

Age and Meteorological Factors in the Occurrence of Spontaneous Intracerebral Hemorrhage in a Metropolitan City

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Objective : The aim of this study was to investigate the correlation between meteorological factors and occurrence of spontaneous intracerebral hemorrhage (ICH) according to age.

Materials and Methods : We retrospectively analyzed the records of 735 ICH patients in a metropolitan hospital-based population. Observed and expected numbers of ICH patients were obtained at 5°C intervals of ambient temperature and a ratio of observed to expected frequency was then calculated. Changes in ambient temperature from the day before ICH onset day were observed. The Wilcoxon-Mann-Whitney test was used to test differences in meteorological variables between the onset and non-onset days. The Kruskal-Wallis test was used for comparison of meteorological variables across gender and age.

Results : ICH was observed more frequently (observed/expected ratio ≥ 1) at lower mean, minimum, and maximum ambient temperature ($p = 0.0002$, 0.0003 , and 0.0002 , respectively). Significantly lower mean, minimum, and maximum ambient temperature, dew point temperature, wind speed, and atmospheric pressure ($p = 0.0003$, 0.0005 , 0.0001 , 0.0013 , 0.0431 , and 0.0453 , respectively) was observed for days on which spontaneous ICH occurred. In the subgroup analysis, the ICH onset day showed significantly lower mean, minimum, and maximum ambient temperature, dew point temperature, relative humidity, and higher atmospheric pressure in the older (≥ 65 years) female group ($p = 0.0093$, 0.0077 , 0.0165 , 0.0028 , 0.0055 , and 0.0205 , respectively).

Conclusion : Occurrence of spontaneous ICH is closely associated with meteorological factors and older females are more susceptible to lower ambient temperature.

J Cerebrovasc Endovasc Neurosurg.
2014 September;16(3):209-215

Received : 19 June 2014

Revised : 31 August 2014

Accepted : 1 September 2014

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Keywords Age, Meteorology, Spontaneous intracerebral hemorrhage

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INTRODUCTION

Spontaneous intracerebral hemorrhage (ICH) is a major cause of mortality and disability in neurosurgical practice. The relationship of meteorological factors

and spontaneous ICH has been studied extensively. However, the results varied according to climate, geography, and study design. Seasonal variation has been found in many countries.²⁾⁶⁾¹⁰⁾¹⁵⁾¹⁹⁾²⁰⁾ Various meteorological factors, including ambient temperature,

atmospheric pressure, and change of diurnal temperature were also demonstrated to show association with the incidence of spontaneous ICH.⁴⁾⁶⁾⁷⁾¹¹⁾¹²⁾ In contrast, a single study failed to show a relationship between meteorological variables and the occurrence of ICH.³⁾

In addition, age is a demographic risk factor in the occurrence of ICH.⁵⁾ As in most developed countries, the general population of South Korea is aging. Therefore, interest in public health is likely to increase in the older age group. It is worthwhile to have knowledge of how the occurrence of ICH is influenced by meteorological factors in the older age group, because it might be helpful in promoting public health in a metropolitan city.

MATERIALS AND METHODS

The study population comprised 735 patients admitted to our institution suffering from spontaneous ICH from January 2000 to December 2009. All patients were evaluated by neurosurgeons and underwent workup by brain computed tomographic (CT) scans or magnetic resonance imaging. We excluded secondary causes of ICH (due to cerebral aneurysm, arteriovenous malformation, moyamoya disease, and trauma) based on CT findings and past medical history. When necessary, cerebral angiography was performed to exclude other etiologies. We also excluded patients for whom the onset day was not known and patients who were transferred in from outside the city. We retrospectively analyzed the clinical data of these patients, including age, sex, past history of hypertension, diabetes mellitus, and the date of ICH onset.

Data on ambient temperature (°C), dew point temperature (°C), daily precipitation amount (mm), wind speed (m/s), relative humidity (%), and atmospheric pressure (hPa) from 1 January 2000 to 31 December 2009 in Seoul, where the hospital was located, were obtained from the Korea Meteorological Administration. The hospital is located in northeast Seoul. The dis-

tance between the Seoul meteorological observatory and the hospital is approximately 12 km. For statistical analysis, we used meteorological data measured at the Seoul meteorological observatory.

Changes in mean, minimum, and maximum temperatures were calculated as the differences between the date of and the day before the onset of ICH, i.e., mean temperature change was calculated as mean temperature of onset day - mean temperature of the day before onset.

Observed and expected numbers of ICH patients were obtained by 5°C intervals of ambient temperature and changes in mean, minimum, and maximum temperature. Subsequently, the ratio of observed to expected frequencies was calculated. Expected frequency was calculated as $E_{ij} = (R_i \times C_j) / T$; where E_{ij} is the expected frequency of the cell in the i th row

Table 1. Clinical data of 735 enrolled patients

Parameters	Values (%)
Age (mean ± SD)	62.4 ± 13.2 (years)
Sex	
male	400 (54.4)
female	335 (45.6)
Hypertension	
Yes	396 (53.9)
No	314 (42.7)
Missing	25 (3.4)
Diabetes mellitus	
Yes	125 (17.0)
No	584 (79.5)
Missing	26 (3.5)
Year of ICH onset	
2000	78 (10.6)
2001	83 (11.3)
2002	96 (13.1)
2003	72 (9.7)
2004	82 (11.2)
2005	71 (9.6)
2006	64 (8.7)
2007	77 (10.4)
2008	62 (8.4)
2009	52 (7.0)

ICH = intra-cerebral hemorrhage

Table 2. Observed versus expected numbers of ICH events by ambient temperature and change in mean, minimum, and maximum temperature

	Mean temperature (°C)							Total	<i>p</i> value
	< 0	0-5	5-10	10-15	15-20	20-25	≥ 25		
No. of days	477	518	471	417	543	798	427	3651	0.0002
Observed	112	121	122	80	89	146	65	735	
Expected	97.2	105.1	96.2	83.6	108.4	159.5	85.0	735	
Observed/Expected	1.15	1.15	1.27	0.96	0.82	0.92	0.76	1.00	
	Minimum temperature (°C)							Total	<i>p</i> value
	< 0	0-5	5-10	10-15	15-20	20-25	≥ 25		
No. of days	842	531	454	494	658	599	73	3651	0.0003
Observed	200	125	104	82	122	91	11	735	
Expected	171.5	107.8	91.7	98.6	131.9	119.1	14.4	735	
Observed/Expected	1.17	1.16	1.13	0.83	0.92	0.76	0.76	1.00	
	Maximum temperature (°C)							Total	<i>p</i> value
	< 0	0-5	5-10	10-15	15-20	20-25	≥ 25		
No. of days	192	406	475	422	413	603	1140	3651	0.0002
Observed	49	98	114	103	75	113	183	735	
Expected	39.6	82.6	96.2	86.4	82.2	120.7	227.3	735	
Observed/Expected	1.24	1.19	1.18	1.19	0.91	0.94	0.81	1.00	
	Change in mean temperature from the day before (°C)					Total	<i>p</i> value		
	≤ -5	-5-0	0-5	> 5					
No. of days	113	1566	1946	24	3649		0.2053		
Observed	31	326	373	5	735				
Expected	23.5	315.2	391.6	4.7	735				
Observed/Expected	1.32	1.03	0.95	1.06	1.00				
	Change in minimum temperature from the day before (°C)					Total	<i>p</i> value		
	≤ -5	-5-0	0-5	> 5					
No. of days	95	1815	1650	89	3649		0.6964		
Observed	22	375	320	18	735				
Expected	19.3	364.9	333.0	17.8	735				
Observed/Expected	1.14	1.03	0.96	1.01	1.00				
	Change in maximum temperature from the day before (°C)					Total	<i>p</i> value		
	≤ -5	-5-0	0-5	> 5					
No. of days	251	1481	1770	147	3649		0.0094		
Observed	71	286	342	36	735				
Expected	51.9	297.9	355.3	30.0	735				
Observed/Expected	1.37	0.96	0.96	1.20	1.00				

Chi-square test was used for evaluations.

Temperatures were measured at the Seoul Meteorological observatory from 2000 to 2009

and the *j*th column, *R_i* is the total number of subjects in the *i*th row, *C_j* is the total number of subjects in the *j*th column, and *T* is the total numbers of the whole table.

Non-parametric significance tests were used for comparison of meteorological differences between

groups, since the meteorological factors were not normally distributed. The chi-square test was used for evaluation of observed versus expected numbers of ICH events by ambient temperature. The Wilcoxon-Mann-Whitney test was used to test differences in meteorological variables between the onset and

Table 3. Relationships between onset of ICH and meteorological factors

Meteorological parameters	Non-onset day (N = 2993)	Onset day* (N = 658)	p value
Mean ambient temperature (°C)	14.9	11.35	0.0003
Minimum ambient temperature (°C)	10.5	7.2	0.0005
Maximum ambient temperature (°C)	19.6	15.85	0.0001
Dew point temperature (°C)	6.2	2.6	0.0013
Daily precipitation amount (mm)	1.5 (N = 1124)	1.1 (N = 264)	0.3462
Wind speed (m/s)	2.1	2.2	0.0431
Relative humidity (%)	62.4	60.8	0.5138
Atmospheric pressure (hPa)	1005.6	1006.7	0.0453
Change in mean temperature from the day before (°C)	0.2	0.2	0.2517
Change in minimum temperature from the day before (°C)	0	-0.2	0.0549
Change in maximum temperature from the day before (°C)	0.2	0.2	0.4068

The Wilcoxon-Mann-Whitney test was used for evaluations.

*Number of meteorological parameters for onset day was 658, because ICH occurred on the same day in 77 patients (735-77=658).

Note: All values are the median, and were measured at the Seoul Meteorological observatory from 2000 to 2009.

non-onset days. The Kruskal-Wallis test was used for comparison of meteorological variables between age groups by gender. Analyses were performed using SAS statistical software (SAS Institute Inc., Cary, NC, United States). A probability value less than 0.05 was considered statistically significant.

RESULTS

We analyzed 735 patients (400 men, 335 women) with a mean age of 62 ± 13 years. Clinical data of 735 enrolled patients are shown in Table 1. ICH was observed more frequently (observed/expected ratio ≥ 1) at lower mean, minimum, and maximum ambient temperature ($p = 0.0002$, 0.0003 , and 0.0002 , respectively). When the change in maximum temperature from the day before was ≤ -5 or $\geq 5^\circ\text{C}$, ICH was observed more frequently ($p = 0.0094$) (Table 2). Significantly lower mean, minimum, and maximum ambient temperature, lower dew point temperature, higher wind speed, and higher atmospheric pressure ($p = 0.0003$, 0.0005 , 0.0001 , 0.0013 , 0.0431 , and 0.0453 , respectively) were observed on days on which spontaneous ICH occurred (Table 3). In the subgroup analysis, the ICH onset day showed significant differences in mean, minimum, and maximum ambient temperature, dew point tem-

perature, relative humidity, and atmospheric pressure in the older (≥ 65 years) female group compared with other groups ($p = 0.0093$, 0.0077 , 0.0165 , 0.0028 , 0.0055 , and 0.0205 , respectively) (Table 4).

DISCUSSION

In this study, we found a significant association between lower ambient temperatures and the occurrence of ICH in a metropolitan city in South Korea. Other meteorological parameters, including dew point temperature, wind speeds, and atmospheric pressure also showed significant association with the occurrence of ICH.

The relevant literature provides inconsistent results for the relationship between ICH and ambient temperature. In a retrospective study, ICH tended to occur more frequently than expected at lower ambient temperatures.¹⁴ A multi-community cohort study conducted in Japan reported an association of low ambient temperature and high annual rainfall with increased risk of stroke incidence in women.¹⁰ A study conducted in Hong Kong showed a strong, negative linear association of daily temperature with hemorrhagic stroke admission.⁴ A population-based study conducted in northern Portugal found that the in-

Table 4. Distributions of meteorological factors between age groups by gender

	Male			Female		
	< 65 yrs (N = 259)	≥ 65yrs (N = 141)	<i>p</i> value	< 65 yrs (N = 152)	≥ 65yrs (N = 183)	<i>p</i> value
Mean ambient temperature (°C)	10.4	8.9	0.3163	14.15	9.9	0.0093
Minimum ambient temperature (°C)	6.3	5	0.3205	10.15	6	0.0077
Maximum ambient temperature (°C)	15.4	14	0.3161	19.1	14.1	0.0165
Dew point temperature (°C)	1.9	1.2	0.4403	7.2	0.9	0.0028
Daily precipitation amount (mm)	1.5	1.25	0.4493	1.5	0.7	0.4065
Wind speed (m/s)	2.2	2.2	0.4728	2.1	2.2	0.1663
Relative humidity (%)	60.1	59.4	0.7651	63.75	57.6	0.0055
Atmospheric pressure (hPa)	1007.4	1007.8	0.6864	1004.9	1007.9	0.0205
Change in mean temperature from the day before (°C)	0	0.2	0.5973	0.3	0.2	0.518
Change in minimum temperature from the day before (°C)	-0.3	-0.2	0.7285	0.1	-0.3	0.1755
Change in maximum temperature from the day before (°C)	-0.2	0.2	0.6074	0.35	0.5	0.7765

The Kruskal-Wallis test was used for evaluations.

All values are the median, and were measured at the Seoul meteorological observatory from 2000 to 2009.

idence of ICH increased by 11.8% for each degree drop in the ambient temperature on the preceding day.⁹⁾ This association is generally explained by the fact that low ambient temperature induces elevation of blood pressure by the constriction of peripheral vessel and changes in blood viscosity, platelet count, and catecholamine secretion.¹⁾⁸⁾ In contrast, Field et al.³⁾ reported that ICH occurrence did not show any association with changes in weather parameters. They concluded that these discrepancies might be explained by confounding factors such as socioeconomic conditions, emotional stress, genetic factors, and study design.

Our study also found an increase in ICH with higher atmospheric pressure. Consistent with our findings, a study from Spain showed that the incidence of ICH was related to the daily increases in atmospheric pressure.⁷⁾ Nakaguchi et al.¹²⁾ showed that the incidence of ICH tends to be high when atmospheric pressure is above the yearly mean value for two-three days preceding the day of onset, or that the pressure is high on the day of onset. They interpreted this finding by suggesting that peripheral vessels are compressed and constricted under high atmospheric pres-

sure causing high blood pressure and increased sympathetic tone. In addition, oxygen saturation can increase when engaging in excessive physical activity. Therefore, high atmospheric pressure is considered to increase the incidence of ICH by these direct and indirect effects. In 2012, Goggins et al.⁴⁾ found an increase in hemorrhagic stroke admissions with increased air pressure from the day before stroke onset. They emphasized the possibility that changes in atmospheric pressure could cause changes in pressure across the vessel wall and in endothelial function, leading to rupture.⁴⁾⁷⁾

One of the interesting results of our study is that ICH tends to occur more frequently on days with faster wind speed. In Seoul, the wind speed is usually faster during the spring and winter than during other seasons.¹³⁾ Therefore, low ambient temperature with faster wind speed may increase the occurrence of ICH in this moderate climate area.

In addition, we found that the older female group is more susceptible to ICH in lower ambient temperature, compared with other groups. Some investigators reported that seasonal and temperature variations were more noticeable in older patients.⁴⁾¹⁶⁾²⁰⁾

There is no clear explanation for this finding; however, Passero et al.¹⁶⁾ suggested that the elderly might have exaggerated physiological responses to cold weather, with greater increases in blood pressure, as shown in studies of myocardial infarction and ischemic stroke.¹⁵⁾¹⁸⁾ A hospital-based study found that the incidence of ICH was prominent in the older population, and hypothesized the possibility that older age is more susceptible to environmental changes.¹⁹⁾ A recent cohort study conducted in Japan reported that females were more susceptible to lower ambient temperature in stroke incidences than males independent of conventional risk factors.¹⁰⁾ There is no clear explanation for this finding, however, this appears to be caused by the difference of cold-induced vascular reaction according to gender.¹⁷⁾

Our study has certain limitations, apart from the inherent limitations of the retrospective design. First, this hospital-based study may have underestimated the true incidence of ICH in the community. Second, the meteorological factors measured in one area could not represent the ambient temperature actually experienced by the ICH patient. Third, it would be difficult to know the exact time of ICH onset and the setting in which the hemorrhagic event began (i.e. indoor or outdoor, cold protection devices, and emotional stress). Fourth, socioeconomic factor of the patients may contribute to the occurrence of ICH in a metropolitan city. In this study, we did not investigate this factor. These limitations may have confounded the study results.

CONCLUSION

The occurrence of spontaneous ICH is closely associated with meteorological factors, and the older females are more susceptible to lower ambient temperature. Further studies with multi-center data may be helpful in elucidating the risk of the occurrence of ICH and promote public health in a community.

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