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Age of onset and cumulative risk of mental disorders

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Articles



M Age of onset and cumulative risk of mental disorders: a cross-national analysis of population surveys from 29 countries

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Summary

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*WHO World Mental Health Survey Collaborators are listed in the appendix (pp 2-4)

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Background Information on the frequency and timing of mental disorder onsets across the lifespan is of fundamental importance for public health planning. Broad, cross-national estimates of this information from coordinated general population surveys were last updated in 2007. We aimed to provide updated and improved estimates of age-of-onset distributions, lifetime prevalence, and morbid risk.

Methods In this cross-national analysis, we analysed data from respondents aged 18 years or older to the World Mental Health surveys, a coordinated series of cross-sectional, face-to-face community epidemiological surveys administered between 2001 and 2022. In the surveys, the WHO Composite International Diagnostic Interview, a fully structured psychiatric diagnostic interview, was used to assess age of onset, lifetime prevalence, and morbid risk of 13 DSM-IV mental disorders until age 75 years across surveys by sex. We did not assess ethnicity. The surveys were geographically clustered and weighted to adjust for selection probability, and standard errors of incidence rates and cumulative incidence curves were calculated using the jackknife repeated replications simulation method, taking weighting and geographical clustering of data into account.

Findings We included 156331 respondents from 32 surveys in 29 countries, including 12 low-income and middleincome countries and 17 high-income countries, and including 85 308 (54.5%) female respondents and 71023 (45.4%) male respondents. The lifetime prevalence of any mental disorder was 28.6% (95% CI 27.9–29.2) for male respondents and 29.8% (29.2-30.3) for female respondents. Morbid risk of any mental disorder by age 75 years was 46.4% (44.9-47.8) for male respondents and 53.1% (51.9-54.3) for female respondents. Conditional probabilities of first onset peaked at approximately age 15 years, with a median age of onset of 19 years (IQR 14-32) for male respondents and 20 years (12-36) for female respondents. The two most prevalent disorders were alcohol use disorder and major depressive disorder for male respondents and major depressive disorder and specific phobia for female respondents.

Interpretation By age 75 years, approximately half the population can expect to develop one or more of the 13 mental disorders considered in this Article. These disorders typically first emerge in childhood, adolescence, or young adulthood. Services should have the capacity to detect and treat common mental disorders promptly and to optimise care that suits people at these crucial parts of the life course.

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Introduction

Age of onset, lifetime prevalence (ie, the proportion of survey respondents with a history of disorder at the time of assessment), and lifetime morbid risk (ie, the projected lifetime prevalence in the sample as of a fixed age) are essential features of epidemiology. Many mental disorders have an onset in the first and second decades of life, $^{\scriptscriptstyle 1\! 2}$ unlike most other non-communicable disorders (eg, respiratory and cardiovascular disorders or cancer), which typically have onsets in late adulthood. Understanding onset patterns is important for several reasons. First, this understanding helps to ensure that the correct mix of services is available to provide prompt treatments to the right groups (eg, early intervention for teenagers with mental disorders). Second, research efforts should focus on understanding risk factors for different types of mental disorders during crucial parts of the lifespan. Third, register-based family pedigree studies3 and genome-wide association studies based on case-control or case-cohort studies4 increasingly use ageof-onset distributions to weight non-cases at the time of sample ascertainment, according to the estimated future

Research in context

Evidence before this study

Age of onset, lifetime prevalence, and lifetime morbid risk of mental disorders are key estimates for service planning as they provide valuable insight into when in the life course disorders first emerge. Previous studies of these estimates often used suboptimal samples (eg, case-only samples or registers of patients who were treated). Few studies present both age of onset and lifetime prevalence for diverse mental disorders based on population-based data from multiple countries. We searched PubMed with the search terms (["age of onset"[TIAB] OR "lifetime prevalence"[TIAB] OR "morbid risk"[TIAB] OR "cumulative incidence" [TIAB]) AND ("mental"[TIAB] OR "psychiatri*"[TIAB]) for articles in any language published between Jan 1, 1966, and June 30, 2022. We identified 4050 articles, including 93 systematic reviews. Most studies focused on one type of mental disorder in one country. A systematic review of age of onset summarised data from 192 studies but did not examine lifetime prevalence. Regarding cross-national studies, one 2007 study reported country-specific age of onset, lifetime prevalence, and morbid risk for 17 countries, but cross-national and sex-specific estimates were not presented.

morbid risk of the disorder of interest, which means having accurate onset information is important, although other factors are also important in controlling for bias in these types of studies. Finally, disease-specific age-of-onset distributions are important inputs for models that are used to estimate the non-fatal burden of disorders.⁵

The most comprehensive data on age of onset, lifetime prevalence, and morbid risk of common mental disorders to date were reported in 2007 by the World Mental Health (WMH) survey collaborators¹ on the basis of data obtained from coordinated community epidemiological surveys in 17 countries. Key features (eg, median and other quantiles) were reported, providing evidence that many mental disorders first emerge between childhood and early adulthood. In 2022, Solmi and colleagues² presented a systematic review and meta-analysis of published literature on age of onset including 192 studies. The authors noted that the studies were heterogeneous, which made pooling the data difficult, but they presented key properties of age-ofonset distributions (eg, peak age of onset and proportion of people with mental disorders with onset by age 25 years) by disorder type and sex.

Epidemiological studies commonly show sex-specific incidence rates by age; the rate of first onset at a specific age is defined as a ratio of the number of disorder onsets divided by the number of people who never had the disorder up to that age and lived through that age. These estimates allow the calculation of cumulative lifetime risk by sex. Whereas incidence rates can either increase or decrease with increasing age, cumulative risk curves

Added value of this study

Using data from community epidemiological surveys in 29 countries, including 16 surveys from the 2007 study and 16 more recent surveys with 13 additional countries, we found a lifetime prevalence for any mental disorder of 28.6% for male respondents and 29.8% for female respondents. The risk for any mental disorder by age 75 years was around one in two people. The peak incidence was at around age 15 years, and the median age of onset was 19 years for male respondents and 20 years for female respondents. In this Article, we provide updated and detailed estimates related to the age of onset, lifetime prevalence, and morbid risk of mental disorders.

Implications of all the available evidence

Mental disorders are common by age 75 years. The updated estimates substantiate that many mental disorders have first onsets during childhood and young adulthood. Health planners should ensure sufficient services to reach out and treat mental disorders among young people. Future research related to causes and prevention can be informed by the distinct age-ofonset curves and sex differences associated with different types of mental disorders.

are always non-decreasing. Univariate statistics can be derived from these cumulative risk curves (eg, median and IQR of the age-of-onset distributions) to estimate lifetime morbid risk. However, the morbid risk does not indicate how many people had the disorder as of their current age, which is known as lifetime prevalence. By comparing the ratio of morbid risk as of some fixed age to lifetime prevalence, we can appreciate how age-related incidence interacts with background population age structure. Disorders with peak incidence in early life will have a lower ratio of morbid risk to lifetime prevalence than will disorders with peak hazard rates later in life.

Existing data on age of onset and morbid risk are prone to under-reporting6 because survey data rely on memory. Recall bias might result in a systematic bias against recalling events in the distant past or telescoping the recalled age (eg, temporally distant events might be incorrectly recalled as having occurred more recently). Although the structured interview used in WMH surveys is designed to reduce recall bias,7 accuracy can also be improved by focusing on respondents who reported more recent onsets (eg, in the past 10 years).8 In this Article, we used this method to update estimates of age of onset and morbid risk for 13 defined mental disorders. Since the 2007 publication,1 WMH surveys have been completed with 74989 respondents, including people in 13 additional countries. We combined these new data with data from the earlier WMH surveys with the aim of providing updated and improved estimates of age-ofonset distributions, lifetime prevalence, and morbid risk.

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| | Survey | Sample characteristics | Survey dates | Age range of participants, years | Sample s | ize | | Sex of particip | ants, n/N (%) | Response rate* |
|-----------------------|----------------------------------|--|-----------------|--|----------|--------|--|--------------------------|------------------------|-------------------|
| | | | | | Part 1 | Part 2 | Part 2 and participants aged ≤44 years† | Male | Female | - |
| Low-income and | d middle-incom | ne countries | | | | | | | | |
| Brazil-São Paulo | São Paulo Megacity | São Paulo metropolitan area | 2005–08 | 18-93 | 5037 | 2942 | NA | 2187/5037 (43·4%) | 2850/5037 (56·6%) | 81.3% |
| Bulgaria | NSHS | Nationally representative | 2002–06 | 18-98 | 5318 | 2233 | 741 | 2430/5318 (45·7%) | 2888/5318 (54·3%) | 72.0% |
| Bulgaria 2 | NSHS-2 | Nationally representative | 2016–17 | 18–91 | 1508 | 578 | NA | 670/1508 (44·4%) | 838/1508 (55·6%) | 61.0% |
| Colombia‡ | NSMH | All urban areas of the country (approximately 73% of the total national population) | 2003 | 18-65 | 4426 | 2381 | 1731 | 1700/4426 (38·4%) | 2726/4426 (61·6%) | 87.7% |
| Colombia- Medellín | MMHHS | Medellín metropolitan area | 2011–12 | 19-65 | 3261 | 1673 | NA | 1120/3261 (34·3%) | 2141/3261 (65·7%) | 97.2% |
| Iraq | IMHS | Nationally representative | 2006–07 | 18-96 | 4332 | 4332 | NA | 2091/4332 (48·3%) | 2241/4332 (51·7%) | 95·2% |
| Lebanon‡ | LEBANON | Nationally representative | 2002-03 | 18-94 | 2857 | 1031 | 595 | 1297/2857 (45·4%) | 1560/2857 (54·6%) | 70.0% |
| Mexico‡ | M-NCS | All urban areas of the country (approximately 75% of the total national population) | 2001–02 | 18-65 | 5782 | 2362 | 1736 | 2285/5782 (39·5%) | 3497/5782 (60·5%) | 76.6% |
| Nigeria‡ | NSMHW | 21 (58%) of 36 states in the country, representing 57% of the national population; surveys were conducted in Yoruba, Igbo, Hausa, and Efk languages | 2002-04 | 18-100 | 6752 | 2143 | 1203 | 3315/6752 (49·1%) | 3437/6752 (50·9%) | 79.3% |
| Peru | EMSMP | Five urban areas of the country (approximately 38% of the total national population) | 2004–05 | 18–65 | 3930 | 1801 | 1287 | 1759/3930 (44·8%) | 2171/3930 (55·2%) | 90-2% |
| China- Shenzhen§ | Shenzhen | Shenzhen metropolitan area; included temporary residents and household residents | 2005–07 | 18-88 | 7132 | 2475 | NA | 3614/7132 (50·7%) | 3518/7132 (49·3%) | 80.0% |
| Romania | RMHS | Nationally representative | 2005-06 | 18-96 | 2357 | 2357 | NA | 1092/2357 (46·3%) | 1265/2357 (53·7%) | 70.9% |
| South Africa‡§ | SASH | Nationally representative | 2002-04 | 18-92 | 4315 | 4315 | NA | 1718/4315 (39·8%) | 2597/4315 (60·2%) | 87.1% |
| Ukraine‡ | CMDPSD | Nationally representative | 2002 | 18-91 | 4725 | 1720 | 541 | 1793/4725 (37·9%) | 2932/4725 (62·1%) | 78·3% |
| Total | | | | | 61732 | 32343 | 7834 | 27 071/61 732 (43·9%) | 34661/61732 (56·1%) | 80.4% |
| High-income co | ountries | | | | | | | | | |
| Argentina | AMHES | Eight largest urban areas of the country (approximately 50% of the total national population) | 2015 | 18-98 | 3927 | 2116 | NA | 1692/3927 (43·1%) | 2235/3927 (56·9%) | 77·3% |
| Australia§ | NSMHWB | Nationally representative | 2007 | 18-85 | 8463 | 8463 | NA | 3843/8463 (45·4%) | 4620/8463 (54·6%) | 60.0% |
| Belgium‡ | ESEMeD | Nationally representative; sample was selected from a national register of residents of Belgium | 2001–02 | 18-95 | 2419 | 1043 | 486 | 1190/2419 (49·2%) | 1229/2419 (50·8%) | 50.6% |
| France‡ | ESEMeD | Nationally representative; sample was selected from a national list of households with listed telephone numbers | 2001–02 | 18-97 | 2894 | 1436 | 727 | 1329/2894 (45·9%) | 1565/2894 (54·1%) | 45·9% |
| Germany‡ | ESEMeD | Nationally representative | 2002-03 | 19-95 | 3555 | 1323 | 621 | 1660/3555 (46·7%) | 1895/3555 (53·3%) | 57.8% |
| lsrael‡ | NHS | Nationally representative | 2003-04 | 21–98 | 4859 | 4859 | NA | 2380/4859 (49·0%) | 2479/4859 (50·9%) | 72.6% |
| Italy‡ | ESEMeD | Nationally representative; sample was selected from municipality resident registries | 2001–02 | 18–100 | 4712 | 1779 | 853 | 2321/4712 (49·3%) | 2391/4712 (50·7%) | 71·3% |
| Japan‡ | WMHJ 2002–2006 | 11 metropolitan areas | 2002-06 | 20–98 | 4129 | 1682 | NA | 1868/4129 (45·2%) | 2261/4129 (54·8%) | 55·1% |
| | (Table 1 continues on next page) | | | | | | | | | |

| Survey | | vey Sample characteristics | | Survey Age range of dates participants, years | | ize | | Sex of participants, n (%) | | Response rate* |
|---------------------|--------------------|---|---------|---|---------|--------|--|----------------------------|---------------------------|-------------------|
| | | | | | Part 1 | Part 2 | Part 2 and participants aged ≤44 years† | Male | Female | - |
| (Continued from | n previous page | 2) | | | | | | | | |
| Netherlands‡ | ESEMeD | Nationally representative; sample was selected from municipal postal registries | 2002-03 | 18–95 | 2372 | 1094 | 516 | 1032/2372 (43·5%) | 1340/2372 (56·5%) | 56.4% |
| New Zealand‡§ | NZMHS | Nationally representative | 2004-05 | 18–98 | 12790 | 7312 | NA | 5537/12 790 (43·3%) | 7253/12790 (56·7%) | 73.3% |
| Northern Ireland | NISHS | Nationally representative | 2005-08 | 18–97 | 4340 | 1986 | NA | 1899/4340 (43·8%) | 2441/4340 (56·2%) | 68-4% |
| Poland | EZOP | Nationally representative | 2010–11 | 18-65 | 10081 | 4000 | 2276 | 4883/10081 (48·4%) | 5198/10081 (51·6%) | 50.4% |
| Portugal | NMHS | Nationally representative | 2008–09 | 18-81 | 3849 | 2060 | 1070 | 1632/3849 (42·4%) | 2217/3849 (57·6%) | 57·3% |
| Qatar | WMHQ | Nationally representative; sample was selected from a national list of mobile telephone numbers and restricted to Qatari nationals and Arab expatriates¶ | 2019-22 | 18-90 | 5195 | 2583 | NA | 3215/5195 (61·9%) | 1980/5195 (38·1%) | 19.2% |
| Saudi Arabia§ | SNMHS | Nationally representative | 2013–16 | 18-65 | 3638 | 1793 | NA | 1719/3638 (47·3%) | 1919/3638 (52·7%) | 61.0% |
| Spain‡ | ESEMeD | Nationally representative | 2001-02 | 18–98 | 5473 | 2121 | 960 | 2421/5473 (44·2%) | 3052/5473 (55·8%) | 78.6% |
| Spain-Murcia | PEGASUS- Murcia | Murcia region; regionally representative | 2010–12 | 18–96 | 2621 | 1459 | NA | 1192/2621 (45·5%) | 1429/2621 (54·5%) | 67.4% |
| USA‡ | NCS-R | Nationally representative | 2001-03 | 18–99 | 9282 | 5692 | 3197 | 4139/9282 (44·6%) | 5143/9282 (55·4%) | 70.9% |
| Total | | | | | 94599 | 52801 | 10706 | 43 952/94 599 (46·5%) | 50 647/94 599 (53·5%) | 56.0% |
| Overall total | | | | | 156 331 | 85144 | 18540 | 71023/156331 (45·4%) | 85 308/156 331 (54·6%) | 63.6% |

Data on income categories are from 2012 <u>World Bank data</u>. Some WMH survey countries have moved into new income categories since the surveys were done; we used income groups at the time of data collection. AMHES=Argentina Mental Health Epidemiologic Survey. CMDPSD=Comorbid Mental Disorders during Periods of Social Disruption. EMSMP=La Encuesta Mundial de Salud Mental en el Peru. ESEMeD=The European Study of The Epidemiology of Mental Disorders. EZOP=Epidemiology of Mental Disorders and Access to Care Survey. IMHS=Iraq Mental Health Survey. LEBANON=LebAnese Evaluation of the Burden of Ailments and Needs of the Nation. M-NCS=The Mexico National Comorbidity Survey. MMHHS=Medellín Mental Health Household Study. NA=not applicable. NCS-R=The US National Comorbidity Survey Replication. NHS=Israel National Health Survey. NISHS=Northern Ireland Study of Health and Stress. NMHS=Portugal National Mental Health Survey. Of Mental Health. NSMHW=The Nigerian Survey of Mental Health and Wellbeing. NSMHWB=National Survey of Mental Health. Survey. PEGASUS-Murcia=Psychiatric Enquiry to General Population in Southeast Spain-Murcia. RMHS=Romania Mental Health Survey. SASH=South Africa Health Survey. SNMHS=Saudi National Mental Health. WMHJ2002-2006=World Mental Health Japan Survey. WMHQ=World Mental Health Qatar Study. *Response rate is the ratio of the number of households or iginally sampled, excluding from the denominator households known not to be eligible either because of being vacant at the time of initial contact or because the residents were unable to speak the designaned survey languages. †Argentina, Australia, Brazil, Bulgaria 2, Colombia-Medellín, Iraq, Israel, Japan, New Zealand, Northern Ireland, China-Shenzhen, Qatar, Romania, Saudi Arabia, South Africa, and Spain-Murcia did not have an age-restricted part 2 sample. All other surveys, were onducted after that report. §For cross-national comparisons, we limited the sample to participants aged 14 years or younger. \$Ivrveys included in the 2007 WMH report; all other su

Table 1: WMH survey sample characteristics by World Bank income categories

Methods

Study design and samples

In this cross-national analysis, we obtained data from 32 WMH surveys conducted in 29 countries. All surveys were based on rigorous, multistage, geographically clustered area probability household sampling designs and were conducted between 2001 and 2022. A detailed description of sample designs is presented in an earlier report.⁹ A discussion of possible reasons for variation in WMH survey response rates is presented elsewhere.¹⁰

Most WMH surveys are based on stratified, multistage, clustered area probability samples of households in the

participating countries. Areas equivalent to counties or municipalities in the USA were selected in the first stage, followed by one or more stages of geographical sampling (eg, towns within counties, blocks [typically 50–100 housing units] within towns, or households within blocks) to establish a nationally representative sample of households. In each household, an attempt was made to obtain a list of all adult (ie, aged \geq 18 years) household members. Then one person (or in surveys conducted in Brazil-São Paulo, Belgium, Bulgaria, France, Germany, Italy, Lebanon, Nigeria, Saudi Arabia, Spain, and the USA, two people) was selected from this (Prof V Kovess-Masfety PhD); Unit of Epidemiological and Evaluation Psychiatry, Istituto di Ricovero e Cura a Carattere Scientifico Istituto Centro San Giovanni di Dio Fatebenefratelli, Brescia, Italy (M Magno MClinPsych); National Institute of Psychiatry Ramón de la Fuente Muñiz, Mexico City, Mexico (Prof M E Medina-Mora DrPh); Institute of Psychiatry and Neurology, Warsaw, Poland (Prof J Moskalewicz PhD);

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Procedures for obtaining informed consent and ethics approval were somewhat different across countries, but ethics approval was always obtained from the institutional review boards of the collaborating organisations in each country before beginning the WMH survey and written or oral informed consent, depending on the country, was obtained from respondents before beginning the interview. Data use agreements allowed only deidentified data to be deposited in the centralised WMH server and required all analyses to be conducted on that server by trained and approved WMH analysts.

Procedures

Interviews were conducted in two parts. Part 1 was administered to one randomly selected adult (or in surveys conducted in some countries, two randomly selected adults) in each sampled household. Part 1 contained assessments of core mental disorders (ie, depression, mania, panic disorder, social phobia, specific phobia, agoraphobia, generalised anxiety disorder, and substance use disorder). A part 1 weight adjusted for differential probabilities of selection within households (on the basis of number of eligible adults in the household) and between households (on the basis of discrepancies between census estimates of the number of households in a sample segment and the number of households found when the interviewers visited that segment). Part 2, which included questions about other mental disorders and correlates (which varied across surveys but were based on the US survey), was then administered to 100% of part 1 respondents who met lifetime criteria for any part 1 disorder and a random subsample, varying across surveys between 20% and

| | Male sex | | | Female sex | | | χ²(p value)* |
|---------------------------------|--|---|--|--|---|--|------------------|
| | Number of lifetime cases at time of interview† | Number of people in sample in whom disorder was assessed‡ | Lifetime prevalence, % (95% CI)§ | Number of lifetime cases at time of interview† | Number of people in sample in whom disorder was assessed‡ | Lifetime prevalence, % (95% CI)§ | |
| Anxiety disorders | | · | | | | | |
| Panic disorder or agoraphobia | 1309 | 67808 | 1.9% (1.8–2.1) | 3303 | 83328 | 3.7% (3.6–3.9) | 289.6 (<0.0001) |
| Generalised anxiety disorder | 1955 | 71023 | 2.7% (2.6–2.9) | 4500 | 85308 | 5.0% (4.8-5.1) | 396.8 (<0.0001) |
| Post-traumatic stress disorder | 1456 | 35 4 4 6 | 2.7% (2.5–2.9) | 3721 | 47 223 | 5.4% (5.2-5.7) | 279.8 (<0.0001) |
| Social phobia | 2254 | 65428 | 3.5% (3.3-3.7) | 3770 | 80849 | 4.6% (4.4-4.8) | 69.5 (<0.0001) |
| Specific phobia | 2825 | 56355 | 5.0% (4.8–5.3) | 6954 | 68781 | 10.0% (9.7–10.2) | 744.6 (<0.0001) |
| Any anxiety disorder | 6559 | 39 060 | 11.3% (10.9–11.7) | 13830 | 50741 | 18.8% (18.3–19.2) | 669.8 (<0.0001) |
| Mood disorders | | | | | | | |
| Major depressive disorder | 5324 | 71023 | 7.5% (7.2–7.7) | 12144 | 85308 | 13.6% (13.3–13.9) | 1160.5 (<0.0001) |
| Bipolar disorder | 1398 | 57 559 | 2.5% (2.4–2.7) | 1593 | 68 307 | 2.3% (2.1-2.4) | 7.2 (0.0074) |
| Any mood disorder | 6674 | 71023 | 9.5% (9.2–9.7) | 13675 | 85308 | 15-4% (15-1–15-7) | 880.6 (<0.0001) |
| Substance use disorders | | | | | | | |
| Alcohol abuse | 7629 | 52757 | 13.7% (13.3–14.1) | 2602 | 67 650 | 3.3% (3.1-3.4) | 2226.0 (<0.0001) |
| Alcohol dependence | 2056 | 52757 | 3.5% (3.3-3.7) | 840 | 67 650 | 0.9% (0.9–1.0) | 516·5 (<0·0001) |
| Drug abuse | 2083 | 46180 | 4.1% (3.9-4.3) | 1204 | 59642 | 1.7% (1.6–1.8) | 337.4 (<0.0001) |
| Drug dependence | 711 | 46180 | 1.4% (1.3–1.6) | 490 | 59642 | 0.6% (0.6–0.7) | 89.9 (<0.0001) |
| Any substance use disorder | 7926 | 46939 | 15.6% (15.2–16.1) | 3181 | 60943 | 4.5% (4.2-4.7) | 1960-2 (<0-0001) |
| Externalising disorders | | | | | | | |
| ADHD | 571 | 19 402 | 2.4% (2.1–2.7) | 575 | 25116 | 1.5% (1.3–1.7) | 26.1 (<0.0001) |
| Intermittent explosive disorder | 1326 | 40 529 | 3.5% (3.2-3.7) | 1267 | 49561 | 2.5% (2.4–2.7) | 39.4 (<0.0001) |
| Any externalising disorder | 1524 | 30 5 2 6 | 4.3% (3.9-4.6) | 1563 | 39231 | 3.1% (2.9–3.3) | 43.0 (<0.0001) |
| Any mental disorder | 14662 | 36700 | 28.6% (27.9–29.2) | 21485 | 48 4 4 4 | 29.8% (29.2–30.3) | 9.2 (0.0024) |

*Wald test for significance of difference between male sex and female sex values. †Observed (ie, unweighted) number of respondents classified as meeting criteria for the disorder. ‡Observed (ie, unweighted) number of respondents in the sample. Scalculated with weighted data.

Table 2: Count and lifetime prevalence of disorders by sex

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33%, of other part 1 respondents. A part 2 weight equal to the inverse of the probability of selection into part 2 was used to restore the representativeness of the part 2 sample, resulting in the prevalence estimates of part 1 disorders in the doubly weighted part 2 sample having the same expected-values-weighted prevalence estimates as in the part 1 sample. A third weight was then applied to the doubly weighted part 2 sample to calibrate discrepancies between sample and population distributions on the cross-classification of census sociodemographic and geographical variables.

Measures

The WHO Composite International Diagnostic Interview (CIDI)11 was used to assess DSM-IV disorders in the WMH surveys (except for the 2019-22 survey in Qatar, in which DSM-5 disorders were assessed). The CIDI is a fully structured diagnostic interview administered by trained interviewers who read questions word for word and record answers in prespecified categories. Sex data were recorded by interviewer observation or, when unsure, by asking respondents if they were male or female. No options were included for gender categories, but all responses other than male or female were coded as other. Consistent interviewer training and quality control monitoring procedures were used across surveys.¹² On the basis of DSM-IV or DSM-5 criteria, 13 specific diagnoses were identified: panic disorder or agoraphobia; generalised anxiety disorder; post-traumatic stress disorder (PTSD); social phobia; specific phobia; major depressive disorder; bipolar disorder; alcohol abuse disorder; alcohol dependence disorder; drug abuse disorder; drug dependence disorder; ADHD; and intermittent explosive disorder. Diagnoses were also pooled for any anxiety disorder, any mood disorder, any substance use disorder, any externalising disorder, and any disorder. We retained the DSM-IV distinction between alcohol and drug abuse and dependence although DSM-5 combines these into single alcohol and drug use disorders. Clinical reappraisal studies indicate that lifetime diagnoses based on the CIDI have good concordance with diagnoses based on masked, semistructured (ie, allowing open-ended clinical probing), clinical research diagnostic interviews.13

Respondents who met lifetime criteria for a specific disorder were asked about their age of onset with a question series designed to review core symptoms and encourage accurate dating. For example, the CIDI question about onset of a major depressive episode was administered after the completion of a question series that focused on symptoms of the respondents' worst lifetime episode, establishing that lifetime criteria were met. The next CIDI question after completing that question series was "Think of the very first time in your life when you had an episode lasting two weeks or longer when most of the day nearly every day you felt (sad/or/ discouraged/or/uninterested) and also had some of the other problems we just reviewed. Can you remember your exact age?". Respondents who could not remember their exact ages were then questioned about the earliest age they could clearly remember having the syndrome with the aim of obtaining upper-bound onset estimates. Symptom-level assessments were only obtained for worst lifetime episodes, not for first onsets, introducing the possibility that age-of-onset estimates were for subthreshold episodes. A more detailed description of the CIDI is presented elsewhere.¹¹

Statistical analysis

We pooled results across surveys with sums of weights equal to numbers of respondents rather than country populations. We estimated lifetime prevalence in the doubly weighted and calibrated part 2 sample as the proportion of respondents who had ever had a specific disorder as of the time of interview. For each disorder, we then estimated the incidence by year of life, by estimating the conditional probability of the first onset in each year of life, *t*, in the subsample of respondents aged at least *t* years and who reported not having had the disorder at Department of Psychiatry and Mental Health and South African Medical Council Research Unit on Risk and Resilience in Mental Disorders, University of Cape Town and Groote Schuur Hospital, Cape Town, South Africa (Prof D J Stein MD); Trimbos-Instituut, Netherlands Institute of Mental Health and Addiction, Utrecht, Netherlands (M ten Have PhD): Center for Excellence on Research in Mental Health. Instituto de Ciencias de la Salud, Medellín, Colombia (Prof Y Torres DraHC); National Institute for Health Services Management, Bucharest, Romania (Prof C Vladescu PhD). Department of Neuroscience, University of Sheffield, Sheffield UK (Prof P W Woodruff PhD); Department of Mental Health,

National Center of Public

| | Morbid risk per 100 age 75 years (95% 0 |) participants at []) | Ratio of morbid risk to lifetime prevalence (95% CI) | | | |
|------------------------------------|--|--------------------------|--|----------------|--|--|
| | Male sex | Female sex | Male sex | Female sex | | |
| Anxiety disorders | | | | | | |
| Panic disorder or agoraphobia | 3.6 (3.2–3.9) | 7·3 (6·8–7·8) | 1.8 (1.7–2.0) | 2.0 (1.9–2.0) | | |
| Generalised anxiety disorder | 6.5 (5.9–7.0) | 12.5 (11.8–13.2) | 2.4 (2.2–2.5) | 2.5 (2.4–2.6) | | |
| Post-traumatic stress disorder | 5.7 (5.1-6.3) | 12.6 (11.7–13.5) | 2.1 (1.9–2.2) | 2·3 (2·2–2·4) | | |
| Social phobia | 4.2 (3.8-4.6) | 6.0 (5.5–6.4) | 1.2 (1.1–1.3) | 1.3 (1.2–1.4) | | |
| Specific phobia | 5.9 (5.3-6.5) | 11.6 (10.9–12.2) | 1.2 (1.1–1.2) | 1.2 (1.1–1.2) | | |
| Any anxiety disorder | 18.3 (17.3–19.4) | 31.0 (29.9–32.2) | 1.6 (1.6–1.7) | 1.7 (1.6–1.7) | | |
| Mood disorders | | | | | | |
| Major depressive disorder | 20.1 (19.2–20.9) | 34.0 (33.2–34.9) | 2.7 (2.6–2.8) | 2.5 (2.4–2.5) | | |
| Bipolar disorder | 6.3 (5.7-6.8) | 6-2 (5-7-6-7) | 2.5 (2.3–2.6) | 2.8 (2.6–2.9) | | |
| Any mood disorder | 24.5 (23.5–25.4) | 37.9 (37.0–38.8) | 2.6 (2.5–2.7) | 2.5 (2.4–2.5) | | |
| Substance use disorders | | | | | | |
| Alcohol abuse | 21.6 (20.6–22.7) | 7.2 (6.7–7.7) | 1.6 (1.5–1.6) | 2.2 (2.1–2.3) | | |
| Alcohol dependence | 6.1 (5.6-6.7) | 2·3 (2·0–2·7) | 1.8 (1.7–1.9) | 2.4 (2.2–2.7) | | |
| Drug abuse | 7.9 (7.2–8.7) | 4.2 (3.8–4.7) | 1.9 (1.8–2.1) | 2.5 (2.3–2.7) | | |
| Drug dependence | 3.0 (2.4–3.5) | 1.6 (1.4–1.9) | 2.1 (1.8–2.3) | 2.5 (2.3–2.8) | | |
| Any substance use disorder | 24.4 (23.3–25.6) | 9.9 (9.2–10.5) | 1.6 (1.5–1.6) | 2·2 (2·1–2·3) | | |
| Externalising disorders | | | | | | |
| ADHD | 3.0% (2.4–3.6) | 1.8% (1.4–2.2) | 1.3% (1.1–1.4) | 1.2% (1.1-1.4) | | |
| Intermittent explosive disorder | 5.9% (5.2–6.5) | 5.2% (4.7–5.8) | 1.7% (1.6–1.8) | 2.1% (1.9–2.2) | | |
| Any externalising disorder | 6.8% (6.0-7.5) | 5.6% (5.0-6.2) | 1.6% (1.5–1.7) | 1.8% (1.7–1.9) | | |
| Any mental disorder | 46.4% (44.9-47.8) | 53·1% (51·9–54·3) | 1.6% (1.6–1.7) | 1.8% (1.8–1.8) | | |

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Health and Analyses, Sofia, Bulgaria (Z Zarkov PhD) Correspondence to: Prof John J McGrath, Queensland Centre for Mental Health Research, Brisbane, QLD 4076, Australia

j.mcgrath@uq.edu.au See Online for appendix For World Bank data see http:// data.worldbank.org/country age t-1 year. We used information only from respondents in the age range between $t\pm 10$ years to minimise recall bias. We then calculated cumulative lifetime disorder risk up to age 75 years (ie, morbid risk) from these incidence data using the standard exponential formula.¹⁴ Smoothing was used with a 5-year bandwidth to reduce instability in estimates.

We calculated standard errors (SEs) of incidence rates and cumulative incidence curves using the jackknife repeated replications simulation method¹⁵ considering both the weighting and geographical clustering of WMH data. In choosing WMH samples with geographical clustering, the typical design began by dividing the population into a series of mutually exclusive and collectively exhaustive geographically clustered strata made up of the equivalent of US counties or metropolitan areas (which could include multiple counties). When a metropolitan area was so populous that it entered the sample with certainty given the sampling fraction, it was referred to as a self-representing stratum. In all other

For the **interactive datavisualisation website** see https://csievert.shinyapps.io/ mental-aoo/





Error bars represent 95% CIs. Incidence of first onset is smoothed to 5-year bandwidths and the hazard rate (ie, incidence rate) curves were calculated per year of age per 10 000 people, defined as the ratio of the number of disorder onsets at an age among people who never had the disorder at any time until that age and who lived through that age. We used information only from respondents in the age range between age of conditional probability of first onset at each year of life (t) up to t+10 years to minimise the effects of recall bias. Cumulative incidence (or morbid risk) curves were calculated on the basis of the age-specific incidence rates at each year of life with the standard exponential formula.¹⁴ cases, a single county or metropolitan area was chosen as the primary sampling unit within each stratum. These non-self-representing primary sampling units were then collapsed into pairs to create pseudostrata, and the primary sampling units within the pseudostrata were used to define sampling error calculation units for the jackknife repeated replications estimation. In the case of self-representing strata, geographically clustered halfsamples were created to define a pair of sampling error calculation units in each stratum.

The approach we used to estimate conditional probability of first disorder onset within specific years of life considered right censoring (ie, the fact that not all respondents had reached age t as of the time of the survey) by excluding (ie, censoring) respondents who were not yet age t years from the denominator. The restriction of estimates to respondents no older than t+10 years reduced the effects of bias due to premature mortality. However, premature mortality bias could still occur if mortality depended on a history of mental disorders and systematic non-response on the basis of factors related to age-specific disorder risks. This possibility of residual bias is inevitable in community surveys, although it can be addressed in population registry data.¹⁶ The direction of this potential bias cannot be assessed rigorously in the absence of information about disorder risk and censoring. However, as people with mental disorders have reduced life spans,17 and people with known histories of being admitted to psychiatric hospitals have been shown to have comparatively low response rates in epidemiological surveys,18 assuming that bias is in the direction of underestimating risk is reasonable.

We compared lifetime prevalence estimates by sex with Wald χ^2 tests and two-sided p values. We show these estimates along with 95% CIs. As WMH survey data are both geographically clustered and weighted, we used the design-based Taylor series linearisation method implemented in SAS version 9.4 to estimate SEs. An interactive data-visualisation website is available; age-specific and sex-specific data underlying the figures can be downloaded from this website.

Missing data at the symptom level were coded conservatively as being absent; subgroup mean imputation was used for other variables.

Role of the funding source

There was no funding source for this analysis.

Results

We obtained data from 32 WMH surveys conducted in 29 countries, including 12 low-income and middleincome countries and 17 high-income countries. These surveys included 156 331 participants (85 308 [54.5%] female respondents and 71 023 [45.4%] male respondents). The overall weighted response rate across all surveys was 63.6% (table 1). 21 (66%) of the 32 surveys

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were based on nationally representative household samples, whereas 11 (34%) were based on regional samples. Ethnicity was not assessed.

Lifetime prevalence of any mental disorder was 28.6% (95% CI 27.9-29.2) for male respondents and 29.8% (29.2-30.3) for female respondents (table 2). Lifetime prevalence of any anxiety disorder was 11.3% (10.9-11.7) for male respondents and 18.8% (18.3-19.2) for female respondents, and of any mood disorder was 9.5% $(9 \cdot 2 - 9 \cdot 7)$ for male respondents and $15 \cdot 4\%$ $(15 \cdot 1 - 15 \cdot 7)$

incidence

Cumulative Median cumulative incidence

А

130

120

110

Male Female 140

Smoothed

hazard rate

for female respondents (table 2). The three mental disorders with highest lifetime prevalence for male respondents were alcohol abuse (13.7%, 13.3-14.1), major depressive disorder (7.5%, 7.2-7.7), and specific phobia $(5 \cdot 0\%, 4 \cdot 8 - 5 \cdot 3)$, and those for female respondents were major depressive disorder (13.6%, 13·3-13·9), specific phobia (10·0%, 9·7-10·2), and PTSD $(5 \cdot 4\%, 5 \cdot 2 - 5 \cdot 7).$

Projected lifetime morbid risk by age 75 years for each mental disorder was higher than the observed lifetime

350

300



В

150

130

400 160

350 140

300 120

Figure 2: Smoothed hazard rates and cumulative incidence by age and sex for any mood disorder (A), any anxiety disorder (B), any substance use disorder (C), and any externalising disorder (D)

Error bars represent 95% CIs. Hazard rate (ie, incidence rate) curves were calculated per year of age per 10 000 people, defined as the ratio of the number of disorder onsets at an age among people who never had the disorder at any time until that age and who lived through that age. We used information only from respondents in the age range between age of conditional probability of first onset at each year of life (t) up to t+10 years to minimise the effects of recall bias. Cumulative incidence (or morbid risk) curves were calculated on the basis of the age-specific incidence rates at each year of life with the standard exponential formula.¹⁴

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⁽Figure 3 continues on next page)

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prevalence at interview. Lifetime morbid risk of any mental disorder as of age 75 years was $46 \cdot 4\%$ ($44 \cdot 9-47 \cdot 8$) for male respondents and $53 \cdot 1\%$ ($51 \cdot 9-54 \cdot 3$) for female respondents (table 3). The three disorders with highest lifetime morbid risk for male respondents were alcohol abuse ($21 \cdot 6\%$, $20 \cdot 6-22 \cdot 7$), major depressive disorder ($20 \cdot 1\%$, $19 \cdot 2-20 \cdot 9$), and drug abuse ($7 \cdot 9\%$, $7 \cdot 2-8 \cdot 7$), and those for female respondents were major depressive disorder ($34 \cdot 0\%$, $33 \cdot 2-34 \cdot 9$), PTSD ($12 \cdot 6\%$, $11 \cdot 7-13 \cdot 5$), and generalised anxiety disorder ($12 \cdot 5\%$, $11 \cdot 8-13 \cdot 2$).

The minimum, maximum, median, and IQR of cumulative lifetime risk as of age 75 years are in the appendix (p 5). For the incidence of first onset of any disorder, peak incidence was approximately at age 15 years. At age 15 years, male respondents had higher incidence (hazard rate $288 \cdot 0$ [95% CI $263 \cdot 7-312 \cdot 3$] per

10000 participants) than female respondents (228.2 [211.8–244.6] per 10000 participants; figure 1). Across the rest of the lifespan, incidence was mostly slightly higher among female than male respondents. Cumulative risk curves (per 1000 people) were higher across the lifespan for female than male respondents. For any mental disorder, the median age of onset of the first disorder was 19 years (IQR 14–32) for male respondents and 20 years (12–36) for female respondents.

The ratio of morbid risk to lifetime prevalence shows the relatively high proportion of mental disorders that first occur in childhood, adolescence, or young adulthood (table 3). Disorders with earlier onsets have ratios closer to 1, whereas disorders with a wider onset distribution have ratios greater than 1. For male respondents, the lowest morbid risk to lifetime prevalence ratios were for



(Figure 3 continues on next page)

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ADHD, social phobia, and specific phobia, and the highest ratio was for major depressive disorder. For female respondents, the lowest ratios were for ADHD, social phobia, and specific phobia, and the highest ratio was for bipolar disorder.

The distributions of incidence rate and cumulative incidence differed when disaggregated by broad general disorder categories (figure 2) or specific disorder types (figure 3). For example, any externalising disorder shows an early peak incidence in the first decade of life, whereas incidence of any substance use disorder peaks around age 15 years. Both types of disorder were more common among male respondents than female respondents. Mood disorders and anxiety disorders were more common among female respondents than male respondents across the lifespan, but the peak incidence of any anxiety disorder was at age 5 years, compared with age 15 years for any mood disorder. The Wald χ^2 test compared the lifetime prevalence for each specific disorder and broad general category (table 2), showing that mood and anxiety disorders were more common among female respondents and substance use disorders and externalising disorders were significantly more common among male respondents.



Figure 3: Smoothed hazard rate and cumulative incidence by age and sex for 13 specific disorders

Error bars represent 95% CIs. Hazard rate (ie, incidence rate) curves were calculated per year of age per 10 000 people, defined as the ratio of the number of disorder onsets at an age among people who never had the disorder at any time until that age and who lived through that age. We used information only from respondents at age of conditional probability of first onset in the age range between each year of life (t) up to t+10 years to minimise the effects of recall bias. Cumulative incidence (or morbid risk) curves were calculated on the basis of the age-specific incidence rates at each year of life with the standard exponential formula.¹⁴

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The WMH surveys were conducted during a wide range of years. To assess whether the results differed depending on time of survey, we replicated all analyses for broad classes of disorders and any disorder separately for the 81342 respondents in the 2007 paper and the 74989 additional respondents from more recent surveys. Results showed broad consistency in relative prevalence across disorders and in comparing male versus female respondents (appendix p 6). There was broad consistency in morbid risk to lifetime prevalence ratios between the earlier and more recent surveys (appendix p 7), as well as for age of onset distributions (appendix p 8–11).

Discussion

Based on WMH surveys in 29 countries, we estimated that by age 75 years, about one in two individuals will develop at least one of the mental disorders considered. We found that the incidence of anxiety and mood disorders was higher in female respondents, and that the incidence of externalising disorders was higher in male respondents. Our findings are based on more accurate estimates of age-of-onset distributions than previous studies, and they support the idea that many mental disorders have their first onset during childhood, adolescence, or young adulthood and that some disorders have earlier ages of onset than others.

Most studies of age-of-onset distributions have used data from register-based studies of treated mental disorders,19 and both commonalities and differences can be seen between those studies and our population-based findings. First, as expected, the lifetime morbid risk of any treated mental disorders based on register data is lower than our estimates. This difference can be explained by the finding that many people with mental disorders never receive treatment. $^{\scriptscriptstyle 20}$ Second, because of the well documented delays in seeking treatment after the first onset of mental disorders,²¹ the age-of-onset distributions for register-based studies tend to be right-shifted (ie, delayed) compared with populationbased studies. This bias is sex dependent, as female respondents are more likely than male respondents to seek treatment and to do so more quickly.²²

A key finding is the substantial proportion of mental disorders that have early first onsets. We found that half of people who develop a mental disorder before age 75 years have their first onset by age 19 years for male respondents or 20 years for female respondents. In addition to the traditional childhood-onset disorders (eg, ADHD, social phobia, and specific phobia), common mental disorders (eg, major depressive disorder, generalised anxiety disorder, panic disorder, and drug use disorders) were found often to have their first onsets between childhood and early adulthood. This observation supports the need to invest in mental health services that have a particular focus on young people.²³ Although the median ages of onset for many disorders in male respondents and female respondents were similar, our

findings indicate sex differences in lifetime prevalence for each of the mental disorders examined, with anxiety disorders and major depressive disorder more common among female respondents compared with impulse control and substance use disorders, which were more common among male respondents. These differences are consistent with previous studies.²⁴

This study has important limitations. First, surveys were conducted over more than two decades, although there was good consistency of results. Second, all data were based on retrospective reports. Recall bias increases with the length of recall and can lead to under-identification of more temporally distant events,⁷ as shown in birth cohort studies.6 We based our estimates on onsets within 10 years of interview to attempt to minimise this bias. Third, our definition of disorder onset was based on single questions rather than detailed diagnostic assessments for first recalled occurrence, leading to possible downwardly biased estimates of age of onset. Fourth, the surveys could have been biased due to differential response. Fifth, although the sample was large enough to generate relatively precise, pooled disorder-specific estimates, it was not large enough to investigate between-country differences. Sixth, not all mental disorders were included in the CIDI. Finally, we did not consider comorbidity. There is increasing understanding that comorbidity is common within different types of mental disorders (eg, individuals with one type of mental disorder are at increased risk of subsequently developing other types of mental disorders)25-27 and that the burden of illness is strongly influenced by comorbidity. Although our analysis provided lifetime estimates for any mental disorder and for specific types of disorders, we did not consider patterns of comorbidity and how they change during the life course.26 We hope to investigate this issue in future studies.

We provided updated estimates of age-of-onset distributions, lifetime prevalence, and morbid risk for a range of mental disorders using improved methods to reduce the effects of recall bias. These estimates will be of value to service planners, researchers interested in the burden of disease, and genetic epidemiologists.

Contributors

JJM and RCK conceptualised and designed the analysis. NAS and RCK supervised the analysis. WTC and NAS analysed the data. JJM, RCK, AA-H, JA, YA, LHA, RB, JMCdA, SC, LD, FF, OG, JMH, EGK, GK, SMK, VK-M, MM, JM, FN-M, DN, JP-V JCS, DJS, CV, PWW, and ZZ provided data. All authors had access to all data for this analysis; JJM, RCK, NAS, and WTC directly accessed and verified the data. JJM and RCK drafted the initial manuscript. All authors reviewed the manuscript and were responsible for the decision to submit for publication.

Declaration of interests

LD receives educational grants from Indivior and Seqirus. OVD receives funding from Kowa Research Institute and has been an advisor in the PROMINENT trial. SMK and PWW receive grant funding from Hamad Medical Corporation through the Cambridgeshire and Peterborough National Health Service Foundation trust and from Qatar University. PWW has received financial support from the Qatar National Research Fund and an honorarium from Gresham College. RCK has been a consultant for Cambridge Health Alliance, Canandaigua Veterans Affairs

Medical Center, Holmusk, Partners Healthcare, RallyPoint Networks, and Sage Therapeutics. He holds stock options in Cerebral, Mirah, Prepare Your Mind, Roga Sciences, and Verisense Health. DN receives honoraria from AIG General Insurance and Takeda Pharmaceutical and financial support from Startia, En-power, and MD.net. DJS receives royalties from American Psychiatric Press, Cambridge University Press, and Elsevier–Academic Press. He has received honoraria from Discovery Vitality, Johnson & Johnson, Kanna, L'Oreal, Lundbeck, Orion, Sanofi, Servier, Takeda, and Vistagen. He was the president of the African College of Neuropsychopharmacology and is financially supported by the Medical Research Council of South Africa. All other authors declare no competing interests.

Data sharing

Access to the cross-national World Mental Health (WMH) data is governed by the organisations responsible for funding and survey data collection in each country. These organisations made data available to the WMH survey consortium through restricted data sharing agreements that do not allow data to be released to third parties. US data are available for secondary analysis via the Inter-University Consortium for Political and Social Research (http://www.icpsr.umich.edu/icpsrweb/ ICPSR/series/00527). A complete list of all within-country and crossnational WMH publications is available (http://www.hcp.med.harvard. edu/wmh/). No additional data or materials will be made available.

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