

Investigation of the characteristics of headache due to unruptured intracranial vertebral artery dissection

Cephalalgia
0(0) 1–11
© International Headache Society 2018
Reprints and permissions:
sagepub.co.uk/journalsPermissions.nav
DOI: 10.1177/0333102418791818
journals.sagepub.com/home/cep



Hiroaki Matsumoto¹, Hiroaki Hanayama¹, Yasuo Sakurai¹,
Hiroaki Minami¹, Atsushi Masuda¹, Shogo Tominaga¹,
Katsuya Miyaji¹, Ikuya Yamaura¹, Yasuhisa Yoshida¹ and
Yutaka Hirata²

Abstract

Background and purpose: It is sometimes difficult to diagnose intracranial vertebral artery dissection in patients with headache as the only symptom. Knowledge of the characteristics of the headache would facilitate the diagnosis. In this study, we aimed to clarify the characteristics of intracranial vertebral artery dissection-related headache using our original self-administered questionnaire.

Methods: Via the questionnaire, we ascertained headache characteristics and investigated whether they differed between two types of unruptured intracranial vertebral artery dissection, headache type and ischemic type, based on analysis of the responses. Then, we tried to validate the consistency of commonly used criteria for intracranial artery dissection by comparing them with our results.

Results: Thirty-seven patients were analyzed. Our results identified the following seven headache characteristics in patients with intracranial vertebral artery dissection: (i) occurring in the occipitounuchal region (89%); (ii) unilateral (81%); (iii) pulsatile (70%); (iv) of acute onset (70%); (v) severe (73%); (vi) without nausea or vomiting (73%); and (vii) with concomitant clinical symptoms unrelated to ischemia (81%). Comparison of headache characteristics between the two types of intracranial vertebral artery dissection headache showed that the pain was significantly more severe in headache type than ischemic type intracranial vertebral artery dissection ($p = 0.01$). Concomitant clinical symptoms occurred significantly more often in ischemic type than headache type intracranial vertebral artery dissection ($p = 0.03$). Our results generally satisfied the established headache diagnostic criteria.

Conclusion: The pain characteristics of headache type and ischemic type intracranial vertebral artery dissection shown in our study may facilitate its diagnosis.

Keywords

arterial dissection, diagnosis, pain

Date received: 9 February 2018; revised: 26 May 2018; 12 June 2018; accepted: 21 June 2018

Introduction

Spontaneous cervicocephalic arterial dissection is well known as an important cause of stroke in young adults (1–5). There are known ethnic differences in the affected arteries: internal carotid artery dissection (ICAD) is common in Caucasian populations, whereas vertebral artery dissection (VAD) is more common in Asian populations (2,6–8). Moreover, the frequency of intracranial VAD (IC-VAD) has been reported to be up to three- to 10-fold greater than that of extracranial VAD in Asian populations (2,6–8). Hence, the majority of cervicocephalic arterial dissections

in Asia are IC-VAD. It is well known that headache is the most common initial symptom in patients with IC-VAD (6,9–13). IC-VAD is divided into three types

¹Department of Neurosurgery, Cerebrovascular Research Institute, Eishokai Yoshida Hospital, Kobe, Japan

²Department of Neurology, Cerebrovascular Research Institute, Eishokai Yoshida Hospital, Kobe, Japan

Corresponding author:

Hiroaki Matsumoto, Daikai-dori9-2-6, Hyogo-ku, Kobe 652-0803, Japan.
Email: hiroaki-matsu@umin.ac.jp

based on the clinical symptoms: hemorrhagic type, ischemic type and headache type (14). In hemorrhagic type, or ruptured IC-VAD, the headache characteristics conform to those of subarachnoid hemorrhage. On the other hand, the headache characteristics of unruptured IC-VAD (i.e. ischemic and headache types) are not well known, and it is difficult to diagnose IC-VAD in patients presenting with only headache to the outpatient clinic or emergency department (6,9,10,15). The fact that it takes a long time to diagnose this condition also reflects the difficulty in its diagnosis (6,16). Understanding the headache characteristics would, thus, facilitate the diagnosis. However, only a few reports have focused on the characteristics of unruptured IC-VAD-related headache. Moreover, questions remain regarding whether the headache characteristics are the same or different between ischemic and headache type IC-VAD because no previous report has investigated the differences in headache characteristics between these two types of IC-VAD.

When patients presenting with headache are evaluated, physicians usually use *The International Classification of Headache Disorders, 3rd edition (beta version)* (ICHD-3 beta) to diagnose the cause (17). However, since the ICHD-3 beta clubs together the criteria for diagnosis of all types of intracranial artery dissection, there are no specific criteria for IC-VAD. The *Clinical Practice Guideline for Chronic Headache 2013*, which was published by the Japanese Headache Society, also lists together the headache characteristics of all types of intracranial artery dissection (18). However, since intracranial ICAD and IC-VAD differ in their pathophysiology (16), their headache characteristics are also likely to be different. Thus, the reliability of these guidelines may be low.

In this study, we aimed to clarify the headache characteristics of unruptured IC-VAD by using our original self-administered questionnaire. We also investigated whether headache characteristics differ between the two types of unruptured IC-VAD. Finally, we tried to validate the consistency of the criteria for intracranial artery dissection in the ICHD-3 beta and Clinical Practice Guideline for Chronic Headache 2013 by comparing them with our results.

Methods

This study was approved by the institutional review board of Eishokai Yoshida Hospital. The prospectively maintained database of our hospital was searched for patients with unruptured intracranial VAD between January 2011 and June 2017. Their medical records and radiographic studies were reviewed retrospectively.

Definition of unruptured intracranial VAD

We diagnosed IC-VAD by magnetic resonance imaging (MRI), such as 3-dimensional time-of-flight MR angiography and basi-parallel anatomical scanning. Unruptured IC-VAD cases were divided into two types: 'headache type' and 'ischemic type'. Headache type was defined as cases presenting with only headache during the entire clinical course, while ischemic type was defined as cases presenting with ischemic symptoms with or without cerebral infarction on MRI during the clinical course. Hence, patients presenting with transient ischemic attack were included in the ischemic group.

Study design

We used our original self-administered questionnaire that consisted of 15 questions (Table 1) in this study. We sent the questionnaire to known cases of unruptured IC-VAD by mail and analyzed the completed questionnaire sent back to us. After investigating the characteristics of the headache due to unruptured IC-VAD, we compared the headache characteristics between headache type and ischemic type IC-VAD. Then, we tried to validate the consistency of the criteria for intracranial artery dissection in the ICHD-3 beta and Clinical Practice Guideline for Chronic Headache 2013 by comparing these criteria with our results. The returned questionnaires were accompanied by the patients' informed consent for study participation.

Definition of headache intensity

We investigated headache intensity in Q5. Headache intensity, which was measured using a Numerical Rating Scale (NRS), was classified as mild (NRS = 1–3), moderate (NRS = 4–6), and severe (NRS = 7–10).

Definition of concomitant clinical symptoms with the headache

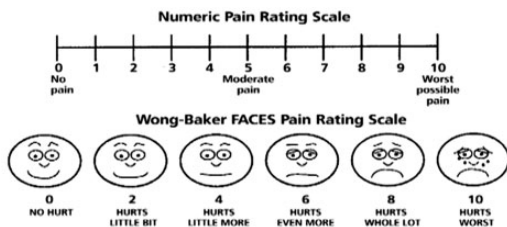
We investigated the presence of concomitant clinical symptoms besides headache in Q8. Clinical symptoms due to ischemia, such as visual field defects, dysarthria, double vision, ataxia, vertigo, and hemiparesis, which could have been indicative of ischemia, were excluded. We selected items such as irritated eyes, giddiness, transient dysesthesia, listlessness, fever, and neck stiffness. We selected these items based on a medical interview with the patients at the time of the initial visit.

Patient population

We treated 63 patients for unruptured IC-VAD at our institution between January 2011 and June 2017.

Table 1. Our original self-administered questionnaire.

- Q1. Where did your headache occur? (multiple answers allowed)
- A. Forehead
 - B. Temporal region
 - C. Parietal region
 - D. Occipitounuchal region
 - E. Face
 - F. None
 - G. Unknown
- Q2. Was your headache unilateral or bilateral?
If unilateral, on which side did your headache occur?
- A. Unilateral: left or right
 - B. Bilateral
 - C. Unknown
- Q3. What was the nature of your headache?
- A. Pulsatile
 - B. Constrictive
 - C. Stabbing
 - D. Dull
 - E. Burning
 - F. Electrical
 - G. Unknown
- Q4. What was the mode of onset of your headache?
- A. Acute onset (developing within 24 hours)
 - B. Gradual onset (developing over more than 24 hours)
 - C. Unknown
- Q5. Was this headache the worst pain you have ever felt?
- A. Yes
 - B. No
 - C. Unknown
- Q6. What was the intensity of your headache? Please show score.



- Q7. Were there any aggravating factors for the pain? (multiple answers allowed)
- A. Body motion
 - B. Drinking alcohol
 - C. Bathing
 - D. Others
 - E. None
 - F. Unknown
- Q8. Did you have any other clinical symptoms apart from headache? (multiple answers allowed)
- A. Irritated eyes
 - B. Giddiness
 - C. Transient dysesthesia
 - D. Listlessness
 - E. Fever
 - F. Stiff shoulder
 - G. Others
 - H. None
 - I. Unknown

Table 1. Continued.

- Q9. How long did your headache continue?
- A. A few days
 - B. One week
 - C. A few weeks
 - D. One month
 - E. Over one month
 - F. Unknown
- Q10. Did you have vomiting or nausea?
- A. Yes
 - B. No
 - C. Unknown
- Q11. What were you doing when your headache occurred?
- A. Sleeping
 - B. Drinking alcohol
 - C. Exercise
 - D. Working
 - E. Other daily life activities
 - F. Unknown
- Q12. Did you take a painkiller?
- A. Yes
 - B. No
 - C. Unknown
- Q13. If you answered Yes to Q12, did it relieve the headache?
- A. Yes
 - B. No
- Q14. Do you have a history of headaches, such as due to migraine?
- A. Yes
 - B. No
- Q15. If you answered Yes to Q14, did this headache differ from previous headaches?
- A. Yes
 - B. No

All cases had unilateral lesions. We sent the questionnaire by mail to these 63 patients. Thirty-nine patients (61%) returned the completed questionnaire. Among these 39 patients, two patients with ischemic type IC-VAD responded that they had no headache and so were excluded. The interval between diagnosis and the current investigation ranged from 1 to 78 months (mean 38.6 months, median 43 months). Finally, 37 patients (59%) were analyzed (Figure 1).

Scoring system for diagnosis of the headache of unruptured IC-VAD

We attempted to establish a scoring system for diagnosis of the headache. We considered items with either a positive or negative response rate of over 65% as the headache characteristics. We scored each characteristic as 1 point and investigated the total score in each patient.

(continued)

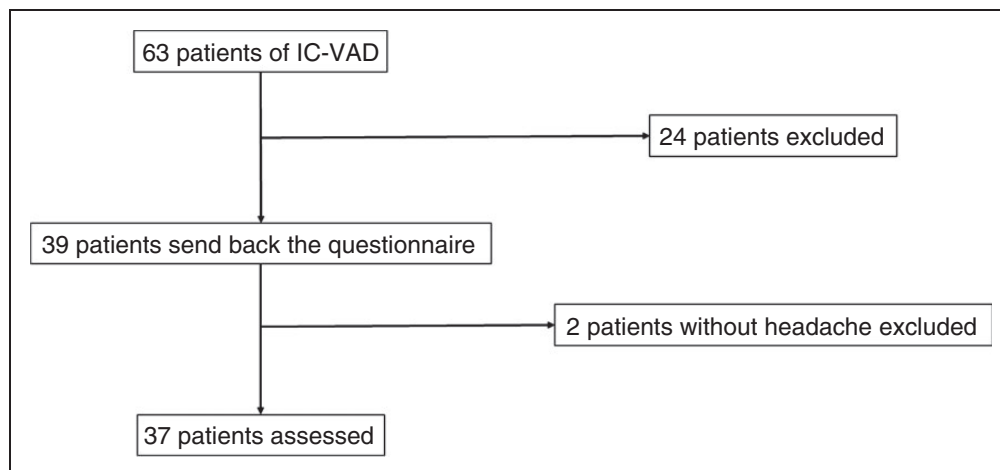


Figure 1. Flow chart showing patient selection.

Statistical analysis

To compare the headache characteristics between headache type and ischemic type IC-VAD, statistical analysis was performed using the Student's *t*-test, Mann-Whitney *U*-test, or chi-squared test of independence. Differences were considered significant for probability values of $p < 0.05$. All statistical analyses were performed using Statcel software, version 4 (OMS Publishing, Saitama, Japan).

Results

Patients ranged in age from 37 to 71 years (mean 48.4 years) and were predominantly male (23 men, 14 women). Twenty-four patients had headache type and 13 patients had ischemic type IC-VAD. Table 2 gives a summary of the clinical characteristics of patients with ischemic type IC-VAD.

General characteristics of headache due to unruptured IC-VAD

Headache occurred mainly in the occipitotemporal region (33/37, 89%), and spread to the temporal and parietal regions, forehead, and face (Figure 2A). However, some patients (4/37, 11%) did not have posterior headache, and instead had pain in the temporal region ($n = 3$) or the face ($n = 1$). Headache was mainly unilateral (30/37, 81%; Figure 2B). The rate of headache ipsilateral to the IC-VAD was 65% (24/37). The pain was most frequently described as pulsatile (26/37, 70%), followed by dull (4/37, 11%), constrictive (3/37, 8%), burning (2/37, 5%), and electrical (2/37, 5%) (Figure 2C). Most of the patients (26/37, 70%) had acute onset headache (Figure 3A). Twenty-four of 37 patients (65%) answered that it was the worst pain they

Table 2. Summary of the clinical characteristics of patients with ischemic type intracranial vertebral artery dissection.

	<i>n</i>
1. Stroke type	
Major stroke (NIHSS > 4)	2
Minor stroke (NIHSS < 3)	10
TIA	1
2. Site of infarction	
Lateral medulla	7
Cerebellum	4
Lateral medulla + cerebellum + occipital lobe	1
Undetermined	1
3. Ischemic symptoms	
Wallenberg syndrome	3
Ataxic gait	4
Sensory disturbance	3
Cranial nerve palsy	1
Vertigo	4
Nystagmus	2
Transient hemiparesis	1

NIHSS: National Institute of Health Stroke Scale; TIA: transient ischemic attack.

had ever felt and 27/37 (73%) rated the pain intensity as severe (Figure 3B and 3C). Fifteen of 37 patients (41%) had aggravating factors for the pain, body motion being a major factor (Figure 4A). Thirty of 37 patients (81%) had concomitant clinical symptoms when the headache occurred, among which neck stiffness was most common (Figure 4B). In almost all patients (35/36, 97%), headache resolved within 1 month of its onset. Duration of headache in the remaining one patient was unknown (Figure 5A). Most of the patients did not have accompanying nausea or vomiting (27/37, 73%, Figure 5B). Although headache usually occurred

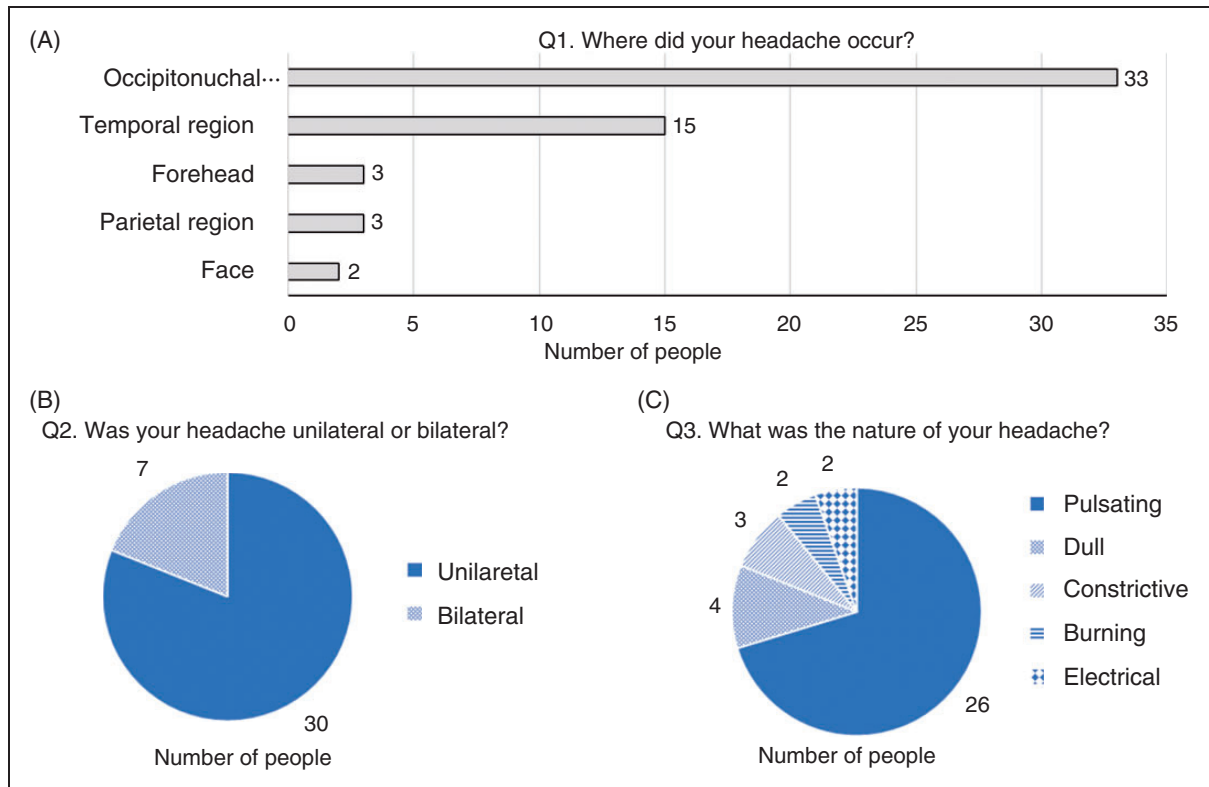


Figure 2. Results of the questionnaire: (A) Q1; (B) Q2; (C) Q3.

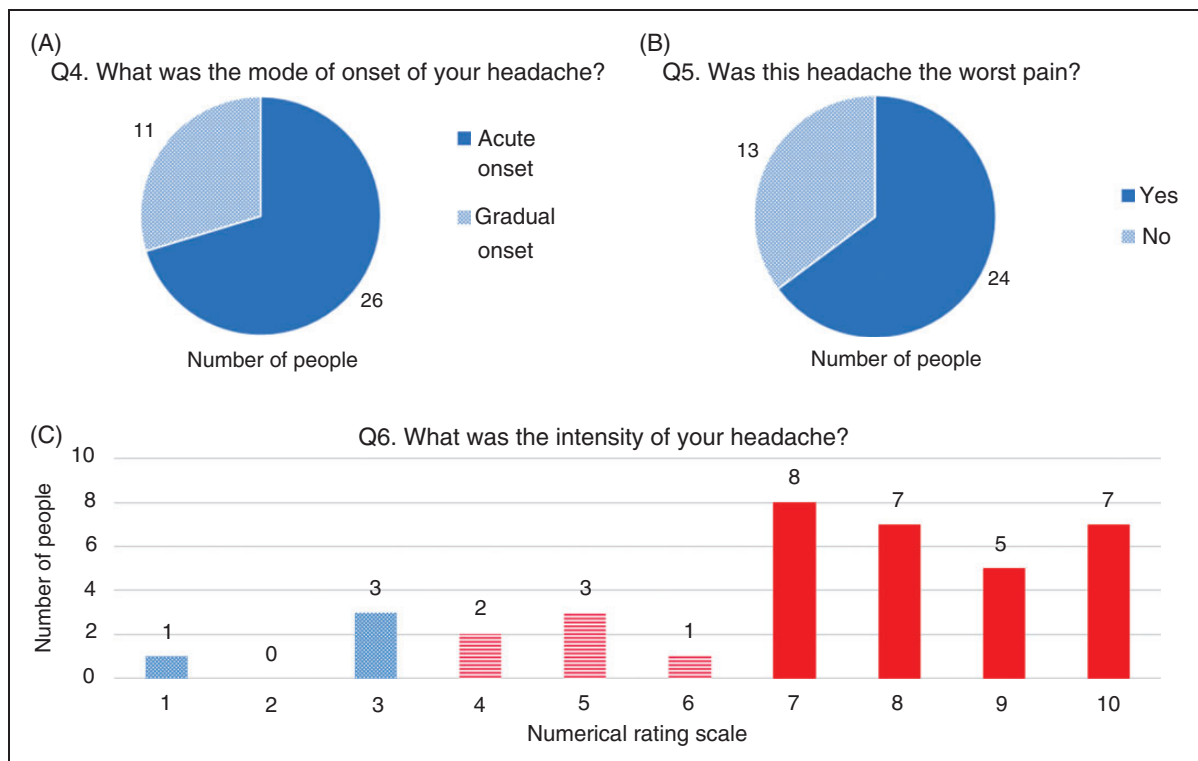


Figure 3. Results of the questionnaire: (A) Q4; (B) Q5; (C) Q6.

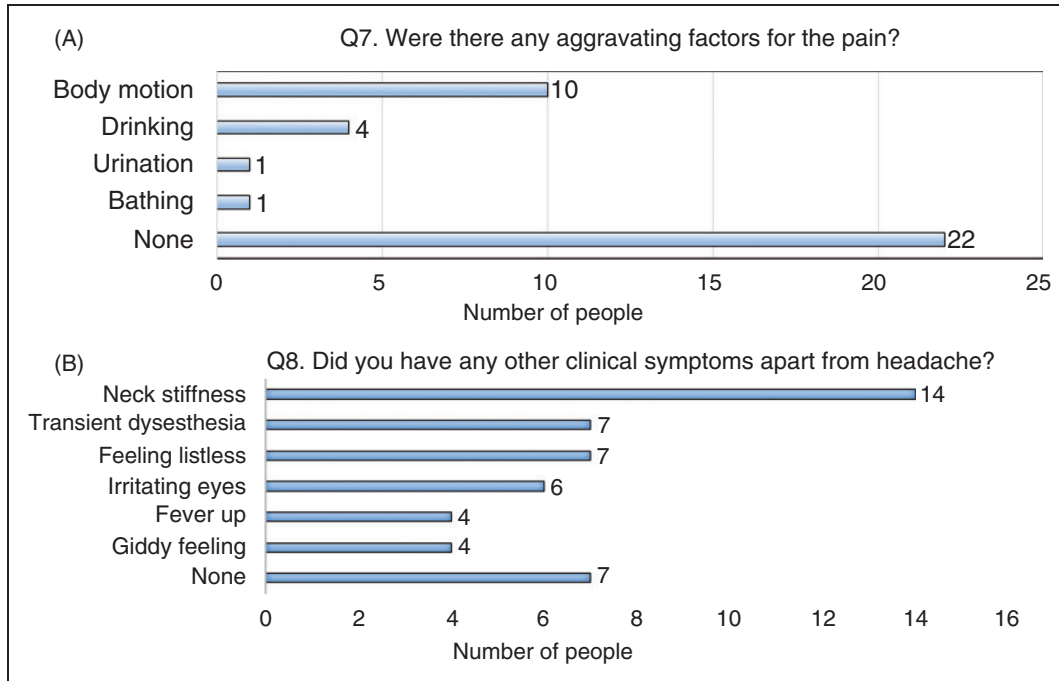


Figure 4. Results of the questionnaire: (A) Q7; (B) Q8.

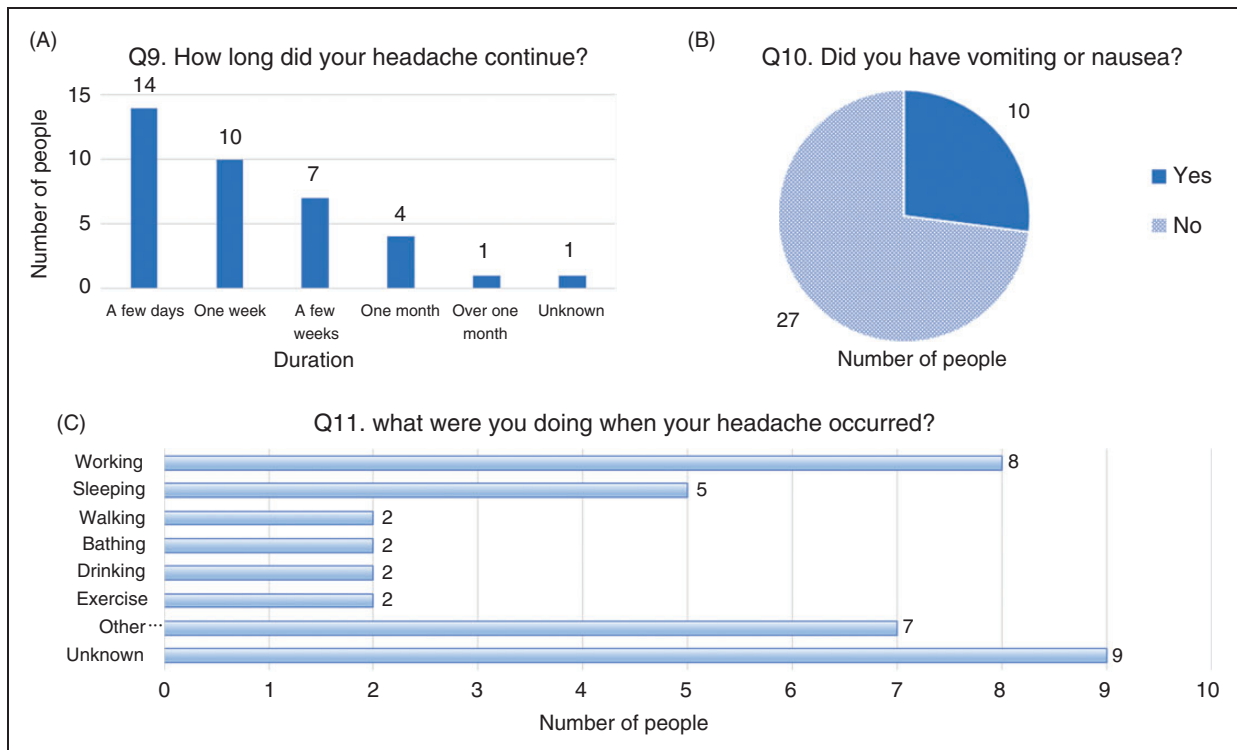


Figure 5. Results of the questionnaire: (A) Q9; (B) Q10; (C) Q11.

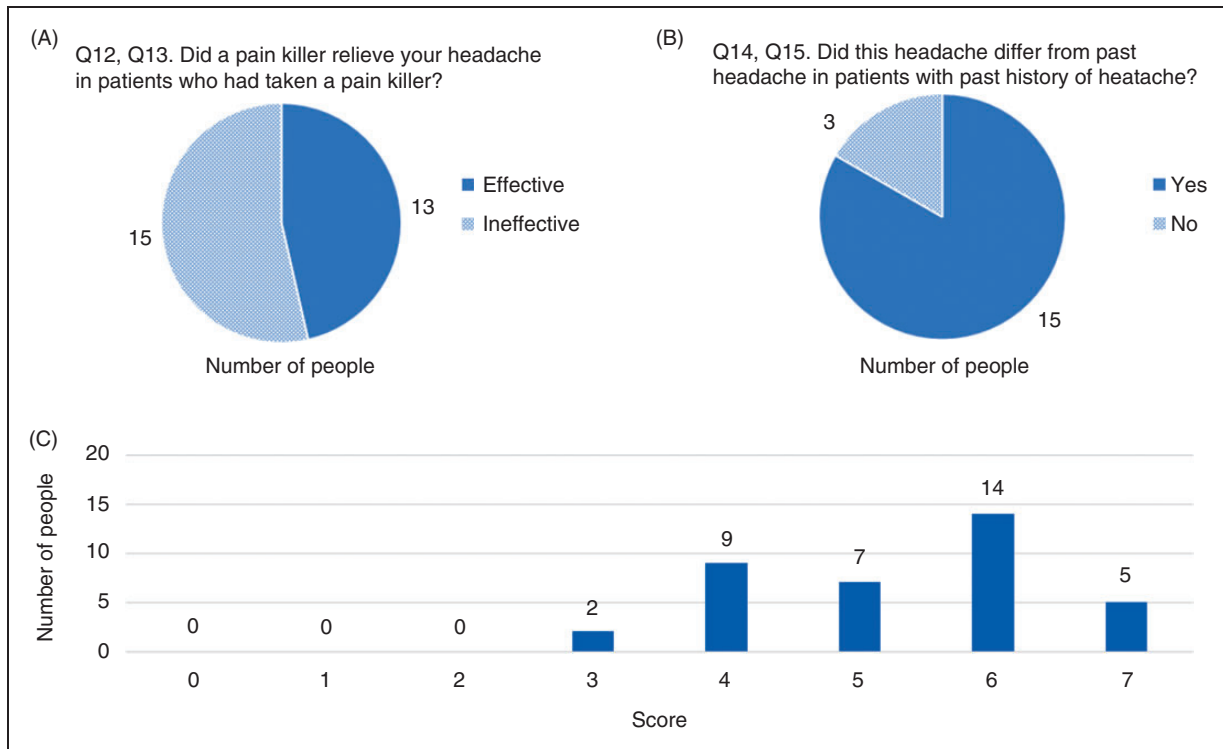


Figure 6. Results of the questionnaire: (A) Q12 and 13; (B) Q14 and 15; (C) patient scores in our scoring system based on the headache characteristics demonstrated by our patients.

while the patients were awake (19/28, 68%), some patients developed headache while sleeping and were woken up by the pain (5/28, 18%). On the other hand, the time of onset of the pain in the remaining nine patients was unknown (Figure 5C). Twenty-eight of 37 patients took painkillers. The number of patients who responded to painkillers was almost the same as those who did not experience a response (effective: 13/28, 46%, ineffective: 15/28, 54%) (Figure 6A). Although 18 of 37 patients (49%) had a previous history of headache, 15 of them (83%) answered that the present headache was different from their previous headache (Figure 6B).

Comparison of headache characteristics between headache type and ischemic type IC-VAD

Comparison of headache characteristics between headache type and ischemic type IC-VAD (Table 3) indicated that the headache was significantly more severe in headache type than ischemic type IC-VAD, based on NRS scores and the rating of this being the worst headache ever experienced by the patient ($p=0.01$). More patients with ischemic type IC-VAD had concomitant clinical symptoms as compared to those with headache type IC-VAD ($p=0.03$). Patients with ischemic type IC-VAD also more often had nausea or vomiting as

Table 3. Comparison of the characteristics of the headache between the two types of unruptured intracranial vertebral artery dissection.

	Headache type (n=24)	Ischemic type (n=13)	p-value
Age	49.5 ± 9.1	46.3 ± 8.1	0.29 ^a
Sex (M:F)	14:10	9:4	0.51 ^b
Headache in the occipitounuchal region	22 (92%)	11 (85%)	0.24 ^b
Acute onset	17 (71%)	9 (69%)	0.91 ^b
Unilateral headache	18 (75%)	12 (92%)	0.19 ^b
Ipsilateral to the IC-VAD	15 (63%)	9 (69%)	0.57 ^b
Pulsatile headache	18 (75%)	8 (62%)	0.39 ^b
Headache intensity (NRS)	7.8 ± 3.8	5.9 ± 6.9	0.01 ^a
Worst headache ever	19 (79%)	5 (38%)	0.01 ^b
Pain aggravating factors	9 (38%)	6 (46%)	0.6 ^b
Nausea or vomiting	4 (17%)	6 (46%)	0.053 ^b
Headache while sleeping	2/16 (13%)	3/12 (25%)	0.39 ^b
Concomitant clinical symptoms	17 (71%)	13 (100%)	0.03 ^b
Effect of painkiller	9/20 (45%)	4/8 (50%)	0.81 ^b
Duration of headache			0.77 ^c
few days	9	5	
1 week	7	3	
few weeks	4	3	

(continued)

Table 3. Continued.

	Headache type (n = 24)	Ischemic type (n = 13)	p-value
1 month	3	1	
over 1 month	1	0	
unknown	0	1	
History of headache	11 (46%)	6 (46%)	0.98 ^b

IC-VAD: intracranial vertebral artery dissection; NRS: Numerical Rating Scale.

^aStudent's *t*-test; ^bchi-squared test for independence; ^cMann-Whitney *U*-test.

compared to those with headache type IC-VAD, although the difference was statistically insignificant. There were no differences in age, sex, other headache characteristics, or history of headache between the two types of IC-VAD.

Validation of the consistency of previous diagnostic criteria for intracranial artery dissection by comparing them with our results

Table 4 summarizes the diagnostic criteria in the ICHD-3 beta and Clinical Practice Guideline for Chronic Headache 2013, which showed consistency with the criteria in our study (17,18). Our results completely satisfied the comprehensive diagnostic criteria of the ICHD-3 beta. Among the required criteria, duration showed the highest consistency (97%). Other required criteria, such as pain intensity and location, were fulfilled in approximately 70% of our patients. However, the mode of onset criterion was not fulfilled by any of our patients (0%). On the other hand, the comprehensive headache characteristics in the Clinical Practice Guideline for Chronic Headache 2013 were fulfilled by 35% of the patients. However, location and duration criteria were fulfilled by 89% and 97% of our patients.

Scoring system for diagnosis of the headache of unruptured IC-VAD

Based on our indicated headache characteristics, a scoring system for diagnosis of the headache was devised, as shown in Table 5. The results are shown in Figure 6C. None of the patients had scores of 0–2, while scores of over 4 points were seen in 95% of the patients.

Discussion

Recently, the detection and diagnosis of unruptured IC-VAD has increased due to

Table 4. Diagnostic criteria in the ICHD-3 beta and Clinical Practice Guideline for Chronic Headache 2013 and their consistency relative to the criteria in our study.

Criteria	Consistency
<i>ICHD-3 beta</i>	
Evidence of causation is demonstrated by the presence of at least two of the following:	100% (37/37)
1. Headache has developed in close temporal relation to other symptoms and/or clinical signs of intracranial dissection, or has led to the diagnosis of intracranial dissection	100% (37/37)
2. Headache resolves within 1 month of its onset	97% (36/37)
3. Headache has either or both of the following characteristics:	73% (27/37)
a) sudden or thunderclap onset	0% (0/37)
b) severe intensity	73% (27/37)
4. Headache is unilateral and ipsilateral to the dissection	65% (24/37)
<i>Clinical Practice Guideline for Chronic Headache 2013</i>	
1. Headache is new, acute onset headache, with facial or neck pain, usually unilateral (ipsilateral to dissection), and severe	35% (13/37)
2. The pain is mostly localized to the back of the head or the neck	89% (33/37)
3. The pain is persistent, but resolves within 1 month	97% (36/37)

ICHD-3 beta: The International Classification of Headache Disorders, 3rd edition (beta version).

advancements in MRI (19,20). A Japanese nationwide study on IC-VAD reported that unruptured IC-VAD accounted for 69.5% of IC-VAD cases (14). In that study, among the unruptured IC-VAD, 47.6% of cases were of the ischemic type and 52.4% of cases were of the headache type. In ischemic type IC-VAD, 56.9% of the cases presented with headache (14). Thus, the majority of unruptured IC-VAD cases experience headaches.

Pain caused by cervicocephalic arterial dissection is thought to be referred pain associated with the nerves innervating the vessels (9,18). The sites of headache and neck pain differ depending on the dissected vessels. In dissections in the vertebro-basilar arterial system, unilateral cervical pain and occipital pain are frequent. These pains can be triggered by stimulation of the artery. Nichols et al. (21) reported that balloon angioplasty in the vertebro-basilar arterial system caused localized ipsilateral occipital and posterior cervical pain. Hence, headache in the occipitonal

region is considered as the major clinical symptom of IC-VAD.

Characteristics of unruptured IC-VAD-related headache

Our study showed that the characteristics of headache due to unruptured IC-VAD are as described below: (i) presenting in the occipitonal region (89%); (ii) unilateral (81%); (iii) pulsatile (70%); (iv) of acute onset (70%); (v) severe (73%); (vi) with no associated nausea or vomiting (73%); and (vii) with the presence of concomitant clinical symptoms (81%). Further, the headache resolved within 1 month of its onset in almost all our patients. Several previous reports have investigated the characteristics of headache due to unruptured IC-VAD (6,9,10). Their results almost completely corresponded with our results in terms of the headache characteristics. Kim et al. (6) reported that headache that is exacerbated or attenuated by head motion or a specific head position is a specific feature of extracranial VAD. However, this feature is not described in other reports, including in the ICHD-3 beta. Our results showed that 15 patients (40%) had factors that aggravated the pain. Although body motion exacerbated the headache in 10 patients, the headache was not exacerbated by head motion in any of our patients.

On the other hand, in our study, 30 patients (81%) had concomitant clinical symptoms when the headache occurred. This feature has not been described in previous reports. Since IC-VAD can cause ischemia in the posterior circulation, ischemic symptoms, such as visual disturbances, vertigo, ataxia, and sensory disturbances, can accompany the headache. However, the associated clinical symptoms in this study were unlikely to have been ischemic symptoms. Although the mechanisms of these clinical symptoms are unclear, asking patients about these symptoms might aid in the diagnosis of unruptured IC-VAD.

Differences in headache characteristics between the two types of unruptured IC-VAD

Our results showed that intensity of the headache in headache type IC-VAD was significantly more severe than in the ischemic type. Moreover, the number of patients describing the headache as the worst they had ever felt was significantly more in the headache type IC-VAD than in the ischemic type. Since headache intensity was measured by the patient's subjective assessment, concomitant ischemic symptoms might have masked the headache intensity in patients with ischemic type IC-VAD. On the other hand,

Table 5. Our headache scoring system.

Headache characteristic	Score
1. Headache in the occipitonal region	1
2. Unilateral headache	1
3. Pulsatile headache	1
4. Acute or sudden onset headache	1
5. Severe headache	1
6. Headache without nausea or vomiting	1
7. Headache with concomitant clinical symptoms, such as irritated eyes, dizziness, transient dysesthesia, listlessness, fever, or neck stiffness	1

concomitant clinical symptoms other than ischemic symptoms were observed significantly more often in patients with ischemic type than in headache type IC-VAD. However, over 70% of patients with headache type IC-VAD also showed concomitant clinical symptoms. Hence, the presence of concomitant clinical symptoms is a common characteristic in both types of dissection, despite the statistically significant difference.

Validation of the consistency of previous diagnostic criteria of intracranial artery dissection by comparing them with our results

The headache characteristics of unruptured IC-VAD shown by us corresponded mostly with those shown in ICHD-3 beta; each criterion of duration, intensity, and laterality was satisfied in over 65% of our patients. However, the mode of onset criteria may be controversial. In ICHD-3 beta, the mode of headache onset is described as 'sudden or thunderclap onset' (17). On the other hand, in the Clinical Practice Guideline for Chronic Headache 2013, 'acute onset headache' is cited as being characteristic (18). This raises the question as to whether the major mode of onset is 'sudden' or 'acute'. Strictly speaking, 'sudden onset headache' means head pain reaching maximal intensity within seconds to a minute. On the other hand, 'acute onset headache' means head pain reaching maximal intensity within a few to several tens of minutes. Our results showed that 70% of the patients had acute onset headache. Maruyama et al. (9) also reported that 80% of their patients had acute onset headache. Kim et al. (6), who also validated the ICHD-3 beta criteria, showed low consistency in mode of onset criteria (7.1%). Their results showed that the headache in most patients had an acute onset (79%), and they suggested that inclusion of 'acute onset' should be considered to achieve better criteria performance. Strictly speaking, 'acute' and 'sudden' onset represents a different time course.

However, the fine distinction between ‘acute’ and ‘sudden’ onset may make its differentiation based only on medical interviews difficult. If the question regarding mode of onset focuses on whether the onset was gradual or not, the description ‘acute or sudden onset’ may be suitable. Hence, we also suggest including ‘acute or sudden onset’ in the criteria.

In the Clinical Practice Guideline for Chronic Headache 2013, mode of onset, laterality, and pain intensity are combined as general headache characteristics (18). However, location and duration are independent characteristics. Our results showed that only 35% of our IC-VAD cases satisfied these general headache characteristics. However, the characteristics of location and duration were satisfied by 89% and 97% of the patients. When the general headache characteristics were considered separately, the criteria of mode of onset, laterality, and pain intensity were satisfied by 70%, 65%, and 73% of IC-VAD cases. Hence, our results indicate that the description of headache characteristics should probably be subdivided to increase the accuracy of the diagnostic criteria like ICHD-3 beta.

Differentiation of IC-VAD headache from primary headaches

It is very important to differentiate headache due to IC-VAD from primary headaches such as migraine, tension-type headache or cluster headache (6,17). In general, severe and unusually persistent occipitonal pain is suggestive of intracranial VAD. However, headache due to IC-VAD without neurological signs are often misdiagnosed as tension-type headache because tension-type headaches commonly occur in posterior locations (6,15). Severe and pulsatile headache in the occipitonal region may be misdiagnosed as atypical migraine. Our results showed that some patients had headache in the temporal region or face. In such cases, physicians may suspect migraine or cluster headache. Moreover, the diagnosis is even more difficult when patients with a past history of headache develop

IC-VAD headache, since, in such cases, physicians may suspect recurrence of the past headache. In fact, there are some reports of difficulty in the diagnosis due to this reason (22,23). However, our results showed that IC-VAD-related headache was different from the previous headache in most patients (83%). This may be helpful in the diagnosis. Based on our indicated headache characteristics, we attempted to establish a scoring system for diagnosis of the headache. In the current study, none of the patients had scores of 0–2, while scores of over 4 points were seen in 95% of the patients. Hence, a score of over 4 points on this scoring system may enable the diagnosis of IC-VAD.

Limitations

This study has some limitations. First, our study involved only a small number of subjects; hence, our findings regarding headache characteristics need to be validated in a larger study. Therefore, we cannot give definitive suggestions for improvement of the ICHD-3 beta, which has been established after much research. Second, because this study covered a long time span, there is a possibility of recall bias. Third, although we suggested a scoring system for the diagnosis, we did not prove its accuracy. The scoring system needs to be tested in a non IC-VAD group with primary headache. Finally, investigation of the sensitivity and specificity of this scoring system using a prospective study is needed.

Conclusion

Our results identified seven headache characteristics in patients with IC-VAD. These headache characteristics almost completely corresponded with those shown in ICHD-3 beta and the Clinical Practice Guideline for Chronic Headache 2013. However, headache with concomitant clinical symptoms that are distinct from ischemic symptoms and headache without nausea or vomiting are not described in these previous tools. We believe that the headache characteristics shown in this study may be helpful in the diagnosis of IC-VAD.

Clinical implications

- We clarified the characteristics of intracranial vertebral artery dissection (IC-VAD)-related headache using our original self-administered questionnaire.
- Our results identified the following seven headache characteristics in patients with IC-VAD: occurring in the occipitonal region, unilateral, pulsatile, of acute onset, severe, without nausea or vomiting, and with concomitant clinical symptoms unrelated to ischemia.
- Based on our evaluation, the presence of four or more of these characteristics may suggest the existence of IC-VAD.

Declaration of conflicting interests

The authors declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

Funding

The authors received no financial support for the research, authorship, and/or publication of this article.

References

1. Arnold M, Boussier MG, Fahrni G, et al. Vertebral artery dissection: Presenting findings and predictors of outcome. *Stroke* 2006; 37: 2499–2503.
2. Kim BM, Kim SH, Kim DI, et al. Outcomes and prognostic factors of intracranial unruptured vertebral artery dissection. *Neurology* 2011; 76: 1735–1741.
3. Mizutani T. Natural course of intracranial arterial dissections. *J Neurosurg* 2011; 114: 1037–1044.
4. Schievink WI. Spontaneous dissection of the carotid and vertebral arteries. *N Engl J Med* 2001; 344: 898–906.
5. Putaala J, Metso AJ, Metso TM, et al. Analysis of 1008 consecutive patients aged 15 to 49 with first-ever ischemic stroke: The Helsinki young stroke registry. *Stroke* 2009; 40: 1195–1203.
6. Kim JG, Choi JY, Kim SU, et al. Headache characteristics of uncomplicated intracranial vertebral artery dissection and validation of ICHD-3 beta diagnostic criteria for headache attributed to intracranial artery dissection. *Cephalalgia* 2015; 35: 516–526.
7. Tsukahara T and Minematsu K. Overview of spontaneous cervicocephalic arterial dissection in Japan. *Acta Neurochir Suppl* 2010; 107: 35–40.
8. Shibahara T, Yasaka M, Wakugawa Y, et al. Improvement and aggravation of spontaneous unruptured vertebral artery dissection. *Cerebrovasc Dis Extra* 2017; 7: 153–164.
9. Maruyama H, Nagoya H, Kato Y, et al. Spontaneous cervicocephalic arterial dissection with headache and neck pain as the only symptom. *J Headache Pain* 2012; 13: 247–253.
10. Nakamura H, Mizuniwa Y, Kouno T, et al. Clinical characteristics of unruptured vertebral artery dissections presenting with headaches. *Jpn J Stroke* 2011; 33: 333–340.
11. Campos CR, Calderaro M, Scaff M, et al. Primary headaches and painful spontaneous cervical artery dissection. *J Headache Pain* 2007; 8: 180–184.
12. Evans RW and Mokri B. Headache in cervical artery dissections. *Headache* 2002; 42: 1061–1063.
13. Sheikh HU. Headache in intracranial and cervical artery dissections. *Curr Pain Headache Rep* 2016; 20: 8.
14. Ono J, Higuchi Y, Tajima Y, et al. Outline of a nationwide study of non-traumatic intracranial arterial dissection in the vertebrobasilar system: A final report. *Surg Cereb Stroke* 2015; 43: 245–251.
15. Silbert PL, Mokri B and Schievink WI. Headache and neck pain in spontaneous internal carotid and vertebral artery dissections. *Neurology* 1995; 45: 1517–1522.
16. Von Babo M, De Marchis GM, Sarikaya H, et al. Differences and similarities between spontaneous dissections of the internal carotid artery and the vertebral artery. *Stroke* 2013; 44: 1537–1542.
17. Headache Classification Committee of the International Headache Society. The International Classification of Headache Disorders, 3rd edition (beta version). *Cephalalgia* 2013; 33: 629–808.
18. Chronic Headache Clinical Practice Guideline Development Committee of the Japanese Headache Society. *Clinical Practice Guideline for Chronic Headache 2013*. Tokyo: Igakusyoin, 2013.
19. Takemoto K, Takano K, Abe H, et al. The new MRI modalities “BPAS and VISTA” for the diagnosis of VA dissection. *Acta Neurochir Suppl* 2011; 112: 59–65.
20. Yamada S, Ohnishi H, Takamura Y, et al. Diagnosing intra-cranial and cervical artery dissection using MRI as the initial modality. *J Clin Neurosci* 2016; 33: 177–181.
21. Nichols FT 3rd, Mawad M, Mohr JP, et al. Focal headache during balloon inflation in the vertebral and basilar arteries. *Headache* 1993; 33: 87–89.
22. Morelli N, Mancuso M, Gori S, et al. Vertebral artery dissection onset mimics migraine with aura in a graphic designer. *Headache* 2008; 48: 621–624.
23. Ohara H, Nakazima T and Kabuto M. A case of vertebral artery dissection that required differentiation from migraine. *J Jpn Soc Pain Clinicians* 2014; 21: 27–30.