

Original Research

Factors associated with adverse drug reactions among residents in nursing home

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Abstract

Background and Aim: Adverse drug reactions (ADRs) are a significant public health concern, especially in the elderly. Factors like multimorbidity, polypharmacy, frailty, and age-related changes in drug metabolism increase variability in drug response and adverse effects. The objective of this research is to examine the prevalence of ADRs in elderly residents of nursing homes and to identify the most significant risk factors contributing to these events. **Material and methods:** This was a retrospective, cross-sectional study that reviewed the medical records of 125 residents in 2 nursing homes. The study population consisted of all residents aged 65 years or older. Resident characteristics included age, gender, clinical diagnoses, number of comorbidities, residents' functional status, and complete medication history were analyzed using general practitioner documentation in residents' records. Relevant clinical information on ADR characteristics was documented through chart review over 12 months. **Results:** The study included 125 residents with a mean age of 76.52 (SD 7.83), the majority of them were women (66%), with polypharmacy of 5 and above (48%). Overall, 10% of elderly took ≥ 10 medications (prescribed, or over the counter), and 38% took 5–9. In addition, the mean number of chronic comorbid conditions was approximately four. Mean Charlson comorbidity index was 1.7. Residents with higher Charlson Comorbidity Index showed increased risk of ADRs, $r = .552$, $p < .001$. We found that patients with non-ADRs used less medications ($M = 3.88$, $SD = 1.56$) compared to those with ADRs, ($M = 7.26$, $SD = 1.27$; $t(123) = -6.51$, $p < .001$). Cardiovascular, gastrointestinal, and anticoagulant medications were associated to ADRs. Finally, using Pearson's r correlation coefficient we found that there is a significant positive relationship between ADRs and specific drug classes use as well as higher polypharmacy and comorbidity. **Conclusion:** This study highlights the high prevalence of ADRs among elderly patients, with a positive association to polypharmacy, comorbidities, and specific drug classes use.

Keywords: adverse drug reactions, risk factors, elderly, nursing home, polypharmacy, comorbidity

INTRODUCTION

Adverse drug reactions (ADRs) represent a significant public health concern, particularly among the elderly population. Older adults are more vulnerable to ADRs due to age-related physiological changes, the presence of multiple comorbidities, and alterations in pharmacokinetics and pharmacodynamics¹. This vulnerability is amplified in nursing home settings, where residents often suffer from multiple chronic conditions and are frequently prescribed numerous medications, a phenomenon known as polypharmacy². Polypharmacy, typically defined as the use of five or more medications concurrently, is a well-established risk factor for ADRs². Studies have shown that up to

91% of nursing home residents are affected by polypharmacy³, significantly increasing the risk of drug-drug interactions, medication errors, and adverse effects⁴. Identifying ADRs in this population is often challenging, as symptoms can be non-specific and are frequently misattributed to normal aging or existing health conditions, leading to underreporting and delayed intervention.

The prevalence of polypharmacy in long-term care facilities has increased substantially over the past two decades^{5,6}. This includes both prescribed medications and over-the-counter products. It is estimated that nearly 50% of older adults take at least one medication that is not medically necessary⁷. While medications are essential for managing chronic illnesses, their inappropriate or excessive use may lead to serious consequences, including increased morbidity, prolonged hospitalizations, and higher healthcare costs⁸. Older adults are up to four times more likely to be hospitalized due to ADRs compared to those under the age of 65 (16.6% vs. 4.1%).⁹ Contributing factors include multimorbidity, polypharmacy, frailty, and age-related changes in drug absorption, distribution, metabolism, and elimination, all of which increase variability in drug response and susceptibility to adverse effects⁹⁻¹¹.

Certain drug classes are particularly associated with a high incidence of ADRs. A systematic review identified six classes of medications—antibiotics, anticoagulants, cardiac glycosides, diuretics, hypoglycemic agents, and non-steroidal anti-inflammatory drugs (NSAIDs) as responsible for 60–70% of ADRs¹². Among hospitalized geriatric patients, common ADRs include falls and fractures, cognitive impairment,

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gastrointestinal bleeding, renal dysfunction, hypoglycemia, constipation, urinary retention, QT interval prolongation, and cardiac arrhythmias¹³. Given the high prevalence of polypharmacy and its associated risks in nursing home populations, the proactive identification and management of these risk factors is essential for reducing the incidence of ADRs and improving patient safety among frail older adults in long-term care settings.

The primary objective of this research is to investigate the prevalence of adverse drug reactions in elderly residents of nursing homes and to identify the most significant risk factors contributing to these events. Specifically, the study aims to assess the relationship between polypharmacy and the occurrence of ADRs, evaluate the impact of multimorbidity on ADR prevalence and identify the most commonly implicated drug classes associated with ADRs in this population.

MATERIALS AND METHODS

Study design

This was a retrospective, cross-sectional study that reviewed the medical records of 125 residents in 2 nursing homes.

Participants

The study population consisted of all residents aged 65 years or older.

Data collection

Data collection and clinical assessments for the present study was conducted between June 1 2024 and October 15 2024.

Outcome measures

Resident characteristics included age, gender, clinical diagnoses, number of comorbidities, patients' functional status, and complete medication history were analyzed using general practitioner documentation in patients' records.

Relevant clinical information relating to the characteristics of the ADRs was also documented. ADRs were identified by chart review during a 12-month period.

The medications were, in all cases, chronic treatments administered by oral, inhalation, or ophthalmic routes. All drugs included in this study required a prescription and had been administered to residents for at least 1 month prior to data collection. Over the counter (OTC) medications, nutritional supplements, and herbal medicines were also considered. We excluded as-needed agents, topical agents, single-dose administration, and vaccinations.

The therapeutic categories of medicines were classified according to the Anatomical Therapeutic Chemical (ATC) classification system of the World Health Organization (WHO) Collaborating Center¹⁰.

Polypharmacy is defined as the use of multiple drugs. We categorized participants into three groups based on the number of medications taken: non-polypharmacy (0–4 medicines), polypharmacy (5–9 medicines), and excessive polypharmacy

(at least 10 medicines).

Adverse drug events were defined as injuries resulting from use of a drug. A pharmacist and general practitioner from the research team assessed ADR by the severity of the event (preventable, serious, or life-threatening). Examples of preventable events included non-urticarial rashes, falls without associated fracture, hemorrhage not requiring transfusion or hospitalization, and oversedation; examples of serious events included urticaria, falls with fracture, hemorrhage requiring transfusion or hospitalization, and delirium¹⁴.

Co-morbidities were classified using International Classification of Diseases (ICD)-10 criteria (as defined by the World Health Organization). To determine the number of comorbidities that participants had, we noted all of their self-reported chronic diseases, which were of the 18 chronic diseases recorded in ELSA, including diabetes, hypertension, stroke, myocardial infarction, congestive heart failure, angina, lung disease, chronic obstructive pulmonary disease, asthma, arthritis, osteoporosis, cancer, hearing problems, Parkinson's, Alzheimer's disease, dementia, macular degeneration, and glaucoma¹⁵.

For quantification of comorbidities the Charlson Comorbidity Index (categorized using scoring as originally developed—0, 1-2, 3-4, and ≥ 5) was used¹⁶.

Data analysis

For the descriptive analysis, we calculated frequencies, percentages, as well as mean with standard deviation (SD) for the distribution of variables (see Table 1 and Table 2, and Figure 1 and Figure 2).

The relationship between quantitative variables were analyzed using the independent samples T-test analyses. We also employed Pearson's *r* correlation coefficient to examine the relationships between main variables (see Table 2).

The significance level for the different analyses was established at least at $p < 0.05$. The data analysis was performed using SPSS Statistics for Windows, version 24.

Ethical Considerations

The study was approved by the University of Prishtina, Faculty of Medicine Ethics Committee and carried out in accordance with the Declaration of Helsinki.

RESULTS

The socio-demographic and clinical characteristics of the residents by total population and ADRs experiences are shown in Table 1. The study included 125 residents with a mean age of 76.52 (SD 7.83), the majority of residents were women (66%), with polypharmacy of 5 and above (48%). Overall, 10% of elderlies took ≥ 10 medications (prescribed, or over the counter), and 38% took 5–9. In addition, the mean number of chronic comorbid conditions was approximately four. Mean Charlson comorbidity index was 1.7. The number of medications taken ranged from 0 to 12 with a median number of 5 drugs. Throughout the study period, ADRs were identified



Table 1. Characteristics of the residents by total population and ADR experiences.

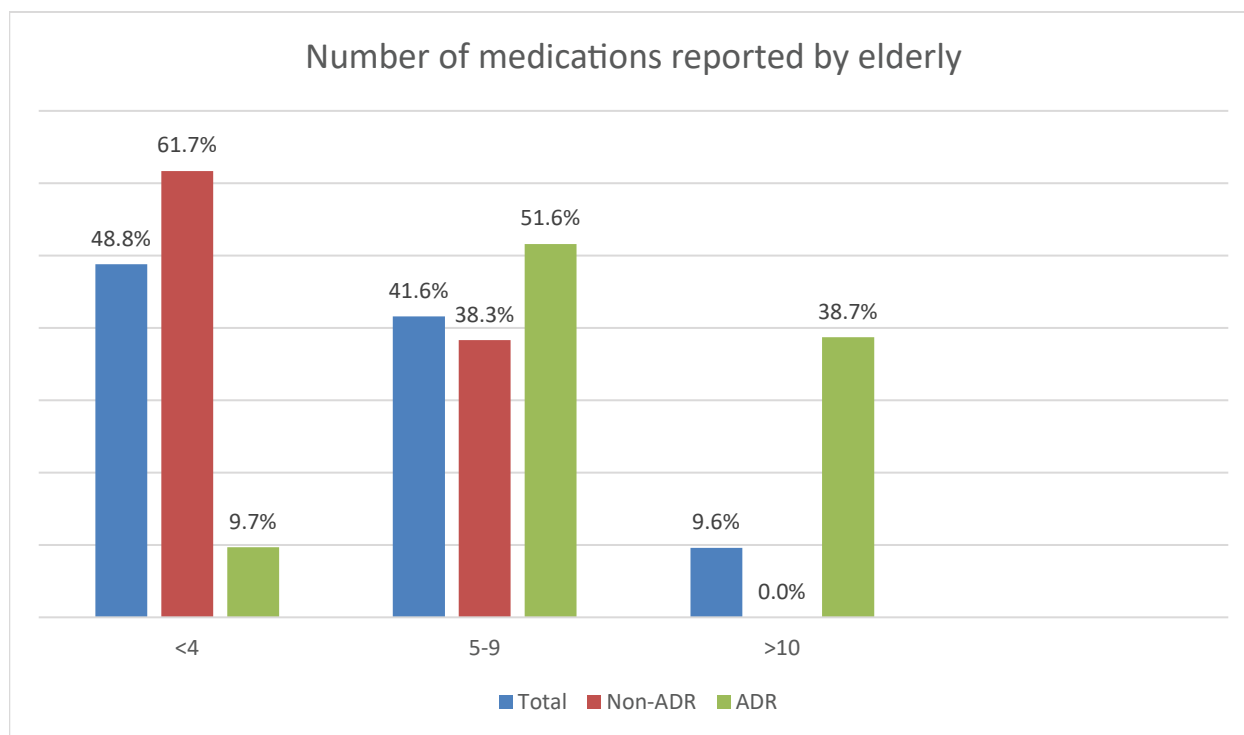
	Total		Residents in non-ADRs group		Residents in ADRs group	
	No	(%)	No	(%)	No	(%)
Characteristics						
Sex						
Male	42	34%	31	33%	11	36%
Female	83	66%	63	67%	20	64%
Age group (years)						
65-74	56		44	47%	12	39%
75-84	44	35%	32	34%	11	36%
85+	26	21%	18	19%	8	26%
Polypharmacy						
0-4	65	52%	49	39.20%	16	12.80%
05-09	48	38%	35	28%	13	10.40%
>10	12	10%	10	8%	2	1.60%
Comorbidities						
01-04	64	51.20%	16	12.80%	48	38.40%
05-08	59	47.20%	14	11.20%	45	36%
>9	2	1.60%	1	0.80%	1	0.80%
CCI						
01-02	9	15%	6	4.80%	3	2.40%
03-04	106	85%	88	70.40%	18	14.40%
>5	-		-		-	
ADR by the severity of the event						
Preventable	-		-		17	13.60%
Serious	-		-		12	9.60%
Life threatening	-		-		2	1.60%

Abbreviations:ADRs-Adverse drug reactions, Non-ADR- Non-Adverse drug reactions
CCI-Charlson Comorbidity Index

Table 2. Means, standard deviations, and correlations among main variables.

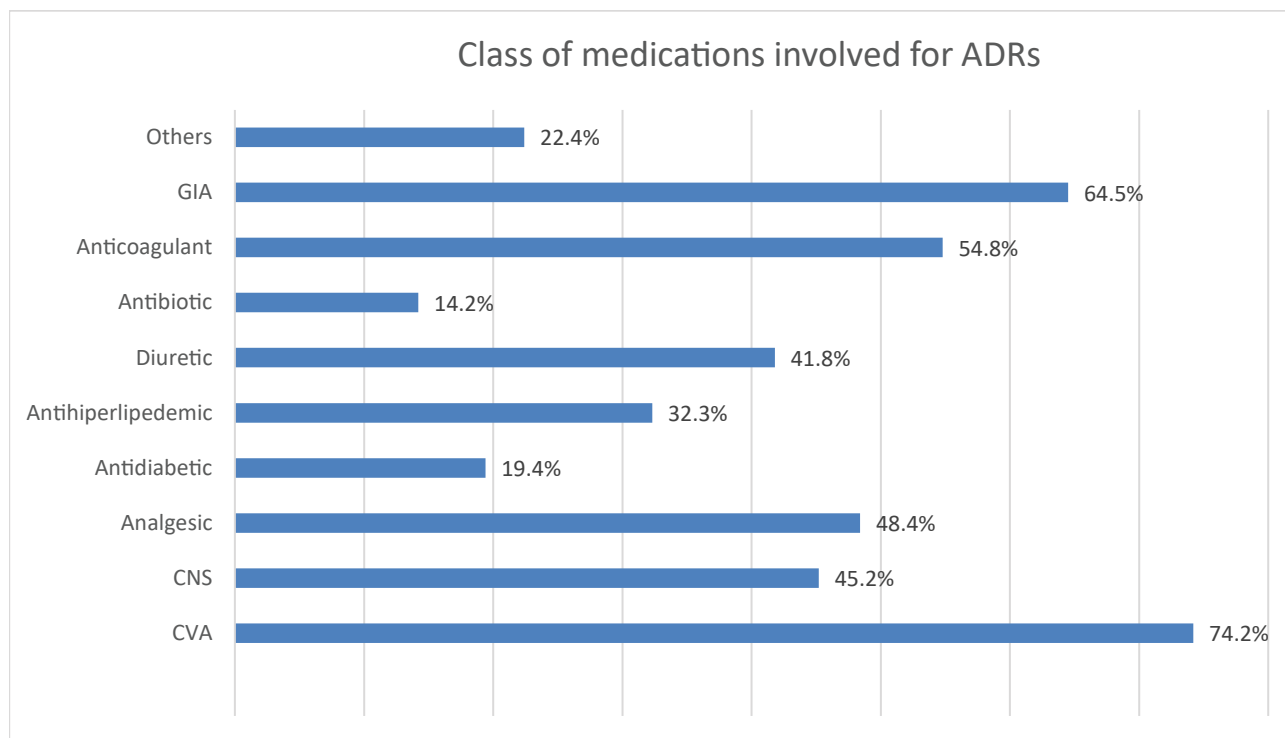
	1	2	3	4	5
1. Gender	-				
2. ADRs	-0.047	-			
3. Polypharmacy	0.082	-0.022	-		
4. Comorbidity	0.037	0.004	.221*	-	
5. Specific drug classes	0.129	.552**	0.011	0.29	-
M	-	1.51	1.58	1.5	4.72
SD	-	2.73	0.66	0.53	2.41

Abbreviations:ADRs-Adverse drug reactions, M-Mean, SD-Standard deviation
Note. Significant effects are in bold. Gender is coded as 1 (male) and 2 (female).
**p < .01, *p < .05



Abbreviations: ADRs-Adverse drug reactions, Non-ADR- Non-Adverse drug reactions

Figure 1. Number of medications reported by elderly.



Abbreviations: ADRs-Adverse drug reactions, GIA-gastrointestinal agents, CNS-central nervous system, CVA-cardiovascular agents

Figure 2. Class of medications involved for ADRs.

in 31 of the 125 cases (24.8%). Overall, 13.6% of the ADRs were considered preventable, 9.6% of the 125 classified as serious events, and 1.6% life-threatening.

Next, we examined whether the number of medications used by residents is related to ADRs. In figure 1 we have presented the prevalence of medication use among residents with adverse and non-ADRs. The figure shows that residents who reported ADRs also reported higher number of medication (5-9 medications, 51.6% ADRs) use compared to who reported no ADRs (38.3%). We performed an independent samples t-test to determine whether number of medications use significantly differs between those with adverse and non-ADRs residents. We found that patients with non-ADRs used less medications ($M = 3.88$, $SD = 1.56$) compared to those with ADRs, ($M = 7.26$, $SD = 1.27$; $t(123) = -6.51$, $p < .001$). This result indicates that increase use of medication increases the ADRs among residents.

We performed another series of independent sample's t-tests to determine the relationship between types of medication and adverse versus non-ADRs. We found that residents who used CVA reported more ADRs, ($M = 2.66$, $SD = 3.26$), compared to those with non-ADRs, ($M = .82$, $SD = 2.08$); $t(123) = 3.46$, $p = .001$. Similarly, residents who used gastrointestinal agents reported more ADRs ($M = 2.24$, $SD = 3.02$) than those with non-ADRs, ($M = 1.10$, $SD = 2.47$); $t(123) = 2.29$, $p = .034$; and residents who used anticoagulant reported more ADRs ($M = 2.16$, $SD = 3.09$) compared to those with non-ADRs, ($M = 1.09$, $SD = 2.40$); $t(123) = 2.18$, $p = .042$. These results indicate that these specific medications, CVA, gastrointestinal agents, and anticoagulant can cause ADRs in residents. In contrast, we found that residents who used diuretics, analgesics, antihyperlipidemics, central nervous system agents, and antidiabetics did not report more ADRs compared to residents with non-ADR (all p values were above .05). The detailed analyses of the medications categories are presented in Figure 2. Cardiovascular agents were most frequently implicated with an ADRs (74.2%), followed by gastrointestinal agents (64.5%) and anticoagulant drugs (54.8%).

Finally, we examined whether higher ADRs relate to gender, polypharmacy, comorbidity, and specific drug classes using Pearson's r correlation coefficient (see Table 2). We found that there is a significant positive relationship between ADRs and specific drug classes use as well as higher polypharmacy and comorbidity. These results indicate that residents with more adverse drug effects also reported more specific medication use, and residents with higher polypharmacy also reported more comorbidity. Residents with higher Charlson Comorbidity Index showed increased risk of ADRs, $r = .552$, $p < .001$. Other relations were not significant.

DISCUSSION

This study revealed a high burden of multimorbidity, polypharmacy, and ADRs among nursing home residents. Nearly half of the cohort (49.0%) had elevated multimorbidity (≥ 5 chronic conditions), while 48% were prescribed five

or more medications, indicating a high prevalence of polypharmacy. A notable 24.8% of residents experienced at least one ADR. The number of regularly scheduled medications was a major risk factor, and certain drug classes—including cardiovascular agents, gastrointestinal drugs, anticoagulants, and psychotropics—were frequently implicated in ADRs. Additionally, a greater number of chronic conditions was associated with increased ADR risk. While sex-based differences were not statistically significant, female residents showed slightly higher rates of ADRs, aligning with previously reported trends. Our findings are consistent with a broad body of literature that underscores the multifactorial nature of ADRs in elderly populations.

The ADR prevalence of 24.8% in this study aligns with prior estimates of 10–30% among elderly nursing home residents^{17,18}. This high prevalence reflects the vulnerability of this population, exacerbated by age-related changes in drug metabolism, polypharmacy, and chronic disease burden. Similar figures were observed in an Irish cohort study where 78% of older adults reported at least one adverse drug event over six months¹⁹.

Consistent with prior studies²⁰⁻²², polypharmacy was a strong and consistent risk factor for ADRs. The threshold of five or more medications significantly increased ADR risk. One study²² found that 44.8% of elderly individuals used nine or more medications, while 35.4% used between five and eight—comparable to the distribution in our cohort. Previous research²¹ also showed that polypharmacy contributes to preventable events such as falls, confusion, and hospitalizations.

Polypharmacy prevalence has risen over recent decades. Between 1995 and 2005, the proportion of individuals taking ≥ 5 medications more than doubled^{5,21}. This trend is further complicated by the concurrent use of non-prescription medications and supplements—nearly half of older adults take OTC drugs and over half use dietary supplements²—heightening the risk of drug-drug and drug-condition interactions.

Our results confirm previous reports^{5,24,25} that specific drug classes are disproportionately associated with ADRs. Cardiovascular drugs, gastrointestinal agents, and anticoagulants were particularly implicated. Cardiovascular and GI drugs are linked to ADRs in older adults because of pharmacodynamic changes that heighten sensitivity and impair compensatory mechanisms²⁶. These age-related factors, combined with polypharmacy and comorbidities, increase the potential for drug-related harm even at standard doses²⁷. Proton pump inhibitors (PPIs), benzodiazepines, antidepressants, and diuretics were also frequently involved—aligning with national pharmacovigilance data, which showed cardiovascular and antithrombotic drugs accounted for 10% of ADR reports, with nearly three-quarters classified as serious or fatal²⁵.

A significant correlation was observed between the number of chronic conditions and ADR risk, reflecting previous findings^{5,21,22,28}. One multivariate analysis²⁸ found that patients with 4–5 or 6–10 diagnoses had odds ratios for ADRs as high as 28.5 and 12.7, respectively. Our cohort's average of four chronic



illnesses aligns with this data, reinforcing the compounded pharmacological risk posed by comorbidity.

Although our study did not find statistically significant sex differences in ADR rates, the slightly higher prevalence among females aligns with Alhawassi et al.²⁹, who found female sex, higher comorbidity burden, and polypharmacy to be significant predictors of ADRs. Possible explanations include sex differences in metabolism, hormonal profiles, and body composition.

These findings carry important implications for clinical care and health policy in long-term care facilities. Regular, structured medication reviews and deprescribing should be prioritized for residents on ≥ 5 medications or high-risk drug classes, as polypharmacy significantly increases ADR risk in older adults³⁰. Caution is especially needed when prescribing cardiovascular drugs, psychotropics, anticoagulants, and PPIs in patients with multimorbidity, given their link to ADR-related hospitalizations³¹. Involving pharmacists and geriatricians can enhance ADR detection and promote safer, individualized prescribing, particularly in resource-limited settings³². Ongoing education for healthcare providers is essential to improve ADR recognition and management³³. These findings support the need for systemic reforms, including prescribing audits, electronic alert systems, and incentives for safe prescribing in aged care³⁴. Although sex differences in ADRs were not conclusive, evidence suggests older women may be at higher risk, warranting further investigation³⁵. Personalized prescribing that accounts for sex, comorbidities, and medication burden may enhance safety and outcomes in older adults³⁶.

Strengths and limitations

Despite these insights, this study has some limitations. The main weaknesses are the relatively small sample size and the fact that data were collected from only two nursing homes, which may affect the generalizability of the results. Additionally, as a retrospective study, the information was limited to what was available in medical charts, and in some cases, it was not possible to obtain further details about the patients' conditions or their pharmacological history prior to admission to the nursing home. Furthermore, ADR detection may be influenced by underreporting, particularly for milder symptoms or those mistakenly attributed to disease progression.

CONCLUSION

This study highlights the high prevalence of ADRs among elderly patients, with a positive association to polypharmacy, comorbidities, and specific drug classes use. Understanding these factors is crucial for enhancing patient safety and optimizing pharmacotherapy. To reduce ADRs, strategies such as minimizing inappropriate prescriptions, improving monitoring, and preventing ADRs are essential. Effective strategies should involve individualized prescribing, deprescribing when necessary, enhanced pharmacovigilance, and clinician education. Future research should explore predictive models incorporating electronic health records and comprehensive approach to proactively identify high-risk patients.

CONFLICTS OF INTEREST

All the authors declare no conflicts of interest.

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