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4 **Risk factors of ischemic cerebrovascular accident: an actual state inPhuTho**  
5 **General Hospital, Vietnam**

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12 **Abstract.** Worldwide, each year the quantity of stroke death equal to5.5 million people. In South-East  
13 Asia, the proposed value is equal to 500 thousand people. In Vietnam the situation of ischemic strokes is  
14 critical. It leads to recently appeared and uncontrolled risk factors. The current work is devoted to the  
15 main risk factors of ischemic cerebrovascular accident.Investigating most important risk factors in  
16 ischemic stroke patients presented in PhuTho Provincial General Hospital. A prospectively observational  
17 study. Under propose research the application (Software SPSS 18.0 and STATA 10.0) had been  
18 initiated.TIA was seen at 13.0% of all cases and risk of progressing to severe or death with OR = 5.2;  
19 63.5% of all patients appeared with hypertension had the risk of being severe or death with OR 3.2. Pre-  
20 exist of heart diseases was available in 28.1% cases with OR = 2.6 in the risk of getting worse or death.  
21 Hyperlipidemia rate was high, risk of deterioration with OR = 2.6. Alcohol and tobacco addiction was  
22 found in 72.6% patients and risk of becoming death or worse 2.4 times higher than alcohol and smoke-  
23 free patients.Some risk factors had good predictive prognosis value in ischemic stroke. The  
24 hypertension, cigarettes and alcohol abuse had been detected as one of the riskiest of the progress  
25 cerebrovascular accident. Most of the patients had pre-history addiction and diseases that can be one the  
26 cause of the Ischemic Cerebrovascular Accident (ICA). The ICA is the most health problems in Vietnam  
27 that need more attention.

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29 **Key words:** Risk factors analysis, Ischemic stroke, Haemorrhagic stroke, ischemic cerebrovascular  
30 accident, stroke risk factors, PhuTho hospital

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## 33 INTRODUCTION

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35 In Europe, it has been estimated that about 1 million cases of cerebrovascular accidents are  
36 presented. In the United States of America (USA), the stroke rate is 794/100.000 of population and  
37 only 400.000 patients are able to be discharged. Current researches shows that in the USA, one  
38 stroke happens every 40 seconds and it is approximately equal to 800 thousand accidents per a year  
39 (Nakajima and Chester, 2019). In Asia, the incidence of stroke is different between different  
40 countries. For example, in Japan the stroke rate is 303/100.000 of the populations, in China this value  
41 equals 115.6/100.000 in which 370/100.000 of the Beijing population only (Nguyen Minh Hien,  
42 2013).

43 Nowadays, more and more people are taking notice about the stroke. The stroke's knowledge  
44 has been significantly improved, especially in fast and accurate diagnostic imaging techniques,  
45 pathology-based treatments and the widespread establishment of stroke centers. Many achievements  
46 were reached in stroke prevention and treatment. Nevertheless, a stroke modality is still high and  
47 stays in third place after cardiovascular diseases and cancer, and most commonly in neurological  
48 diseases, which contributed 20% of all internal medical conditions.

49 The solidus includes Acute Ischemic (AIS) and Acute Hemorrhagic (AHS) strokes. The AIS  
50 have a more common incidence, which value is about 80 - 85%. In the European Union (EU), the  
51 USA and other first developed countries, the AHS had been observed in a range of 10-15%. In Asia,  
52 the AHS could be increased, but never higher than AIS. In Vietnam, the researcher Le Van Thanh  
53 investigated 2962 stroke patients: 40.42% of them had the AHS and 59.58% the AIS. In another  
54 study of Hoang Khanh in Hue city, the AHS counted 39.42% and the AIS were about 60.58% (Trinh  
55 Viet Thang, 2011).

56 An average life expectancy in Vietnam is currently increased due to developed social-  
57 economic conditions, but unfortunately, at the same time more strokes are observed (Dao  
58 ThiBichHoa, 1996; Dang Quang Tam, 2005). Many studies were conducted by focusing on the AIS,  
59 but most of them were done in the national hospitals. Risk factors of AIS weren't noticed  
60 appropriately in accordance with a lack of money and bad habits (drinking alcohol, smoking) directly  
61 connected to its traditional and social factors. On the other hand, stroke patients are cared for in  
62 different facilities: the Intensive Care Unit (ICU), emergency room, Internal medicine wards,  
63 cardiology wards, neurology wards etc. Therefore, a study of the AIS in PhuTho was never  
64 conducted before and highly recommended being done to investigate the actual state of the AIS and  
65 to prepare resources in the diagnosis and treatment of the AIS.

66 The aim of the following work leads to evaluate all-important risk factors of the AIS in  
67 PhuTho Provincial Hospital.

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## 69 MATERIAL AND METHODS

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71 Between October 2014 and October 2016 years, more than 190 AIS patients received medical  
72 care in PhuTho provincial general hospital. The World Health Organisation (WHO) diagnostic  
73 criteria of the AIS in 1990: sudden symptoms (minutes to hours, days) which prolong more than 24  
74 hours, with focal symptoms (belonging to impair arteries) without a correlation to a trauma (Roth et  
75 al., 2015). The Brain Magnetic Resonance Imaging (MRI) showing at least one of the 2 signs:

76 Homogeneous intensity on T1W, a T2W hyperintensity in subcortical area and loss of  
77 differences between the parenchyma and cortex. A hypodense signal on T1W and a hyperintense  
78 signal on T2W (Hoang Duc Kiet, 2004). The Transient Ischemic Attack (TIA), ischemic stroke in a  
79 patient with the prehistory of head trauma or blunt trauma. Comorbidity of encephalitis, meningitis  
80 or brain tumor; ischemic stroke converted to haemorrhage; relapsed ischemic stroke or normal  
81 finding on MRI. Statistical Analysis: Software SPSS 18.0 and STATA10.0

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## 83 RESULTS

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The following section shows a practically obtained statistical data. An investigation of the issue has analysed the data of 192 patients. All respondents agreed on the process of the data in accordance with the ethical standards of the responsible committee on human experimentation (institutional or regional) and with the Helsinki Declaration of 1975, as revised in 2000.

It is important to consider that AIS and AHS have similar risk factors, but there are some notable differences. Both types of stroke are characterized by non-modifiable and modifiable stroke risk. Non-modifiable risks similar for both types are age, gender, race and ethnicity (Boehme et al., 2017). Similar modifiable factors are Hypertension (HTN), Transient Ischemic Attack (TIA), genetic heritage, blood pressure, bad habits (alcohol, tobacco etc.) (Bhatt et al., 2008). Different factors that cause Ischemic stroke are heart diseases, hyperlipidemia, physical inactivity, diabetes (Boehme et al., 2017). In the following work, all of this risk had been analyzed in accordance to patient history.

### A pre-history of an addiction

As it has been mentioned above, alcohol and tobacco initiate top cases of the Ischemic Cerebrovascular Accident (ICA) (Han and Berry, 2019). In the following work, the statistical data of the main factors that reveals the precursors of the ischemic disease had been collected and shown in Table 1.

Table 1. Tobacco and alcohol abuse

History	Group II (n = 64)		Group I (n = 128)		n (%) n = 192
	n	%	n	%	
Alcohol abuse	22	34.4	27	21.1	49 (25.8%)
Tobacco addiction	27	42.2	26	20,3	52 (27.4%)
Alcohol and tobacco addiction	17	25.5	21	16.4	37 (19.4%)

As it has been observed in Table 1, it is possible to conclude that in both groups the constituent part of the alcohol abuse equals to 25.8%. The Tobacco addiction is a bit higher and equals to 27.4%. Because of the death rate and morbidity related to both tobacco and alcohol abuse, in Table 1 also addressed both addictions. The rate of both is lower and relates to 19.4% of the total number tested patients.

### Other pre-medical history

In the course of the research, other diseases had been determined like: the HTN, the diabet, heart diseases and the TIA. The proposed data had been fulfilled in Table2.

Table 2. Pre-medical history playing role of risk factors

History	Group II (n = 64)		Group I (n = 128)		n (%) (n = 192)
	N	%	n	%	
Hypertension (HTN)	52	80,2	70	54.7	122 (63.5%)
Diabetes	12	19,8	18	14.1	30 (15.6%)
Heart diseases (Heart failure, AF, mitral stenosis)	21	33.3	33	25.8	54 (28.1%)

TIA	18	28.6	07	5.4	25 (13.0%)
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Diagnostic results showed that a major risk rate corresponds to a patient group of heart diseases and equals to 28.1%. The intermediate and lower rates correspond to the patients with diabetes and the TIA. The critical risk factoring of 63.5% is associated with the HTN. The following results show that in Vietnam a real quantity of people with HTM disease is higher than it had been mentioned in statistical data received through the 2005 National Adult Obesity Survey by the Do Phi Phuong Ha (Do, 2014). A patient consultation was conducted in order to initiate lifestyle modification, promotion of physical exercises and healthy food during a day. In Table 3, a pressure measurement of the HTN group has been fulfilled.

Table 3. Blood pressure measurement in HTN group

Blood pressure	Minimum	Maximum	Mean	SD
Systolic BP (mmHg)	140	180	152	10.2
Diastolic BP (mmHg)	70	120	94	8.9
Years of HTN	1	20	7	4.7

Also, all patients from the first and second groups had been qualified due to the risk factor number and the following data had been represented in the Table 4.

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Table 4. Patients Classification based on risk factors number

		Number of risk factors						Total
		0	1	2	3	4	5	
Group II	n	4	14	12	9	15	10	64
	%	6.2	21.8	18.7	14.1	23.4	15.8	100
P		< 0.05						
Group I	n	11	45	38	12	15	5	128
	%	8.5	35.2	29.7	9.4	11.7	5.5	100

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The predictive value of some risk factors: An expression of the likelihood that had been given on the basis of investigation, which correlates with the positive or negative test results of the mentioned diseases had been shown in Tables 5, 6 and 7.

Table 5. Patient distribution based on severity

	Number of patients, n	Ratio, %
Group I (mild and medium)	128	66.7
Group II (severe – death)	64	33.3
Total amount	192	100

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Table 6. Predictive value of pre-medical historical risk factors

Diseases	Yes	No	Total
Hypertension			
Group II	52 (81.0%)	12 (19.0%)	64
Group I	71 (55.4%)	57 (44.6%)	128
OR = 3,2	95% CI = 1,3 – 7,8		p = 0,009
Heart diseases (Heart failure, AF, mitral valve stenosis)			
Group II	22 (34.3%)	42 (65.6%)	64 (100%)
Group I	34 (26.5%)	94 (73.5%)	128 (100%)
OR=26	95% CI = 1.1 – 6.1		p = 0.034
Diabetes			
Group II	13 (20.3%)	51 (79.7%)	64 (100%)
Group I	19 (14.8%)	109(85.2%)	128 (100%)
OR = 2.2	95% CI = 0.9 – 5.1		p = 0.073

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TIA			
Group II	19(29.7%)	45 (70.3%)	64 (100%)
Group I	08 (6.2%)	120(93.8%)	128 (100%)
OR = 5.2	95% CI = 1.7 – 15.8		p = 0.004
Tobacco addiction			
Nhóm II	28(43.7%)	36 (56.3%)	64 (100%)
Nhóm I	24(18.8%)	104(81.2%)	128 (100%)
OR = 2,4	95% CI = 1,0 – 5,6		p = 0,044
Alcohol abuse			
Nhóm II	23(35.9%)	41 (64.1%)	64 (100%)
Nhóm I	28(21.9%)	100(78.1%)	128 (100%)
OR = 2,5	95% CI = 1,1 – 6,0		p = 0,038
Hyperlipidemia			
Group II	42 (65.6%)	22 (34.4%)	64 (100%)
Group I	53 (41.4%)	75 (58.9%)	128 (100%)
OR = 2.6	95% CI = 1.2 – 5.9		P = 0.02

Table 7. Predictive values of MRI finding

Severity	Size of ischemic area		n (%)
	< 3cm	≥ 3cm	
Group II	27 (42.2%)	37 (57.8%)	64 (100%)
Group I	81 (63.2%)	47 (36.8%)	128 (100%)
OR = 3.4	95% CI = 1.5 – 7.4		p = 0.002
Severity	Number of ischemic area		n (%)
	1	>1	
Group II	48 (75.0%)	16 (25.0%)	64 (100%)
Group I	94 (73.5%)	34 (26.5%)	128 (100%)
OR = 0.7	95% CI = 0.3 – 1.7		p = 0.536

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## DISCUSSION

In research were identified risk factors of ischemic cerebrovascular accident such as tobacco addiction, alcohol abuse, hypertension, diabetes mellitus, previous heart diseases and TIA.

### Tobacco addiction

Smoking is becoming a crucial issue in modern societies. Vietnam is one of the countries in which the smoking rate is highest in all over the world: 50% of the adult male have a smoking habit, which is about 17 million smoking people. Many studies before show the strong correlation between smoking and stroke. In this study, 27.4% of patients smoked (table 1) which was higher than the results of Phan ThiHuong (16.0%, Dinh Van Thang 17.9%). These results from the smoking rate of

178 Vietnamese increased significantly recently.

179 Patients with a pre-social habit of smoking have an increased risk of converting to severe  
180 condition or death in AIS with OR = 2.4 (p = 0.044; CI: 1.02 – 5.6). The results are considered  
181 similar to other Vietnamese and international authors. In Le Quang Cuong study, in females smoking  
182 increased the risk of the AIS in 1.9 times and 2.5 times in males while Wolf P.A et al. (1988)  
183 concluded that smoking rose up the risk of stroke about 40% of male and 60% in females. The study  
184 of Wolf P.A. et al. conducted within 26 years shows that heavy addiction (more than 40 cigarettes a  
185 day) had a risk of stroke 2 times higher than lighter ones (less than 10 cigarettes a day) and this risk  
186 reduced greatly 5 years after the smoke quitting. Therefore, smoking is seriously considered a risk of  
187 AIS (Bui My Hanh et al., 2019).

### 188 **Alcohol abuse**

191 More than 25.8% of the study population is alcoholic which was similar to Le Ngoc Trong  
192 (26.9%) and Dang Quang Tam (31%); and higher than Pham Thi Thu Ha (6.62%) and Phan Thi  
193 Huong (7.06%) (Dang Quang Tam, 2005; Phan Thi Huong, 2004; Wolf P.A. et al., 1988).

194 The Odd ratio of alcohol abuse on severity and death was 2.5 (p = 0.038, CI = 1.1  
195 – 6.0) compared to non-alcohol use. Many other studies impressed the role of alcohol in the  
196 deterioration of stroke. Zhang's study in China found that heavy alcoholic people had a 1.9 times  
197 higher risk of stroke than others (Zhang et al., 2014). Mukamalet al. (2003) observed alcohol addicts  
198 during 14 years expressed that the habit of drinking 10-29 gram ethanol a day, 3 – 4 days a week  
199 induced the lowest risk of stroke whereas people consumed more than 2 glasses of whiskey a day had  
200 the highest risk. Le Quang Cuong reported that ethanol consumption increased 10% in the population  
201 would raise the stroke morbidity up to 29% and mortality up to 16% (Nguyen and Hoang, 2018).

202 Because alcoholism significantly induced hypertension, increased platelet cell adherence,  
203 fulminant atrial fibrillation which all contributed to AIS. Hence, ethanol abuse, especially in such a  
204 long duration, had a bad effect on the severity of stroke, especially AIS.

### 205 **Hypertension (HTN) history**

207 In accordance with the results of the proposed investigation, the HTN was seen in 63.5% of  
208 the population that was higher than Dao Thi Bich Hoa (40%) (Dao Thi Bich Hoa, 1996). This result  
209 from the HTN became more popular, especially in developed countries. In a report by the WHO in  
210 2000 year it is estimated that 972 million people had the HTN and this number was predicted to be  
211 1.5 trillion people in 2025 (World Health Organization, 2003). In the United States of America, the  
212 medical system is considered the best in the world and only 77.6% people recognized the HTN.  
213 Since, the result was collected by asking the patients directly or their family member so that the  
214 actual number of the HTN in the proposed population should be much higher than 63.7% that in the  
215 proposed work had been mentioned.

217 Other studies show that both systolic and diastolic blood pressure independently induced  
218 different types of stroke (Webb et al., 2018). It had been found that systolic BP higher than 160 mm  
219 Hg and the BP diastolic is higher than 95 mm Hg could in 3.1 times increase the risk of the AIS in  
220 men and 2.9 times in women. If the BP systolic stayed in the range of 140 and 159 mm Hg and the  
221 BP diastolic stayed in the range of 90 and 94 mm Hg would raise the risk of stroke up to 50% (Ngo  
222 Duc Vuong, 2010).

223 The longest duration with the HTN, the more severe symptoms of stroke patients could  
224 suffer. In the proposed study, the mean duration of the HTN is 7 years. Consequently, chronic HTN  
225 patients ought to pay a higher warning attention to the AIS. In accordance to the proposed findings,  
226 the HTN put the AIS patient at the risk of getting worse or even death 3.2 times higher than non-  
227 HTN with the CI: 1.3 – 7.8 and p = 0.009). Other authors estimated that a 10 mm Hg reduction in  
228 systolic BP in adults would decrease 30% in the risk of death due to heart diseases and also drop 40%  
229 risk of mortality due to the stroke. Hence, early recognition and regular treatment of the HTN and  
230 community education for this issue are extremely necessary. However, even in the United States of

America, only 67.9% in 73 million and only 44.1% of those were tightly controlled (Webb et al., 2018). So, the difficulty in the HTN management does not just belong to developing countries, but also of the developed ones.

### **Diabetes mellitus**

There has been a huge amount of trials worldwide affirmed that diabetes mellitus or diabetes is a crucial risk of atherosclerosis. It played an important role in embolism appearance and the AIS. In the United States in 1976-1980 years, diabetes patients had a risk of the AIS from 2.5 to 4 times higher than people who didn't have diabetes (Deshpande et al., 2008). In the Honolulu Cardiovascular Program, the Japanese Hawaii with diabetes had 2 times risk of the DVT compared to others without diabetes. In another investigation Framingham, although the most influence of diabetes was on microvascular diseases, although that still had a bad effect on coronary and cerebral vascular (Wolf et al., 1988).

In the study of proposed work, the Odd ratio of diabetes on the AIS is 2.2 but with condition when  $p > 0.05$ . It had been suggested that diabetes could increase the risk of getting the AIS or being worse with the AIS, however the p value suggested that it is possible to try with a larger population to confirm this number. Within the proposed AIS population, diabetes took 15.6%, which was higher than previous studies such as Phan Thi Huong 6.2% (Phan Thi Huong, 2004). One of the reasons was the diabetes rate in the proposed country increased in the last few years. The recent report found that diabetes morbidity rose from 8% to 20% annually putting Vietnam in the list of countries, which had a fastest increased number of diabetes worldwide.

Explaining the sharply elevated diabetes rate recently, it had been considered the highest standard of living, imbalance between nutrition and physical activities. These should be considerable factors to be strictly justified to reduce diabetes and the AIS morbidity.

### **Previous Heart Diseases**

In the proposed study, the 28.1% of patients had previous heart diseases similar to result from Dang Quang Tam 21.2%, Dinh Van Thang 6.5%, and Nguyen Xuan Than 6.1%. Those patients had an increased risk of severe and death with  $OR = 2.6$  ( $CI = 1.1 - 6.1$ ;  $p = 0.034$ ) (Dang Quang Tam, 2005). This factor had a high predictive value with  $p < 0.05$ . In a patient with pre-exist cardiovascular diseases, the risk of the AIS was higher and when they suffered from AIS, the possibility of deterioration was higher because cardiovascular conditions could appear embolism from the heart and reduced peripheral and cerebral perfusion. The Atrial Fibrillation (AF) was a common impairment and was a strong risk of the AIS, a well-cared for the AF therefore would prevent AIS's appearance effectively.

A Framingham study showed that there was a clear elevation of the AIS in the AF patients: 1.5% at the age of 50-59 and 23.5% at the age of 80-89. The author also impressed that 8% of men and 11% of women could have AIS within 6 years after an acute Myocardial Infarction (MI). Besides the AF, other cardiovascular conditions such as dilated Myocardiopathy (DMC), Mitral Valve Prolapse (MVP), Mitral Valve Stenosis (MVS), Heart Failure (HF), artificial heart valve, acute endocarditis and other congenital heart defects could induce the AIS with different risk level.

The proposed study didn't focus on the detailed diagnosis of heart diseases due to the small number of this group. Furthermore, heart disease has been just a limited part between other important factors needed to be investigated. However, from received limited data, the actual information from the patients who had previous heart diseases did improve the belief and efficacy in the treatment of AIS patients.

### **Previous Transient Ischemic Attack**

The TIA is an important factor influenced on the risk of the AIS but also the severity of AIS,

283 especially the death risk in the first few days after the AIS on set. The TIA was combined with  
284 neurological deficits that existed only within 24 hours and recovered completely, but tended to  
285 relapse multiple times later. However, the TIA highly warned about a widespread atherosclerosis and  
286 it easily became worse, which could result in the AIS and Multifocal ischemic stroke. In Fleming et  
287 al. (2015) study, in the first month, 8% of the TIA patients suffered from actual AIS, 20% of these  
288 patients had a risk of acute MI, stroke or sudden death in the first 12months.

289 In proposed study it was found that the TIA equals 13.0%. It is similar to LeThiHoaBinh  
290 12.8%, Le Van Thinh 11% and Dinh Van Thang 8.9% (Le Van Thinh, 2004). When a patient  
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293 had the TIA, risk of the AIS severe was highest with OR = 5.2, p = 0.004). So then, if the patient has  
294 previous TIA, patients would have the risk of becoming severely or death 5.2 times higher than  
295 patients without the TIA before. The TIA was a clear sign of “nearly” obstruction in the cerebral  
296 vascular system, which raised up an important alarm of further poor-prognosis complications after.

297 In the studies of Fleming, the TIA had the highest risk of the AIS severe compared to all other  
298 risk factors such as the HTN, diabetes and heart diseases (Fleming et al., 2015). Because of this, TIA  
299 treatment could help to prevent the AIS in the future and was considered more crucial than  
300 reperfusion therapies after AIS happened already. From those results are strongly recommended  
301 investigating and closely observe the TIA signs to achieve an adequate diagnosis and treatment.  
302 Therefore, it could prevent poor-outcome, complications and reduce the AIS severe or death.

### 303 304 **Total Number of Risk factors in a patient**

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306 In the mild and medium group, patients had only one or maximum two risk factors while in  
307 the severe or death group most patients had at least 3 risk factors, the difference was significant.  
308 Therefore, the more risk factors a patient had previously, the more severe symptoms patients could  
309 suffer. In the study of Le Thi Thanh Tuyen and Le Tu Phuong Thao, most patients had 2 risk factors,  
310 rarely worn, especially many patients brought 3 and more risk factors (Le Thi Thanh Tuyen, 2010).  
311 This rang an alarming notice because uncontrolled hyperlipidemia, the HTN, diabetes would  
312 contribute directly to the severity of the AIS. Each factor had a special effect on each other. For  
313 example, patients with a hyperlipidemia tend to have atherosclerosis and the HTN whereas  
314 uncontrolled diabetes surely increased atherosclerosis and therefore increased the risk of  
315 ischemicstroke.

### 316 317 **CONCLUSION**

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319 The research shows that with the increasing number of diseases such as diabetes,  
320 hypertension, various heart diseases, as well as bad habits and inactive lifestyles can lead to the ICA.  
321 As it had been shown in proposed analysis, when more risk factors the patient have, more likely it  
322 had be fatal forhim.

323 The TIA was found in 13.2% of patients which could progress to a severe group with OR =  
324 5.2. Hypertension was seen in 63.7% patients and had a risk of being severe 3.2 times higher than  
325 normotensive cases. Patients with heart diseases contributed 28.4% of the population and risk of  
326 getting worse with OR = 2.6. The highest rate of hyperlipidemia was caught with risk of  
327 deterioration with OR = 2.6). Cigarettes and alcohol abuse was 72.6%; smoke addiction had a risk of  
328 being worse with OR = 2.4 while the risk of alcohol abuse was with OR = 2.5. The problem is actual  
329 and needs furtherinvestigation.

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## CONFLICTS OF INTEREST

The authors declare no conflict of interest.

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