

26 of delusions, and lower rate of dyskinesia were associated with nursing home
27 placement.

28 *Conclusions and Implications:* These clinical characteristics suggest that in patients with
29 Parkinsonism who are nursing home residents presence of cognitive impairment and
30 delusions particularly add to the higher overall symptom burden, and more often require
31 specific treatments, including clozapine. Despite similar LEDD, motor severity is higher
32 and dyskinesias, indicative of a response to levodopa, are less common. Falls however
33 also occur less commonly, and dopamine agonists are less frequently used, with lower
34 rates of ICD.

35

36 **Introduction**

37 The global burden of Parkinson disease (PD) has more than doubled over the past
38 generation, and as populations age the number of people with late stage PD will
39 continue to increase.¹ As many as 48 percent of those with late stage PD reside in
40 institutions.² This has economic as well as social consequences; a study in the UK found
41 that those with late-stage PD who reside in nursing homes are the most expensive
42 group to treat, and that accommodation in a nursing home costs approximately four
43 times more than living in one's own home.³ Clinical features and healthcare needs may
44 vary considerably in this population from those of patients living at home. However,
45 relatively little is known on the clinical features and complications of late-stage PD to
46 guide management and address the care needs of this population.

47 Current data on PD patients in nursing homes is country-specific. A study of Medicare
48 records in the US showed PD patients in nursing homes are more likely to be female,
49 have dementia, and be of Afro-Caribbean race.⁴ A prospective study in Norway found
50 age, dependence, dementia, and hallucinations assessed on the UPDRS to be
51 predictors of nursing home admission.⁵ This is in keeping with studies reporting
52 hallucinations to be the main predictor of nursing home placement in the US.⁶ However,
53 a study in Northumberland, UK found the same rate of hallucinations among those living
54 in nursing homes and those living in their own homes.⁷

55 We used data from a large multinational study on the care of late stage Parkinsonism
56 (the CLaSP study) to examine the socio-demographic and clinical data associated with
57 residence in nursing homes. These data may be useful to better address the needs of
58 this population and to improve management guidelines.

59

60 **Methods**

61 The CLaSP study includes 692 people with Parkinsonism in the late stage of the disease
62 (defined as a disease duration of at least 7 years and Hoehn and Yahr stage IV or V in
63 the “On” state or Schwab and England stage 50% or less) recruited by eight centers
64 across six European countries (UK, Germany, Portugal, Sweden, The Netherlands and
65 France). Patients were excluded if: dementia had clearly preceded the onset of motor
66 symptoms; if they were in stages I-III in the “On” state and had Schwab and England
67 50% or greater; or if they had a diagnosis of potentially reversible Parkinsonism such as
68 normal pressure hydrocephalus or drug-induced Parkinsonism, except if persisting
69 following discontinuation of the causative drug. Further study details were previously
70 reported.⁸ In order to include patients not under regular follow up in specialist centers,
71 patients were recruited from specialist and non-specialist settings, including general
72 practitioners, hospitals, nursing homes, patient advocate groups as well as self-help
73 groups. Data were collected in face-to-face interviews with participants and their
74 caregiver. Interviewers attempted to minimize curtailed interviews and missing data by
75 providing appropriate breaks and undertaking repeat visits if required.

76

77 *Assessments*

78 A battery of tests were administered to participants, detailed elsewhere.⁸ We calculated
79 the Charlson Comorbidity index, a measure of 16 comorbidities (including dementia)
80 adjusted for age, which is correlated to life expectancy.⁹ The levodopa equivalent daily
81 dose (LEDD) was calculated from dopaminergic medications.¹⁰ Motor features and
82 complications were assessed using the Unified Parkinson’s Disease Rating Scale
83 (UPDRS) on-state part III and part IV. Activities of daily living were assessed using
84 UPDRS part II and the Schwab and England scale. Dopamine dysregulation
85 syndrome/impulse control disorder was assessed by the relevant question from the

86 MDS-UPDRS.¹¹ Non-motor symptoms were assessed with the non-motor symptom
87 scale (NMSS).¹² Neuropsychiatric symptoms were assessed using the neuropsychiatric
88 inventory (NPI).¹³ Participants were asked to rate how satisfied they were with their
89 overall care on a Likert scale, with 1 corresponding to “very satisfied” and 5 to “very
90 dissatisfied”. The ESAS-PD questionnaire, which aims to form a holistic picture of a
91 patient, was used to assess overall late stage symptom burden.¹⁴
92 Cognitive impairment was assessed in several ways, including presence of an existing
93 diagnosis of dementia, and the mini mental state examination (MMSE), with <24 as a
94 cut-off for cognitive impairment. Additionally, we calculated the level 1 Movement
95 Disorder Society definition of PDD, using MMSE<26, lexical fluency test, and the pill
96 questionnaire were used.¹⁵ We excluded those with a score of 4 on the UPDRS question
97 on depression as severe depression precludes a diagnosis of dementia in these criteria).
98 As well as the raw MMSE score, most centers recorded how many questions a
99 participant was able to attempt (given other non-cognitive comorbidities). In the 627
100 cases where this information was available, we calculated how many mistakes a
101 participant had made. Therefore, in the calculation of the MDS-PDD criteria, those with
102 greater than 4 mistakes were counted as equivalent to having MMSE<26. If the number
103 of questions attempted was not available (n=65), we used the raw MMSE score.

104

105 *Statistical analysis*

106 Differences in continuous variables were analysed by ANOVA or, for non-normally
107 distributed data, the Mann-Whitney U test. Normality was assessed visually. Differences
108 in categorical variables were analysed by the chi-square test. For the univariate
109 analysis, missing data were excluded; we report the numbers of missing data for each
110 variable. One site did not collect comorbidity data.

111 One important difference between patients from different sites was whether participants
112 had idiopathic or atypical Parkinsonism. Some sites recruited specifically from units that
113 focussed on atypical Parkinsonism, and at one site (Lisbon) 53.2% of participants had
114 atypical Parkinsonism. We therefore performed a sensitivity analysis for the univariate
115 analysis, including only patients with idiopathic PD.

116 A logistic regression model was then built using backward stepwise selection, with
117 residential status as dependent variable, and potential contributors to nursing home
118 placement with $p < 0.1$ in univariate analysis as independent variables. Medications were
119 not included in this model, as they may be outcomes rather than predictors of residential
120 status. Similarly, impulse control disorders were not included, as they are likely an
121 outcome of medication use. Comorbidities were not entered as they were not
122 consistently collected at all sites. We did not include the sexual function domain of the
123 NMSS as sexual performance was often not applicable to patients with very severe
124 disease (the relevant NMSS item was then recorded as 0). We imputed missing data
125 using multivariate imputation in chained equations (MICE).¹⁶ Statistical analyses were
126 performed using R.¹⁷

127

128 **Results**

129 All 692 participants fulfilling inclusion criteria had data on current residence. The
130 baseline characteristics of the cohort across sites are shown in the supplementary data.
131 Table 1 shows differences in participant characteristics between those living in their own
132 homes and in nursing homes. Those in nursing homes were significantly less likely to be
133 married, slightly older, and less likely to have idiopathic PD. There was a trend towards
134 a higher comorbidity burden on the Charlson comorbidity index.

135

136 *Motor problems and disability*

137 Table 1 shows differences in disease characteristics between groups. Those in nursing
138 homes had significantly worse motor function and activities of daily living scores.

139 Amongst individual items of the UPDRS part III (data not shown), the greatest difference
140 was seen amongst the items that are important for safe standing, e.g. leg agility, arising
141 from chair, postural instability and gait (all $p < 0.001$). Nevertheless, nursing homes
142 residents experienced fewer falls than those in their own homes, as assessed on
143 question 13 of the UPDRS. Those with greater disability (Schwab and England score
144 $< 50\%$) in the overall group were less likely to fall ($p = 0.001$).

145 Patients in nursing homes reported spending slightly more time in off-periods
146 (supplementary table 2) with no difference in the rate of early morning dystonia. Nursing
147 home residents also had fewer dyskinesias, and found dyskinesias less disabling and
148 less painful.

149

150 *Dementia*

151 Nursing home residents were more likely to have an existing diagnosis of dementia
152 (table 1), had worse MMSE scores, and were more likely to meet the criteria for MDS-
153 PDD. Forty-eight participants with a diagnosis of dementia did not meet the MDS-PDD
154 criteria; three of these had severe depression. Conversely, 70 of those who met the
155 MDS criteria did not have a diagnosis of dementia.

156

157 *Non motor symptoms*

158 Nursing home residents had a significantly higher non-motor symptom burden, as
159 assessed by the NMSS (table 2). They had significantly higher scores in the domains of

160 mood/cognition, perceptual problems/hallucinations, attention/memory, sexual function,
161 and urinary symptoms.

162 Nursing home residents had significantly more neuropsychiatric symptoms as assessed
163 on the NPI (table 2). They had a significantly higher rate of delusions, hallucinations, and
164 depression, with similar findings as on the NMSS. In contrast, dopamine dysregulation
165 syndrome was less common in nursing home residents. Those with impulse control
166 disorders were much more likely to be on a dopamine agonist than those without,
167 (59.2% vs 39.1%, $p<0.001$) and were younger (73.5yrs vs 76.6yrs, $p<0.001$) with no
168 difference in overall LEDD.

169 The ESAS-PD scale assesses palliative symptom burden, with higher scores
170 corresponding to worse feelings or situations. There was no difference in total symptom
171 burden between those in their own home than nursing homes (42.3 vs 43.5, $p=0.54$),
172 although in the domains of confusion (2.5 vs 3.5, $p<0.001$) and stiffness (4.5 vs 5.1,
173 $p=0.036$), those in nursing homes had worse scores, corresponding to our other
174 findings. Wellbeing was similar between the two groups (4.9 vs 4.4, $p=0.048$). There
175 was no difference in satisfaction with care between those in nursing homes and their
176 own homes on the Likert scale (2.1 vs 2.3, $p=0.052$).

177

178 *Medications*

179 There was no significant difference in LEDD between the two groups (table 3). Nursing
180 home residents were less likely to be on dopamine agonists, but were more likely to be
181 on hypnotics, anxiolytics, and antipsychotics. Those on hypnotics and anxiolytics were
182 more depressed, with worse scores in the depression domain of the NPI (for hypnotics,
183 2.54 vs 3.38, $p=0.013$; for anxiolytics, 2.52 vs 4.31, $p<0.001$). Of those on antipsychotics,
184 only three participants were not on quetiapine or clozapine: two on risperidone, one not

185 recorded. Those in nursing homes were 2.5 times more likely to be on clozapine. There
186 was no difference in total NPI score or individual NPI domains between those on
187 clozapine and those on quetiapine (19.2 vs 19.2, $p=0.17$), with variability in prescription
188 rates between countries (see supplementary table 1).

189

190 *Variations between countries*

191 We compared markers of disease severity between nursing home residents in different
192 countries (see supplementary table 3). Motor disease severity, Schwab and England
193 scales, and the NMSS were worse in nursing home residents in all countries, although
194 the size of these differences varies. CLaSP did not prospectively examine admission to
195 nursing homes and these differences may reflect social and cultural differences between
196 countries, or recruitment methods in different countries.

197

198 *Sensitivity analysis*

199 In a sensitivity analysis including only those with idiopathic PD ($n=592$, 85.1%), most of
200 the results of the univariate analysis were unchanged. The difference in prescription rate
201 of antipsychotics failed to reach significance (20.4% vs 25.0%, $p=0.25$) although rates of
202 clozapine prescription remained significantly higher in nursing homes (6.5% vs 14.4%,
203 $p=0.003$).

204

205 *Multivariate analysis*

206 The following factors were included as independent variables in the multivariate
207 analysis: marital status, diagnosis of idiopathic PD, age, current diagnosis of dementia,
208 MDS-PDD status, MMSE <24 , UPDRS section 2 and 3, Hoehn and Yahr stage, NPI
209 domains for delusions, hallucinations and dysphoria/depression; NMSS domains for

210 mood/cognition, perceptual problems/hallucinations, attention/memory, GI tract and
211 urinary tract; and the UPDRS questions on dyskinesias, painful dyskinesias, and
212 disabling dyskinesias.

213 The factors included in the model predicting nursing home status were marital status
214 and presence of cognitive impairment as assessed by MMSE score <24, with risk also
215 increased by severity of delusions on the NPI and worse motor function on the UPDRS
216 part 3 motor and part 2 ADL score. Presence of painful and of disabling dyskinesias
217 were negatively and independently associated with nursing home residence (table 4).
218 Nagelkerke's R² showed the model accounted for 26.7% of the variability in place of
219 residence.

220

221 **Discussion**

222 In this large study of a difficult-to-access group with late-stage Parkinsonism, we report
223 significant differences in the clinical profile of those living in nursing homes and those
224 residing at home, which we have summarised in table 5. These differences are in
225 addition to marital status, a factor known to be a strong predictor of nursing home status
226 in the general population.^{18, 19, 20} WHO estimate that 40% of people with any form of
227 dementia are cared for mainly by their spouse,²¹ and when patients do not have a
228 family to care for them, the role falls to institutions. In our study, those in nursing homes
229 were slightly older but, in contrast to previous studies, gender was not a determining
230 factor for nursing home placement, and differences in general medical comorbidities
231 were less important than PD-related symptoms.

232

233 Nursing home residents had more advanced motor disease severity. However, falls, a
234 major motor complication in advanced disease, were less prevalent among those in

235 nursing homes. This may be due to the lower rate of mobilization due to severity of
236 motor and non-motor problems in those in the most advanced stages, as well as
237 appropriate supervision and protective measures being implemented more easily in a
238 nursing home environment.

239

240 Despite worse motor disease scores and no difference in LEDD, nursing home residents
241 had fewer, less disabling and less painful dyskinesias, and spent slightly more time in off
242 periods. The non-motor side effect profile does not appear to be limiting L-dopa
243 treatment; for example, those on a higher LEDD did not have higher scores in the
244 hallucinations/delusions domain of the NMSS (5.16 for LEDD<600mg vs 6.18, $p=0.093$).
245 One interpretation of these results is that this group of older patients with severe disease
246 may be undertreated. There are other possible reasons for our findings: dyskinesias and
247 motor fluctuations may be less prominent in this population with late-stage disease,
248 there could be increasing unresponsiveness at the late disease stages, or the presence
249 of other non-motor features may have necessitated cautious approaches to the
250 treatment of motor problems.

251

252 As expected, nursing home residents were much more likely to have cognitive
253 impairment. In addition, application of the MDS-PDD criteria in our cohort indicated that
254 70 participants (10.1%) with dementia in this cohort were previously undiagnosed with
255 dementia. The MDS-PDD criteria have previously been found to be more sensitive than
256 DSM IV criteria for patients with PDD.²² Conversely, not all participants with an existing
257 diagnosis of dementia fulfilled these criteria. Although we did not systematically collect
258 information on how our participants received a diagnosis of dementia, and clinical
259 diagnosis of dementia using different criteria or in-depth neuropsychological assessment

260 may provide different results from that using the MDS-PDD criteria, it is also possible
261 that there is overdiagnosis of dementia in patients with severe disability due to other
262 features of Parkinsonism in the very advanced stage.

263

264 Psychiatric complications were more prevalent in those in nursing homes. Goetz et al
265 have previously found hallucinations/delusions, as measured by the thought-disorder
266 question on the UPDRS, to be a predictor of nursing home admission in a case-control
267 study in the US;⁶ Aarsland et al echoed this finding in their prospective study in Norway.⁵
268 Using the NPI, which allows assessment of neuropsychiatric features in greater depth,
269 we found that delusions are more strongly associated with nursing home placement than
270 hallucinations, a clinically important difference as delusions with firmly held beliefs may
271 be more distressing than hallucinations, and may require more aggressive intervention.
272 Correspondingly, antipsychotics were used more often in nursing homes (23.1% vs
273 31.4%, $p=0.03$), and in particular clozapine use was nearly 2.5 times higher in nursing
274 homes. In Europe, clozapine is the most effective antipsychotic available for PD
275 psychosis. Its efficacy has been demonstrated in placebo controlled trials.^{23,24} However,
276 because of monitoring requirements, clozapine is sometimes reserved for more severe
277 cases of psychosis not responding to other strategies.²⁵ The differences we observed in
278 prescribing patterns of clozapine may therefore either be due to greater severity of
279 delusions in those in nursing homes, or greater monitoring ability in this setting.
280 Prescription rates varied by country, with clozapine use highest in Portugal (20%) and
281 the Netherlands (15%). In both countries, any doctor can prescribe clozapine and any
282 pharmacist can dispense clozapine.²⁶ Clozapine use was lowest in the UK (2%), where
283 clozapine can only be dispensed from registered pharmacies and prescribed by

284 registered prescribers, who are not typically involved in the care of patients with late-
285 stage PD.

286

287 In the US, pimavanserin, an inverse agonist of the 5HT-2A receptor, is licensed for
288 psychosis in PD; clozapine use for PD psychosis is off-label, although the Movement
289 Disorder Society describe both clozapine and pimavanserin as efficacious.²³ There are
290 no head-to-head trials of clozapine and pimavanserin, but pimavanserin does not require
291 specialized monitoring, which may make it more appropriate for frail patients.²⁷
292 However, the manufacturer does not yet have European licence.

293

294 Nursing home residents had a lower rate of impulse control disorders than those
295 residing at home. This is likely to primarily be related to their lower rate of dopamine
296 agonist prescriptions,²⁸ although additional factors such as restricted access cannot be
297 excluded and nursing home residents were also older.

298

299 *Limitations*

300 The key limitation to our study is that participating sites were heterogeneous and
301 participants were recruited in different ways in different sites and from different
302 healthcare systems. However, we aimed to gain a comprehensive picture of patients
303 with late-stage Parkinsonism across countries and included patients from various
304 settings, reflective of clinical practice. In addition, we did not restrict our analysis to those
305 with idiopathic PD as the clinical features and needs of patients with advanced stage
306 parkinsonism are often similar. A sensitivity analysis restricted to those with idiopathic
307 PD reflected the overall findings. Participants had significant disability and were not
308 always able to complete all parts of the study, and there were therefore missing data in

309 some variables. We have imputed missing data in the multivariate model, but cannot
310 eliminate bias in the pattern of missingness.

311

312 **Conclusions and implications**

313

314 Our analysis shows that PD patients in nursing homes have more severe motor
315 symptoms, psychiatric symptoms, and higher rates of dementia. We may be
316 undertreating the motor symptoms of those in nursing homes; conversely, psychiatric
317 symptoms appear to be treated more appropriately in nursing homes. Nonetheless,
318 nursing homes are safe places to live; participants living there had fewer falls, and
319 participants had same satisfaction with care in and out of nursing homes..

320 Given that those living in nursing homes have more severe disease, are at greatest
321 need, and are less likely to attend outpatient clinics or to have family input, efforts should
322 be made to provide specialist input for these patients in their place of care.

323

324 **References**

- 325 1. Ray Dorsey E, Elbaz A, Nichols E, et al. Global, regional, and national burden of
326 Parkinson's disease, 1990–2016: a systematic analysis for the Global Burden of
327 Disease Study 2016. *Lancet Neurol.* 2018;17(11):939-953. doi:10.1016/S1474-
328 4422(18)30295-3
- 329 2. Hely MA, Reid WGJ, Adena MA, et al. The Sydney Multicenter Study of
330 Parkinson's disease: The inevitability of dementia at 20 years. *Mov Disord.*
331 2008;23(6):837-844. doi:10.1002/mds.21956

- 332 3. Findley L, Aujla M, Bain PG, et al. Direct economic impact of Parkinson's disease:
333 A research survey in the United Kingdom. *Mov Disord*. 2003;18(10):1139-1145.
334 doi:10.1002/mds.10507
- 335 4. Safarpour D, Thibault DP, Desanto CL, et al. Nursing home and end-of-life care in
336 Parkinson disease. *Neurology*. 2015;85(5):413-419.
337 doi:10.1212/WNL.0000000000001715
- 338 5. Aarsland D, Larsen JP, Tandberg E, Laake K. Predictors of nursing home
339 placement in Parkinson's disease: A population-based, prospective study. *J Am*
340 *Geriatr Soc*. 2000;48(8):938-942. doi:10.1111/j.1532-5415.2000.tb06891.x
- 341 6. Goetz CG, Stebbins GT. Risk factors for nursing home placement in advanced
342 Parkinson's disease. *Neurology*. 2012;43(11):2227-2227.
343 doi:10.1212/wnl.43.11.2227
- 344 7. Porter B, Henry SR, Gray WK, Walker RW. Care requirements of a prevalent
345 population of people with idiopathic Parkinson's disease. *Age Ageing*.
346 2009;39(1):57-61. doi:10.1093/ageing/afp199
- 347 8. Balzer-Geldsetzer M, Ferreira J, Odin P, et al. Study protocol: Care of Late-Stage
348 Parkinsonism (CLaSP). *BMC Neurol*. 2018;18(1):185. doi:10.1186/s12883-018-
349 1184-3
- 350 9. Charlson ME, Pompei P, Ales KL, MacKenzie CR. A new method of classifying
351 prognostic comorbidity in longitudinal studies: development and validation. *J*
352 *Chronic Dis*. 1987;40:373-83
- 353 10. Parkinson's measurement levodopa equivalent dose calculator,
354 <https://www.parkinsonsmeasurement.org/toolBox/levodopaEquivalentDose.htm>
355 [Accessed 12th July 2019](#)

- 356 11. Goetz CG, Tilley BC, Shaftman SR, et al. Movement Disorder Society-Sponsored
357 Revision of the Unified Parkinson's Disease Rating Scale (MDS-UPDRS): Scale
358 presentation and clinimetric testing results. *Mov Disord*. 2008;23(15):2129-2170.
359 doi:10.1002/mds.22340
- 360 12. Chaudhuri KR, Martinez-Martin P, Brown RG, et al. The metric properties of a
361 novel non-motor symptoms scale for Parkinson's disease: Results from an
362 international pilot study. *Mov Disord*. 2007;22(13):1901-1911.
363 doi:10.1002/mds.21596
- 364 13. Cummings JL, Mega M, Gray K, et al. The neuropsychiatric inventory:
365 comprehensive assessment of psychopathology in dementia. *Neurology*.
366 1994;44:2308–14.
- 367 14. Miyasaki JM, Long J, Mancini D, et al. Palliative care for advanced Parkinson
368 disease: An interdisciplinary clinic and new scale, the ESAS-PD. *Park Relat*
369 *Disord*. 2012;18(SUPPL. 3):S6-S9. doi:10.1016/j.parkreldis.2012.06.013
- 370 15. Dubois B, Burn D, Goetz C, et al. Diagnostic procedures for Parkinson's disease
371 dementia: Recommendations from the Movement Disorder Society Task Force.
372 *Mov Disord*. 2007;22(16):2314-2324. doi:10.1002/mds.21844
- 373 16. van Buuren S, Groothuis-Oudshoorn K. mice: Multivariate Imputation by Chained
374 Equations in R. *Journal of Statistical Software*, 2011;45(3), 1-
375 67. <https://www.jstatsoft.org/v45/i03/>.
- 376 17. R Core Team (2019). R: A language and environment for statistical computing. R
377 Foundation for Statistical Computing, Vienna, Austria. URL [https://www.R-](https://www.R-project.org/)
378 [project.org/](https://www.R-project.org/).

- 379 18. Thomeer MB, Mudrazija S, Angel JL. Relationship status and long-term care
380 facility use in later life. *Journals Gerontol - Ser B Psychol Sci Soc Sci*.
381 2016;71(4):711-723. doi:10.1093/geronb/gbv106
- 382 19. Hedinger D, Hämmig O, Bopp M. Social determinants of duration of last nursing
383 home stay at the end of life in Switzerland: A retrospective cohort study Health
384 services research. *BMC Geriatr*. 2015;15(1):1-8. doi:10.1186/s12877-015-0111-3
- 385 20. Everink IHJ, Van Haastregt JCM, Van Hoof SJM, et al. Factors influencing home
386 discharge after inpatient rehabilitation of older patients: A systematic review
387 Health services research. *BMC Geriatr*. 2016;16(1). doi:10.1186/s12877-016-
388 0187-4
- 389 21. Ander Wimo, Serge Gauthier, Martin Prince M. Global estimates of informal care.
390 2018:Alzheimer's Disease International. [https://www.alz.co.uk/adi/pdf/global-
391 estimates-of-informal-care.pdf](https://www.alz.co.uk/adi/pdf/global-estimates-of-informal-care.pdf).
- 392 22. Martinez-Martin P, Falup-Pecurariu C, Rodriguez-Blazquez C, et al. Dementia
393 associated with Parkinson's disease: Applying the Movement Disorder Society
394 Task Force criteria. *Park Relat Disord*. 2011;17(8):621-624.
395 doi:10.1016/j.parkreldis.2011.05.017
- 396 23. Seppi K, Ray Chaudhuri K, Coelho M, et al. Update on treatments for nonmotor
397 symptoms of Parkinson's disease—an evidence-based medicine review. *Mov
398 Disord*. 2019;34(2):180-198. doi:10.1002/mds.27602
- 399 24. Frieling H, Hillemacher T, Ziegenbein M, Neundörfer B, Bleich S. Treating
400 dopamimetic psychosis in Parkinson's disease: Structured review and meta-
401 analysis. *Eur Neuropsychopharmacol*. 2007;17(3):165-171.
402 doi:10.1016/j.euroneuro.2006.08.007

- 403 25. Grosset DG, Macphee GJA, Nairn M. Diagnosis and pharmacological
404 management of Parkinson's disease: Summary of SIGN guidelines. *BMJ*.
405 2010;340(7739):206. doi:10.1136/bmj.b5614
- 406 26. Nielsen J, Young C, Ifteni P, et al. Worldwide differences in regulations of
407 clozapine use. *CNS Drugs*. 2016;30(2):149-161. doi:10.1007/s40263-016-0311-1
- 408 27. Hawkins T, Berman B. Pimavanserin: A novel therapeutic option for Parkinson
409 disease psychosis. *Neurol Clin Pract*. 2017;7(4):282.1-282.
410 doi:10.1212/cpj.0000000000000380
- 411 28. Vargas AP, Cardoso FEC. *Impulse Control and Related Disorders in Parkinson's*
412 *Disease*. Vol 76. 1st ed. Elsevier Inc.; 2018. doi:10.1590/0004-282X20180052
413

414 *Table 1: Participant demographics and disease characteristics by place of residence*
 415 *(mean (SD) or n (%)).*

	Nursing home (N=194)	Own home (N=498)	Total (N=692)	Missing (n)	p value
Female	98 (50.5%)	221 (44.4%)	319 (46.1%)	0	0.15
Age	78.1 (7.6)	75.4 (8.5)	76.1 (8.4)	1	< 0.001
Married	80 (41.5%)	356 (71.9%)	436 (63.4%)	4	< 0.001
Years of education	9.9 (4.6)	10.0 (3.7)	10.0 (3.9)	24	0.78
Charlson comorbidity index	4.9 (1.3)	4.7 (1.5)	4.8 (1.4)	109	0.09
Idiopathic PD	146 (76.0%)	443 (89.0%)	589 (85.4%)	2	<0.001
Disease duration (years)	15.9 (8.3)	15.2 (7.4)	15.4 (7.7)	7	0.28
UPDRS II	29.6 (7.3)	26.4 (7.8)	27.3 (7.8)	3	< 0.001
UPDRS III	53.0 (16.4)	45.1 (15.6)	47.3 (16.2)	6	< 0.001
UPDRS IV	4.7 (3.2)	5.3 (3.7)	5.1 (3.5)	3	0.26
Schwab and England	27.7 (14.1)	36.3 (16.1)	33.9 (16.0)	0	< 0.001
H&Y stage 5	109 (56.2%)	120 (24.1%)	229 (33.1%)	0	< 0.001
DDS/ICD	0.2 (0.7)	0.4 (0.9)	0.3 (0.8)	93	0.002
MMSE<24	111 (65.3%)	182 (39.1%)	293 (46.1%)	57	< 0.001
Dementia diagnosis	99 (51.0%)	156 (31.4%)	255 (36.9%)	1	< 0.001
MDSPDD	81 (47.9%)	152 (32.8%)	233 (36.9%)	60	< 0.001
Falls (any) (UPDRS item 13)	130 (68.8%)	389 (79.1%)	519 (76.2%)	11	0.005

416

417 *Table 2: Non-motor symptom scale domain scores and neuropsychiatric inventory*
 418 *domain scores (mean (SD))*

	Nursing home (N=194)	Own home (N=498)	Total (N=692)	Missing (n)	p value
<i>Non-motor symptom scale</i>					
D1 Cardiovascular	4.2 (6.1)	3.0 (4.6)	3.3 (5.1)	37	0.21
D2 Sleep/fatigue	14.7 (10.0)	14.9 (10.6)	14.8 (10.4)	40	0.97
D3 Mood/cognition	22.6 (18.2)	17.7 (16.3)	19.0 (17.0)	38	0.002
D4 Perceptual problems/hallucinations	7.1 (8.1)	5.3 (7.6)	5.8 (7.8)	39	0.002
D5 Attention/memory	17.4 (13.2)	13.5 (12.0)	14.6 (12.4)	38	0.001
D6 GI tract	12.6 (8.1)	11.5 (8.5)	11.8 (8.4)	34	0.06
D7 Urinary	19.3 (13.2)	16.2 (12.7)	17.0 (12.9)	45	0.006
D8 Sexual function	11.7 (10.0)	8.9 (10.1)	9.7 (10.1)	74	<0.001
D9 Miscellaneous	9.7 (8.0)	10.4 (8.6)	10.2 (8.4)	43	0.46
NMSS total	118.2 (52.7)	102.3 (51.5)	106.4 (52.3)	100	0.001
<i>Neuropsychiatric inventory</i>					
A: Delusions	1.6 (3.1)	0.9 (2.4)	1.2 (2.7)	73	<0.001
B: Hallucinations	2.2 (3.1)	1.6 (2.9)	1.8 (3.0)	70	0.005
C: Agitation/aggression	1.1 (2.5)	1.1 (2.3)	1.1 (2.3)	73	0.23
D: Dysphoria/depression	3.0 (3.4)	2.5 (3.3)	2.7 (3.3)	75	0.03
E: Anxiety	2.0 (3.2)	1.9 (2.9)	1.9 (3.0)	73	0.92
F: Euphoria/elation	0.2 (0.8)	0.1 (0.7)	0.1 (0.7)	70	0.80
G: Apathy/indifference	3.5 (4.3)	2.9 (3.9)	3.1 (4.0)	70	0.19
H: Disinhibition	0.3 (1.1)	0.4 (1.6)	0.3 (1.4)	74	0.42
I: Irritability/lability	1.1 (2.4)	1.0 (2.2)	1.1 (2.2)	72	0.84
J: Aberrant motor	1.6 (3.3)	1.3 (2.7)	1.4 (2.9)	76	0.34
K: Nighttime behaviour	2.0 (3.3)	2.3 (3.4)	2.2 (3.4)	83	0.23
L: Appetite/Eating	1.8 (3.1)	1.68(2.9)	1.8 (3.0)	81	0.40

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420 *Table 3: Medications in nursing home residents vs participants residing at home*

	Nursing home (N=194)	Own home (N=498)	Total (N=692)	Missing (n)	p value
LD dose (mg)	749.8 (441.6)	711.1 (565.9)	722.0 (533.6)	15	0.39
LD dose >600mg	120 (62.5%)	271 (55.9%)	391 (57.8%)	15	0.12
On dopamine agonist	53 (28.0%)	224 (45.4%)	277 (40.6%)	10	< 0.001
On hypnotic	51 (27.0%)	79 (16.1%)	130 (19.1%)	13	0.001
On antipsychotic	59 (31.4%)	113 (23.1%)	172 (25.4%)	14	0.026
On antidepressant	81 (42.9%)	170 (34.6%)	251 (36.9%)	12	0.046
On anxiolytic	33 (17.5%)	37 (7.6%)	70 (10.3%)	13	< 0.001
On clozapine	34 (17.5%)	35 (7.0%)	69 (10.0%)		< 0.001
On quetiapine	23 (11.9%)	77 (15.5%)	100 (14.5%)		0.23

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422 *Table 4: Logistic regression of factors contributing to residential status, with backwards*
 423 *stepwise selection performed for model reduction. The model accounts for 26.7% of the*
 424 *variability in residential status (Nagelkerke's R^2).*

	Odds ratio	2.5% confidence interval	97.5% confidence interval	p value
Unmarried	4.86	3.31	7.24	<0.001
MMSE<24	2.56	1.70	3.89	<0.001
UPDRS II	1.03	0.99	1.06	0.13
UPDRS III	1.02	1.00	1.03	0.03
NPI: delusions	1.08	1.01	1.15	0.03
Disabling dyskinesias	0.81	0.64	1.01	0.06
Painful dyskinesias	0.65	0.41	0.97	0.05

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426

427 **Table 5: Key findings and implications for care**
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Key findings:

- *Being older, unmarried and having worse cognition are associated with living in a nursing home in patients with PD*
- *Patients with PD in nursing homes have worse motor severity, but do not have worse dyskinesia and do not receive higher dopaminergic treatment.*
- *Those in nursing homes also have more hallucinations, delusions and depression and also have higher treatment rates with psychotropic medications, including clozapine.*
- *Falls are less common in patients with nursing homes*
- *Satisfaction with care was similar between those living at home and in nursing homes*

Implications for care:

- *The more severe motor disease of those in nursing homes may be undertreated.*
 - *Psychiatric symptoms are also higher in those living in a nursing home but they are also more frequently treated than those living in their own homes.*
 - *Nursing homes are safe places with lower rates of falls and satisfaction with care is similar in and out of nursing homes*
 - *Efforts should be made to provide specialist input to patients, regardless of their place of care.*
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