#### Iran J Public Health, Vol. 48, No.11, Nov 2019, pp.1997-2006



# **Original Article**

# Correlation between Metabolic Syndrome and Intracranial versus Extracranial Arteriosclerosis among Chinese Patients with Stroke

### \*Chunyu LIU, Xiaodan YANG, Changqing CHEN

Department of Neurology, PLA 254 Hospital, Tianjin 300141, P.R. China

\*Corresponding Author: Email: c1239837zongla@163.com

(Received 09 Oct 2018; accepted 11 Dec 2018)

#### Abstract

**Background:** We aimed to explore the correlation between metabolic syndrome and intracranial and extracranial arteriosclerosis. Overall, 318 over 60-yr-old patients with cerebral infarction or TIA who were examined by digital subtraction angiography (DSA) in our hospital were enrolled in the study.

**Methods:** Overall, 192 patients with intracranial and extracranial arteriosclerosis were admitted to the case group (the intracranial and extracranial arteriosclerosis group). Also, 196 patients, suffering from the same condition, were selected from our outpatient clinic and enrolled in the control group.

**Results:** The prevalence of metabolic syndrome was 31.4%. The prevalence of each metabolic syndrome component in the intracranial arteriosclerosis group was higher than those of the extracranial arteriosclerosis and the control groups. The average component values in the intracranial arteriosclerosis group was higher than those observed in other groups. The prevalence rate of metabolic syndrome had no significant difference among different degrees of stenosis for extracranial arteriosclerosis group. There was a remarkable correlation between intracranial arteriosclerosis and metabolic syndrome (P<0.001), while no correlation was detected between extracranial arteriosclerosis and metabolic syndrome (P<0.001). We concluded that metabolic syndrome may increase the prevalence risk of intracranial arteriosclerosis. There was a significant correlation between intracranial arteriosclerosis and metabolic syndrome components including hyperglycemia and hypertension. Also, there was a significant correlation between extracranial arteriosclerosis and metabolic syndrome components including hyperglycemia.

**Conclusion:** We believe that at least three components of metabolic syndrome can obviously increase the risk of intracranial arteriosclerosis.

Keywords: Metabolic syndrome; Intracranial arteriosclerosis; Extracranial arteriosclerosis

#### Introduction

Acute cerebrovascular disease is one of the important causes of mortality and disability all over the world. There are 7 million known cases of acute cerebrovascular in China, with annul prevalence rate of 260-719 for 10 million people. Every year, 1.65 million patients die from acute cerebrovascular disease (1). The disability caused by

cerebrovascular disease is around 75%. At present, the incidence rate of acute cerebrovascular disease is increasing by 8.7% every year. Acute cerebrovascular disease featuring high incidence rate, death rate and disability rate brings a burden to society and families. 87% of patients with acute cerebrovascular disease suffer from ischemic stroke (2). Intracranial and extracranial arteriosclerosis (AS) is an important pathological basis of ischemic stroke (3). Back in 1990s, several studies were conducted on the distribution of intracranial and extracranial arteriosclerosis for different races, and results showed that the distribution of intracranial and extracranial arteriosclerosis had a racial difference. Asian arteriosclerosis lesion always happens to intracranial artery, which is the main reason of ischemic stroke, while the cases in west are just the opposite (4-8). Due to the difference in risk factors and mechanism that can induce intracranial and extracranial arteriosclerosis (9-11), similar treatment methods may have dissimilar effects on the changes to intracranial and extracranial arteriosclerosis.

Metabolic syndrome describes a phenomenon that multiple pathogenic factors aggregate in an individual. As a group of metabolic disorder syndrome, it consists of obesity, hypertension, pathoglycemia and dyslipidemia. In recent years, with the aging population in China as well as the lifestyle of surfeit and sedentariness, the prevalence rate of metabolic syndrome is gradually increasing. The results of the latest nationwide metabolic syndrome research baseline (Inter Asia, 2010-2012) showed that the prevalence rate of metabolic syndrome is 11.0 % (12). All components of metabolic syndrome can increase the risk of arteriosclerosis, at the same time, the risk of arteriosclerosis occurrence is higher when many abnormities combine. There is a difference between intracranial and extracranial arteriosclerosis and metabolic syndrome, as well as the correlation among difference components (13, 14). However, the relevant reports on this difference for Chinese population are very rare.

We aimed to discuss the correlation between intracranial and extracranial arteriosclerosis and metabolic syndrome as well as its relevant components respectively. We conducted crosssectional study on the patients with intracranial and extracranial arteriosclerosis, in order to provide objective basis for prevention and treatment of ischemic cerebrovascular disease in future.

### Materials and Methods

#### Source of Crowd

From September. 2015 to September 2017, 318 over 60-yr-old patients with cerebral infarction or TIA were examined by digital subtraction angiography (DSA) in our hospital. According to the inspection result of DSA that distinguished intracranial arteriosclerosis and extracranial arteriosclerosis, 192 patients with intracranial and extracranial arteriosclerosis were enrolled in this study as the case group (intracranial and extracranial arteriosclerosis group). This included 105 patients with intracranial arteriosclerosis and 87 patients with extracranial arteriosclerosis. Overall, 196 patients, suffering from the same condition, were selected from our outpatient clinic and enrolled in the control group. Clinical general information and blood vessel image data were collected from all patients. Blood samples were used for metabolic syndrome predication.

### Criterion of Intracranial and Extracranial Arteriosclerosis

The stenosis of intracranial and extracranial arterial lumen  $\geq$  30% was deemed to be arteriosclerosis lesion. The calculation of arterial stenosis rate referred to the measurement standard of North American Symptomatic Carotid Endarterectomy Trial (NASCET) (15): arterial stenosis rate= [1 - (inside diameter of blood vessel at the most narrow / inside diameter of normal blood vessel at the far-end stenosis] ×100%.

#### Exclusion Criteria

1.Cerebrovascular malformation, aortic dissection, vasculitis;

2.Potential cardiac embolism;

3.Cerebral infarction with unknown cause;

4.Degree of artery stenosis less than 30%, combined with intracranial and extracranial arteriosclerosis;

5. Tumor, infection and hypercoagulation;

6.Severe visual and auditory dysfunction and aphasia (RVR<6 points)

7.Incomplete medical history;

8.Refusal for informed consent signature.

### Diagnostic Criteria

1. Diagnostic Criteria of Metabolic Syndrome IDF standard (16) was applied to metabolic syndrome diagnosis which met these 3 items: 1) Central obesity: the waistline of male  $\geq$ 90cm, the waistline of female  $\geq$ 80cm; 2) Triglyceride (TG) level increasement: > 1.7 mmol/ L or acceptance of special treatment on the lipid abnormity; 3) High density lipoprotein cholesterol (HDL-C) level reduction: male < 1.03 mmol/ L, female < 1.29 mmol/L or acceptance of special treatment on the lipid abnormity; 4) Elevated blood pressure: systolic pressure  $\geq$ 130 mmHg or diastolic pressure  $\geq$ 85 mmHg, or acceptance of treatment due to the previous diagnosis as high blood pressure; 5) Fasting blood-glucose increasement: Fasting bloodglucose≥100 mg/ dl (5. 6 mmol/ L) or diagnosis as type 2 diabetes mellitus.

2. Hypertension: with a chronic hypertension history, systolic pressure  $\geq$ 140 mmHg, diastolic pressure  $\geq$ 90 mmHg; or only systolic pressure  $\geq$ 140 mmHg after admission to hospital.

3. Glycuresis: with a diabetes history, fasting blood-glucose  $\geq$ 7.0 mmol/L, random blood glucose  $\geq$ 11.1 mmol/L after admission to hospital.

4. History of Coronary Heart Disease: with an atrial fibrillation and ischemic heart disease history, or prompt of atrial fibrillation and myocardial ischemia by electrocardiography after admission to hospital to make a definite diagnosis.

5. Smoking: smoke  $\geq 10$  cigarettes, for  $\geq 5$  years, and still smoke now.

6. Hyperhomocysteinemia: homocysteine ≥15umol/l.

### Main Examination Methods

1. Measurement of Abdominal Girth and Blood Pressure:

When patients stood still and kept eupnea, the diameter from 1 cm above navel was measured horizontally; then patients took a sitting position twice for blood pressure measurements by cuff type mercurial sphygmomanometer.

2. Blood Extraction and Laboratory Examination Methods:

Two samples of 5 ml fast venous blood were extracted from all patients within 24 hours after admission to hospital, and two samples of 5 ml fast venous blood were extracted during medical examination. Oxidase method was used to measure TG/CHO; for the measurement of HDL, precipitator was applied for precipitation and cholesterol levels were measured in supernatant using enzymic method. LDL levels were measured using Friedwald formula (LDL=CHO-HDL-TG/5) and glucose oxidase method was used to measure fasting blood glucose.

3. DSA Examination: AdvantxLCA+/LCV+/LC+ angiography machine from America GE company was used and voltage was set as 80 KV, magnitude of current was 500mA and F1 was 75 KV. Seldinger method was adopted to femoral artery puncture tube, with contrast medium of iohexo 300 mgI/ml, to inject by high pressure injector. The condition of common carotid artery arteriography was: flow velocity 5 ml/s, flow volume 7 ml and pressure at 150 Psi.

#### Statistical Process

SPSS 18.0 (Chicago, IL, USA) was applied for statistical analysis. Baseline population statistics and the continuous variable of laboratory data was expressed by Mean±standard deviation, and absolute variable was showed by frequency. The comparison of continuous variable among groups was detected by *t* or analyzed by variance, while the comparison of absolute variable was tested by Chi-square. Multi-factor Logistic analysis of regression was used for the research on correlation between intracranial and extracranial arteriosclerosis and metabolic syndrome as well as its components. The results were signified by OR value and 95% confidential intervals. P < 0.05 meant a statistical significance.

### Results

### Basic Information of Research Population

Among 388 patients, there were 250 males and 138 females with average age of  $65.8\pm9.3$  years old. The comparison of the number the male patients between the intracranial and extracranial arteriosclerosis and the control groups showed

no significant difference. There was also no significant difference between the intracranial arteriosclerosis group and the extracranial arteriosclerosis group in terms of the number the male patients. In the intracranial and extracranial arteriosclerosis group, age, smoking, BMI, hypertension, diabetes and prevalence rate of metabolic syndrome were obviously higher than those of the control group (P < 0.05, P < 0.01). There was no significant difference in age, smoking pattern, BMI, hypertension, diabetes and prevalence rate of metabolic syndrome between the intracranial arteriosclerosis group and the extracranial arteriosclerosis group. The prevalence rate of metabolic syndrome was 31.4%. The prevalence rate of each metabolic syndrome component in the intracranial arteriosclerosis group was higher than those of other groups. The average component number of metabolic syndrome in the intracranial arteriosclerosis group was higher than those of the extracranial arteriosclerosis and the control groups (Table 1).

Table 1: Baseline Features of Research Population with Intr	racranial and Extracranial Arteriosclerosis
---	---

Variable	Control Group n=196 (%)	Intracranial and Ex- tracranial Arterioscle- rosis n=192 (%)	Intracranial Arte- riosclerosis n=105 (%)	Extracranial Arte- riosclerosis n=87 (%)	P Value	
Age (vr)	$62.67 \pm 10.6$	67.9±9.5	$67.7 \pm 8.8$	$68.2 \pm 10.4$	0.009	
Gender, Male	119 (60.7)	131(68.2)	73 (69.5)	58 (66.7)	0.376	
Smoking	41(20.9)	65(33.9)	33 (31.6)	32 (36.8)	0.012	
Coronary Heart Disease	4 (2.0)	18 (9.4)	5 (4.8)	13 (14.9)	< 0.001	
Hypertension	17 (8.7)	117(60.9)	70 (66.7)	47 (54.0)	< 0.001	
Diabetes	13 (6.6)	97(50.5)	57 (54.3)	40 (46.0)	< 0.001	
Metabolic Syndrome	28 (14.3)	94(49.0)	56 (53.3)	38 (43.7)	< 0.001	
Components of Metabolic Syndrome				~ /		
Hypertension	23 (11.7)	136(70.8)	80 (76.2)	56 (64.4)	< 0.001	
Hyperglycemia	48 (24.5)	117(60.9)	68 (64.8)	49 (56.3)	< 0.001	
Hypertriglyceridemia	50 (25.5)	74(38.5)	45 (42.9)	29 (33.3)	0.008	
Low High- density Lipoprotein	35 (17.9)	97(50.5)	59 (56.2)	38 (43.7)	< 0.001	
Abdominal Obesity	75 (38.3)	119(62.0)	75 (71.4)	44 (50.6)	< 0.001	
Metabolic Syndrome (Number of	$1.66 \pm 1.32$	2.31±1.56	$2.55 \pm 1.44$	$2.11 \pm 1.52$	< 0.001	
Component)						
1	59 (30.1)	33(170.2)	14 (13.3)	19 (21.8)		
2	50 (25.5)	38(19.8)	24 (22.9)	14 (16.1)		
3	25 (12.8)	48(25.0)	28 (26.7)	20 (23.0)		
4	16 (8.2)	31(16.1)	18 (17.1)	13 (14.9)		
5	6 (3.1)	15(7.8)	10 (9.5)	5 (5.7)		

#### Different Metabolic Syndrome Distribution of Different Degrees of Intracranial and Extracranial Arteriosclerosis Stenosis

In intracranial Arteriosclerosis group, there were 26 patients (52.6%) with metabolic syndrome among patients with degrees of stenosis  $\geq$ 70%. There were 30 patients (44.8%) with metabolic syndrome among patients with degrees of stenosis between 30% and 69%. The prevalence rate of metabolic syndrome had a significant difference among different degrees of stenosis (P < 0.05). Number of patients with degrees of steno-

sis  $\geq$ 70% suffered from hypertension, hyperglycemia, hypertriglyceridemia, low high-density lipoprotein and abdominal obesity were 34, 30, 19, 23 and 28 respectively. Number of patients with degrees of stenosis between 30% and 69% suffered from hypertension, hyperglycemia, hypertriglyceridemia, low high-density lipoprotein and abdominal obesity were 46, 38, 26, 36 and 47 respectively. Finally, number of patients with degrees of stenosis  $\geq$ 70% suffered from hypertension and hyperglycemia was obviously more than that of patients with degrees of stenosis between 30% and 69%, and the number of patients with degrees of stenosis  $\geq$ 70% suffered from 3, 4 and 5 components of metabolic syndrome were apparently higher than those of patients with degrees of ste-

nosis between 30% and 69% (P < 0.05).

Extracranial Arteriosclerosis Group: The prevalence rate of metabolic syndrome had no significant difference among different degrees of stenosis and the number of patients with degrees of stenosis  $\geq$ 70% suffered from hypertension, hyperglycemia, hypertriglyceridemia, low highdensity lipoprotein cholesterol and abdominal obesity were 12, 15, 7, 8 and 9 respectively. Number of patients with degrees of stenosis between 30% and 69% suffered from hypertension, hyperglycemia, hypertriglyceridemia, low highdensity lipoprotein cholesterol and abdominal obesity were 44, 34, 22, 30 and 35 respectively (Table 2).

 Table 2: Metabolic Syndrome Distribution of Different Degrees of Intracranial and Extracranial Arteriosclerosis

 Stenosis

Variable		Arteriosclerosis 05 (%)	Extracranial Arteriosclerosis n=87 (%) Degrees of Stenosis		
	Degrees	of Stenosis			
	30%-69%	≥70%	30%-69% n= 68	≥70%	
	n= 67	n= 38		n= 19	
Metabolic Syndrome	30 (44.8)	26 (52.6)	30 (44.1)	8 (42.1)	
Components of Metabolic Syndrome	. ,			. ,	
Hypertension	46 (68.7)	34 (89.5)	44 (63.8)	12 (63.2)	
Hyperglycemia	38 (56.7)	30 (78.9)	34 (50.0)	15 (78.9)	
Hypertriglyceridemia	26 (38.8)	19 (50.0)	22 (32.4)	7 (36.8)	
Low High- density Lipoprotein	36 (53.7)	23 (60.5)	30 (44.1)	8 (42.1)	
Abdominal Obesity	47 (70.1)	28 (73.7)	35 (51.5)	9 (47.4)	
Metabolic Syndrome (Number of Component)	· · · ·				
1	10 (14.9)	4 (10.5)	15 (22.1)	4 (21.1)	
2	16 (23.9)	8 (21.1)	9 (13.2)	5 (26.3)	
3	13 (19.4)	15 (39.5) 🔺	14 (20.6)	6 (31.6)	
4	7 (10.4)	11 (28.9)	8 (11.8)	5 (26.3)	
5	3 (4.5)	7 (18.4)	3 (4.4)	2 (10.5)	

 $\blacktriangle$  *P* < 0.05, compared with degree of 30%-69%

#### Correlation Analysis of Intracranial and Extracranial Arteriosclerosis and Metabolic Syndrome as well as Related Risk Factors

Intracranial Arteriosclerosis Group: After adjusting the factors including gender, smoking and coronary heart disease, a remarkable correlation between intracranial arteriosclerosis and metabolic syndrome (P<0.001) was detected. There was a correlation between intracranial arteriosclerosis and some components of metabolic syndrome such as hypertension, hyperglycemia and low high- density lipoprotein. Additionally, age and hyperhomocysteinemia were significantly related to intracranial arteriosclerosis (OR, 1.96 P= 0.01; OR, 1.89 P=0.007).

Extracranial Arteriosclerosis Group: After adjusting the factor of gender, no correlation between extracranial arteriosclerosis and metabolic syndrome was detected. There was a correlation between intracranial arteriosclerosis and hyperglycemia among the components of metabolic syndrome. Additionally, age, smoking habits and hyperhomocysteinemia were significantly related to intracranial arteriosclerosis as well. There was no correlation between extracranial arteriosclerosis and the number of components of metabolic syndrome (Table 3).

	OR	95% IC				
$\langle \rangle$	1.07	20/010	Р	OR	95% IC	Р
ge(yr)	1.96	1.2-3.99	0.01	2.01	1.23-2.97	0.013
ender	1.46	0.85-2.50	0.172	1.28	0.73-2.24	0.395
noking	1.13	0.94-2.12	0.056	1.68	1.24-3.00	0.005
oronary Heart Disease	1.63	0.54-5.00	0.184	1.73	1.16-4.79	0.001
yperhomocysteinemia	1.89	1.23-2.56	0.007	1.93	1.34-3.65	0.006
letabolic Syndrome	3.22	2.03- 6.12	< 0.001	1.42	0.91-2.97	0.072
omponents of Metabolic Syndrome						
ypertension	2.05	1.78-5.13	0.012	1.57	0.94-3.23	0.065
vperglycemia	2.54	1.83-5.27	0.006	1.89	1.35-4.13	0.035
ypertriglyceridemia	1.42	0.89-2.17	0.092	1.01	0.56-1.67	0.910
ow High-Density Lipoprotein	1.57	1.19-4.10	0.012	1.47	0.84-2.63	0.091
bdominal Obesity	1.28	0.76-2.67	0.09	1.56	0.89-2.74	0.054
letabolic Syndrome (Number of						
omponent)						
	0.65	0.19-1.23	0.241	0.53	0.36-1.17	0.142
)	0.77	0.42-1.46	0.511	0.41	0.26-1.06	0.151
)	2.15	1.17-4.35	0.007	1.94	0.93- 3.83	0.062
)	2.27	1.03- 4.54	0.022	1.83	0.84-4.21	0.091
,	3.12	1.15445	0.023	1.79	0.54-5.45	0.288

 

 Table 3: Logistic Analysis of Intracranial and Extracranial Arteriosclerosis and Metabolic Syndrome as well as Related Risk Factors

a, control group as reference group; b, 0 component of metabolic syndrome as reference group

### Discussion

Among 142 cases with recent cerebral infarction, 55% had a corresponding intracranial arteriosclerosis lesion (17). Extracranial arteriosclerosis was common for the western population, while the Asian, Blacks and Hispanic always suffered from intracranial arteriosclerosis (18-22). By continuously observing 318 patients with cerebral infarction and TIA, we discovered that the occurrence rate of intracranial and extracranial arteriosclerosis was 60.4%, among which intracranial arteriosclerosis accounted for 33.1% and extracranial arteriosclerosis accounted for 27.4%. Compared with existing results, the incidence rate of intracranial and extracranial arteriosclerosis was somewhat inconsistent. Though the lesion of intracranial arteriosclerosis is more than that of extracranial arteriosclerosis, the incidence rate of extracranial arteriosclerosis is on the rise, and the reason may be explained by the fact that in recent years we observed a significant improvement in life standard and the lifestyle (23).

We, showed that the comparison of gender between the intracranial and extracranial arteriosclerosis group and the control group had no statistical difference. The intracranial arteriosclerosis group and the extracranial arteriosclerosis group results were not in line with previous research results (24), because of the selected population. Patients enrolled in this study were over 60 years old (average ages were 67.9 years for male patients and 66.8 years for female patients). Estrogen in female patients could inhibit the arteriosclerotic progress. A lack of estrogen due to surgery or other reasons is the most important element leading to arteriosclerosis for reproductiveaged females. Before 65 years old, the incidence rate of arteriosclerosis for females are obviously lower than males, however, after 65 years old, the incidence rate of arteriosclerosis for females and males are equal. Once stroke occurs in female patients, the results and prognosis are usually worse than those of the male patients belonging to the same age group (25). Here we showed that after losing the protection of estrogen, the risk of

arteriosclerosis lesion for females is equivalent to males. Previous studies showed that smoking is an independent predictive factor for intracranial and extracranial arteriosclerosis (26), which can lead to arteriosclerosis by inducing vascular inflammation and oxidative stress (36). Meanwhile, vasomotor dysfunction, coagulation and fibrinolysis disorders and lipid deformation may appear, which can indirectly aggravate angiopathy (27). Results of the present study showed that smoking did not have any significant correlation with intracranial arteriosclerosis, instead, there was a significant correlation between it and extracranial arteriosclerosis, which was consistent with those results obtained from previous studies (27, 28). For the patients with ischemic heart disease, it was very common to suffer from intracranial and extracranial arteriosclerosis, and a significant correlation was detected between the occurrence rate of extracranial arteriosclerosis and the severity of coronary arteriosclerosis (29). Coronary heart disease was not an independent risk factor of intracranial arteriosclerosis, instead, it was related to extracranial arteriosclerosis. The results were not only consistent with the results obtained in the studies conducted in Japan and Korean (30, 31). These results may explain the consistency of the correlation between extracranial arteriosclerosis and coronary heart disease exists among Asian population.

Among Asian and white patients, the relation between metabolic syndrome and intracranial arteriosclerosis is closer than extracranial arteriosclerosis (13, 14, 32, 33). We did not detect any correlation between different degrees of extracranial arteriosclerosis stenosis and metabolic syndrome. The reason why intracranial arteriosclerosis showed a significant correlation with metabolic syndrome might be as follows: 1. Under normal conditions, antioxidant enzyme contained in intracranial artery is superior to that of extracranial artery. Antioxidant enzyme in intracranial artery can decrease quickly with the age, which is more obvious compared with extracranial artery (34). It means that intracranial arteriosclerosis can speed up with age. This can explains the fact that age is an independent risk factor in intracranial arteriosclerosis in our study consistent with another study (35); 2. The main pathophysiology basics of metabolic syndrome is insulin resistance (IR) that can damage intracranial arteriectasia function (36) and lead to an increase in vascular wall stiffness and reduction of buffer ability. This increases the susceptibility of artery oxidative stress (37), and can speed up the formation of intracranial arteriosclerosis.

Hypertension is a clear risk factor that accelerates arteriosclerosis. The increase in systolic pressure and pulse pressure may increase the vascular shear stress and blood flow fluctuation. Therefore, the endotheliocyte can secrete NO, PGI, ET, leading to creation of an unbalance in angiokinesis material and exacerbation of endothelial cell damage. Compared with extracranial artery, the thickness and elasticity of tunica media in intracranial artery is the worse, so it can be more easily influenced by the vascular stress and blood flow caused by hypertension (38). Results obtained from other studies confirmed that the relationship between intracranial arteriosclerosis and hypertension is much closer, compared with extracranial arteriosclerosis (28, 39). The chief reason for diabetic attack is insulin resistance, and it has been verified that insulin resistance facilitate the incidence of arteriosclerosis. In the north of Manhattan and in Spain, diabetes was closely associated with intracranial arteriosclerosis (40, 41). Also, other studies (4, 24) on Asian population produced similar results. Hyperglycemia is an independent risk factor of extracranial arteriosclerosis, but no correlation between hypertension and extracranial arteriosclerosis was detected. This, suggested that pathoglycemia was a more important risk factor for extracranial arteriosclerosis compared with dysarteriotony.

High triglyceride and low high-density lipoprotein levels can promote the formation of lowdensity lipoprotein particles. Hypercholesteremia is a risk factor of intracranial arteriosclerosis (24, 42). In our study, high triglyceride levels did not show any correlation with intracranial arteriosclerosis, but low levels of high-density lipoprotein had an obvious correlation with intracranial arteriosclerosis. It has been shown that dyslipidemia has a close relation with intracranial arteriosclerosis, especially in Chinese population. There are also other studies showing that hypercholesteremia has a close relation with extracranial arteriosclerosis (43-45)

Visceral obesity is the basic constitution of abdominal obesity, which can result in an increase in free fatty acid release. An increase in plasma free fatty acid level plays an important role in occurrence mechanism of insulin resistance, induction of oxidative stress, and reduction of inflammation and blood vessel responsiveness. Moreover, visceral obesity can increase the secretion of very low-density lipoproteins, inhibit apolipoprotein B degradation and contribute to the formation of smaller low-density lipoproteins, which are closely related with the incidence of arteriosclerosis. Abdominal obesity showed a correlation with intracranial arteriosclerosis (14), however we did not find any correlation between abdominal obesity and intracranial and extracranial arteriosclerosis.

The results of Logistic regression in this study showed that the risk of intracranial arteriosclerosis occurrence would increase with an increase in the number of components of metabolic syndrome. This rising trend was not found in extracranial arteriosclerosis group, which is relatively consistent with the previous results (14, 33).

The drawback of this study lies in cross-sectional study, which only described the incidence rate of metabolic syndrome and the prevalence rate of intracranial and extracranial arteriosclerosis of patients who were admitted to our hospital from Sep. 2015 to Sep. 2017. The related risk between metabolic syndrome and intracranial and extracranial arteriosclerosis revealed in this study was very meaningful, providing a basis for target prevention and treatment on intracranial and extracranial arteriosclerosis lesion in the future.

# Conclusion

The high prevalence rate, death rate and disability confirmed the importance of the studies on intracranial and extracranial arteriosclerosis. This includes multicenter, large sample and prospective studies on intracranial and extracranial arteriosclerosis and risk factors, as well as the analysis of plaque image and pathology. Besides, intracranial arterial stenosis is an important reason for cerebral arterial thrombosis in China, therefore, seeking a reasonable treatment on intracranial arteriosclerosis has a far-reaching significance.

# Ethical considerations

Ethical issues (Including plagiarism, informed consent, misconduct, data fabrication and/or falsification, double publication and/or submission, redundancy, etc.) have been completely observed by the authors.

### Acknowledgements

There was no financial source of this study.

### **Conflicts of interests**

The authors declare that there is no conflict of interests.

### References

- Wang LD, Wang JH, Peng B, Xu YM (2017). Report on the Chinese Stroke Prevention 2016 Summay. *Chinese Journal of Cerebrovascular Diseases*, 14 (04): 217-24.
- Lam RC, Lin SC, DeRubertis B, Hynecek R, Kent KC, Faries PL (2007). The impact of increasing age on anatomic factors affecting carotid angioplasty and stenting. J Vast Surg, 45 (5): 875-80.
- Sitzer M, Skutta M, Siebler M, Sitzer G, Siegrist J, Steinmetz H (1998). Modifiable stroke risk factors in volunteers willing to participate in a prevention program. *Neuroepidemiology*, 17 (4): 179-87.
- Leung SY, Ng TH, Yuen ST, Lauder IJ, Ho FC (1993). Pattern of cerebral atherosclerosis in Hong Kong Chinese. Severity in intracranial and extracranial vessels. *Stroke*, 24 (6): 779-86.
- Kasner SE, Chimowitz MI, Lynn MJ, et al (2006). Predictors of ischemic stroke in the territory of a symptomatic intracranial arterial stenosis. *Cinculation*, 113 (4): 555-63.

- Higashida RT, Meyers PM, Connors JJ, et al (2009). Intracranial angioplasty and stenting for cerebral atherosclerosis: a position statement of the American Society of Interventional and Therapeutic Neuroradiology, Society of Interventional Radiology, and the American Society of Neuroradiology. J Vasc Interv Radiol, 20 (7 Suppl): S312-6.
- Sacco RL, Adams R, Albers G, et al (2006). Guidelines for prevention of stroke in patients with ischemic stroke or transient ischemic attack: a statement for healthcare professionals from the American Heart Association/American Stroke Association Council on Stroke: co-sponsored by the Council on Cardiovascular Radiology and Intervention: the American Academy of Neurology affirms the value of this guideline. *Cinulation*, 113 (10): e409-49.
- Wong KS, Li H, Chan YL, et al (2000). Use of transcranial Doppler ultrasound to predict outcome in patients with intracranial large-artery occlusive disease. *Stroke*, 31 (11): 2641-7.
- Kim JS, Kim YJ, Ahn SH, Kim BJ (2018). Location of cerebral atherosclerosis: Why is there a difference between East and West. *Int J Stroke*, 13 (1): 35-46.
- Kalita J, Misra UK, Kumar B, Somarajan BI, Kumar S, Mittal B (2018). ACE and ADD1 gene in extra and intracranial atherosclerosis in ischaemic stroke. *Neural Res*, 35 (4): 429-34.
- Kim BJ, Lee KM, Lee SH, et al (2018). Ethnic Differences in Intracranial Artery Tortuosity: A Possible Reason for Different Locations of Cerebral Atherosclerosis. J Stroke, 20 (1): 140-1.
- He YN, Zhao WH, Zhao LY, (2017). [Prevalence of metabolic syndrome in Chinese adults in 2010-2012]. Zhonghua Liu Xing Bing Xue Za Zhi, 38 (2): 212-5.
- De Silva DA, Woon FP, Lee MP, Chen CL, Chang HM, Wong MC (2009). Metabolic syndrome is associated with intracranial large artery disease among ethnic Chinese patients with stroke. J Stroke Cerebronas Dis, 18 (6): 424-7.
- Park JH, Kwon HM, Roh JK (2007). Metabolic syndrome is more associated with intracranial atherosclerosis than extracranial atherosclerosis. *Eur J Neurol*, 14 (4): 379-86.
- 15. North American Symptomatic Carotid Endarterectomy Trial Collaborators, HJM B, Taylor DW, et al (1991). Beneficial effect of carotid endarter-

ectomy in symptomatic patients with high-grade carotid stenosis. N Engl J Med, 325 (7): 445-53.

- Alberti KG, Zimmet P, Shaw J (2005). The metabolic syndrome-a new worldwide definition. *Lancet*, 366 (9491): 1059-62.
- Moossy J (1966). Cerebral infarction and intracranial arterial thrombosis. Necropsy studies and clinical implications. *Anh Neurol*, 14 (2): 119-23.
- Suemoto CK, Grinberg LT, REP L, et al (2018). Morphometric measurements of extracranial and intracranial atherosclerotic disease: A population-based autopsy study. *Atherosclerosis*, 270: 218-23.
- Nishimaru K, McHenry LC, Toole JF (1984). Cerebral angiographic and clinical differences in carotid system transient ischemic attacks between American Caucasian and Japanese patients. *Stroke*, 15 (1): 56-9.
- Sacco RL, Kargman DE, Gu Q, Zamanillo MC (1995). Race-ethnicity and determinants of intracranial atherosclerotic cerebral infarction. The Northern Manhattan Stroke Study. *Stroke*, 26 (1): 14-20.
- Suh DC, Lee SH, Kim KR, et al (2003). Pattern of atherosclerotic carotid stenosis in Korean patients with stroke: different involvement of intracranial versus extracranial vessels. *AJNR Am J Neuroradiol*, 24 (2): 239-44.
- Wong KS, Huang YN, Yang HB, et al (2007). A door-to-door survey of intracranial atherosclerosis in Liangbei County, China. *Neurology*, 68 (23): 2031-4.
- Jung KH, Lee SH, Kim BJ, et al (2012). Secular trends in ischemic stroke characteristics in a rapidly developed country: results from the Korean Stroke Registry Study (secular trends in Korean stroke). *Circ Cardiovasc Qual Outcomes*, 5 (3): 327-34.
- 24. Lei C, Wu B, Liu M, Chen Y (2014). Risk factors and clinical outcomes associated with intracranial and extracranial atherosclerotic stenosis acute ischemic stroke. *J Stroke Cerebrovasc Dis*, 23 (5): 1112-7.
- Zou C, Wei C, Wang Z, Jin Y (2017). Sex differences in outcomes and risk factors among elderly patients with ischemic stroke. *Oncotarget*, 8 (61): 104582-93.
- Higashi Y, Miyata T, Shigematsu H, et al (2017). Baseline Characterization of Japanese Peripheral Arterial Disease Patients - Analysis of Surveillance of Cardiovascular Events in Antiplatelet-

Treated Arteriosclerosis Obliterans Patients in Japan (SEASON). *Cin J*, 80 (3): 712-21.

- 27. Csordas A, Bernhard D (2017). The biology behind the atherothrombotic effects of cigarette smoke. *Nat Rev Cardiol*, 10 (4): 219-30.
- Chen H, Hong H, Xing S, et al (2015). Intracranial versus extracranial artery dissection cases presenting with ischemic stroke. J Stroke Cerebrovasc Dis, 24 (4): 852-9.
- Uehara T, Tabuchi M, Mori E, Yamadori A (2003). Evolving atherosclerosis at carotid and intracranial arteries in Japanese patients with ischemic heart disease: a 5-year longitudinal study with MR angiography. *Eur J Neurol*, 10 (5): 507-12.
- Seo WK, Yong HS, Koh SB, et al (2008). Correlation of coronary artery atherosclerosis with atherosclerosis of the intracranial cerebral artery and the extracranial carotid artery. *Eur Neurol*, 59 (6): 292-8.
- 31. Bae HJ, Yoon BW, Kang DW, et al (2006). Correlation of coronary and cerebral atherosclerosis: difference between extracranial and intracranial arteries. *Cerebrovasc Dis*, 21 (1-2): 112-9.
- Park JH, Kwon HM (2008). Association between metabolic syndrome and previous ischemic lesions in patients with intracranial atherosclerotic stroke. *Clin Neurol Neurosurg*, 110 (3): 215-21.
- Bang OY, Kim JW, Lee JH, et al (2005). Association of the metabolic syndrome with intracranial atherosclerotic stroke. *Neurology*, 65 (2): 296-8.
- 34. D'Armiento FP, Bianchi A, de Nigris F, et al (2001). Age-related effects on atherogenesis and scavenger enzymes of intracranial and extracranial arteries in men without classic risk factors for atherosclerosis. *Stroke*, 32 (11): 2472-9.
- Huang HW, Guo MH, Lin RJ, et al (2007). Prevalence and risk factors of middle cerebral artery stenosis in asymptomatic residents in Rongqi County, Guangdong. *Cerebrovasc Dis*, 24 (1): 111-5.
- Campia U, Sullivan G, Bryant MB, Waclawiw MA, Quon MJ, Panza JA (2004). Insulin impairs endothelium-dependent vasodilation independent

of insulin sensitivity or lipid profile. *Am J Physiol Heart Circ Physiol*, 286 (1): H76-82.

- Palmieri VO, Grattagliano I, Portincasa P, Palasciano G (2006). Systemic oxidative alterations are associated with visceral adiposity and liver steatosis in patients with metabolic syndrome. J Nutr, 136 (12): 3022-6.
- Ritz K, Denswil NP, Stam OC, van Lieshout JJ, Daemen MJ (2014). Cause and mechanisms of intracranial atherosclerosis. *Cinculation*, 130 (16): 1407-14.
- Caplan LR, Gorelick PB, Hier DB (1986). Race, sex and occlusive cerebrovascular disease: a review. *Stroke*, 17 (4): 648-55.
- 40. Rincon F, Sacco RL, Kranwinkel G, et al (2009). Incidence and risk factors of intracranial atherosclerotic stroke: the Northern Manhattan Stroke Study. *Cerebrovasc Dis*, 28 (1): 65-71.
- López-Cancio E, Galán A, Dorado L, et al (2012). Biological signatures of asymptomatic extra- and intracranial atherosclerosis: the Barcelona-AsIA (Asymptomatic Intracranial Atherosclerosis) study. *Stroke*, 43 (10): 2712-19.
- Wong KS, Ng PW, Tang A, Liu R, Yeung V, Tomlinson B (2007). Prevalence of asymptomatic intracranial atherosclerosis in high-risk patients. *Neurology*, 68 (23): 2035-8.
- 43. Kim JS, Nah HW, Park SM, et al (2012). Risk factors and stroke mechanisms in atherosclerotic stroke: intracranial compared with extracranial and anterior compared with posterior circulation disease. *Stroke*, 43 (12): 3313-18.
- Uehara T, Tabuchi M, Mori E (1998). Frequency and clinical correlates of occlusive lesions of cerebral arteries in Japanese patients without stroke. Evaluation by MR angiography. *Cerebrovasc Dis*, 8 (5): 267-72.
- Kim YD, Choi HY, Jung YH, et al (2009). Classic risk factors for atherosclerosis are not major determinants for location of extracranial or intracranial cerebral atherosclerosis. *Neuroepidemiology*, 32 (3): 201-7.