ORIGINAL CONTRIBUTION

Association Between Male Sex and Increased Mortality After Falls

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Abstract

Objectives: Whether sex affects the mortality of trauma patients remains unknown. The hypothesis of this study was that sex was associated with altered mortality rates in trauma.

Methods: A retrospective review of trauma patients' records in the Japan Trauma Data Bank was conducted (n = 80,813) from 185 major emergency hospitals across Japan. The primary outcome variable was in-hospital mortality within 28 days. Secondary outcome variables included serious injuries to different body regions with an Abbreviated Injury Scale of ≥3.

Results: In the analysis of 80,813 trauma patients, males had significantly greater 28-day mortality compared to females (adjusted p = 0.0072, odds ratio [OR] = 1.14, 95% confidence interval [CI] = 1.06 to 1.23) via logistic regression analysis adjusted for age, mechanism, Injury Severity Score, Revised Trauma Score, and potential preexisting risk factors. Of 10 injury categories examined, sex significantly affected in-hospital 28-day mortality rate in falls (adjusted p < 0.0001, OR = 1.34, 95% CI = 1.19 to 1.52). Further analysis of three fall subcategories by falling distance revealed that male patients who fell from ground level had significantly higher 28-day mortality (adjusted p < 0.0001, OR = 1.75, 95% CI = 1.43 to 2.14) and a significantly greater frequency of serious injury to the head, thorax, abdomen, and spine, but a lower frequency of serious injury to the extremities, compared to female patients.

Conclusions: Compared to female trauma patients, male trauma patients had greater 28-day mortality. In particular, ground-level falls had a significant sex difference in mortality, with serious injury to different body regions. Sex differences appeared to be important for fatalities from ground-level falls.

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rauma is a leading cause of overall population mortality and a major health problem.^{1,2} In the United States, road injury, one of the major trauma categories, is the fifth leading cause of years of life lost, ranking immediately below ischemic heart disease, lung cancer, stroke, and chronic obstructive pulmonary disease, while ranking above colorectal and breast cancer.¹ Falls, transport injuries, burns, and exposures to mechanical forces have also been mentioned in global public health studies aiming at reducing premature mortality in public health.^{1,2}

Differences in clinical outcomes by patient sex have been extensively reported in common diseases or conditions, including heart diseases,^{3,4} lung diseases,⁵ stroke,⁶ and critical illnesses.⁷ Furthermore, recent proposals on sex-specific approaches to improve clinical

outcomes, including sex-specific surveillance, screening, and interventions, have highlighted the importance of investigating sex differences.^{3,5,8} Sex differences have been previously studied in trauma, focusing on dissimilar responses by sex to hemorrhagic shock^{9,10} or altered risks of trauma-associated lung complications. 11,12 However, data on whether sex affects the mortality of trauma patients remain inconsistent.7,13-16 Therefore, in this study, we tested the hypothesis that sex was associated with mortality rate in trauma patients using a large cohort across Japan. Because clinical outcomes are associated with not only the type of injury but also its severity, we investigated the differences in mortality by sex according to the type of injury with adjustment for severity. The primary outcome was in-hospital 28-day mortality, and the

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secondary outcome variables included serious injuries to various body regions.

Japan Trauma Data Bank

The Japan Trauma Data Bank (JTDB) is a trauma databank supported by the Japanese Association for the Surgery of Trauma and the Japanese Association for Acute Medicine, similar to trauma databases in North America, Europe, and Oceania. The JTDB aims to improve the quality of trauma care and provides annual reports. The JTDB data were registered by 185 major emergency medical hospitals across Japan.

The JTDB captures trauma cases with data on age, sex, mechanism of injury, Abbreviated Injury Scale (AIS; 1998 version), Injury Severity Score (ISS), Revised Trauma Score (RTS), and in-hospital 28-day mortality. Patients are categorized by institutions into one of 10 categories according to their mechanism of injury. which includes eight blunt trauma categories (road injury, fall, compression injury, sports injury, machinery injury, falling object injury, railway injury, and other), penetrating trauma (stab, gunshot, impalement injury, and other), and burns. Similarly, patients who experience falls are divided into three subcategories according to the height of fall: low (falls from ground level), medium (falls from stairs), and high (falls from higher levels than stairs). Preexisting conditions such as cerebrovascular disease (hemiplegia, etc.), ischemic heart disease, liver cirrhosis, and dementia, as well as alcohol use on day of injury and use of anticoagulants, are also extracted from the medical record and entered into the JTDB by participating institutions.

METHODS

Study Design

This was a retrospective cohort study using the JTDB. The institutional review board at the Senshu Trauma and Critical Care Center approved this study, and because the JTDB is an anonymous registry, the institutional review board waived the need for patient consent.

Study Setting and Population

In this study, we analyzed the data set of 80,813 patients in Japan nationwide who experienced trauma between January 2004 and December 2012 and were registered in the JTDB.

Study Protocol

The primary outcome variable was in-hospital mortality within 28 days. We first focused on all trauma patients and tested for sex differences using a binary logistic regression test, which allowed for the correction of potential confounding factors including age, mechanism, ISS (anatomically based severity score), RTS (physiologically based severity score), and potential preexisting risk factors of trauma morality, such as preexisting cerebrovascular disease (hemiplegia, etc.), ischemic heart disease, liver cirrhosis, and dementia. ²¹ For the fall category, we further examined sex difference in the three height-of-fall subcategories mentioned above using the logistic regression method. Because

alcohol use on day of injury and use of anticoagulants are potential factors for mortality risk in ground-level fall, 22 we added these as covariates in the logistic regression analysis of the ground-level fall.

The secondary outcome variables were serious injuries (AIS \geq 3) to six body regions: head, thorax, abdomen, spine, upper, and lower extremities. We used the chi-square or Fisher's exact test for analysis. We further investigated sex differences in mortalities using a stratified analysis by anatomic injury location.

Data Analysis

Differences were considered significant with a two-tailed p-value of <0.05. Multiple comparisons were adjusted using the Bonferroni procedure. Absolute standardized differences in characteristics between male and female patients of ground-level falls were calculated. All analyses were performed using SPSS (version 21)

RESULTS

We first examined data on 80,813 trauma patients for differences in 28-day in-hospital mortality by sex. In the logistic regression test, adjusting for potential confounding factors including age, mechanism, ISS, RTS, and potential preexisting risk factors of trauma morality such as preexisting cerebrovascular disease (hemiplegia, etc.), ischemic heart disease, liver cirrhosis, and dementia, males had significantly greater 28-day mortality compared to females (adjusted p = 0.0072; odds ratio [OR] = 1.14; 95% confidence interval [CI] = 1.06 to 1.23).

Sex differences in 28-day in-hospital mortality in each injury category are shown in Table 1. Of the 10 injury categories, injuries from falls were significantly associated with altered 28-day in-hospital mortality by sex. Male patients who had fall injuries had significantly greater 28-day mortality compared to female patients (adjusted p < 0.0001, OR = 1.34, 95% CI = 1.19 to 1.52, via logistic regression analysis followed by Bonferroni correction; Table 1).

Patients experiencing falls were assigned to the three subcategories by the height: low (falls from ground level, n = 15,207), medium (falls from stairs, n = 8,553), and high (fall from levels higher than stairs, n = 9,725). We determined sex difference in mortality for each fall subcategory. No significant sex difference in in-hospital 28-day mortality was observed in the high (adjusted p = 0.21, OR = 1.15, 95% CI = 0.92 to 1.44) or medium (adjusted p = 0.76, OR = 1.04, 95% CI = 0.82 to 1.31) fall subcategories via logistic regression analysis adjusted by age, ISS, RTS, and preexisting risk factors. In contrast, male patients who fell from ground level had a significantly greater mortality rate compared to female patients (in-hospital 28-day mortality, male vs. female [total], 0.069 vs. 0.029 [0.045]; p < 0.0001, OR = 1.66, 95% CI = 1.35 to 2.03), via logistic regression analysis adjusted for age, ISS, and RTS, preexisting cerebrovascular disease (hemiplegia, etc.), ischemic heart disease, liver cirrhosis, and dementia; Table 2, Figure 1). Alcohol use on day of injury and use of anticoagulant are potential factors for mortality risk in ground-level falls.²¹ Repeating the logistic regression analysis adding alco-

Table 1 In-hospital 28-day Mortality Rate by Sex in Ten Different Injury Categories

	Mortality (Death/Total cases)			
	Male	Female	Adjusted p-value	OR (95% CI)
Blunt trauma				
Road injury	0.120 (2,962/24,728)	0.136 (1,462/10,741)	0.85	1.01 (0.90–1.13)
Falls	0.121 (2,334/19,233)	0.086 (1,227/14,252)	<0.0001*	1.34 (1.19–1.52)
Compression injury	0.139 (143/1,030)	0.099 (10/101)	0.49	1.58 (0.44–5.73)
Sports injury	0.022 (20/892)	0.006 (1/166)	0.15	34.0 (0.28-4200)
Machinery injury	0.033 (25/749)	0.036 (4/111)	0.19*	0.09 (0.01-0.67)
Falling object injury	0.089 (57/643)	0.093 (5/54)	0.11	0.31 (0.08-1.28)
Railway injury	0.367 (106/289)	0.370 (61/165)	0.78	0.90 (0.42-1.92)
Others	0.080 (120/1,503)	0.075 (43/576)	0.21	1.47 (0.81–2.69)
Penetrating trauma	0.086 (179/2,090)	0.088 (71/804)	0.84	1.06 (0.62-1.82)
Burn	0.168 (291/1,735)	0.171 (163/951)	0.48	1.11 (0.83-1.50)
Total	0.118 (6,237/52,892)	0.109 (3,047/27,921)	0.0072*	1.14 (1.06–1.23)

*When the logistic regression p-value was <0.05, Bonferroni correction was subsequently performed; the Bonferroni corrected p-value is shown.

Table 2 Characteristics of Patients With Injury Due to Falls From Ground Level

Characteristics	Male (n = 6,385)	Female (<i>n</i> = 8,822)	Absolute Standardized Difference, %
Age, yr	67.2 (20.2)	79.1 (14.5)	67.7
Systolic blood pressure, mm Hg	142.4 (32.6)	147.7 (30.6)	16.8
Heart rate, beats/min	84.9 (20.5)	83.0 (16.8)	10.1
Respiratory rate, breaths/min	19.9 (5.5)	19.7 (4.8)	3.9
Glasgow Coma Scale score	13.3 (3.1)	14.2 (2.0)	34.5
Injury Severity Score	13.4 (7.3)	10.5 (5.0)	46.4
Revised Trauma Score	7.416 (1.011)	7.670 (0.633)	30.1
Alcohol use on day of injury, n (%)	1,332 (20.9)	215 (2.4)	60.2
Preexisting conditions, n (%)			
Cerebrovascular disease	826 (12.9)	1,002 (11.4)	4.6
Ischemic heart disease	425 (6.7)	655 (7.4)	2.7
Liver cirrhosis	120 (1.9)	85 (1.0)	7.5
Dementia	466 (7.3)	1,490 (16.9)	29.8
Use of anticoagulant	177 (2.8)	183 (2.1)	4.5

hol use on day of injury and use of anticoagulant as covariates yielded the same conclusions (p < 0.0001, OR = 1.75, 95% CI = 1.43 to 2.14).

In addition, we also examined sex differences in seriously injured body regions (AIS \geq 3) among patients experiencing ground-level falls. Compared to female patients, male patients had a significantly greater frequency of serious injury to the head, thorax, abdomen, and spine, resulting in more surgical interventions to the head and abdomen (Table 3). However, female patients had a higher frequency of serious injury to the upper and lower extremities, resulting in more bone fixation procedures compared to those performed in male patients (Table 3). We next investigated sex differences in mortality using a stratified analysis by injury location. There was no significant sex difference in mortality in serious head, thorax, abdomen, or spine injury, while female patients who had serious lower extremity injury had significantly lower mortality compared to male patients (Table 4). It seems probable that the higher prevalence of injuries to the head, thorax, and spine in

males, as well as higher prevalence of lower extremity with lower mortality in females, might explain the overall greater mortality in males.

DISCUSSION

Overall sex differences in mortality among trauma patients have been tested previously in multiple cohorts. In this study, when analyzing overall trauma mortality, we observed that male patients had a greater mortality rate compared to female patients. In accordance with our findings, a univariate analysis of all trauma patients admitted to three Level I trauma centers in the United States revealed that male patients had a significantly higher mortality rate than female patients (mortality, male vs. female, 0.07 vs. 0.05, p < 0.002 by chi-square test, patients younger than 50 years of age, n = 15,839). However, two other single-institution studies from Level I trauma centers in Denver, Colorado, and Maryland reported no significant differences in mortality by sex via univariate analyses

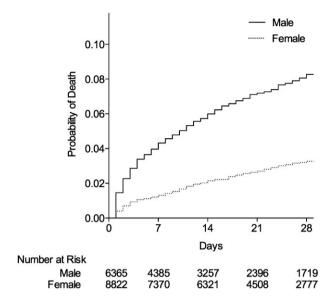


Figure 1. Survival curves by sex during a 28-day period for patients with ground-level falls.

Table 3 Frequencies of Serious Injury (Abbreviated Injury Scale [AIS] of \geq 3) and Surgical Intervention in the Cohort of Ground-level Falls

Variable	Male (n = 6,385)	Female (<i>n</i> = 8,822)	p-value
AIS score ≥ 3 Head Thorax Abdomen Spine Upper extremity Lower extremity Surgical intervention Head Thorax Abdomen Bone	0.456	0.185	<0.0001
	0.071	0.021	<0.0001
	0.017	0.009	<0.0001
	0.101	0.031	<0.0001
	0.026	0.039	<0.0001
	0.248	0.659	<0.0001
	0.092	0.036	<0.0001
	0.002	0.001	0.067
	0.005	0.003	0.010
	0.232	0.556	<0.0001

(Denver, ISS > 15, n = 545—mortality, male vs. female, 0.093 vs. 0.096, p = 0.87 by chi-square test; Maryland, all trauma patients, n = 18,892—0.048 vs. 0.047, p = 0.93 by chi-square test). Nonetheless, these studies showed that male trauma patients had a significantly increased risk of infections. A recent multivariate logistic regression analysis in a statewide trauma cohort demonstrated a nonsignificant sex difference (p = 0.093, n = 22,324). Because the above-mentioned published studies did not include analysis by injury type, our significant findings on mortality difference by sex after falls has not been described.

We screened 10 injury categories and identified that a mortality difference by sex was only observed in the fall category, including its three subcategories according to the height of such falls (Table 1). Falls are the number one cause of geriatric trauma,²⁶ and surpassing traffic crashes, falls suffered by seniors both are the leading cause of spinal cord injury in the United States and have a trend of rising incidence.²⁷ In accordance with our results in the fall category, the Centers for Disease Control and Prevention (CDC) reported that male patients experiencing falls had a higher mortality rate than did female patients,²⁸ although the CDC report did not include subcategory analysis like ours. We found that, of the three fall subcategories, sex difference in mortality was significant only in ground-level falls.

Spaniolas et al.²⁹ studied ground-level falls using the National Trauma Databank of the American College of Surgeons and reported that fall patients older than 70 years of age with GCS scores of <15 had an increased mortality rate. Since the severity and hospital mortality rate of the American study²⁹ were slightly different from our Japanese study (Japanese vs. American, average ISS 12 vs. 8, hospital mortality 0.056 vs. 0.032), our study results are not generalizable to U.S. mechanical falls victims. However, interestingly, higher mortality in males after ground-level falls was similarly observed in the American study (mortality, male vs. female, 0.048% vs. 0.023%).

Ground-level falls have characteristics of lower magnitude of injury energy,²⁹ and higher occurrence in elderly patients,³⁰ compared to medium- or high-level

Table 4 28-day Mortality of Patients Who Had Serious Injury (Abbreviated Injury Scale [AIS] of ≥ 3) and Surgical Intervention in the Cohort of Ground-level Falls

Measure, Injury Location	Mortality (Death/Total Cases)			
	Total (<i>N</i> = 15,207)	Male (n = 6,385)	Female Mortality (n = 8,822)	p-value
AIS score ≥ 3				
Head	0.120 (545/4,543)	0.125 (364/2,910)	0.111(181/1,633)	0.16
Thorax	0.038 (24/635)	0.047 (21/451)	0.016 (3/184)	0.11
Abdomen	0.049 (9/185)	0.037 (4/109)	0.066 (5/76)	0.49
Spine	0.017 (16/917)	0.020 (13/642)	0.011 (3/275)	0.42
Upper extremity	0.012 (6/511)	0.006 (1/168)	0.015 (5/343)	0.67
Lower extremity	0.014 (101/7,403)	0.028 (44/1,586)	0.010 (57/5,817)	< 0.0001
Surgical intervention				
Head	0.166 (150/905)	0.176 (104/590)	0.146 (46/315)	0.24
Thorax	0.267 (4/15)	0.300 (3/10)	0.200 (1/5)	1.0
Abdomen	0.105 (6/57)	0.029 (1/34)	0.217 (5/23)	0.034
Bone	0.007 (42/6,380)	0.010 (15/1,479)	0.006 (27/4,901)	0.053

falls. We speculate that there are possible links between these two characteristics and the sex differences in mortality of ground-level falls.

Female patients have been reported to more likely fall first on their hips, and sex differences in fall direction have also been reported.³¹ In accordance with these observations, we found that female patients experienced more serious injuries to the lower extremities than did male patients. In addition, a slow gait speed was more likely to result in an impact on the hip, 32 and gait speed was reported to be lesser in women than in men.³³ Thus, gait speed might be a potential factor for the mortality difference by sex. Male patients experienced more serious injuries to the head and spine than did female patients in this study. Similar to our findings, male sex was reported to be a risk factor for head injury in patients with cervical spine fracture resulting from ground-level falls.³⁴ Our results suggest the possibility that the kinematics of ground-level falls may differ between sexes and deserves further exploration.

LIMITATIONS

The investigation was designed as a retrospective study. Additionally, although our results indicate that compared to female patients, male patients have an increased mortality rate and a higher frequency of serious injury to the head, thorax, and spine, these two significant findings do not prove causation. Another consideration might be that death may not have been directly related to the trauma. In addition, there is a possibility that the trauma database did not capture patients who received significant blunt trauma to any part of their body and eventually fell, in the "fall" trauma category. Further studies including detailed subcategories of fall would strengthen the study results.

CONCLUSIONS

Our analysis revealed that compared to female trauma patients, male trauma patients have a greater 28-day mortality rate, particularly in patients with injuries caused by ground-level falls. Significant differences in seriously injured body regions were observed between male and female patients experiencing ground-level falls. Compared to female patients, male patients had an increased frequency of serious injuries to the head, thorax, and spine, and more surgical intervention of head and abdomen, yet a decreased frequency of injuries to the extremities. These results suggested the potential importance of sex in fatalities resulting from falls.

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