



# **Chronic Migraine and complications**



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### Case

- 23-year-old lady presented to ED for 7 days history of gradual onset of visual field defect in the right eye.
- She also experiences difficulties findings the right words and slurred speech.
- These symptoms did not resolve since 7 days before.









- Daily severe headache & Persistent nausea & vomiting for a week
- She described character as a throbbing pain over the left forehead and behind her left eye.
- Associated with phonophobia & photophobia
- Worse with movement
- Aura





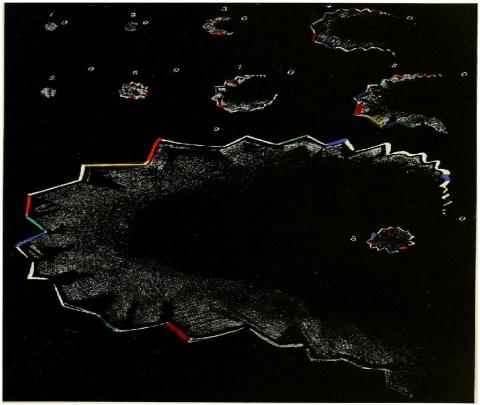
- Flashes of light and fortifications and zigzag lines
- aura symptom spreads gradually over ≥5 min
- symptoms occurred in succession
- aura symptoms lasted more than 60 min for this time



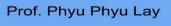












• She has similar headaches for **10 years**, intermittent, unilateral, throbbing , occurring 2-3 times a month.

Positive visual aura occurred < 60 mins</li>

 Last year, headaches were more frequent: > 15 headache days per month and Migrainous attacks were > 8 days per month for > 3 months. prescribed NSAIDs to be used as needed as abortive therapy.

• She was given long acting **propranolol** at 80 mg/day as a prophylactic medication

Usual treatment was not effective for this time





- Neurological examination revealed
- right visual field hemianopia,
- mild expressive aphasia and
- ☐ hypoesthesia of the right arm.

 Arterial blood pressure was 120/80 mmHg, heart rate was 84/min, SR and temperature was normal.



# **Differential diagnosis**

- Migraine with typical aura and it's complications
- Stroke
- ICH/SAH
- RCVCS/PRES
- Arterial dissection
- CVST
- AVM
- Meningitis
- leptomeningeal angiomatosis





- CT head with CTA was performed, but did not showed any signs of acute cerebral infarct, SAH, or other intracranial pathology.
- Routine blood test including CBC, ESR, CRP, D-dimer and standard biochemistry was unremarkable.
- Cerebrospinal fluid analysis was normal.
- She was diagnosed as complicated migraine.





She was treated with intravenous analgesics and supportive therapy.

• The symptoms slowly improved over the next few days and there was no residual neurology deficits 3 weeks later.

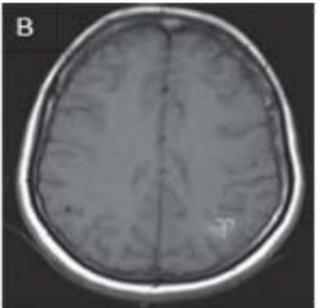
 MRI was performed on follow-up and showed cortical laminar necrosis in left occipital lobe indicative of migrainous infarction





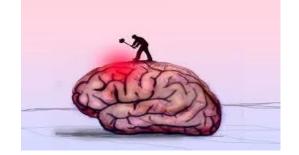
# MRI(Brain)











 Chronic migraine is a neurologic condition associated with individual, societal, and economic burden

It's estimated that up to 148 million people in the world live with chronic migraine.







# **Chronic Migraine**

- A. Migraine-like or tension-type-like headache on ≥15 days/month for >3 months that fulfill criteria B and C
- B. Occurring in a patient who has had at least five attacks fulfilling criteria B–D for migraine without aura and/or criteria B and C for migraine with aura
- C. On ≥8 days/month for >3 months, fulfilling any of the following:
  - Criteria C and D for migraine without aura
  - Criteria B and C for migraine with aura
  - Believed by the patient to be migraine at onset and relieved by
  - a triptan or ergot derivative
- D. Not better accounted for by another diagnosis



ICHD 3. Cephalalgia 2018; 38: 1–211. © 2018 International Headache Society



# Features of migraine and chronic migraine

#### MIGRAINE



#### Headache frequency o to 14 days per month

#### Diagnosis

At least 5 migraine attacks over the course of a lifetime



### CHRONIC MIGRAINE



### **Headache frequency**

15 or more days per month

#### Diagnosis

A history of 15 or more headache days a month, at least 8 with migraine characteristics, for at least 3 months

#### MIGRAINE SYMPTOMS AND CHARACTERISTICS

- Headache pain on one side
- Pulsating feeling
- Moderate to severe pain intensity
- Pain that gets worse with physical activity
- Nausea and/or vomiting
- Sensitivity to lights, sounds, or smells

SOURCE: International Classification of Headache Disorders, 3rd edition





# **Complications of Migraine**

Debilitating migraine attack that lasts more than 72

| 8 | 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5                                       |
|---|---|
|   | hours   |
|   | <ul> <li>Remissions of up to 12 hours related to medication or</li> </ul>   |
|   | sleep accepted  |
|   | <ul> <li>Status migrainosus can last several weeks</li> </ul>               |
|   | <ul> <li>Precipitating factors are stress/anxiety, menstruation,</li> </ul> |
|   | and lack of sleep   |

- Persistent aura without infarction Aura symptoms that last over 1 week without evidence of infarction on neuroimaging
- Migraine aura status
   Multiple auras (at least 2) occurring per day for more than 3 days in patients with migraine with aura



Status migrainosus



# **Complications of Migraine**

| Migrainous infarction           | • | with an ischemic cerebral lesion on neuroimaging Infarct develops in the course of the migraine attack Aura symptoms last over 60 min Lesions mostly located in the posterior circulation Younger women are often affected |
|---------------------------------|---|--|
| Migraine aura-triggered seizure | • | Occurs within 1 hour after or during a migraine  |

Agostoni E, Aliprandi A. The complications of migraine with aura. Neurol Sci. 2006 May;27 Suppl 2:S91-5. doi: 10.1007/s10072-006-0578-y. PMID: 16688637.

with aura attack



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# Other complications

- Work disability
- Loss of work

- Depression, anxiety and panic disorders
- Stroke, CVD, HT





### **Risk factors for CM**

### **Modifiable**

- Overuse of acute migraine medication,
- Ineffective acute migraine treatment,
- Obesity,
- Depression
- Stressful life events
- Caffeine intake
- Sleep disorders: insomnia, snoring, sleep apnea
- Chronic pain conditions: low back pain, neck pain, arthritis

## Non-Modifiable

- age
- female sex
- low educational status
- Head injury
- cutaneous allodynia,
- lower socioeconomic status





## **Co morbidities of Chronic Migraine**

- depressive disorders
- Anxiety
- post-traumatic stress disorder
- back pain, fibromyalgia & other musculoskeletal pain conditions)
- Hypertension & cardiovascular disease
- Allergies & asthma

- restless leg syndrome
- sleep disorders
- irritable bowel syndrome
- Epilepsy
- skin conditions
- Anemia
- Genetics- CADASIL





# **Triggers**

- Hormone changes in women- estrogen
- Alcohol, wine, caffeine
- Stress
- Sensory stimuli
- Sleep changes
- Weather changes
- Medications
- Foods & sweeteners
- Intense physical exertion





### **Treatment of CM**

The mainstay of chronic migraine treatment is prevention.

 Abortive Treatments - non-steroidal anti-inflammatory drugs, triptans, ergotamines

 Prophylaxis Treatments- Topiramate, Valproic/A, propranolol/ atenolol, telmisartan, flunarizine, amitriptyline, venlafaxine





• chemodenervation with onabotulinumtoxin A - FDA-approved preventive treatment for chronic migraine.

• CGRP receptor antagonists - erenumab, fremanezumab, galcanezumab, and eptinezumab.

rimegepant

lasmitidan



### **Treatment of CM**

- Non-medication preventive treatments
  - neuromodulation
  - cognitive behavioral therapy, biofeedback, accupuncture and relaxation treatments
  - The combination of new treatments including neuromodulation, behavioral approaches, and other interventions such as nerve blocks  $\rightarrow$  more effective
- co morbidities, risk factors such as depression, anxiety, snoring, obesity, etc. and cost should be considered.
- Getting enough sleep & Drinking plenty of water





# The diagnostic criteria of migrainous infarction

(International Classification of Headache Disorders, 3rd edition beta version) (ICHD-3β)

A. A migraine attack fulfilling criteria B and C

B. Occurring in a patient with Migraine with aura and typical of previous attacks except that one or more aura symptoms persists for > 60 minutes

- C. Neuroimaging demonstrates ischemic infarction in a relevant area
- D. Not better accounted for by another diagnosis





# Suggested mechanisms of ischemic stroke in migraineurs

Genetic predisposition MTHFR C677T polymorphism ACE-DD polymorphism rs7698623 in *MEPE* rs4975709 in *IRX4* 

Endothelial dysfunction Impaired endothelium-dependent vasodilation Endothelial cell damage

Coagulation abnormalities Increased PAF, vWF, prothrombin factor 1.2 (F1.2) Decreased resistance to activated protein C Protein S deficiency

Arterial dissection
Elevated serum elastase activity
MTHFR C677T polymorphism
PHACTR1 polymorphism (protective)

Patent foramen ovale (controversial)
Increased shunt amount in patients with migraine and stroke
Associated with juxtacortical white matter

Other comorbidities
Antiphospholipid antibody syndrome
Sneddon syndrome
Systemic lupus erythematosus

MTHFR, methylenetetrahydrofolate reductase; ACE-DD, angiotensin-converting enzyme deletion/deletion; PAF, platelet-activating factor; vWF, von Willebrand factor.

Journal of Stroke 2016;18(2):146-156 http://dx.doi.org/10.5853/jos.2015.01683





# **Take Home Messages**

• Chronic migraine is defined as having headache on at least 15 days per month, with 8 of these having migraine symptoms, for at least 3 months.

 Treatment strategies for chronic migraine include risk factor modification, identification of triggers, co morbidity management and the use of acute and prophylactic pharmacotherapy.

Chronic migraineurs invariably require prophylactic treatment.





## **Take Home Messages**



- Migraines generally don't get worse over time, but they can lead to more serious complications.
- These complications are quite rare but careful evaluation is needed to distinguish its from other serious debilitating diseases
- Atypical or prolonged aura must be specified and demands further analysis
- In persistent aura, neuro-imaging should always be in the workup











## References

- Elizabeth W. Loder, Rebecca C. Burch et al(2014). Common Pitfalls in the Evaluation and Management of Headache, Case based learning.
- Aksel Siva, Christian Lampl(2015). Case-Based Diagnosis and Management of Headache Disorders
- Marco A. Pescador Ruschel; Orlando De Jesus. Last Update: November 27, 2022.
   Migraine Headache StatPearls NCBI Bookshelf https://www.ncbi.nlm.nih.gov > books > NBK560787
- <u>Janie McQueen</u>. **Complications of Migraines**. <a href="https://www.webmd.com/migraines-headaches/headaches-migraine-complications">https://www.webmd.com/migraines-headaches-migraine-complications</a>
- <u>Dr. Emma Foster</u>. Migraine Complications. 17 May 2021. https://headacheaustralia.org.au/headachetypes/migraine-complications/
- Clinical Review; Management of chronic migraine. BMJ 2022; 379 doi: <a href="https://doi.org/10.1136/bmj-2021-067670">https://doi.org/10.1136/bmj-2021-067670</a> (Published 10 October 2022) Cite this as: BMJ 2022; 379:e067670









