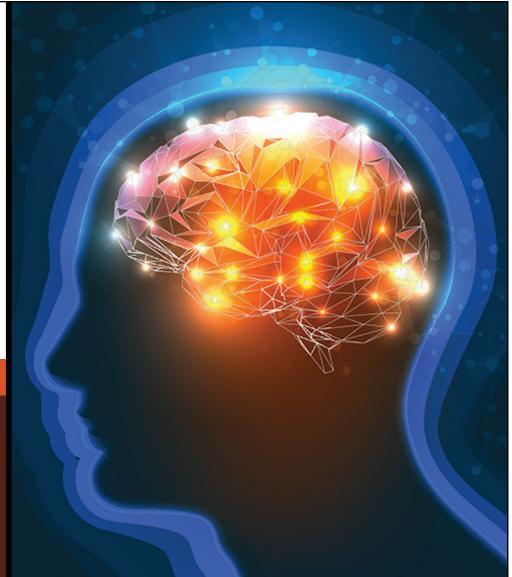
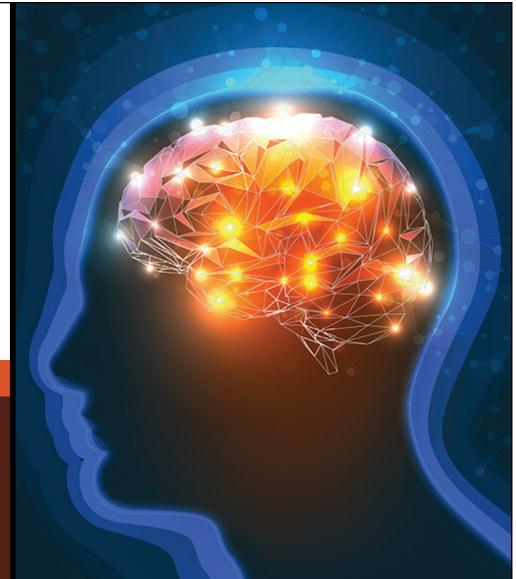


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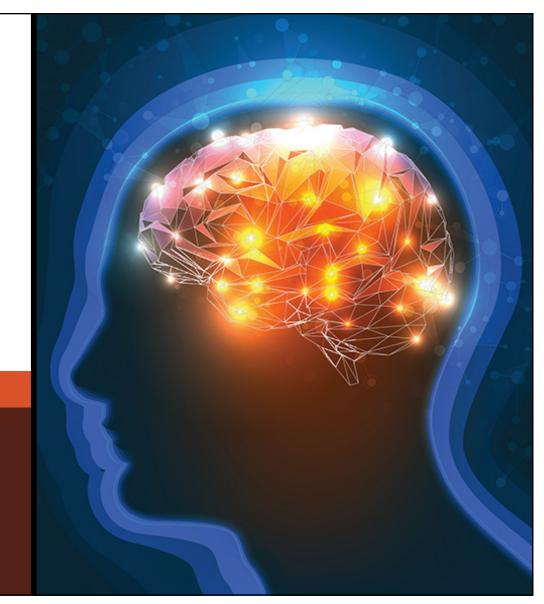
### Creating Individualized Treatment Plans for Patients with Migraine

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## Learning Objective

Identify factors that determine treatment decision-making for specific patient subgroups with migraine.



## **Case Presentation**

- I saw Donald in clinic initially on December 27, 2016. He is a 79-year-old male with a history of headaches for about 11 years. He tells me that the headaches first became problematic in February 2016, when they first became persistent. He had magnetic resonance imaging (MRI) back in 2014 that was done for the indication of intractable episodic headaches; however, the pain is now without relief. He was previously evaluated by another neurologist who recommended vitamin D and B12 supplementation as well as rheumatology evaluation for mildly elevated C-reactive protein (CRP) with normal sedimentation rate. He was seen by a rheumatologist, who thought this was not of rheumatological etiology.
- The patient's severe headaches are at a pain intensity of 8-9/10 twice a week, and on other days his severity is rated at a 5-7/10. He describes pain in the bilateral frontal regions that radiate backward and then also involves his neck and shoulders. The pain is continuous, throbbing, and tightening. In addition to sensitivity of light and sound, he has occasional vertigo, lightheadedness, and difficulty concentrating. He does not have any visual, sensory, motor, or language disturbances for auras.

## **Case Presentation**

### TRIGGERS/LIFESTYLE FACTORS/RISK FACTORS:

There are no provoking factors other than probably stress, flashing lights, sound, and weather changes. Alcohol does not provoke head pain. There have been a number of stressors, including taking care of an ill grandson for a number of weeks, which was emotionally exhausting. He also experienced home foreclosure in September and for a period of time was homeless, followed by the loss of his furniture, and an ill dog. He reports some depression related to head pain but denies a prior history of psychiatric illness. He is particularly bothered by not being able to enjoy activities due to head pain, including playing cards and a new-found desire for acting. Obstructive sleep apnea is not treated with continuous positive airway pressure (CPAP). He no longer thinks it helps headaches.

**PAST MEDICAL HISTORY:** Current medical problems include history of transient ischemic attack (described as wobbling for 24-48 hours), which resolved; asthma; hypertension; cervical spondylosis; gastroesophageal reflux disease; hiatal hernia; obstructive sleep apnea, which is associated with worsening headaches after a nap in the daytime, which is improved with daytime compliance to CPAP.

**PAST PSYCHIATRIC HISTORY:** He reports some depression related to head pain, but denies a prior history of psychiatric illness.

**PAST SURGICAL HISTORY:** Right knee surgery, right-sided ankle surgery, and cataract surgery

FAMILY HISTORY: His mother has a prior history of headache disorders.

**CURRENT MEDICATIONS:** Metoprolol, tizanidine, terazosin, potassium, multivitamins, omega-3, clopidogrel, atorvastatin

**ACUTE TREATMENT:** Excedrin was used daily and is now used in alternation with acetaminophen daily. Excedrin daily, headaches go away in 15 minutes. Acetaminophen 975 mg helps in 1-2 hours and is taken every day.

**PREVENTIVE TREATMENT:** Botox was tried once (17 injections?). Acupuncture and chiropractic manipulation were of no benefit.

**PAST TRIALS:** In the past, he has tried amitriptyline (which caused confusion) and topiramate (caused side effects); propranolol was not tried (history of bradycardia). Duloxetine 30 mg does not help. Sumatriptan and rizatriptan caused headaches. Acupuncture and chiropractic manipulation were of no benefit. Botox was tried once (17 injections?).

**ALLERGIES:** Include codeine. In the past, he has tried amitriptyline (which caused confusion) and topiramate (caused side effects); propranolol was not tried.

**SOCIAL HISTORY:** He has been retired since 2005. Married. Quit smoking 25 years ago. Previously worked as a chemist, testing pharmaceutical compounds. Drinks alcohol about once per week and coffee 3 times per day.

**REVIEW OF SYSTEMS:** Confusion; poor memory, forgetting appointments; change in sleep habits; excessive daytime drowsiness; loss of interest in activities; loss of consciousness; spinning sensation; muscle pain, mostly in the right hip, arms, and shoulder; balance difficulties; tremor; difficulty walking; lightheadedness; weight gain; shortness of breath; and joint pain. He reports right shoulder pain as well as right-sided hip pain, which have been recent and for several days. There is no pain with chewing.

#### **PHYSICAL EXAMINATION:**

Blood Pressure: 136/53 Pulse: 52 Respirations: 22 Temperature: 97.7 °F (36.5 °C) Body Mass Index: 33 General: Distress, normal tempo

General: Distress, normal temporal artery pulsations, without palpable cord or tenderness, normal range of neck motion

#### **NEUROLOGICAL EXAMINATION:**

Mental status: Alert, fluent, and follows commands. Good fund of knowledge, although admits to difficulty with naming and some forgetfulness. Cranial nerves: Disc clear. No cranial nerve deficits. Pupils equally round, reactive to light. Temporal arteries appear normal.

Motor: Full strength throughout.

Sensation: Intact to light touch. Reflexes: 1 to 2+. Toes downgoing. Coordination: Finger-tonose intact. Heel-to-shin intact.

Gait: Steady. Normal Romberg. Mildly impaired tandem.

#### **DIAGNOSTIC TESTING:**

The patient had MRI of the brain in August 2014 that showed stable white matter disease.

C-spine showed mild cervical multilevel spondylitic changes as well as in the T-spine.

Sedimentation rate 1 (normal), C-reactive protein 0.13 (mildly elevated).

## **Headache Assessment Scales**

#### MIDAS (Migraine Disability Assessment)

1-Miss work/school (days) 0
2-Productivity reduced >50% (days) 10
3-Housework not done (days) 0
4-Housework reduced >50% (days) 0
5-Miss activities family/social/leisure (days) 10

MIDAS Total (days) 20 MIDAS Grade III = Moderate

#### HIT-6 (Headache Impact Test) Score: 71

| 1-Severe Pain              | 11 |
|----------------------------|----|
| 2-Limit daily activities   | 10 |
| 3-Wish to lie down         | 13 |
| 4-Too tired for activities | 13 |
| 5-Fed up or irritated      | 13 |
| 6-Limit concentration      | 11 |
| HIT-6 Total                | 71 |

# **Clinical Considerations**

- 1. What is Donald's diagnosis?
- 2. What is the next diagnostic step?
- 3. What is the next best step to manage his headaches?
- 4. What important factors should be considered?
- 5. How may he benefit from an integrated treatment plan?

# **Risk Factors for Migraine Progression**

#### Demographics

- Female gender
- Low socioeconomic state
- Obesity

#### History of trauma

- Head and/or neck injury
- Traumatic life event

### Sleep disorders

- Insomnia
- Snoring

### Psychiatric comorbidities

- Anxiety
- Depression

### Pain disorders

- Arthritis
- Medication factors
  - Medication overuse of acute analgesics
  - Inadequate treatment/ineffective response
  - Caffeine overuse

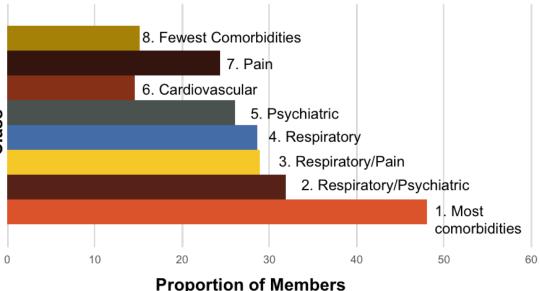
### Attack characteristics

- High frequency
- Allodynia
- Frequent nausea

Buse DC, et al. Headache. 2018. https://onlinelibrary.wiley.com/doi/epdf/10.1111/head.13459.

### Are There Clinical Subgroups of Migraine Based on Comorbidity and Condition Profiles?

- Real-world clinical trials are needed to fully assess
  - In the Chronic Migraine Epidemiology and Outcomes Study, a prospective web-based survey study of 12,810 respondents with migraine, patients were divided into:
    - Comparing Class 1 (Most Comorbidities) and Class 8 (Fewest Comorbidities), Class 1 had a greater proportion of individuals with severe disability (Migraine Disability Assessment grade IV; 48.1% vs. 22.3% of overall individuals) and higher rates of allodynia (67.6% vs. 47.0%), medication overuse (36.4% vs. 15.0%), chronic migraine (23.1% vs. 9.1%), and aura (40.1% vs. 28.8%).

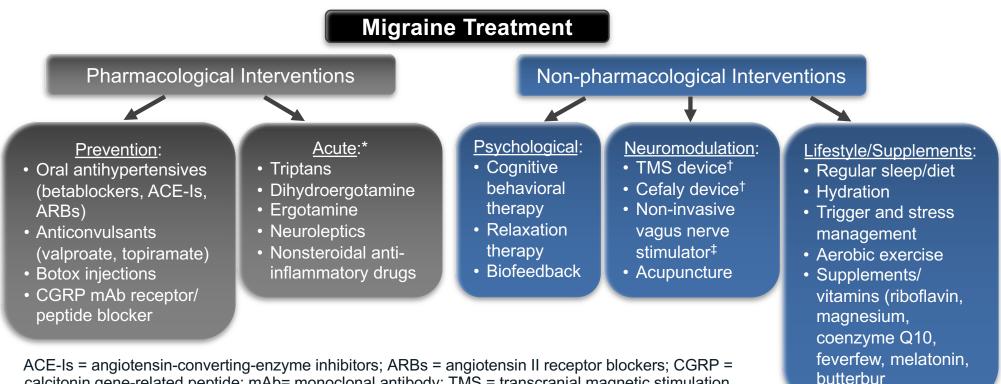


Lipton RB, et al. Headache. 2018;58(7):933-947.

### Shared Decision-Making: Values, Goals, Preferences



# **Migraine Treatment Options**



ACE-Is = angiotensin-converting-enzyme inhibitors; ARBs = angiotensin II receptor blockers; CGRP = calcitonin gene-related peptide; mAb= monoclonal antibody; TMS = transcranial magnetic stimulation. \*Limit medication overuse and tailor acute medications.

<sup>‡</sup>Approved by Food and Drug Administration (FDA) for acute and preventive indications.

<sup>‡</sup>GammaCore device cleared by FDA for acute attacks.

### **FDA-Approved Neuromodulation Devices for Migraine**

| Target  | Acute Pain Relief  | Preventive  | Device |  |
|---|--|---|--------|--|
| Transcutaneous<br>supraorbital<br>stimulation | Chou D, et al.<br>2018   | Schoenen J, et al.<br>2013                            | CEFAIN |  |
| Transcranial<br>magnetic stimulation<br>(TMS) | Lipton RB, et al. 2010   | Starling AJ, et al.<br>2018                           |        |  |
| Non-invasive vagus nerve stimulation          | Tassorelli C, et al.<br>2018<br>Approved by FDA for<br>cluster | Under investigation<br>Approved by FDA for<br>cluster |        |  |

Chou D, et al. *Cephalalgia*. 2018;333102418811573; Schoenen J, et al. *Neurology*. 2013;80;697-704. Lipton RB, et al. *Lancet Neurol*. 2010;9:373-380. Starling AJ, et al. *Cephalalgia*. 2018;38:1038-1048. Tassorelli C, et al. *Neurology*. 2018;91:e364-e373.

### Novel Migraine-Specific Preventive Treatments: mABs to CGRP or Its Receptor

|                                       | Erenumab             | Galcanezumab         | Fremanezumab              | Eptinezumab                                |
|---------------------------------------|----------------------|----------------------|---------------------------|--|
| Target                                | CGRP receptor        | CGRP ligand          | CGRP ligand               | CGRP ligand                                |
| Route of Administration               | SC                   | SC                   | SC                        | IV   |
| T 1/2                                 | Not disclosed        | 25-35 days           | Around 45 days            | Around 32 days                             |
| Frequency                             | Monthly              | Monthly              | Monthly<br>every 3 months | Every 3 months IV and<br>monthly SC        |
| Migraine<br>prophylaxis<br>indication | Episodic and chronic | Episodic and chronic | Episodic and chronic      | Episodic and chronic                       |
| Status                                | FDA approved         | FDA approved         | FDA approved              | Episodic and chronic<br>phase III complete |
| Safety                                | Safe and tolerated   | Safe and tolerated   | Safe and tolerated        | Safe and tolerated                         |

Drugs@FDA: FDA Approved Drug Products. https://www.accessdata.fda.gov/scripts/cder/daf. Saper J, et. Al. Neurology. 2018; 90 (15 Suppl) S20.001.

### SMART Goals Specific, Measurable, Attainable, Relevant, Timely

- Carefully listen to patient concerns, preferences, and goals
- Identify high-quality treatment based on side effect profile, comorbidities, and potential effectiveness



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