

# **Core Curriculum on Headache for Neurologists**

Hayrunnisa Bolay, Nooshin Yamani, Sait Ashina, Fabíola Dach, Allan Purdy, Espen Kristoffersen, Michalis Vikelis, Michele Viana, Henrik Schytz, on behalf of the IHS Education Committee

The aim of the Curriculum is to define the minimum knowledge about headache required by neurologists for qualification as a specialist in neurology, and is based on information collected from different regions worldwide. This Curriculum allows for flexibility, and the exact content should be further defined by local headache specialists in the country in which it is used. In the Core Curriculum each section includes a list of articles relevant for further reading.

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### I. Anatomy and Pathophysiology

- Processing and modulation of nociceptive input from head and neck in the peripheral and central structures (trigeminovascular system, meninges, trigeminocervical junction, pons, periaqueductal grey matter, parabrachial nucleus, amygdala, hypothalamus, first order and higher order thalamic nuclei and sensorimotor cortical areas, cingulate and insular cortex, sympathetic and parasympathetic fibres).
  - Examination of the peripheral nerves, muscles, ligaments and other soft tissue structures of the head and neck, including tenderness of the greater occipital nerve, lesser occipital nerve, supraorbital nerve, supratrochlear nerve, temporal artery, tenderness of cervical structures and range of motion in the cervical area, and examination of the temporomandibular joints, to identify peripheral sources of headache pain.
- Disrupted sensory input processing during migraine attack, to understand the concepts of allodynia, photophobia, phonophobia, and osmophobia.
  - Examination of allodynia in the head and neck.
- The pathophysiology of a migraine attack, role of cortical spreading depression, peripheral and central mechanisms involved in the chronicity process.
- The molecular imprints of migraine, neuropeptides, genetic models and therapeutic targets.
- The pathophysiology of tension-type headache.
- The pathophysiology of trigeminal autonomic cephalalgias (TACs).
  - Examination of autonomic features, and location of the sphenopalatine ganglia and vagal nerve.
- The pathophysiology of neuropathic pain, trigeminal and other cranial neuralgias and persistent pain in the face and/or head.
  - Location of infraorbital nerve, mental nerve and auriculotemporal nerve etc, for peripheral interventions.

Akerman S, Goadsby PJ. A novel translational animal model of trigeminal autonomic cephalalgias. Headache 2015;55(1):197-203. https://doi.org/10.1111/head.12471

Ashina M, Hansen JM, Do TP et al. Migraine and the trigeminovascular system-40 years and counting. Lancet Neurol 2019;18(8):795-804. https://doi.org/10.1016/S1474-4422(19)30185-1

Bolay H, Messlinger K, Duox M et al. Anatomy of Headaches, Pathophysiology of Headaches. Eds. Ashina M, Geppetti P. Springer International Publishing, Switzerland. Page:1-31, 2015.

Bolay H, Vuralli D, Goadsby PJ. Aura and head pain: relationship and gaps in the translational models. J Headache Pain 2019;20(1):94. <a href="https://doi.org/10.1186/s10194-019-1042-8">https://doi.org/10.1186/s10194-019-1042-8</a>

Edvinsson L, Haanes KA, Warfvinge K et al. CGRP as the target of new migraine therapies - successful translation from bench to clinic. Nat Rev Neurol 2018;14(6):338-350. <a href="https://doi.org/10.1038/s41582-018-0003-1">https://doi.org/10.1038/s41582-018-0003-1</a>

Gambeta E, Chichorro JG, W Zamponi G. Trigeminal neuralgia: an overview from pathophysiology to pharmacological treatments. Mol Pain 2020;16:1744806920901890.

https://doi.org/10.1177/1744806920901890

Goadsby PJ, Holland PR. An update: pathophysiology of migraine. Neurol Clin 2019;37(4):651-671. https://doi.org/10.1016/j.ncl.2019.07.008

Pearl TA, Dumkrieger G, Chong CD et al. Sensory hypersensitivity symptoms in migraine with vs without aura: results from the American Registry for Migraine Research. Headache 2020;60(3):506-514. https://doi.org/10.1111/head.13745

# II. Epidemiology, Environment, Disability and Genetics

- Epidemiology, burden, costs and impact of migraine, tension-type headache, cluster headache, and other trigeminal autonomic cephalalgias, trigeminal neuralgia, persistent idiopathic facial pain, and the most important secondary headache disorders (e.g. medication overuse headache, post-traumatic headache).
- The incidence, prevalence and burden of headaches in children and adolescents, pregnancy and older subjects with or without dementia.
- Physical, psychological, social and environmental factors contributing to headache.
- Modifiable and non-modifiable risk factors for chronification of primary headaches
- Underlying genetic and insights into molecular and biological aspects of migraine and cluster headache and other special headache disorders (e.g. headache in mitochondrial disorders).

Ashina H, Porreca F, Anderson T et al. Post-traumatic headache: epidemiology and pathophysiological insights. Nat Rev Neurol 2019;15(10):607-617. https://doi.org/10.1038/s41582-019-0243-8

Buse DC, Greisman JD, Baigi K et al. Migraine progression: a systematic review. Headache 2019;59(3):306-338. https://doi.org/10.1111/head.13459

Gibson KF, Santos AD, Lund N et al. Genetics of cluster headache. Cephalalgia 2019;39(10):1298-1312. https://doi.org/10.1177/0333102418815503

Gormley P, Anttila V, Winsvold BS et al. Meta-analysis of 375,000 individuals identifies 38 susceptibility loci for migraine. Nat Genet 2016;48(8):856-866. <a href="https://doi.org/10.1038/ng.3598">https://doi.org/10.1038/ng.3598</a>

Lyngberg AC, Rasmussen BK, Jørgensen T et al. Has the prevalence of migraine and tension-type headache changed over a 12-year period? A Danish population survey. Eur J Epidemiol 2005;20(3):243-249. https://doi.org/10.1007/s10654-004-6519-2

Lyngberg AC, Rasmussen BK, Jørgensen T et al. Incidence of primary headache: a Danish epidemiologic follow-up study. Am J Epidemiol 2005;161(11):1066-1073. https://doi.org/10.1093/aje/kwi139

Lyngberg AC, Rasmussen BK, Jørgensen T et al. Prognosis of migraine and tension-type headache: a population-based follow-up study. Neurology 2005;23;65(4):580-585. https://doi.org/10.1212/01.wnl.0000172918.74999.8a

Marmura MJ. Triggers, protectors, and predictors in episodic migraine. Curr Pain Headache Rep 2018;22(12):81. <a href="https://doi.org/10.1007/s11916-018-0734-0">https://doi.org/10.1007/s11916-018-0734-0</a>

Pellegrino AB, Davis-Martin RE, Houle TT et al. Perceived triggers of primary headache disorders: a meta-analysis. Cephalalgia 2018;38(6):1188-1198. https://doi.org/10.1177/0333102417727535

Philipp J, Zeiler M, Wöber C et al. Prevalence and burden of headache in children and adolescents in Austria—a nationwide study in a representative sample of pupils aged 10–18 years. J Headache Pain 2019;20(1):101. https://doi.org/10.1186/s10194-019-1050-8

Pohl H, Gantenbein AR, Sandor PS et al. Interictal burden of cluster headache: results of the EUROLIGHT cluster headache project, an internet-based, cross-sectional study of people with cluster headache. Headache 2020;60(2):360-369. https://doi.org/10.1111/head.13711

Stovner LJ, Nichols E, Steiner TJ et al. Global, regional, and national burden of migraine and tension-type headache, 1990–2016: a systematic analysis for the Global Burden of Disease Study 2016. Lancet Neurol 2018;17(11):954-976. https://doi.org/10.1016/S1474-4422(18)30322-3

Sutherland HG, Griffiths LR. Genetics of migraine: insights into the molecular basis of migraine disorders. Headache 2017;57(4):537-569. <a href="https://doi.org/10.1111/head.13053">https://doi.org/10.1111/head.13053</a>

# **III. Clinical Investigation for Headache**

- To understand the importance and significance of proper history taking.
- To understand the significance of neurological examination, including fundoscopy (specifically signs of papilledema) and pericranial muscle examination.
- How and when to order and how to interpret blood tests, lumbar puncture, neuroimaging, angiography and other laboratory testing.
- Understand the significance of blood tests, such as erythrocyte sedimentation rate and Creactive protein.
- Understand the significance of lumbar puncture in secondary headaches, such as headaches secondary to intracranial hypo/hypertension.
- Understand and interpreting the significance of neuroimaging findings in migraine, such as white matter lesions.
- Understand the significance of magnetic resonance imaging in secondary headaches, including magnetic resonance angiography, magnetic resonance venography and other techniques for elucidating causes, such as idiopathic intracranial hypertension, spontaneous intracranial hypotension, mitochondrial, vascular and neoplastic diseases.
- Understand the significance of cervical and temporomandibular imaging.
- Understand the significance of neuroimaging in cranial neuralgias.

Cain MR, Arkilo D, Linabery AM et al. Emergency department use of neuroimaging in children and adolescents presenting with headache. J Pediatr 2018;201:196-201.

https://doi.org/10.1016/j.jpeds.2018.05.023

Chaudhry P, Friedman DI. Neuroimaging in secondary headache disorders. Curr Pain Headache Rep 2015;19(7):30. https://doi.org/10.1007/s11916-015-0507-y

Dainese F, Avanzini G, La Neve A et al. Proposal guidelines for epilepsy and headache. J Headache Pain 2015;16(Suppl 1):A193. https://doi.org/10.1186/1129-2377-16-s1-a193

Detsky ME, McDonald DR, Baerlocher MO et al. Does this patient with headache have a migraine or need neuroimaging? JAMA 2006;296(10):1274-1283. https://doi.org/10.1001/jama.296.10.1274

Do T, Remmers A, Schytz HW et al. Red and orange flags for secondary headaches in clinical practice: SNNOOP10 list. Neurology 2019;92(3):134-144. https://doi.org/10.1212/WNL.0000000000006697

Evans RW, Burch RC, Frishberg BM et al. Neuroimaging for migraine: the American Headache Society systematic review and evidence-based guideline. Headache 2020;60(2):318-336.

https://doi.org/10.1111/head.13720

Jang YE, Cho EY, Choi HY et al. Diagnostic neuroimaging in headache patients: a systematic review and metaanalysis. Psychiatry Investig 2019;16(6):407-417. <a href="https://doi.org/10.30773/pi.2019.04.11">https://doi.org/10.30773/pi.2019.04.11</a>

Kraya T, Deschauer M, Joshi PR et al. Prevalence of headache in patients with mitochondrial disease: a cross-sectional study. Headache 2018;58(1):45-52. <a href="https://doi.org/10.1111/head.13219">https://doi.org/10.1111/head.13219</a>

Kuruvilla DE, Lipton RB. Appropriate use of neuroimaging in headache. Curr Pain Headache Rep 2015;19(6):17. https://doi.org/10.1007/s11916-015-0490-3

Ling ML, Yosar J, Lee BW et al. The diagnosis and management of temporal arteritis. Clin Exp Optom 2019. https://doi.org/10.1111/cxo.12975

Mollan SP, Davies B, Silver NC et al. Idiopathic intracranial hypertension: consensus guidelines on management J Neurol Neurosurg Psychiatry 2018;89(10):1088-1100. <a href="https://doi.org/10.1136/jnnp-2017-317440">https://doi.org/10.1136/jnnp-2017-317440</a>

Sandrini G, Friberg L, Coppola G et al; European Federation of Neurological Sciences. Neurophysiological tests and neuroimaging procedures in non-acute headache (2nd edition). Eur J Neurol 2011;18(3):373-381. https://doi.org/10.1111/j.1468-1331.2010.03212.x Spears RC. Low-pressure/spinal fluid leak headache. Curr Pain Headache Rep 2014;18(6):425.

https://doi.org/10.1007/s11916-014-0425-4

Young NP, Elrashidi MY, McKie PM et al. Neuroimaging utilization and findings in headache outpatients: significance of red and yellow flags. Cephalalgia 2018;38(12):1841-1848.

https://doi.org/10.1177%2F0333102418758282

# IV. Diagnosis and Classification of Headache Disorders According to the International Classification of Headache Disorders, 3rd Edition (ICHD-3)\*

General concepts, major groups and subgroups.

- Diagnostic entities and criteria.
  - Migraine subtypes including chronic migraine.
  - Tension-type headache.
  - Trigeminal Autonomic Cephalalgias.
  - Other common primary headaches.
  - Secondary headache disorders.
  - Medication-overuse headache.
  - Headache attributed to head and/or neck trauma; trigeminal neuralgia & other cranial neuralgias.
  - Headache attributed to temporomandibular joint (TMJ) disorders.
  - Trigeminal neuralgia & other cranial neuralgias.
- Differential diagnoses of the above diagnostic entities.
- Concepts of chronicity within the classification.
  - Chronic vs. episodic migraine and tension-type headache.
  - Chronic vs. episodic TACs.
  - Chronic vs. episodic secondary headaches.
- Diagnostic criteria for the most common paediatric headache diagnoses and differences in presentation between children, adolescents and adults.
- To be able to use the classification to diagnose a given case of a common primary headache, an unusual or a secondary headache; including a case with multiple headache disorders\*.
- The role of history, examination and appropriate investigations in the diagnosis of primary and secondary headache disorders.
- Warning signs (red flags) suggestive of secondary headache disorders for various clinical situations (e.g. thunderclap headache, headaches with focal neurological deficits) and how to evaluate and treat them.

- Clinical characteristics, diagnostic criteria and treatment of the following secondary headache syndromes:
  - Persistent post-traumatic headache.
  - Headache attributed to non-traumatic intracranial haemorrhage.
  - Headache attributed to venous sinus thrombosis (CVT), arterial and vertebral dissection and other cranio-cervical vascular disorders.
  - Headache attributed to giant cell arteritis (temporal arteritis).
  - Headache attributed to reversible cerebral vasoconstriction syndrome (RCVS)
  - Headache attributed to idiopathic intracranial hypertension (IIH).
  - Headache attributed to low cerebrospinal fluid (CSF) pressure.
  - Medication-overuse headache.
  - Headache attributed to bacterial meningitis and meningoencephalitis.
  - Headache attributed to viral meningitis or encephalitis
  - Headache attributed to intracranial neoplasia.
  - Sleep apnoea headache.
  - Headache attributed to acute angle-closure glaucoma.
- \* Although in clinical practice headache patients can be diagnosed by a skilled practitioner without the use of criteria, the members of this committee recommend using the common language of ICHD-3 criteria as it is important both in clinical practice and for research reasons. It is important to remember that a patient can have more than one headache disorder and thus separate diagnoses can be applied as per ICHD-3 criteria. On-line versions of the ICHD-3 are available (https://ichd-3.org) as is a Pocket Version for clinical usage. Visit the IHS website (http://www.ihs-headache.org).

Headache Classification Committee of the International Headache Society (IHS). The International Classification of Headache Disorders, 3rd edition. Cephalalgia 2018;38:1–211.

https://doi.org/10.1177/0333102417738202

### V. Therapy

- Non-pharmacological and behavioural therapy.
  - Trigger factor awareness and avoidance.
  - Headache diaries (calendars) and their importance as part of follow-up and identification of possible triggers of headaches in headache patients.
  - Behavioural medicine therapies (biofeedback training, neurofeedback, relaxation techniques, visual imagery, cognitive restructuring, psychotherapy, counselling, etc.).
  - Physical techniques, such as routine exercise, physical therapy etc., including evidence for efficacy.
  - Natural substances, such as vitamins, minerals, herbs, and supplements (vitamin B2, magnesium, feverfew, butterbur, coenzyme Q10, melatonin, etc.), including evidence for efficacy.
  - Complementary therapy of headache, including evidence for efficacy.
- Acute pharmacotherapy of migraine, chronic migraine, status migrainosus, tension-type headaches, trigeminal autonomic cephalalgias (TACs), and miscellaneous headaches.
  - Evidence-based guidelines for acute therapy.
  - Evidence base for use of all classes and individual drugs within a class.
  - Evidence base for use of neuromodulation for acute therapy.
  - Mode of action, properties, dosages, mode of administration, drug—drug
    interactions, adverse events, and contraindications of preventive drugs.
  - Therapeutic strategies such as step care and stratified care, and the importance of correct timing and dosing.
  - Rescue and backup treatments when the first-line therapy is ineffective.
  - Limits of acute headache treatment, the risk of developing medication overuse headache, including the relative risk of various categories as causative factors.
  - Therapeutic options in menstrual or menstrually-related migraine.
- Preventive pharmacotherapy of migraine, chronic migraine, tension-type headache, TACs and miscellaneous headaches.
  - Evidence-based guidelines for prevention by drugs, interventional treatments
     (onabotulinum toxin injections, nerve blocks, etc.) and neuromodulation

- Evidence base for use of all classes and individual drugs within a class
- Mode of action, properties, dosages, mode of administration, drug-drug interactions, adverse events and contraindications of preventive drugs.
- Preventive therapeutic strategies, such as the selection of agents in the context of comorbidities according to their effect and side-effect profile, and the necessity of concomitant withdrawal of acute care medications in the context of medication overuse headache.
- Withdrawal therapy in the presence of acute headache medication overuse
  - Outpatient and inpatient options.
- Therapeutic options during pregnancy and lactation.
- Therapeutic options in children, adolescents, and elderly patients with primary headache disorders.
- Intravenous protocols for refractory patients: inpatient treatment and management, criteria for hospital admission.
- Surgical and interventional therapies (indications and limitations).
  - Evidence base for use of onabotulinum toxin A treatment for chronic migraine and other minimally invasive procedures, such as occipital nerve block and sphenopalatine ganglion block for cluster headache and migraine.
  - Trigeminal neuralgia, chronic and intractable migraine, cluster headache, other TACs.

American Headache Society. The American Headache Society position statement on integrating new migraine treatments into clinical practice. Headache 2019;59(1):1-18. https://doi.org/10.1111/head.13456

Bendtsen L, Sacco S, Ashina M et al. Guideline on the use of onabotulinumtoxinA in chronic migraine: a consensus statement from the European Headache Federation. J Headache Pain 2018;19(1):91. https://doi.org/10.1186/s10194-018-0921-8

Berk T, Ashina S, Martin V et al. Diagnosis and treatment of primary headache disorders in older adults. J Am Geriatr Soc 2018;66(12):2408-2416. https://doi.org/10.1111/jgs.15586

Marmura MJ, Silberstein SD, Schwedt TJ. The acute treatment of migraine in adults: the American Headache Society evidence assessment of migraine pharmacotherapies. Headache 2015;55(1):3-20. https://doi.org/10.1111/head.12499

Oskoui M, Pringsheim T, Billinghurst L et al. Practice guideline update summary: pharmacologic treatment for pediatric migraine prevention: report of the Guideline Development, Dissemination, and Implementation Subcommittee of the American Academy of Neurology and the American Headache Society. Neurology 2019;93(11):500-509. https://doi.org/10.1212/WNL.00000000000008105

Sacco S, Bendtsen L, Ashina M et al. European Headache Federation guideline on the use of monoclonal antibodies acting on the calcitonin gene related peptide or its receptor for migraine prevention. J Headache Pain 2019;20(1):6. <a href="https://doi.org/10.1186/s10194-018-0955-y">https://doi.org/10.1186/s10194-018-0955-y</a>

Sacco S, Merki-Feld GS, Ægidius KL et al; European Headache Federation (EHF) and the European Society of Contraception and Reproductive Health (ESC). Hormonal contraceptives and risk of ischemic stroke in women with migraine: a consensus statement from the European Headache Federation (EHF) and the European Society of Contraception and Reproductive Health (ESC). J Headache Pain 2017;18(1):108. https://doi.org/10.1186/s10194-017-0815-1

Silberstein SD, Holland S, Freitag F et al; Quality Standards Subcommittee of the American Academy of Neurology and the American Headache Society. Evidence-based guideline update: pharmacologic treatment

for episodic migraine prevention in adults: report of the Quality Standards Subcommittee of the American Academy of Neurology and the American Headache Society. Neurology 2012;78(17):1337-1345. https://doi.org/10.1212/WNL.0b013e3182535d20

Wells RE, Beuthin J, Granetzke L. Complementary and integrative medicine for episodic migraine: an update of evidence from the last 3 years. Curr Pain Headache Rep 2019;23(2):10. <a href="https://doi.org/10.1007/s11916-019-0750-8">https://doi.org/10.1007/s11916-019-0750-8</a>

# VI. Comorbidities, Course and Prognosis of Headache Disorders

- Psychosocial factors in the context of migraine, tension-type headache, TACs, as well as
  other primary, and the secondary disorders. The impact of medication overuse on headache
  progression and outcomes.
- The natural course of primary and secondary headache disorders.
- Non-psychiatric comorbidities.
- Psychiatric comorbidities of migraine, tension-type headache, cluster and other primary and secondary headaches, including depression, anxiety, panic disorder, psychosis, and others.
- Depression and anxiety questionnaires.
- Most used headache-related disability questionnaires (e.g. MIDAS, HIT-6, HURT-index).
- Migraine as a risk factor for stroke.
- Estrogen and progesterone treatment in migraineurs with and without aura.

Bottiroli S, Galli F, Viana M et al. Traumatic experiences, stressful events, and alexithymia in chronic migraine with medication overuse. Front Psychol 2018;9:704. <a href="https://dx.doi.org/10.3389%2Ffpsyg.2018.00704">https://dx.doi.org/10.3389%2Ffpsyg.2018.00704</a>

Burch RC, Buse DC, Lipton RB. Migraine: epidemiology, burden, and comorbidity. Neurol Clin 2019;37(4):631-649. <a href="https://doi.org/10.1016/j.ncl.2019.06.001">https://doi.org/10.1016/j.ncl.2019.06.001</a>

Diener HC, Holle D, Solbach K et al. Medication-overuse headache: risk factors, pathophysiology and management. Nat Rev Neurol 2016;12(10):575-583. https://doi.org/10.1038/nrneurol.2016.124

Gryglas A, Smigiel R. Migraine and stroke: What's the link? What to do? Curr Neurol Neurosci Rep 2017;17:22. https://dx.doi.org/10.1007%2Fs11910-017-0729-y

Lampl C, Thomas H, Tassorelli C et al. Headache, depression and anxiety: associations in the Eurolight project. J Headache Pain 2016;17:59. https://doi.org/10.1186/s10194-016-0649-2

Lee MJ, Choi HA, Shin JH et al. Natural course of untreated cluster headache: a retrospective cohort study. Cephalalgia 2018;38:655-661. https://doi.org/10.1177/0333102417706350

Lipton RB, Fanning KM, Buse DC et al. Migraine progression in subgroups of migraine based on comorbidities: results of the CaMEO study. Neurology 2009;93(4): e2224-e2236. https://doi.org/10.1212/wnl.0000000000008589

Minen MT, Begasse De Dhaem O et al. Migraine and its psychiatric comorbidities. J Neurol Neurosurg Psychiatry 2016;87(7):741-749. https://doi.org/10.1136/jnnp-2015-312233

Nappi RE, Merki-Feld GS, Terreno E et al. Hormonal contraception in women with migraine: is progestogen-only contraception a better choice? J Headache Pain 2013;14:66. https://doi.org/10.1186/1129-2377-14-66

Raggi A, Giovannetti AM, Quintas R et al. A systematic review of the psychosocial difficulties relevant to patients with migraine. J Headache Pain 2012;13(8):595-606. <a href="https://dx.doi.org/10.1007%2Fs10194-012-0482-1">https://dx.doi.org/10.1007%2Fs10194-012-0482-1</a>

Sacco S, Merki-Feld GS, Ægidius KL et al; European Headache Federation (EHF), the European Society of Contraception and Reproductive Health (ESCRH). Effect of exogenous estrogens and progestogens on the course of migraine during reproductive age: a consensus statement by the European Headache Federation (EHF) and the European Society of Contraception and Reproductive Health (ESCRH). J Headache Pain 2018;19:76. https://doi.org/10.1186/s10194-018-0896-5

Viana M, Bottiroli S, Sances G et al. Factors associated to chronic migraine with medication overuse: a cross-sectional study. Cephalalgia 2018;38:2045-2057. <a href="https://doi.org/10.1177/0333102418761047">https://doi.org/10.1177/0333102418761047</a>

### VII. Miscellaneous

- To understand the basic methodological issues of clinical studies in headache. In particular:
  - Selection of patients.
  - Trial design, including, blinding, use of placebo and/or active comparator.
  - Evaluation of endpoints.
  - Report of adverse events.
  - Sample size.
- To understand and promote the global, regional, and local interests of people with headache disorders, as well as challenge their pervasive stigma.
- To understand the ethical issues arising in headache research, headache management and relationships with the pharmaceutical industry.

Abu-Arafeh I, Hershey AD, Diener HC et al. Guidelines of the International Headache Society for controlled trials of preventive treatment of migraine in children and adolescents, 1st edition. Cephalalgia 2019;39(7):803-816. https://dx.doi.org/10.1177/0333102419842188.

Diener HC, Tassorelli C, Dodick DW et al. Guidelines of the International Headache Society for controlled trials of acute treatment of migraine attacks in adults: fourth edition. Cephalalgia 2019;39(6):687–710. https://doi.org/10.1177/0333102419828967

Dodick D, Edvinsson L, Makino T et al. Vancouver Declaration on Global Headache Patient Advocacy 2018. Cephalalgia 2018;38(13):1899-1909. https://doi.org/10.1177/0333102418781644

Dodick DW, Ashina M, Sakai F et al. Vancouver Declaration II on Global Headache Patient Advocacy 2019. Cephalalgia 2020 https://doi.org/10.1177/0333102420921162

Ethical Issues in Headache Research and Management: Report and Recommendations of the Ethics Subcommittee of the International Headache Society. Cephalalgia 1998;18:505-529. https://doi.org/10.1111/j.1468-2982.1998.1807505.x

Ethical issues arising from commercial sponsorship and from relationships with the pharmaceutical industry—report and recommendations of the Ethics Subcommittee of the International Headache Society. Cephalalgia 2008;28(Suppl 3);1–25. https://doi.org/10.1111/j.1468-2982.2007.01402.x

Hougaard A, Tfelt-Hansen P. General lack of use of placebo in prophylactic, randomised, controlled trials in adult migraine. A systematic review. Cephalalgia 2016;36(10):960-969.

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Hougaard A, Tfelt-Hansen P. Are the current IHS guidelines for migraine drug trials being followed? J Headache Pain 2010;11(6):457-468. https://doi.org/10.1007/s10194-010-0257-5

Tassorelli C, Diener HC, Dodick DW et al. Guidelines of the International Headache Society for controlled trials of preventive treatment of chronic migraine in adults. Cephalalgia 2018;38(5):815-832. https://dx.doi.org/10.1177/0333102418758283

Tfelt-Hansen P, Bjarnason NH, Dahlöf C et al. Evaluation and registration of adverse events in clinical drug trials in migraine. Cephalalgia 2008;28(7):683-688. https://dx.doi.org/10.1111/j.1468-2982.2008.01600.x

Tfelt-Hansen P, Pascual J, Ramadan N et al. Guidelines for controlled trials of drugs in migraine: third edition. A guide for investigators. Cephalalgia 2012; 32 (1), 6-38. <a href="https://doi.org/10.1177/0333102411417901">https://doi.org/10.1177/0333102411417901</a>