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3 **Access to means of lethal overdose among psychiatric patients with co-morbid physical**
4 **health problems: analysis of national suicide case series data from the United Kingdom**

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6 Alexandra Pitman^{1,2}, Su-Gwan Tham³, Isabelle M. Hunt³, Roger T. Webb^{3,4,5}, Louis
7 Appleby³, Nav Kapur^{3,4,5,6}

8 ¹UCL Division of Psychiatry, University College London, UK

9 ²Camden & Islington NHS Foundation Trust, St Pancras Hospital, London NW1 0PE,
10 UK

11 ³Centre for Mental Health and Safety, University of Manchester, UK

12 ⁴Manchester Academic Health Sciences Centre (MAHSC), University of
13 Manchester, UK

14 ⁵NIHR Greater Manchester Patient Safety Translational Research Centre, UK

15 ⁶Greater Manchester Mental Health NHS Foundation Trust, Manchester, UK

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18 **Corresponding author:** Dr Alexandra Pitman MRCPsych PhD, Associate Professor, UCL
19 Division of Psychiatry, University College London, Maple House, 149 Tottenham Court Rd,
20 London W1W 7NF, UK a.pitman@ucl.ac.uk

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23

24 **Abstract**

25

26 **Background:** Many physical health problems are associated with elevated suicide risk whilst
27 also providing access to means of overdose. We aimed to investigate whether psychiatric
28 patients with physical co-morbidities who die by suicide were more likely than those without
29 co-morbidities to self-poison with non-psychotropic medications.

30 **Methods:** We analysed data on 14,648 psychiatric patients who died by suicide in England
31 & Wales during 2004-2015, as recorded by the National Confidential Inquiry into Suicide and
32 Safety in Mental Health. Using logistic regression models adjusted for age, gender, ethnicity,
33 and primary drug dependence/misuse we compared patients diagnosed with physical co-
34 morbidities *versus* those without to assess whether a greater proportion of the former had
35 died by overdose, and medication prescribed to treat such disorders (e.g. opioids, insulin).

36 **Results:** 24% (n=3525) were recorded as having physical co-morbidity. A greater proportion
37 of these individuals died by self-poisoning than those without physical co-morbidity (37% vs.
38 20%, $p<.001$; adjusted OR 2.47; 95% CI 2.26-2.70), and they were more likely to have used
39 medications for a physical health disorder in overdose (50% vs. 34%; adjusted OR 2.10; 95%
40 CI 1.80-2.46), particularly opioids (30% vs. 22%; $p<0.001$), paracetamol/opioid compounds
41 (11% vs. 7%, $p<.001$) and insulin (4% vs. 1%, $p<.001$).

42 **Limitations:** Use of survey data may have resulted in under-reporting of physical health
43 problems and/or overdose medications.

44 **Conclusions:** Overdose, rather than hanging, is the leading cause of suicide among
45 psychiatric patients with physical co-morbidities, particularly using non-psychotropic
46 medications. There is potential for means restriction in preventing suicide among these
47 patients.

48

49 **Key words:** suicide; means restriction; overdose; physical health co-morbidities; prescribing

50 Background

51 Restricting access to lethal means is the suicide prevention intervention with the best
52 evidence for effectiveness (Zalsman et al., 2016). Means restriction has most public
53 health impact in relation to common, high-lethality suicide methods. After hanging,
54 poisoning is the second commonest method of suicide in England, Scotland and Wales;
55 accounting for 18% of male and 36% of female suicides in 2016 (Office for National
56 Statistics (ONS), 2016a). Restricting means of overdose entails impeding access to the
57 medication load available to at-risk persons to a level that, even if taken in one dose,
58 will not pose serious harm (Hawton et al., 2013). Usually this involves adjusting the
59 frequency (and therefore volume) of medication prescribed or available over-the-
60 counter. The value of this approach is exemplified in the significant reduction in fatal
61 paracetamol overdoses associated with UK legislation restricting pack size of over-the-
62 counter analgesics (Barber & Miller, 2014). Where methods are not easily substituted
63 by others, means restriction does not necessarily prompt means substitution
64 (Sarchiapone et al., 2011). Indeed, the UK withdrawal of co-proxamol was associated
65 with a significant reduction in deaths involving co-proxamol poisoning but no
66 corresponding increase in deaths involving analgesics (Hawton et al., 2009). Physical
67 disorders such as cancer (Henson et al., 2019; Ahmedani et al., 2017), osteoporotic
68 fracture (Chang et al., 2018; Webb et al., 2012), back pain (Ahmedani et al., 2017),
69 diabetes (Ahmedani et al., 2017; Webb et al., 2014), and heart disease (Ahmedani et
70 al., 2017; Wu et al., 2018) are associated with an increased risk of suicide, and may
71 provide affected individuals access to potentially lethal doses of prescribed medication
72 (Gorton et al., 2016). In a Swedish sample, 9% of patients diagnosed with diabetes who
73 died from fatal poisoning had taken overdoses of diabetic drugs (Webb et al., 2014).
74 For people with pain conditions, particularly chronic pain (Petrosky et al., 2018),
75 opioids are a key target for means restriction, especially as the association of non-
76 cancer pain and suicide risk is independent of psychiatric illness (Ilgen et al, 2013). In
77 2016 opioids accounted for 54% of all fatal drug poisonings (suicides and accidental
78 overdoses) in England & Wales (ONS, 2016b). The most common opioid responsible
79 was heroin and/or morphine (ONS, 2016b), although available data do not indicate
80 what proportion involved 'street' opioids or those prescribed for chronic pain. With
81 approximately 6,000 people dying by suicide in the UK annually (ONS, 2016c), there is

82 great interest among both clinicians and policymakers in the potential to restrict the
83 volume of potentially lethal medication available to patients with physical illnesses.
84 However, an improved understanding is needed regarding the role of access to these
85 medications in pathways to suicide.

86 Our research question was whether a greater proportion of psychiatric patients also
87 diagnosed with physical illnesses who die by suicide poison themselves compared to
88 individuals without physical co-morbidities, and whether they are more likely to self-
89 poison using medication prescribed to treat their physical health problems. We
90 thereby aimed to explore the potential for means restriction interventions in a sub-
91 group of psychiatric patients with co-morbid physical illnesses. Using national suicide
92 case series data on psychiatric patients who died by suicide in England & Wales during
93 2004-2015, we aimed to describe the sociodemographic and clinical characteristics of
94 psychiatric patients with a diagnosis of a co-morbid physical illness. We tested the
95 hypotheses that:

- 96 • a greater proportion of deceased patients diagnosed with co-morbid
97 physical illness fatally poisoned themselves than such patients without co-
98 morbidity
- 99 • a greater proportion of deceased patients diagnosed with physical illness
100 who fatally poisoned themselves overdosed on medication used for
101 physical health problems *versus* such patients who died by intentional self-
102 poisoning without co-morbidity
- 103 • among deceased patients with co-morbid physical illness who fatally
104 poisoned themselves with medications prescribed to treat these
105 conditions, a higher proportion had been prescribed the medication taken
106 in overdose *versus* those without physical health disorders
- 107 • among deceased patients diagnosed with cancer, diabetes, and pain
108 conditions the proportion who fatally self-poisoned using physical health
109 medications was greater than among such patients diagnosed with other
110 physical illnesses. These conditions have been linked with elevated suicide
111 risk (Henson et al., 2019; Ahmedani et al., 2017; Webb et al., 2012), whilst

112 also providing access to medications that are highly toxic in overdose
113 (Gorton et al, 2016).

114

115 **METHODS**

116 **Study dataset**

117 Questionnaire data were collected as part of the National Confidential Inquiry into
118 Suicide and Safety in Mental Health (Appleby et al., 1999). This database provides a
119 national case series of patients under the care of mental health services who have
120 died by suicide across the UK (i.e. England, Scotland, Wales and Northern Ireland). A
121 detailed description of the National Confidential Inquiry's methodology is available
122 elsewhere (Windfuhr et al., 2008). In brief, firstly, data on all deaths in England &
123 Wales receiving a verdict of suicide or unnatural death of undetermined intent ('open'
124 verdict) at coroner's inquest were received from the Office for National Statistics
125 (ONS). Suicide research conducted in the UK conventionally includes open verdicts to
126 avoid underestimating the number of suicide deaths (Linsley et al., 2001). Second,
127 administrative contacts at NHS Trusts or Health Boards in the deceased person's
128 district of residence identified whether contact had been made with secondary mental
129 health services in the 12 months prior to death. Third, for those individuals with
130 psychiatric contact, detailed data were collected via a questionnaire sent to the
131 clinicians who had been responsible for that psychiatric patient's care. The
132 questionnaire captured information on suicide method, demographic details, clinical
133 characteristics, including any major physical illness at the time of death, aspects of
134 care and treatment received.

135 **Ethical approvals**

136 The National Confidential Inquiry has research ethics approval from North West - GM
137 South REC (reference: ERP/96/136) and Section 251 Approval under the NHS Act 2006
138 (reference: PIAG 4-08(d)/2003), allowing collection of patient identifiable data for
139 medical research.

140 **Measures**

141 We defined physical health conditions on the basis of responses to the questionnaire
142 item: *“Did the patient have a major physical illness at the time of death? (include
143 conditions even if well controlled by treatment)”*. Free text responses to a further
144 specifier permitted categorisation of conditions into those corresponding to
145 International Classification of Diseases (ICD-10, 1992) categories (diseases of the
146 musculoskeletal system, circulatory system, nervous system, digestive system, and
147 endocrine disease). We used clinician-derived search terms to identify conditions with
148 heterogeneous descriptors. For our sub-analyses we defined a specific diabetes
149 category and overlapping categories for pain conditions and cancer.

150

151 We categorised the substances used in self-poisoning on the basis of fixed-choice
152 responses to the questionnaire item: *“If self-poisoning, specify substance (if more than
153 one substance, select most likely cause of death)”*, to develop a categorical measure of
154 whether or not these drugs are prescribed to treat physical illnesses. This was coded
155 by a psychiatrist (AP), including free text responses to the *“Other drug (please specify)”*
156 category. Categories within the physical illness treatment group were: opioids
157 (morphine, codeine and methadone), paracetamol/opioid compounds, other
158 analgesics, insulin, cardiac medications, and other specified drugs for physical
159 conditions (Box 1).

160

161 We categorised the source of substances used in self-poisoning cases using fixed-
162 choice responses to the relevant questionnaire item (prescribed for the patient;
163 prescribed for someone else; not prescribed). For data collected from 2012, where
164 opioids were reported in self-poisoning cases, further detail was available on whether
165 these were prescribed for the patient for treatment of pain or for the treatment of
166 drug misuse, prescribed for someone else, obtained illicitly, or obtained over-the-
167 counter. We analysed drugs used for physical health conditions in cases of self-
168 poisoning, irrespective of whether they had been prescribed for the patient or for
169 someone else, or obtained illicitly.

170

171 **Statistical analysis**

172 Chi-square tests (with a 2-sided p-value threshold of <.05) were used to compare
173 proportional distributions of sociodemographic and clinical characteristics between
174 psychiatric patients with *versus* without diagnosed physical illness. We fitted logistic
175 regression models to estimate the strength of these associations, with and without
176 adjustment for age, gender, ethnicity, and presence of a primary drug
177 dependence/misuse disorder (which may itself be associated with chronic pain
178 conditions). Odds ratios (ORs) and their 95% confidence intervals (CIs) were presented.
179 Pairwise deletion was applied to address missing data; ie. if an item of information was
180 unknown, the case was removed from the analyses of that variable. All analyses were
181 conducted using Stata version 15.0 (StataCorp, 2017).

182 **Sensitivity analyses**

183 We conducted sensitivity analyses to assess robustness of findings when using a more
184 stringent definition of medications that may have been prescribed to treat physical
185 health conditions. This excluded drugs that can be used to treat psychiatric conditions
186 (e.g. gabapentin and pregabalin for anxiety) or to address the side effects of
187 psychotropics (e.g. metformin for antipsychotic-induced weight gain). We also
188 repeated our analysis for data from 2012-2015 excluding opioids not prescribed for
189 pain, medications prescribed for someone else, and non-prescribed medications
190 (including over-the-counter paracetamol/opioid compounds). In a *post hoc* sensitivity
191 analysis we tested whether our findings were accounted for by the older age of those
192 with co-morbid physical illness, and their greater prevalence of affective disorder.

193 **RESULTS**

194 **Descriptive statistics and prevalence of physical illnesses**

195 Between 1st January 2004 and 31st December 2015 inclusive, the National Confidential
196 Inquiry was notified of 57,863 suicides in England & Wales (43,539 cases with a suicide
197 verdict; 14,324 with an open verdict). Of these, 15,934 (28%) people had been in
198 contact with secondary mental health services in the 12 months before they died.
199 Questionnaires were returned on 15,662 patients, a response rate of 98%. We
200 excluded 6% (1,014 cases) with missing data for presence/absence of physical co-
201 morbidities, leaving a final dataset for analysis of 14,648 patients. Of these, 3,525

202 (24%) had a recorded diagnosis of one or more co-morbid physical illness, most
203 commonly diseases of the musculoskeletal (884, 25%); circulatory (822, 23%);
204 endocrine (646, 18%); nervous (608, 17%); and digestive systems (580, 16%). Overall,
205 66% had a condition from a single major category of physical illness, 25% from two
206 major categories, and 9% from three or more. Overlying these diagnostic categories,
207 16% (546 patients) had a pain condition and 9% had a cancer diagnosis.

208 **Patient characteristics of those with a co-morbid physical illness**

209 The median age of psychiatric patients who died by suicide and had a co-morbid
210 physical illness was 53 years (interquartile range (IQR) 43-64); significantly older than
211 those without a physical health condition (median age 44, IQR 33-54; $p < 0.001$).
212 Patients with a physical illness were more likely to be female, white, widowed, and to
213 live alone than other patients (Table 1). They were less likely to be unemployed,
214 unmarried or homeless. Whilst the proportions with a history of self-harm did not
215 differ (around 68% in both groups), those with a physical health condition less often
216 had a history of violence (19% v. 22%; $p < 0.001$) or of alcohol (42% v. 46%; $p < 0.001$) or
217 drug misuse (27% v. 35%; $p < 0.001$). Patients with a physical illness were more likely
218 than those without to have a primary psychiatric diagnosis of affective disorder, and
219 less likely to have schizophrenia (including other delusional disorders) or personality
220 disorder (Table 1). They were less likely to have been a psychiatric in-patient at the
221 time of death, to have recently (<3 months) been discharged from psychiatric in-
222 patient care, or to have been under the care of a crisis resolution/home treatment
223 team. They had more often attended their last contact with mental health services and
224 were more likely to have been adherent with medication treatment compared with
225 patients with mental illness alone. Nearly half (47%) had been in contact with services
226 in the week before death, which was significantly fewer than for patients without a
227 physical condition (51%; $p < 0.001$), with 68% exhibiting psychiatric symptoms at this
228 appointment, proportionally more than other patients (63%; $p < 0.001$). However, these
229 differences were unlikely to be clinically significant.

230

231 **Method of suicide and substances used in self-poisoning**

232 A significantly greater proportion of psychiatric patients who had been diagnosed with
233 a physical illness died by self-poisoning compared to those without physical co-
234 morbidity (37% v. 20%, $p<.001$; AOR 2.47, 95% CI 2.26-2.70; Tables 2 & 3). The
235 proportions who died by hanging/strangulation (33% v. 47%; $p<.001$),
236 jumping/multiple injuries (12% v. 16%; $p<.001$), and gas inhalation (1% v. 3%; $p<.001$)
237 (Table 2) were significantly lower in the physical co-morbidity group, although some of
238 these differences were unlikely to be clinically significant.

239 It was possible to classify the specific drugs used in cases of self-poisoning in 3,283
240 (86%) of cases; in 445 patients (12%) the data were missing and in 77 (2%) the
241 substances were described as “multiple toxicity”. More patients with a physical illness
242 were described as using multiple drugs in the overdose compared to those without a
243 physical illness (37, 3% v. 37, 2%; $p=0.02$), although this difference was unlikely to be
244 clinically significant. Opioids were the most common type of drug used in all cases of
245 self-poisoning, but particularly for those with a physical illness, nearly a third (30%) of
246 whom died by opioid overdose compared with those with mental illness alone (22%;
247 $p<0.001$) (Table 2). Patients with physical illness were also more likely to use
248 paracetamol/opioid compounds (11% v. 7%; $p<.001$) and insulin (4% v. 1%; $p<.001$)
249 and less likely to use SSRIs/SNRIs (7% v. 11%; $p<.001$) or antipsychotics (8% v. 13%;
250 $p<.001$) in self-poisoning.

251 Overall, half (586, 50%) of psychiatric patients with a co-morbid physical illness who
252 died by self-poisoning had used medications for a physical health disorder (i.e. opioids,
253 paracetamol/opioid compounds, other analgesics, insulin, cardiac medications, and
254 other specified drugs for physical conditions). This compared to a third (680, 34%) of
255 those without a physical illness ($p<.001$) (AOR 2.10, 95% CI 1.80-2.46; Table 3). The
256 majority (436; 64%) of this latter group had used opioids in overdose.

257 **Sub-group analyses**

258 **Method of obtaining medication**

259 Details of how the substances were obtained were available for 2,097 (55%) of the
260 3,805 patients who died by self-poisoning, before excluding cases without data on

261 physical illness. For the 1,306 with a physical illness who died by overdose with any
262 medication, data were available on how they obtained the drugs in 727 (56%), of
263 whom 523 (72%) were prescribed those drugs, 20 (3%) used drugs prescribed for
264 someone else, and 184 (25%) used unprescribed drugs.

265 Focussing specifically on non-psychotropics, of the 586 patients with a comorbid
266 physical illness who overdosed using a medication for a physical disorder, 246 (74%
267 when excluding unknowns) had been prescribed this medication (Table 2). This
268 compared to 102 (27%) of those without a documented physical illness who overdosed
269 using prescribed non-psychotropics ($p < .001$) (AOR 7.14, 95% CI 4.98-10.24; Table 3).
270 The main substances used in the 102 cases without documented physical illness were
271 opioids (52%), paracetamol/opioid compounds (24%), other substances, e.g.
272 propranolol (15%), and other analgesics (6%). A minority (14%) of this group had a
273 diagnosis of drug dependence/misuse, and 44% had a history of drug misuse; these
274 patients may have been prescribed opioids for drug misuse. Others may have been
275 prescribed medication for a health condition not viewed by the clinician completing
276 the questionnaire as a major physical illness.

277 A quarter of patients with comorbid physical illness who overdosed using a physical
278 health medication had not been prescribed it. A clinically significant minority had
279 overdosed on prescription-only medications not prescribed for them. Insulin had been
280 prescribed to 32 (86%) of the 37 patients with diabetes who self-poisoned using
281 insulin. Of the 12 patients diagnosed with cardiovascular conditions who self-poisoned
282 using cardiac medications, these were prescribed for 8 (67%). However, it was more
283 common for patients without a documented co-morbid physical health problem to
284 have used medications for a physical disorder prescribed for someone else (13% v. 5%;
285 $p < .001$) or obtained elsewhere (60% v. 21%; $p < .001$), presumably over-the-counter or
286 illicitly.

287 **Sub-analyses: patients with cancer, diabetes, and pain conditions**

288 When repeating the analysis for patients diagnosed with cancer compared to those
289 with other physical illnesses, there was no association of death by self-poisoning with
290 medication used for treating physical disorders (49% v. 50%; $p = .973$) (Table 3).

291 Substances used most commonly in overdose in patients with cancer were: opiates
292 (29%), paracetamol/opiate compounds (16%), and paracetamol (12%).

293 Similarly, there was no association of death by self-poisoning with substances for
294 physical disorders for patients with diabetes (54% v. 49%; $p=.203$) compared to those
295 with other physical illnesses. Substances used most commonly in overdose among
296 patients with diabetes were: insulin (21%), opiates (18%), and tricyclic antidepressants
297 (11%).

298 However, patients with a pain condition (the largest sub-group) were significantly
299 more likely to overdose with drugs for non-psychiatric conditions compared to other
300 patients with a physical condition (63% v. 46%; $p<.001$; AOR 2.12, 95% CI 1.56-2.88).
301 The majority (67%) of substances used in overdose in patients with a pain condition
302 were pain medications (opioids 46%; paracetamol/opiate compounds 12%;
303 paracetamol 6%; any other pain meds 3%), whilst 9% used tricyclic antidepressants.

304 **Sensitivity analyses**

305 The above associations remained unchanged in sensitivity analyses using a more
306 stringent definition of drugs that could have been prescribed for treating physical
307 health problems (Supplementary file). A *post hoc* sensitivity analysis to test whether
308 our findings partly reflected the older age of those with co-morbid physical illness and
309 their greater prevalence of affective illness, we found no association between older
310 age or affective disorder and self-poisoning.

311 **Discussion**

312 **Main findings**

313 We found that almost a quarter of psychiatric patients who died by suicide over the
314 period 2004 to 2015 had a co-morbid physical health condition, and that over a third
315 of this group died by self-poisoning. Our findings of an association between physical
316 health problems and fatal overdose among psychiatric patients suggest that access to
317 means is a key explanation. We found striking differences in the suicide methods used
318 by psychiatric patients with and without physical health problems. Hanging (followed
319 by overdose) was the most common method used by those with no physical co-
320 morbidities; matching the national picture for psychiatric patients (NCISH, 2017), and

321 the general population (ONS, 2016c). However, self-poisoning (followed by hanging)
322 was the leading method used by patients with physical health problems, suggesting
323 that overdose is the most accessible approach for this patient group if contemplating
324 suicide. Restricting access to this method is more feasible than for hanging.

325 The substances used in overdose by patients with a co-morbid physical health
326 condition were more likely to be medications prescribed to treat physical health
327 problems, and less likely to be psychotropics, even though these patients probably had
328 access to both. Nearly half of those with a co-morbid physical health condition who
329 died by self-poisoning did so using a medication for such a condition. Of specific sub-
330 groups, patients with pain conditions, for whom chronic pain is itself a risk factor for
331 suicide (Racine, 2018) were most likely to overdose with drugs for physical disorders.
332 This was likely due to a high proportion of this group using toxic pain medications in
333 overdose. The tendency of patients with physical co-morbidities to overdose using
334 non-psychotropics rather than psychotropics may relate to perceived lethality of non-
335 psychotropics, potentially greater lethality of non-psychotropics, or to prescribers
336 being more primed to consider overdose potential when issuing and monitoring
337 potentially cardiotoxic psychotropic drugs (Hawton et al., 2010) than medications used
338 for physical health problems. Whilst acknowledging the poor predictive value of
339 suicide risk classification scales (Steege et al., 2018), our findings suggest that needs-
340 based assessments of psychiatric patients with physical health problems should focus
341 on addressing modifiable risk factors such as reviewing the need for more toxic
342 medications, particularly opioids (Ilgen et al., 2016), considering safer transdermal
343 routes for opioid administration (Coplan et al., 2017), and addressing inadequately-
344 treated pain (Yarborough et al., 2016). Guidelines on safe prescribing aim not to
345 compromise on optimal pain management, but to reduce the potential for opioid
346 addiction, diversion and fatalities (Volkow et al., 2019).

347 **Findings in the context of other studies**

348 No other studies have sought to investigate this research question among psychiatric
349 patients. More widely, a systematic review of studies investigating the association
350 between non-psychotropic medications and attempted suicide found cardiovascular
351 medications not to be associated with any increased risk, but concluded that

352 associations with other medications remained inconclusive (Gorton et al.,
353 2016). Separately, two studies of US veterans with non-cancer pain found an
354 association between dose of opioids and risk of suicide (Ilgen et al., 2016), presumably
355 with dose a marker of pain severity, but no clear excess risk of overdose in these
356 patients over other methods (Ilgen et al., 2013).

357 **Strengths and limitations**

358 We examined a national, comprehensive case series of all suicides amongst patients
359 with recent contact with psychiatric services over a 12 year period. Consultants
360 completing the questionnaire were unaware of the study's hypotheses, so it was
361 unlikely that clinicians' recall bias for overdose using physical health medications might
362 explain our findings. Our categorisation of physical illnesses was systems-based but
363 also acknowledged the overlapping categories of cancer and pain conditions. We
364 adjusted our models for variables identified as potential confounders *a priori*, such as
365 drug dependence/misuse. Alternative explanations for associations identified are the
366 under-identification of drug dependence/misuse, and the assumption that opioids
367 used in overdose were obtained for a physical health problem rather than for abuse or
368 intentional overdose. We had access to data on how medications were obtained for
369 only 55% of the case series, but findings were similar in a sensitivity analysis confined
370 to those who died from 2012-2015.

371 The study's main limitation is that its use of survey data captured only those co-
372 morbid physical health problems and overdose medications of which the responding
373 consultant was aware. Under reporting of physical health problems is likely to have
374 occurred where the patient was only briefly under their care (particularly in liaison
375 settings), where clinical notes were unclear regarding physical health conditions or
376 medications, or where the clinician did not judge the condition to be a 'major physical
377 illness'. This may have excluded conditions like acne that contribute significant clinical
378 distress and for which medications prescribed to treat it have been linked with suicide
379 risk (Sundström et al., 2010). Under reporting of specific physical health medications
380 used in overdose is likely to have occurred where the completing clinician's response
381 denoted multiple unspecified drugs. Over half of all general population drug poisoning
382 deaths involve more than one drug and/or alcohol and the substance primarily

383 responsible for the death is not identifiable (ONS, 2016b). We could also not be certain
384 that medications used in overdose had been specifically issued to treat that patient's
385 physical health problem, as opposed to being obtained specifically to attempt suicide.
386 We did not have data specifying whether onset of physical illness had preceded
387 psychiatric illness or vice versa, and it was possible in some cases that patients had
388 been diagnosed with a physical health problem some time before their psychiatric
389 illness commenced. This preceding physical illness may have also influenced some
390 patients in their choice of self-poisoning agent.

391 Detailed data on how opioids and paracetamol/opioid compounds were obtained
392 were only available from 2012 onwards, but we addressed this in our sensitivity
393 analysis. This extra analysis also ruled out the older age of those with co-morbid
394 physical illness, and their greater prevalence of affective illness, as an explanation for
395 our findings. Finally, by examining a national case series design, without living controls,
396 we could estimate proportional contrasts between the groups but not incidence, or
397 absolute/relative risks.

398 **Clinical and policy implications**

399 These findings provide evidence to suggest that access to means of lethal overdose
400 may contribute to suicide risk in psychiatric patients with physical co-morbidities,
401 particularly those with chronic pain. Such patients would be more likely than other
402 psychiatric patients to have supplies of prescribed non-psychotropics at home,
403 particularly patients in chronic pain. Such availability creates the potential for suicide
404 attempts with high lethality, particularly during a flare-up of a physical condition. All
405 clinicians involved in the care of these patients should ensure careful prescribing for
406 this patient group, with clear risk management. This could include regular reviews to
407 check that indications remain, referral to pain clinics to consider transdermal opioid
408 administration, and raised frequency of issuing pain medication prescriptions,
409 although the latter may compromise patient convenience and therapeutic alliance.
410 Assertive pain management is critical because inadequately-treated pain is itself a risk
411 factor for suicide (Yarborough et al., 2016). Future research should seek to evaluate
412 the effect of improved pain management pathways and prescribing guidelines on risk
413 of overdose among psychiatric patients.

414 Restricting access to non-prescribed medications has been partly addressed at the
415 population level (Hawton et al., 2013; Hawton et al., 2009) with a restriction on
416 analgesic pack size, but there is also a role for community pharmacists in responding
417 to customers trying to purchase over-the-counter analgesics above recommended
418 limits (MHRA, 2014). A non-confrontational approach that responds to distress, and
419 shows awareness of local service provision is more likely to be acceptable to patients.
420 Our findings also suggest that access to medications prescribed for household
421 members should be considered for psychiatric patients with or without physical illness.
422 Carers have a role in safeguarding their own medications, as well as those of a
423 psychiatric patient at risk.

424 Finally, our findings show that opioids are a substance commonly used in lethal
425 overdose among psychiatric patients, whether they have physical health problems
426 (30%) or not (22%). Access to naloxone for carers and professionals, accompanied by
427 training, is a high-risk intervention worth considering among some psychiatric patients
428 (Ashrafioun et al., 2016). Qualitative work is needed with carers regarding their
429 attitudes towards such a safeguarding role.

430 **Conclusions**

431 Overdose, rather than hanging, is the leading method of suicide in the 24% of
432 psychiatric patients who die by suicide and have co-morbid physical health problems;
433 accounting for over a third of cases. In such patients, particularly for those in chronic
434 pain, the medications used in overdose are more likely to be those for a physical
435 health disorder; primarily opioids. Psychiatric patients with physical health co-
436 morbidities therefore require careful needs-based risk assessment, with clinicians
437 reducing access to the means of overdose where possible. Optimal care includes
438 addressing inadequately-treated pain, reviewing the need for more toxic medications,
439 considering transdermal routes, and involving carers in safeguarding household
440 medications.

441

442

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453

454 **Declaration of interest:**

455 LA and NK are members of the Department of Health's (England) National Suicide
456 Prevention Advisory Group, of which LA is Chair, for which he has received personal fees. LA
457 is a board member of the CQC, for which he has received personal fees. NK is Chair of the
458 Guideline Development Group for the NICE guideline on depression in adults and is a topic
459 expert for the NICE suicide prevention guideline. All authors declare that there are no other
460 conflicts of interest.

461 **Data availability:**

462 The National Confidential Inquiry case series database is not publically available, but
463 requests to conduct analyses in collaboration with the Centre for Mental Health and Safety
464 team are granted, subject to internal peer review.

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