Original Research

Potentially inappropriate medications prescribing according to Beers criteria among elderly outpatients in Jordan: a cross sectional study

Ahmad AL-AZAYZIH, Rawan ALAMOORI, Shoroq M. ALTAWALBEH

Received (first version): 28 Dec 2018  Accepted: 26 May 2019 Published online: 5 Jun 2019

Abstract

Background: Due to aging, along with its associated physiological changes, older adults are extremely vulnerable to be afflicted with multiple chronic conditions (multimorbidity). Accordingly, prescribing a large number of drugs to older adults would be inevitable. Resulted complex drug regimens can lead to prescribing of Potentially Inappropriate Medications (PIMs) with subsequent negative health and economic outcomes.

Objectives: The main objective of this study is to investigate the prevalence and predictors of PIMs prescribing among Jordanian elderly outpatients, using the last updated version of the American Geriatrics Society (AGS) Beers Criteria (2015 version).

Methods: A Unicenter, cross-sectional study were data was assessed using medical records of included study subjects conducted over three months period from beginning of October to the end of December 2016 at King Abdullah University Hospital, Al Ramtha, Jordan. Our study included patients aged 65 years or above who visited the outpatient clinics at King Abdullah University hospital (KAUH) and were prescribed at least one oral medication during the study period. PIMs were identified for these patients and further classified according to the 2015 AGS Beers Criteria. We measured the prevalence of PIMs prescribed among elderly outpatients in Jordan.

Results: A total of 4622 eligible older adults were evaluated in this study, of whom 62.5% (n=2891) were found to have at least one PIM prescribed during the three months study period. 69% of identified PIMs were medications to be used with caution in elderly, 22% were medications to avoid in many or most older adults, 6.3% were medications to be avoided or have their dosage adjusted based on kidney function in older adults, 2.04% medications were to avoid in older adults with specific diseases/syndromes, and 1.6% were potentially clinically important non-anti-infective drug-drug interactions to be avoided in older adults. Female gender and polypharmacy were found to be significant predictors of PIMs use among elderly.

Conclusions: Potentially Inappropriate Medication prescribing is common among Jordanian elderly outpatients. Female gender and polypharmacy are associated with more PIMs prescribing and so need further attention.

Keywords

Potentially Inappropriate Medication List; Inappropriate Prescribing; Polypharmacy; Aged; Outpatients; Risk Factors; Pharmacists; Cross-Sectional Studies; Jordan

INTRODUCTION

Potentially inappropriate medications (PIMs) among elderly is defined as “medications or medication classes which generally should be avoided in patients aged 65 years or older because of being either ineffective or pose potential high risk for such age group while safer alternatives are available”1. PIMs use, where adverse effects exceed its health benefits, is a highly witnessed issue among older adults ranging from a prevalence of 11.5 - 85.1% among community dwelling and hospitalized elderly patients in various countries.2-4 Data from numerous studies performed at various settings revealed PIM prescribing with prevalence of up to 40% in nursing home residents and up to 28% among community-based older patients.5-23 Healthcare utilization, including: hospitalizations, healthcare visits (Inpatient, Outpatient and Emergency Department), re-admissions and length of stays, was shown to be significantly associated with PIM use among elderly patients.24

Detecting adverse drug reactions among geriatric patients is challenging, as they often exhibit non-specific symptoms such as constipation, lethargy, lightheadedness, confusion, falls and depression.25 Nonetheless, adverse drug reaction rates among elderly patients are at least three-times that of the younger as well as general population.25 Several studies performed in order to investigate the association between PIMs use and developing unwanted Adverse Drug Reactions (ADRs) and Adverse Drug Events (ADEs) among elderly patients at different settings (hospitalized, outpatient and even nursing home residents) concluded that using PIMs is significantly associated with an increased risk of developing ADRs and ADEs.26-28

Screening tools for PIMs use have been developed over the past two decades, with the latest updated version of the Beers Criteria was released by the American Geriatrics Society (AGS) in October, 2015.29 In addition to updating existing criteria (i.e., 2012), the 2015 version has come up with two new major aspects: 1) Drugs for which dosage adjustment is required based on the patient’s kidney function, and 2) Drug-Drug Interactions (DDIs).29 The goal of the 2015 update of AGS Beers Criteria continues to be improving geriatric care by reducing their exposure to PIMs. Prudent application of the criteria as a non-punitive
Educational tool and quality improvement measure is intended to achieve a closer monitoring of medication usage among older adults. Overall, PIM use among geriatric patients has become a global concern, and studies were conducted in many countries using globally developed tools for screening this phenomenon. However, no study has yet systematically examined PIMs prescribing among Jordanian elderly patients. Accordingly, this study is aiming to investigate the prevalence and predictors of PIM prescribing among Jordanian elderly outpatients, using the last updated version of the American Geriatrics Society (AGS) 2015 Beers Criteria. The main objective of this study is to investigate the prevalence and predictors of PIMs prescribing among Jordanian elderly outpatients, using the last updated version of the American Geriatrics Society (AGS) Beers Criteria (2015 version).

METHODS

Ethics Approval

This study was approved by the Institutional Review Board (IRB) of King Abdullah University Hospital (KAUH). (Approval Number: 16/100/2016)

Study design and population

This study was a cross-sectional study where data was assessed using medical records of included study subjects. In this study, patients aged 65 years and older who visited the King Abdullah University Hospital (KAUH) located at Al-Ramtha, Jordan, and were ordered at least one outpatient prescription containing at least one prescribed medication during the period from October 1, 2016 to December 31, 2016 were identified as the study population. Electronic outpatient medical records were reviewed and data including demographics and clinical characteristics including age, gender, comorbidities related to AGS Beers Criteria, medications prescribed, and the latest measured serum Creatinine level were extracted. Latest measured serum creatinine level for each patient was used to calculate creatinine clearance using Jelliffe equation, which could be used and reasonably accurate when height and weight are not available in average-sized patients. Polypharmacy was investigated and defined as simultaneous use of five or more different prescribed medications.

Primary outcomes

Potentially Inappropriate Medications (PIMs) among participants were identified and further classified according to the 2015 American Geriatrics Society (AGS) Beers Criteria in the following classes:

1) Medications to avoid in many or most geriatrics (Class I). Prescribing any of the listed medications among the study population was considered as a PIM.

2) Medications to avoid in older adults with certain diseases/syndromes (Class II). Drug-disease interactions were evaluated for each patient who was ordered any of the listed medications. All related conditions/diseases have been investigated and confirmed for those patients using their medical records. Subsequently, patients exposed to PIM were

Table 1. Potentially inappropriate medications to avoid in many or most older adults (22% of total PIMs)

| Class/Medication                  | % of total | QOE | SOR  |
|-----------------------------------|------------|-----|------|
| Pain medications                  |            |     |      |
| Orphenadrine                      | 31.2       | Moderate | Strong |
| Non-COX-Selective NSAIDs          |            |     |      |
| Gastrointestinal                  |            |     |      |
| Metoclopramide                    | 25.3       | High | Strong |
| Antispasmodics                    |            |     |      |
| Scopolamine (Hyoscine butylbromide)| 13.5     | Moderate | Strong |
| Endocrine                         |            |     |      |
| Glyburide (Glibenclamide)         | 10.8       | High | Strong |
| Estrogens with or without progester| 9.8       | Moderate | Strong |
| Central Nervous System (CNS)      |            |     |      |
| Benzodiazepines                   |            |     |      |
| Antidepressants                   |            |     |      |
| Antipsychotics                    |            |     |      |
| Cardiovascular                    |            |     |      |
| Digoxin                           | 5.4        | Moderate | Strong |
| Central alpha blockers (Methylidopa)|        |     |      |
| Amiodarone                        |            |     |      |
| Nifedipine, IR                    |            |     |      |
| Peripheral alpha blockers (Doxazosin)|    |     |      |
| Anticholinergics (1st generation antihistamines)| 2.6 | Moderate | Strong |
| Chlorpheniramine                   |            |     |      |
| Metcizine                          |            |     |      |
| Antithrombotics                   |            |     |      |
| Dipyridamole, SA                  | 1.2        | Moderate | Strong |
| Genitourinary                     |            |     |      |
| Desmopressin                      | 0.15       | Moderate | Strong |

Non-COX-Selective NSAIDs: non-cyclooxygenase selective non-steroidal anti-inflammatory drugs, PPIs: Proton Pump Inhibitors, QOE: Quality of Evidence, SOR: Strength of Recommendation.
then highlighted based on related recommendation. (e.g., patients ordered the oral decongestant pseudoephedrine were evaluated for the presence of insomnia; if such drug-disease interaction was confirmed then the patient was considered to have been exposed to a PIM).

3) Medications that should be used with caution among elderly patients (Class III). Prescribing any of the listed medications among the study population was considered as a PIM.

4) Potentially clinically important non-anti-infective drug-drug interactions (Class IV). Interacted medications were identified and highlighted for each subject.

5) Medications to avoid or their dosage should be adjusted based on patient kidney function (Class V). Using at least one of the listed medications along with patient creatinine clearance below indicated thresholds was considered as a PIM (e.g. patient with creatinine clearance <30ml/min and was concurrently prescribed enoxaparin was considered to have a PIM).

6) Medications belong to more than one class of the above were counted as more than one PIMs (for example, if a drug belong to class I and II, it will be considered as two PIMs)

Table 2. Medications to avoid for older adults with specific diseases or syndromes (2.04% of total PIMs)

| Object System/Disease | Interacting class/Medication | % of total | QOE | SOR |
|-----------------------|-----------------------------|------------|-----|-----|
| Central Nervous System (CNS) | | | | |
| History of falls or fractures | | | | |
| Anticonvulsants | – | 81.3 | Moderate | Strong |
| Antipsychotics | – | Moderate | Strong |
| Benzodiazepines | – | Moderate | Strong |
| TCAs | – | Moderate | Strong |
| SSRIs | – | Moderate | Strong |
| Dementia or cognitive impairment | | | | |
| Anticholinergics | – | Moderate | Strong |
| Benzodiazepines | – | Moderate | Strong |
| Antipsychotics | – | Moderate | Strong |
| Chronic seizures or epilepsy | | | | |
| Insomnia | | | | |
| Tramadol | | – | Low | Strong |
| Theophylline | | – | Moderate | Strong |
| Kidney and Urinary tract | | | | |
| Lower Urinary Tract Symptoms, BPH | | | | |
| Anticholinergics (Strong) | | – | Moderate | Strong |
| Cardiovascular | | | | |
| Heart failure | | | | |
| NSAIDs | | – | Moderate | Strong |
| TCA | – | Moderate | Strong |
| Anticholinergics | – | Moderate | Strong |
| Non-steroidal Anti-inflammatory Drugs, QOE: Quality of Evidence, SOR: Strength of Recommendation

The quality of evidence (QOE) and strength of recommendation (SOR) for each encountered Beers PIM or criterion were also reported according to The Grades of Recommendation, Assessment, Development and Evaluation (GRADE: “a common, sensible and transparent approach to grading quality or certainty of an evidence as well as strength of recommendation”).

Statistical analysis

Descriptive statistics, such as percentages and arithmetic means, were used to describe patient characteristics and to estimate the prevalence of PIM use among studied population. Multivariable logistic regression model was conducted to investigate potential risk factors associated with PIM use. P value of less than 0.05 was considered statistically significant. All analyses were conducted using Stata v14 (StataCorp. 2015. Stata Statistical Software: Release 14. College Station, TX: StataCorp LP).

RESULTS

Eligible 4,622 geriatric patients were included in the study. Mean age (standard deviation) of studied population was 73.2 (SD 6.3) years, and 50.6 % (n=2340) of subjects were females. After excluding over the counter products, 4,356

Table 3. Medications to be used with caution in older adults (69% of total PIMs)

| Class/Medication | % of total | QOE | SOR |
|------------------|------------|-----|-----|
| Diuretics | 45.1 | Moderate | Strong |
| Aspirin for primary prevention of cardiac events | 38.8 | Moderate | Moderate |
| Vasodilators | 12.3 | Moderate | Weak |
| Antidepressants | | | |
| SSRIs* | 1.7 | Moderate | Strong |
| TCAs* | – | Moderate | Strong |
| SNRIs* | – | Moderate | Strong |
| Antipsychotics | | | |
| | 1.1 | Moderate | Strong |
| Carbamazepine | 0.63 | Moderate | Strong |
| Chemotherapy | 0.19 | Moderate | Strong |

SSRIs: Selective Serotonin Reuptake Inhibitors, TCAs: Tri-cyclic Antidepressants, SNRIs: Serotonin-Norepinephrine Reuptake Inhibitors, QOE: Quality of Evidence, SOR: Strength of Recommendation
patients were found to be prescribed at least one medication, with average of 5.6 (SD 3.9) prescribed medications per patient. Overall, 53.7% of women and 51.7% of men aged 65 years and older were ordered at least five prescription medications (i.e. exposed to “polypharmacy”).

Of the total geriatric sample evaluated, 2891 subjects (62.5%) were prescribed at least one PIM according to 2015 Beers Criteria. The majority of these subjects were prescribed one (39.6%) or two (23.3%) PIMs, however, prescribing of three (15.8%) or four (7.6%) PIMs were also not uncommon. Of the identified PIMs, 69% were medications to be used with caution in older adults (class III), whereas 22% were medications to avoid in many or most older adults (class II), 6.3% were medications to avoid or their dosage should be adjusted based on the kidney function (class V), 2.1% were medications to avoid for older adults with specific diseases/syndromes (class II), and 1.6% were potentially clinically important non-anti-infective drug-drug interactions (class IV).

Most commonly prescribed medication classes considered as PIMs to avoid in majority of elderly patients were pain medications (31.2%), followed by gastrointestinal medications (25.3%) and antispasmodics (13.5%), Table 1.

Potential drug-disease/syndrome interactions were 2.04% of total identified PIMs. Subjects diagnosed with Central Nervous System (CNS)-related conditions (including history of falls or fractures, dementia, chronic seizures or epilepsy and insomnia) were most commonly prescribed PIMs (81.3%) in this class, Table 2.

The majority (69%) of total PIMs prescribed were medication classes to be used with caution among elderly. Diuretics (45.1%), aspirin for primary prevention of cardiac events among older adults aged ≥80 years (38.8%) and vasodilators (12.3%) were most commonly prescribed PIMs in this class, Table 3.

Only (1.6%) of total prescribed PIMs were due to drug-drug interactions (the 1st newly added component by the 2015 AGS Beers Criteria). Almost (91.5%) of detected interactions resulted from the simultaneous prescribing of at least two different medications with anticholinergic properties including first generation antihistamines (cholorpheniramine and triprolidine), antimuscarinic (oxybutynin), antispasmodics (clidinium-chlordiazepoxide and hyoscyamine), Table 4.

Drugs to be avoided or have their dosage adjusted based on varying levels of kidney function in older adults (the 2nd newly added component by the 2015 AGS Beers Criteria) have contributed to almost 6.3% of total PIMs encountered. Prescribing gabapentin when CrCl <60 ml/min was responsible for the majority of PIMs in this class, Table 5.

Having at least one identified PIM was significantly associated with female gender (adjusted OR: 1.33, 95% CI: 1.14-1.55) and polypharmacy (adjusted OR: 28.39, 95%CI: 23.83-33.81), Table 6.

**DISCUSSION**

To the best of our knowledge, this is the first study examining PIM prevalence using Beers Criteria as a screening tool among geriatric patients in Jordan. Moreover, it considered the first study investigating and presenting the 2015 AGS Beers Criteria PIM classes in details among elderly outpatients.

Results from this study showed a high prevalence of PIM prescribing among Jordanian elderly outpatients (62.5%) according to the most updated version of the AGS Beers Criteria (2015). Female gender and polypharmacy were identified as significant predictors of PIM prescribing in this population. Previous iterations of Beers Criteria were used by a wide variety of studies examining the prevalence of PIM prescribing among older adults at various populations and settings with estimated ranges between 11.5% and 85.1%.4,5,10,15,20 A good number of these studies have assessed prevalence of PIM prescribing based on Beers Criteria (mostly 2012 update) among community-dwelling elderly patients (outpatients) with total prevalence ranges between (23% and 81%).4,9,11,13,18 The updated AGS Beers Criteria were released in 2015 and so still less thoroughly used in the available literature. In the United States, a study was conducted using Medicare Current Beneficiary Survey (MCBS) data set to identify PIMs according to 2015 AGS Beers Criteria.26 Findings from this study showed that around 57% of older adults with dental care visits were

**Table 4. Potential clinically important drug-drug interactions in older adults (1.6% of total PIMs)**

| Object Drug/Class | Interacting Drug/Class | % of total | QOE | SOR |
|------------------|------------------------|-----------|-----|-----|
| Anticholinergic   | Anticholinergic         | 91.5      | Moderate | Strong |
| CNS-active drug* | ≥ 2 other CNS-active drugs | 4.3      | High | Strong |
| ACEs             | Amiloride              | 2.1       | Moderate | Strong |
| Peripheral Alpha-1 blockers | Loop diuretics | 1.1      | Moderate | Strong |
| Corticosteroids, oral or parenteral | NSAIDs | 1.1      | Moderate | Strong |
| CNS-active drugs: SSRI, TCAs, Antipsychotics, Benzodiazepines, Opioids, ACEIs: Angiotensin-Converting Enzyme Inhibitors, NSAIDs: non-steroidal anti-inflammatory drugs |

**Table 5. Drugs to be avoided or have their dosage adjusted with varying levels of kidney function in older adults (6.3% of total PIMs)**

| Medication | CrCl (ml/min) | Action Required | QOE | SOR | % of total |
|------------|---------------|----------------|-----|-----|-----------|
| Gabapentin | <60           | Reduce dose    | Moderate | Strong | 71.7 |
| Levetiracetam | ≤80         | Reduce dose    | Moderate | Strong | 13.9 |
| Ranitidine | <50           | Reduce dose    | Moderate | Strong | 5 |
| Pregabalin | <60           | Reduce dose    | Moderate | Strong | 5 |
| Famotidine | <50           | Reduce dose    | Moderate | Strong | 4.5 |

CrCl: Creatinine Clearance (ml/min), QOE: Quality of Evidence, SOR: Strength of Recommendation.
prescribed at least one Beers Criteria medication. Another study conducted in Brazil found that 50% of community-dwelling elderly patients were prescribed at least one PIM according to the 2015 update of AGS Beers Criteria. Accordingly, PIM prevalence in the current study (62.5%) is relatively comparable with available literature which investigated geriatric PIMs using the updated Beers Criteria in the outpatient settings. Nevertheless, it is an apparently high prevalence in Jordan. The vast majority of PIMs identified in the current study were drugs to be used with caution (69%). This finding is consistent with recent study in Qatar where 65% of PIMs were drugs to be used with caution.

Female gender was significantly associated with PIM use. This finding is consistent with various published studies. Females generally are at higher risk for developing multiple chronic conditions than males, consequently being more susceptible for drug-disease and drug-drug interactions which will lead to more inappropriate prescribing. Polypharmacy was the most significant predictor of PIM prescribing in the current study. Increasing age was reported by some studies as a significant risk factor of PIM use. However, others have found no association between age and PIM use. In the current study, age was not a significant predictor of PIM use after adjusting for polypharmacy and gender. The effect of age on PIM prescribing in older adults may be mediated by the multi-comorbidity and the resultant polypharmacy in this population.

This study has some limitations. As this was a retrospective study, all data were obtained from patient medical records and were as accurate as available documentation. Due to the large study sample, some patients’ characteristics (for example, patients’ weight and height) and clinical indicators were not extracted and so their association with PIM prescribing was not evaluated in this study. Similarly, complete comorbidity profiles were not obtained for all patients, however, all conditions related to PIMs were carefully evaluated for the relevant patients. In addition, because of the retrospective nature of the study, we were not able to capture the dose reduction if happened; however, we considered all those on regular maintenance dose with varying kidney functions to have potential inappropriate use. This was challenging since Beers criteria did not provide a cut off dose for these meds to be considered as PIMs. Finally, all identified PIMs are not conclusive without the clinical judgment, the criteria serve only as a warning light to raise attention to clinical evaluation that further assess risks and benefits associated with these PIMs. Overall, this study has the strength of evaluating a large number of elderly patients, enabling generalizability and reliability of findings. In addition, this is the first study-examining trend of prescribing among elderly patients in Jordan using the most updated version of Beers Criteria (2015) for PIMs identification.

CONCLUSIONS
This study demonstrates that PIM prescribing is common among Jordanian elderly outpatients. Female patients and those with polypharmacy are the most vulnerable groups that need further attention. Strategies to monitor, review, and prevent inappropriate prescribing should be supported.

CONFLICT OF INTEREST
The authors declare no conflict of interest does exist.

FUNDING
This work was funded by the Jordan University of Science and Technology – Deanship of Research. The results and contents of this work are solely the responsibility of the authors. The funders had no role in study design, data collection and analysis, and decision to publish.

Table 6. Predictors of potentially inappropriate medications prescribing among elderly patients.

| Variable          | OR (95% CI)  | P-value |
|-------------------|-------------|---------|
| Age               | 0.99 (0.98-1.0) | 0.14    |
| Gender (Female)   | 1.33(1.14-1.55) | *<0.001 |
| Polypharmacy #    | 28.39 (28.83-33.81) | *<0.001 |

*p-value is significant at <0.05
#: Polypharmacy was defined as ≥ 5 prescribed medications
CI, confidence interval; OR, odds ratio; PIM, potentially inappropriate medication

References
1. Beers MH, Ouslander JG, Rollingher I, Reuben DB, Brooks J, Beck JC. Explicit criteria for determining inappropriate medication use in nursing home residents. UCLA Division of Geriatric Medicine. Arch Intern Med. 1991;151(9):1825-1832. https://doi.org/10.1001/archie.1991.0400090107019
2. Guaraldo L, Cano FG, Damasco GN, Rozenfeld S. Inappropriate medication use among the elderly: a systematic review of administrative databases. BMC Geriatr. 2011;11:79. https://doi.org/10.1186/1471-2318-11-79
3. da Costa FA, Periquito C, Carneiro MC, Oliveira P, Fernandes AI, Cavaco-Silva P. Potentially inappropriate medications in a sample of Portuguese nursing home residents: Does the choice of screening tools matter? Int J Clin Pharm. 2016;38(5):1103-1111. https://doi.org/10.1007/s11096-016-0337-y
4. Zeenny R, Wakim S, Kuyumjian YM. Potentially inappropriate medications use in community-based aged patients: a cross-sectional study using 2012 Beers criteria. Clin Interv Aging. 2017;12:65-73. https://doi.org/10.2147/CIA.S87564
5. Li H, Pu S, Liu Q, Huang X, Kuang J, Chen L, Shen J, Cheng S, Wu T, Li R, Li Y, Mo L, Jiang W, Song Y, He J. Potentially inappropriate medications in Chinese older adults: The beers criteria compared with the screening tool of older persons’ prescriptions criteria. Geriatr Gerontol Int. 2017;17(11):1951-1958. https://doi.org/10.1111/ggi.12999
6. Hudhra K, Begi E, Petrela E, Xhafa D, Garcia-Caballos M, Bueno-Cavanillas A. Prevalence and factors associated with potentially inappropriate prescriptions among older patients at hospital discharge. J Eval Clin Pract. 2016;22(5):707-713. https://doi.org/10.1111/jep.12521

7. Nam YS, Han JS, Kim JY, Bae WK, Lee K. Prescription of potentially inappropriate medication in Korean older adults based on 2012 Beers Criteria: a cross-sectional population based study. BMC Geriatr. 2016;16:118. https://doi.org/10.1186/s12877-016-0285-3

8. Lim YJ, Kim HY, Choi J, Lee JS, Ahn AL, Oh EJ, Cho DY, Kweon HJ. Potentially inappropriate medications by beers criteria in older outpatients: prevalence and risk factors. Korean J Fam Med. 2016;37(6):329-333. https://doi.org/10.4092/kjfim.2016.37.6.329

9. Osei EK, Berry-Cabán CS, Haley CL, Rhodes-Pope H. Prevalence of Beers criteria medications among elderly patients in a military hospital. Gerontol Geriatr Med. 2016;2:e2333721416637790. https://doi.org/10.1177/2333721416637790

10. Hudhra K, García-Caballos M, Casado-Fernandez E, Jucja B, Shabani D, Bueno-Cavanillas A. Polypharmacy and potentially inappropriate prescriptions identified by Beers and STOPP criteria in co-morbid older patients at hospital discharge. J Eval Clin Pract. 2016;22(2):189-193. https://doi.org/10.1111/jep.12452

11. Davidoff AJ, Miller GE, Sarpong EM, Yang E, Brandt N, Fick DM. Prevalence of potentially inappropriate medication use in older adults using the 2012 Beers Criteria. J Am Geriatr Soc. 2015;63(3):486-500. https://doi.org/10.1111/jgs.13320

12. Almohm E, Khalifa S, Bahi AA. Prevalence and predictors of potentially inappropriate medications among home care elderly patients in Qatar. Int J Clin Pharm. 2015;37(5):815-821. https://doi.org/10.1007/s11096-015-0125-0

13. Baidoni Ade O, Ayres LR, Martinez EZ, Dewulf Nde L, Dos Santos V, Pereira LR. Factors associated with potentially inappropriate medications use by the elderly according to Beers criteria 2003 and 2012. Int J Clin Pharm. 2014;36(2):316-324. https://doi.org/10.1007/s11096-013-9880-y

14. Ubeda A, Ferrández L, Maicas N, Gomez C, Bonet M, Peris JE. Potentially inappropriate prescribing in institutionalised older patients in Spain: the STOPP-START criteria compared with the Beers criteria. Pharm Pract (Granada). 2012;10(2):83-91.

15. Aparasu RR, Mort JR. Inappropriate prescribing for the elderly: beers criteria-based review. Ann Pharmacother. 2000;34(3):338-346. https://doi.org/10.1345/aph.19006

16. Rothberg MB, Pekow PS, Liu F, Korc-Grodzicki B, Brennan MJ, Bellantoni S, Heelon M, Lindenzauer PK. Potentially inappropriate medication use in hospitalized elders. J Hosp Med. 2008;3(2):91-102. https://doi.org/10.1002/jhm.290

17. Tragni E, Casula M, Pieri V, Favato G, Marcobelli A, Trotta MG, Catapano AL. Prevalence of the prescription of potentially interacting drugs. PLoS One. 2013;8(10):e78827. https://doi.org/10.1371/journal.pone.0078827

18. Barry PJ, Gallagher P, Ryan C. Inappropriate prescribing in geriatric patients. Curr Psychiatry Rep. 2008;10(1):37-43.

19. Cahir C, Fahey T, Teeling M, Teljeur C, Feely J, Bennett K. Potentially inappropriate prescribing and cost outcomes for older people: a national population study. Br J Clin Pharmacol. 2010;69(5):543-552. https://doi.org/10.1111/j.1365-2125.2010.03628.x

20. Harlon JT, Schmader KE, Boult C, Arzt MB, Gross CR, Fillenbaum GG, Ruby CM, Garrard J. Use of inappropriate prescription drugs by older people. J Am Geriatr Soc. 2002;50(1):26-34. https://doi.org/10.1046/j.1532-5415.2002.50004.x

21. Giron MS, Wang HX, Bernsten C, Thorslund M, Winblad B, Fastbom J. The appropriateness of drug use in an older nondemented and demented population. J Am Geriatr Soc. 2001;49(3):277-283. https://doi.org/10.1046/j.1532-5415.2001.4930277.x

22. Blozik E, Rapold R, von Overbeck J, Reich O. Polypharmacy and potentially inappropriate medication in the adult, community-dwelling population in Switzerland. Drugs Aging. 2013;30(7):561-568. https://doi.org/10.1007/s40266-013-0073-0

23. Galvin R, Moriarty F, Cousins G, Cahir C, Motterlini N, Bradley M, Hughes CM, Bennett K, Smith SM, Fahey T, Kenny RA. Prevalence of potentially inappropriate prescribing and prescribing omissions in older Irish adults: findings from The Irish LongitudinalDinal Study on Ageing study (TILDA). Eur J Clin Pharmacol. 2014;70(5):599-606. https://doi.org/10.1007/s00228-014-1851-3

24. Hyttinen V, Jyrkkä J, Valtonen H. A systematic review of the impact of potentially inappropriate medication on health care utilization and costs among older adults. Med Care. 2016;54(10):950-964. https://doi.org/10.1097/MLR.0000000000000587

25. Gallagher P, Barry P, O'Mahony D. Inappropriate prescribing in the elderly. J Clin Pharm Ther. 2007;32(2):113-121. https://doi.org/10.1080/136527107007093.x

26. Chang CM, Liu PY, Yang YH, Yang YC, Wu CF, Lu FH. Use of the Beers criteria to predict adverse drug reactions among first-visit elderly outpatients. Pharmacotherapy. 2005;25(6):831-838.

27. Onder G, Landi F, Lipietz R, Fialova D, Gambassi G, Bernabei R. Impact of inappropriate drug use among hospitalized older adults. Eur J Clin Pharmacol. 2005;61(5-6):453-459. https://doi.org/10.1007/s00228-005-0928-3

28. Perri M 3rd, Menon AM, Deshpande AD, Shinde SB, Jiang R, Cooper JW, Cook CL, Griffin SC, Lorys RA. Adverse outcomes associated with inappropriate drug use in nursing homes. Ann Pharmacother. 2005;39(3):405-411. https://doi.org/10.1345/aph.1E230

29. By the American Geriatrics Society Beers Criteria Update Expert Panel. (2015). American Geriatrics Society 2015 Updated Beers Criteria for Potentially Inappropriate Medication Use in Older Adults. J Am Geriatr Soc. 2015;63(11):2227-2246. https://doi.org/10.1111/jgs.13702

30. Huikins D, Macleod U, Boland JW. Identifying potentially inappropriate prescribing in older people with dementia: a systematic review. Eur J Clin Pharmacol. 2019;75(4):467-481. https://doi.org/10.1007/s00228-018-02612-x

31. Motter FR, Fritzen JS, Hilmer SN, Paniz EV, Paniz VM. Potentially inappropriate medication in the elderly: a systematic review of validated explicit criteria. Eur J Clin Pharmacol. 2018;74(6):679-700. https://doi.org/10.1007/s00228-018-2446_0
32. Storms H, Marquet K, Aertgeerts B, Claes N. Prevalence of inappropriate medication use in residential long-term care facilities for the elderly: A systematic review. Eur J Gen Pract. 2017;23(1):69-77. https://doi.org/10.1080/13814788.2017.1288211

33. O'Connell MB, Dwinell AM, Bannick-Mohrand SD. Predictive performance of equations to estimate creatinine clearance in hospitalized elderly patients. Ann Pharmacother. 1992;26(5):627-635. https://doi.org/10.1177/10600280922600503

34. Prithviraj GK, Koroukian S, Margevicius S, Berger NA, Bagai R, Owusu C. Patient characteristics associated with polypharmacy and inappropriate prescribing of medications among older adults with cancer. J Geriatr Oncol. 2012;3(3):228-237. https://doi.org/10.1016/j.jgo.2012.02.005

35. Guyatt GH, Oxman AD, Vist GE, Kunz R, Falck-Ytter Y, Alonso-Coello P; GRADE Working Group. GRADE: an emerging consensus on rating quality of evidence and strength of recommendations. BMJ. 2008;336(7650):924-926. https://doi.org/10.1136/bmj.39489.470347.AD

36. Fick DM, Cooper JW, Wade WE, Waller JL, Maclean JR, Beers MH. Updating the Beers criteria for potentially inappropriate medication use in older adults: results of a US consensus panel of experts. Arch Intern Med. 2003;163(22):2716-2724. https://doi.org/10.1001/archinte.163.22.2716

37. American Geriatrics Society 2012 Beers Criteria Update Expert Panel. American Geriatrics Society updated Beers Criteria for potentially inappropriate medication use in older adults. J Am Geriatr Soc. 2012;60(4):616-631. https://doi.org/10.1111/j.1532-5415.2012.03923.x

38. Buck MD, Atreja A, Brunker CP, Jain A, Suh TT, Palmer RM, Dorr DA, Harris CM, Wilcox AB. Potentially inappropriate medication prescribing in outpatient practices: prevalence and patient characteristics based on electronic health records. Am J Geriatr Pharmacother. 2009;7(2):84-92. https://doi.org/10.1016/j.amjpharm.2009.03.001

39. Skaar DD, O'Connor H. Using the Beers criteria to identify potentially inappropriate medication use by older adult dental patients. J Am Dent Assoc. 2017;148(5):298-307. https://doi.org/10.1016/j.adaj.2017.01.020

40. Novaes PH, da Cruz DT, Lucchetti ALG, Leite ICG, Lucchetti G. Comparison of four criteria for potentially inappropriate medications in Brazilian community-dwelling older adults. Geriatr Gerontol Int. 2017;17(10):1628-1635. https://doi.org/10.1111/ggi.12944

41. Fabbri E, Zoli M, Gonzalez-Freire M, Salive ME, Studenski SA, Ferrucci L. Aging and multimorbidity: new tasks, priorities, and frontiers for integrated gerontological and clinical research. J Am Med Dir Assoc. 2015 Aug 1;16(8):640-647. https://doi.org/10.1016/j.jamda.2015.03.013

42. Charlesworth CJ, Smit E, Lee DS, Aliramadhan F, Odden MC. Polypharmacy Among Adults Aged 65 Years and Older in the United States: 1988-2010. J Gerontol A Biol Sci Med Sci. 2015;70(8):989-995. https://doi.org/10.1093/gerona/glv013

43. Santos AP, da Silva DT, dos Santos Junior GA, Silvestre CC, Nunes MA, Lyra DP Jr, Antonioli AR. Evaluation of the heterogeneity of studies estimating the association between risk factors and the use of potentially inappropriate drug therapy for the elderly: a systematic review with meta-analysis. Eur J Clin Pharmacol. 2015;71(9):1037-1050. https://doi.org/10.1007/s00228-015-1891-2

44. Kondo N, Nakamura F, Yamazaki S, Yamamoto Y, Akizawa T, Akiba T, Saito A, Kurokawa K, Fukushima S. Prescription of potentially inappropriate medications to elderly hemodialysis patients: prevalence and predictors. Nephrol Dial Transplant. 2015;30(3):498-505. https://doi.org/10.1093/ndt/gfu070