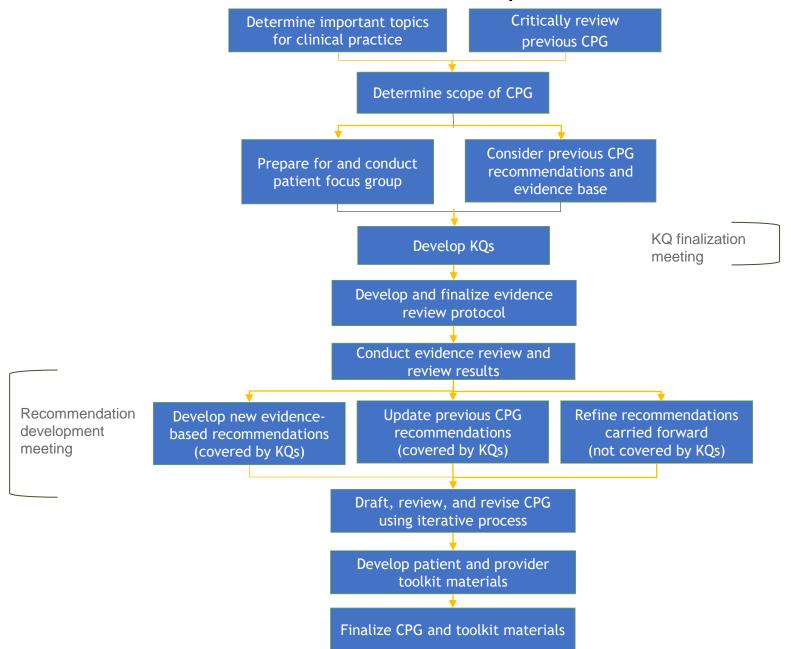


Introduction to VA/DoD Clinical Practice Guideline for the Management of Headache Natasha Antonovich, PharmD, BCPS Aven W. Ford, M.D. Lt Col, USAF, MC, FS Jason J. Sico, MD, MHS, FAHA, FACP, FAAN, FAHS Rebecca Vogsland, DPT, OCS

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Overview of CPG Development Process





Grading Recommendations - GRADE

- Evidence-based clinical practice recommendations were developed based on the:
 - Evidence review, which was informed by 12 key questions
 - GRADE (Grading of Recommendations Assessment, Development and Evaluation) methodology and use of four decision domains to determine strength (*Strong* or *Weak*) and direction (*For* or *Against*) of each recommendation:
 - Confidence in the quality of evidence
 - Balance of desirable and undesirable outcomes
 - Values and preferences
 - Other implications, as appropriate (e.g., resource use)



Strength of a Recommendation

- Strength of a recommendation on a continuum:
 - Strong for (or "We recommend...")
 - Weak for (or "We suggest...")
 - **Neither for nor against** (or "There is insufficient evidence...")
 - Weak against (or "We suggest against...")
 - Strong against (or "We recommend against...")



Headache is extremely common

- Headache is the second leading cause of years lived with disability across all age groups, trailing only low back pain.
- Ten percent of people living with headache report having multiple different types of headaches each week, and 3% report having some type of headache daily.

Headache is disabling

- More disability-adjusted life years (DALY) are attributable to headache than all other neurological disorders combined.
- Headache disability is linked to headache characteristics (e.g., throbbing, stabbing), frequency (e.g., hundreds of times a day, annually), associated features (e.g., nausea, photophobia, unilateral weakness), and conditions highly comorbid with headache (e.g., depression, stroke).
- Furthermore, health-related QoL scores, a measure of an individual's perceived mental and physical health over time, might decrease during a headache and in periods between headaches.
- · Headaches negatively affect family life, group activities, relationships, and financial stability.



Headache carries stigma

- Stigma, or "a set of negative and often unfair beliefs that a society or group of people have about something," is commonly experienced by those living with migraine and other types of headache and is increasingly being recognized as an important contributor to headache disability.
- Stigma can worsen headache symptoms and is associated with impaired QoL.
- Stigma contributes to many patients **not reporting their headaches** to healthcare providers, causing **significant delay** in diagnosis and treatment.

Language to reduce stigma

- For example, the term *"attack"* is frequently used within the headache community to distinguish among the acute symptoms an individual is experiencing from the chronic disease itself, yet it **can hold negative connotations**, often signifies an external locus of control, and might have strong connotations for Veterans and active duty Service members.
- Although communicating the severity and fluctuating nature of headache diseases is essential, you might consider using phrases such as "symptom onset" or "symptom escalation" rather than the word attack.



· Headache is associated with other conditions.

- Studies have found significant relationships between migraine and placental abruption, preeclampsia, and stroke during pregnancy.
 - Given the high prevalence and increased risk of adverse outcomes related to migraine in individuals of childbearing age, discussion regarding contraception and early treatment to reduce the burden of disease while minimizing teratogenic effects should be considered among this population.
- Conditions that more frequently co-occur with migraine than in those without include:
 - Insomnia, depression, anxiety, gastric ulcers/GI bleeding, chest pain, and epilepsy
- Conditions that were associated with a higher degree of migraine headache pain include:
 - Inflammatory conditions, psychiatric disorders, and sleep disorders.
- Conditions that are associated with higher migraine headache frequency include:
 - Gastric ulcers/GI bleeding, diabetes, anxiety, depression, insomnia, asthma, and allergies/hay fever
- Though most headaches are not a sign of another illness, headache can be the presenting symptom of a more serious systemic or neurologic condition.
 - Even when red flag features are absent and evaluations are appropriate significant concern may linger for patients and providers.



Headache is costly

- The estimated direct and indirect medical costs of caring for people with migraines in the U.S. is approximately **\$36 billion annually**.
- Sixty percent of costs are accounted for by physician office visits.
- The indirect annual cost is largely attributed to missed days of work (i.e., **absenteeism**) and impaired work function when people come to work while impaired by their headache (i.e., **presenteeism**).



• Disparities in headache care are poorly understood

- Understanding health disparities as they relate to headache care <u>has largely been unexplored</u>. Men are historically underrepresented in headache research, including clinical trials and epidemiological work. For example, historically more than 80% of subjects enrolled in migraine clinical trials have been women, whereas 43% of women and 18% of men have migraine sometime during their lifetime.
- Beyond gender differences in headache care, limited evidence suggests poorer health care utilization, more inaccurate diagnoses, and poorer care quality among Black patients compared with White, non-Hispanic patients with migraine.
- Other marginalized and underserved groups also seem to bear disproportionate burden of migraine, including Hispanics and Latinos, people with low socioeconomic status, and persons living in rural areas.



Headache in VA patients

- Headache appears to be getting more common.
 - From 2008 to 2019, the 1-year prevalence of Veterans **diagnosed with migraine has steadily increased**, from 8.5% to 13.0% and from 1.1% to 2.5%, for women and men respectively.
- Headache in VA patients is complex.
 - More than one half of veterans diagnosed with migraine have multiple headache diagnoses, compared to ten percent of the general headache population.
 - In fiscal year 2017, approximately 380,000 Veterans sought care in the VA system for a headache disorder and more than 75% of headache management occurred within primary care.
- History of traumatic brain injury (TBI) is common.
 - TBI is a strong predictor of headache as a symptom in the first year of care for a Veteran within VA, with the severity of TBI and history of recurrent TBIs associated with greater headache severity.
- Psychiatric comorbidities are common.
 - Psychiatric comorbidities increase the likelihood of headache among those with a TBI diagnosis.





Headache in DoD patients

- Headaches are common in active duty Service members and their dependents
 - In a longitudinal study including a large cohort of 77,000 participants (active duty Service members, Reservists, and National Guard), the self-reported prevalence of provider-diagnosed migraine was 6.9% in males and 20.9% in females.
- Posttraumatic headache (PTH) is common, mirroring increased rates of mild TBI in the DoD population
 - Mild traumatic brain injury can result in a complex set of physical, behavioral, and cognitive symptoms.
 - The incidence of **mTBI** and concurrent headache in the DoD population is **four to five times higher** than that in the general U.S. population.
- DoD patients may be especially prone to healthcare avoidance
 - Active duty patients, especially those with special duties, **may hesitate to report headaches** or other painful conditions to their health care providers. This hesitation might stem from not wanting to appear "weak," not wanting to be taken off special duties or assignments, or both.
 - A recent study showed that military pilots self-report healthcare avoidance because of fear of loss of flight status.
 - This avoidance causes significant delay in accurately diagnosing and treating headache conditions and might lead to additional care challenges for the patients and providers.



Active duty, Guard, and Reserve patients may have occupationally specific concerns

- Providers must be diligent in reviewing their patient's occupational history and job-specific duties before developing a treatment plan.
- A clinically indicated and medically appropriate medication **might lead to duty limitations** or even a review for military retention.
- Providers should strongly consider placing **duty and mobility restrictions** on active duty Service members when starting new medications, even if only temporarily, to allow for evaluation of treatment effect and assessment of potential side effects.
- **Occupational medicine** and flight medicine specialists can be consulted by primary care as needed for guidance and recommendations.





		Tension-Type Headache ª	Migraine Headache ^b	Cluster Headache °
Headache	Duration	30 minutes to 7 days	4–72 hours	15–180 minutes
Symptom Duration and Frequency	Frequency	Variable	Variable	Once every other day to eight per day; often occurring at the same time of day
	Severity	Mild to moderate	Moderate to severe	Severe or very severe
Headache	Location	Bilateral Unilateral		Unilateral orbital, supraorbital, or temporal pain or any combination of such pain
Characteristics	Quality	Ility Pressing or tightening, non- pulsating Throbbing or pulsating		Stabbing, boring
	Aggravated by routine physical activity	Not aggravated by routine activity	Aggravated by routine activity	Causes a sense of agitation or restlessness; routine activity might improve symptoms
Associated	Photophobia and phonophobia	Can have one but not both	Both	Variably present
Features	Nausea, vomiting, or both	Neither	Either or both	Might be present
Other Features Autonomic features ^d None		None	Might occur but are often subtle and not noticed by the patient	Prominent autonomic features ipsilateral to the pain (see <u>Appendix B</u>)

Headache types

- Tension-type headache is the most common headache type the general population; however, migraines tend to be more debilitating and are more common in patients presenting for medical care.
- Within the U.S., the prevalence of selfreported migraine, severe headache, or both ranges between 15–18% in women and 6–10% in men; nearly one-half of women and men experience TTH.
 - However, migraine is often misdiagnosed in men and 6-10% may be a significant underestimation.



- Tension-type headache
 - At least 10 headaches lasting 30 minutes to 7 days
 - At least two defining characteristics
 - Bilateral location
 - Non-pulsating quality
 - Mild to moderate intensity
 - Not aggravated by routine physical activity
 - Both associated features
 - No nausea or vomiting
 - Either photophobia or phonophobia, but not both
 - If headaches fulfill all but one of the TTH criteria (e.g., having both photophobia and phonophobia), the diagnosis would be probable TTH.
 - Chronic vs. episodic
 - Chronic refers to having frequent headaches occurring on 15 or more days per month for more than 3 months.





- Migraine without aura
 - At least five headaches lasting 4–72 hours
 - At least two defining headache characteristics:
 - Unilateral
 - Throbbing or pulsating
 - Moderate or severe intensity
 - Aggravated, or caused by routine physical activity
 - At least one associated feature
 - Nausea, vomiting, or both
 - Both photophobia and phonophobia
 - If headaches fulfill all but one of the migraine criteria (e.g., photophobia or phonophobia but not photophobia and phonophobia), the diagnosis would be probable migraine.





- Migraine with aura vs. without aura
 - Aura; Recurrent attacks, lasting minutes, of unilateral fully-reversible visual, sensory or other central nervous system symptoms that usually develop gradually and are usually followed by headache and associated migraine symptoms.
- Chronic vs. episodic
 - Headache occurring on 15 or more days/month for more than 3 months, which, on at least 8 days/month, has the features of migraine headache.



- Cluster headache
 - · At least five attacks fulfilling the criteria below
 - Severe or very severe unilateral orbital, supraorbital and/or temporal pain lasting 15-180 minutes (when untreated)
 - Either or both of the following:
 - at least one of the following symptoms or signs, ipsilateral to the headache:
 - conjunctival injection and/or lacrimation
 - nasal congestion and/or rhinorrhoea
 - eyelid oedema
 - forehead and facial sweating
 - miosis and/or ptosis
 - a sense of restlessness or agitation
 - Occurring with a frequency between one every other day and 8 per day
 - Not better accounted for by another ICHD-3 diagnosis.





- Cluster headache
 - Called "Episodic" when there have been at least two cluster periods lasting from 7 days to 1 year (when untreated) and separated by pain-free remission periods of ≥3 month.
 - Clusters usually last between 2 weeks and 3 months.
 - Called "Chronic" when attacks occur without a remission period, or with remissions lasting <3 months, for at least 1 year.



Primary Headache Disorders Criteria*			
	Tension-type headache	Migraine headache	Cluster headache
	Attack	duration and frequency	
Duration	30-minutes – 7-days	4 – 72 hours	15 – 180 minutes
Frequency	Variable	Variable	Once every other day to eight per day; often occurring at the same time of day
	Head	ache characteristics	
Severity	Mild to moderate	Moderate to severe	Severe or very severe
Location	Bilateral	Unilateral	Unilateral orbital, supraorbital, and/or temporal
Quality	Pressing or tightening, non- pulsating	Throbbing or pulsating	Stabbing, boring
Aggravated by routine physical activity	Not aggravated by routine activity	Aggravated by routine activity	Causes a sense of agitation or restlessness; routine activity may improve symptoms
Associated features			
Photophobia and phonophobia	Can have one but not both	Both	Variably present
Nausea and/or vomiting	Neither	Either or both	May be present
Other features			
Autonomic features	None	May occur, but are often subtle and not noticed by the patient	Prominent autonomic features ipsilateral to the pain



• Secondary headache:

- When a new headache occurs for the first time in close temporal relation to another disorder that is known to cause headache, or fulfils other criteria for causation by that disorder, the new headache is coded as a secondary headache attributed to the causative disorder. This remains true even when the headache has the characteristics of a primary headache (migraine, tension-type headache, cluster headache or one of the other trigeminal autonomic cephalalgias).
- When a pre-existing primary headache becomes chronic or is made significantly worse (usually meaning a twofold or greater increase in frequency and/or severity) in close temporal relation to such a causative disorder, both the primary and the secondary headache diagnoses should be given, provided that there is good evidence that the disorder can cause headache.



General diagnostic criteria for secondary headaches:

- Any headache fulfilling criterion below
- Another disorder scientifically documented to be able to cause headache has been diagnosed with evidence of causation demonstrated by at least two of the following:
 - Headache has developed in temporal relation to the onset of the presumed causative disorder
 - Either or both of the following:
 - Headache has significantly worsened in parallel with worsening of the presumed causative disorder
 - Headache has significantly improved in parallel with improvement of the presumed causative disorder
 - · Headache has characteristics typical for the causative disorder
 - Other evidence exists of causation
- Not better accounted for by another ICHD-3 diagnosis.



• Types of secondary headache:

- Headache attributed to trauma or injury to the head and/or neck.
- Headache attributed to cranial or cervical vascular disorder.
- Headache attributed to non-vascular intracranial disorder.
- Headache attributed to a substance or its withdrawal.
- Headache attributed to infection.
- Headache attributed to disorder of homeostasis.
- Headache or facial pain attributed to disorder of the cranium, neck, eyes, ears, nose, sinuses, teeth, mouth or other facial or cervical structure.
- Headache attributed to psychiatric disorder.



Medication-overuse headache

- Headache occurring on ≥15 days/month in a patient with a pre-existing headache disorder
- Regular overuse for >3 months of one or more drugs that can be taken for acute and/or symptomatic treatment
 of headache
- Not better accounted for by another ICHD-3 diagnosis.



Medication-overuse headache

Medication Overuse Headache Type	Medication Overuse Frequency	
Butalbital overuse	≥5 days/month for >3 months	
Opioid overuse	≥8 days/month for >3 months	
Triptan overuse		
Ergotamine overuse		
Combination-analgesic overuse (any combination of classes, not to include combinations that only include non-opioid analgesics) ^c	≥10 days/month for >3 months	
Non-opioid analgesic overuse (e.g., aspirin, NSAIDs, acetaminophen, steroids, and combinations of non- opioid analgesics)	≥15 days/month for >3 months	

^c Combination-analgesic refers to a headache abortive medication that contains more than one active ingredient and may refer to over-the-counter or prescription agents.



Screening and Diagnosis

Recommendation			Category ^b
	Medication Overuse Headache Screening and Other Cons	iderations	
factor order • H • N • N • N • N • N • N • N • N • N • N	uggest providers assess for and consider the following high-risk rs for medication overuse headache in patients with headache (in of relative impact): deadache frequency (greater than or equal to 7 days per month) <i>A</i> ligraine diagnosis <i>M</i> edication use: frequent use of anxiolytics, analgesics (for any condition, including use of opioids or non-opioid analgesics for acute reatment of migraine), or sedative hypnotics distory of anxiety or depression, especially in combination with nusculoskeletal complaints or gastrointestinal complaints Physical inactivity Sick leave of greater than 2 weeks in the last year	Weak for	Not reviewed, Amended
• S	Self-reported whiplash		
• S	Smoking (tobacco use)		



Pharmacotherapy Recommendations – overarching principles

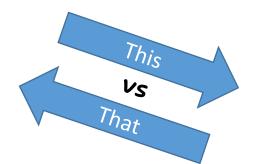
- Keep in mind recommendations are made **recent literature pull** (2016 – 2022). See Appendix A
- Some pharmacotherapies indicated for headache treatment are not in recommendations
 - This does not mean these are not options for patients
 - Examples: almotriptan, dihydroergotamine, zavegepant
- Full list of commonly used outpatient pharmacotherapies are in Appendix G of CPG
- We will review select pharmacotherapies' VA National Criteria for Use. The VA Formulary Advisor is available to search formulary status and, if applicable, Criteria For Use for any drug: <u>https://www.va.gov/formularyadvisor/</u>

		VA/DoD Clinical Practice (Guideline for Management of Headach	he
The foll prescrib	bing information for fu	ize pharmacotherapy optio	ns for preventive and abortive tr	reatment. Refer to each drug's
Туре	Drug	Initial Dose	Usual Range	Comments
υ	Atenolol	50 mg/day	50–200 mg/day	
Beta-Adrenergic Antagonists	Metoprolol tartrate and metoprolol succinate	100 mg/day in divided doses	100–200 mg/day in divided doses	Dose should be titrated and
-Adre tago	Nadolol	4080 mg/day	80–240 mg/day	maintained for at least 3 months before assessment of response
An	Propranolol	40 mg/day in divided doses	40–160 mg/day in divided doses]
ň	Timolol	20 mg/day in divided doses	20-60 mg/day in divided doses	1
ants	Amitriptyline	10 mg at bedtime	20–50 mg at bedtime	Use slow titration to reduce sedation.
Antidepressants	Nortriptyline	10 mg daily or at bedtime	20–50 mg daily or at bedtime	Use slow titration to reduce sedation.
Antie	Venlafaxine	37.5 mg/day	75–150 mg/day	Titrate dose weekly, as tolerated
convulsants	Topiramate	25 mg/day	50-200 mg/day in divided doses	 Increase by 25 mg/week.
	Valproic acid/ divalproex sodium	250–500 mg/day in divided doses or daily for extended	500–1,500 mg/day in divided doses or daily for extended release	May monitor levels if adherence an issue





New in 2023 update: Comparative Pharmacotherapy Evidence



- With all the new and existing pharmacotherapies approved for headache treatment, which is more effective?
- 50. There is **insufficient evidence** to recommend for or against any specific medication over another for the acute treatment of migraine.
- 51. There is insufficient evidence to recommend for or against any specific medication over another for the prevention of migraine headache, tension headache, or cluster headache.
- What this means there is no evidence that exists to suggest that any pharmacotherapy with a "for" recommendation is any better than another



Pharmacologic "For" Migraine Recommendations

- Prevention
 - 4. We recommend **candesartan or telmisartan** for the prevention of episodic migraine
 - 5. We recommend **erenumab**, **fremanezumab**, **or galcanezumab** for the prevention of episodic or chronic migraine
 - 6. We suggest intravenous eptinezumab for the prevention of episodic or chronic migraine
 - 7. We suggest lisinopril for the prevention of episodic migraine
 - 8. We suggest oral magnesium for the prevention of migraine
 - 9. We suggest **topiramate** for the prevention of episodic and chronic migraine.
 - 10. We suggest propranolol for the prevention of migraine.
 - 11. We suggest valproate for the prevention of episodic migraine.
 - 12. We suggest **memantine** for the prevention of episodic migraine.
 - 13. We suggest **atogepant** for the prevention of episodic migraine.



Pharmacologic "For" Migraine Recommendations

Prevention

- 4. We recommend candesartan or telmisartan for the prevention of episodic migraine
- 5. We recommend **erenumab**, **fremanezumab**, **or galcanezumab** for the prevention of episodic or chronic migraine
- 6. We suggest intravenous **eptinezumab** for the prevention of episodic or chronic migraine
- 7. We suggest **lisinopril** for the prevention of episodic migraine
- 8. We suggest oral magnesium for the prevention of migraine
- 9. We suggest **topiramate** for the prevention of episodic and chronic migraine.
- 10. We suggest **propranolol** for the prevention of migraine.
- 11. We suggest valproate for the prevention of episodic migraine.
- 12. We suggest **memantine** for the prevention of episodic migraine.
- 13. We suggest **atogepant** for the prevention of episodic migraine.



Erenumab: VA's Formulary with Prior Authorization CGRP antibody.

What prior medication trials are required for chronic migraine in Criteria for Use?

- 9. We suggest **topiramate** for the prevention of episodic and chronic migraine.
- 10. We suggest **propranolol** for the prevention of migraine.
- 11. We suggest **valproate** for the prevention of episodic migraine.

Contraindication, intolerance, or lack of therapeutic response after at least 12 weeks of a therapeutic dose of **one** beta blocker (e.g. metoprolol 50-100 mg BID, propranolol 20-80 mg BID)¹

- Contraindication, intolerance, or lack of therapeutic response after at least 12 weeks of a therapeutic dose of topiramate 50-200 mg BID¹
- Contraindication, intolerance, or lack of therapeutic response after at least 12 weeks of a therapeutic dose of divalproex 500-1000 mg daily (divalproex is not recommended in patients who can become pregnant)¹

This is an **excerpt**. Full CFU for erenumab at: <u>https://www.va.gov/</u> formularyadvisor/

Erenumab-aooe CFU



Pharmacologic "For" Migraine Recommendations

- Acute
 - 19. We recommend eletriptan, frovatriptan, rizatriptan, sumatriptan (oral or subcutaneous), the combination of sumatriptan and naproxen, or zolmitriptan (oral or intranasal) for the acute treatment of migraine
 - 20. We recommend **aspirin/acetaminophen/caffeine** for the acute treatment of migraine
 - 21. We **suggest** acetaminophen, aspirin, ibuprofen, or naproxen for the acute treatment of migraine
 - 22. We suggest rimegepant or ubrogepant for the acute treatment of migraine



Pharmacologic "For" Migraine Recommendations

• Acute

- 19. We **recommend** eletriptan, frovatriptan, rizatriptan, sumatriptan (oral or subcutaneous), the combination of sumatriptan and naproxen, or zolmitriptan (oral or intranasal) for the acute treatment of migraine
- 20. We **recommend** aspirin/acetaminophen/caffeine for the acute treatment of migraine
- 21. We suggest **acetaminophen**, **aspirin**, **ibuprofen**, or **naproxen** for the acute treatment of migraine
- 22. We suggest rimegepant or ubrogepant for the acute treatment of migraine



Ubrogepant: VA Non-formulary gepant. What prior medication trials are required in Criteria for Use?

 19. We recommend eletriptan, frovatriptan, rizatriptan, sumatriptan (oral or subcutaneous), the combination of sumatriptan and naproxen, or zolmitriptan (oral or intranasal) for the acute treatment of migraine.

Inclusion Criteria

The answers to **all** of the following must be fulfilled in order to meet criteria.

- Treatment initiated by a VA/VA Community Care neurologist or locally designated headache expert
- Diagnosis of migraine, with or without aura, per the International Classification of Headache Disorders (ICHD-3)
-] Moderate to severe migraine intensity
- Currently receiving preventive therapy for migraine if indicated
- Contraindication¹, intolerance, or lack of response to trial of two different triptans at a clinically effective dose.

This is an **excerpt.** Full CFU for ubrogepant at: <u>https://www.va.gov/</u> formularyadvisor/



Pharmacologic "Neither For Nor Against" and "Against" Migraine Recommendations

- Preventive
 - 16. There is insufficient evidence to recommend for or against rimegepant for the prevention of episodic migraine
 - 18. There is insufficient evidence to recommend for or against levetiracetam for the prevention of episodic migraine
 - 17. We suggest against the use of **gabapentin** for the prevention of episodic migraine
- Acute
 - 24. There is insufficient evidence to recommend for or against **lasmiditan** for the acute treatment of migraine
 - 23. We suggest against intravenous ketamine for the acute treatment of migraine



Pharmacologic "Neither For Nor Against" and "Against" Migraine Recommendations

Preventive

- 16. There is insufficient evidence to recommend for or against **rimegepant** for the prevention of episodic migraine
- 18. There is insufficient evidence to recommend for or against levetiracetam for the prevention of episodic migraine
- 17. We <u>suggest against</u> the use of **gabapentin** for the prevention of episodic migraine

• Acute

- 24. There is insufficient evidence to recommend for or against **lasmiditan** for the acute treatment of migraine
- 23. We <u>suggest against</u> intravenous ketamine for the acute treatment of migraine



Pharmacologic Tension-Type Headache Recommendations

- Preventive
 - 25. We <u>suggest</u> **amitriptyline** for the prevention of chronic tension-type headache.
- Acute
 - 27. We <u>suggest</u> **ibuprofen** (400 mg) or **acetaminophen** (1,000 mg) for the acute treatment of tension-type headache.



- Preventive
 - 28. We <u>suggest</u> galcanezumab for the prevention of episodic cluster headache
 - 30. There is insufficient evidence to recommend for or against verapamil for the prevention of episodic or chronic cluster headache
 - 29. We <u>suggest against</u> galcanezumab for the prevention of chronic cluster headache.
- Acute
 - 31. We <u>suggest</u> subcutaneous sumatriptan (6 mg) or intranasal zolmitriptan (10 mg) for the acute treatment of cluster headache.
 - 32. We <u>suggest</u> the use of **normobaric oxygen therapy** for the acute treatment of cluster headache



Preventive

- 28. We <u>suggest</u> galcanezumab for the prevention of episodic cluster headache
- 30. There is <u>insufficient evidence</u> to recommend for or against **verapamil** for the prevention of episodic or chronic cluster headache
- 29. We <u>suggest against</u> galcanezumab for the prevention of chronic cluster headache
- Acute
 - 31. We <u>suggest</u> subcutaneous sumatriptan (6 mg) or intranasal zolmitriptan (10 mg) for the acute treatment of cluster headache
 - 32. We <u>suggest</u> the use of normobaric oxygen therapy for the acute treatment of cluster headache



Preventive

- 28. We <u>suggest</u> galcanezumab for the prevention of episodic cluster headache
- 30. There is insufficient evidence to recommend for or against verapamil for the prevention of episodic or chronic cluster headache
- 29. We <u>suggest against</u> galcanezumab for the prevention of chronic cluster headache

• Acute

- 31. We <u>suggest</u> subcutaneous sumatriptan (6 mg) or intranasal zolmitriptan (10 mg) for the acute treatment of cluster headache
- 32. We <u>suggest</u> the use of normobaric oxygen therapy for the acute treatment of cluster headache



Preventive

- 28. We <u>suggest</u> galcanezumab for the prevention of episodic cluster headache
- 30. There is insufficient evidence to recommend for or against verapamil for the prevention of episodic or chronic cluster headache
- 29. We suggest against galcanezumab for the prevention of chronic cluster headache

• Acute

- 31. We <u>suggest</u> subcutaneous **sumatriptan** (6 mg) or intranasal **zolmitriptan** (10 mg) for the acute treatment of cluster headache
- 32. We <u>suggest</u> the use of normobaric oxygen therapy for the acute treatment of cluster headache



Pharmacologic recommendations for Headache

- Evidence for medications in these recommendations included multiple types of headache.
- Preventive:
 - 2. There is <u>insufficient evidence</u> to recommend for or against co enzyme Q10, feverfew, melatonin, omega-3, vitamin B2, or vitamin B6 for the prevention of headache
 - 3. There is <u>insufficient evidence</u> to recommend for or against **fluoxetine** or venlafaxine for the prevention of headache



Medication Overuse Headache

 33. There is <u>insufficient evidence</u> to recommend for or against the addition of any specific preventive agent or withdrawal strategy to guide the treatment of medication overuse headache.



Injections, Procedures, and Invasive Interventions for the Acute Treatment of Headache

- Migraine
 - 34 We <u>suggest</u> greater occipital nerve block for the acute treatment of migraine
 - 36 There is <u>insufficient evidence</u> to recommend for or against supra orbital nerve block for the acute treatment of migraine
 - 37 There is <u>insufficient evidence</u> to recommend for or against intravenous **antiemetics** (i.e., chlorpromazine, metoclopramide, prochlorperazine), **intravenous magnesium** or **intranasal lidocaine** for the acute treatment of headache



Injections, Procedures, and Invasive Interventions for the Prevention of Headache

- Migraine
 - 14 We <u>suggest</u> onabotulinumtoxinA injection for the prevention of chronic migraine
 - 15 We suggest against onabotulinumtoxinA injection for the prevention of episodic migraine
 - 35 There is <u>insufficient evidence</u> to recommend for or against greater occipital nerve block for the prevention of chronic migraine

- Tension-Type Headache
 - 26 We suggest <u>against</u> botulinum/neurotoxin injection for the prevention of chronic tension-type headache



Non-Pharmacologic Therapies – Neuromodulation

- 41 We suggest non-invasive vagus nerve stimulation for the treatment of episodic cluster headache
- 48 There is insufficient evidence to recommend for or against any form of neuromodulation for the treatment and/or prevention of migraine:
 - Non-invasive vagus nerve stimulation
 - Supraorbital, or external trigeminal nerve, nerve stimulation
 - Remote electrical neurostimulation
 - External combined occipital and trigeminal neurostimulation system
 - Repetitive transcranial magnetic stimulation
 - Transcranial direct current stimulation



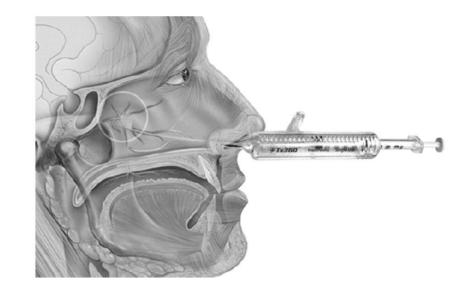
Injections, Procedures, and Invasive Interventions for the Prevention of Headache

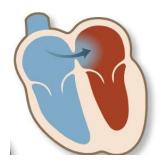
- Migraine
 - 14 We suggest onabotulinumtoxinA injection for the prevention of chronic migraine
 - 15 We suggest against onabotulinumtoxinA injection for the prevention of episodic migraine
 - 35 There is <u>insufficient evidence</u> to recommend for or against greater occipital nerve block for the prevention of chronic migraine

- Tension-Type Headache
 - 26 We suggest <u>against</u> botulinum/neurotoxin injection for prevention of chronic tension-type headache

Injections, Procedures, and Invasive Interventions for the Prevention of Headache

- 38 There is insufficient evidence to recommend for or against greater pulsed radiofrequency procedure of the upper cervical nerves of sphenopalatine ganglion block for the treatment of chronic migraine
- 39 We <u>suggest against</u> implantable sphenopalatine ganglion stimulator for the treatment of cluster headache
- 40 We <u>suggest against</u> patent foramen ovale closure for the treatment or prevention of migraine







Non-Pharmacologic Therapies – Rehabilitation Approaches and Exercise

- 42 We <u>suggest</u> **physical therapy** for the management of tension-type, migraine, or cervicogenic headache
- 43 We <u>suggest</u> aerobic exercise or progressive strength training for the prevention of tension-type and migraine headache
- 45 There is <u>insufficient evidence</u> to recommend for or against acupuncture, dry needling, or yoga for the treatment and/or prevention of headache







Non-Pharmacologic Therapies – Behavioral Interventions

- 44 There is <u>insufficient evidence</u> to recommend for or against the following **behavioral** interventions for the treatment and/or prevention of headache:
 - Biofeedback and smartphone application-based heartrate variability monitoring
 - Cognitive behavioral therapy
 - Mindfulness-based therapies
 - Progressive muscle relaxation





Non-Pharmacologic Therapies – Dietary Triggers and Testing



- 46 There is <u>insufficient evidence</u> to recommend for or against the **dietary trigger avoidance** for the prevention of headache
- 47 We suggest against immunoglobulin G antibody testing for dietary trigger avoidance for the prevention of headache

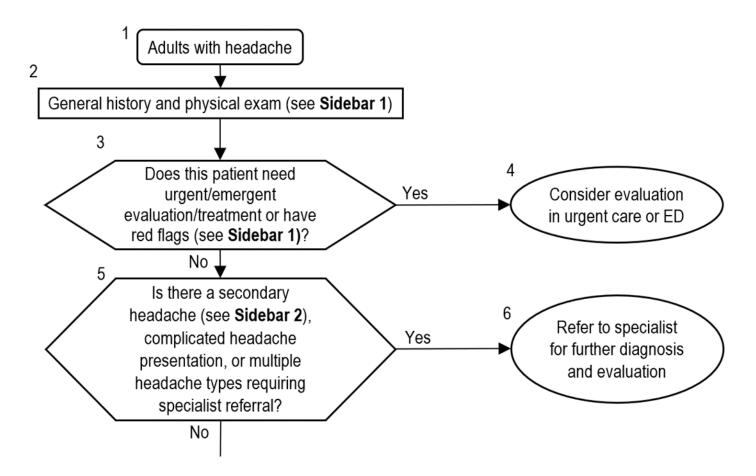


Comparative Effectiveness and Combination Therapies

- 49 to 51 There is insufficient evidence to recommend for or against:
 - Any specific medication over another for the acute treatment of migraine
 - Any specific medication over another for the prevention of migraine, tension type, or cluster headache
 - Choosing a specific treatment strategy for posttraumatic headache
 - 52 There is insufficient evidence to recommend for or against:
 - Combination of therapies for the prevention of headache



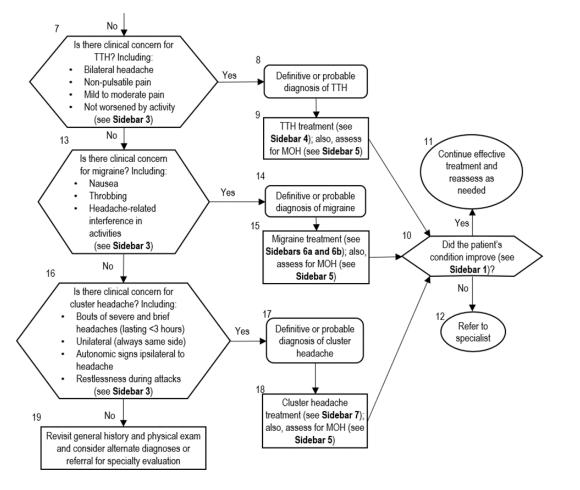
Algorithm – Module A: Evaluation and Treatment of Headache



Note: Box 5 connects to Box 7 on the next slide



Algorithm – Module A: Evaluation and Treatment of Headache (cont.)



Note: Box 7 connects to Box 5 on the previous slide



About the CPG

The guideline is formatted as a single clinical algorithm and 27 evidence-based recommendations:

Questions about the Headache Guideline

Guideline Links	Patient Provider Tools	Related Guidelines					
Headache Full Guideline (2020) 📩	Headache Patient Summary (2020) 📆	Concussion-mTBI					
Headache Provider Summary (2020)	Headache Diagnosis Coding Tool (2021)	Posttraumatic Stress Disorder (PTSD)					
Headache Pocket Card (2020) 📩	Headache Patient 7 Day Diary (2021)	Opioid Therapy (OT) for Chronic Pain					
	Headache Patient 3 Months Diary (2021)						
	Types of Headache Handout (2021)						

Webinars	Headache in Peer Reviewed Publications
Introduction to the New VA/DOD CPG: The Primary Care Management of Headache Webinar (2020)	Synopsis of the 2020 Headache CPG (2022)



https://www.healthquality.va.gov/guidelines/pain/headache/

Patient Provider Tools	
Headache Patient Summary (2020)	因
Headache Diagnosis Coding Tool (2021)	因
Headache Patient 7 Day Diary (2021)	因
Headache Patient 3 Months Diary (2021)	因
Types of Headache Handout (2021)	因

7-DAY HEADACHE DIARY This form can be printed and filled in manually, or completed on a computer. Write down your headache information DAILY to share with your headaches correctly. Check the boxes of the topic that apply to you each day.

ame: _				Prophyla	dis:		Clear Button
Date	Prevention	Headache	Symptoms	Warning Signs	Medication / Device	Lifestyle	Behavioral Coping
	Medication Device Behaviors	Pain (0-10): Start time: End time:	Sensitive to: Light Sound Nausea Vomiting Worse with activity	Aura	Medication: Time: Dose: Device: Time:	Stress (0-10): Headache interference (0-10): Hours slept: Sleep quality: Physically active Skipped meal Hydration Caffeine	-
	Medication Device Behaviors	Pain (0-10): Start time: End time:	Sensitive to: Light Sound Nausea Vomiting Worse with activity	Aura	Medication: Time: Dose: Device: Time:	Stress (0-10): Headache interference (0-10): Hours slept: Sleep quality: Physically active Skipped meal Hydration Gaffeine	-
	Medication Device Behaviors	Pain (0-10): Start time: End time:	Sensitive to: Light Sound Nausea Vomiting Worse with activity	Aura	Medication: Time: Dose: Device: Time:	Stress (0-10):	-
	Medication Device Behaviors	Pain (0-10): Start time: End time:	Sensitive to: Light Sound Nausea Vomiting Worse with activity	Aura	Medication: Time: Dose: Device: Time:	Stress (0-10): Headache interference (0-10): Hours slept: Sleep quality: Physically active Skipped meal Hydration Gaffeine	
	Medication Device Behaviors	Pain (0-10): Start time: End time:	Sensitive to: Light Sound Nausea Vomiting Worse with activity	Aura	Medication: Time: Dose: Device: Time:	Stress (0-10): Headache interference (0-10): Hours slept: Sleep quality: Physically active Skipped meal Hydration Caffeine	-
	Medication Device Behaviors	Pain (0-10): Start time: End time:	Sensitive to: Light Sound Nausea Vomiting Worse with activity	Aura	Medication: Time: Dose: Device: Time:	Stress (0-10): Headache interference (0-10): Hours slept: Sleep quality: Physically active Skipped meal Hydration Caffeine	-
	Medication Device Behaviors	Pain (0-10): Start time: End time:	Sensitive to: Light Sound Nausea Vomiting Worse with activity	Aura	Medication: Time: Dose: Device: Time:	Stress (0-10): Headache interference (0-10): Hours slept: Sleep quality: Physically active Skipped meal Hydration Caffeine	-

VA/DDD Clinical Practice Guideline for Primary Care Management of Headache. For more information, https://www.healthquality.va.gov/guidelines/pain/headache/VADoDHeadacheCPGPatientSummaryFinal508.pdl



https://www.healthquality.va.gov/guidelines/pain/headache/

Patient Provider Tools	
Headache Patient Summary (2020)	
Headache Diagnosis Coding Tool (2021)	Z
Headache Patient 7 Day Diary (2021)	X
Headache Patient 3 Months Diary (2021)	X
Types of Headache Handout (2021)	内

3-MONTH HEADACHE DIARY

This form can be printed and filled in manually, or completed on a computer. Save the file for future reference. Daily, write down all your headaches according to their severity. (1=mild, 2=moderate, 3=severe). Write the names of your acute medications in the IX squares on the left. Put a check if you used them for each day. Write down the efficacy of the medications (meds) (0=none, 1=partial, 2=success, attack was controlled). If you have notes, mark the square then add notes on the back or on the notes page.

Name: _____ Prophylaxis:

Month:	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23	24	25	26	27	28	29	30	31
Headache 0 1 2 3																															
Aura																															Γ
Menses																															F
Notes				⊢									H				Η													Η	┢
Tx				\vdash													H														F
Тх			\vdash	⊢		\vdash	\vdash	Η			\vdash		H	\vdash		\vdash	Η					Η	H		\square	\vdash	\vdash	Η	\vdash	Η	┢
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Effect of meds 0 1 2																															
Month:	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23	24	25	26	27	28	29	30	3
Headache 0 1 2 3																															
Aura																															Γ
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Effect of meds 0 1 2																															
Month:	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23	24	25	26	27	28	29	30	3
Headache 0 1 2 3																															
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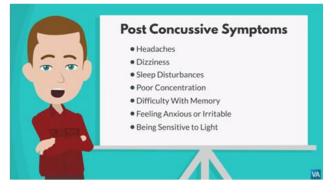


https://www.healthquality.va.gov/guidelines/pain/headache/

Health Psychology for Headache – YouTube



Post-Traumatic Headache - YouTube



Nutrition & Headache - YouTube



Exercise for Headache (Video) - YouTube





- Annie is named after Lieutenant Annie G. Fox, the first woman to receive the Purple Heart for combat. She was Chief Nurse in the Army Nurse Corps at Hickman Field, Pearl Harbor
- Annie uses SMS text messaging to promote self care in Veterans. Patients receive automated prompts to track/monitor health as well as motivational/educational messages.
- Headache protocol is assigned by provider/designee or veteran can self subscribe
- Annie Headache Protocol includes:
 - Headache diary:
 - Asks patient daily about presence of headache
 - If patient answers Yes to headache questions, 5 more questions are asked about pain, disability, symptoms and medications
 - 5 month duration
 - Educational tips:
 - Releases two educational tips per week for 6 months
 - Tips relate to non-pharmacological information on headache (acupressure, nutrition, sleep, health psychology, relaxation)
 - Some tips include links to learn more





ANNIE HEALTH SUBSCRIPTIONS ARE DIVIDED INTO SEVERAL CATEGORIES

Select the plus sign to reveal the available health subscriptions in a category. To subscribe to any of these, text **SUB** and the associated **KEYWORD** to Annie or 75338. For example, in the Diabetes category, to subscribe to the subscription named Diabetes Foot Care Reminder, text **SUB FOOT**

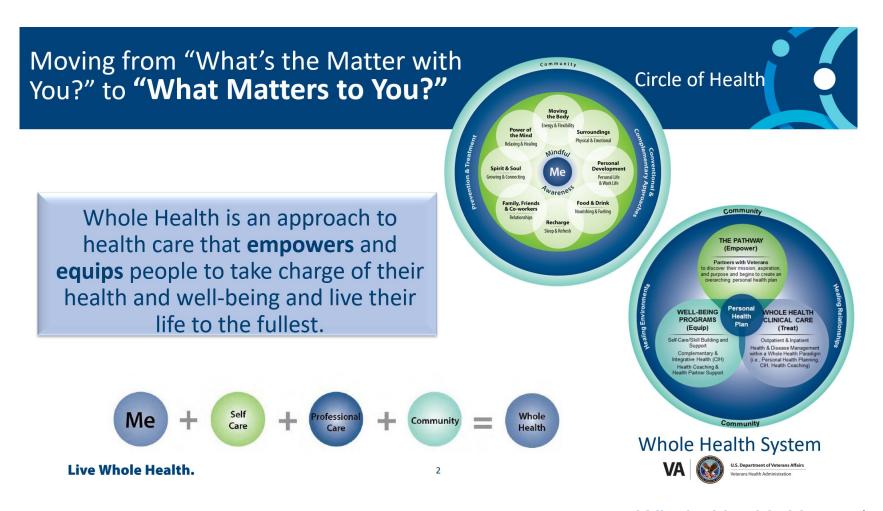
+ Cancer

+ COVID

+ Diabetes					
+ General Wellness	Subscription Name	Description	Duration	Approximate Frequency of Texts	Keyword to Subscribe
+ Hypertension	Pain Self-	This subscription provides Veterans with techniques	26 days	7/week	SUB PAIN
+ Hearing Difficulty	Management Support	to self-manage pain.			
+ Infectious Disease	Post Battlefield Acupuncture	This subscription provides Veterans with follow-up instructions after the placement of Battlefield	4 days	Up to 7/week	SUB BFA
+ Mental Health	Instructions	Acupuncture (BFA) needles, tacks, and seeds.			
- Pain	Headache Diary and Self-Management	This subscription provides Veterans with tips to improve headache control and an electronic diary to better understand how headaches impact their lives.	6 months	Up to 12/week	SUB HEADACHE



https://mobile.va.gov/app/annie-app-veterans#protocol





Whole Health Home (va.gov)



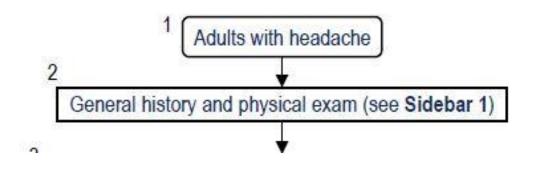
Headache Case Studies

DOD Patient

Case Example #1

DoD Patient

- 28 y/o active duty male,
 - MOS: 11B Infantry



- Presents with frequent headaches, increasing for the past 6 months, after returning from deployment
- Command directed evaluation patient does not like docs and meds



History and Physical

- Migraines as a teen
- Recurrent headaches after enlistment, 1-2 per month
- Post-deployment readjustment disorder (follows with mental health) following complicated deployment
- No history of TBI

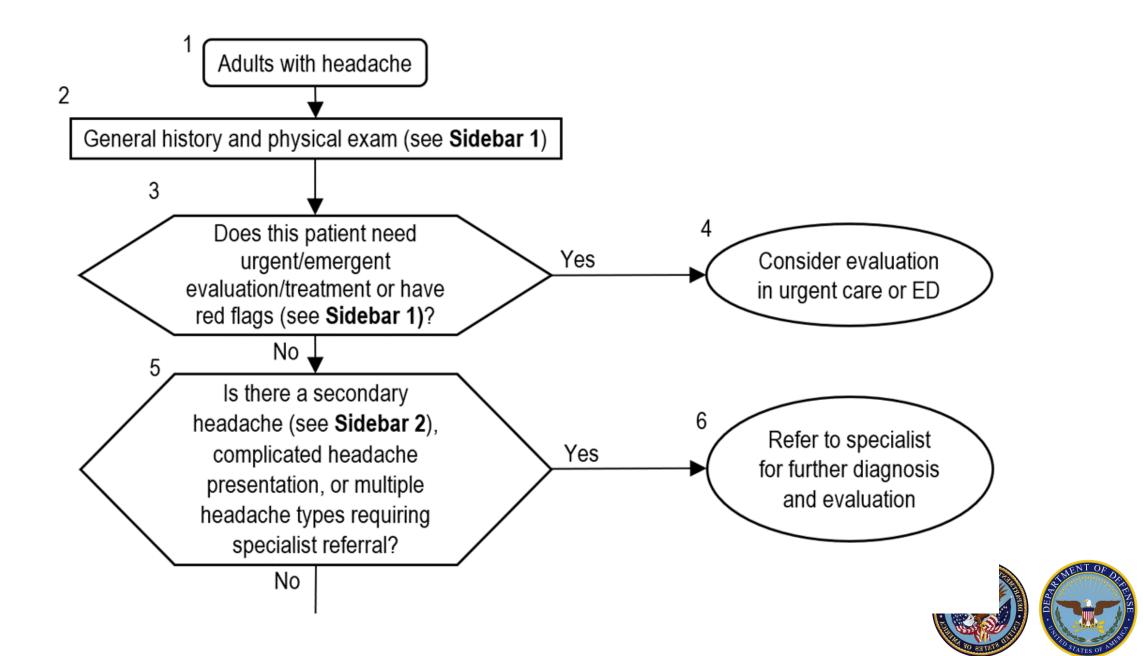


History and Physical

- Medications
 - OTC 1-2 x's week (Excedrin Migraine)
 - No prescription medications
- Physical limitations on physical profile due to daily physical training aggravating headaches
- Misses work once a month from a headache

- Frustrated as he uses exercise to help with his stress (work and relationships) now short tempered & gaining weight
- Vaping or tobacco dipping daily
- Social History
 - Married (separated) with 2 young children (ages 3 and 5)





Headache Description

Historical Feature	Description
Frequency	1-2 times/month
Duration	One hour if treated; 5 hours if untreated
Location	Right temporal
Quality	Throbbing
Intensity	6-8/10
Prodrome	No visual aura
Associated features	Nausea, vomiting, photophobia, phonophobia, osmophobia
Dysautonomia	Absent
Triggers	Decreased sleep, fluorescent lights from his office/computer
Worsened by	Exercise, household chores, driving home at the end of the day
Alleviated by	Sitting in a dark room; OTC medications "takes the edge off"
Debilitating	Yes



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Alleviated by	Sitting in a dark room; OTC medications "takes the edge off"
Debilitating	Yes



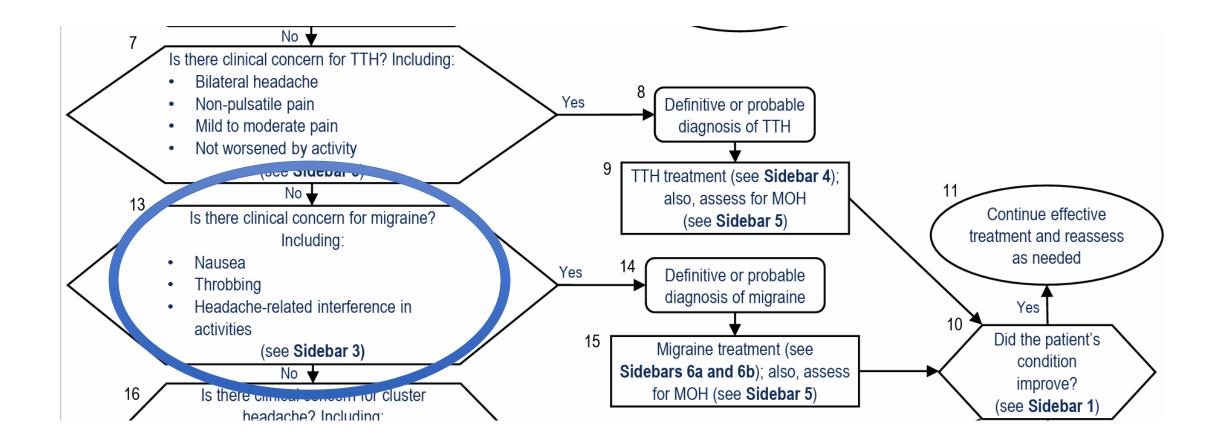


Sidebar 1: General History and Physical Exam								
 History Frequency, character Onset, prodrome/aura Location, duration Relieving or exacerbating factors 	 Red flags SNOOP(4)E (84) Systemic symptoms, illness, or condition (e.g., fever, chills, myalgias, night sweats, weight loss or gain, cancer, infection, giant cell arteritis, pregnancy or 	 sical Exam Examination Blood pressure General neurologic (upper extremities reflexes, sensation, strength, UMN, pathologic reflexes) Cranial nerves (including funduscopic exam) 						
 Associated symptoms Autonomic symptoms Jaw symptoms Neck symptoms Visual deficits/changes Dizziness and imbalance Current medications, abortive dose and frequency per month, prophylactic dose Prior medication trials Diet and nutrition, hydration Alcohol, caffeine intake Sleep Exercise 	 postpartum, or an immunocompromised state—including HIV) <u>Neurologic</u> symptoms or abnormal signs (e.g., confusion, impaired alertness or consciousness, changes in behavior or personality, diplopia, pulsatile tinnitus, focal neurologic symptoms or signs, meningismus, or seizures, ptosis, proptosis, pain with eye movements) <u>Onset</u> (e.g., abrupt or "thunderclap" where pain 	 Cervical spine and surrounding musculature (palpation, ROM, Spurling's sign test) Temporomandibular joint (palpation, ROM, symmetry, jaw claudication) Pericranial muscle palpation Temporal artery palpation; pertinent findings might include tenderness, cord-like artery, or lack of pulse Standardized headache assessments MIDAS (migraine-related disability) (85) HIT-6 (impact of headache on daily 						
 Aggravated by routine physical activity Sense of restlessness Foreign body sensation in the eye Nicotine and other stimulant use Risk factors for MOH History of trauma to the head, neck, or both Other comorbid conditions that might contribute to or exacerbate headaches Mental health (e.g., depression, anxiety, PTSD) Menstrual cycle and proximity to menopause 	reaches maximal intensity immediately or within minutes after onset; first ever, severe, or "worst headache of life") • <u>Older onset (age ≥50 years)</u> • <u>Progression or change in pattern (e.g., in headache frequency, severity, clinical features) • <u>Precipitated by Valsalva (e.g., coughing, bearing down)</u> • <u>Postural aggravation</u> • <u>Papilledema</u> • <u>Exertion</u></u>	 life and pain severity) (86) MSQL (quality of life) (87) ID Migraine (migraine) (88) Patient Headache Diary (7 day, 3 months)^a Additional screening tools PHQ-2 and PHQ-9 (depression) (89, 90) GAD-2 and GAD-7 (anxiety) (91, 92) CAGE (ethanol overuse headache) (93) AUDIT-C (ethanol overuse headache) (94) PC-PTSD (PTSD) (95) STOP-BANG (sleep) (96) 						



	Primary Headache Disorders Criteria*								
	Tension-type headache	Migraine headache	Cluster headache						
	Attack du	ration and frequency							
Duration	30-minutes – 7-days	4 – 72 hours	15 – 180 minutes						
Frequency	Variable	Variable	Once every other day to eight per day; often occurring at the same time of day						
	Headac	he characteristics							
Severity	Mild to moderate	Moderate to severe	Severe or very severe						
Location	Bilateral	Unilateral	Unilateral orbital, supraorbital, and/or temporal						
Quality	Pressing or tightening, non-pulsating	Throbbing or pulsating	Stabbing, boring						
Aggravated by routine physical activity	Not aggravated by routine activity	Aggravated by routine activity	Causes a sense of agitation or restlessness; routine activity may improve symptoms						
	Asso	ciated features							
Photophobia and phonophobia	Can have one but not both	Both	Variably present						
Nausea and/or vomiting	Neither	Either or both	May be present						
	0	ther features							
Autonomic features	None	May occur, but are often subtle and not noticed by the patient	Prominent autonomic features ipsilateral to the pain						







CPG Application: Assess for Risk Factors MOH

□Headache frequency >/= 7d/mo

✓ Migraine diagnosis

- Medication use: frequent use of anxiolytics, analgesics, or sedative hypnotics
- History of anxiety or depression with or without musculoskeletal complaints and/or gastrointestinal complaints

✓ Physical inactivity

✓ Sick leave of greater than two weeks in the last year

□Self-reported whiplash

✓ Smoking (tobacco use)



Sidebar 5: Medication Overuse Headache Criteria

ICHD-3 diagnostic criteria include:

- A. Headache occurring on 15 or more days per month in a patient with a preexisting headache disorder
- B. Regular overuse for more than 3 months of one or more drugs that can be taken for the acute or symptomatic treatment of headache (see table below)
- C. No better accounted for by another ICHD-3 diagnosis

Medication Overuse Headache Type	Medication Overuse Frequency	
Butalbital overuse ^a	≥5 days/month for >3 months	
Opioid overuse ^a	≥8 days/month for >3 months	
Triptan overuse		
Ergotamine overuse	≥10 days/month for >3 months	
Combination-analgesic overuse (any combination of classes, not to include combinations that include only non-opioid analgesics) ^{a, b}		
Non-opioid analgesic overuse (e.g., aspirin, NSAIDs, acetaminophen, steroids, and combinations of non-opioid analgesics)	≥15 days/month for >3 months	



Diagnosis and Concerns

Migraine without aura, episodic

At risk for Medication Overuse Headache



Veteran Patient in VA CARE

Case Example #2

Veteran is a 36-year-old right-handed gentleman with a history of episodic migraine.

He reports that his first, more severe type of headache is consistent with his episodic migraine attacks, though have become increasingly more severe, frequent, disabling, and less responsive to OTC medications.

His second type of headache began more than 6-months ago.



- Medications
 - OTC multiple times daily (Excedrin Migraine)
 - No prescription medications (previously on topiramate)
- Physical limitations has reduced physical activity because of headaches
- Misses work 2-3 times every month from a headache

Social History

- Vaping or tobacco dipping daily
- Divorced



- Mental Status: AAOx4, fluent speech
- Cranial Nerves: no ptosis; increased sensitivity over Right V1 without trigger points along V2
- Motor: 5/5 UE/LE; normal tone and bulk
- Sensory: intact UE/LE
- DTRs: 3+ UE/LE; toes down-going
- Cerebellar: intact
- Gait: normal

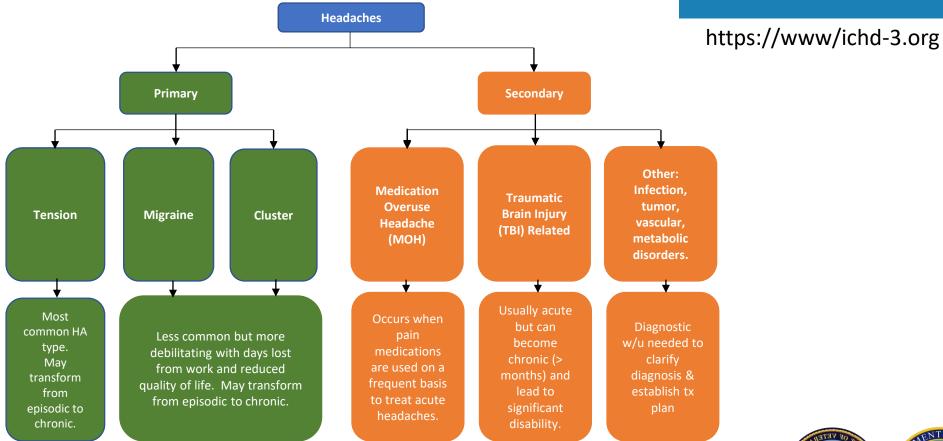


Historical feature	Headache 1	Headache 2
Frequency	4-5 times/week	Daily
Duration	Two hours if treated; 24 hours + if untreated	Continuous
Location	Right temporal	Holocranial
Quality	Throbbing	Aching
Intensity	8-9/10	2-3/10
Prodrome	No visual aura	No visual aura
Associated features	Nausea, vomiting, photophobia, phonophobia, osmophobia	None
Dysautonomia	Absent	Absent
Triggers	Decreased sleep, fluorescent lights	None
Worsened by	Exercise, household chores, driving	Going more than 12 hours without taking OTC medication
Alleviated by	Sitting in a dark room	None
Debilitating	Yes	No



Headache Classification









Migraine without aura, chronic

Medication Overuse Headache



http://www.healthquality.va.gov





Audience Q&A



