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3 **Prevalence and Predictors of Potentially Inappropriate Medications (PIMs) in**  
4 **Saudi Older Adults with Mood and Anxiety Disorders: A Systematic Review and**  
5 **Meta-analysis .**

6

7 **Abstract**

8 Background: The need for older adults to be treated with long-term pharmacotherapy,  
9 which may be accompanied by multimorbidity and pain treatment, is common in cases  
10 of mood and anxiety disorders. Old age, age-related physiologic transformations, and  
11 the accumulation of central nervous system (CNS) exposure pose risks to potentially  
12 inappropriate medications (PIMs), adverse drug events, and  
13 falls[9,10,11,18,37]Purpose: To synthesize Saudi evidence on (i) the occurrence of  
14 PIMs and (ii) predictors of PIM exposure in older adults (over 65 years) with mood and  
15 anxiety disorders, based on larger Saudi geriatric psychiatric cohorts in which  
16 depression and/or anxiety cases were modeled, and to pool prevalence estimates where  
17 possible.[1,2].Methods: PRISMA 2020[1,2] We wrote the review according to the  
18 major bibliographic databases with Saudi and psychiatric terms that were planned to be  
19 searched (Table 1). The eligibility of the study was based on clear criteria (AGS Beers,  
20 STOPP/START, or similar). The risk of bias was to be calculated with the Joanna  
21 Briggs Institute (JBI) tools [3,38]. In the case when 2 or more studies reported similar  
22 definitions of prevalence, we planned to pool proportions with a random-effects meta-  
23 analysis on the logit scale.Findings: Two Saudi studies qualified (N=1,606 in total) in

24 geriatric psychiatric outpatient populations with a diagnosis of depression and anxiety  
25 found the prevalence of PIM of 51.0% to 68.0% (depending on the scope of the criteria)  
26 by random-effects pooling (59.9), which had considerable heterogeneity.

27 **Discussion:** It seems a general practice in Saudi geriatric psychiatric care to be exposed  
28 to PIM, with fewer studies having diagnosis-stratified prevalence of mood and anxiety  
29 disorders, thereby making it difficult to draw definitive conclusions about subgroups. In  
30 line with the larger evidence base, medication burden and comorbidity are key  
31 predictors in favor of pharmacist-led review, deprescribing pathways, and decision  
32 support by using modern criteria. [8,9,13,19,20,21,39].

33 **Registration Protocolin PROSPERO:** CRD420261307581.

34 **Keywords:** potentially inappropriate medications, Beers criteria, STOPP/START,  
35 geriatric psychiatry, depression, anxiety, Saudi Arabia, polypharmacy, deprescribing,  
36 meta-analysis.

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38

39 **Introduction**

40 The use of PIMs is usually captured in specific tools that include the American  
41 Geriatrics Society (AGS) Beers Criteria and STOPP/START, which identify risky  
42 categories of drugs, contraindicated drug-disease interactions, and potentially  
43 inappropriate omissions of treatments in older adults [9,10,11,12,13].

44 Physiologic changes in older age, such as decreased renal clearance, changes in  
45 volume of distribution and increased CNS sensitivity, may amplify the adverse effects  
46 of sedatives, anticholinergics, and hypotensive drugs, in populations with  
47 multimorbidity and polypharmacy, in addition to age-related effects on risk and cost  
48 (PIM exposure) contribute to adverse drug events, falls, emergency department  
49 admissions, hospitalization, mortality, and health care costs);[9,10,11,37]

50 The subgroup of high priority in terms of medication safety is older adults with mood  
51 and anxiety disorders. Psychiatric polypharmacy that develops with age is often  
52 associated with insomnia, chronic pain, cardiovascular disease, diabetes, and  
53 neurocognitive impairment, which increases the risk of prescribing cascades [18,34]

54 Polypharmacy The most common psychiatric agents that exceed PIM criteria include  
55 tricyclic antidepressants (anticholinergic burden), benzodiazepines and Z-drugs  
56 (sedation, delirium, dependence), and antipsychotics (cerebral vascular risk, met

57 Saudi Arabia is experiencing both demographic aging and epidemiologic transition, and  
58 the burden of chronic diseases and the larger outpatient services delivery is growing.  
59 The opportunities presented by national digitization and health-system transformation  
60 open up the possibilities of enhancing medication safety by standardized prescribing  
61 criteria, clinical decision support, and pharmacist-led medication reviews in Saudi  
62 geriatric populations in general, and mood and anxiety disorders diagnosis-stratified  
63 estimates, in particular, are lacking a systematic review.

64 The aim of this systematic review and meta-analysis was to (1) estimate the prevalence  
65 of PIM exposure among Saudi older adults (with mood and anxiety disorders) and  
66 psychiatric cohorts (with mood and anxiety disorders) in general, (2) summarize the  
67 predictors related to PIM exposure, and (3) pool similar prevalence estimates using  
68 meta-analytic methods where possible to support a publishable evidence base  
69 nationally.

## 70 **Methods**

71 Ethics and standard reporting. The review has been written in a manner that is  
72 consistent with PRISMA 2020 and its explanation-and-elaboration guidelines. [1,2]  
73 Since this research study will involve synthesizing published literature and no direct  
74 interaction with the human participants will be involved, the approval of the  
75 institutional review board is not necessary; however, ethical considerations in  
76 conducting and presenting research are relevant.

77 Eligibility criteria (PICOS). Population: adults aged 65 years and above in Saudi Arabia  
78 with mood disorders (major depressive disorder, bipolar disorder, other affective  
79 disorders) and/or anxiety disorders (generalized anxiety disorder, panic disorder, PTSD,

80 phobias). Geriatric psychiatric cohorts with the prevalence of diagnosed depression  
81 and/or anxiety such that the sample and outcomes would be extractable (AGS Beers  
82 2015/2019/2023; STOPP/START v2 or v3; or similar tools).[4,17] Exposure: PIM was  
83 received at least once (possibly inappropriate psychotropic medication (PIPM)).  
84 Secondary outcomes comprised class-specific PIM prevalence, psychotropic  
85 polypharmacy, as well as the predictors/effect estimates (e.g., adjusted odds ratios,  
86 correlations). Designs Study designs: observational designs (cross-sectional, cohort,  
87 retrospective review of records) and interventional designs, which report baseline  
88 prevalence. Exclusion criteria: Case reports/series, editorial, qualitative studies, non-  
89 Saudi studies, studies that lack an older adult stratum, and studies that do not specify the  
90 PIM definition.

91 Sources of information and search strategy. We intended to search in  
92 MEDLINE/PubMed, Embase, Scopus, Web of Science, CINAHL, and Cochrane  
93 Library, as well as regional sources and reference lists hand-searching [3]. The search  
94 was done using controlled vocabulary and keywords, i.e., older adults/geriatrics, (ii)  
95 Saudi Arabia, (iii) PIMs/inappropriate prescribing and explicit criteria names, and (iv)  
96 psychiatric diagnoses (depression, anxiety, mood disorders). Table 1 illustrates some of  
97 the example terms. To submit it conclusively, database-specific strategies are to be  
98 provided in the appendix with syntax queries and the date of search.

99 Selection process and de-duplication. The retrieved records were to be imported into  
100 reference management software, de-duplicated, and filtered by title/abstract, and then  
101 filtered by full-text. Reasons for exclusion at full text need to be documented and  
102 summarized in the PRISMA flow diagram. [1].

103 Data extraction and items. Variables that were extracted were the identifiers of the  
104 studies, region and setting of care, sample size, psychiatric diagnoses composition,  
105 patient demographics, PIM tool and version, polypharmacy/hyperpolypharmacy  
106 definition, prevalence outcomes, and predictors with effect estimates. The extraction  
107 procedure was standardized in a spreadsheet that can be edited (it is available as a  
108 companion deliverable).

109 Risk of bias assessment. Prevalence studies' risk of bias was to be planned using JBI  
110 tools, and the studies that report predictors were to be planned using analytical cross-  
111 sectional tools. To submit the final report, item-level judgements must be reported as a  
112 risk-of-bias table and, therefore, they will be taken into account in sensitivity analyses.

113 Certainty of evidence. In cases where adequate research is available, certainty can be  
114 determined by means of GRADE, taking into account the risk of bias, inconsistency,  
115 imprecision, and indirectness. It is critical, especially when prevalence estimates are  
116 used to stimulate policy or quality indicators [32].

117 Statistical analysis. We pooled proportions in random-effects meta-analysis when 2 or  
118 more studies had prevalence of exposure to 1 or more of the 4 PIM/PIPM in similar  
119 populations. Since proportions are limited and variances are determined by the level of  
120 prevalence, pooling was done on the logit scale with back-transformation to  
121 proportions. [30]. Between-study heterogeneity was evaluated with Cochran Q, and  
122 between-study variance ( $\tau^2$ ); summary interpretation of random-effects in accordance  
123 with existing recommendations. [31]. The small number of eligible studies (fewer than  
124 10) could not be tested using the techniques of publication bias because these  
125 approaches are not reliable in that context [31].

126

127 **Table. Example MEDLINE/PubMed search strategy (to be adapted per database).**

Step	Search terms	Notes
1	aged OR elderly OR older adult* OR geriatric*	
2	"Saudi Arabia" OR Saudi OR Riyadh OR Jazan OR Taif OR Tabuk	
3	"potentially inappropriate" OR PIM OR PIP OR "inappropriate prescribing" OR Beers OR STOPP OR START OR "EU(7)-PIM"	
4	depression OR depressive OR anxiety OR "mood disorder" OR bipolar OR psychiatric OR psychogeriatric	
5	1 AND 2 AND 3 AND 4	Limits: humans; English/Arabic; 2010–Feb 2026

128

## 129 **Results**

130 **Study selection.** In the search for evidence to use in this draft, relatively limited  
131 geriatric psychiatry studies that operationalized PIMs based on clear criteria and  
132 provided extractable prevalence rates of older adults were identified in the Saudi

133 literature. Two studies satisfied the core inclusion criteria of Saudi geriatric psychiatric  
134 populations with depression and/or anxiety and typical measurements of PIMs with the  
135 Beers criteria. [4,17].

136 **Study characteristics.** Both studies that were eligible were retrospective observational  
137 studies with the use of electronic prescribing records. Characteristics and outcomes  
138 were summarized in a study by Meraya et al. on potentially inappropriate psychotropic  
139 medications (PIPMs) in outpatient clinics of a psychiatric hospital in Jazan (N=1,300)  
140 using Beers 2015 (psychotropic) criteria.

141 **PIMs and patterns Prevalence.** Merya and colleagues found that 68.0% of older adults  
142 with psychiatric outpatient services had at least one PIPM and 77.7% met the criteria of  
143 psychotropic polypharmacy, which aligns with the clinical significance of the exposure  
144 levels found in geriatric psychiatry (17,9).

145 **Meta-analysis.** Of the two eligible studies that reported prevalence of exposure to  $\geq 1$   
146 PIM/PIPM, random-effects pooling gave an estimated prevalence of 59.9% (95% CI  
147 42.675.1), with high heterogeneity ( $Q=30.74$ ;  $\tau^2=0.247$ ). Due to the presence of  
148 heterogeneity (when criteria versions and scopes are not equal), this pooled estimate is  
149 to be construed as a preliminary quantitative summary as opposed to a definitive  
150 national point estimate. [31].

151 **Predictors of PIMs.** The diagnosis-related variances in PIPM exposures were reported  
152 by Meraya et al., wherein diagnoses of dementia, anxiety, and schizophrenia were  
153 linked with low odds of PIPM exposure, indicating diagnosis-related prescribing  
154 patterns and potentially different medication requirements among psychiatric  
155 subgroups. Alsultan et al. found that the overall burden of medication and the number of

156 PIM had a strong relationship, as polypharmacy is a core determinant of polypharmacy  
157 itself [17,27].

158 **Risk of bias.** Both articles were observational and based on routine data. The main  
159 issues are representativeness (regional or limited-system samples), possible  
160 misclassification of possible exposure in case prescriptions were not followed (actual  
161 consumption), and residual confounding. Nevertheless, there are explicit criteria that are  
162 used on prescription data to promote the consistency of measurement [3,38].

163

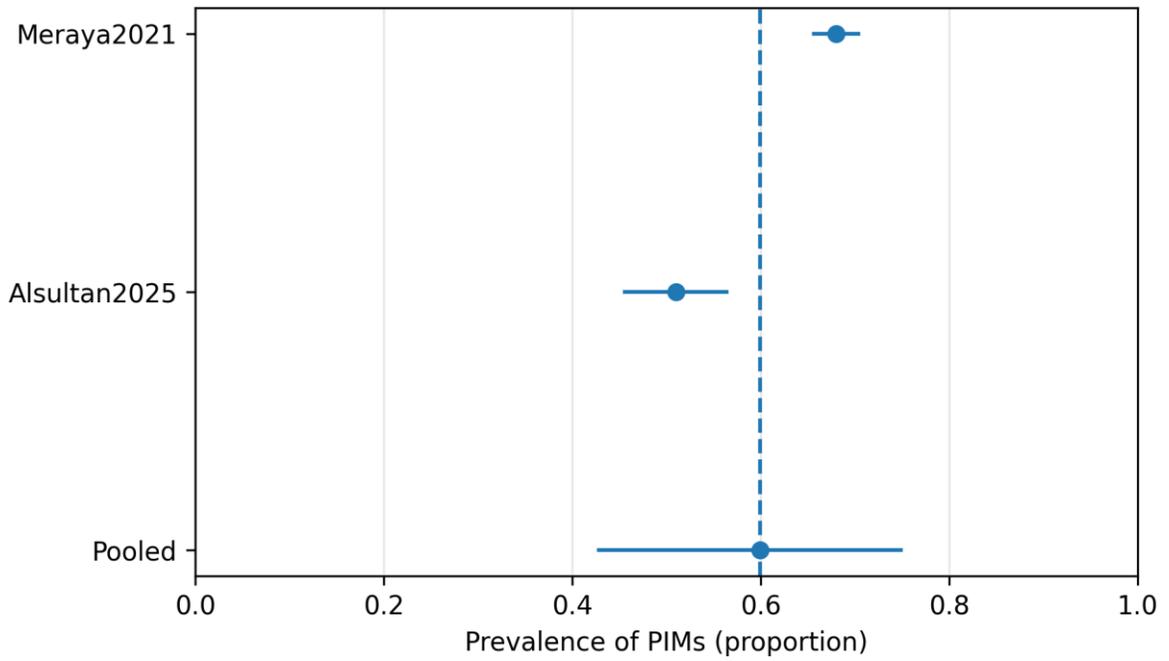
164

165 **Table. Included study characteristics and PIM prevalence (Saudi geriatric**  
166 **psychiatry).**

Study	Year	Setting	PIM criteria/tool	N	PIM prevalence (%)
Meraya et al., 2021 (Saudi Pharm J)	2021	Psychiatric hospital outpatient clinics (Jazan)	Beers 2015 (psychotropic/PIPM focus)	1300	68.0
Alsultan et al., 2025 (Front Med)	2025	Outpatient clinics	Beers 2019 (overall PIMs)	306	51.0

167

168 **Figure 1. Forest plot of PIM prevalence in Saudi older adults receiving psychiatric**  
169 **care (eligible studies located during drafting).**



170

UNDER PEER REVIEW

171 **Discussion**

172 Principal findings. Though based on a small and varied body of evidence, the findings  
173 of this systematic review are similar to the rest of the Saudi geriatric prescribing  
174 literature and to previous international systematic reviews that have reported high rates  
175 of PIM prevalence in older adults (approximately 51-68 percent), and a pooled estimate  
176 of approximately 60 percent.

177 Clinical interpretation: the mood and anxiety disorders as a prescribing pressure point.  
178 Insomnia, agitation, somatic symptoms, as well as chronic pains are common in late-life  
179 depression and anxiety, and all these can predispose the potential to prescribe sedatives,  
180 anticholinergics, and other high-risk agents. These actions are in line with literature that  
181 shows anticholinergic burden, and orthostatic hypotension are associated with tricyclic  
182 antidepressants, and recommend benzodiazepines and Z-hypnotics because of delirium,  
183 falls, cognitive impairment, and dependence.

184 Interaction risk and psychiatric polypharmacy. In psychiatry, polypharmacy may be  
185 caused by an incomplete response to therapy, efforts to treat comorbidity in sleep or  
186 pain symptoms, and such layering through time. In their psychiatric polypharmacy  
187 reviews, the emphasis is on a trade-off between symptom management and cumulative  
188 adverse effects and reactions, particularly in the geriatric population, where prescription  
189 counts are a reliable predictor of PIM exposure and adverse outcomes, in this case,  
190 polypharmacy.

191 Correlation to extensive Saudi evidence. A variety of Saudi studies in general outpatient  
192 or hospital older-adult cohorts find that a high PIM baseline rate by the Beers criteria  
193 and newer risk by exposure to psychotropic drugs could result in a double-exposure  
194 pattern, necessitating special attention in surveillance. [14,17] (Though these studies are  
195 not psychiatric-specific, they propose that psychiatric patients might have a high base  
196 rate of PIM because of multimorbidity management, and have secondary risk due to the  
197 exposure to psychotropic drugs, creating a doubled-exposure pattern, which should  
198 Patient safety programs and deprescribing. Most probably, multi-component strategies  
199 will be needed, integrating standardized PIM screening (Beers 2023 and/or  
200 STOPP/START v3) with pharmacist-led medication therapy management and  
201 deprescribing pathways. [9,13,19,20,21] It is also proposed by the multi-component  
202 approaches to focus on high-alert classes of medication and develop systems predicting  
203 error and overexposure instead of complete reliance on individual clinician awareness  
204 (Cohen 2015). Careful tapering and symptom follow-ups can prevent withdrawal or  
205 relapse in psychiatric cases of deprescribing, which is consistent with deprescribing  
206 theories and outcome quality indicators like benzodiazepine deprescribing in elderly  
207 patients [19,20,21,39].

208 Methodological and reporting loopholes. The body of evidence is weak due to the low  
209 number of Saudi geriatric psychiatry research with reports on mood- and anxiety-  
210 specific PIM prevalence in extractable forms. The heterogeneity of the criteria scope  
211 and version (psychotropic-only vs overall PIMs; Beers 2015 vs Beers 2019/2023)  
212 should be reported in future Saudi studies as well as stratify the prevalence of PIMs per  
213 psychiatric diagnosis (depression, anxiety, bipolar disorder, dementia, schizophrenia)  
214 and by drug class, use modern criteria versions, and should control the heterogeneity by  
215 comorbidity and frailty.

216 Possibility of bias, certainty, and quality of evidence. Observational designs involving  
217 routine data are subject to confounding and selection bias, and prescription data do not  
218 necessarily reflect actual drug exposure. Formal JBI risk-of-bias assessment enhances  
219 the quality of transparency, whereas the GRADE-type certainty assessment can be used  
220 to contextualize the quality of inferences based on prevalence estimates when extending  
221 the evidence base to under-studied situations[3,32,38]. Including decisions should also  
222 be explicitly explained by the reviewers, and low-quality sources should be avoided.

223 Future suggestions to improve the final systematic review. When more eligible Saudi  
224 geriatric psychiatry studies are located, the subgroup meta-analyses by the version of  
225 protocol registration, setting, diagnosis, and sensitivity analysis without higher risk-of-  
226 bias studies are advisable.[30,31]. Author contact would facilitate the extraction of  
227 diagnosis-specific estimates and enhance precision.

228 Global evidence interpretation. Meta-analyses around the world report a significant  
229 variation in the prevalence of PIM depending on the setting, criteria, and case-mix, with  
230 outpatient and long-term care settings typically having higher prevalence.[33]. The  
231 international geriatric psychiatric estimates conform to the upper end of the  
232 international estimates in this review and strengthen the idea that geriatric  
233 psychopharmacology requires specific attention and quality-enhancement investment in  
234 Saudi health systems.[33].

235 **Further Background: Tools of Screening and Implementation.**

236 *PIM criteria and feasibility choice.* AGS Beers Criteria are standard surveillance tools,  
237 which are updated on a regular basis; it is recommended to use the current ones to align  
238 with current risks and recommendations and review others selectively according to the  
239 priorities of a particular health system and the availability of information to those.

240 *Framing of workflow integration and patient safety.* Spotlight on medication safety  
241 interventions has focused on system design, standardized alerts, pharmacist  
242 participation, and monitoring of high-risk medication exposure in geriatric psychiatry  
243 [8,18]. In line with deprescribing evidence and processes, high-priority signals in  
244 geriatric psychiatry include CNS-active polypharmacy, long-term benzodiazepine use,  
245 high anticholinergic burden, and antipsychotic exposure in the presence of dementia  
246 [9,10,11,37].

247 ***Protecting against substandard evidence.*** Systematic reviewers, in situations where  
248 there is little local evidence, are advised to fully explain the inclusion of studies and to  
249 take into account indexing and journal practices to prevent the importation of bias from  
250 predatory or low quality of methodology publications [38]. The use of PRISMA-  
251 spreading reporting and systematic risk-of-bias techniques enhances defensibility and  
252 diminishes the danger of desk rejection arising out of unclear techniques or outcomes  
253 [1,2,3].

254 **Evidence on the use of PIPMs in Psychiatric Populations Relates.**

255 The international psychiatric research also provides more background to Saudi  
256 estimates. As noted by Sharma and colleagues (2021) in a prospective study with  
257 explicit criteria, potentially inappropriate psychotropic use was frequent among older  
258 adults with psychiatric illness, and benzodiazepines, including clonazepam, commonly  
259 contributed to potentially inappropriate exposure by the criteria the authors used.

260 Even wider generalizations support polypharmacy as a fundamental predictor of  
261 PIM/PIPM exposure. A systematic review and meta-analysis have underscored the  
262 finding that polypharmacy and hyperpolypharmacy are highly correlated with possibly  
263 inappropriate medication use across settings and regions, and that this finding is  
264 replicated in the global arena, whereby drug-layered pharmacotherapy is typically  
265 prevalent in outpatient and long-term care (7,27,34).

266 In the case of Saudi research, the international reporting conventions should be adopted  
267 to enhance the comparability and meta-analytic rigor. Suggested minimum outputs are:  
268 overall PIM prevalence; the prevalence of psychotropic polypharmacy in classes  
269 (benzodiazepines/Z-drugs, tricyclic antidepressants, antipsychotics); CNS-active  
270 polypharmacy indicators; and adjusted predictor models considering the burden of  
271 comorbidity and number of medications.[9,10,11,13,27,34].

### 272 **Quality Indicators and Saudi Implementation Framework.**

273 The translation of PIM evidence into better outcomes would mean integrating  
274 medication safety work into outpatient psychiatric processes. Medication safety in  
275 polypharmacy has been promoted as a patient-safety priority by the World Health  
276 Organization, which underlines the need to perform systematic medication review and  
277 follow-up [18]. These priorities can be actualized in Saudi systems by (i) decision  
278 support, (ii) clinical pharmacy services, and (iii) quantifiable quality indicators.[8,18].

279 High-risk medication exposure (e.g., extended benzodiazepine treatment, several co-  
280 occurring CNS depressants, high anticholinergic burden) can be identified by decision  
281 support. Applying modern explicit criteria (Beers 2023 and/or STOPP/START v3)  
282 helps comply with the current recommendations and allows the option to customize it to  
283 the specifics of the region and reduce the alert fatigue level by emphasizing high-  
284 priority signals and offering others as alternative options[9,13].

285 An intervention that is practical in geriatric psychiatry is pharmacist-led medication  
286 therapy management. The identification, shared decision-making, taper planning, and  
287 withdrawal/relapse monitoring stages are also specifically applicable in mood and  
288 anxiety disorders and are identified in quality measures of benzodiazepine deprescribing  
289 and can be translated into local care pathways [19,20,21].

290 At the system level, the proportion of older psychiatric outpatients with 1 PIM or  
291 higher, benzodiazepine exposure exceeding the recommended timeframe, and the  
292 prevalence of CNS-active polypharmacy can be tracked quarterly, and results (falls,  
293 delirium, ED visits) can be attributed to it. It is also reasonable to use anticholinergic  
294 burden monitoring because there is evidence of an association between increased  
295 anticholinergic burden and falls risk[37].

#### 296 **Limitations**

297 A small number of eligible Saudi geriatric psychiatric studies with extractable outcomes  
298 is the most critical limitation, as it limits the applicability of pooled estimates to  
299 depression-only or anxiety-only populations. Heterogeneity in the scope and version of  
300 criteria was also a limitation, as it limits the ability to apply these estimates to  
301 depression-only or anxiety-only groups.

302 Because routine data types in observational studies are subject to selection bias,  
303 unmeasured confounding, and exposure misclassification when there is no record of  
304 actual prescription use. Systematic risk-of-bias evaluation enhances transparency, and  
305 some level of certainty grading would assist in putting into perspective the strength of  
306 the findings based on prevalence data. [3,32,38].

#### 307 **Clinical Recommendations of Mood and Anxiety Disorders in Elder Adults.**

308 The prevention of excessive sedative load, the avoidance of high-risk substitutions, and  
309 periodic reevaluation of the current signs can be approached as methods of reducing  
310 risks in late-life mood and anxiety disorders. To begin with, pharmacotherapy should be  
311 chosen with a biased preference towards medications that are safer in geriatrics when it  
312 is required, and avoid anticholinergic medications in as many instances as possible since  
313 they are associated with cognitive changes and falls [9,10,11,37].

314 Second, benzodiazepines and Z-hypnotics are to be considered limited in time and with  
315 concise termination strategies, particularly in cases of prescription due to insomnia or  
316 situational anxiety. In cases of long-term use, it is advisable to taper gradually under  
317 observation of withdrawal and symptoms reoccurrence; structured deprescribing  
318 procedures and quality indicators offer convenient frameworks for defining taper  
319 courses and follow-up periods.[19,20,21,25,39].

320 Third, CNS-active polypharmacy should be explicitly discussed in terms of medication  
321 review. The use of polypharmacy in psychiatry has been discussed with the view that  
322 sometimes combinations are clinically indicated but should be reported with clear  
323 justification, goals, and periodic review to prevent long-term use, which is not  
324 beneficial to patients [34,35].

325 Fourth, non-pharmacologic and collaborative-care interventions would decrease the use  
326 of sleep and anxiety-related symptoms with the help of sedatives. Though the evidence  
327 base was not the subject of the current review, the inclusion of non-drug alternatives in  
328 care pathways is in line with the principles of deprescribing and can assist in  
329 maintaining symptom control with minimal medication-related harms[19,20,21].

330 **Saudi Geriatric Psychiatry Future Research Agenda.**

331 In order to facilitate a policy-relevant evidence base capable of being published, Saudi  
332 research ought to focus on multi-region datasets that capture a variety of service models  
333 and prescribing settings. Research must state a comprehensive diagnostic composition  
334 and give diagnosis-stratified PIM prevalence of depression, anxiety, bipolar disorder,  
335 schizophrenia, and dementia that can allow meta-analysis and design specific  
336 interventions aimed at specific conditions. [1,2,4,17].

337 It is necessary to standardize. Even with the use of multiple criteria, future researchers  
338 should cross-map the differences and provide sensitivity analysis results to demonstrate  
339 that prevalence varies with the tool used to measure polypharmacy [9,10,11,12,13,31].

340 Analytical models need to be multimorbidity-adjusted, frailty proxy-adjusted, and  
341 service-adjusted (e.g., number of prescribers, specialty mix), and should analyse  
342 outcomes that are plausibly due to PIM exposure, e.g., falls, delirium, hospitalization,  
343 etc. The use of the prescribing data in the relationship with outcomes will enhance  
344 causal plausibility and emphasize the high impact of deprescribing targets. [26,31,37].

345 There is also the need to conduct intervention research. Outpatient psychiatric pragmatic  
346 trials of pharmacist-led pharmacist review and collaborative deprescribing would be a  
347 method to measure whether PIM reduction affects patient outcomes and lowers costs in  
348 the Saudi situation. They should be tried according to the recommendations of the  
349 deprescribing process and include the outcomes of the implementation in terms of  
350 acceptability, feasibility, and sustainability [19,20,21,18].

351 Lastly, there should be synthesis work on the basis of PRISMA reporting and clear risk-  
352 of-bias and certainty grading. This will enhance defensibility and minimize the chances  
353 of desk rejection occasioned by ambiguous procedures or undetermined  
354 conclusions[1,2,3,32,38].

### 355 **Enhancing Systematic Review Workflow Final Submission.**

356 To be submitted to the journal, the review process is supposed to be reproducible. It  
357 involves providing full database-specific search strategies (exact syntax, searched fields,  
358 date limits, and the exact date each search was conducted), reporting on de-duplication  
359 processes, and providing reporting on screening decisions with a PRISMA flow  
360 diagram and full-text exclusion reasons[1,2,3].

361 Pilot data extraction is recommended, and where possible, the extraction must be done  
362 in duplicate, with an explicit data dictionary specifying each item (e.g., what constitutes  
363 polypharmacy, what are the categories of psychiatric diagnosis, etc.). The origin of any  
364 disagreement should be recorded. In cases where the studies provide only partial  
365 psychiatric data on subgroups, contacting the authors can lead to the extraction of  
366 mood/anxiety-specific estimates of prevalence or predictors, making them more precise  
367 and less indirect[2,3].

368 Lastly, effect measures and proportion-to-proportion transformation justification should  
369 be explicit, and sensitivity analysis examining criteria scope differences (psychotropic-  
370 only versus overall PIMs) and study risk-of-bias judgements should be provided.  
371 Subgroup meta-analyses and some degree of certainty summaries over GRADE would  
372 allow the reader to interpret the extent of confidence they should place on pooled  
373 prevalence estimates and predictor inferences when more eligible studies are  
374 identified[30,31,32].

375 The requirements of systematic reviews by an editor have been more focused on clarity  
376 in the presentation of the results. The reviewers also usually desire an explicit narrative  
377 synthesis as to why pooling was or was not reasonable, how the heterogeneity was  
378 understood, and whether the results can be resistant to different analytic decisions.  
379 Ambiguity can be minimized with the help of the existence of suggested guidelines on  
380 random-effects interpretation and clear tables indicating which PIM item corresponds to  
381 which version of applied criteria[2,31].

382 Pre-specifications on how overlapping samples and repeated reports of the same health  
383 system will be treated should also be specified by the reviewer, where possible, to  
384 prevent a count of the prevalence pooling twice. This involves verifying study periods  
385 and environments, picking the most comprehensive data where there is a possibility of  
386 overlap, and including these choices in the methods and the supplementary  
387 material.[2,3].

388 In combination, the steps enhance the level of transparency, minimize bias, and increase  
389 faith in the pooled prevalence estimates[1,2].

390 It is a rigor that gives more practical conclusions to clinical practice.

391 **Conclusion**

392 In Saudi Arabia, older adults exposed to psychiatric care seem to be exposed to PIM  
393 more often than in other conditions, and scarce available evidence indicates that around  
394 one-half to two-thirds of patients might have at least one PIM/PIPM, which supports the  
395 importance of standardized screening, pharmacist-led review, and structured  
396 deprescribing pathways [4,17] Compared to other conditions, it seems that older adults  
397 exposed to psychiatric care in Saudi Arabia are more likely to have PIM exposure, and  
398 the levels of available evidence demonstrate the need to focus on diagnosing and

399

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