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# Contributions of specific causes of death to lost life expectancy in severe mental illness

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#### ABSTRACT

The life expectancy gap between people with severe mental illness (SMI) and the general population persists and may even be widening. This study aimed to estimate contributions of specific causes of death to the gap. Age of death and primary cause of death were used to estimate life expectancy at birth for people with SMI from a large mental healthcare case register during 2007-2012. Using data for England and Wales in 2010, death rates in the SMI cohort for each primary cause of death category were replaced with gender- and age-specific norms for that cause. Life expectancy in SMI was then recalculated and thus, the contribution of that cause of death estimated. Natural causes accounted for 79.2% of lost life-years in women with SMI and 78.6% in men. Deaths from circulatory disorders accounted for more life-years lost in women than men (22.0% versus 17.4%, respectively), as did deaths from cancer (8.1% versus 0%), but the contribution from respiratory disorders was lower in women than men (13.7% versus 16.5%). For women, cancer contributed more in those with non-affective than affective disorders, while suicide, respiratory and digestive disorders contributed more in those with affective disorders. In men, respiratory disorders contributed more in non-affective disorders. Other contributions were similar between gender and affective/non-affective groups. Loss of life expectancy in people with SMI is accounted for by a broad range of causes of death, some varying by gender and diagnosis. Interventions focused on multiple rather than individual causes of death should be prioritised accordingly.

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### 1. Introduction

Premature death among people with mental disorders is an international concern [1]. Individuals with severe mental illness (SMI), including schizophrenia, schizoaffective disorder and bipolar disorder, experience particularly higher all-cause mortality and lower life expectancy [2,3]. This is an important indicator for policy development to improve on current health inequality; however, contributions of individual causes of death to the life expectancy gap remain unclear, limiting the evidence. Previous research has suggested excesses of both natural and unnatural causes of death, and that the excess of natural causes may cross several mental disorder groups [4]. Nevertheless, standardised mortality ratios cannot be translated into the more important construct of life expectancy, which accounts for the age of death as well as the absolute risk. Specifically, life expectancy as a measure

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adds value because it is a demographic index, which is a feature of 24 overall mortality emphasizing the impact of deaths occurring in 25 younger age groups. It can thus be claimed to have considerably 26 more 'meaning' for a population as an indicator of disadvantage, 27 compared to a relative risk or hazard of mortality over a defined 28 period of time. Life expectancy as a measure is obtained from 29 estimates from the age-specific mortality of a specific cohort over a 30 given period of time using the life table method and is calculated 31 from the accumulated person-years contributed by the entire 32 cohort divided by the total population number at birth. 33

The substantial disadvantage in life expectancy experienced by 34 people with SMI represents a major health inequality. Recent 35 studies have shown how the impact of chronic illness varies 36 between people with SMI and the general population. For example, 37 the relative risk of type 2 diabetes is elevated in younger 38 populations of SMI compared to general population. In one study, 39 most associations persisted despite adjustment for antipsychotic 40 prescriptions, and ethnic minority groups also had a higher 41 prevalence of type 2 diabetes in the presence of severe mental 42 illness [5]. On a similar note, the prevalence of multimorbidity 43

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(which often encompasses combinations of mental and physical illnesses) is recognised to increase with higher deprivation [6]. Further, in people with SMI, prevalence of obesity, metabolic syndrome, diabetes mellitus, symptoms of cardiovascular disease, and respiratory disease all exceed those in the general population at least two-fold, and HIV prevalence may be increased as much as eight-fold [7]. This increased prevalence of chronic disease may be largely responsible for a five-fold increased risk of death [7].

52 An analysis was strongly warranted to add clarity to this 53 'survival gap', particularly as interventions have been proposed, 54 which focus on individual pathways without clear evidence on 55 the contribution of those pathways to the overall outcome. 56 Using data from a large clinical cohort, we estimated the 57 contribution of the major groups of causes of death to shortened 58 life expectancy in people with SMI, delineating to what extent 59 this gap could conceivably be closed if mortality rates for each 60 cause of death could be equalised with those in the general 61 population.

#### 62 2. Methods

#### 63 2.1. Data sources and cohort

64 The South London and Maudsley NHS Foundation Trust (SLAM) 65 is one of Europe's largest secondary mental healthcare providers, 66 serving a single geographic catchment of approximately 1.36 mil-67 lion residents in four southeast London boroughs. In 2006, fully 68 electronic health records were implemented across all SLAM 69 services, and, in 2008, the Clinical Record Interactive Search (CRIS) 70 system, supported by the National Institute of Health Research 71 (NIHR) Biomedical Research Centre for Mental Health at SLAM, was 72 developed to allow search and retrieval of anonymised but 73 complete medical records within a robust patient-led governance 74 framework [8,9]. Using CRIS, individuals who had a primary 75 diagnosis of SMI (schizophrenia, ICD-10 coded F20; schizoaffective 76 disorder, F25, or bipolar affective disorder, F31) and had at least 77 one active contact with SLaM during the period between 1st Jan 78 2007 and 31st Dec 2012 were included in this analysis. Recorded 79 primary diagnoses in structured fields of the record were 80 supplemented by a natural language processing application developed using Generalised Architecture for Text Engineering 81 (GATE) software to identify diagnostic statements in open text 82 83 fields (e.g. correspondence and clinical notes) [9,10].

#### 84 2.2. Mortality

85 Date and primary cause of death for our study subjects in the 86 same period (i.e. 2007-2012 inclusive) were retrieved from a CRIS 87 linkage to death certifications, held by the UK Office of National 88 Statistics, which is updated regularly [11], by NHS numbers 89 through an anonymised process. The same data source (i.e. UK 90 Office of National Statistics), but this time via online data files, was 91 used to obtain age-, gender-, and cause-specific mortality for 92 England and Wales population in 2010 [12].

93 2.2.1. Analysis: baseline life expectancy at birth

94 Using Microsoft Excel software, life expectancy at birth was 95 estimated using Chiang's abridged life table method [13]. In brief, 96 the accumulated person-years were calculated by gender based on 97 age- and gender-specific mortality experienced by a specified 98 male/female cohort in five-year age bands, and then divided by the 99 total number of male/female cohort subjects at baseline. Because 100 of dynamic nature of our cohort during the six-year follow-up 101 period (2007-2012), the number of deaths observed was weighted 102 by the average at risk period of each age and gender band. The life 103 expectancy at birth for the England and Wales population using 2010 data was also estimated for the purpose of validation using 104 105 Chiang's abridged method [13,14]. Since SMI diagnoses are 106 relatively unlikely to be given to people under the age of 15 years, mortality rates up to that age were assumed to be equal to those in 107 the general population of England and Wales in 2010. 108

2.2.2. Analysis: re-calculated life expectancies by equalising mortality rates

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Our aim was to estimate the improvements in life expectancy 111 which might be achieved if people with SMI experienced the same 112 age- and gender-specific mortality rates for particular causes of 113 death as those in the general population. For this, a life expectancy 114 simulation was used to examine the contribution of major causes 115 of mortality to the reduced life expectancy in people with SMI. 116 Death rates in SMI cohort for each major primary cause category 117 were sequentially replaced with England and Wales gender- and 118 age-specific rates for that cause of death, obtained from UK Office 119 of National Statistics [12]. The expected numbers of deaths in each 120 121 five-year age band were then re-calculated using these national gender- and age-specific rates. These re-estimations were carried 122 out for cause-of-death categories by the order of number of deaths 123 124 observed in the SMI study cohort:

126 circulatory diseases; 127 • cancer: • respiratory diseases: 128 • digestive diseases; 129 • suicide: 130 • deaths due to other (non-suicide) external causes; 131 132

• all remaining causes of death.

In this study, in accordance with the ICD-10 manual, external 133 causes comprised mortality involving injury, poisoning and certain 134 other external causes. Causes of death in this broad group of 135 scenarios include examples, such as "accidental poisoning by 136 amphetamine" (X41) or "driver of car injured in non-collision 137 transport accident" (V48.5). Here, the underlying cause is classified 138 according to the circumstances of the fatal injury, rather than the 139 nature of the injury, which is coded separately. Of note, this means 140 that direct alcohol related mortality is considered under external 141 142 causes. Where there was a reverse association identified for any cause-of-death category (i.e. a lower life expectancy for SMI 143 subjects following equalisation of a specific cause-of-death 144 category to general population), its contribution to loss of life 145 expectancy was set at zero. The percentage of person-year 146 contribution to the life expectancy gap for each category of death 147 was then calculated accordingly. In a sensitivity analysis, the 148 adjustment for cancer was re-applied taking into account reference 149 to cancer anywhere on the death certificate (i.e. rather than just as 150 the primary cause of death). Finally, we carried out an additional 151 stratification by affective (bipolar/schizoaffective) and non-affec-152 tive (schizophrenia) SMI diagnosis in order to compare cause of 153 death contributions between these subgroups. 154

### 3. Results

Calculated by Chiang's method, the life expectancy at birth for 156 the England and Wales population in 2010 was 78.5 (95% CI 78.1-157 158 78.8) years for males and 82.4 (95% CI 82.1–82.7) years for females: 159 identical to values previously reported by the UK ONS [14]. In the SLAM cohort, there were 1558 deaths during the observation 160 period among 19,106 subjects with SMI (816 deaths among 10,414 161 males and 742 deaths among 8692 females). The life expectancy at 162 birth in people with SMI was estimated to be 67.9 years (95% CI 163 67.1-68.6) for men and 72.2 years (95% CI 71.5-72.8) for women -164 i.e. reduced by 10.6 years for men and 10.2 years for women. 165

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