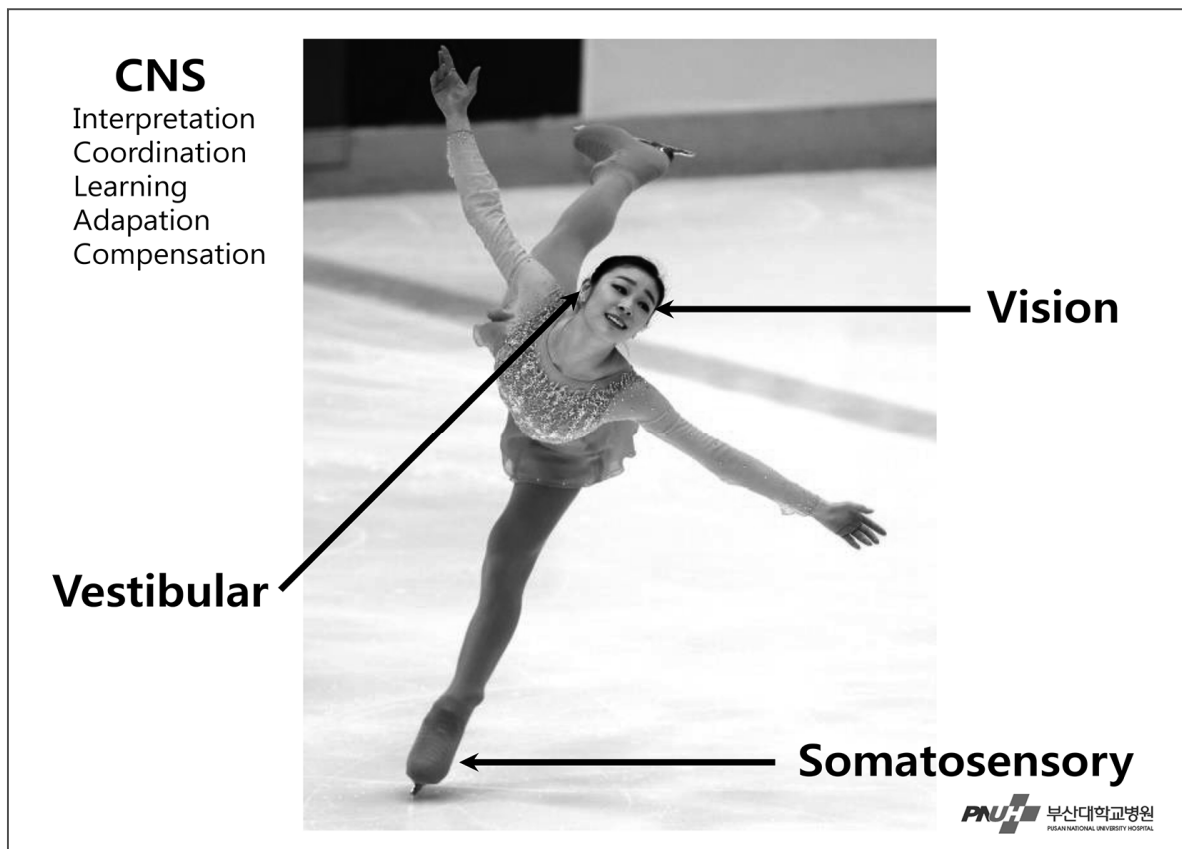


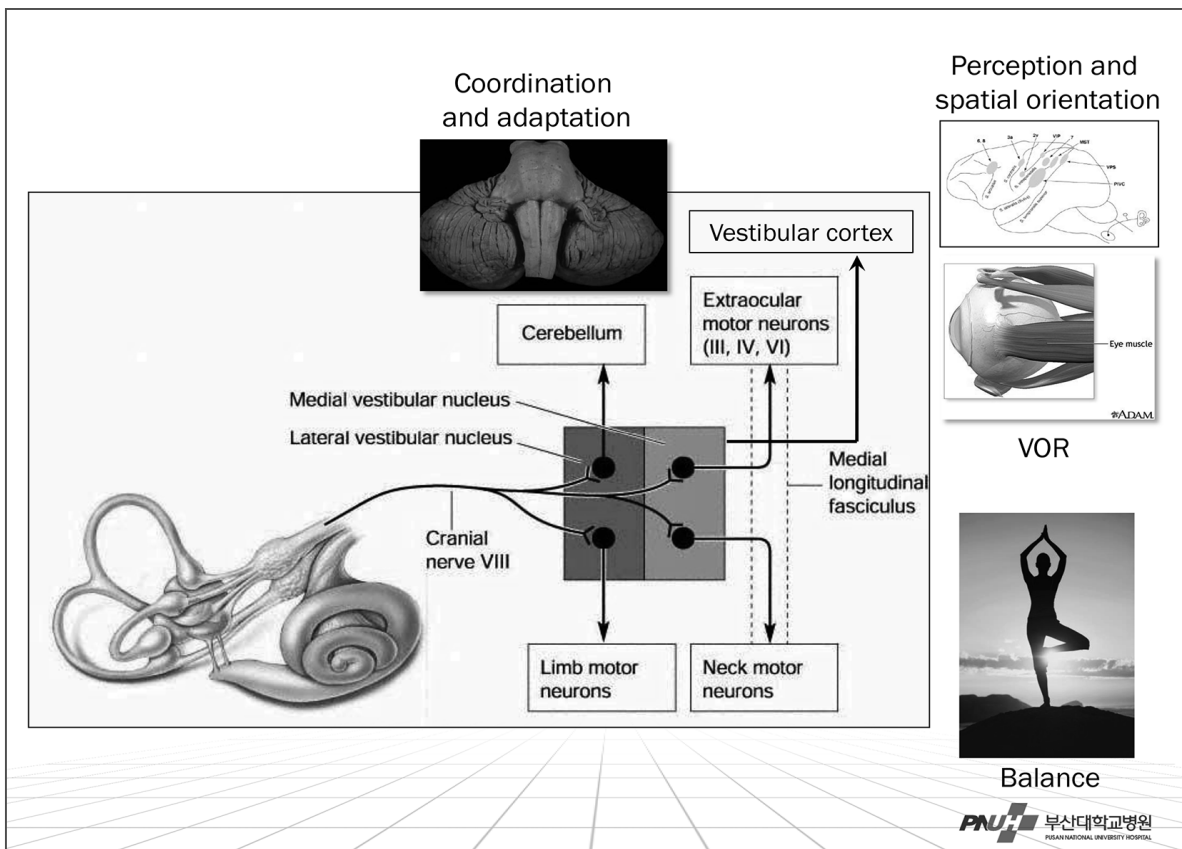
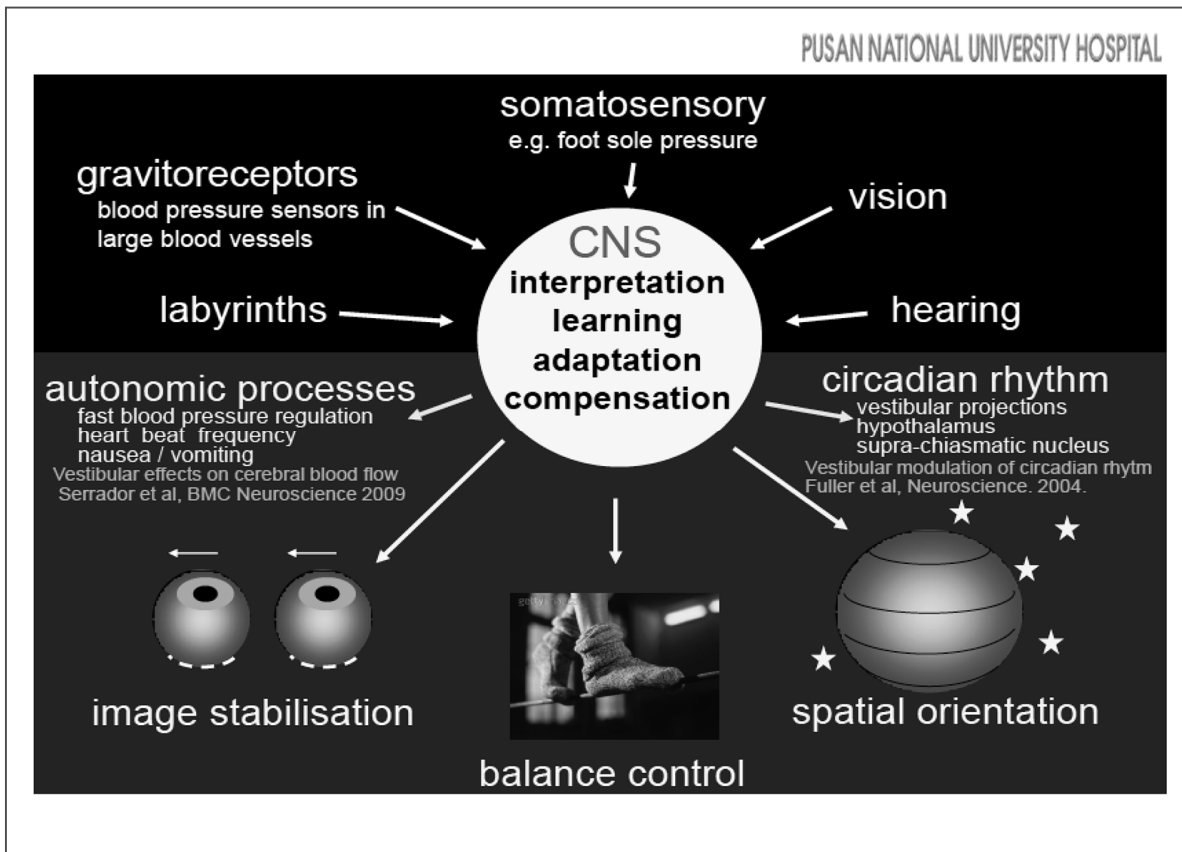
중추성현훈의 감별진단



최 광 등

부산대학교 의과대학 신경과학교실





Approach to dizzy patients

- History taking
- ↓
- Bedside evaluation
- ↓
- Laboratory evaluation

The 'Quality-of-Symptoms' Approach

Type of dizziness implies pathophysiologic mechanism

- Vertigo = vestibular = ear/ENT problem
- Presyncope = cardiovascular = Med/Cards problem
- Disequilibrium = neurologic = Neuro problem
- Non-specific dizziness = psychiatric = NP problem

→ Old approach
Quality approach doesn't work!

PUSAN NATIONAL UNIVERSITY HOSPITAL

The Old Approach Quality Doesn't Work

- “vestibular” system
 - peripheral vestibular (benign inner ear)
 - central vestibular (bleed, stroke, TIA)
 - cortical projections (migraine, psych)
 - spinal cord projections (cord compression)

non-specific dizziness
presyncope

ANY OF THE FOUR QUALITATIVE SYMPTOMS

vertigo

imbalance

PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

PUSAN NATIONAL UNIVERSITY HOSPITAL

The Old Approach Quality Approach Doesn't Work

Cerebrovascular Dizziness is of All 4 Types

- 59% of cerebellar strokes present with vertigo (n=66) (Kase, 1993)
- 41% of cerebellar strokes have non-specific dizziness or disequilibrium, without vertigo (n=66) (Kase, 1993)
- 32% of vertebrobasilar insufficiency patients have pre-syncope/syncope (n=39) (Toursarkissian, 1998)

PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

ORIGINAL ARTICLE

Vertigo and nystagmus in orthostatic hypotension

J.-H. Choi^a, J.-D. Seo^{b,c}, M.-J. Kim^b, B.-Y. Choi^b, Y. R. Choi^b, B. M. Cho^d, J. S. Kim^e and K.-D. Choi^b

^aDepartment of Neurology, Pusan National University School of Medicine, Research Institute for Convergence of Biomedical Science and Technology, Pusan National University Yangsan Hospital, Yangsan; ^bDepartment of Neurology, Pusan National University Hospital, Pusan National University School of Medicine and Biomedical Research Institute, Busan; ^cDepartment of Neurology, Bonhosipital, Busan; ^dDepartment of Preventive Medicine, Pusan National University School of Medicine, Research Institute for Convergence of Biomedical Science and Technology, Pusan National University Yangsan Hospital, Yangsan; and ^eDepartment of Neurology, Seoul National University Bundang Hospital, Seongnam, Korea

Keywords:

cerebral ischaemia, nystagmus, orthostatic hypotension, orthostatic vertigo, vertebrobasilar insufficiency

Received 29 June 2014
Accepted 7 October 2014

European Journal of
Neurology 2015, 22: 648–655

doi:10.1111/ene.12622

Background and purpose: Generalized cerebral ischaemia from cardiovascular dysfunction usually leads to presyncopal dizziness, but several studies reported a higher frequency of rotatory vertigo in cardiovascular patients. Whether generalized cerebral ischaemia due to cardiovascular disorders may produce objective vestibular dysfunction was investigated.

Methods: Thirty-three patients with orthostatic dizziness/vertigo due to profound orthostatic hypotension and 30 controls were recruited. All participants underwent recording of eye movements during two orthostatic challenging tests: the Schellong and the squatting–standing tests. Most patients had neuroimaging, and patients with abnormal eye movements were subjected to follow-up evaluations.

Results: Symptoms associated with orthostatic dizziness/vertigo included blurred vision, fainting and tinnitus. Ten (30%) of 33 patients developed rotatory vertigo and nystagmus during the Schellong ($n = 5$) or squatting–standing test ($n = 5$). Four of them showed pure downbeat nystagmus whilst five had downbeat and horizontal nystagmus with or without torsional component. Patients with orthostatic nystagmus had shorter duration of orthostatic intolerance than those without nystagmus (1.0 ± 1.6 vs. 11.0 ± 9.7 months, $P < 0.001$). In two patients, orthostatic nystagmus disappeared during follow-up despite the persistence of profound orthostatic hypotension.

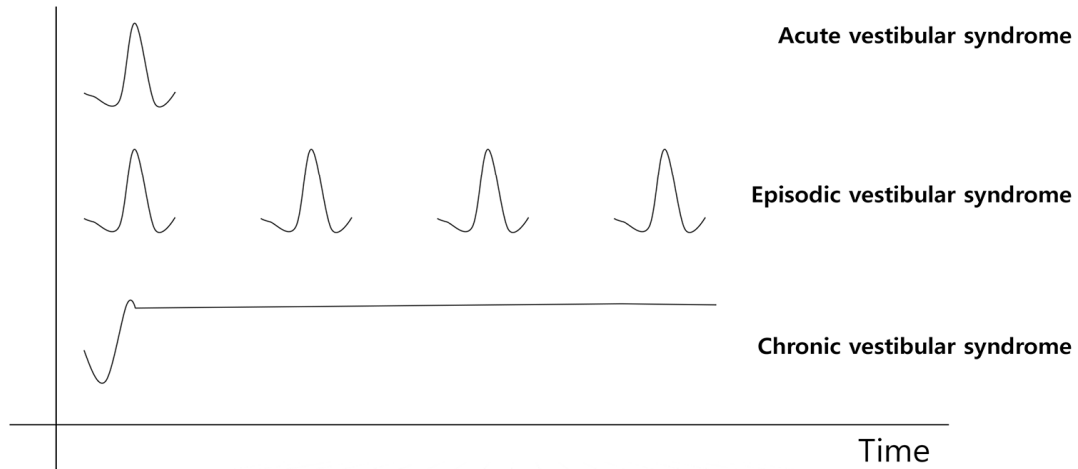
Conclusions: Generalized cerebral ischaemia caused by orthostatic hypotension induces rotatory vertigo due to objective vestibular dysfunction. The presence of orthostatic vertigo and nystagmus has an association with the duration of orthostatic intolerance.

A new approach!!

✓Timing

✓Triggers and duration

Vestibular Syndromes (Timing)



PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

Vestibular Syndromes (Timing)

Acute vestibular syndrome

- Vestibular neuritis/labyrinthitis
- Brainstem or cerebellar stroke

Episodic vestibular syndrome

- Vestibular migraine
- Meniere's disease
- TIA
- BPPV
- Central positional vertigo
- Rotational vertebral artery syndrome
- Orthostatic hypotension
- Cardiogenic vertigo

Chronic vestibular syndrome

- Chronic subjective dizziness
- Cerebellar ataxia
- PSP
- Post-concussion syndrome
- Bilateral vestibulopathy
- Drug-induced dizziness

PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

Acute vestibular syndrome



Vestibular Syndromes (Timing)

Acute vestibular syndrome

- Vestibular neuritis/
labyrinthitis
- Brainstem or cerebellar
stroke

Acute vestibular syndrome (AVS)

- ✓ Rapid onset (over seconds to hours)
- ✓ Vertigo, N/V, gait unsteadiness
- ✓ Head motion intolerance and nystagmus
- ✓ Lasting more than 24 hours

Acute vestibular syndrome (AVS)

- ✓ Acute peripheral vestibulopathy vs central vestibulopathy
- ✓ Previous studies
 - : 20-60% of AVS – posterior circulation stroke

Acute vestibular syndrome (AVS)

- ✓ CT scans : low sensitivity (16%) for acute infarction
- ✓ MRI : high cost,
false-negative for acute infarction (12-20%)
- ✓ Long-tract or cerebellar signs
: fewer than half of AVS
: isolated vertigo without neurological signs??
- ✓ Careful eye movement assessment !!

Acute vestibular syndrome ; HINTS examination (Head impulse test, direction changing nystagmus, skew deviation)

HINTS to Diagnose Stroke in the Acute Vestibular Syndrome

Three-Step Bedside Oculomotor Examination More Sensitive Than Early MRI Diffusion-Weighted Imaging

Jorge C. Kattah, MD; Arun V. Talkad, MD; David Z. Wang, DO;
Yu-Hsiang Hsieh, PhD, MS; David E. Newman-Toker, MD, PhD

Background and Purpose—Acute vestibular syndrome (AVS) is often due to vestibular neuritis but can result from vertebrobasilar strokes. Misdiagnosis of posterior fossa infarcts in emergency care settings is frequent. Bedside oculomotor findings may reliably identify stroke in AVS, but prospective studies have been lacking.

Methods—The authors conducted a prospective, cross-sectional study at an academic hospital. Consecutive patients with AVS (vertigo, nystagmus, nausea/vomiting, head-motion intolerance, unsteady gait) with ≥ 1 stroke risk factor underwent structured examination, including horizontal head impulse test of vestibulo-ocular reflex function, observation of nystagmus in different gaze positions, and prism cross-cover test of ocular alignment. All underwent neuroimaging and admission (generally < 72 hours after symptom onset). Strokes were diagnosed by MRI or CT. Peripheral lesions were diagnosed by normal MRI and clinical follow-up.

Results—One hundred one high-risk patients with AVS included 25 peripheral and 76 central lesions (69 ischemic strokes, 4 hemorrhages, 3 other). The presence of normal horizontal head impulse test, direction-changing nystagmus in eccentric gaze, or skew deviation (vertical ocular misalignment) was 100% sensitive and 96% specific for stroke. Skew was present in 17% and associated with brainstem lesions (4% peripheral, 4% pure cerebellar, 30% brainstem involvement; χ^2 , $P=0.003$). Skew correctly predicted lateral pontine stroke in 2 of 3 cases in which an abnormal horizontal head impulse test erroneously suggested peripheral localization. Initial MRI diffusion-weighted imaging was falsely negative in 12% (all < 48 hours after symptom onset).

Conclusions—Skew predicts brainstem involvement in AVS and can identify stroke when an abnormal horizontal head impulse test falsely suggests a peripheral lesion. A 3-step bedside oculomotor examination (HINTS: Head-Impulse—Nystagmus—Test-of-Skew) appears more sensitive for stroke than early MRI in AVS. (*Stroke*. 2009;40:3504-3510.)

Acute peripheral vestibulopathy

- Contralateral horizontal-torsional spontaneous nystagmus
 - unidirectional
 - suppression by visual fixation
- Positive head impulse test
- No skew deviation

Acute vestibular syndrome ; dangerous HINTS examination

Direction changing nystagmus

Normal head impulse test

Skew deviation

Acute vestibular syndrome ; HINTS examination

(Head impulse test, direction changing nystagmus, skew deviation)

- HINTS examination is more sensitive than early DWI in excluding stroke maintaining a high specificity.

Table 4. Bedside Signs and Initial MRI With DWI Test Properties for Ischemic Stroke in AVS

	Sensitivity (n=69)	Specificity (n=25)	NLR Stroke (95% CI)
General neurological signs*	19%	100%	0.81 (0.72–0.91)
Obvious oculomotor signs	28%	100%	0.72 (0.63–0.84)
Severe truncal ataxia	33%	100%	0.67 (0.56–0.79)
Any obvious signs	64%†	100%	0.36 (0.27–0.50)
Initial MRI with DWI	88%‡	100%	0.12 (0.06–0.22)
Dangerous bedside HINTS	100%	96%	0.00 (0.00–0.12)

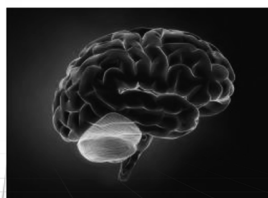
Isolated vestibular syndrome in posterior circulation stroke

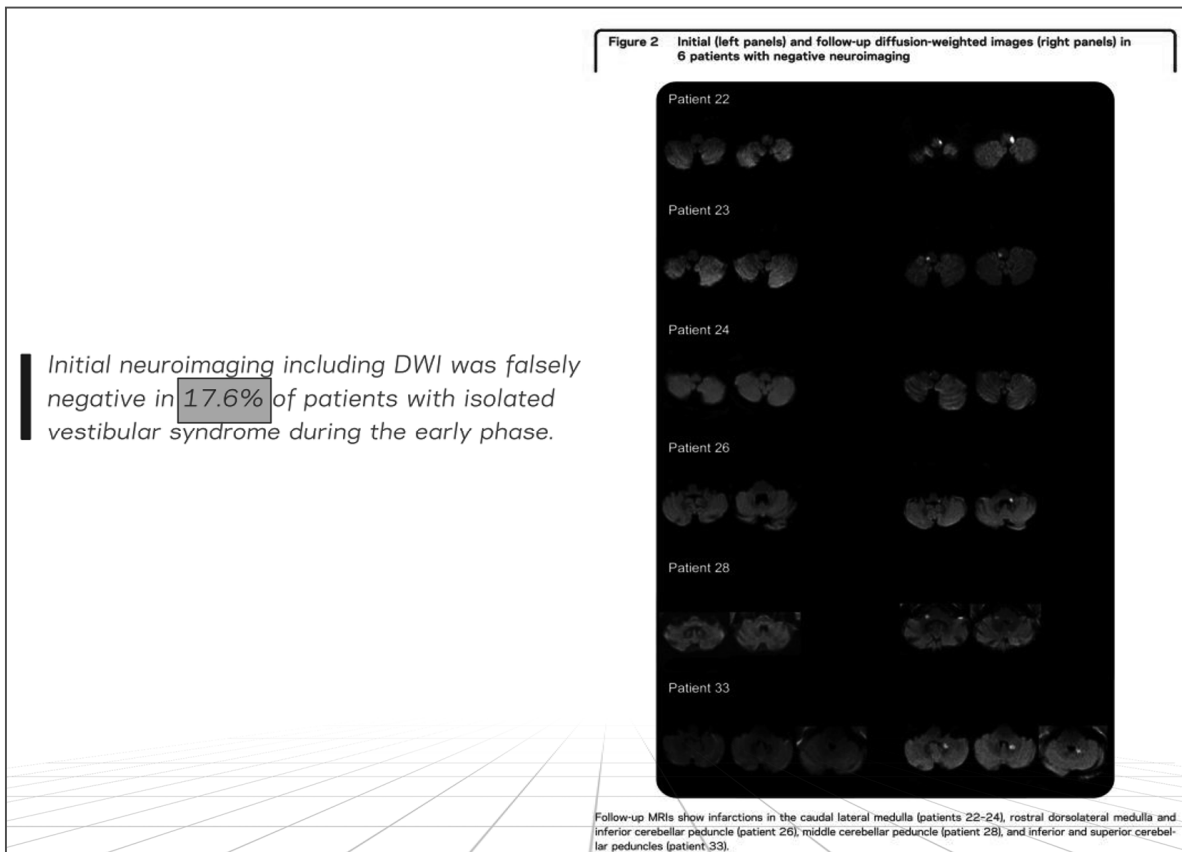
Frequency and involved structures

Jae-Hwan Choi, MD*
Hyun-Woo Kim, MD*
Kwang-Dong Choi, MD
Min-Ji Kim, BSc
Yu Ri Choi, BSc
Han-Jin Cho, MD
Sang-Min Sung, MD
Hak-Jin Kim, MD
Ji-Soo Kim, MD
Dae-Soo Jung, MD

Summary

Dizziness/vertigo is a common symptom of posterior circulation stroke and usually accompanies other neurologic symptoms and signs. Although strokes involving the brainstem or cerebellum may produce isolated vestibular syndrome (isolated vertigo or imbalance), the overall frequency and involved structures of isolated vestibular syndrome in the posterior circulation stroke remain uncertain. Isolated vestibular syndrome occurs in approximately 25% of the patients with posterior circulation stroke, and mostly involves the cerebellum, inferior or superior cerebellar peduncles, and caudal lateral or rostral dorsolateral medulla. The occasional negative neuroimaging in patients with acute isolated vascular vertigo highlights the importance of appropriate bedside evaluation in acute vestibular syndrome.

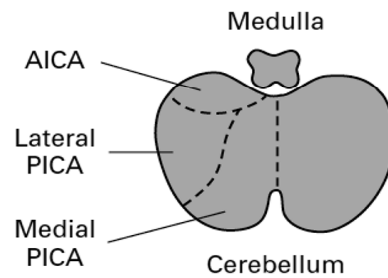
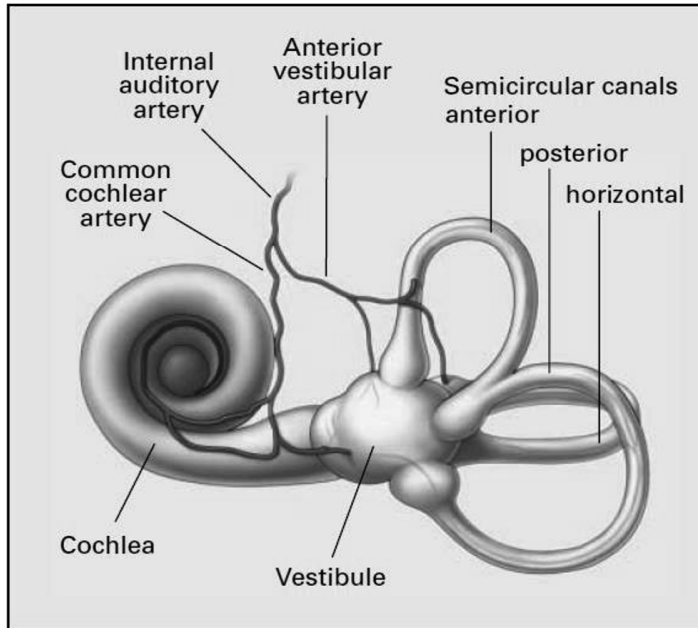




Limitation of HINTS

✓ AICA infarction : benign HINTS

Blood supply to labyrinth



Limitation of HINTS

- ✓ AICA infarction : benign HINTS
- ✓ AVS without nystagmus
- ✓ Rapid resolution of AVS within 24 hours
- ✓ False negative stroke

Acute transient vestibular syndrome (ATVS)

- ✓ Rapid onset (over seconds to hours)
- ✓ Vertigo, N/V, gait unsteadiness
- ✓ Head motion intolerance and nystagmus?
- ✓ Resolution within 24 hours
- ✓ Vertebrobasilar TIA????

Acute transient vestibular syndrome (ATVS)

- ✓ During one year (2014)

713 pts. with first dizziness presentations



86 pts. with ATVS



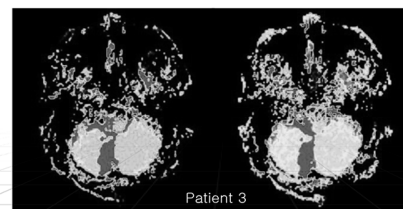
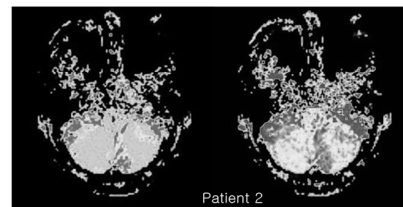
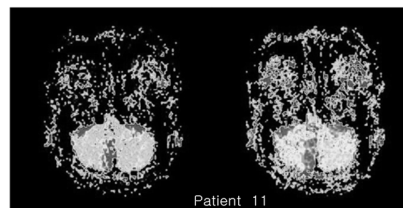
HINTS, MRI, PWI

	ATVS		Total (n=86)
	Nystagmus (n=22, 26%)	No nystagmus (n=64, 74%)	
APV, n (%)	15 (68)	0 (0)	15
Stroke, n (%)	2 (9)	21 (33)	23
cerebral infarction, n (%)	2 (9)	11 (17)	13
cerebellar hypoperfusion, n (%)	0 (0)	10 (16)	10
Undetermined, n (%)	5 (23)	43 (67)	48
HINTS plus			
sensitivity (%)	100	NC	NC
specificity (%)	75	NC	NC
MRI			
sensitivity (%)	100	52	57
specificity (%)	100	100	100

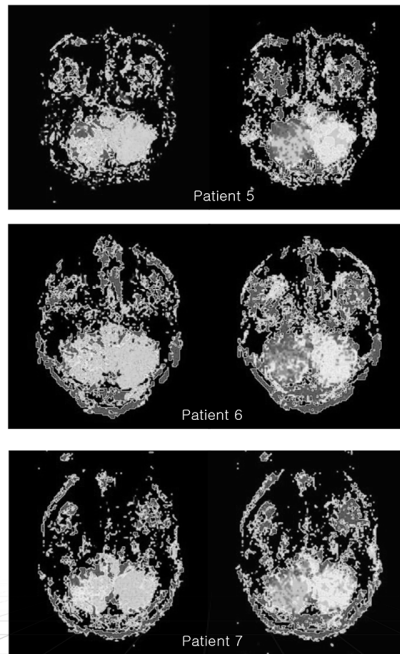
Data are n (%) unless otherwise indicated.

ATVS = acute transient vestibular syndrome; APV = acute peripheral vestibulopathy; NC=not calculated.

➤ Perfusion delay at medial cerebellum (n=3)



➤ Perfusion delay at whole cerebellum (n=7)

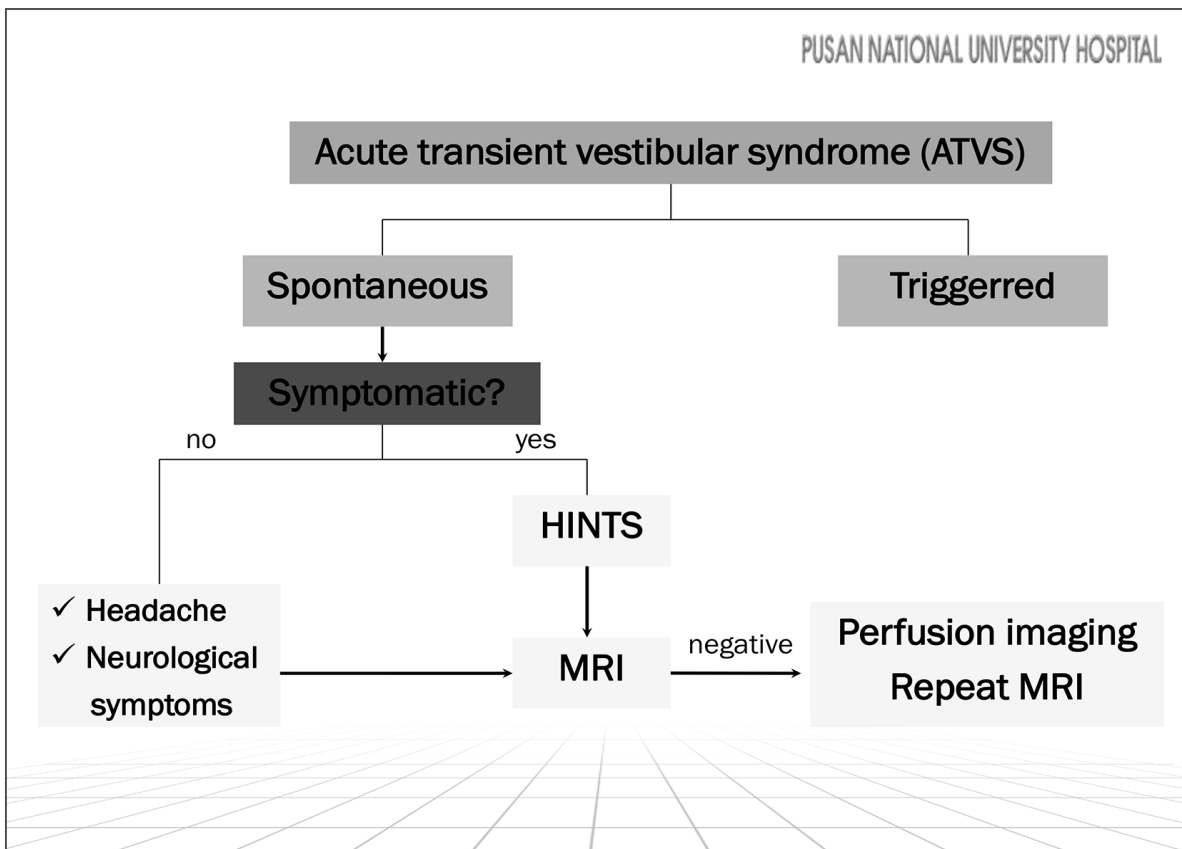
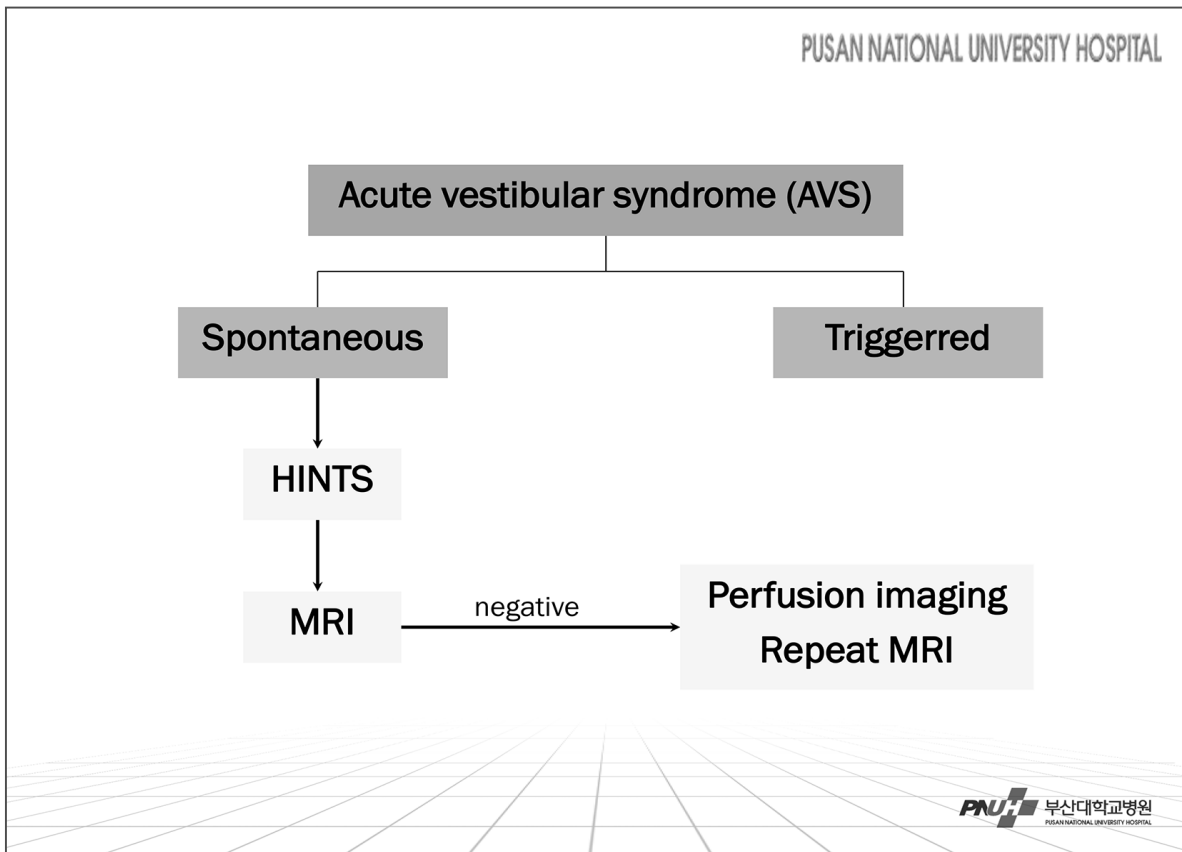


Acute transient vestibular syndrome (ATVS)

Table 5. Results of multivariable logistic regression analysis with dependent variable of stroke in ATVS.

Primary variables	HR (95% CI)	P value
Male gender	4.0 (0.9~17.2)	0.066
Headache	9.6 (2.0~45.2)	0.004
Focal neurologic symptoms/signs	15.2 (2.5~93.8)	0.003
Symptom duration of minutes	2.9 (0.7~11.7)	0.142
VA stenosis or hypoplasia	7.0 (1.7~29.4)	0.008

ATVS = acute transient vestibular syndrome; VA=vertebral artery



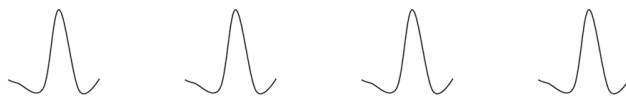
PUSAN NATIONAL UNIVERSITY HOSPITAL



PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

PUSAN NATIONAL UNIVERSITY HOSPITAL

Episodic vestibular syndrome



PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

Vestibular Syndromes (Timing)

Episodic vestibular syndrome

- Vestibular migraine
- Meniere's disease
- TIA
- BPPV
- Central positional vertigo
- Rotational vertebral artery syndrome
- Orthostatic hypotension
- Cardiogenic vertigo

Episodic vestibular syndrome (EVS)

✓ Mostly benign

✓ Caution !!

1. episodic vertigo lasting minutes

VBI (TIA)

2. episodic vertigo lasting seconds

Cardiogenic vertigo

3. episodic vertigo with unilateral hearing loss

Acoustic neuroma

Episodic vestibular syndrome (EVS)

Common

- Meniere's disease
- Vestibular migraine
- Vertebrobasilar insufficiency (VBI)
- BPPV
- Orthostatic hypotension

Uncommon

- Acoustic neuroma
- Episodic ataxia type 2
- Autoimmune inner ear disease
- Otosyphilis
- Multiple sclerosis
- Vestibular epilepsy
- Perilymph fistula/SCD
- Cardiogenic vertigo

Episodic vestibular syndrome (EVS)

Common

- Meniere's disease
- Vestibular migraine
- Vertebrobasilar insufficiency (VBI)
- BPPV
- Orthostatic hypotension

Uncommon

- Acoustic neuroma
- Episodic ataxia type 2
- Autoimmune inner ear disease
- Otosyphilis
- Multiple sclerosis
- Vestibular epilepsy
- Perilymph fistula/SCD
- Cardiogenic vertigo

Episodic vestibular syndrome (EVS)

- Triggers
- Duration
- Auditory symptoms
- Headache
- Neurological symptoms
- Family history

History taking



Interictal exam.



Ictal exam.

- Normal or abnormal

- The most important !!

Triggers – position

Video 1

Video 2

Video 3

ORIGINAL ARTICLE

Vertigo and nystagmus in orthostatic hypotension

J.-H. Choi^a, J.-D. Seo^{b,c}, M.-J. Kim^b, B.-Y. Choi^b, Y. R. Choi^b, B. M. Cho^d, J. S. Kim^e and K.-D. Choi^b

^aDepartment of Neurology, Pusan National University School of Medicine, Research Institute for Convergence of Biomedical Science and Technology, Pusan National University Yangsan Hospital, Yangsan; ^bDepartment of Neurology, Pusan National University Hospital, Pusan National University School of Medicine and Biomedical Research Institute, Busan; ^cDepartment of Neurology, Bonhosipital, Busan; ^dDepartment of Preventive Medicine, Pusan National University School of Medicine, Research Institute for Convergence of Biomedical Science and Technology, Pusan National University Yangsan Hospital, Yangsan; and ^eDepartment of Neurology, Seoul National University Bundang Hospital, Seongnam, Korea

Keywords:

cerebral ischaemia, nystagmus, orthostatic hypotension, orthostatic vertigo, vertebrobasilar insufficiency

Received 29 June 2014
Accepted 7 October 2014

European Journal of Neurology 2015, **22**: 648–655

doi:10.1111/ene.12622

Background and purpose: Generalized cerebral ischaemia from cardiovascular dysfunction usually leads to presyncopal dizziness, but several studies reported a higher frequency of rotatory vertigo in cardiovascular patients. Whether generalized cerebral ischaemia due to cardiovascular disorders may produce objective vestibular dysfunction was investigated.

Methods: Thirty-three patients with orthostatic dizziness/vertigo due to profound orthostatic hypotension and 30 controls were recruited. All participants underwent recording of eye movements during two orthostatic challenging tests: the Schellong and the squatting–standing tests. Most patients had neuroimaging, and patients with abnormal eye movements were subjected to follow-up evaluations.

Results: Symptoms associated with orthostatic dizziness/vertigo included blurred vision, fainting and tinnitus. Ten (30%) of 33 patients developed rotatory vertigo and nystagmus during the Schellong ($n = 5$) or squatting–standing test ($n = 5$). Four of them showed pure downbeat nystagmus whilst five had downbeat and horizontal nystagmus with or without torsional component. Patients with orthostatic nystagmus had shorter duration of orthostatic intolerance than those without nystagmus (1.0 ± 1.6 vs. 11.0 ± 9.7 months, $P < 0.001$). In two patients, orthostatic nystagmus disappeared during follow-up despite the persistence of profound orthostatic hypotension.

Conclusions: Generalized cerebral ischaemia caused by orthostatic hypotension induces rotatory vertigo due to objective vestibular dysfunction. The presence of orthostatic vertigo and nystagmus has an association with the duration of orthostatic intolerance.

Triggers – position

Video 1

Video 2

Video 3

Rotational Vertebral Artery Occlusion Mechanisms and Long-term Outcome

Kwang-Dong Choi, MD*; Jae-Hwan Choi, MD*; Ji-Soo Kim, MD; Hyo Jung Kim, MSc; Min-Ji Kim, BSc;
Tae-Hong Lee, MD; Hyung Lee, MD; In Soo Moon, MD; Hui Jong Oh, MD; Jae-Il Kim, MD

Background and Purpose—To elucidate the mechanisms and prognosis of rotational vertebral artery occlusion (RVAO).

Methods—We analyzed clinical and radiological characteristics, patterns of induced nystagmus, and outcome in 21 patients (13 men, aged 29–77 years) with RVAO documented by dynamic cerebral angiography during an 8-year period at 3 University Hospitals in Korea. The follow-up periods ranged from 5 to 91 months (median, 37.5 months). Most patients (n=19; 90.5%) received conservative treatments.

Results—All the patients developed vertigo accompanied by tinnitus (38%), fainting (24%), or blurred vision (19%). Only 12 (57.1%) patients showed the typical pattern of RVAO during dynamic cerebral angiography, a compression of the dominant vertebral artery at the C1-2 level during contralateral head rotation. The induced nystagmus was mostly downbeat with horizontal and torsional components beating toward the compressed vertebral artery side. None of the patients with conservative treatments developed posterior circulation stroke, and 4 of them (21.1%) showed resolution of symptoms during the follow-ups.

Conclusions—RVAO has various patterns of vertebral artery compression, and favorable long-term outcome with conservative treatments. In most patients with RVAO, the symptoms may be ascribed to asymmetrical excitation of the bilateral labyrinth induced by transient ischemia or by disinhibition from inferior cerebellar hypoperfusion. Conservative management might be considered as the first-line treatment of RVAO. (*Stroke*. 2013;44:1817-1824.)

Key Words: mechanism ■ prognosis ■ rotational vertebral artery occlusion ■ stroke

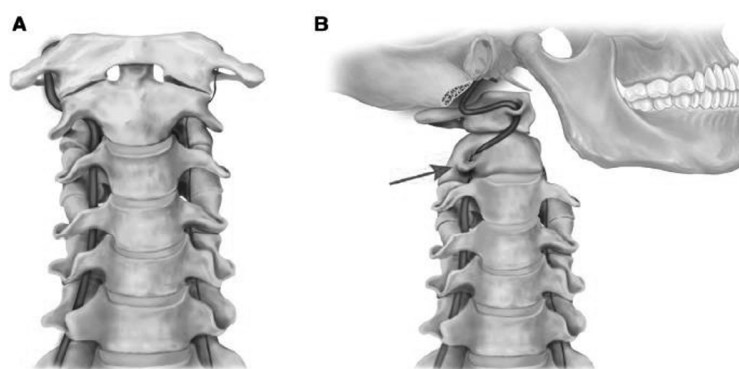


Figure 1. Typical rotational vertebral artery occlusion. A typical pattern of rotational vertebral artery occlusion shows a stenosis or anomaly of the vertebral artery on 1 side (A) and compression of the dominant vertebral artery at the C1-2 level during contraversive head rotation (B).

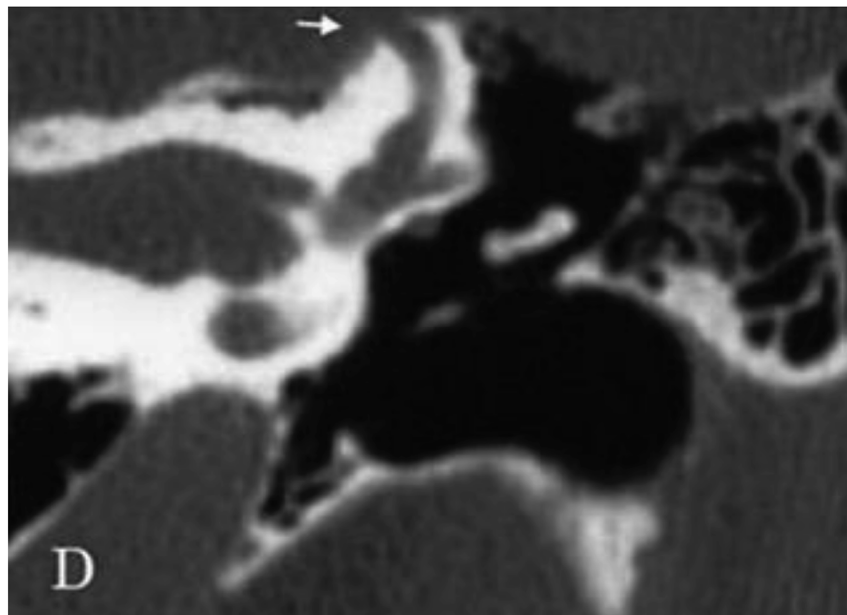
Triggers

Sound

Mastication

PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

PUSAN NATIONAL UNIVERSITY HOSPITAL



PNU 부산대학교병원
PUSAN NATIONAL UNIVERSITY HOSPITAL

Triggers

Sound

Mastication

J Neurol (2014) 261:480–489
DOI 10.1007/s00415-013-7221-7

ORIGINAL COMMUNICATION

PUSAN NATIONAL UNIVERSITY HOSPITAL

Mastication-induced vertigo and nystagmus

Seong-Ho Park · Hyo-Jung Kim · Ji-Soo Kim ·
Ja-Won Koo · Seo Won Oh · Dong-Uk Kim · Joon-Tae Kim ·
Miriam Welgampola · Franca Deriu

Received: 3 September 2013 / Revised: 13 December 2013 / Accepted: 13 December 2013 / Published online: 31 December 2013
© Springer-Verlag Berlin Heidelberg 2013

Abstract Even though trigeminovestibular connections are well established in animals, mastication-induced dizziness has been described only as a vascular steal phenomenon in humans. We determined induction or modulation of nystagmus in two index patients with mastication-induced vertigo, 12 normal controls, and 52 additional patients with peripheral ($n = 38$, 26 with vestibular neuritis/labyrinthitis and 12 with Meniere's disease) or central ($n = 14$, 11 with Wallenberg syndrome, two with cerebellar infarction, and one with pontine infarction) vestibulopathy during their acute or compensated phase. Both index patients developed mastication-induced vertigo after near complete resolution of the spontaneous vertigo from presumed acute unilateral peripheral vestibulopathy. The nystagmus and vertigo gradually built up during mastication and dissipated slowly after cessation of mastication. Brain MRI and cerebral angiography were normal in these patients. Mastication did not induce nystagmus in

normal controls. However, mastication induced nystagmus in five (24 %) of the 21 patients without spontaneous nystagmus (SN) but with a previous history of a vestibular syndrome, and either increased (21/31, 68 %) or decreased (7/31, 23 %) the SN in almost all the patients (28/31, 90 %) with SN. Mastication may induce significant vertigo and nystagmus in patients with a prior history of acute vestibulopathy. The induction or modulation of nystagmus by mastication in both peripheral and central vestibulopathies supports trigeminal modulation of the vestibular system in human. The gradual build-up and dissipation suggest a role of the velocity storage mechanism in the generation of mastication-induced vertigo and nystagmus.

Keywords Nystagmus · Vertigo · Mastication · Velocity storage · Vestibular neuritis · Cerebral infarction

Duration

✓ Days

: **acute vestibular syndrome** (vestibular neuritis, stroke...)

✓ Hours

: **meniere's disease**, vestibular migraine

✓ Minutes

: **VBI**, vestibular migraine

✓ Seconds

: **Cardiogenic vertigo, vestibular paroxysmia**, vestibular epilepsy, psychogenic dizziness

Episodic vestibular syndrome (EVS)

- Triggers
- Duration
- Auditory symptoms
- Headache
- Neurological symptoms
- Family history

- Normal or abnormal

- The most important !!

History taking



Interictal exam.



Ictal exam.

**Exercise-induced downbeat nystagmus in a Korean family with a nonsense mutation in *CACNA1A***Jae-Hwan Choi¹ · Jae-Deuk Seo² · Yu Ri Choi³ · Min-Ji Kim³ · Jin-Hong Shin¹ · Ji Soo Kim⁴ · Kwang-Dong Choi³Received: 23 December 2014 / Accepted: 6 March 2015 / Published online: 18 March 2015
© Springer-Verlag Italia 2015

Abstract Episodic ataxia type 2 (EA2) is characterized by recurrent attacks of vertigo and ataxia lasting hours triggered by emotional stress or exercise. Although interictal horizontal gaze-evoked nystagmus and rebound nystagmus are commonly observed in patients with EA2, the nystagmus has been rarely reported during the vertigo attack. To better describe exercise-induced nystagmus in EA2, four affected members from three generations of a Korean family with EA2 received full neurological and neuro-otological evaluations. Vertigo was provoked in the proband with running for 10 min to record eye movements during the vertigo attack. We performed a polymerase chain reaction-based direct sequence analysis of all coding regions of *CACNA1A* in all participants. The four affected members had a history of exertional vertigo, imbalance, childhood epilepsy, headache, and paresthesia. The provocation induced severe vertigo and imbalance lasting several hours, and oculography documented pure downbeat

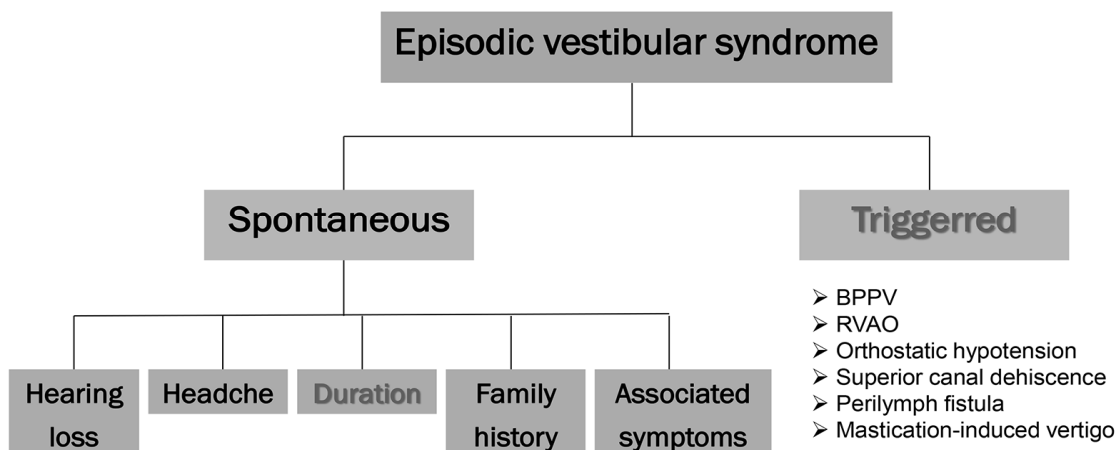
nystagmus during the attack. Genetic analyses identified a nonsense mutation in exon 23 which has been registered in dbSNP as a pathogenic allele (c.3832C>T, p.R1278X) in all the affected members. Ictal downbeat nystagmus in the studied family indicates cerebellar dysfunction during the vertigo attack in EA2. In patients with episodic vertigo and ataxia, the observation of exercise-induced nystagmus would provide a clue for EA2.

Keywords Episodic ataxia · Exercise · Downbeat nystagmus · *CACNA1A* mutation · Calcium channel · Cerebellar dysfunction

Introduction

Episodic ataxia type 2 (EA2) is an autosomal-dominant inherited disorder caused by mutations in *CACNA1A*, which encodes Cav2.1, the main subunit of the P/Q-type

Episodic vestibular syndrome (EVS)



Chronic vestibular syndrome (CVS)



Vestibular Syndromes (Timing)

Chronic vestibular syndrome

- Chronic subjective dizziness
- Cerebellar ataxia
- PSP
- Post-concussion syndrome
- Bilateral vestibulopathy
- Drug-induced dizziness

