Analysis of medication and prescription background risk factors contributing to oral medication administration errors by nurses
A case–control study

Ryohei Suzuki, BS
Takamasa Sakai, PhD
Mariyo Kato, MS
Masaaki Takahashi, PhD
Akira Inukai, MD, PhD
Fumiko Ohtsu, PhD

Abstract
Medication errors, including overdose and underdose, have a significant impact on patients and the medical economy. We need to prevent or avoid recurring medication errors. Therefore, we conducted a survey to identify medication and prescription background risk factors contributing to the administration of medication by nurses. This study surveyed cases of medication administration errors. This study was conducted at Higashinagoya National Hospital from April 1, 2018, to October 31, 2019. Patients' backgrounds and medication and prescription background risk factors were investigated. Three control cases were randomly selected for each medication error case. We defined the group of medication error cases as the medication error group and the group of control cases as the no-medication-error group. A logistic regression analysis was performed for factors related to medication errors. A total of 202 patients were included in the medication error group. The median age and number of medications were 78 years and 7, respectively. A total of 606 cases were included in the no-medication-error group. The median age and number of medications were 77 years and 6, respectively. The factors that exhibited a relationship with the medication error group were the number of administrations per day, dosing frequency on indicated days, prescription and start dates were the same, medications from multiple prescriptions, and continuous use of a medication received prior to admission.

This study identified existing medication and prescription background risk factors. Overlapping risk factors from these groups might contribute to medication administration errors. Therefore, reviewing these factors is necessary to avoid recurring medication administration errors.

Abbreviation: MRCI = Medication Regimen Complexity Index.

Keywords: administration error, case-control study, contributing factor, medication factor, prescription factor

1. Introduction
Medication errors are the most commonly reported errors in hospitals.[1] The most frequently reported medication errors, including incorrect dose, incorrect medication, and omitted administration, occur during medication administration.[12–15] Medication overdose, possibly due to administration of an incorrect dose, enhances the effect of the medication and sometimes leads to adverse effects, unnecessary hospitalization, and increased medical costs.[6–8] Conversely, medication underdose, possibly due to omitted administration or administration of an incorrect dose, delays the patient’s treatment, deteriorates the condition, and prolongs hospital stay. Therefore, medication errors have a significant impact on patients and the medical economy; thus, it is necessary to avoid them and prevent their recurrence.

Medical professionals, mainly nurses, often manage oral medications in hospitals. This is because patients with deteriorated condition or decline in cognitive function are unable to self-manage medication. Furthermore, the administration of medication, such as narcotics, requires careful management. Common medications whose incorrect administration can result in medication errors include cardiovascular drugs, neurological drugs, anticoagulants, nonsteroidal anti-inflammatory drugs, and methotrexate.[8–11] The association between the number of medications and medication errors has also been reported.[6–9]

The authors have no conflicts of interest to disclose.

The datasets generated during and/or analyzed during the current study are available from the corresponding author on reasonable request.

This study was approved by the Medical Ethics Committee of the National Hospital Organization Higashinagoya National Hospital (no. 2-17). All methods were performed in accordance with relevant guidelines and regulations.

* Correspondence: Ryohei Suzuki, Graduate School of Pharmacy, Meijo University, 150 Yagotoyama, Tempaku-ku, Nagoya, Aichi, 468–8503, Japan (e-mail: 194331506@ccmailg.meijo-u.ac.jp).

Copyright © 2022 the Author(s). Published by Wolters Kluwer Health, Inc. This is an open access article distributed under the Creative Commons Attribution License 4.0 (CCBY), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

How to cite this article: Suzuki R, Sakai T, Kato M, Takahashi M, Inukai A, Ohtsu F. Analysis of medication and prescription background risk factors contributing to oral medication administration errors by nurses: A case–control study. Medicine 2022;101:33(e30122).

Received: 14 February 2022 / Received in final form: 29 June 2022 / Accepted: 1 July 2022
http://dx.doi.org/10.1097/MD.00000000000030122
Risk factors, such as identical prescription and treatment initiation dates, as well as medication that were prescribed by >1 physician and/or by the same physician at different times are important risk factors during medication administration errors in clinical practice. However, studies on the analysis of these risk factors have not been sufficiently conducted.

Clarifying the risk factors of medication errors is important for preventing the recurrence of medication administration errors, ensuring patient safety, and improving healthcare quality. Therefore, this study was conducted in an effort to identify the existing medication and prescription background risk factors related to medication administration errors.

2. Methods

2.1. Reporting system for medication administration error cases

At Higashinagoya National Hospital, all nurses who encountered or were related to a medication error promptly reported errors to their supervisor or departmental risk manager. They also wrote a medication error report.

We collected medication error reports of 2 types: for cases where patients were not subjected to incorrect actions by nurses, and for cases where patients were subjected to incorrect actions by nurses, regardless of their impact. The medication error reports included the date of occurrence, patient age, sex, specific details of the case, and countermeasures.

2.2. Surveyed cases

We surveyed cases of oral medication administration errors between April 1, 2018, and October 31, 2019. We excluded the same case reported in duplicate. The American Society of Health-System Pharmacists defines medication administration errors as the same case reported in duplicate. The American Society of Health-System Pharmacists defines medication administration errors as the same case reported in duplicate. We excluded the following medication and prescription background factors.

2.3. Research items

This study retrospectively surveyed medication error factors such as patient background and medication, prescription background based on medication error reports, electronic medical records, and dispensing records such as prescriptions.

2.3.1. Patient background. This study surveyed the following parameters for patient background from medication error reports and electronic medical records: age, sex, primary medical department, modified Rankin Scale score, and route of administration.

2.3.2. Medication and prescription background. This study surveyed the following medication and prescription background factors from medication errors based on medication error reports, electronic medical records, and dispensing records: number of medications, number of administration per day, dosing frequency on indicated days, prescription and start dates were the same, orders prescribed by >1 physician and/or by the same physician at different times (medications from multiple prescriptions), continuous use of a medication received prior to admission, and not 1 package or 1 tablet at each dosage.

2.4. Research design and statistical methods

We conducted a case–control study to analyze the factors associated with medication administration errors. A patient who was confirmed to have a medication administration error was considered in the medication error case. The corresponding control cases were randomly selected from patients in the same ward on the same day that the medication administration error case was confirmed in the medication error cases. Additionally, we excluded self-administered cases, in which nurses did not manage medications for the corresponding control cases. Three control cases were randomly selected for each medication error case. The group with medication error cases was defined as the medication error group, and the group with control cases was defined as the no-medication-error group.

We compared medication error factors between the 2 groups using univariate logistic regression analysis. In addition, multivariate logistic regression analysis was performed by selecting patient background, medication and prescription background factors (P < .20 in univariate logistic regression analysis). The number of samples required for logistic regression analysis is >10 times the number of explanatory variables to be included in the prediction model.[13] Then, we conducted a logistic regression analysis, provided that the sample size was at least 10 times the number of explanatory variables. Additionally, we calculated the correlation coefficients of each factor to check multicollinearity. Subsequently, multivariate logistic regression analysis was performed after confirming the absence of any correlation between each factor.

We also calculated medication and prescription background factors that showed a relationship in multivariate logistic analysis for medication error group and no-medication-error group. We compared the number of medication factors in the medication error group and no-medication-error group using the Mann–Whitney U test. The number of medications was a medication factor if the number of medications was 5 or more from the aspect of polypharmacy,[14] and if the number of administrations per day was 3 or more from the aspect of adherence.[15]

The significance level was set at 5%. IBM SPSS Statistics Version 27 (IBM Corporation, Armonk, NY, USA) was used to analyze the data.

2.5. Ethical considerations

This study was approved by the Ethical Review Committee of the National Hospital Organization Higashinagoya National Hospital.

3. Results

3.1. Summary of medication administration error cases

There were 362 cases of medication errors. This study excluded 79 cases of inadequate medication management, 9 cases caused by medication to be administered as needed, 42 cases of inadequate prescription order or dispensing error, 17 cases of insufficient information, and 13 cases of incorrect patient error; thus, a total of 202 medication administration error cases were analyzed in the medication error group. There were 606 cases in the no-medication-error group (Fig. 1).

3.2. Patient background, medication, and prescription background

Table 1 shows the patient background, medication, and prescription background factors for each group. The median age
Medication error cases related to oral medication
n=362

Medication error cases not related to oral medication administration or
medication and prescription background factors

Cases of inadequate medication management: n=79
Cases of inadequate prescription order or dispensing error: n=42
Cases of insufficient information: n=17
Cases of incorrect patient error: n=13
Cases caused by medication to be administered as needed: n=9

Oral medication administration error cases reported
by nurses n=202

Random selection of controls

Control cases were patients whose medication was being managed
by nurses in the same ward on the same day the medication error was
confirmed in the medication case.

1 : 3

Medication error group
n=202

No-medication-error group
n=606

Figure 1. Summary of data extraction.

Table 1
Patient background, medication and prescription background factors.

|                               | Medication error group (n = 202) | No-medication-error group (n = 606) | P value |
|-------------------------------|----------------------------------|-----------------------------------|---------|
| Age, median (IQR), yr         | 78 (67–83)                      | 77 (67–84)                        | .9*     |
| Sex, n (%)                    |                                  |                                   |         |
| Men                           | 112 (55)                        | 282 (47)                          | .03†    |
| Women                         | 90 (45)                         | 324 (53)                          |         |
| Primary medical department, n (%)|                                 |                                   |         |
| Neurology                     | 99 (49)                         | 288 (48)                          | .71†    |
| Respiratory                   | 29 (14)                         | 91 (15)                           | .82†    |
| Pediatrics                    | 22 (11)                         | 63 (10)                           | .84†    |
| Neurosurgery                  | 20 (10)                         | 63 (10)                           | .84†    |
| Orthopedic surgery            | 19 (9)                          | 64 (11)                           | .64†    |
| Others                        | 13 (6)                          | 37 (6)                            | .87†    |
| mRS, n (%)                    |                                  |                                   |         |
| 0–3: Able to walk independently| 52 (26)                         | 92 (15)                           | <.01†   |
| 4–5: Unable to walk independently| 150 (74)                      | 514 (85)                          |         |
| Route of administration, n (%)|                                  |                                   |         |
| Oral                          | 156 (77)                        | 432 (71)                          | .10†    |
| Not oral                      | 46 (23)                         | 174 (29)                          |         |
| Number of medications, median (IQR) | 7 (6–10)                     | 6 (4–6)                           | <.01*   |
| Number of administrations per day, median (IQR) | 4 (3–5)                      | 3 (3–4)                           | <.01*   |
| Dosing frequency on indicated days, n (%) | 6 (4)                        | 5 (1)                             | <.01†   |
| Prescription and start dates were the same, n (%) | 38 (19)                       | 34 (6)                            | <.01†   |
| Medications from multiple prescriptions, n (%) | 136 (67)                      | 271 (45)                          | <.01†   |
| Continuous use of a medication received prior to admission, n (%) | 74 (37)                       | 110 (18)                          | <.01†   |
| Not 1 package or 1 table at each dosage, n (%) | 189 (94)                      | 525 (87)                          | <.01†   |

IQR = interquartile range, mRS = modified Rankin Scale.

* Mann-Whitney U test.
† χ² test.
of the medication error group was 78 years (112 men and 90 women). The median age of the no-medication-error group was 77 years (282 men and 324 women). The primary medical departments included neurology and respiratory in both groups. The median number of medications was 7 in the medication error group and 6 in the no-medication-error group. The median number of administrations per day was 4 in the medication error group and 3 in the no-medication-error group.

The medication error factors that exhibited $P < .2$ in univariate logistic regression analysis were sex, modified Rankin Scale, route of administration, number of medications, number of administrations per day, dosing frequency on indicated days, prescription and start dates were the same, medications from multiple prescriptions, continuous use of a medication received prior to admission, and not 1 package or 1 tablet at each dosage.

Table 2 shows the results of multivariate logistic regression analysis. Medication error factors that exhibited a relationship with the medication error group were the number of dose administrations per day, dosing frequency on indicated days, prescription and start dates were the same, medications from multiple prescriptions, and continuous use of a medication received prior to admission. The medication error group in this study included 202 cases, which was 10 times more than the explanatory variables in the multivariate logistic analysis; thus, the required sample size was satisfied. Moreover, the correlation coefficients were <0.6 for each factor, and multicollinearity was not observed.

### 3.3. Number of medication and prescription background factors that showed a relationship with medication administration error

The medication and prescription background factors that showed a relationship with medication errors were number of administrations per day, dosing frequency on indicated days, prescription and start dates were the same, medications from multiple prescriptions, and continuous use of a medication received prior to admission. The median number of medication error factors that showed a relationship with the medication error group was 2, and that with the no-medication-error group was 1 and was significantly higher in the medication error group (Table 3).

### 4. Discussion

This study identified medication and prescription background risk factors in medication administration errors. It was found that risk factors, such as the number of administrations per day, dosing frequency on indicated days, prescription and start dates were the same, medications from multiple prescriptions, continuous use of a medication received prior to admission showed a relationship with medication administration errors.

The number of administrations per day showed a relationship with medication administration errors. There are multiple parameters that need attention during medication administration, such as understanding the medication instructions, confirming the medication route, dose, and medication name, and administering the medication.[13,16,17] When the number of administrations per day became high, the chances for omissions and mistakes by nurses in each check also increased. This may have been a factor in medication administration errors.

The presence of prescription and start dates were the same showed a relationship with medication administration errors. It is necessary to add or review medication when there are changes in the patient’s condition and when the prescription is suddenly changed. Therefore, insufficient communication between physicians and nurses, and among nurses may have had an effect. It has been empirically understood that prescription date and start date were the same factors in medication administration errors. However, actual data are not shown. This study was able to clarify that prescription and start dates were the same factors in medication error. Medications with prescription and start dates were same and must be administered promptly to patients. This is because they are urgent medications for the patient’s condition. Therefore, it is difficult to avoid medications with prescription and start dates were the same in actual clinical practice. The importance of communication between medical professionals has also been suggested to avoid the occurrence of medication errors.[18] Therefore, it is necessary to accurately communicate information and confirm the latest prescription details of patients, especially when the medication prescription and start dates were the same.

The presence of medications from multiple prescriptions showed a relationship with medication administration errors. The reason for this is thought to be the addition or change of medications owing to diagnosis by multiple department physicians or changes in medical conditions. This may result in multiple prescription configurations at each time of administration, making drug management more complex. This also supports the WHO’s Patient Safety Curriculum Guide for Medical Schools and points out that multiple prescriptions from different physicians should be considered as a background factor for patients and should be considered when a medication error occurs.[19] Moreover, the dosing frequency on indicated days of medication showed a relationship with medication administration errors. Administering daily medications is an integral part of

### Table 2

Results of multivariate logistic regression analysis of patient background, medication and prescription background factors.

|änge | Medication error group (n = 202) | No-medication-error group (n = 606) | Odds ratio | 95% CI | P value |
|---|---|---|---|---|---|
| Sex, n (%) | | | | | |
| Men | 112 (55) | 282 (47) | 1.4 | 1.0–2.0 | .06 |
| mRS, n (%) | | | | | |
| 0–3 | 52 (26) | 92 (15) | 1.4 | 0.9–2.2 | .13 |
| Route of administration, n (%) | | | | | |
| Oral | 156 (77) | 432 (71) | 0.9 | 0.6–1.4 | .74 |
| Number of medications, median (IQR) | 7 (6–10) | 6 (4–8) | 1.1 | 1.0–1.1 | .11 |
| Number of administrations per day, median (IQR) | 4 (3–5) | 3 (3–4) | 1.2 | 1.1–1.4 | <.01 |
| Dosing frequency on indicated days, n (%) | 8 (4) | 5 (1) | 6.3 | 1.9–21.0 | <.01 |
| Prescription and start dates were the same, n (%) | 38 (19) | 34 (6) | 2.8 | 1.7–4.8 | <.01 |
| Medications from multiple prescriptions, n (%) | 136 (67) | 271 (45) | 1.7 | 1.2–2.4 | <.01 |
| Continuous use of a medication received prior to admission, n (%) | 74 (37) | 110 (18) | 2.0 | 1.3–3.0 | <.01 |

CI = confidence interval, IQR = interquartile range, mRS = modified Rankin Scale.
daily practice. However, irregularly administered medications, such as medications administered on alternate days or less frequently, are different from routine daily doses. As a result, nurses tend to omit tasks such as conformation, thus leading to medication errors. Therefore, it is necessary to share information on patients’ latest prescriptions and dose timings to prevent medication errors.

It has been reported that the number of medications is a factor in medication errors. However, the results of this study did not show a relationship with the number of medications used. Previous studies have focused on the number of medications. However, the prescription context, such as medications from multiple prescriptions or prescription and start dates were the same, has not been sufficiently examined. In view of this, it is necessary to confirm not only the number of medications but also the medication factors, such as the prescription background, when the number of medications is high. These medication factors need to be reduced.

The number of medication and prescription background factors was significantly higher in the medication error group than in the no-medication-error group. The Medication Regimen Tors was significantly higher in the medication error group than in the no-medication-error group. The IQR = interquartile range. *Mann–Whitney U test.

| Medication error group | No-medication-error group | P value |
|------------------------|---------------------------|--------|
| Number of medication and prescription background factors, median (IQR) | 2 (1–3) | 1 (1–2) | <.01* |

5. Conclusions

This study identified the medication and prescription background risk factors contributing to medication administration errors: the number of administrations per day, dosing frequency on indicated days, prescription and start dates were the same, medications from multiple prescriptions and continuous use of a medication received prior to admission. A few studies have analyzed medication administration errors from the viewpoint of medication and prescription background factors. The results of this study are expected to contribute to the prevention of medication administration errors.

Acknowledgments

We thank the participants of this study for their time and thoughtful contributions as well as Sanae Niwa (Department of Nursing) and Yuko Inoue (Department of Nursing) for their comments on earlier drafts of this manuscript.

Author contributions

Conceptualization: Ryohei Suzuki, Takamasa Sakai, Fumiko Ohtsu.
Data curation: Ryohei Suzuki, Mariyo Kato, Masaaki Takahashi.
Formal analysis: Ryohei Suzuki, Takamasa Sakai, Fumiko Ohtsu.
Investigation: Ryohei Suzuki, Mariyo Kato, Masaaki Takahashi.
Methodology: Ryohei Suzuki, Takamasa Sakai, Akira Inukai, Fumiko Ohtsu.
Project administration: Ryohei Suzuki, Masaaki Takahashi, Akira Inukai, Fumiko Ohtsu.
Supervision: Masaaki Takahashi, Akira Inukai, Fumiko Ohtsu.
Visualization: Suzuki Ryohei, Takamasa Sakai, Mariyo Kato, Fumiko Ohtsu.
Writing – original draft: Suzuki Ryohei, Takamasa Sakai, Mariyo Kato, Fumiko Ohtsu.
Writing – review & editing: Masaaki Takahashi, Akira Inukai, Fumiko Ohtsu.

References

[1] Japan Council for Quality Health Care. Project to collect medical near-miss/ adverse event information 2019 annual report Available at: https://www.med-safe.jp/pdf/year_report_enlish_2019.pdf [access date June 19, 2021].
[2] Cottell M, Wätterby-Jörg I, Halleberg Nyman M. Medication-related incidents at 19 hospitals: a retrospective register study using incident reports. Nurs Open. 2020;7:1526–35.
[3] Tang FI, Sheu S-J, Yu S, et al. Nurses relate the contributing factors involved in medication errors. J Clin Nurs. 2007;16:447–57.
[4] Fathi A, Sheu SJ, Yu S, et al. Medication errors among nurses in teaching hospitals in the west of Iran: what we need to know about prevalence, types, and barriers to reporting. Epidemiol Health. 2017;39:e2017022.
[5] Kraehnibuhl-Melcher A, Schlienger R, Lampert M, et al. Drug-related problems in hospitals: a review of the recent literature. Drug Saf. 2007;30:379–407.
[6] Morimoto T, Sakuma M, Matsu K, et al. Incidence of adverse drug events and medication errors in Japan: the JADE study. J Gen Intern Med. 2011;26:148–53.
[7] Walsh EK, Hansen CR, Sahn LJ, et al. Economic impact of medication error: a systematic review. Pharmacoepidemiol Drug Saf. 2017;26:481–97.
[8] Damen NL, Baines R, Wagner C, et al. Medication-related adverse events during hospitalization: a retrospective patient record review study in the Netherlands. Pharmacoepidemiol Drug Saf. 2017;26:32–9.
[9] Berdot S, Sabatier B, Gillaizeau F, et al. Evaluation of drug administration errors in a teaching hospital. BMC Health Serv Res. 2012;12:60.
[10] Härkänen M, Paananen J, Murrells T, et al. Descriptions of incident reports. BMJ Health Serv Res. 2019;19:791.
[11] Saeddeker EA, Brock B, Nielsen LP, et al. Identifying high-risk medication: a systematic literature review. Eur J Clin Pharmacol. 2014;70:637–45.
[12] Billsle-Beleer M, Carrillo CJD, Cassano AT, et al. ASHP guidelines on preventing medication errors in hospitals. Am J Hosp Pharm. 1993;50:305–14.
[13] Peduzzi P, Concato J, Kemper E, et al. A simulation study of the number of events per variable in logistic regression analysis. J Clin Epidemiol. 1996;12:1373–9.
[14] Masnoon N, Shabib S, Kalisch-Ellelt L, et al. What is polypharmacy? A systematic review of definitions. BMC Geriatr. 2017;17:230.
[15] Claxton AJ, Cramer J, Pierce C. A systematic review of the associations between dose regimens and medication compliance. Clin Ther. 2001;23:1296–310.

[16] Pape TM. Applying airline safety practices to medication administration. Medsurg Nurs. 2003;12:77–93.

[17] Elliott M, Liu Y. The nine rights of medication administration: an overview. Br J Nurs. 2010;19:300–5.

[18] Wittich CM, Burkle CM, Lanier WL. Medication errors: an overview for clinicians. Mayo Clin Proc. 2014;89:1116–25.

[19] World Health Organization. WHO Patient Safety Curriculum Guide for Medical Schools. Geneva, Switzerland: WHO Press; 2009.

[20] George J, Phun YT, Bailey MJ, et al. Development and validation of the medication regimen complexity index. Ann Pharmacother. 2004;38:1369–76.