1	Causes of death up to ten years after hospitalisation for self-
2	inflicted, drug/alcohol-related, or violent injury during
3	adolescence: a nationwide cohort study
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21	Tables: 3
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24	Supplementary figures: 1
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- 26 Summary
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**Background:** Emergency hospital admission with adversity-related injury (self-inflicted, drug/alcohol-related, violent) affects 4% of 10-19 year olds. Their risk of death in the decade after discharge is twice as high compared to adolescents hospitalised for accidentrelated injury. We determined how cause of death varied between these groups.

Methods: We compared risks of death in five causal groups (suicide, drug/alcohol-related, homicide, accidental, 'other') up to ten years after discharge following adversity-related or accident-related injury. We used linked hospital admission (to the National Health Service) and mortality data for England (1997-2012) to determine cause-specific risks of death for 10-19 year olds, and to compare risks between adversity- and accident-related index injury after adjustment for age-group, socio-economic status, and chronic conditions.

38 Findings: Among 333,009 adolescents admitted with adversity-related injury (girls 39 181,926, boys 181,053), and 649,818 with accident-related injury (girls 166,462, boys 40 483,356), 4,782 died in the ten years post-discharge (girls 1,312, boys 3,470). Adolescents 41 discharged after adversity-related injury had higher risks of suicide and of drug/alcohol-42 related death in the next decade than after accident-related injury (adjusted hazard ratios 43 [aHRs] varied from 3.2 [95% CI: 2.7, 3.6] for suicide in boys to 4.7 [3.3, 6.8] for 44 drug/alcohol-related death in girls). Risks of suicide were increased following self-inflicted 45 injury, drug/alcohol related injury, and violent injury (e.g. boys, aHR: 6.2 [5.3, 7.3], 4.5 46 [3.9, 5.2], 1.4 [1.2, 1.8], respectively vs. accident-related injury). Following each type of 47 index injury, risks of suicide and risks of drug/alcohol-related death were increased by 48 similar magnitudes (e.g. boys with self-inflicted injury vs. accident-related injury, aHR of 49 suicide: 6.2 [5.3, 7.3], drug/alcohol-related injury death: 5.9 [5.0, 7.0]).

50 **Interpretation:** Risks of suicide increased after all types of adversity-related injury, as did 51 risks of drug/alcohol-related death by a similar magnitude. Current practice to reduce risks 52 of harm after self-inflicted injury should be extended to drug/alcohol-related and violent 53 injury in adolescence. Prevention should address the substantial risks of drug/alcohol-54 related death alongside risks of suicide.

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## 57 Introduction

58 Evidence from population-based cohort studies suggests that different types of 59 'adversity-related injury' (self-inflicted [including poisonings], drug/alcohol-related, or 60 violent injury) during adolescence are associated with similar underlying psychosocial problems, including adverse experiences (e.g. maltreatment), poor mental health (e.g. 61 anxiety, depression), and poor social circumstances (e.g. poverty).<sup>1-3</sup> Among the 4% of 62 63 adolescents (10-19 year olds) who are admitted to hospital with one of these types of 64 adversity-related injury in England, approximately three-quarters of girls and one-third of boys are admitted with injuries related to multiple types of adversity.<sup>4</sup> Despite this 65 66 apparent overlap between self-inflicted, drug/alcohol-related, and violent injury, most research in these adolescents has focused on specific types of adversity-related injury. 67 68 A previous study of adolescents admitted to hospital in England as an emergency with any adversity-related injury reported that 1 in 136 girls (7.3 per 1,000) and 1 in 64 boys 69 70 (15.6 per 1,000) died within the ten years after discharge, and that these risks were 71 similar whether the initial injury was self-inflicted, drug/alcohol-related, or violent.<sup>5</sup> 72 These ten-year risks were approximately twice the risks for adolescents discharged after 73 accident-related injury (girls 3.8 per 1,000 and boys: 6.0 per 1,000) or for the general 74 population of adolescents (girls 3.0 and boys: 3.0).

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76 Despite common underlying psychosocial problems and elevated mortality risks among 77 adolescents with any of these three types of adversity-related injury, UK national 78 clinical guidelines recommend different approaches to psychosocial assessment and intervention to reduce future harm.<sup>6-8</sup> For example, guidelines for managing self-79 80 inflicted injury presenting to hospital recommend admission of patients younger than 16 years and assessment of psychosocial circumstances and suicide risk at all ages.<sup>6,7</sup> 81 82 Guidelines for drug- or alcohol- related presentations do not specifically address psychosocial needs of adolescents.<sup>8</sup> No UK guidelines exist for responding to violent 83 84 injury. A further issue is that clinical management to reduce the risk of further harm 85 after self-inflicted injury focuses on risks of recurrent self-harm, despite evidence for increased risks of other adverse outcomes.<sup>9</sup> A cohort study of 15-24 year olds 86 presenting to a hospital in Oxford with self-inflicted injury in 1978-1997 reported 87 88 increased mortality due to respiratory disorders, circulatory disorders, and accidents, as well as suicide, during the subsequent 20 years.<sup>9</sup> No comparable estimates have been
published for risks of harm following drug/alcohol-related or violent injury (see panel
'Research in Context').

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93 This study aims to inform preventive strategies for reducing risks of future harm for 94 adolescents who are discharged from hospital after self-inflicted, drug/alcohol-related, 95 or violent injury. Given standard practice to reduce risks of repeated self-harm or 96 suicide after discharge following self-inflicted injury, we examined, for girls and boys 97 separately, whether risks of suicide difference between adolescents discharged 98 following drug/alcohol-related and violent injury. Second, among girls and boys 99 respectively, we compared risks of cause-specific death (suicide, drug/alcohol-related, 100 homicide, accidental, and other) up to ten years from discharge after each type of index 101 injury, including accident-related injury.

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#### 104 Methods

### 105 Study design

We used Hospital Episode Statistics (HES) data, which contain all emergency (acute, unplanned) admissions to the National Health Service (NHS) in England (April 1997-March 2012), including to independent sector providers paid for by the NHS.<sup>10</sup> Approximately 98-99% of hospital activity in England is funded by the NHS,<sup>11</sup> and so these data captured nearly all admitted adolescents. As we used a standard, de-identified HES extract from NHS Digital (formerly known as the Health and Social Care Information Centre), ethics approval was not required.<sup>12</sup>

We derived a cohort of adolescents (aged 10-19 years inclusive) who were admitted for injury (the index injury), and categorised them as 'adversity-related injury' (comprising non-mutually exclusive groups of self-inflicted, drug/alcohol-related, or violent injury; irrespective of whether the injury was also accident-related) or 'accident-related injury' (where there was no recorded adversity-related injury). Therefore, adversity-related injury and accident-related injury were two mutually exclusive groups. Deaths within the cohort were evaluated in five 'causal' groups: suicide, drug/alcohol-related, homicide, accidental, or 'other'. We compared risks of death (total and by cause) up to
ten years following discharge from admission for adversity-related injury (exposure)
with risks after accident-related injury (comparator).

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We excluded adolescents who did not have sex recorded (885; 0.08%), died during the index admission (1,877; 0.17%), had no valid discharge date (372; 0.03%), or were admitted with injury related to neither adversity nor accidents (94,407; 8.9%; the majority of these latter adolescents were admitted primarily for chronic conditions or complications of surgery).<sup>4</sup>

## 130 Study cohort and exposures

131 Self-inflicted, drug/alcohol-related, violent, and accident-related injuries were 132 identified using ICD-10 codes in HES data (i.e. characteristics that were identified and 133 recorded by clinicians). Details of classification of injury and descriptive statistics of the cohort have been reported elsewhere.<sup>4,5</sup> Briefly, 333,009 adolescents who had at 134 135 least one adversity-related injury (181,926 girls, 151,083 boys; 20.3% and 24.0% of 136 which had an injury that was also accident-related), and 649,818 with at least one accident-related injury but no adversity-related injury (166,462 girls, 483,356 boys) 137 were identified (Supplementary Figure S1).<sup>5</sup> 138

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140 Table 1 shows that the most frequent type of injury among girls and boys was 141 drug/alcohol-related followed by self-inflicted injury in girls and violent injury in boys. 142 We have previously reported that the peak age group for adversity-related injury was 143 15-17 years old for girls (47%) and 18-19 years old for boys (62.%), but for accident-144 related injury it was 10-14 years for girls (54%) and boys (62%). Compared with adolescents admitted with accident-related injury, those admitted for adversity-related 145 146 injury were more likely to be in the in the most deprived category, or to have a chronic condition (Herbert et al 2015, Table 1).<sup>5</sup> 147

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#### 149 **Outcomes**

150 The primary outcome was cause-specific death between one day and ten years after 151 discharge from the index injury admission. We identified deaths using Office for 152 National Statistics (ONS) mortality data linked to HES (within NHS Digital). We used any ICD-9 or -10 codes in the mortality data (based on the underlying and up to 15 153 154 other contributing causes recorded in the death certificate) to categorise deaths into five 155 'causal' groups (Supplementary Table S1). As Figure 1 illustrates, suicide, 156 drug/alcohol-related, and homicide were not mutually exclusive, but these three groups 157 (i.e. adversity-related deaths), accidental (no codes for adversity-related death, but 158 codes for accidental causes) and 'other' deaths (no codes for adversity-related or 159 accidental deaths) were mutually exclusive. As advised by the ONS, undetermined causes of death (codes E980-E989, Y1-Y34; n=483) were classified as suicide 160 (accounting for 38.1% of all suicides).<sup>13</sup> Deaths with codes indicating an adjourned 161 inquest (U50.9; n=130) were categorised as homicide (80.2% of all homicides). 162

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164 Demographic and clinical factors

Covariates were included in the analyses, based on previous findings of their 165 166 relationship with adversity-related injury and death, including sex, age, socio-economic status (SES), and chronic conditions.<sup>5</sup> Age was grouped (10-15, 16-17, 18-19 years) to 167 reflect different recommendations in UK national guidelines for management of self-168 harm or alcohol misuse according to age, and different stages of social development.<sup>6-</sup> 169 <sup>8</sup> SES was categorised according to Index of Multiple Deprivation scores based on 170 residential postcode,<sup>14</sup> using quintile cut-off values for England. An adolescent was 171 172 classified as having an underlying chronic condition if HES records for the index injury 173 admission or any admissions in the previous year included one of a cluster of ICD-10 codes for chronic conditions (Hardelid *et al*, 2013; Appendix Table  $6 \cdot 3 \cdot 2$ ).<sup>15</sup> Of the 174 117,453 adolescents with adversity-related or accident-related injury who had a chronic 175 176 condition, 93,592 (79.7%) had a physical condition (data not shown). The most 177 common physical condition was chronic respiratory disorder (e.g. asthma, 39.8% to 55.4% by sex and type of injury).<sup>5</sup> 178

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### 180 Statistical analyses

181 All analyses were conducted in Stata/SE 12 (StataCorp), and separately for girls and182 boys.

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We first derived numbers (and proportions) of deaths (total and by cause) in the ten years post-discharge after adversity-related (self-inflicted, drug/alcohol-related, or violent) or accident-related index injury. As statistical disclosure rules required us not to publish counts <10 we did not present exact numbers of homicides for certain groups.

189 We determined unadjusted cumulative risks and 95% confidence intervals (CIs) of 190 deaths for each cause of death over the ten years following discharge from the index 191 injury admission. The cumulative risk of death by cause of death was estimated as a 192 cumulative incidence function, which accounted for other 'competing' causes (e.g. for 193 suicide, competing causes included homicide, drug/alcohol-related, accidental and 194 other).<sup>16</sup> For reference, we present unadjusted ten-year cumulative risks and 95% CIs by cause of death and type of index injury, sex, and age-group (Supplementary Table 195 196 S2). We also estimated total and cause-specific risks of death for the general population 197 of 10-19 year olds in England in 1997-2012, using publicly available ONS life-tables 198 for total mortality and suicide, and bespoke life-tables for drug/alcohol-related and 199 accidental deaths provided to us by the ONS (according to ICD codes in Supplementary 200 Table S1).17 -19

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We fitted Fine & Gray models<sup>16</sup> to estimate the relative risks of total and cause-specific 202 mortality following adversity-related index injury, adjusted for covariates and taking 203 204 into account competing risks of other causal groups. The exposure was type of index 205 injury, and covariates included age-group, SES, and chronic condition status. 'Sub-206 hazard ratios' (SHRs) of each cause of death were estimated for adversity-related injury 207 (vs. accident-related injury), age-groups 16-17 and 18-19 years (vs. 10-15 years), each 208 level of SES (vs. least deprived), and chronic condition (vs. none). To compare risks 209 following each type of adversity-related injury, we fitted the models as above but where 210 the exposure was self-inflicted, drug/alcohol-related, and violent injury, respectively 211 (each vs. accident-related injury).

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Finally, we assessed whether the finding that increased risks of suicide and drug/alcohol-related deaths following self-inflicted or drug/alcohol-related injury was due to the 'overlap' between these two types of index injury (73% of girls and 44% of boys with either type had both types),<sup>5</sup> or the overlap between suicide and drug/alcoholrelated deaths (~12% of deaths that were either suicide or drug/alcohol-related, were both [Figure 2]). We fitted the Fine & Gray models as above, but where the exposure was the three different combinations of self-inflicted and drug/alcohol-related injury (vs. accident-related injury), and the outcome was suicide, drug/alcohol-related death, and each combination of these types of death, respectively (further details within footnotes of Supplementary Table S3).

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We checked model assumptions using log-log plots of the Kaplan-Meier estimate of the survival function and the link test, and assessed their goodness-of-fit using plots of the Nelson-Aalen estimate of the cumulative hazard function against Cox-Snell residuals.<sup>16</sup>

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### 230 **Results**

231 By ten years after discharge from admission for the index injury, there were 2,415 232 deaths (girls 873, boys 1,542) after adversity-related injury and 2,367 deaths (girls 439, 233 boys 1,928) after accident-related injury (Figure 1, Table 1). After adversity-related 234 index injury, nearly two-thirds (63.9%, n=1,046) of the deaths were related to suicide, 235 drug/alcohol use, or homicide, compared with only one-third (33.6%, n=796) after 236 accident-related index injury (Figure 1, Table 1). The proportions of deaths related to 237 suicide, drug/alcohol use, or homicide were similar between girls and boys after 238 adversity-related injury (girls 59.3% [n=518], boys 66.5% [n=1,025]), but lower for 239 girls than boys after accident-related injury (girls 19.4% [n=85], boys 36.9% [n=711]) 240 (Table 1). The most frequent causes of death after accident-related index injury were 241 'other' (overall 37.1% [n=877]; girls 59.2% [n=260], boys 32.0% [n=617]) and 242 accidental (29.3% [n=694]; girls 21.4% [n=94], boys 31.1% [n=600]) (Figure 1, Table 243 1).

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Two thirds of all accidental deaths, 67.8% (n=759) were recorded as transport accidents; this proportion did not differ according to type of index admission (data not shown). Among deaths due to other causes, the most common causes were related to

- 248 neurological conditions (30.9%, n=473) or cancer/blood disorders (25.1%, n=384; of
- 249 nine possible groups of ICD codes relating to systems within the body).<sup>15</sup>
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# 251 Risks of total and cause-specific deaths by type of index injury

### 252 Adversity-related vs. accident-related index injury

- Ten-year cumulative risks of total death after adversity related index injury were 7.3 per 1,000 (or 1 per 137) girls (95% CI: 6.8 to 7.8 per 1,000) and 15.6 per 1,000 (or 1 per 64) boys (14.8 to 16.4 per 1,000) (Figure 2, Supplementary Table S2). Cumulative risks were lower after accident-related index injury (girls 3.7 per 1,000, 3.4 to 4.1; boys 6.0, 5.7 to 6.3).
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The increased risks of death after an adversity-related compared with accident-related injury were due to substantially higher risks of suicides and drug/alcohol-related deaths at all time-points after the index injury (Figure 2). After adjustment for other covariates, risks of suicides and drug/alcohol-related deaths were three to five times higher following discharge from adversity-related injury admission (Table 2).

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## 265 Self-inflicted, drug/alcohol-related, and violent index injury

266 Ten-year risks of suicide were similar after hospital discharge following self-inflicted 267 index injury and drug/alcohol-related index injury (girls 2.9 vs. 2.5 per 1,000; boys 9.8 vs. 7.2; Figure 3, Supplementary Table S2). Compared with adolescents discharged 268 269 after accident-related injury, risks of suicide were increased five- to six-fold for 270 adolescents discharged after self-inflicted or drug/alcohol-related injury (Table 3 shows 271 sub-hazard ratios adjusted for covariates; e.g. for boys the adjusted SHR of suicide after 272 self-inflicted injury was 6.20 [5.27, 7.30] and after drug/alcohol-related injury 4.51 273 [3.89, 5.24]). Risks of suicide were increased after self-inflicted and after drug/alcohol-274 related injury, whether the index injury was for either one of these types of injury only, 275 or both (Supplementary Table S3; i.e. comparing between rows, per sex).

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277 Ten-year risks of suicide and of drug/alcohol-related death were similar after each type278 of index injury. These risks were highest after self-inflicted or drug/alcohol-related

279 index injury (Figure 3; Supplementary Table S2). For example, after self-inflicted injury, the ten-year risk of suicide for girls was 2.9 per 1,000, whereas the ten-year risk 280 281 of drug/alcohol-related death was 2.7 per 1,000 (Figure 3, Supplementary Table S2). 282 After adjustment for covariates, the increased risks of suicide after self-inflicted and 283 after drug/alcohol-related index injury (vs. accident-related injury) were similar to the 284 risks of drug/alcohol-related death. For example, among boys discharged after self-285 inflicted injury compared with after accident-related injury, the adjusted SHR was 6.20 286 [5.27, 7.30] for suicide and 5.91 [4.96, 7.03], for drug/alcohol-related death) (Table 287 3). These adjusted SHRs were similar whether the death was related to suicide but not 288 drugs/alcohol, drugs/alcohol but not suicide, or both causes (Supplementary Table S3; 289 i.e. comparing between columns).

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## 291 Socio-demographic and clinical covariates

Boys aged 18-19 years who were discharged after self-inflicted injury or drug/alcoholrelated injury had the highest risks of death due to any cause (ten-year risks: 30.4 per 1,000, or 1 per 33, after self-inflicted injury, 25.1 per 1,000, or 1 per 40, after drug/alcohol related injury; Supplementary Table S2). These risks were substantially higher than after accident-related injury (8.8 per 1,000) or for the general population of 18-19 year old boys (8.9 per 1,000). These risks were driven by high risks of suicide and drug/alcohol-related death.

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300 Adolescents aged 18-19 years had twice the mortality risk compared with 10-15 year

301 olds, due to increased risks of suicide and drug/alcohol-related deaths among older

302 girls and boys, and increased risks of accidental deaths among older boys (Table 2;

303 Table 3). Low SES (i.e. most deprived) was associated with increased risks of total

and cause-specific mortality, apart from suicide in boys, in whom low SES was

305 associated with a decreased risk of suicide.

Adolescents with a chronic condition (vs. none) had a 3- to 4-fold increased risk of death due to any cause, and a 10- to 12-fold increased risk of death due to causes other than adversity or accidents, regardless of the type of index injury (Table 2). For example, for 18-19 year old boys discharged after an adversity-related index injury, the 310 ten-year risk of death due to any cause was 37.5 per 1,000 given a chronic condition

- 311 and 14.8 per 1,000 given none (data not shown). For 18-19 year old boys discharged
- after accident-related injury, these risks were 17.5 and 8.8 per 1,000 respectively.
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# 314 **Discussion**

315 This retrospective cohort study determined cause-specific risks of death up to ten years 316 after adolescents were discharged from the NHS in England following injury related to 'adversity' (self-harm, drug/alcohol misuse, violence) or accidents. Within ten years 317 318 after discharge following adversity-related injury 1 per 137 girls and 1 per 63 boys had 319 died. We found that suicide, drug/alcohol-related deaths, and a small number of 320 homicides accounted for 61% of all deaths ten years after adversity-related injury, but 321 only 35% of deaths after accident-related injury. Second, we showed that risks of 322 suicide were all increased following self-inflicted injury, drug/alcohol-related injury, 323 and following violent injury. These risks were highest for 18-19 year old boys. Third, 324 the risks of suicide were similar to those of drug/alcohol-related deaths regardless of 325 whether the adversity-related index injury was self-inflicted, drug/alcohol-related, or 326 violent. Fourth, adolescents with an underlying chronic condition at the index injury admission (10-15%)<sup>5</sup> were at increased risk of all causes of death, independently of the 327 328 type of adversity or accident-related injury or age at admission.

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## **330** Strengths and limitations

331 The main strength of our study is the use of linked NHS emergency admissions and 332 mortality data, which included all injury admissions in England linked to subsequent mortality reacords in England and Wales over 15 years.<sup>13</sup> The population-based cohort 333 334 of nearly one million 10-19 year olds allowed us to compare risks of cause-specific 335 mortality between different types of index injury admissions. We used time-to-event 336 statistical methods to estimate risks whilst taking into account censoring of outcomes and competing risks of different causes of death.<sup>16</sup> Although we combined index injury 337 admissions across a 15-year period, our conclusions were not sensitive to calendar 338 339 period (e.g. boys in 1997, adjusted SHR of suicide for adversity-related vs. accidentrelated injury [95% CI]: 2.6 [1.7, 3.9]; corresponding SHR for boys in 2012: 3.2 [2.2, 340 341 4.7]; data not shown).

343 One limitation is that ICD codes used to define adversity-related injury and deaths tend to have high specificity but low sensitivity.<sup>20-22</sup> The potential misclassification of 344 exposure (i.e. self-inflicted, drug/alcohol-related, or violent injury, misclassified as 345 346 accident-related injury) and outcomes (i.e. suicide, drug/alcohol-related deaths, or 347 homicides, misclassified as accidental or other deaths) may induce bias in the estimates 348 of their associations, which is likely to under-estimate the increased risks of suicide and 349 drug/alcohol-related deaths after adversity-related injury relative to after accident-350 related injury. To minimise this potential bias we included codes for undetermined 351 intent and adjourned inquests in the definitions of suicide and homicide, respectively. 352 The prevalence of chronic conditions recorded by codes at the index injury admission 353 or at hospitalisation during the previous year may be under-ascertained, particularly to 354 the presence of chronic mental health conditions.

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A further limitation is potential linkage error between HES and ONS mortality data. One of the few studies that have investigated linkage errors in HES data showed high missed match rates (4.1%) that were higher for males and ethnic minorities.<sup>22</sup> Linkage error between HES and ONS mortality data would favour underestimation of mortality rates.<sup>22</sup> Lastly, the study was likely under-powered to detect differences in the risks of homicide between index injury groups.

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## 363 **Comparison with other studies**

364 Our main finding of similarly increased risks of suicide death following self-inflicted 365 injury and following drug/alcohol related injury has not been previously reported. We 366 report lower ten-year risks of death after admission with self-inflicted injury (girls: 7.7 367 per 1,000, boys: 24.1 per 1,000; Supplementary Table S2) than the 20-year mortality rates after presentation with self-inflicted injury reported by Hawton et al (girls: 17 per 368 1,000, boys: 50 per 1,000).<sup>9</sup> These differences may be explained by different lengths of 369 370 follow-up between the two studies, and different age-ranges for exposure (current 371 study: 10-19y vs. Hawton et al: 15-25y) and for deaths (10-29y vs. 15-44y). In Hawton et al's study, 60.0% of deaths in girls and 45.6% of those in boys were from suicide 372 (including deaths of undetermined intent and drug/alcohol-related suicides),<sup>9</sup> compared 373 374 with 39.8% and 43.2% in our study (Table 1).

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#### 375

#### 376 Implications for practice, policy, and research

377 Our findings suggest that specialist psychosocial assessment by a child and adolescent 378 mental health professional, which is part of recommended standard practice for self-379 inflicted injury in the UK, should be considered for adolescents presenting with 380 drug/alcohol-related or violent injury. The need for a consistent approach targeting all 381 three adversity-related injury groups is supported by previous evidence of their 382 common underlying psychosocial problems, the overlap among the same admitted adolescents,<sup>4</sup> and the inter-relationship between related behaviours into young 383 adulthood, particularly self-harm and drug/alcohol use.<sup>24-26</sup> Clinical and public health 384 strategies need to be extended to include reducing risks of death related to 385 386 drugs/alcohol, which are just as high as risks of suicide death. If it were possible to 387 completely eradicate the excess mortality risk associated with adversity-related injury 388 among hospitalised adolescents, we could have expected 857 fewer suicide and 389 drug/alcohol-related deaths in our cohort (girls: 392 [219 drug/alcohol-related deaths], 390 boys: 683 [394]; based on the estimated relative risks in Table 2). Among 16-19 year 391 olds, the burden of suicides in the decade after adversity-related injury represented 392 approximately 10-25% of suicides expected in the general population during the same follow-up (based on ten-year risks in Supplementary Table S2, and ~3-4% of the 393 general population of 16-19 year olds being admitted with adversity-related injury).<sup>4</sup> 394

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Findings from the current study may be generalised to other UK countries that have similar rates of hospitalisations during adolescence for adversity-related injury,<sup>15</sup> and similar rates of mortality through intentional injuries.<sup>29</sup>

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400 There needs to be more investment in interventions for reducing harm after all types

401 of adversity-related injury, whether self-inflicted, drug/alcohol-related, or violent.

402 Risks of deaths through causes both related to mental health (suicide, drugs/alcohol)

403 and potentially not related to mental health (accidents, other) are substantially

- 404 increased in adolescents admitted with chronic conditions, and mechanisms of
- 405 effective interventions may differ for this sub-group. The evidence base for how
- 406 public health bodies and health services should respond to the common manifestation
- 407 of distress in vulnerable adolescents of adversity-related injury is weak,<sup>28-30</sup> and there

- 408 is a need for the development of potentially effective interventions and then
- 409 evaluation through large trials to determine what works and for whom.

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- 415

# 416 **Contributors**

AH, RG, and LL conceived and designed the study. AH analysed the data and drafted
the first version of the article. AH, RG, DC, and LL interpreted the data, revised the
article critically for important intellectual content, and approved the final version to be
published.

421

# 422 **Conflicts of interest**

423 None to declare.

424

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# 435 Data sources

436 Hospital Episode Statistics data can be accessed by researchers applying to NHS

437 Digital (previously the Health and Social Care Information Centre for England).

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- 439 Bespoke extracts and tabulations of mortality data for England and Wales are
- 440 available to order from the ONS (subject to legal frameworks, disclosure control,
- 441 resources and agreement of costs, where appropriate). Such enquiries should be made
- to the mortality team at <u>mortality@ons.gsi.gov.uk</u>.

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#### 529 Figure 1: Numbers and proportions of deaths by reported cause 530 531 Circles represent proportions and are drawn to scale within each figure (i.e. type of injury). Accidental death: codes for accidents and no codes for adversity in death certificate; Other death: no codes for accidents or adversity in death certificate. 532 533 Figure 2: Cumulative risk of cause-specific death over time, by sex and adversity-related or accident-related index injury at admission 534 535 Drug/alc = Drug/alcohol-related; Acc = Accidental; Oth = Other; 'Suicide' includes all suicides, whether homicide or drug/alcohol-related death were also implicated or not; Drug/alc death includes only drug/alcoholrelated deaths where suicide was not also implicated; 'Homicide' includes only where suicide or drug/alcohol-related death was not also implicated. Here cumulative risks are cumulative incidence functions. 536 537 Figure 3: Ten-year cumulative risk of cause-specific deaths, by sex and type of index injury 538 Drug/alc = Drug/alcohol-related; Acc = Accidental; Oth = Other; 539 540 **Table 1:** Number and proportion of cause-specific deaths within ten years after index injury admission, by sex and type of index injury

Type of injury at index admission	Discharged	Total deaths	Adversity-related*	Suicide	DA	Accidental	Other
Girls	348 388	1 312 (100·0)	603 (46·0)	361 (27·5)	319 (24·3)	228 (17·4)	481 (36·7)
Accident-related	166 462	439 (100·0)	85 (19·4)	47 (10·7)	41 (9·3)	94 (21·4)	260 (59·2)
Adversity-related	181 926	873 (100·0)	518 (59·3)	314 (36·0)	278 (31·8)	134 (15·3)	221 (25·3)
Self-inflicted	131 739	651 (100·0)	408 (62·7)	259 (39·8)	210 (32·3)	93 (14·3)	150 (23·0)
DA	163 888	776 (100·0)	464 (59·8)	283 (36·5)	250 (32·2)	117 (15·1)	195 (25·1)
Violent	13 262	54 (100·0)	25 (46·3)	10 (18·5)	16 (29·6)	11 (20·4)	18 (33·3)
Boys	634 439	3 470 (100.0)	1 736 (50·0)	903 (26·0)	861 (24·8)	891 (25·7)	843 (24·3)
Accident-related	483 356	1 928 (100·0)	711 (36·9)	375 (19·5)	311 (16·1)	600 (31·1)	617 (32·0)
Adversity-related	151 083	1 542 (100·0)	1 025 (66·5)	528 (34·2)	550 (35·7)	291 (18·9)	226 (14·7)
Self-inflicted	44 621	704 (100·0)	526 (74·7)	304 (43·2)	276 (39·2)	92 (13·1)	86 (12·2)
DA	85 421	1 112 (100·0)	775 (69·5)	418 (37·6)	424 (38·1)	183 (16·5)	154 (13·8)
Violent	70 594	460 (100·0)	268 (58·0)	120 (26·1)	135 (29·2)	122 (26·4)	70 (15·2)

#### Numbers of deaths by cause (row %)

541 DA = Drug/alcohol-related \*Suicides, drug/alcohol-related

42 \*Suicides, drug/alcohol-related deaths, and homicides. These deaths were not mutually exclusive. Numbers and proportions are not reported for homicides due to small counts.

# **Table 2:** Relative risk of cause-specific death within 10 years after adversity-related index injury (vs. accident-related injury), adjusted for age-group, socio-

544 economic status, and chronic condition status, by sex (multivariable analyses)

	Cause of death, Adjusted sub-hazard ratio (95% Confidence Interval)										
Characteristic at index injury admission	All deaths			Suicide		DA death		Accidental death		Other death	
irls											
Adversity- (vs. accident-related) injury	1.51	(1·34 to 1·71)	4.54	(3·25 to 6·36)	4.71	(3·28 to 6·76)	1.21	(0.90 to 1.63)	0.64	(0.53 to 0.77)	
Age-group (vs. 10-15y)											
16-17у	1.40	(1·21 to 1·61)	2.30	(1·63 to 3·25)	1.88	(1·35 to 2·63)	1.13	(0.80 to 1.58)	1.08	(0.87 to 1.35)	
18-19y	2.10	(1·82 to 2·42)	4.34	(3·10 to 6·07)	2.76	(1·98 to 3·86)	1.60	(1.13 to 2.26)	1.44	(1.16 to 1.79)	
Socio-economic status (vs. least deprived)											
Second least	1.17	(0·89 to 1·54)	0.81	(0·55 to 1·18)	1.13	(0·70 to 1·84)	1.28	(0·80 to 2·04)	1.15	(0·80 to 1·65)	
Middle	1.19	(0·92 to 1·56)	0.69	(0·47 to 1·01)	1.29	(0·82 to 2·04)	1.12	(0·70 to 1·79)	1.28	(0·90 to 1·81	
Second most	1.53	(1·20 to 1·95)	0.89	(0·64 to 1·25)	1.44	(0·94 to 2·22)	0.97	(0·61 to 1·55)	1.48	(1∙07 to 2∙06	
Most deprived	1.57	(1·24 to 1·98)	0.78	(0·57 to 1·08)	1.64	(1·09 to 2·47)	1.02	(0·67 to 1·57)	1.59	(1·17 to 2·16	
Chronic condition (vs. none)	3.77	(3·38 to 4·20)	1.91	(1·54 to 2·36)	2.53	(2·02 to 3·16)	2.35	(1.80 to 3.07)	10.14	(8.29 to 12.4	
bys											
Adversity- (vs. accident-related) injury	1.94	(1·80 to 2·08)	3.15	(2·73 to 3·63)	3.53	(3·04 to 4·09)	1.26	(1·09 to 1·47)	0.99	(0·84 to 1·17	
Age-group (vs. 10-15y)											
16-17у	1.73	(1·58 to 1·89)	2.70	(2·21 to 3·30)	3.05	(2·41 to 3·84)	1.60	(1·35 to 1·89)	1.14	(0·97 to 1·35	
18-19y	2.23	(2·04 to 2·44)	3.48	(2·83 to 4·26)	5.04	(4·03 to 6·31)	1.91	(1·61 to 2·27)	1.22	(1·02 to 1·45	
Socio-economic status (vs. least deprived)											
Second least	1.24	(1·08 to 1·42)	1.17	(0·89 to 1·54)	1.19	(0·86 to 1·63)	1.62	(1·25 to 2·09)	1.14	(0·88 to 1·47	
Middle	1.28	(1·13 to 1·46)	1.19	(0·92 to 1·56)	1.66	(1·24 to 2·22)	1.62	(1·25 to 2·07)	1.02	(0·79 to 1·32	
Second most	1.42	(1·26 to 1·61)	1.53	(1·20 to 1·95)	1.86	(1·41 to 2·45)	1.29	(1·00 to 1·66)	1.33	(1·05 to 1·68	
Most deprived	1.63	(1·45 to 1·83)	1.57	(1·24 to 1·98)	2.17	(1·66 to 2·82)	1.72	(1·36 to 2·18)	1.26	(1·01 to 1·58	

Chronic condition (vs. none)	2·63 (2·45 to 2·82)	1·26 (1·08 to 1·47)	1·81 (1·56 to 2·09)	1·62 (1·39 to 1·88)	11·72 (10·09 to 13·61)
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Each column (by sex) represents a separate Fine & Gray's competing risks model. Adversity (vs. accident-related) injury, age-group, socio-economic status, and chronic condition status were entered as covariates

545 546 547

simultaneously, per model. DA = Drug/alcohol-related

## 548 **Table 3:** Relative risk of cause-specific death within 10 years after each type of adversity-related index injury (vs. accident-related injury), adjusted for age-

549 group, socio-economic status, and chronic conditions, by sex (multivariable analyses)

	Cause of death, Sub-hazard ratio (95% Confidence Interval)									
Type of adversity-related index injury (vs. accident-related)	All deaths		Suicide		DA death		Accidental death		Other death	
Girls										
Self-inflicted	1.52	(1·33 to 1·73)	5.11	(3·61 to 7·23)	5.14	(3·50 to 7·55)	1.17	(0.85 to 1.63)	0.59	(0.48 to 0.72)
Drug/alcohol-related	1.45	(1·28 to 1·64)	4.55	(3·23 to 6·39)	4.52	(3·14 to 6·51)	1.20	(0.88 to 1.64)	0.62	(0.51 to 0.75)
Violent	1.24	(0·93 to 1·66)	1.48	(0·73 to 2·98)	2.75	(1·47 to 5·17)	1.34	(0.71 to 2.55)	0.76	(0.46 to 1.23)
Boys										
Self-inflicted	2.83	(2·58 to 3·10)	6.20	(5·27 to 7·30)	5.91	(4·96 to 7·03)	1.31	(1.05 to 1.64)	1.07	(0.84 to 1.35)
Drug/alcohol-related	2.46	(2·27 to 2·66)	4.51	(3·89 to 5·24)	4.91	(4·24 to 5·73)	1.40	(1.18 to 1.67)	1.11	(0.92 to 1.34)
Violent	1.25	(1·13 to 1·39)	1.43	(1·15 to 1·78)	1.78	(1·44 to 2·19)	1.10	(0.90 to 1.35)	0.76	(0.59 to 0.97)

Each cell represents a separate Fine & Gray's competing risks model, where the corresponding type of adversity-related index injury (vs. accident-related injury), age-group, socio-economic status, and chronic condition status were entered as covariates simultaneously, per model. Sub-hazard ratios for age-group, socio-economic status, and chronic condition status, for each of the thirty models are not presented here but were very similar to those presented in Table 2 (conditional on sex and cause of death).

DA = Drug/alcohol-related

## 558 Panel: Research in context

#### 559 Systematic review

560 We searched for studies (including reviews) of cause-specific death after hospital attendance for any adversity-related injury published from Jan 1995-May 2016. We 561 562 searched Google Scholar, Scopus, PubMed, and Web of Science using terms 563 "adolescents", "injury", "hospital", "self-harm", "drug or alcohol use", "violence", and "mortality". We found six studies (seven articles), but no relevant 564 systematic review. Five (European) studies reported risks of death due to suicide, and 565 566 some also reported risks of deaths due to drug/alcohol use (n=2), homicide (n=2), 567 undetermined/accidental causes (n=3), and chronic conditions (n=3), in up to 15 years 568 after adolescents presented to hospital with self-inflicted injury. One (US) study 569 reported frequencies of deaths from homicide, drug overdose, and traffic accidents in 570 the two years after discharge following violent injury in 559 adolescents. We did not 571 identify any studies that reported rates of cause-specific death following hospital 572 presentation or admission for drug/alcohol-related injury, or compared risks of cause-573 specific deaths after discharge following any adversity-related injury with those

574 following accident-related injury.

#### 575 Interpretation

576 Our study adds new evidence on the risks of cause-specific death up to ten years after 577 discharge following adversity-related and accident-related injury among young 578 people. Our finding of elevated risks of suicide following all types of adversity-579 related injury (versus accident-related injury) suggests that clinical and public health 580 strategies need to be extended to reduce harm after all types of adversity-related 581 injury, whether self-inflicted, drug/alcohol-related or violent. Similar risks of suicide 582 and drug/alcohol-related deaths following discharge from any type of index injury 583 found in our study also stress the need of preventive strategies, both within and 584 outside the healthcare sector, to reduce public health burden of suicide and 585 drug/alcohol-related deaths.