OBSERVATIONAL STUDY

OPEN

Association Between IV Contrast Media Exposure and Acute Kidney Injury in Patients Requiring Emergency Admission: A Nationwide Observational Study in Japan

OBJECTIVE: This study aimed to elucidate the association between IV contrast media CT and acute kidney injury (AKI) and in-hospital mortality among patients requiring emergency admission.

DESIGN: In this retrospective observational study, we examined AKI within 48 hours after CT, renal replacement therapy (RRT) dependence at discharge, and inhospital mortality in patients undergoing contrast-enhanced CT or nonenhanced CT. We performed 1:1 propensity score matching to adjust for confounders in the association between IV contrast media use and outcomes. Subgroup analyses were performed according to age, sex, diagnosis at admission, ICU admission, and preexisting chronic kidney disease (CKD).

SETTING AND PATIENTS: This study used the Medical Data Vision database between 2008 and 2019. This database is Japan's largest commercially available hospital-based claims database, covering about 45% of acute-care hospitals in Japan, and it also records laboratory results.

INTERVENTIONS: None.

MEASUREMENTS AND MAIN RESULTS: The study included 144,149 patients with (49,057) and without (95,092) contrast media exposure, from which 43,367 propensity score-matched pairs were generated. Between the propensity score-matched groups of overall patients, exposure to contrast media showed no significant risk of AKI (4.6% vs. 5.1%; odds ratio [OR], 0.899; 95% Cl, 0.845–0.958) or significant risk of RRT dependence (0.6% vs. 0.4%; OR, 1.297; 95% Cl, 1.070–1.574) and significant benefit for in-hospital mortality (5.4% vs. 6.5%; OR, 0.821; 95% Cl, 0.775–0.869). In subgroup analyses regarding preexisting CKD, exposure to contrast media was a significant risk for AKI in patients with CKD but not in those without CKD.

CONCLUSIONS: In this large-scale observational study, IV contrast media was not associated with an increased risk of AKI but concurrently showed beneficial effects on in-hospital mortality among patients requiring emergency admission.

KEYWORDS: acute kidney injury; contrast media; contrast-induced nephropathy; emergency department; postcontrast acute kidney injury

ontrast-enhanced CT plays a crucial role in the prompt diagnosis of life-threatening conditions such as bleeding in trauma (1), infections in sepsis (2, 3), and critical diagnoses like aortic dissection (4, 5). Timely and accurate diagnosis is mandatory in emergency settings, where patients often face potentially fatal situations. However, the use of contrast media has been associated with nephrotoxicity, including acute kidney injury (AKI) and death (6, 7). Some studies have revealed the mechanisms by which contrast Ryo Hisamune, MD¹ Kazuma Yamakawa[®], PhD¹ Yutaka Umemura, PhD² Noritaka Ushio, MD¹ Katsunori Mochizuki, PhD¹ Ryota Inokuchi, PhD³ Kent Doi, PhD³ Akira Takasu, PhD¹

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DOI: 10.1097/CCE.000000000001142

KEY POINTS

Question: How does the risk of contrast-enhanced CT relate to kidney dysfunction and mortality compared with nonenhanced CT in patients requiring emergency admission?

Findings: In this large retrospective study of 144,149 emergently admitted patients in a Japanese database who underwent CT scans, exposure to IV contrast media was not associated with an increased risk of acute kidney injury. However, it was significantly associated with lower hospital mortality compared with nonexposure, even after adjustment for important confounders.

Meaning: Our findings suggest that contrastenhanced CT may pose a limited risk for renal impairment in many emergency department patients.

agents cause renal injury (8, 9). Two pathways have been suggested: direct damage to renal tubular epithelial cells by contrast agents and indirect damage via renal ischemia due to intrarenal vasoconstriction and reduced renal glomerular blood flow (8, 9). Clinical studies have shown that contrast media administration causes contrasted-induced nephropathy (CIN), especially in high-risk patients, who are the elderly (10), have chronic kidney disease (CKD) (10–12), diabetes mellitus (10, 13, 14), chronic heart failure (10, 14), anemia (14), and take nephrotoxic drugs (14). The occurrence rate of contrast-induced AKI reportedly varies widely, ranging from 3.3% to 14.5% (15–17).

However, it has been suggested that the perceived risk of AKI associated with contrast media use may have been overestimated in previous reports due to methodological limitations such as the absence of adequate control groups (18, 19). Recent observational studies and meta-analyses reported that exposure to contrast media was not significantly associated with the incidence of AKI (20–23). Also, studies in patients with renal impairment have indicated no significant difference in AKI incidence between patients who underwent contrast-enhanced CT and those who did not (24). Similar findings extend to critically ill patients who had a 20–50% chance of developing AKI regardless of contrast media use in ICUs (19). In addition, AKI and mortality do not increase with contrast media use, even in sepsis patients who were introduced to renal replacement therapy (RRT) at an early stage (25).

Despite these findings across various conditions, little research has been conducted on patients visiting emergency departments. Existing clinical evidence in this area is limited due to small-scale studies, particularly with almost no research focusing specifically on patients requiring emergency admission. Therefore, we aimed to elucidate the potential risks associated with contrast-enhanced CT, including AKI, induction of RRT, and in-hospital mortality, using comprehensive clinical big data from patients requiring emergency admission.

METHODS

Data Source

We performed a retrospective observational cohort study using the Medical Data Vision (MDV) database, Japan's largest commercially available hospitalbased claims database. As of April 1, 2022, the MDV database contained over 35 million inpatients and outpatients treated at any of the 793 hospitals, covering about 45% of acute-care hospitals in Japan (26). This database is composed of administrative claims data originating from Japan's healthcare system and includes data on each patient's age, sex, admission and discharge dates, discharge status, primary diagnoses, comorbidities at admission, postadmission complication diagnoses, types of surgical procedures coded with the original Japanese operation codes, daily records of procedures, diagnostic examinations, drug administrations, and results of laboratory tests. The diagnoses in this database are recorded using the International Classification of Diseases, 10th revision (ICD-10) codes. From the original MDV database, we extracted the clinical data of 534,739 patients who required emergency admissions to 42 hospitals that recorded laboratory data from April 2008 to December 2019.

This study was conducted in compliance with the principles of the Declaration of Helsinki of 1975 and according to the ethical standard of our institutional committee (the institutional review board [IRB] for Clinical Research of Osaka Medical Center) and the Strengthening the Reporting of Observational Studies in Epidemiology guideline. Because of the anonymous

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nature of the data, the requirement for informed consent from patients was waived according to our institutional committee (IRB number S201916015, study title: "Evaluation of clinical issues in emergency and intensive care medicine using big data," approved on July 7, 2020).

Study Participants and Exposure Variable

Patients were eligible for this study if they met the following criteria: patients 1) required emergency admission, 2) received a CT scan with or without IV contrast media at admission, and 3) were older than 18 years at the time of admission. We excluded patients who underwent coronary angiography or percutaneous coronary intervention at admission, who had end-stage renal disease based on the ICD-10 code of N18.0 or N18.5 at admission, who died within 24 hours after admission, and who did not have laboratory test results for serum creatinine level at admission and within 48 hours after admission. We also excluded data from the second and subsequent hospitalizations of patients who were hospitalized multiple times during the study period and used only the data from their first hospitalization. Patients who received a CT scan with IV contrast media were classified into the enhanced CT group, and those who received a CT scan without IV contrast media were classified into the nonenhanced CT group based on the procedure code. Only low-osmolar contrast media were used for contrast-enhanced CT in this study.

Definition of Outcomes

The primary outcome was the incidence of AKI. In this study, we evaluated three different definitions of AKI. AKI (stages 1–3) was defined as an increase in serum creatine level of more than 0.3 mg/dL or a rise of at least 1.5 times from the result of the baseline serum creatinine level within 48 hours after admission, following the criteria of the Kidney Disease Improving Global Outcomes (KDIGO). AKI stage 3 was defined as an increase in serum creatinine level of more than 4.0 mg/dL or a rise of more than 3.0 times from the result of the baseline creatinine level within 48 hours of admission and the use of RRT within 48 hours of admission, following stage 3 of the KDIGO criteria. CIN was diagnosed based on serum creatine level when serum creatinine increases by more than 0.5 mg/dL

or rises to at least 1.25 times from the baseline creatinine level within 72 hours of contrast-enhanced CT scan. The secondary outcomes included RRT during hospitalization, RRT dependence at discharge, and in-hospital mortality. RRT during hospitalization was defined as receiving RRT at least once during hospitalization. RRT dependence was defined as receiving RRT within 2 days before discharge. In Japan, patients dependent on RRT at discharge and who require outpatient RRT after discharge are scheduled to undergo RRT thrice weekly as conventional intermittent dialysis (27). Therefore, the definition of RRT dependence as the requirement for RRT "within 2 days before discharge" appears reasonable (22).

Data Collection

We collected the following data: age, sex, height, weight, hospital volume, comorbidities, laboratory tests, Charlson Comorbidity Index (CCI) (28), use of nephrotoxic medications, primary diagnoses at admission, ICU admission, and emergency operation. We used predefined ICD-10 coding algorithms for evaluating CCI (29). Based on the ICD-10 coding, we identified a recorded diagnosis of congestive heart failure, diabetic nephropathy, CKD, end-stage renal disease, and hypertension as a complication disease, and trauma, macrovascular disease, cerebrovascular disease, myocardial infarction, and sepsis as the primary diagnosis. We defined emergency operation as any surgery using general anesthesia within 2 days of admission. We identified the following treatments performed at admission: vasoactive agents, including dopamine, dobutamine, norepinephrine, epinephrine, or vasopressin, and nephrotoxic medications, such as nonsteroidal anti-inflammatory drugs and diuretics. Details on the definition of the ICD-10 codes used in this study are provided in Supplemental Table 1 (http://links.lww.com/CCX/B386).

Statistical Analysis

Due to the nature of a retrospective study, it was assumed that an imbalance existed among the covariates in the enhanced CT group and the nonenhanced CT group at baseline. Propensity score matching was thus performed between each group (30). The propensity scores were determined using logistic regression with 19 variables assumed to be associated with IV contrast media used as covariates (Supplemental Table 2, http://links.lww.com/CCX/B386). We chose these covariates to calculate the propensity scores based on the risk factors of contrast-inducing AKI and the clinical importance of enhanced CT (31). C-statistics were evaluated as a measure of discrimination. The distribution of propensity scores for each group was compared graphically to assess the overlap assumption. We conducted a one-to-one analysis using nearest-neighbor matching based on the estimated propensity scores for each patient. A match occurred when a patient in the enhanced CT group had an estimated score within 0.2 sDs of a patient in the nonenhanced CT group. Standardized mean difference was calculated for the balance assessment between each group before and after matching, and we considered values less than 0.1 acceptable (32).

We performed propensity score matching for the subgroups of age (\geq 65 yr old vs. < 65 yr old), sex (male vs. female), diagnosis at admission (trauma vs. sepsis vs. cerebrovascular disease vs. macrovascular disease), preexisting CKD, ICU admission (ICU vs. general ward), and admission patient's condition with or without organ dysfunction.

We also performed a sensitivity analysis using logistic regression analysis. In total, 21 independent variables were included, of which 19 were the same factors as in the propensity score matching, and the newly added variables were enhanced CT and an interaction term between CKD and enhanced CT. The final model estimated the relative risk of event occurrence for each variable by calculating the odds ratio (OR) and 95% CI.

Continuous variables are reported as means with sp and categorical variables as numbers with percentages. We analyzed continuous outcomes using the *t*-test and categorical outcomes using the Fisher exact test and the chi-square test. All statistical inferences were two-sided; a p value less than 0.05 indicated statistical significance. Data were analyzed using R software, version 4.2.3 (R Foundation for Statistical Computing, Vienna, Austria).

RESULTS

Study Population

We identified 314,335 patients who fulfilled the inclusion criteria and subsequently excluded 170,186 patients who met the exclusion criteria. The remaining 144,149 eligible patients were divided into the enhanced CT group (n = 49,057) and the nonenhanced CT group (n = 95,092). **Supplemental Figure 1** (http://links.lww. com/CCX/B386) shows the distributions of propensity scores in the two groups. The *C*-statistic for the fit of the propensity scores was 0.727 (95% CI, 0.724–0.729). After matching, 43,367 propensity score-matched pairs were generated (**Fig. 1**).

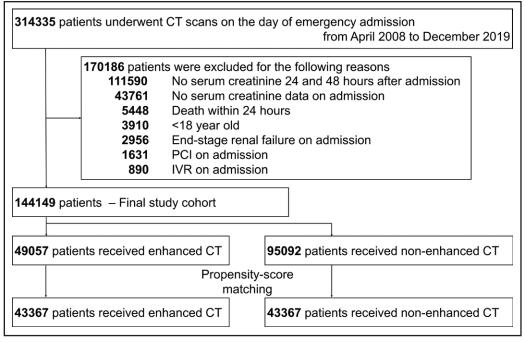
Table 1 shows the baseline characteristics of the two groups before and after propensity score matching. In the unmatched groups, patients in the enhanced CT group tended to be younger, have lower rates for several comorbidities and better renal function, and be more often diagnosed as having macrovascular disease than those in the nonenhanced CT group. Also, the enhanced CT group received more intensive treatment, including catecholamines, ICU admission, and emergency operation, than the nonenhanced CT group. After propensity score matching, the two groups had no significant differences in inpatient characteristics.

Effect of Contrast Media on AKI and Mortality

After propensity score matching, there was a significantly lower risk of AKI (stages 1–3, stage 3) and CIN associated with IV contrast media administration (**Table 2**). The incidence of RRT during hospitalization and RRT dependence at discharge were significantly higher, and in-hospital mortality was significantly lower in the enhanced CT group (**Table 3**).

Effect of Contrast Media by Subgroups

All outcomes in the subgroups using propensity score matching are shown in Supplemental Tables 3-8 (http://links.lww.com/CCX/B386). The risk of AKI stages 1-3 associated with enhanced CT by the clinically relevant subgroups is summarized in Figure 2. There were significant associations between the incidence of AKI stages 1-3 and enhanced CT in the subsets of patients with CKD (OR, 2.082; 95% CI, 1.571-2.769), and in-hospital mortality showed no significant difference between the groups. Among patients without CKD, the risk of AKI stages 1-3 was significantly lower (OR, 0.833; 95% CI, 0.781-0.889), and in-hospital mortality was also significantly lower (OR, 0.796; 95% CI, 0.751-0.843) in the enhanced CT group (Supplemental Table 6, http://links.lww.com/ CCX/B386). According to the diagnostic disease, the



consistent with those obtained from the subgroup analysis based on CKD.

DISCUSSION

We used a nationwide, large-scale inpatient database in Japan to evaluate the risk of AKI (stages 1-3, stage 3, CIN) and clinical inhospital outcomes between patients requiring emergency admissions who received contrastenhanced CT or noncontrast-enhanced CT. To our knowledge, this study is the largest

Figure 1. Flowchart for patient selection. IVR = interventional radiology, PCI = percutaneous coronary intervention.

risk of AKI stages 1-3 in each disease showed no significant difference between the groups, whereas the inhospital mortality in trauma patients was significantly higher in the enhanced CT group (OR, 1.530; 95% CI, 1.162–2.021). However, among the patients with sepsis, there was no significant difference in the risk of AKI stages 1-3, whereas the in-hospital mortality was significantly lower in the enhanced CT group (OR, 0.796; 95% CI, 0.675–0.938) (Supplemental Table 5, http://links.lww.com/CCX/B386). Among patients with organ dysfunction, the risk of AKI stages 1-3 and in-hospital mortality did not show statistical significance, but a trend toward lower incidence in the enhanced CT group was revealed (Supplemental Table 8, http://links.lww.com/CCX/B386). A similar trend was observed among the patients without organ dysfunction, with a significant difference noted in inhospital mortality (OR, 0.834; 95% CI, 0.765–0.910).

Results of Sensitivity Analysis

All outcomes in the multiple logistic regression analysis are shown in **Supplemental Table 9** (http://links. lww.com/CCX/B386). Sensitivity analyses confirmed that there were no significant changes in the adjusted ORs for all outcomes. Additionally, the results for the interaction term of CKD and enhanced CT were observational study regarding this clinically relevant research topic. The main finding of this study was that the use of contrast media did not show a significant difference in the incidence of AKI (stages 1–3, stage 3, CIN) among patients requiring emergency admission. Additionally, the in-hospital mortality in the enhanced CT group was significantly lower than in the nonenhanced CT group. These findings suggested that withholding enhanced CT may not be necessary when an accurate diagnosis is required to succeed in treating patients requiring emergency admission.

This study also elucidated the risk of mortality and revealed a low proportion of patients admitted to the ICU, which reflects that the patients were not critically ill. Consequently, the study included patients with an overall lower mortality. However, AKI is reportedly associated with risks for mortality and morbidity (33–35), but in patients who develop CIN, there is a debate about whether this form of AKI is associated with these clinically adverse outcomes (36). Recent studies found that an amount exceeding 100 mL of contrast medium used per patient was associated with in-hospital death (37) and that mortality following AKI was often due to the time required for recovery of renal function (38). The maximum serum creatinine level within 24 hours of AKI diagnosis was also an independent predictor of short-term prognosis

TABLE 1.

Baseline Characteristics of Patients in Unmatched and 1:1 Propensity Score-Matched Groups

	Unmatched Groups			Matched Groups			
Characteristics	Nonenhanced CT	Enhanced CT	SMD	Nonenhanced CT	Enhanced CT	SMD	
Patient number	95,092	49,057		43,367	43,367		
Age, yr	73.29 (16.65)	65.62 (18.07)	0.441	66.41 (18.84)	66.91 (17.33)	0.028	
Sex, female	44,867 (47.2)	21,951 (44.7)	0.049	19,377 (44.7)	19,510 (45.0)	0.006	
Hospital volume			0.201			0.033	
199 beds	4,233 (4.5)	1,179 (2.4)		1,124 (2.6)	987 (2.3)		
200–499 beds	66,081 (69.5)	31,087 (63.4)		27,716 (63.9)	28,340 (65.3)		
500 beds	24,778 (26.1)	16,791 (34.2)		14,527 (33.5)	14,040 (32.4)		
Comorbidities							
Congestive heart failure	17,664 (18.6)	4663 (9.5)	0.263	3,661 (8.4)	4,080 (9.4)	0.034	
Chronic kidney disease	5,210 (5.5)	639 (1.3)	0.232	512 (1.2)	590 (1.4)	0.016	
Hypertension	37,444 (39.4)	17,033 (34.7)	0.097	14,488 (33.4)	14,844 (34.2)	0.017	
Diabetic nephropathy	1,394 (1.5)	249 (0.5)	0.097	191 (0.4)	242 (0.6)	0.017	
Myocardial infarction	5,118 (5.4)	1,949 (4.0)	0.067	1,765 (4.1)	1,799 (4.1)	0.004	
Laboratory test results at admission	on						
Glomerular filtration rate, mL/ min/1.73 m ²	60.98 (34.19)	73.99 (27.39)	0.420	73.20 (38.95)	72.18 (24.96)	0.031	
Creatinine, mg/dL	1.20 (1.26)	0.84 (0.55)	0.370	0.96 (0.91)	0.85 (0.57)	0.138	
Hemoglobin $< 11 \text{ g/dL}$	25,096 (26.4)	9,840 (20.1)	0.150	8,664 (20.0)	8,936 (20.6)	0.016	
Charlson Comorbidity Index	2.40 (2.49)	2.36 (2.68)	0.014	2.11 (2.41)	2.43 (2.73)	0.124	
Nephrotoxic medication							
Nonsteroidal anti-inflammatory drugs	23,616 (24.8)	16,317 (33.3)	0.186	12,906 (29.8)	13,180 (30.4)	0.014	
Catecholamine	1,756 (1.8)	1,404 (2.9)	0.067	981 (2.3)	1,046 (2.4)	0.010	
Diuretic medication	14,607 (15.4)	4,840 (9.9)	0.166	3,561 (8.2)	3,996 (9.2)	0.036	
Antibiotic (aminoglycoside)	31,639 (33.3)	12,027 (24.5)	0.039	422 (1.0)	634 (1.5)	0.045	
Antibiotic (anti-methicillin- resistant <i>Staphylococcus aureus</i> drug)	889 (0.9)	661 (1.3)	0.014	567 (1.3)	498 (1.1)	0.014	
Diagnosis at admission							
Trauma	12,561 (13.2)	3,155 (6.4)	0.229	3,110 (7.2)	3,127 (7.2)	0.002	
Sepsis	9,084 (9.6)	2,845 (5.8)	0.141	2,584 (6.0)	2,741 (6.3)	0.015	
Cerebrovascular disease	9,659 (10.2)	4,326 (8.8)	0.046	3,742 (8.6)	4,017 (9.3)	0.022	
Macrovascular disease	441 (0.5)	2,772 (5.7)	0.305	437 (1.0)	673 (1.6)	0.048	
Organ dysfunction	31,639 (33.3)	12,027 (24.5)	0.194	12,280 (28.3)	10,757 (24.8)	0.080	
ICU admission	5,133 (5.4)	4,734 (9.6)	0.162	3,002 (6.9)	3,230 (7.4)	0.020	
Emergency operation	7,453 (7.8)	9,573 (19.5)	0.345	5,794 (13.4)	6,662 (15.4)	0.057	

 $\mathsf{SMD} = \mathsf{standardized} \text{ mean difference}.$

Continuous variables are presented as means with SD, and categorical variables are presented as numbers with percentages.

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TABLE 2.Risk of Acute Kidney Injury in the Propensity Score-Matched Groups

Primary Outcomes	Nonenhanced CT, n (%)	Enhanced CT, n (%)	OR	95% Cl	р
AKI stages 1-3	2199 (5.1)	1989 (4.6)	0.899	0.845-0.958	0.001
AKI stage 3	427 (1.0)	493 (1.1)	1.156	1.013-1.320	0.031
Contrast-induced nephropathy	3889 (9.0)	3669 (8.5)	0.938	0.895-0.984	0.008

AKI = acute kidney injury, OR = odds ratio.

TABLE 3. Risk of Secondary Patient-Centered Outcomes in the Propensity Score-Matched Groups

Secondary outcomes	Nonenhanced CT, n (%)	Enhanced CT, n (%)	OR	95% CI	p
RRT during hospitalization	408 (0.9)	543 (1.5)	1.335	1.171-1.523	< 0.001
RRT dependence	193 (0.4)	250 (0.6)	1.297	1.070-1.574	0.008
In-hospital mortality	2812 (6.5)	2335 (5.4)	0.821	0.775-0.869	< 0.001

RRT = renal replacement therapy, OR = odds ratio.

		OR	95% CI	OR with	95% CI
Primary analysis		0.899	0.845-0.958	-	
Subgroup analysis					
Age	≥ 65	0.928	0.861-0.999	1	
	< 65	0.744	0.658-0.841		
Sex	Men	0.933	0.860-1.013	- +=::	
	Women	0.831	0.753-0.918	- H ⊞ H	
Diagnosis on	Trauma	1.239	0.955-1.610		
admission	Sepsis	0.984	0.838-1.156	-	н
	Macrovascular	1.103	0.761-1.600		■
	Cerebrovascular	1.054	0.848-1.312	- i	∎1
Comorbidities	With CKD	2.082	1.571-2.769		⊢∎
	Without CKD	0.833	0.781-0.889		
Admission site	ICU	0.917	0.790-1.065	- ⊢ ∎ :	1
	General ward	0.881	0.820-0.945	- 1	
Organ dysfunction	With	0.930	0.849-1.018	H a ti	
	Without	0.921	0.840-1.010	- H eni	
				0.3 1	3
			Fav	vors enhanced CT	Favors non-enhanced

Figure 2. Association between IV contrast media administration and the risk of acute kidney injury (AKI) in several subsets based on patient characteristics. The risk of AKI between: 1) overall, 2) elderly patients (65 yr and older) and younger patients (younger than 65 yr), 3) men and women, 4) primary diagnoses such as trauma, sepsis, macrovascular disease, and cerebrovascular disease, 5) patients with chronic kidney disease (CKD) and without CKD, 6) admission to ICU and general ward, and 7) patients with organ dysfunction and without organ dysfunction was evaluated by propensity score-matched analysis. OR = odds ratio.

(39). In the present study, which focused on patients diagnosed early after the administration of contrast media, the risk of death was observed to decrease,

subgroup analysis of various diagnoses at admission. In trauma patients, the amount of blood transfusion (40), injury severity score, and transcatheter arterial

regardless

organ

of

dysfunction,

increased risk of RRT. As this trend was also observed in subgroup analysis among more severe patients with

using contrast media for accurate diagnosis at the time of admission is considered beneficial for improving patient outcomes.

Emergency patients are characterized by a variety of diseases and poor general health and often have mul-

tiple organ damage, in-

cluding renal failure. It

is hard for physicians

to instantly ascertain a

patient's condition and

clinical risk of CIN. We,

therefore, performed a

an

embolization (41) are also known to be associated with the incidence of AKI. In the present study, in which we were unable to adjust for these factors, we found that the short-term prognosis in trauma patients deteriorated in the enhanced CT group compared with the nonenhanced CT group. In sepsis, AKI has generally been associated with high mortality (42, 43); however, assessment of sources of infection with enhanced CT is critical in the initial management of sepsis (44). Miyamoto et al (25) reported there to be no association between IV contrast media use and adverse in-hospital outcomes in patients with septic AKI requiring dialysis. Our findings also corroborated their findings.

Contrast media can cause potential toxicity to the kidney through direct toxicity on the renal tubules and indirect toxicity induced by renal ischemia due to vasospasm (8, 9). Patients with CKD are particularly susceptible to this adverse effect (45, 46). Observational studies showed that enhanced CT was associated with AKI in emergency patients with renal impairment (12, 47). In our results, the risks of RRT induction and RRT dependence were higher in the enhanced CT group. Among the patients who underwent RRT in the enhanced CT group, 27% had CKD, and of those who became RRT dependent, 40% had CKD. These proportions were approximately two to three times higher than those in the nonenhanced CT group. In the subgroup analysis, patients with CKD were at higher risk for AKI, RRT induction, and RRT dependence. Therefore, the use of contrast media in patients with CKD may pose more harm than benefit.

Limitations

Our study has several limitations. First, this was a retrospective observational study. Therefore, there may be potential unmeasured confounders between those patients with and without contrast media use, although we adjusted for differences in baseline characteristics using propensity score matching. For example, information on the site of the CT scan, periprocedural IV hydration, and vital signs was lacking in the dataset used in this study. However, we were not able to completely remove residual confounding due to unmeasurable confounding factors. Second, the accuracy of diagnoses recorded in administrative claims databases is typically lower than that recorded in prospective studies. Physicians often register diagnosis codes based on clinical judgment, and there are no established accurate criteria. Third, we considered only the risk of AKI within 48–72 hours and mortality during hospitalization. We were unable to assess long-term adverse events following hospital discharge because the dataset used in this study comprised only hospitalized patients. Finally, we were unable to collect information on the criteria for using contrast media as its use was determined by the treating medical team based on factors such as the patient's condition and renal function. Therefore, we could not adjust for this factor.

CONCLUSIONS

Our study using a nationwide inpatient database suggested that the use of contrast-enhanced CT for diagnostic purposes may not be associated with a significantly increased risk of AKI and might be associated with a mortality benefit in patients requiring emergency admission. Even though the patient's condition and renal function must be considered, contrast-enhanced CT may be considered necessary for accurate diagnosis and treatment in the emergency department.

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Supplemental digital content is available for this article. Direct URL citations appear in the printed text and are provided in the HTML and PDF versions of this article on the journal's website (http://journals.lww.com/ccejournal).

Drs. Hisamune, Yamakawa, and Umemura designed the study. Drs. Hisamune, Umemura, and Ushio were involved in acquiring data. Drs. Hisamune, Yamakawa, Umemura, and Mochizuki were responsible for the statistical analysis. Drs. Hisamune and Yamakawa interpreted the data and wrote the first draft of the report. Drs. Umemura, Inokuchi, Doi, and Takasu revised the report critically for important intellectual content. All authors approved the final version of the article. The corresponding author confirms to have had full access to the data in the study and final responsibility for the decision to submit for publication.

The authors have disclosed that they do not have any potential conflicts of interest.

Ethics approval and consent for each institution to participate in the study were approved by the institutional research ethics committee in accordance with local ethical regulations.

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Informed consent was not required due to the observational and anonymous nature of data collection.

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