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# RESIDENTS AND FELLOWS

# The unforgivable curse of Harry Potter's thunderclap headaches

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Several attempts to pigeonhole Harry Potter's headaches into a Muggles' Headache Classification System have been done, examining the characteristics of pain and the evolution of Harry's attacks through the entire saga. The diagnoses proposed so far include migraine,<sup>1</sup> supraorbital neuralgia,<sup>1,2</sup> chronic headache attributed to head trauma,<sup>1,3</sup> symptomatic trigeminal neuralgia,<sup>3</sup> and nummular headache.<sup>4</sup> A secondary life-threatening headache has been repeatedly excluded since all of Harry's attacks resolved without consequences. However, we invite the readers to put themselves in the shoes of a doctor who examines Harry's case for the first time.

An 11-year-old boy presents to the emergency room because of recurrent attacks of severe headache. He describes "a pain like he's never felt before" that "pierced his head," "so bad he fell to his knees" and each time "the pain had gone as quickly as it had come." The general examination is unremarkable, except for a "curiously shaped cut, like a bolt of lightning" over his forehead. Considering the abrupt onset of pain, its severity, and the rapid escalation to peak intensity, thunderclap headache (TH) should be the first suspect that comes to your mind.

TH is defined as an excruciating headache with an abrupt onset, reaching maximal intensity within 1 min.<sup>5</sup> In patients presenting with such a clinical scenario, a comprehensive and urgent evaluation is mandatory to rule out the underlying disorders associated with high mortality and morbidity, mostly subarachnoid hemorrhage (SAH). Several disorders are potentially responsible for TH, yet five of them represent the most common and threatening ones.<sup>6</sup> Table 1 summarizes the main clinical and investigative findings of those conditions. Medical history and neurological examination may reveal distinctive

findings, pointing toward a specific etiology. For instance, recurrent short-lasting THs are suggestive of reversible cerebral vasoconstriction syndrome (RCVS),<sup>7</sup> whereas orthostatic headache is the hallmark of spontaneous intracranial hypotension.<sup>8</sup> Nonetheless, clinical features alone are unable to discriminate with certainty among different causes and a rapid stepwise diagnostic approach is always recommended (Figure 1).<sup>6,9,10</sup>

Noncontrast brain CT should be obtained promptly as its sensitivity for SAH decreases progressively (95%-99% at Day 1; 90% at Day 2: 75% at Day 3: 50% at Day 5).<sup>11-14</sup> Whenever a brain CT is unrevealing, a lumbar puncture (LP) should be performed to fully rule out SAH. Although its sensitivity increases progressively over time, as opposed to brain CT, reaching maximal sensitivity from 12 h to 2 weeks since disease onset, it must not be delayed. Cerebrospinal fluid analysis should include opening pressure, blood cell count, and visual inspection for xanthochromia. Four tubes are recommended in order to distinguish SAH from traumatic LP; indeed, red blood cell count remains unchanged in all tubes in the former. Additionally, immediate centrifugation usually aids to this end, revealing a xanthochromic supernatant only in SAH. Whenever available, spectrophotometry should also be used to measure products of red cell lysis (bilirubin), because it increases sensitivity for SAH. Negative CT brain and LP, if performed within a few days, reasonably exclude SAH. However, patients should be further investigated with contrast-enhanced brain magnetic resonance imaging and/or noninvasive vascular imaging to rule out other potential threatening etiologies.<sup>13</sup> Additionally, whenever a high suspicion of RCVS remains despite negative diagnostic workup (e.g., recurrent, triggered

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Abbreviations: RCVS, reversible cerebral vasoconstriction syndrome; SAH, subarachnoid hemorrhage.

LP findings Brain MRI findings Angiography findings	Elevated Subarachnoid blood Ruptured aneurysm, blood cells, vasospasm xanthochromia	LP findings Brain MRI findings Angiography findings	Normal Normal, sulcal SAH, Multifocal, ischemic stroke multivessel vasoconstriction	Normal Normal, ischemic Dissected artery stroke		Elevated opening Venous Venous sinus pressure, hemorrhages, thrombosis elevated protein evidence of level intraluminal thrombus
Brain CT findings LP f	Subarachnoid Elev blood I	Brain CT findings LP f	Normal, sulcal Nor SAH, ischemic stroke	Normal, ischemic Nor stroke		Dense triangle Elev sign, cord I sign, venous 6 hemorrhages I
Associated clinical features	Impaired consciousness, neck stiffness, seizure, focal defects	Associated clinical features	Transient or persistent focal neurological defects, seizures	Focal neurological defects, Horner syndrome, muleatile tinnitus audible	pursance minutes, addition bruit, lower cranial neuropathies, amaurosis fugax	providence uningual additional pruit, lower cranial neuropathies, amaurosis fugax Papilledema, impaired consciousness, seizures, focal neurological defects
TH-specific features	May be preceded by physical exertion	TH-specific features	Recurrent short-lasting THs, triggered by physical exertion, Valsalva maneuvers, or hot/cold water exposure	Pain located in unilateral ear and face (carotid) or	occipitai/nucnai region (vertebral)	occipital/nucnal region (vertebral) May worsen in supine position in response to Valsalva maneuvers
TH frequency at presentation	50%	TH frequency at presentation	94% <sup>19</sup>	9.2% and 3.6%		5% <sup>20</sup>
Etiology	Subarachnoid hemorrhage (SAH)	Etiology	Reversible cerebral vasoconstriction syndrome	Carotid and vertebral artery dissection		Cerebral venous thrombosis

TABLE 1 Clinical and investigative findings of the most common causes of TH

b 20 D colloid cyst.

Abbreviations: CT, computer tomography; LP, lumbar puncture; MRI, magnetic resonance imaging; TH, thunderclap headache. *Source*: Modified from Schwedt (2015).<sup>6</sup>



**FIGURE 1** Diagnostic algorithm for patients presenting with thunderclap headache. Contrast-enhanced brain magnetic resonance imaging and vascular neuroimaging should be performed based on clinical suspicion and not routinely [Color figure can be viewed at wileyonlinelibrary.com]

TH attacks), vascular imaging tests should be repeated after a few weeks.<sup>7,15</sup> Indeed, vasoconstriction usually starts distally and only later involves medium-large cerebral vessels detectable on radiological investigations.<sup>15</sup>

Primary TH remains a diagnosis of exclusion, and some experts believe that such an entity does not exist and those patients reflect poor diagnostic investigations or unrevealed structural changes/ small blood seepage of cerebral unruptured aneurysms.<sup>5</sup>

Magnetic resonance vessel wall imaging is an emerging technique that provides details of the vessel walls, different from conventional neuroimaging for which the assessment is limited to the vessel's lumen.<sup>16</sup> Such properties make this technique promising for the evaluation of patients presenting with TH, showing distinctive features across RCVS, arterial dissections, and unstable intracranial aneurysms.<sup>16-18</sup>

# CONCLUSIONS

Secondary headaches may be challenging for physicians, especially trainees and residents, but should always be excluded prior to diagnosing a primary headache. TH represents the most worrisome clinical feature in headache medicine, unfolding threatening underlying etiologies of secondary headaches. Therefore, it should not be underestimated and all patients must be expeditiously and comprehensively investigated; famous magicians are no exceptions. Unfortunately, we doctors can not rely on magical powers to appropriately diagnose and manage our patients, nonetheless remembering *HARRY POTTER* headaches and their strict association with serious danger, namely Voldemort to Harry, may aid us in this particularly difficult diagnostic challenge, reminding us of the five most important underlying disorders and the diagnostic



FIGURE 2 Main underlying etiologies and diagnostic workup suggested by the HARRY POTTER acronym

workup to perform (Figure 2). The present work is not meant to be an exhaustive review, but an attempt to discuss with a humorous tone, yet a clear didactic purpose, the main causes of TH, and its initial diagnostic approach. Indeed, the range of differential diagnosis is wider than we described, and controversy still exists over its best management.

Regarding Harry Potter's recurrent headaches, considering his long follow-up, we may now reasonably exclude threatening underlying causes. Therefore, a primary headache diagnosis might be sought, as previously reviewed. However, it remains uncertain if Muggles' headache classification may be extendable to the wizard world, and a headache related to an *unforgivable curse* remains the most likely diagnosis.

#### CONFLICT OF INTEREST

The authors report no relevant conflict of interest.

#### AUTHOR CONTRIBUTIONS

Study concept and design: Umberto Pensato. Acquisition of data: Umberto Pensato, Eleonora Matteo. Analysis and interpretation of data: Umberto Pensato, Eleonora Matteo, Sabina Cevoli. Drafting of the manuscript: Umberto Pensato. Revising it for intellectual content: Umberto Pensato, Eleonora Matteo, Sabina Cevoli. Final approval of the completed manuscript: Umberto Pensato, Eleonora Matteo, Sabina Cevoli.

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