

Cerebrovascular Disease and Stroke in Korean Male Adolescent

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Purpose: This study aimed to estimate the prevalence and incidence of cerebrovascular disease (CVD) and stroke in Korean male adolescents. Materials and Methods: The authors reviewed all medical certificates, medical records, and radiologic images from the examinations of Korean military conscription from January 2008 to May 2011. Results: Of the 101156 examinees, 40 had CVD and stroke during adolescence. The overall prevalence and incidence of CVD and stroke was 39.54 cases per 100000 adolescents and 2.08 cases per 100000 adolescents per year, respectively and these were similar to the worldwide data. There were 3 cases of aneurysm, 3 cases of dural arteriovenous fistula, 11 cases of arteriovenous malformation, 4 cases of cavernous hemangioma, 4 cases of cerebrovascular infarction, 16 cases of Moyamoya disease, and 1 case of missing data. The incidence of arteriovenous malformation (0.57 cases per 100000 adolescents per year) was lower than the incidence for the worldwide general population. The incidence of Moyamoya disease was higher than that in any other country (15.82 cases per 100000 adolescents, vs. 0.83 cases per 100000 adolescents per year). Conclusion: We observed ischemic and hemorrhagic stroke, each accounting for approximately half of cases, and high incidence of Moyamoya disease with low incidence of arteriovenous malformation in Korean male adolescents.

Key Words: Cerebrovascular disease, stroke, adolescent, male, Korea, conscription

INTRODUCTION

With a worldwide incidence of 1.3 to 13 per 100000 adolescents per year, cerebrovascular disease (CVD) and stroke has become an important cause of morbidity and mortality in adolescents; it is already one of the top ten causes of death in adolescents.^{1.4} The incidence has increased due to increased recognition, less invasive vascular diagnostics [magnetic resonance images (MRI), magnetic resonance angiography or computed tomography (CT) angiography], and therapeutic advances which allow adolescents with predisposing conditions to survive.

Clinically, CVD and stroke present as either hemorrhagic or ischemic, but clinicians may also find incidentally. Hemorrhagic presentations include intracerebral hemorrhage (the commonest form in children under 10) and subarachnoid hemorrhage (the commonest form in children and adolescents over 10).⁵ These clinical presentations in adolescents are usually caused by arteriovenous malformation

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This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (http://creativecommons.org/ licenses/by-nc/3.0) which permits unrestricted noncommercial use, distribution, and reproduction in any medium, provided the original work is properly cited. (AVM), cerebral cavernous malformation (CCM), cerebral aneurysm, venous sinus thrombosis (VST), or Movamova disease (MMD). Ischemic stroke is commonly arterial ischemic stroke (AIS) or cerebral VST. The underlying cause or trigger may be a diagnostic clue to the distinction between AIS, VST and stroke mimics.⁶⁻¹¹ Adolescent CVD and stroke are strongly associated with death critically, and also with disability.¹² Unfortunately, however, there have been few epidemiologic studies on CVD and stroke in Korean adolescents. Only a few studies have reported some of ischemic stroke's characteristics in Korean young adults compared to the number of western studies.¹³ Herein, we reviewed 101156 examinees (examined for conscription into the Korean military) to estimate prevalence and incidence of the cerebrovascular disease and stroke in the Korean male adolescent population.

MATERIALS AND METHODS

Korea has a conscription system, and all men at age 19 are examined for the purposes of conscription at the Military Manpower Administration. We conducted this cross-sectional survey at the Korean Military Manpower Administration in Seoul, from January 2008 to May 2011, with the approval of the Military Manpower Administration Committee. There were 101156 examinees, aged 19 to 20 years. During this period, 45 men had medical records of CVD and stroke, and 5 cases were excluded from this study because the diagnosis age exceeded 19 years old. We retroactively reviewed all medical records and radiographic images (skull radiographs and CT, MRI, or MR angiographies) of CVD and stroke. Most such cases were rechecked by CT or MRI angiography. And the data were categorized to 6 groups; aneurysm, dural arteriovenous fistula, arteriovenous malformation, cavernous hemangioma, cerebrovascular infarction, and Moyamoya disease. Total and each estimated prevalence and incidence were calculated.

The prevalence of CVD and stroke in a population is defined as the total number of cases in the population (total conscription examinees aged 19 to 20), divided by the number of individuals in the population (total conscription examinees aged 19 to 20). The incidence rate is the number of new cases per population in a given time period, when the denominator is the sum of the person-time of at risk population. It's reasonable to include the mortality and morbidity cases in this study; these cases could not be investigated because the data were derived from the military conscription.

The statistical significances of differences were calculated via Chi-square tests. *p*-values less than 0.05 were regarded as statistically significant. Statistical comparisons were analyzed with SPSS 12.0 (SPSS Inc., Chicago, IL, USA).

RESULTS

In this study, 45 men had medical records of CVD and stroke, and 5 cases were excluded in this study because the diagnosis age exceeded 19 years old (1 cases of aneurysm with 22 years old of diagnostic age, 2 cases of AVM with 20 and 22 years old, 2 cases of cavernous malformation with 20 and 22 year old). Therefore, 40 cases (0.04%) of 101156 conscription examinees in Korea received diagnoses of CVD and stroke under the age of 19. Among them, we could not find any of medical records and image studies in 1 case. Nevertheless, his medical certificate showed that the examinee did have CVD, and we could not categorize the result to any the type of CVD. The overall prevalence and incidence rate of CVD and stroke was 39.54 and 2.08 cases per 100000 adolescent per year, respectively. There were 3 cases of aneurysm, 3 cases of dural arteriovenous fistula, 11 cases of arteriovenous malformation, 4 cases of cavernous hemangioma, 4 cases of cerebrovascular infarction, 16 cases of Moyamoya disease, and 1 case of missing data. Two cases of aneurysm coexisted with dural arteriovenous fistula or Moyamoya disease.

Cerebral aneurysm

This study found 3 cases of cerebral aneurysm, representing a prevalence of 2.97 cases per 100000 adolescents, or an incidence of 0.16 cases per 100000 adolescents per year, in Korean male adolescents. Table 1 shows clinical information on the cerebral aneurysms. There were 2 cases combined with other cerebrovascular diseases, such as dural arteriovenous fistula (DAVF) or MMD. A case with a ruptured pseudoaneurysm presented with mental changes. In 1 case of ruptured pseudoaneurysm, clinicians observed vasospasm in the left distal internal cerebral artery, left proximal middle cerebral artery (M1), and proximal left anterior cerebral artery (A1). We did not observe the multiple aneurysm cases in this study.

Cerebral dural arteriovenous fistula

We also found 3 cases of DAVF, representing a prevalence

Case	Case <u>Age</u> Symptom		Sumptom	Diagnosis (size)	Location	Treatment
Case			Symptom	Diagnosis (size)	Location	meatment
1	19	18	Incidental	Unruptured pseudoaneurysm with DAVF	Right middle meningeal a.	Embolization
2	19	9	Incidental	Unruptured pseudoaneurysm with Moyamoya disease	Left A1 segment	Observation
3	19	17	Mental change	Ruptured pseudoaneurysm	Right distal ICA	Embolization

Table 1. The Clinical Information of 3 Cerebral Aneurysms

Age of Ex, the age with the examination for conscription; Age of Dx, the age with the diagnosis. DAVF, dural arteriovenous fistula; ICA, internal cerebral artery.

Table 2. The Clinical Information of 3 Dural Arteriovenous Fistulas

Case –	Age		Presentation	Diagnosis	Angiographic findings	
	Ex Dx		Fiesemation	Diagnosis		
1	19	5	Exophthalmos	DAVF	Feeder: carotid artery, right drain: orbital vein, right	
2	19	6	Seizure	DAVF	Feeder: posterior cerebral artery, right drain: vein of Galen & transverse sinus	
3	19	18	Raccoon sign	DAVF+pseudoaneurysm	Feeder: middle meningeal artery, right drain: sphenoparietal sinus	

Age of Ex, the age with the examination for conscription; Age of Dx, the age with the diagnosis; DAVF, dural arteriovenous fistula.

of 2.97 cases per 100000 adolescents and an incidence of 0.16 cases per 100000 adolescents per year. Table 2 shows clinical information on the DAVFs. One case presented with right exophthalmos with 6th nerve palsy after penetration of by a wooden chopstick at age 5. In conventional cerebral angiography and CT, its feeder appeared as the right carotid artery, and it drained into the ipsilateral orbit vein. Another case presented with general tonic type seizures at age 6, without any definite trauma history. It had a feeder in the right posterior cerebral artery, and it drained into the vein of Galen and right transverse sinus. The last case presented with right periorbital ecchymosis ("raccoon eye") after a traffic accident. In conventional cerebral angiography, its feeder appeared as the right middle meningeal artery, and it drained into the right sphenoparietal sinus. Incidentally, we found that this case also had a pseudoaneurysm of the middle meningeal artery. All cases were treated by embolization. In the DAVF case with pseudoaneurysm, the DAVF recurred 1 year later after embolization and re-embolization was performed.

Cerebral arteriovenous malformation

We found 11 cases of cerebral arteriovenous malformation, giving a prevalence of 10.87 cases per 100000 adolescents and incidence of 0.57 cases per 100000 adolescents per year. Table 3 shows the AVM clinical information. The median age when the AVM symptoms presented was 16 years (range, 6-19 years, with 1 case of missing data). Among the 11 cases, 5 presented with ruptured AVMs (headache, weakness, seizure, and mental changes), and 6 cases presented

with unruptured AVMs (headache, seizure, syncope, and memory impairment). Additionally, 6 AVM cases were Spetzler-Martin (SM) grade 2, 2 cases were SM grade 1, and 3 cases were not calculable due to insufficient medical information. With regard to treatment, 3 cases received decompression operations, 4 cases underwent operations for nidus removal, 5 cases received radiotherapy, and 3 cases received embolization. Conventional cerebral angiographies were performed in 7 cases (Table 4). However, we could not be certain whether the other 4 cases underwent conventional cerebral angiography. The records showed that 4 AVM cases experienced significant complications: 2 cases had visual disturbances, 1 case had memory impairment, and 1 case experienced speech impediment and motor weakness.

Cerebral cavernous malformation

We found 4 cases of cavernous hemangioma among the conscription, giving a prevalence of 3.95 cases per 100000 adolescents and an incidence of 0.21 cases per 100000 adolescents per year. Table 4 shows clinical information. Presentation symptoms were weakness and seizure. Of these cases, 3 CCMs were located on the subcortical area, and 1 was on the insular. Recurrent bleeding was observed in all these CCMs.

Cerebrovascular infarction

In this study, 4 cases of cerebovascular infarction was found, and the prevalence and incidence rate were 3.95 cases per 100000 adolescents, or an incidence of 0.21 cases per 100000 adolescents per year. Table 5 shows clinical information about the cerebovascular infarction. Presenting symptoms were weakness in 3 cases and disequilibrium in 1 case. The infarcted lesions were randomly distributed as 2 cases of MCA territory, 1 cases of centrum semioval, and 1 cases of cerebellum. All cases were induced by arterial ischemic stroke. Other systemic review was not presented by the medical records.

Moyamoya disease

Finally, we found 16 cases of MMD, giving a prevalence of 15.82 cases per 100000 adolescents and an incidence of 0.83 cases per 100000 adolescents per year. Table 6 shows MMD clinical information. The median age when the symptoms presented was 11 years (range, 8-18 years). Among these 16 cases, 15 (94%) had initial symptoms caused by decreased cerebral perfusion (transient ischemic

Table 3. The Clinica	al Information of 11	Cerebral Arteriovenous	Malformation
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Case	Age		Sumptom	AVM rupturo	SM	Treatment	Angiographic findings	
Case	Ex	Dx	Symptom	AVM rupture	SIVI	Treatment	Angiographic findings	
1	19	6	Mental change	Rupture	2	Decompression Nidus removal	Feeder: left MCA, left PCA Drain: SSS	
2	19	7	Seizure	Rupture	-	Decompression	Unknown	
3	19	10	Mental change	Rupture	2	Nidus removal	Feeder: left MCA Drain: straight sinus	
4	19	11	Headache	Rupture	2	Nidus removal	Feeder: left post choroidal a. Drain: SSS	
5	19	11	Headache	Rupture	-	Decompression Radiotherapy	Unknown	
6	19	16	Memory impairment	No	2	Radiotherapy	Feeder: left MCA Drain: sigmoid sinus	
7	19	17	Syncope	No	1	Radiotherapy	Feeder: left MCA Drain: transverse sinus	
8	19	17	Seizure	No	-	Embolization Radiotherapy	Unknown	
9	19	19	Visual disturbance	No	2	Embolization Radiotherapy	Unknown	
10	20	19	Seizure	No	2	Embolization Nidus removal	Feeder: both ACA, left MCA Drain: SSS, pterygoid plexus	
11	19	-	Headache	No	1	Radiotherapy	Feeder: right PCA Drain: not definite	

Age of Ex, the age with the examination for conscription; Age of Dx, the age with the diagnosis; AVM, arteriovenous malformation; SM, Spetzler-Martin grading system; ACA, anterior cerebral artery; MCA, middle cerebral artery; PCA, posterior cerebral artery; SSS, superior sagittal sinus.

Table 4. The Clinical Information of 4 Cerebral Cavernous Malformation

Case	A	ge	Presentation	Location	Treatment	Recurrent bleeding	
Case	Ex	Dx	Flesentation	Location	meatiment		
1	19	4	Weakness	Right insular	Hematoma removal	Yes	
2	19	13	Seizure	Right frontal lobe	Conservative	Yes	
3	19	15	Seizure	Right occipital lobe	Excision	Yes	
4	19	17	Seizure	Right parietal lobe	Excision	Yes	

Age of Ex, the age with the examination for conscription; Age of Dx, the age with the diagnosis.

Table 5. The Clinical Information of 4 Cerebrovascular Infarction

Case	А	ge	Symptom	Infarcted location	Cause	
Case	Ex Dx		Symptom	inital cied location	Cause	
1	19	13	Disequilibrium	Cerebellum, right	Arterial ischemic stroke	
2	19	15	Left side weakness	Centrum semiovale, left	Arterial ischemic stroke	
3	19	16	Left side weakness	MCA territory, right	Arterial ischemic stroke	
4	19	18	Right side weakness	MCA territory, left	Arterial ischemic stroke	

Age of Ex, the age with the examination for conscription; Age of Dx, the age with the diagnosis; MCA, middle cerebral artery.

attack, dysarthria, and seizure), and 1 case presented with mental changes due to intraventricular hemorrhage from the Moyamoya vessels. In review of image study, the brain parenchymal ischemic encephalomalatic change was observed in 9 cases (60%, among 15 cases confirmed by image study, 1 case with missed image data) (Table 6). All cases had been operated [enecephaloduroarteriosynangiogenesis (EDAS), enecephaloduroarteriomyosynangiogenesis (EDAMS), extra-intracranial arterial bypass (EIAB), or combined operations]. We observed 30 MMD locations among these cases; Suzuki grade 4 was in 9 locations, and Suzuki grades 3 and below were in 21 locations. During the imaging studies of the conscription, clinicians observed postoperative encephalomalatic changes in 9 cases and 11 locations (37%, we could not check 1 case's imaging study from the conscription data). The encephalomalatic changes were found mostly in the frontal lobe. This change did not correlate with the Suzuki grade (*p*-value=0.722). One case was diagnosed with MMD with pseudoaneurysm (Table 6). This 2 mm pseudoaneurysm was in the proximal segment of the left anterior cerebral artery (A1 segment). In this case, the MMD was treated with EDAS and the pseudoaneurysm with observation. The medical records did not indicate whether pseudoaneurysm's disappearance was checked after treatment for MMD.

Table 6. The Clinical Information of 16 Moyamoya Disease

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DISCUSSION

Cerebrovascular disease and stroke have been important causes of morbidity and mortality in adolescents; they are already among the top ten causes of death in adolescents.¹⁴ Hemorrhagic stroke has been reported to be more common than ischemic stroke in children and young adults in the United States,^{3,14,15} however, more recent studies have found different results. Two large hospital discharge studies in the United States reported higher rates of ischemic than hemorrhagic stroke,^{16,17} and a study of a California-wide hospital discharge database found an incidence rate of 1.1 cases per 100000 person-years for hemorrhagic stroke. These rates and proportions are similar to those in the present study.

In this study, the prevalence of CVD and stroke was 39.54 cases per 100000 adolescents, or an incidence of 2.08 cases per 100000 adolescents per year, respectively. The incidence of hemorrhagic stroke was 1.09 cases per 100000 adolescents per year (3 cases of aneurysm, 3 DAVFs, 11 AVMs, and 4 CCMs were included in the hemorrhagic stroke) and the incidence of ischemic stroke was 1.04 cases per 100000 adolescents per year (4 cases of cerebral infarction, and 16 Moyamoya diseases were included in the ischemic stroke).

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Case	Age Ex Dx		- Symptom	Treatment –	Suzuki	grade	Encephalomalatic
Case			Symptom	medunent	Left	Right	changes in brain
1	19	8	TIA	EDAS, both	Grade 4	Grade 2-3	Both frontal
2	19	9	TIA	EDAS, both	Grade 3	Grade 2-3	Unknown
3	19	9	TIA*	EDAS, both	Grade 4	Grade 3	Not observed
4	22	10	TIA	EDAS, right	Grade 2	Grade 2-3	Right frontal
5	23	10	TIA	EIAB and EDAMS, right	Grade 2	Grade 4	Right frontal
6	20	10	TIA	EDAS, both	Grade 2	Grade 2	Not observed
7	19	11	TIA	EDAS, both	Grade 2-3	Grade 4	Right frontal
8	19	11	TIA	EDAMS, right	Grade 1	Grade 4	Right frontal
9	19	11	TIA	EDAS, both	Grade 2-3	Grade 4	Not observed
10	19	13	TIA	EDAS, both	Grade 3	Grade 2-3	Left frontal
11	19	16	TIA	EDAMS, both	Grade 2-3	Grade 4	Both frontal
12	20	16	TIA	EIAB, left	Grade 3	-	Not observed
13	19	17	TIA	EDAS, both	Grade 4	Grade 2-3	Left frontal
14	19	17	TIA	EDAS, right	Grade 3	Grade 4	Right frontal
15	19	17	Mental change [†]	EDAS, right	-	Grade 3	Not observed
16	19	18	TIA	EDAS, both	Grade 2-3	Grade 3	Not observed

Age of Ex, the age with the examination for conscription; Age of Dx, the age with the diagnosis; EDAS, enecephaloduroarteriosynangiogenesis; EDAMS, enecephaloduroarteriomyosynangiogenesis; EIAB, extra-intracranial arterial bypass; TIA, transient ischemic attack.

*A case of Moyamoya disease with pseudoaneurysm.

[†]Symptom was developed by the intraventricular hemorrhage.

Thus, nearly a half of adolescent CVD and stroke was hemorrhagic or ischemic stroke. Regarding hemorrhagic stroke, the incidence worldwide is shown in Table 7. The incidence and proportion of aneurysm, AVM, and CCM are not significantly different to worldwide data.

Cerebral aneurysm

Intracranial aneurysms in adolescents are rare, most studies quoting an incidence of 5% or less.²⁶ In one cooperative study, there was only one child less than 4 years of age with an intracranial aneurysm.²⁷ In the 1000-human cadaver study, just one person under 20 years old harbored an intracranial aneurysm.²⁸ Nevertheless, the actual incidence is not yet clear in the literature review. Generally, females are more common than males, but the study about gender predilection among adolescent is rare.²⁹ In a recent study, simple prevalence and incidence proportions of cerebral aneurysms in adolescents were found to be 2.97 cases per 100000 adolescents per year.

In this study, we found only one case of an actual, truetype aneurysm in a person under 20 years old, and the other cases were combined with other diseases such as DAVF or MMD. Aneurysms with DAVF or MMD may arise because of hemodynamic stress. The combination of aneurysm and DAVF is not well known, but aneurysm with MMD or AVM is not uncommon. In one report, 3 cases among 40 patients (7.5%) had an accompanying aneurysm.³⁰ Aneurysm with MMD tends to occur in the posterior circulation. The most commonly involved location seems to be the basilar bifurcation. However, in this study, we found an aneurysm with MMD in the proximal anterior cerebral artery (A1). Generally, the incidence of multiple aneurysms appears low, and we observed no such cases in this study.^{31,32}

Cerebral dural arteriovenous fistula

The DAVF prevalence is unknown, as some DAVFs remain asymptomatic for years. The estimated incidence is 0.17 per 100000 persons, but this is likely underestimated, because of number of asymptomatic, unreported lesions.³³ We found 3 cases of DAVF in the conscription in Korea. This incidence is similar to that in the general population, but it may not be exact because of underestimation among our examinees. No gender predilection was reported.²⁹

Cerebral arteriovenous malformation

Cerebral AVMs constitute a complex tangle of abnormal arteries and veins. The anatomical absence of a capillary bed in the AVM nidus leads to high-flow arteriovenous shunting through one or more fistulas.³⁴ Generally, the AVM detection rate is 1.11-1.21 per 100000 people per year, and the incidence of AVM hemorrhage is 0.42 per 100000 people per year.^{33,35} In our literature review, we found scant data on AVMs among adolescents. In this study, we found 11 AVM cases (10.87 cases per 100000 adolescents, or 0.57 cases per 100000 adolescents per year) and also found that the incidence in Korean adolescents was lower than the general, worldwide incidence. No gender predilection was reported in the literature.²⁹

In AVMs that present with hemorrhage, the rate for first hemorrhage is 2-4% and the rate for recurrent hemorrhage is 6-18% in the initial year, declining to pre-hemorrhage rate over 5 years.³³ Kondziolka, et al. approximated the chances

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Authors	Country	Age	Incidence	Case	Proportion (%)		
Autions	Country	range	(case/100000/yrs)	number	Aneurysm	AVM	CCM
Schoenberg, et al. ¹⁴	USA	0-14 y	1.89	31	19	42	-
Eeg-Olofsson and Ringheim ¹⁸	Sweden	6 m-15 y	2.1	14	29	57	-
Broderick, et al. ¹⁵	USA	0-14 y	1.5	9	11	33	22
Visudhiphan, et al. ¹⁹	Thailand	6 m-15 y	-	30	17	40	3
Giroud, et al. ²⁰	France	1 m-15 y	5.11	11	9	46	27
Lin, et al. ²¹	Taiwan	0-15 y	-	42	0	45	5
Lanthier ²²	Canada	1 m-18 y	-	21	5	38	19
Blom, et al. ²³	Netherlands	0-15 y	-	56	9	41	0
Meyer-Heim and Boltshauser ²⁴	Switzerland	0-18 y	-	34	15	53	6
Liu, et al. ²⁵	Great Britain	1 m-15 y	-	50	8	26	18
Oh, et al. (current study)	Korea	19 y	1.09	21*	14	52	19

 Table 7. Comparison of Incidence and Proportion of Hemorrhagic Stroke, Aneurysm, Arteriovenous Malformation and Cerebral Cavernous Malformation, among Reported Studies and Recent Study

m, month old; y, years old; AVM, arteriovenous malformation; CCM, cerebral cavernous malformation.

*Study for only male adolescents.

of an AVM hemorrhage according to age by an equation: Risk of hemorrhage=1 - (risk of no hemorrhage)^{expected years of remaining life .36} In a group of 19-year-olds the approximate 19-yearolds' risk of hemorrhage is 64%, if the expected years of remaining life is 60, and the risk of hemorrhage is 1.7% per year. In this study, we found 11 AVM cases, of which 5 cases had ruptured AVM (45%), ant that all well-formed AVMs were SM grades 1 or 2. The recommended general treatment strategy for SM grades 1 and 2 AVMs is surgery or focused irradiation. In this review, 6 among the 8 cases of known SM grade 1 or 2 received surgical treatment and/or radiosurgery, and 2 cases received surgical treatment or radiosurgery after embolization.

Cerebral cavernous malformation

The exact incidence and prevalence of CCMs are unknown, as many CCMs are asymptomatic. A populationbased study showed an incidence of 0.15 per 100000 persons per year.³³ In some autopsy and MRI studies, the prevalence was 0.4-0.9%.^{37,39} No gender predilection was reported.²⁹ In this study, we found 4 cases of CCM, giving a prevalence of 3.95 cases per 100000 adolescents and an incidence of 0.21 cases per 100000 adolescents per year. Generally, evidence of previous hemorrhage is present at every lesion, regardless of clinical history. In addition, CCMs have persistent intralesional microhemorrhages, which occur over time. We found such previous hemorrhages in all CCM cases in this study.

Cerebrovascular infarction

Adolescent ischemic stroke is not as rare as one might surmise. Incidence estimates vary, depending on definition and inclusion criteria. AIS ranges from 0.6 to 7.9 per 100000 adolescents under age 18 per year in North America and Europe.^{3,4} Arterial ischemic stroke is defined as an acute focal neurological deficit lasting longer than 24 hours with neuroimaging evidence of cerebral infarction. There is no gender predilection in cerebrovascular infarction, but AIS is more common in male than female.²⁹ Arterial ischemic stroke is a complex disease with many risk factors. The most frequently reported risk factors for AIS in adolescent are cardiac disorders, hematological disorders, metabolic disorders, arteriopathies, and infection.^{40,41} Unfortunately, however, systemic review of risk factors was not investigated in this study. Total 4 cases of cerebovascular infarction by AIS were found in this study, and the prevalence and incidence rate was 3.95 cases per 100000 adolescents, or an incidence of 0.21 cases per 100000 adolescents per year, respectively. The incidence is lower than that in any other reports, but similar to that including Moyamoya disease (other reports included Moyamoya disease to AIS as arteriopathies).

Moyamoya disease

Moyamoya, meaning a "hazy puff of smoke" in Japanese, is a chronic, occlusive cerebrovascular disease involving bilateral stenosis or occlusion of the terminal portion of the internal carotid arteries and/or the proximal portions of the anterior cerebral arteries and middle cerebral arteries.⁴² The MMD incidence is high in East Asian countries, such as Japan and Korea (Table 8). In Japan, the estimated annual prevalence and incidence are 6.03 and 0.54 per 100000 persons, respectively.⁴³ MMD's clinical background in Korea is essentially similar to that in Japan.⁴⁴ In our series, we

Table 8. Epidemiologic Studies or	Case Series	Involving Mov	amova Disease acc	ording to High Prevalence

A (1	Derived a Cart of (are)	Diana Catal	Prevalence	Prevalence Incidence (per 100000 persons)	
Authors	Period of study (yrs)	Place of study	(per 100000		
Oh, et al. (current study)	2008-2011	Korea (Seoul)	15.82	0.83	t
Baba, et al.47	2002-2006	Japan (Hokkaido)	10.5	0.94	2.18
Kuriyama, et al.43	2003	Japan (nationwide)	6.03	0.54	1.8
Wakai, et al.48	1994	Japan (nationwide)	3.16	0.35	1.8
Su, et al. ⁴⁹	1978-1993	Taiwan (Hualien)	-	0.024	1.21
Ikezaki, et al.44	-1995	Korea (nationwide)	451 cases		1.3
Choi, et al.50	1978-1993	Korea (Seoul)	72 cases duri	ing 16 yrs*	1.81
Marcinkevicius, et al.51	1995-2005	Lithuania (Kaunas)	14 cases duri	ing 11 yrs*	0.75
Kraemer, et al.52	1996-2007	Germany (Essen)	21 cases duri	ing 12 yrs*	4.25
Shoukat, et al.53	1988-2006	Pakistan (Karachi)	13 cases duri	ing 19 yrs*	0.86

*Hospital-based study.

[†]Study for only male adolescents.

found 16 MMD cases among the young conscripted men in Korea (15.85 cases per 100000 adolescents, 0.83 cases per 100000 adolescents per year). Our literature review revealed that this is the highest prevalence and second-highest incidence, next to the report by Baba, et al. In Korea and Japan, the number of women with MMD is higher than that of men.⁴⁷ Because we based our present study on the examination of conscripts, we did not include female adolescents. Therefore, if this epidemiological study extends to all male and female adolescents, the prevalence and incidence proportions would be higher than those shown in this study.

In this study, mean age at diagnosis was 11.5 years (range, 8-18 years), and 15 (93.5%) of 13 MMD cases presented with decreased perfusion; 1 case presented with hemorrhage by MMD. Clinically, young patients with MMD present with ischemic attacks, and adult patients present with either ischemic or hemorrhagic events.⁴⁵ Although this point has recently been debated. Generally, most patients present with bilateral involvement; up to 18% of patients have angiographically documented unilateral involvement.⁴⁶ In this study, we observed unilateral involvement in 2 (12.5%) of 16 cases. In addition, 1 case had a pseudoaneurysm, located in the proximal anterior cerebral artery (A1).

Limitation of this study

This study was based on the data of Korean military conscription. Therefore, a few limitations in the epidemiologic study were present. First, pass away or severely disabled examinees were excluded from the conscription, so mortality or morbidity cases were not included in this study. Nevertheless, according to the data of Seoul regional Military Manpower Administration, less than 0.01% of cases were excluded from the conscription by severe disability, and these morbidity cases do not seem to affect the result. Second, only male adolescent was included in this study. Thus, overall prevalence and incidence calculated could underappreciate or not indicate the general population rate by different gender factors. The authors used reference data to supplement this problem, however, it was not enough. The general population study would be needed to this problem. Finally, by using medical certificate and record, not all conscription cases were checked for image study. The general symptom of CVD and stroke is headache and other popular symptom, therefore, undiagnosed incidental CVD and stroke cases could be present. However, this limitation was applicable not only to this study, but also to all epidemiologic study. To solve this limitation, well designed large population study would be needed; also it is very difficult.

Regardless of limitations, this study is a large cross-sectional survey with 101156 examinees with localized geographical distribution, same ethics, and specific 19 years old aged males. Consequently, this study is very useful for estimating the prevalence rates and types of CVD in Korean male adolescents.

In conclusion, in Korean male adolescents, the incidence of cerebrovascular disease and stroke (2.08 cases per 100000 adolescents per year) and the proportions of hemorrhagic and ischemic stroke are similar to those worldwide. However, the incidence of arteriovenous malformation (0.57 cases per 100000 adolescents per year) was lower than the incidence of worldwide, general population. The incidence of Moyamoya disease was higher than that in any other country (for a prevalence of 15.82 cases per 100000 adolescents and an incidence of 0.83 cases per 100000 adolescents per year).

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