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# Characteristics and Prognosis of Hospitalized Patients at High Risk of Deterioration Identified by the Rapid Response System: a Multicenter Cohort Study

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The authors have no potential conflicts of interest to disclose.

#### **Authors Contributions**

Conceptualization: Kim SH. Formal analysis: Kim SH. Supervision: Kim Y. Validation: Kim

# ABSTRACT

We aimed to investigate the characteristics and prognosis of high risk hospitalized patients identified by the rapid response system (RRS). A multicentered retrospective cohort study was conducted from June 2019 to December 2020. The National Early Warning Score (NEWS) was used for RRS activation. The outcome was unexpected intensive care unit (ICU) admission within 24 hours after RRS activation. The 11,459 patients with RRS activations were included. We found distinct clinical characteristics in patients who underwent ICU admission. All NEWS parameters were associated with the risk of unexpected ICU admission except body temperature. Body mass index, pulmonary disease, and cancer are related to the decreased risk of unexpected ICU admission. In conclusion, there were differences in clinical characteristics among high risk patients, and those differences were associated with unexpected ICU admissions. Clinicians should consider factors relating to unexpected ICU admission in the management of high risk patients identified by RRS.

Keywords: Early Warning Score; Risk Management; Hospital Mortality

Despite the frequent occurrence of an unexpected deterioration in hospitalized patients, it was not monitored adequately due to insufficient human resources and equipment.<sup>1,2</sup> A concept of the rapid response system (RRS) for the early identification of hospitalized patients expected to experience serious adverse events emerged in the 2000s.<sup>3,4</sup> According to a growing body of evidence, the RRS reduced serious adverse events such as unexpected intensive care unit (ICU) admission and cardiac arrest.<sup>5-7</sup> Following the current trend, a RRS pilot program including more than 40 hospitals has been conducted in South Korea since May 2019.<sup>8</sup> However, few studies reported the results of the RRS pilot program. In addition, the clinical features of high-risk patients identified by the RRS are rarely studied. Therefore, this study aimed to investigate the characteristics and prognosis of hospitalized patients at high risk of deterioration identified by the RRS.

A multicenter-based, retrospective, consecutive cohort study from June 2019 to December 2020 was conducted at five hospitals in Hallym Medical Center. Patients were classified

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as high risk with the National early warning score (NEWS) over seven regardless of the single item score. After RRS activation, rapid response team members were immediately summoned to assess the patient's condition. Also, they prescribed appropriate medications or escalated the level of care based on the clinical assessment. When the RRS was activated at different times in the same patient, we considered those activations as separate cases.

Of the 18,131 patients identified by RRS activations, we excluded 480 identified after the ICU admission. We further excluded 6,154 with missing data and 38 with incorrectly recorded NEWS scores. Finally, 11,459 patients with RRS activations were included in the analysis. Comorbidities were defined using the 10<sup>th</sup> International Classification of Disease codes and consisted of cardiovascular disease, pulmonary disease, gastrointestinal disease, cerebrovascular disease, genitourinary disease, and cancer.<sup>9</sup> The outcome of this study was unexpected ICU admission within 24 hours after the RRS activation. Data were described as medians (interquartile ranges) for continuous variables and numbers (percentages) for categorical variables. In high-risk patients, factors relating to unexpected ICU admission were further evaluated using the multivariable logistic regression analysis. All factors were included in the analysis. Statistical analyses were performed using R 4.0.3 (R Foundation for Statistical Computing, Vienna, Austria).

Clinical characteristics of the RRS activation events were described in **Table 1**. Of 11,459 patients, the incidence of unexpected ICU admission was 7.4%, and the mean RRS activation was 3.1 times per patient. Patients who underwent unexpected ICU admission were heavier (22.2 [19.5–25.1] vs. 21.4 [18.5–24.2] kg/m<sup>2</sup>, *P* < 0.001) and had higher total NEWS score (9 [8–10] vs. 8 [7–9] kg/m<sup>2</sup>, *P* < 0.001) than patients who did not undergo unexpected ICU admission.

Table 1. Clinical characteristics of	f patients identified b	by the rapid response system
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Variables	Total (n = 11,459)	Cases who did not undergo unexpected ICU admission (n = 10,614)	Cases who underwent unexpected ICU admission (n = 845)	P value
Age, yr	74 (64-81)	74 (64–81)	74 (64–81)	0.522
Male	7,061 (61.6)	6,551 (61.7)	510 (60.4)	0.454
Body mass index, kg/m <sup>2</sup>	21.5 (18.6-24.2)	21.4 (18.5–24.2)	22.2 (19.5-25.1)	< 0.001
NEWS parameter				
Systolic blood pressure, mmHg	100 (100–115)	100 (100–110)	103 (90–130)	0.051
Heart rate, rates/min	103 (93–116)	102 (92–116)	113 (98–128)	< 0.001
Respiratory rate, rates/min	22 (21–26)	22 (21–26)	25 (22–30)	< 0.001
Body temperature, °C	36.9 (36.6-37.6)	36.9 (36.6-37.5)	37.1 (36.6–37.8)	< 0.001
Oxygen saturation, %	94 (91–96)	95 (91–96)	91 (87–95)	< 0.001
Need for supplemental oxygen	10,131 (88.4)	9,463 (89.2)	668 (79.1)	< 0.001
Altered mental status	23 (0.2)	16 (0.2)	7 (0.8)	< 0.001
NEWS parameter score				
Systolic blood pressure	2 (0-2)	2 (1–2)	2 (0-3)	0.064
Heart rate	1 (1–2)	1 (1–2)	2 (1–2)	< 0.001
Respiratory rate	2 (2-3)	2 (2-3)	3 (2-3)	< 0.001
Body temperature	0 (0)	0 (0)	0 (0-1)	0.003
Oxygen saturation	1 (0-3)	1 (0-3)	3 (1–3)	< 0.001
NEWS total score	8 (7-9)	8 (7-9)	9 (8–10)	< 0.001
Etiology				
Pulmonary disease	5,492 (47.9)	5,186 (48.9)	306 (36.2)	< 0.001
Gastrointestinal disease	979 (8.5)	893 (8.4)	86 (10.2)	0.089
Genitourinary disease	476 (4.2)	420 (4.0)	56 (6.6)	< 0.001
Cerebrovascular disease	262 (2.3)	228 (2.1)	34 (4.0)	< 0.001
Cancer	3,374 (29.4)	3,218 (30.3)	156 (18.5)	< 0.001

Data were expressed as medians (interquartile ranges) for continuous variables and numbers (percentages) for categorical variables. When the rapid response system was activated at different times in the same patient, we considered this as a separate case.

ICU = intensive care unit, NEWS = national early warning score.

Factors	OR (95% CI, <i>P</i> )	Outcome: unexpected ICU admission
Age	1.00 (0.99–1.00, 0.235)	•
Male	1.07 (0.92-1.25, 0.372)	⊢,,
Body mass index	1.02 (1.01-1.03, 0.002)	•
NEWS scores		
Systolic blood pressure	1.16 (1.09–1.24, < 0.001)	· · · · · · · · · · · · · · · · · · ·
Heart rate	1.62 (1.49–1.76, < 0.001)	HEH
Respiratory rate	1.42 (1.31–1.53, < 0.001)	HEH
Body temperature	0.99 (0.88-1.12, 0.907)	H
Oxygen saturation	1.59 (1.49–1.70, < 0.001)	HEH
Oxygen supplement	0.90 (0.81-0.99, 0.034)	HEH
Level of consciousness	2.20 (1.57-2.99, < 0.001)	⊢ <b>∎</b>
Comorbidities		
Pulmonary disease	0.59 (0.50-0.70, < 0.001)	HEH
Gastrointestinal disease	1.15 (0.88–1.48, 0.309)	⊢┼═──┤
Genitourinary disease	1.19 (0.87-1.62, 0.261)	⊢ <b>∔ ∎</b> 1
Cerebrovascular disease	1.35 (0.90–1.98, 0.133)	
Cancer	0.39 (0.32-0.47, < 0.001)	H
		0 1 2
		OR (95% CI)

**Fig. 1.** Factors associated with unexpected intensive care unit admission among patients with a high risk of deterioration. The OR, 95% CI, and *P* value were calculated using multivariate logistic regression analysis. OR = odds ratio, CI = confidence interval, ICU = intensive care unit, NEWS = national early warning score.

Significantly different NEWS parameters were observed between two groups, except systolic blood pressure (P=0.051); heart rate (103 [90–130] vs. 100 [100–110] rates/min), respiratory rate (25 [22–30] vs. 22 [21–26]), body temperature (37.1 [36.6–37.8] vs. 36.9 [36.6–37.5]), oxygen saturation (91 [87–950] vs. 95 [91–96]), need for supplemental oxygen (79.1% vs. 89.2%), and altered mental status (0.8% vs. 0.2%). Among etiologies, there was a low proportion of pulmonary disease (36.2% vs. 48.9%), genitourinary disease (6.6% vs. 4.0%), cerebrovascular disease (4.0% vs. 2.1%), and cancer (19.8% vs. 39.2%) in patients who underwent unexpected ICU admission. This difference was also confirmed in the NEWS score.

In the analysis for exploring factors relating to unexpected ICU admission, body mass index, pulmonary disease, and cancer were associated with the decreased risk of unexpected ICU admission (**Fig. 1**). Of NEWS parameter scores, high scores of systolic blood pressure, heart rate, respiratory rate, oxygen saturation, and level of consciousness were related to the increased risk of unexpected ICU admission. However, a high oxygen supplement score was associated with the decreased risk of unexpected ICU admission.

We found distinct clinical characteristics in patients who underwent unexpected ICU admission among high-risk patients identified by RRS. All NEWS scores were associated with the unexpected ICU admission, except body temperature. Low body mass index, pulmonary disease, and cancer were associated with the decreased risk of unexpected ICU admission.

Among high-risk patients identified by the RRS, there were differences in clinical characteristics according to the experience of unexpected ICU admission. In particular, almost all NEWS parameters were worse in patients who underwent unexpected ICU admission. It means that even among detected high risk patients, there are more severe patients who required immediate management.<sup>10</sup> Our finding suggested that it will be possible to prioritize patient management in high risk patients identified by the RRS. Interestingly, patients who underwent unexpected ICU admission were heavier and had a low proportion of pulmonary disease and cancer. It may be because a large number of

patients with pulmonary disease and cancer were chronically ill. In these patients, the heart rate or respiratory rate was high because of their underlying conditions.<sup>11,12</sup> Also, they are malnourished and often require long-term oxygen therapy.<sup>13,14</sup> The discrimination ability of RRS for identifying high risk patients may be insufficient in chronic patients who had a high baseline early warning score. Therefore, the etiology of the patient must be considered in the RRS implementation.<sup>15</sup>

Most of the high NEWS scores were associated with the increased risk of unexpected ICU admission. Notably, systolic blood pressure showed no difference between those who underwent unexpected ICU admission and those who did not. However, systolic blood pressure was related to the increased risk of unexpected ICU admission. On the other hand, an inversed relationship was observed on the body temperature. The body temperature, which was higher in those who underwent unexpected ICU admission, did not show significance in exploring risk factors. These results indicate that systolic blood pressure is a more important parameter for predicting acute deterioration than body temperature in managing high-risk patients. A previous study has reported that adverse events can be accurately predicted using systolic blood pressure added to other indicators without body temperature.<sup>16,17</sup> Therefore, clinicians should cautiously interpret the details of NEWS parameters.

There are some limitations to this study. First, a causal relationship could not be confirmed because this is a retrospective study. Also, this study was conducted based on the retrospective review of the patient's records. Thus, some parameters, such as level of consciousness, might be inaccurate. Despite these limitations, we should address the strength of this study. First, to the best of our knowledge, this is the first study that reported characteristics and prognosis of the high-risk patients based on the results of the RRS pilot program in Korea. Also, our study can be used for prioritizing management among a large number of high-risk patients identified by RRS. In a resource-limited environment such as Korea, prioritizing patients who need immediate management will be of great help in the operation of RRS.<sup>18</sup>

In conclusion, there were differences in clinical characteristics among high risk patients, and those differences were associated with unexpected ICU admissions. Clinicians should consider factors relating to unexpected ICU admission in the management of patients identified by the RRS.

# **Ethics statement**

We conducted this retrospective cohort study in accordance with the Helsinki declaration. We obtained approval from the Institutional Review Board (IRB) of Hallym University Chuncheon Sacred Heart Hospital (IRB number: 2021-02-007). The need for written informed consent was waived because this was a retrospective cohort study.

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