# Risk-prediction model for acute myocardial infarction using atmospheric pressure data

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Prediction of high-risk days for onset of acute myocardial infarction (AMI) would aid prevention by taking specific actions on such days. To construct a predictive model, we investigated meteorological conditions related to high risk, with particular attention to atmospheric pressure.

The data used were records of conveyance by ambulances in the city of Hiroshima from January 1993 to December 2002, and corresponding meteorological data in the area. We used a Poisson regression model and several variables representing different critical conditions of atmospheric pressure decline. Finally, we selected the best model according to Akaike's Information Criterion (AIC).

A prediction model using a continuous variable of daily mean atmospheric temperature was established as the baseline model. Among models using different variables, one using weather pattern variables achieved the lowest AIC, showing it to be the best choice. In this model, strong winter patterns on the previous day were correlated with high risk of AMI.

The following meteorological factors were particularly related to high AMI risk: 1) A weather chart showing a strong winter pattern on the previous day, and 2) a decline in atmospheric pressure  $\geq 16$  hPa in addition to low atmospheric temperature. Such a strong winter pattern is easy to use and will improve performance of the Hiroshima prefectural AMI alert system.

## Introduction

It has been reported that meteorological conditions are environmental risk factors for acute myocardial infarction (AMI)<sup>1,2)</sup>. Human physical conditions are maintained relatively constant, independent of surrounding environmental changes, because of "homeostasis"<sup>3)</sup>. This property may fail if the magnitude of environmental change exceeds the limit of controllability. Diseases affected by changes in daily weather conditions are called meteorotropic. AMI is one such disease<sup>4)</sup>.

Japan is in a temperate zone and has four distinctive seasons. Thus, meteorological conditions have great seasonal change. Low atmospheric pressure systems frequently traverse the country and residents must adapt to their changes<sup>5)</sup>. Therefore, meteorotropic diseases are of major concern in Japan. AMI is one of the most important meteorotropic diseases, because of its frequency and lethality.

There are many studies on the relationship between AMI onset and meteorological conditions. However, these studies mainly address the relationship with seasonal variation<sup>6-9)</sup> or temperature<sup>10-15)</sup>. There has been very little investigation of other meteorological conditions. Some studies that considered the effect of atmospheric pressure suggest that mean daily atmospheric pressure does not have a significant effect on the AMI onset<sup>14, 16-18)</sup>. In contrast, other works indicate that AMI is frequent with a rapid decline of atmospheric pressure<sup>1,19-21)</sup>. However, there has been no clear evidence of the effect of atmospheric pressure.

http://home.hiroshima-u.ac.ip/hews

<sup>・</sup>気圧を考慮した急性心筋梗塞リスクの予測モデル

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In Hiroshima Prefecture, warnings of high-risk days of AMI are announced via the mass media<sup>22)</sup>. An alert for stroke is now provided in the same way (Fig. 1). The AMI warning has three risk levels, namely, "caution," "attention," and "small risk." "Caution" carries the highest risk, the criterion for which is that average temperature and pressure are low (less than  $6^{\circ}$ C and and 1013 hPa, respectively)<sup>23)</sup>. The present criterion does not use information on decline of atmospheric pressure, leaving room for improvement. The purpose of the present study is to clarify the impact of pressure decline on AMI incidence, for improving the AMI alert system.



Fig. 1. AMI and stroke alert presented on home page of Hiroshima Medical Association.

# Materials and Methods

#### Materials

The data used were records of conveyance by ambulances in the city of Hiroshima during the 10-year period from January 1993 to December 2002, along with corresponding meteorological data in the area. The ambulance data were provided by the Hiroshima Municipal Fire Department. The data were completely anonymous and included no personal information, so no ethical review was required. The data contained the date and time for each conveyance. Daily numbers of AMI incidence were calculated from the data. AMI for each patient was diagnosed by physicians in the emergency department to which patients were conveyed. AMI diagnosis does not include similar but distinct cardiac diseases such as heart failure. The data were collected independent of any history of heart failure. Meteorological data were provided by the Japan Weather Association. The data included daily mean atmospheric temperature, daily mean atmospheric pressure, weather type, hourly atmospheric temperature and pressure, relative humidity, rainfall, and snowfall.

# Model construction

# Probability distributions of AMI incidence and model selection

When a response variable in the model is count data like numbers of occurrence, the variable has nonnegative integer values. Therefore, it is inappropriate to assume that error has a normal distribution, as in an ordinary multiple regression model. We used a Poisson regression model, assuming that the response variable has a Poisson distribution, for which the logarithmic transformation of the Poisson parameter (mean value,  $\lambda$ ) is expressed by linear combination of explanatory variables.

The risk prediction model for AMI is

 $log (\lambda) = \beta_0 + \beta_1 X_1 + \beta_2 X_2 + \beta_3 X_3 \cdots$  $\lambda : Daily average events of AMI$  $X_1 : Atmospheric pressure$ 

 $X_2$ : Atmospheric temperature

 $X_3$ : Other meteorological condition

Coefficients of the linear model are estimated by the maximum likelihood method. To evaluate the aptness of the Poisson distribution, we examined observed and predicted values from January and June separately, when daily average AMI events were greatest or smallest, respectively.

#### Criterion of decline in atmospheric pressure

Decline in daily atmospheric pressure may take various forms; there is considerable variation of steepness and duration. We need a criterion for defining the decline, to detect the specific type related to AMI occurrence<sup>15,19,20,24-26)</sup>. Consequently, we created five types (Type 1 - Type 5) of variables defining specific pressure declines (Table 1). Next, we analyzed the relationship between AMI onset and these decline variables using the Poisson regression models. We explored which type of declines are most related to the

Table <sup>-</sup>	<ol> <li>Definition of</li> </ol>	variables	used in	models

Variable		Туре	Definition of values
Response variable			
AMI	Events of AMI	discrete (count)	daily average events of AMI
Explanatory variables:	temperature and pressure		
Т	Daily mean atmospheric temperature	daily mean atmospheric temperature-daily mean atmospheric temperature in ten year (= 16.5°C)	
		discrete (trinary)	<ol> <li>low atmospheric temperature group (T &lt; 10°C)</li> <li>moderate atmospheric temperature group (10 ≤ T &lt; 20°C)</li> <li>high atmospheric temperature group (20°C ≤ T)</li> </ol>
		discrete (binary)	1: low atmospheric temperature group $(T < 10^{\circ}C)$ 2: moderate or high atmospheric temperature group $(10^{\circ}C \le T)$
Р	Daily mean atmospheric pressure	continuous	daily mean atmospheric pressure-daily mean atmospheric pressure in ten years $({=}1008.9\mathrm{hPa})$
		discrete (trinary)	<ol> <li>low of atmospheric pressure group (P &lt; 1006hPa)</li> <li>moderate of atmospheric pressure (1006 ≤ P &lt; 1013hPa)</li> <li>high of atmospheric pressure group (1013hPa ≤ P)</li> </ol>
Explanatory variables:	decline of atmospheric press	ure	
Type 1 (=the sum of t	the hourly declines in a 24-hou	ır period)	
dec	The decline of atmospheric pressure	continuous	the sum of differeces if pressure values were decreasing (hPa)
dec11, dec12, dec13, dec14, dec15, dec16		discrete (binary)	0: dec < 11hPa; 1: 11hPa ≤ dec (dec11), 0: dec < 12hPa; 1: 12hPa ≤ dec (dec12) 0: dec < 13hPa; 1: 13hPa ≤ dec (dec13), 0: dec < 14hPa; 1: 14hPa ≤ dec (dec14) 0: dec < 15hPa; 1: 15hPa ≤ dec (dec15), 0: dec < 16hPa; 1: 16hPa ≤ dec (dec16)
Type 2 (= the variable	es representing 21 different w	eather patterns)	
tnk1-tnk21 (lagtnk1-lagtnk21)	Weather pattern (previous day, the day)	discrete (binary)	tnk=1 (if weather pattern $=1),=0$ (otherwise) and so on.
Type 3 (= the rate of	change per hour from maximu	ım atmospheric p	pressure to minimum atmospheric pressure within 24 hours)
slope	The rate of change per hour of the atmospheric pressure $^{a \lambda}$	continuous	the rate of change per hour of the atmospheric pressure $\left(h\mathrm{Pa}\right)$
slope3, slope4, slope5, slope6, slope7, slope8, slope9		discrete (binary)	$\begin{array}{l} 0: \mbox{ slope < 3hPa; 1: 3hPa \le slope (slope3), 0: \mbox{ slope < 4hPa; 1: 4hPa \le slope (slope4)} \\ 0: \mbox{ slope < 5hPa; 1: 5hPa \le slope (slope4), 0: \mbox{ slope < 6hPa; 1: 6hPa \le slope (slope6)} \\ 0: \mbox{ slope < 7hPa; 1: 7hPa \le slope (slope7), 0: \mbox{ slope < 8hPa; 1: 8hPa \le slope (slope8)} \\ 0: \mbox{ slope < 9hPa; 1: 9hPa \le slope (slope9)} \end{array}$
Type 4 (= binary varia	ables representing whether or	not continuous o	decline of atmospheric pressure occurred between 0-6, 0-8, 0-12, 0-18, or 0-24)
deck1-deck5 (lagdeck1-lagdeck5)	Continued atmospheric pressure decline	discrete (binary)	$0 \ (\mathrm{if} \ \mathrm{no})  ,  1 \ (\mathrm{if} \ \mathrm{yes}) \ \mathrm{between} \ 0 \ \mathrm{a.m.} \ \mathrm{to} \ 6 \ \mathrm{a.m.} \ \mathrm{for} \ \mathrm{deck1} \ \mathrm{and} \ \mathrm{so} \ \mathrm{on}.$
Type 5 (= variables r	epresenting the sun of decline	of atmospheric	pressure within a definite time intervals)
lagdec 1-5 daydec 1-5 leaddec 1-5	Atmospheric pressure decline of definite period of time interval (previous day, day, next day)	continuous	time intervals: 0-3, 3-6, 6-9, 9-12, 12-15, 15-18, 18-21, 21-0 for lagdeck1 (hPa) time intervals: 0-4, 4-8, 8-12, 12-16, 16-20, 20-0 for lagdeck2 (hPa) time intervals: 0-6, 6-12, 12-18, 18-0 for lagdeck3 (hPa) time intervals: 0-8, 8-16, 16-0 for lagdeck4 (hPa) time intervals: 0-12, 12-0 for lagdeck5 (hPa) and so on.
a)			difference of pressure in 24 hours
The rate of change per hour of the atmospheric = -		<sup>heric</sup> = -	(= maximum pressure - minimum pressure)
		(:	continued time of pressure decline = observation time of maximum pressure – observation time of minimum pressure)

onset. The decline variables of atmospheric pressure were selected in the model when the risk ratio of onset was larger than 1.0 and p < 0.10.

#### Selection of the optimal model

The selection of the models was performed based on the Akaike Information Criterion (AIC)<sup>27)</sup>. AIC can evaluate goodness of fit of a model with consideration of its complexity. Models with smaller AIC values are deemed superior to others. We established the baseline model as one containing only temperature as an explanatory variable, because temperature is known as the most influential factor in AMI onset. Next, we comprehensively evaluated goodness of fit for candidate models with various atmospheric pressure declines, and determined which had optimum goodness of fit using AIC. Finally, we checked model consistency using fundamental knowledge of AMI risk factors. We performed statistical analyses using PASW Statistics 18 (SPSS Japan Inc.). We considered a P-value less than 0.05 as statistically significant.

# Results

#### Descriptive statistics of data used

The study period spanned 3652 days, during which 3755 AMI events occurred on 2312 days (63.33% of total days), with an average of 1.03 events per day (range 0-6).

The monthly distribution of AMI events and meteorological conditions from 1993 to 2002 in Hiroshima are shown in Fig. 2. There were more daily events with statistical significance from October through April, relative to those in June. Daily mean temperature was highest in August ( $28.4 \pm 1.8^{\circ}$ C) and lowest in January ( $5.6 \pm 2.3^{\circ}$ C). Daily mean atmospheric pressure was highest in December ( $1016.0 \pm 4.4$  hPa) and lowest in July ( $1001.9 \pm 3.9$  hPa). These daily means had strong negative correlation (r = -0.676; p < 0.001). There were only a few days in winter when the daily mean atmospheric pressure was lower than that in summer. In contrast, the sum of all hourly atmospheric pressure declines in a 24-hour period was largest in March ( $5.4 \pm 3.6$  hPa) and smallest in July ( $3.4 \pm 1.7$  hPa).

#### Fit of observed AMI incidence to Poisson distribution

We statistically tested whether the observed values at AMI onset in January and June would fit the Poisson distribution. Figure 3 shows that those values conformed to the distribution (daily incidence was 1.24 in January and 0.86 in June).

#### Atmospheric pressure decline affecting AMI onset

Among the variables of atmospheric pressure decline, Type 1 sum of hourly declines in a 24-hour period) was selected as influential if its value was  $\geq 16$ hPa (risk ratio = 1.44 (p = 0.05), 95% CI = (0.99, 2.08)).

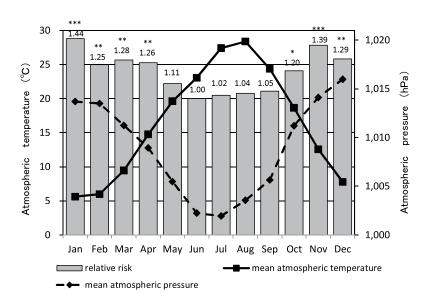


Fig. 2. Relative risk of AMI incidence by month, with monthly variation of atmospheric temperature and pressure. Relative risk (risk ratio) was estimated by Poisson regression model (reference was set as June).
\*\*\*\*p < 0.001, \*\*p < 0.01, \*p < 0.05</p>

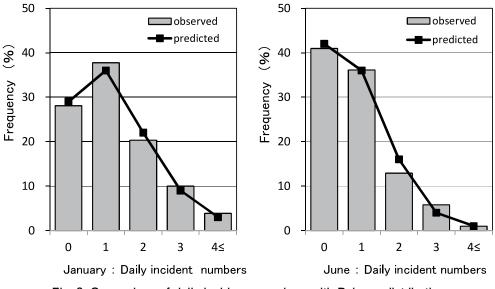


Fig. 3. Comparison of daily incidence numbers with Poisson distribution. Daily incidence was 1.24 in January and 0.86 in June.

Among the 21 variables for Type 2 (those representing 21 different weather patterns), only one, representing the strong winter type (strong western high and eastern low) of the previous day was statistically significant (risk ratio = 1.32 (p < 0.01), 95% CI = (1.08, 1.61)). Weather charts depicting strong winter patterns, in which there is low pressure moving east of the Japanese islands and continental high pressure emerging from the west, cause high pressure in the west and low in the east. Neither

Type 3 (the rate of change per hour from maximum to minimum pressure within a 24-hour period), Type 4 (binary variables representing whether there was continuous decline of pressure in the periods 0-6, 0-8, 0-12, 0-18, or 0-24 hours), or Type 5 (variables representing the sum of decline of pressure within a definite period, i.e., 3, 4, 6, 8, 12 hours intervals) variables contributed much to AMI onset.

Variable	Coefficient	Risk	ratio (95%CI)	p-value	AIC	ΔAIC
Baseline model						
Daily mean atmospheric temperature (continuous variable)	-0.013	0.99	(0.98 - 0.99)	<0.001***	9694.76	(-)
Model 1						
Daily mean atmospheric temperature (continuous variable)	-0.013	0.99	(0.98 - 0.99)	<0.001***	9690.31	-4.45
Strong winter pattern on the previous day	0.153	1.17	(1.00 - 1.35)	$0.045^{*}$		
High pressure pattern on the previous day	-0.073	0.96	(0.86 - 1.01)	0.069		
Rear of strong low pressure pattern on the previous day	-0.398	0.67	(0.39 - 1.16)	0.152		
Model 2						
Daily mean atmospheric temperature (continuous variable)	-0.013	0.99	(0.98 - 0.99)	<0.001***	9693.32	-1.44
low atmospheric temperature group ★decline in atmospheric pressure (16hPa≤)	0.488	1.63	(1.01 - 2.63)	0.024*		

Table 2. Parameter estimate of baseline model and optimal models

Relative risk (risk ratio) was estimated by Poisson regression model

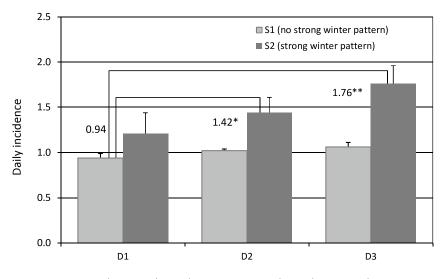
\*\*\*\*p < 0.001, \*p < 0.05

 $\Delta\,{\rm AIC}:$  baseline AIC – optimal model AIC

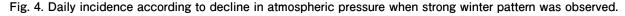
#### Selection of optimal model

We first investigated how to construct the best baseline model, by considering which was more suitable for use with atmospheric temperature and pressure as continuous or discrete variables (Table 1). As shown in Table 2, the best baseline model was determined as that using only the continuous variable of daily mean temperature (AIC = 9694.76). Next, as shown in Table 2, we created candidate models using variables of pressure decline from Type 1 to Type 5, and compared them with the baseline model according to AIC. Model 1, which uses the variable of Type 2 (different weather patterns), achieved the lowest AIC (=9690.31, baseline AIC -4.45) and was therefore the best. In this model, the variable representing a strong winter weather pattern of the previous day most strongly affected the risk of AMI. Over the 10-year period, there were 143 days with such a weather pattern. The relationship between this pattern and pressure decline was further investigated. We found that more AMI patients were transported by ambulance on days with strong winter weather patterns and greater pressure declines (Fig. 4).

Another remarkable model that had the second lowest AIC was one using variables of atmospheric pressure decline within a 24-hour period, daily mean temperature, and an interaction term of low temperature and  $\geq$  16 hPa decrease in pressure (model 2 in Table 2). This model predicted a "super-risky" day for AMI (risk ratio = 2.10 (p < 0.01), 95% CI = (1.30, 3.38)) (Fig. 5). There were only 9 such days in 10 years, and these were between December and March. The probability of AMI occurrence three or more times per day was 55%. The probability was 64% after 18:00, while AMI usually takes place more frequently in the morning. We also assessed weather characteristics on these super-risky days. On such days, weather changed suddenly from the previous evening. In most cases, it changed to rain and sleet all day long and was likely to continue into the following day. In addition, there were rapid declines in atmospheric pressure from the previous night (18.7  $\pm$ 2.4 hPa decline within 24 hours). Temperature was low  $(6.9 \pm 2.2$ °C daily mean) and relative humidity high (81.7  $\pm$  4.2% daily mean).



 $D1(D \le hPa)$ ,  $D2(5hPa \le D \le 10hPa)$ ,  $D3(10hPa \le D)$ 



#### \*\*p < 0.01, \*p < 0.05

D1, D2 and D3 show low, moderate and high decline groups of atmospheric pressure, respectively. Poisson regression model was used to test interaction between various meteorological groups with strong winter pattern groups. D1 and no strong winter pattern groups were set as reference.

Daily events corresponding to moderate declines of atmospheric pressure groups (D2) with strong winter pattern groups were significantly greater than those with small declines of atmospheric pressure groups (D1) with no strong winter pattern groups (p < 0.05).

Daily events corresponding to strong declines in atmospheric pressure groups (D3) with strong winter pattern groups were significantly greater than those with small declines of atmospheric pressure groups (D1) with no strong winter pattern groups (risk ratio = 1.87 = 1.76/0.94 (p < 0.01), 95% CI = (1.29, 2.71)).

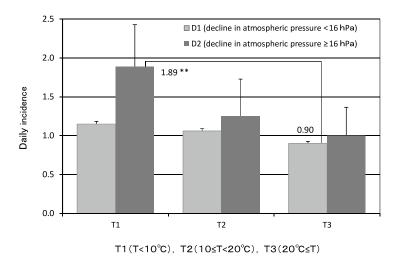


Fig. 5. Daily incidence according to atmospheric pressure decline and temperature.  $**_p < 0.01$ 

T1, T2 and T3 show low, moderate, and high atmospheric temperature groups. Poisson regression model was used to test interaction between various meteorological groups with  $\geq 16$  hPa atmospheric pressure decline. T3 and < 16 hPa decline in atmospheric pressure groups were set as reference.

Daily events in low atmospheric temperature groups with  $\geq 16$  hPa atmospheric pressure decline were significantly greater than those in high atmospheric temperature groups with  $\leq 16$  hPa decline in atmospheric pressure groups (risk ratio = 2.10 = 1.89/0.90 (p < 0.01), 95% CI = (1.30, 3.38)).

# Discussion

Prediction models were obtained for meteorological conditions related to high AMI risk, particularly atmospheric pressure. Advantages of the model and analysis limitations are discussed below.

# Contribution of meteorological conditions to high risk of AMI onset

We found that a strong winter weather pattern was one of the most influential factors for AMI occurrence. When the weather chart shows such a pattern, it is often cloudy and snowy in the region fronting the Sea of Japan, and daylight hours are short. In contrast, on the opposite side of the mountains where Hiroshima is located, cold and dry winds blow frequently on days with fine weather<sup>28)</sup>. This indicates that snow and rain are rarely observed there. The Siberian high-pressure system that strongly affects area weather fluctuates in strength within almost a week, reducing the pressure. It has been reported that pressure declines more than 10 hPa are likely on such days, with "high in the west and low in the east"<sup>29)</sup>. Rapid drops in pressure are reported to reduce autonomic control function for maintaining physical balance, by exceeding the limit of homeostatic regulation<sup>3)</sup>. It rarely snows or rains in Hiroshima during winter but, if it does, it is considered a day with

a strong winter weather pattern.

We also discovered that the AMI risk significantly increased in cases of low temperature and a pressure decline  $\geq$  16 hPa in a 24-hour period. Previous studies of daily incidence showed that on days of low temperature and pressure, there were 37% more cases of AMI than on days of high temperature and moderate pressure<sup>15,26)</sup>. Sama et al.<sup>25)</sup> showed that in Finland, AMI increased because of conditions of low temperature and low pressure. This study showed a super-risky day of AMI with a pressure decrease  $\geq$  16 hPa combined with low temperature. This result supported earlier studies and revealed more concrete meteorological conditions related to high risk. Super-risky days of AMI were all from December through March, a season with the largest pressure difference in a 24-hour period <sup>29)</sup>. Previous studies showed that all days with low temperature and low pressure had rainfall or accumulating snow, accompanied by low temperature and high relative humidity. This condition corresponded to the superrisky days. In addition to these features, it was shown that weather suddenly deteriorated from the evening of the previous day<sup>15,25)</sup>.

The present study showed that AMI is frequent under conditions of low temperature and rapid drop in pressure. However, it is not clear how such meteorological conditions affect AMI onset. One study reported that histamine or a similar substance in the body is released during decompression accompanying the approach of low atmospheric pressure. This triggers meteorotropic diseases because histamine can increase fluid retention within the body, contraction of smooth muscles, permeability of blood vessels, and inflammatory reaction<sup>30)</sup>. Sato<sup>31)</sup> supported this by experimentation with rats. That study showed that rats exhibited increased blood pressure and heart rate, indicating excitement of the sympathetic nervous system while acting freely under declining atmospheric temperature and pressure, which is often experienced in everyday life. Houck et al.<sup>19)</sup> reported that the strong influence of atmospheric pressure change damages coronary arteries by formation of plaques. In an alpine environment of low atmospheric pressure, it has been reported that the functions of breathing and blood circulation change rapidly because of oxygen and carbon dioxide partial pressures in the lung, leading to altitude sickness. However, the magnitude of this effect is unclear for changes in atmospheric pressure commonly observed near sea level<sup>28)</sup>. It has been shown that even a slight change of pressure may influence the condition of the human body<sup>32,33)</sup>.

Thus, the effects of atmospheric pressure decline are wide ranging. They involve direct and indirect impacts on the living body, and work independently or in conjunction with other meteorological conditions such as temperature decrease. The finding that low temperature and rapid pressure declines are important risks for AMI onset is undeniable, although the mechanism is unclear. Further study is required.

#### Usefulness of prediction model

AMI onset is related to multiple factors, beyond that of atmospheric pressure<sup>34,35)</sup>. Temperature change is another important risk factor for AMI onset<sup>1, 36)</sup>. However, we focused only on pressure, leaving temperature to be analyzed in the future.

The prediction models analyzed herein involve weather patterns of the previous day and pressure decline to predict the risk of AMI onset. Various criteria for defining the decline were evaluated. As a result, we found that the best model was that using a weather chart indicative of a strong winter pattern on the previous day, or a decrease in pressure  $\geq 16$  hPa. Using these variables considerably improved the goodness of fit. We must also consider the availability of predictive variables when deciding the AMI alert level. Given this consideration, it is easy to use the strong winter pattern of the weather chart. In contrast, it is difficult to use atmospheric pressure decline over a 24-hour period, because this information is unavailable at the time of alert issuance. Use of the strong winter pattern should therefore be convenient and useful toward improving the performance of practical criteria within the alert system.

# Conclusion

The goodness of fit was considerably improved by use of the weather chart showing a strong winter pattern of the previous day, or by a decrease of atmospheric pressure  $\geq 16$  hPa in combination with low temperature. The former usage is considered easy to execute. This suggests that the model would be useful for improving the present Hiroshima prefectural alert system.

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