Comparing the neurologic outcomes of patients with out-of-hospital cardiac arrest according to prehospital advanced airway management method and transport time interval

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Objective The incidences of prehospital advanced airway management by emergency medical technicians in South Korea are increasing; however, whether this procedure improves the survival outcomes of patients experiencing out-of-hospital cardiac arrest remains unclear. The present study aimed to investigate the association between prehospital advanced airway management and neurologic outcomes according to a transport time interval (TTI) using the Korean Cardiac Arrest Research Consortium database.

Methods We retrospectively analyzed the favorable database entries that were prospectively collected between October 2015 and December 2016. Patients aged 18 years or older who experienced cardiac arrest that was presumed to be of a medical etiology and that occurred prior to the arrival of emergency medical service personnel were included. The exposure variable was the type of prehospital airway management provided by emergency medical technicians. The primary endpoint was a favorable neurologic outcome provided by emergency medical technicians.

Results Of 1,871 patients who experienced out-of-hospital cardiac arrest, 785 (42.0%), 121 (6.5%), and 966 (51.6%) were managed with bag-valve-mask ventilation, endotracheal intubation (ETI), and supraglottic airway (SGA) devices, respectively. SGAs and ETI provided no advantage in terms of favorable neurologic outcome in patients with TTIs ≥ 12 minutes (odds ratio [OR], 1.37; confidence interval [CI], 0.65–2.87 for SGAs; OR, 1.31; CI, 0.30–5.81 for ETI) or in patients with TTI < 12 minutes (OR, 0.57; CI, 0.31–1.07 for SGAs; OR, 0.63; CI, 0.12–3.26 for ETI).

Conclusion Neither the prehospital use of SGA nor administration of ETI was associated with superior neurologic outcomes compared with bag-valve-mask ventilation.

Keywords Airway management; Emergency medical services; Intubation, intratracheal; Out-of-hospital cardiac arrest

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INTRODUCTION

During cardiopulmonary resuscitation (CPR) on a patient experiencing out-of-hospital cardiac arrest (OHCA), it is recommended that healthcare providers maintain bag-valve-mask (BVM) ventilation before performing advanced airway management. BVM ventilation has the disadvantage of producing complications such as aspiration and pneumonia due to gastric expansion or regurgitation. After the return of spontaneous circulation (ROSC), performing endotracheal intubation (ETI) elicits controlled and effective ventilation that is necessary while transporting the patient with OHCA to the emergency department. Advanced airway management including ETI or the use of supraglottic airway (SGA) devices can facilitate the performance of high-quality CPR without interrupting chest compressions (as would be required with BVM ventilation) and increases the rates of short-term survival, ROSC, and survival to admission. ETI, which prevents airway obstruction and facilitates airway management, is preferred by hospitals, although performing it requires high skill. Various SGAs such as the esophageal obturator airway, laryngeal mask, laryngeal tube, and I-gel device are being disseminated to assist healthcare providers to maintain open airways with minimal training.

With the increase in the proportion of level 1 emergency medical technicians (EMTs) among the 119 emergency medical service (EMS) agencies in South Korea, there has been an increasing trend toward performing advanced airway management, including ETI and SGA use, during the prehospital phase. However, the effects of prehospital airway management on the neurologic outcomes of patients experiencing OHCA remain unclear despite previous investigations. Previous studies showed that the direct transfer of OHCA patients to a percutaneous coronary intervention-capable center or a critical care center improved survival outcomes; as a result, strategies aimed at the regionalization of postresuscitation care for OHCA patients are emerging. A previous meta-analysis revealed that the transport time interval (TTI) was not associated with neurologic outcomes or survival to discharge among OHCA patients. Advanced airway management is associated with an improved no-flow ratio when performing CPR, and can reduce complications commonly associated with BVM ventilation such as pneumonia, gastric expansion, and regurgitation. Therefore, it is possible that prehospital advanced airway management may improve the outcomes of patients requiring extended transport times. Nevertheless, the association between prehospital airway management and survival outcomes according to TTI remains unclear. Additionally, there have been no studies in South Korea on the effects of prehospital airway management and ventilation methods on survival rates and neurologic outcomes according to TTI in patients experiencing OHCA.

Considering that prehospital airway management can minimize chest compression interruptions, provide effective ventilation, and reduce airway complications in patients with OHCA, we hypothesized that prehospital advanced airway management will have advantages over BVM ventilation with respect to neurologic outcomes, and that the beneficial effect of advanced airway management will increase with the length of the TTI. The present study aimed to investigate the association between the prehospital airway management method (BVM, ETI, or SGA) and neurologic outcomes in patients with OHCA using data from the Korean Cardiac Arrest Research Consortium (KoCARC) database.

METHODS

Study design, setting, and subjects
We retrospectively analyzed data from the KoCARC database that was recorded between October 2015 and December 2016. Patients aged 18 years or older who experienced cardiac arrest that was presumed to be of a medical etiology and that occurred prior to
the arrival of EMS personnel on the scene were included in this study. The KoCARC registry excluded OHCA patients with terminal illnesses documented in their medical records as well as those under hospice care, those who were pregnant, and those with “do not resuscitate” directives. Furthermore, we also excluded the following groups: patients with OHCA verifiably caused by nonmedical etiologies such as trauma, drowning, poisoning, burn, asphyxia, and hanging; patients who transferred from other hospitals and had insufficient prehospital time variables and/or unclear prehospital airway management methods; and patients who experienced cardiac arrest during transport to the hospital (whether in an ambulance or otherwise). In South Korea, if three or more EMTs are dispatched to the location of a patient with cardiac arrest, they are required to perform ETI or use SGAs while administering effective chest compressions in the field according to field treatment guidelines. This study was approved by the institutional review board of Kyungpook National University Hospital (2015-11-013-007). The requirement for informed consent was waived owing to the retrospective nature of this study.

Data source
The KoCARC registry is a multicenter nationwide network of 64 participating institutions for data collection and collaborative research from across South Korea. Data are collected according to standardized Utstein-style templates for OHCA to facilitate uniform reporting using precisely defined variables and outcomes. Data are error-checked prior to consolidation with the master dataset. The quality management committee provides feedback regarding quality management processes to the research coordinators and investigators.

Outcomes and variables
The exposure variable in this study was the type of prehospital airway management provided by EMTs. The subjects were divided into three groups according to the prehospital airway management method (BVM, ETI, or SGA); patient demographics, arrest characteristics, and survival outcomes were compared among the three groups.

Neurologic status was assessed using cerebral performance category scores, which are based on a five-point scale in which scores of 1 (good recovery) and 2 (moderate disability) indicated favorable neurologic outcome. The primary endpoint was a favorable neurologic outcome, whereas the secondary endpoint was survival to hospital discharge. We used the following a favorable registry core variables: (1) patient demographics (i.e., sex, age, medical history of hypertension, diabetes, and dyslipidemia); (2) community resuscitation (i.e., ‘witnessed’ status, bystander CPR, and location of cardiac arrest occurrence); (3) EMS resuscitation (i.e., initial electrocardiogram rhythm, prehospital defibrillation, prehospital epinephrine administration, prehospital ROSC, elapsed time from the initial call to ambulance arrival at the scene [response time interval], elapsed time from arrival at the scene to departure [scene time interval], and elapsed time from departure to arrival at the emergency department [TTI]); and (4) hospital resuscitation and postresuscitation care (i.e., performed extracorporeal membrane oxygenation, coronary angiography, and target temperature management).

Statistical analysis
The significance of differences between groups was tested using analysis of variance with post-hoc analysis (with multiple com-
Effect of prehospital advanced airway management

Table 1. Baseline characteristics and outcomes of the study population based on the prehospital airway management method

|                  | Total (n = 1,871) | Prehospital airway management | P-value |
|------------------|-------------------|-------------------------------|---------|
|                  | BVM (n = 785)     | SGA (n = 965)                | ETI (n = 121) |
| Sex              |                   |                               |          |
| Male             | 1,223 (65.4)      | 479 (61.0)                    | 673 (69.7) | 71 (58.7) | < 0.001 |
| Female           | 648 (34.6)        | 306 (39.0)                    | 292 (30.3) | 50 (41.3) |          |
| Age (yr)         |                   |                               |          |
| < 65             | 758 (40.5)        | 335 (42.7)                    | 391 (40.5) | 32 (26.4) | 0.003    |
| ≥ 65             | 1,113 (59.5)      | 450 (57.3)                    | 574 (59.5) | 89 (73.6) |          |
| Age (yr)         | 70 (56–78)        | 68 (55–78)                    | 70 (56–78) | 74 (63–81) | 0.026*   |
| RTI (min)        | 7 (6–10)          | 7 (5–10)                      | 8 (6–10)   | 7 (5–10)   | 0.655    |
| STI (min)        | 11 (8–17)         | 9 (6–13)                      | 14 (10–20) | 14 (10–20) | < 0.001* |
| TTI (min)        | 10 (6–14)         | 10 (7–14)                     | 9 (6–13)   | 10 (6–15)  | 0.003**  |
| Location         |                   |                               |          |
| Home             | 1,261 (67.4)      | 478 (60.9)                    | 689 (71.4) | 94 (77.7) | < 0.001 |
| Public place     | 419 (22.4)        | 218 (27.8)                    | 186 (19.3) | 15 (12.4) |          |
| Other/unknown    | 191 (10.2)        | 89 (11.3)                     | 90 (9.3)   | 12 (9.9)   |          |
| Comorbidities    |                   |                               |          |
| Hypertension     | 768 (41.0)        | 302 (38.5)                    | 406 (42.1) | 60 (49.6) | 0.045    |
| Diabetes mellitus| 482 (25.8)        | 178 (22.7)                    | 261 (27.0) | 43 (35.5) | 0.005    |
| Hyperlipidemia   | 96 (5.1)          | 48 (6.1)                      | 39 (4.0)   | 9 (7.4)   | 0.073    |
| Witnessed cardiac arrest | 1,112 (59.2) | 502 (64.0) | 536 (55.5) | 70 (57.9) | 0.002    |
| Bystander CPR    | 961 (51.4)        | 375 (47.8)                    | 524 (54.3) | 62 (51.2) | 0.025    |
| Initial shockable rhythm | 366 (19.6) | 138 (17.6) | 207 (21.5) | 21 (17.4) | 0.104    |
| Prehospital defibrillation | 483 (25.8) | 185 (23.6) | 271 (28.1) | 27 (22.3) | 0.066    |
| Prehospital epinephrine administration | 293 (15.7) | 38 (4.8) | 226 (23.4) | 29 (24.0) | < 0.001 |
| Prehospital ROSC | 242 (12.9)        | 91 (11.6)                     | 134 (13.9) | 17 (14.0) | 0.339    |
| Hospital treatment |                |                               |          |
| CAG              | 204 (10.9)        | 80 (10.2)                     | 108 (11.2) | 16 (13.2) | 0.559    |
| TTM              | 143 (7.6)         | 56 (7.1)                      | 73 (7.6)   | 14 (11.6) | 0.230    |
| ECMO             | 46 (2.5)          | 19 (2.4)                      | 19 (2.0)   | 8 (6.6)   | 0.008    |
| Survival to discharge |          |                               |          |
| Yes              | 192 (10.3)        | 93 (11.8)                     | 87 (8.9)   | 13 (10.7) | 0.130    |
| No               | 1,679 (89.7)      | 692 (88.2)                    | 888 (91.1) | 109 (89.3)|          |
| Good neurologic outcome |        |                               |          |
| Yes              | 130 (6.9)         | 68 (8.7)                      | 56 (5.8)   | 6 (5.0)   | 0.044    |
| No               | 1,741 (93.1)      | 717 (91.3)                    | 909 (94.2) | 115 (95.0)|          |

Values are presented as number (%) or median (interquartile range).

BVM, bag-valve-mask; SGA, supraglottic airway; ETI, endotracheal intubation; RTI, response time interval; STI, scene time interval; TTI, transport time interval; CPR, cardiopulmonary resuscitation; ROSC, return of spontaneous circulation; CAG, coronary angiography; TTM, targeted temperature management; ECMO, extracorporeal membrane oxygenation.

*Differences were significant in post-hoc analysis of a one-way ANOVA test including BMV vs. ETI, and SGA vs. ETI. **Differences were significant in post-hoc analysis of a one-way ANOVA test including BMV vs. SGA, and BMV vs. ETI. ***Differences were significant in post-hoc analysis of a one-way ANOVA test including BMV vs. SGA.
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rhythm, ‘witnessed’ status, bystander CPR, prehospital defibrillation, and prehospital epinephrine administration. The results are expressed as odds ratios (ORs) and 95% confidence intervals (CIs). All statistical tests were performed using SAS software ver. 9.4 (SAS Institute Inc., Cary, NC, USA), and P-values < 0.05 were considered statistically significant.

RESULTS

Baseline characteristics and outcomes of patients according to prehospital airway management method

Among the 3,187 OHCA subjects registered in the KoCARC database between October 1, 2015 and December 31, 2016, 94 patients aged < 18 years were excluded. Additionally, 231 patients who were transferred from a different hospital prior to admission to the participating hospital, 912 who had unclear or unknown time variables between prehospital EMS site arrival and hospital arrival, 9 whose prehospital airway management methods were unknown, and 70 who experienced cardiac arrest in the ambulance were excluded. Ultimately, 1,871 patients were included in the final analysis. The subjects were divided into the following three groups according to the prehospital airway management method: BVM (n = 785), SGA (n = 965), and ETI (n = 121) (Fig. 1 and Table 1).

Patients in the ETI group were the oldest among the three, with subjects older than 65 years of age (median, 74 years) accounting for 73.6% (89 subjects) of the entire group. The BVM group had the lowest median on-scene time at 9 minutes, the highest rate of cardiac arrest occurring in a public place at 27.8% (218 subjects), and the highest witnessed cardiac arrest rate at 64% (502 subjects). Moreover, 23.4% and 24% of subjects in the SGA and ETI groups, respectively, required prehospital epinephrine administration; these rates were higher than that for the BVM group. Among the three groups, the BVM group had the highest survival to discharge rate and the highest proportion of patients with good neurologic outcomes at 11.8% and 8.7%, respectively.

Analysis of the association between prehospital airway management and outcomes

After adjustment for potential confounders, neither SGA nor ETI use was associated with better neurologic outcomes than BVM ventilation (OR, 0.59; CI, 0.33–1.06 for SGA; OR, 0.77; CI, 0.21–2.83 for ETI). Similarly, the use of SGA or ETI was not associated with improved survival to discharge compared to BVM ventilation (OR, 0.88; CI, 0.57–1.38 for SGA; OR, 1.52; CI, 0.66–3.47 for ETI) (Table 2).

Table 2. Effect of prehospital advanced airway management on survival outcomes

| Prehospital airway management | Good neurologic outcome | Survival to discharge |
|------------------------------|-------------------------|-----------------------|
|                              | Crude OR (95% CI)       | Adjusted OR\(^a\) (95% CI) | Crude OR (95% CI) | Adjusted OR\(^a\) (95% CI) |
| BVM                          | Reference               | Reference             | Reference         | Reference         |
| SGA                          | 0.65 (0.45–0.94)        | 0.59 (0.33–1.06)      | 0.73 (0.53–0.99)  | 0.88 (0.57–1.38)  |
| ETI                          | 0.55 (0.23–1.30)        | 0.77 (0.21–2.83)      | 0.90 (0.48–1.66)  | 1.52 (0.66–3.47)  |

OR, odds ratio; CI, confidence interval; BVM, bag-valve-mask; SGA, supraglottic airway; ETI, endotracheal intubation.

\(^{a}\)Adjusted for sex, age, response time interval, scene time interval, transport time interval, location of cardiac arrest occurrence, shockable rhythm, witnessed status, bystander cardiopulmonary resuscitation, prehospital defibrillation, and prehospital epinephrine administration.

Fig. 2. Rates of good neurologic outcome (A), and survival to discharge (B) according to the transport time interval and prehospital airway management method. BVM, bag-valve-mask; SGA, supraglottic airway; ETI, endotracheal intubation.
Analysis of the association between prehospital airway management and outcomes according to prehospital TTI

Among patients in whom the TTI was < 12 minutes, those in the SGA and ETI groups had good neurologic outcome rates of 4.4% and 2.9% (P = 0.561), as well as survival to discharge rates of 7.0% and 7.3% (P = 0.950), respectively (Fig. 2). After adjustment for potential confounders, neither the use of SGA nor ETI was associated with improved neurologic outcomes in patients with TTIs ≥ 12 minutes (684 subjects) (OR, 1.37; CI, 0.65–2.87 for SGA; OR, 1.31; CI, 0.30–5.81 for ETI). In patients with TTIs < 12 minutes (1,197 subjects), the use of SGA and ETI were also not associated with improved neurologic outcomes (OR, 0.57; CI, 0.31–1.07 for SGA; OR, 0.63; CI, 0.12–3.26 for ETI) or survival to discharge (OR, 0.79; CI, 0.48–1.31 for SGA; OR, 1.11; CI, 0.38–3.25 for ETI) compared with BVM ventilation (Table 3).

DISCUSSION

Our study revealed that the prehospital use of SGA and the performance of ETI were not associated with improved neurologic outcomes or survival to discharge compared to BVM ventilation regardless of transport time.

Conversely, a previous study by Kang et al. based on a cardiovascular disease surveillance database in South Korea showed significant differences in survival to discharge and good neurologic outcomes between subjects who underwent ETI and those who received BVM; their results were thus inconsistent with ours. The percentages of subjects who underwent ETI and SGA in Kang et al’s study study were 3.7% and 5.0%, respectively, whereas the corresponding percentages in our study were 6.5% and 51.6%, respectively. Furthermore, the percentage of subjects who underwent bystander CPR in their study (8.3%) was far lower than in ours (51.4%). With the gradual increase in the percentage of bystander CPR performed in South Korea, the rates of survival to discharge and good neurologic outcome in our study were relatively higher than those in the previous study. The proportion of EMS personnel who are level 1 EMTs has increased, and a videolaryngoscope to assist EMTs performing ETI has been deployed in ambulances. Furthermore, as the use of I-gel has also increased, the implementation rate of prehospital airway ventilation provided through ETI or the placement of an SGA has also increased. The higher percentage of patients undergoing ETI or SGA placement in our study than in Kang et al’s study may have reduced the selection bias.

A recent randomized controlled trial (RCT) on prehospital ventilation support methods based on advanced life support performed in Belgium and France found that BVM was neither inferior nor non-inferior to ETI in terms of survival with favorable 28-day neurologic function. Our present data are consistent with those of the aforementioned RCT, as we found that prehospital advanced airway management is not associated with improved neurologic outcomes and survival to discharge compared to BVM ventilation. Another RCT conducted in England that compared I-gel and ETI found no difference in the rate of favorable neurologic outcomes on the 30th day or at the time of discharge. However, each country may have differences in the scope of duty and performance level of EMTs. South Korea implements a single-tiered system according to which each ambulance has two or three EMTs. Furthermore, as the use of I-gel has also increased, the use of I-gel has also increased.

Table 3. Effect of prehospital advanced airway management on survival outcomes according to transport time interval

| Good neurologic outcome | Survival to discharge |
|-------------------------|-----------------------|
|                         | Crude OR (95% CI)     | Adjusted OR* (95% CI) |
|                         | Reference             | Reference              |
| TTI ≥ 12 min (n = 684)  | BVM                   | SGA                    | ETI                    |
|                         | Reference             | Reference              | Reference              |
| Crude OR (95% CI)       | Adjusted OR* (95% CI) |
| BVM                     | Reference             | Reference              |
| SGA                     | 1.16 (0.65–2.06)      | 1.37 (0.65–2.87)       |
| ETI                     | 1.03 (0.34–3.10)      | 1.31 (0.30–5.81)       |
| TTI < 12 min (n = 1,187) | Crude OR (95% CI)     | Adjusted OR* (95% CI) |
|                         | Reference             | Reference              |
| Crude OR (95% CI)       | Adjusted OR* (95% CI) |
| BVM                     | Reference             | Reference              |
| SGA                     | 0.44 (0.27–0.72)      | 0.57 (0.31–1.07)       |
| ETI                     | 0.29 (0.07–1.22)      | 0.63 (0.12–3.26)       |

OR, odds ratio; CI, confidence interval; TTI, transport time interval; BVM, bag-valve-mask; SGA, supraglottic airway; ETI, endotracheal intubation.

*Adjusted for sex, age, response time interval, scene time interval, location of cardiac arrest occurrence, shockable rhythm, witnessed status, bystander cardiopulmonary resuscitation, prehospital defibrillation, and prehospital epinephrine administration.
Advanced airway management was performed longer on-scene times than did their subjects who underwent BVM. It is presumed that the presence of EMTs who provided advanced care, including advanced airway management and prehospital epinephrine administration, influenced the on-scene time.

We also compared TTI to reflect differences according to region (urban vs. rural) and to medical care accessibility. We hypothesized that prehospital advanced airway management will have advantages over BVM ventilation with respect to neurologic outcomes, and that the benefits of advanced airway management for patients with OHCA will increase with the length of the TTI because it can provide effective ventilation and reduce airway complications without interrupting chest compressions. After adjusting for confounding variables, there were no differences in rates of favorable neurologic outcomes among the BVM, SGA, and ETI groups regardless of transport time. A previous study by Wang et al. found that ETI performance was associated with chest compression interruptions, and that the no-flow time was actually extended in situations where CPR was required. Jarman et al. reported no differences in CPR interruption time according to the prehospital airway management method, and showed that the failure of the first ETI attempt via direct laryngoscopy actually increased the duration of interrupted chest compressions. Therefore, we must consider the possibility that interrupted chest compressions do not differ according to the airway management method and may actually increase for subjects who undergo advanced airway management; this would explain why advanced management did not affect the neurologic outcomes in real-world situations.

A previous study showed that if the TTI is expected to be under 14 minutes, it is reasonable to redirect patients experiencing OHCA to percutaneous coronary intervention–capable hospitals. Although we considered the possibility that the benefits of prehospital advanced airway management during prolonged transfer times may affect the outcomes, we did not identify an association between prehospital advanced airway management and survival outcomes according to TTI. Previous studies found that excessive hyperventilation after advanced airway management should be avoided because it can cause increased intrathoracic pressure leading to decreased coronary and cerebral perfusion pressure in patients with OHCA. Kilgannon et al. also reported that hypoxia that occurs in patients following resuscitation from cardiac arrest is associated with increased mortality. It is, therefore, possible that these unexpected physiologic effects may offset the potential benefits of proper advanced airway management during transport.

According to a previous study conducted in Japan, a group of subjects in whom advanced airway management was performed within four minutes of CPR initiation showed better neurologic outcomes than did the group in whom advanced airway management was performed after four minutes. However, conflicting opinions exist regarding the effects of advanced airway management and advanced CPR (including drug use) on site on survival outcomes. Japan implements a single-tiered EMS system similar to that in South Korea; however, emergency life-saving technicians have different ETI certification criteria. Moreover, the ability to identify the timing of advanced airway management or the cause of its delay in the data registry that we used is limited. Thus, a well-designed follow-up study adapted for the South Korean system is required. In our current study, we were able to identify the association between advanced airway management methods and survival outcomes under the real-life emergency medical system in present-day South Korea. This can help improve future prehospital advanced airway management methods and advanced care directions in multiple-dispatch situations managed by the South Korean emergency medical system.

Our study had a number of limitations that may restrict the generalizability of our results. First, the integrity and validity of the data may be subject to the constraints inherent in multi-institutional observational studies. However, data were collected based on Utstein-style guidelines, and efforts were made to reduce the possibility of potential biases through quality control. Second, the competence levels of prehospital EMTs, regional differences, in-hospital CPR performance levels, and differences in post-cardiac arrest treatments were not reflected in our data; such variables may have influenced our results. Third, although multivariate analyses were performed to adjust for potential confounding variables, the possibility of selection bias still exists. The ETI group comprised the oldest subjects, and differences in the locations of cardiac arrest occurrence were observed; hence, there may have been bias in determining the EMT’s attempt and success rates when delivering prehospital advanced airway management to patients with OHCA. Fourth, the actual use of BVM in the field is difficult to identify via hospital records; this may also have contributed to reporting bias. Fifth, the KoCARC registry had insufficient information on the EMTs’ level of expertise, patients’ initial mental status upon admission to the emergency department, number of advanced airway management attempts, time of intervention, and type of SGA. Therefore, we could not investigate these parameters even though they may influence survival outcomes.

In conclusion, our study revealed no differences in the rates of favorable neurological outcomes between patients with OHCA who were managed using BVI, ETI, or SGA, regardless of transport time.
CONFLICT OF INTEREST

No potential conflict of interest relevant to this article was reported.

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