ARTICLE

Medications Prescribed and Occurrence of Falls in General Medicine Inpatients

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ABSTRACT

Background: Although falls are multifactorial, medications are a key risk factor that may be modifiable. Falls were among the most common occurrences entered into a risk identification system at the authors' hospital.

Objectives: To identify whether general medicine inpatients who had experienced a fall were taking any medications known to be associated with falls.

Methods: The literature was reviewed to develop a list of high-risk medications that have been associated with falls. In a retrospective quality-improvement database-based study, information from the risk identification system was merged with data from the pharmacy dispensing system for general medicine inpatients who had experienced a fall. The primary end point was the percentage of patients with a documented fall who had a prescription for a high-risk medication. The number of such medications that had been prescribed for patients who fell was also calculated.

Results: Eighty-one unique medications were found to be associated with falls. During the study period (April 1, 2008, to March 31, 2009), 151 patients experienced a fall. Of those, 144 (95.4%) were taking at least one high-risk medication. The mean number of high-risk medications per patient who experienced a fall was 2.2. Of all documented falls, a new high-risk medication had been started within 7 days before the fall for 74 (49.0%) and within 24 h before the fall for 17 (11.3%). The most commonly prescribed drugs during all time periods (i.e., within 24 h or 7 days before the fall or since the patient's admission) were lorazepam and zopiclone. The pharmacy database did not track administration of medications, so it is possible that some of the drugs prescribed were not actually taken by the patient.

Conclusion: Almost all inpatients who experienced a fall during the hospital stay had a prescription for at least one medication associated with a high risk for falls. Lorazepam and zopiclone were the drugs most commonly associated with falls in this hospital, and their use should be reviewed.

Key words: accidental falls, pharmaceutical preparations, inpatients

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RÉSUMÉ

Contexte : Bien que la cause des chutes soit multifactorielle, les médicaments en sont un facteur de risque clé pouvant être modifié. Les chutes sont l'un des événements les plus fréquents saisis dans le système d'identification des risques à l'hôpital des auteurs.

Objectifs : Déterminer si les patients hospitalisés en médecine générale qui ont subi une chute prenaient des médicaments reconnus pour être associés aux chutes.

Méthodes : On a effectué une revue de la littérature pour établir une liste des médicaments à haut risque associés aux chutes. Une étude rétrospective fondée sur une base de données visant l'amélioration de la qualité a fusionné les données tirées du système d'identification des risques aux données issues du système de distribution de la pharmacie portant sur les patients hospitalisés en médecine générale ayant subi une chute. Le paramètre d'évaluation principal était le pourcentage de patients ayant subi une chute constatée à qui l'on avait prescrit un médicament à haut risque. Le nombre de médicaments à haut risque ayant été prescrits aux patients qui avaient subi une chute a également été calculé.

Résultats : On a dénombré 82 médicaments différents associés à des chutes. Durant la période de l'étude (du 1^a avril 2008 au 31 mars 2009), 151 patients ont subi une chute. De ceux-ci, 144 (95,4 %) prenaient au moins un médicament à haut risque. Le nombre moyen de médicaments à haut risque par patient ayant subi une chute était de 2,2. De tous les cas de chutes consignés, on avait commencé à administrer un nouveau médicament à haut risque dans les sept jours précédant la chute chez 74 (49,0 %) des patients. Les médicaments les plus souvent prescrits pour toutes les périodes de temps (c.-à-d. moins de 24 heures ou de sept jours avant la chute ou depuis l'admission du patient) étaient le lorazépam et la zopiclone. La base de données de la pharmacie ne permet pas de savoir si les médicaments ont été réellement administrés et il est donc possible que certains des médicaments prescrits n'aient pas été pris par quelques patients.

Conclusion : Presque tous les patients ayant subi une chute durant leur hospitalisation avaient reçu une ordonnance d'au moins un des médicaments associés à un risque élevé de chute. Le lorazépam et la zopiclone étaient les médicaments les plus souvent associés à des chutes dans cet hôpital et leur utilisation devrait donc être examinée.

Mots clés : chutes accidentelles, préparations pharmaceutiques, patients hospitalisés

[Traduction par l'éditeur]

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INTRODUCTION

A fall is a sudden, unintended change in position causing an individual to land on a lower level.¹ In the United States, a total of 10 300 fatal fall injuries occurred in 2000, and they incurred an estimated total cost of US\$179 million.² In the same year, an estimated 2.6 million nonfatal fall injuries resulted in a total cost of US\$19 billion.² Although falls are often multifactorial, the risk of falls can be lessened by addressing any factors that are modifiable. Because falling is a common adverse event related to medications,^{3,4} modification of a patient's medication regimen is one possible method of reducing the risk of falls.

At the Red Deer Regional Hospital Centre, a nonteaching referral hospital (with about 400 beds) in rural Alberta, falls constituted the second most common event recorded in the risk identification system. The risk identification system was available at all computer terminals in the hospital, and any event that might cause patient harm could be entered into this system by any employee. From April 1, 2008, to March 31, 2009, a total of 287 falls were recorded in the system. Of these falls, 173 (60.3%) occurred on 3 general medicine inpatient units.

Given the high costs associated with falls and the frequency of falls that occur in the hospital setting, this study was performed to identify whether general medicine inpatients who had experienced a fall were taking any medications known to be associated with falls.

METHODS

Literature Search

Two individuals (including M.Y.) independently conducted a comprehensive search of Ovid MEDLINE (1950 to May Week 3, 2009) to identify high-risk medications that have been statistically significantly associated with falling. The search strategy was based on the Medical Subject Headings (MeSH) "accidental falls" and "pharmaceutical preparations" combined with the keywords "falls", "medications", and "drugs". Articles retrieved were limited to studies in adults over 18 years of age with publication in English. One of the authors (M.Y.) screened the articles by reading the abstracts; when there was any doubt about suitability for inclusion, the full text was retrieved. Studies were included if the primary end point was identification of a statistically significant relation between the medication and an increased risk of falls. Studies were excluded if they specifically examined the adverse effects of a single medication. The medications identified through this literature search were then divided into categories created by the author team, as outlined in Table 1. High-risk medications associated with falling were defined as medications falling into categories A and B in Table 1. For situations in which only a class of medications was identified in the literature, a list of possible medications in that class was determined from the British National Formulary⁵ and the World Health Organization classification.⁶ These classes were then limited to products available in Canada.

Inclusion and Exclusion Criteria for Patients with a Fall

All patients for whom a fall had been entered into the risk identification system from April 1, 2008, to March 31, 2009, were eligible for inclusion in the study. Because of time constraints, the review was limited to patients whose falls occurred on the 3 units of the hospital with the highest frequency of falls. These data were cross-referenced with the pharmacy database of medications ordered and dispensed for each patient by matching on the basis of the patient's name and date of birth. For patients with multiple hospital stays, the correct visit was selected by matching the appropriate admission and discharge date with the date of the fall as recorded in the risk identification system. Patients who were still in hospital at the time the analysis was conducted (in July 2009), those who had been admitted to the hospital for less than 48 h, outpatients, and patients under 18 years of age were excluded.

Data Analysis and End Points

The risk identification system was used to identify all patients who fell between April 1, 2008, and March 31, 2009. The severity of injury, as determined by the individual who entered data about the fall, was also recorded. The severity of injury was categorized as follows: level 0 = near miss ("a situation that may potentially result in an incident or harm", as defined in the risk identification system), level 1 = an incident

Category	Definition
A	Medication(s) or medication class reported as significantly associated with an increased risk of falls in a meta-analysis, systematic review, or more than 2 unique observational studies
В	Medication(s) or medication class reported as significantly associated with an increased risk of falls in only 2 observational studies
С	Medication(s) or medication class reported as significantly associated with an increased risk of falls in only 1 observational study

Table 1. Categories of High-Risk Medications

with no adverse outcomes, level 2 = an incident with minor adverse outcomes (i.e., patient experienced minor harm requiring minor treatment, additional monitoring, and follow-up), level 3 = an incident with moderate adverse outcomes (i.e., patient sustained harm that was reversible by clinical interventions and that was anticipated to last less than 6 months), level 4 = anincident with a serious or major adverse outcome (i.e., patient sustained harm that was not immediately reversible by clinical interventions and that resulted in a disability anticipated to last more than 6 months), and level 5 = an incident that resulted in death. A fall was considered to have associated morbidity if the person entering the information into the risk information system classified the severity as level 2 or higher.

For each fall in the risk identification system, all high-risk medications for which the patient had a prescription were recorded and categorized in relation to 3 time periods: new medications that had been prescribed in hospital in the 24-h period before the fall, new medications prescribed in hospital in the 7-day period before the fall, and any medications that had been prescribed for the patient since admission. No analysis was performed for discontinued medications and their potential relation to falls. For patients with prescriptions for more than one high-risk medication, each medication was analyzed separately in terms of time since initiation.

During data collection, no patient identifiers were retained, and all data were analyzed as aggregates. The primary end point of the study was the percentage of falls in which the patient had a prescription for one or more high-risk medications associated with falls, for each of the 3 prespecified time frames. Other end points were a comparison of the number of falls associated with single versus multiple high-risk medications; identification and ranking of medications according to which were most commonly prescribed for patients who fell; determination of average length of stay before the time of the fall; and determination of the number of falls with no injury, any injury, or death.

RESULTS

The literature search yielded 16 unique articles, consisting of 2 meta-analyses,^{7,8} 2 systematic reviews,^{9,10} and 12 observational or retrospective studies.¹¹⁻²² As anticipated, most of the articles identified classes of medications, not individual drugs. A list of 81 unique medications associated with increased risk of a fall was extrapolated from the published information, as presented in Table 2. This list of medications was used in reviewing the medication profiles of patients who experienced a fall.

Over the 1-year study period, the 3 patient care units with the most falls accounted for 173 (60.3%) of all documented falls. These 3 units were all general medicine inpatient areas. As mentioned earlier, because of time constraints, data collection was limited to this sample of falls. Of the 173 falls, 10 were excluded from analysis because the patients had been admitted to hospital for less than 48 h, 8 falls were excluded because

Medication Class	Medications		
Benzodiazepines ^{7,9–17,19,22}	Alprazolam, bromazepam, chlordiazepoxide, clobazam, clonazepam, clorazepate, diazepam, flurazepam, lorazepam, midazolam, nitrazepam, oxazepam, temazepam, triazolam		
Hypnotics, sedatives, and anxiolytics ^{7,9–11,14,15,17,22}	Barbiturates: pentobarbital, thiopental, phenobarbital, primidone Aldehydes: paraldehyde Other: buspirone, chloral hydrate, hydroxyzine, meprobamate, scopolamine, valerian, zopiclor		
Antidepressants ^{7,9,10,12,15-19,21,22}	 Tricyclic antidepressants: amitriptyline, clomipramine, desipramine, doxepin, imipramine, nortriptyline, trazodone, trimipramine Selective serotonin reuptake inhibitors: citalopram, escitalopram, fluoxetine, fluvoxamine, paroxetine, sertraline Monoamine oxidase inhibitors: maprotiline, moclobemide, phenelzine, quinupramine, tranylcypromine Other antidepressants: bupropion, desvenlafaxine, duloxetine, mirtazapine, tryptophan, venlafaxine 		
Antipsychotics and neuroleptics ^{7,13,14,16,18-20,22}	Phenothiazines: chlorpromazine, fluphenazine, methotrimeprazine, perphenazine, pipotiazine, prochlorperazine, thioproperazine, trifluoperazine Butyrophenones: droperidol, haloperidol Diphenylbutylpiperidine: pimozide Thioxanthenes: flupenthixol, zuclopenthixol Atypical antipsychotic drugs: clozapine, olanzapine, quetiapine, risperidone Indole derivative: ziprasidone Other: lithium, loxapine, paliperidone		
Antiepileptics9,21	Phenytoin, carbamazepine		
Cardiac medications ^{8,14}	Thiazide diuretics: hydrochlorothiazide, mebutizide, trichlormethiazide Furosemide Type la antiarrhythmics: procainamide, quinidine Other: digoxin		

Table 2. Medications Classified as High Risk

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there was no patient match between the risk identification system and the pharmacy database, and 4 falls were excluded because they involved patients who were still in hospital at the time the analysis was conducted. Therefore, 151 falls were included in the analysis. The patients included in the review experienced their falls between 2 and 178 days after admission, with a mean length of stay between admission and the fall of 25.5 days. The mean age of patients who had a fall was 73.5 years, and almost equal numbers of men and women had falls (Table 3).

The percentages of patients with prescriptions for one or more high-risk medications who had a fall are shown in Figure 1. Nearly all patients who fell (144 or 95.4%) had a prescription for at least one high-risk medication, and most patients who fell had prescriptions for more than one high-risk medication (mean number of high-risk medications per patient 2.2 [Table 3]). The number of high-risk medications prescribed differed with the time interval analyzed: the shorter the time frame before the fall, the greater the chance that a single high-risk medication had been started during that time frame (Figure 1).

Regardless of the time frame, the most common medications prescribed to patients who fell were lorazepam and zopiclone. When the high-risk medications were grouped into classes, as in Table 4, benzodiazepines were the medications most commonly prescribed before a fall. Antipsychotics constituted the second most common class of medications prescribed before a fall for all time periods, despite the fact that only one antipsychotic (haloperidol) ranked among the top 5 medications prescribed before a fall.

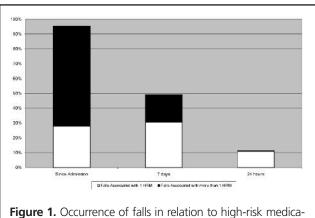
Of the 151 falls analyzed, 91 (60.3%) resulted in no injury, and 60 (39.7%) resulted in an injury with severity of at least level 2. No falls resulted in death, according to the information recorded in the risk identification system.

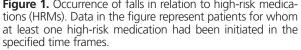
DISCUSSION

Even with the strict criteria established for "high-risk" medications in this study, there were many classes of medications associated with falling. The large number of medications

Table 3. Demographic Characteristics of Patients Who Fell

Characteristic	No. (%) of Patients or Mean (Range)			
No. of falls	151			
Age, years	73.5 (25–95)			
Sex, no. (%) male	78 (51.7%)			
High-risk medications				
≥ 1	144 (95.4%)			
≥ 2	102 (67.6%)			
Time to fall (days)				
Mean	25.5 (2–178)			
Median	14			
Mean no. of high-risk medications				
Since admission	2.2			
Started within 7 days before fall	0.8			
Started within 24 h before fall	0.1			





presents a challenge to practitioners, as it may difficult to recall such a lengthy list in day-to-day clinical practice. When we extrapolated a list of available medications from the classes of medications identified, the list consisted of 81 unique medications. What is not known at present is whether each medication in each class carries a risk of falling, or if certain medications

	Table 4. Medication C	lasses Most Often Pre	escribed among Patier	nts Who Had a Fall*
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	Time Frame of Initiation of Drug; No. (%) of Falls†			
Medication Class	24 h Before Fall	7 Days Before Fall	Since Admission	
Benzodiazepines	6 (4.0)	28 (18.5)	83 (55.0)	
Antipsychotics	5 (3.3)	23 (15.2)	65 (43.0)	
Hypnotics	3 (2.0)	19 (12.6)	58 (38.4)	
Diuretics	2 (1.3)	14 (9.3)	47 (31.1)	
SSRIs	2 (1.3)	17 (11.3)	42 (27.8)	

SSRI = selective serotonin reputable inhibitor.

*Some patients were receiving drugs from more than 1 class of high-risk medications at the time of the fall. +For analysis of 151 falls over a 1-year period. within a class are more likely to be associated with a fall. Although medications within a class may have different pharmacokinetics, their pharmacodynamics are generally similar. This similarity makes it plausible that all medications within a class would carry a similar risk for falls; however, we recognize that some medications may be more likely to be associated with falls than others. It is possible that applying a different method to identify high-risk medications might have yielded medications that were not included in our analysis but that might have an even stronger association with falls. Although the number of medications that may contribute to a fall is large, practitioners need to be able to recognize medications associated with falls when they are completing a patient's medication review upon admission to an acute care institution, so that interventions may be undertaken to lessen the chance of patients falling.

These data show that almost all patients who experienced a fall had a prescription for at least one high-risk medication. Only 49.0% of falls were associated with prescription of a new high-risk medication within the preceding 7 days, and many of the high-risk medications were prescribed on admission. Many medications prescribed at the time of admission are simply continuations of the patient's home medications, but it cannot be assumed that these medications are safe for the patient during a hospital stay. By virtue of their admission to an acute care institution, it is reasonable to assume that many patients will experience a change in pharmacokinetic parameters. In acute illness, there may be changes in absorption (e.g., decreased perfusion at the site of absorption or motility dysfunction), distribution (e.g., pH changes, fluid shifts, or changes in plasma protein binding), metabolism (e.g., disruption to hepatic blood flow), and excretion (e.g., decreased renal function).23 In addition, the disease state or disease states for which the patient is admitted may also be associated with an increased risk of falling. Although causality between medications prescribed and occurrence of falls cannot be determined from the analysis reported here, it appears that reviewing patients' medication regimens with the goal of reducing the use of medications associated with falling may be beneficial. Given that the majority of patients who fell were taking more than one high-risk medication, there is a possibility of undertaking several interventions to decrease the chance of a fall. The literature does not support the notion of an increased risk of falling with more than one high-risk medication, but expert opinion suggests that unnecessary medications be removed from the patient's regimen and that the lowest possible effective dose be used for those medications that are deemed necessary.²⁴

In this sample of falls, almost all occurred in patients with a prescription for a high-risk medication, lorazepam and zopiclone being the individual drugs associated with the greatest number of falls. Given that zopiclone is indicated only for the treatment of insomnia²⁵ and given that a common use of lorazepam is for nighttime sedation, it is possible that a number of falls may have been linked to the use of medications for this indication. Further research toward a safer procedure for nighttime sedation is warranted and may result in fewer falls.

Overall, harm was documented for about 40% of the falls, but this may be an underestimation. The severity of each fall was determined by the user of the risk identification system. It is possible that some people had injuries affecting their prognosis or leading to a longer length of stay that were missed by the reporter at the time of the incident report. The extent of such morbidity was not available from the database used. However, this possibility does suggest that actions to reduce the occurrence of falls are needed.

Many of the studies identified in the literature search involved geriatric patients, and it is uncertain whether their conclusions can be extended to the general population. However, in the present study, the mean age of patients experiencing falls was 73.5 years; as such, we believe that the method used to determine the list of high-risk medications was appropriate for the patient population studied. Another limitation of this analysis was that both aspects of data extraction (identification of high-risk medications from the literature and determination of use of high-risk medications by patients who fell) was done by a single reviewer (M.Y.). This drawback was lessened by having a librarian complete an independent literature review and provide relevant sources to the main reviewer.

The prescription data were extracted from a pharmacy database and not a computerized medication administration record. It is therefore possible that some patients were not taking the high-risk medications that had been prescribed for them. In this regard, medications that are prescribed for use on an as-needed basis are most likely to affect the results. Consequently, the data are presented as high-risk medications prescribed before a fall, and we acknowledge that some patients who had a fall may not have received all of the medications prescribed. Ideally, the timing of medication doses in relation to the fall would have been assessed for any potential correlation.

Although the reporting of incidents and near misses in the risk identification system is supported by the hospital's administration, it is possible that this system did not capture all of the events that occurred within the time frame of the study and that more falls occurred than were documented.

CONCLUSIONS

Many medications have been associated with falls, as documented in the literature, and nearly every patient who experienced a fall in 3 general medicine inpatient units in the study hospital had a prescription for at least one of these medications. Given that many such falls could cause some degree of morbidity, a review of patients' medications, to assess the risk of falling, is needed as part of a multidisciplinary fall prevention strategy. The best time for a medication review cannot be identified from this study. However, because many of the high-risk medications associated with falls appear to be started at the time of admission, this may be a reasonable time to review medications with the specific goal of decreasing the number of medications with a documented association with falling. Medications for nighttime sedation appear to be the most frequently prescribed medications before a fall, and a review of nighttime sedation procedures should also be undertaken.

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