

# Effect of contributing factors on the incidence of non-communicable diseases among adults with common mental health disorders: a systematic review

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### Abstract

**Background:** Individual, social and environmental factors play a dynamic role in determining mental health outcomes. The linkage between mental health and non-communicable disease is widely noted, but the mechanisms are poorly understood. The current systematic review aims to identify common contributing factors linking mental health to non-communicable disease incidence among adults to inform planned preventive interventions for high-risk noncommunicable disease and mental ill-health populations.

**Methods:** MEDLINE, PsycINFO, EMBASE and CINAHL were searched from February to August 2019 for casecontrol and longitudinal studies of adults with common mental health disorders (depression and anxiety) assessing the causal effect of individual, environmental and social factors on the incidence of common non-communicable diseases (cancers, cardiovascular diseases, chronic obstructive pulmonary disease and diabetes mellitus). There were no geographical restrictions for the selected studies and the results were generated utilising a narrative synthesis.

**Results:** Of 15,266 unique documents identified by search terms, 419 met criteria for full-text review and 11 studies met inclusion criteria for data extraction. None of the identified studies had the onset of chronic obstructive pulmonary disease as an outcome. The majority of the studies showed a significant effect of depression and/or post-traumatic stress disorder on non-communicable disease incidence. The chronicity of mental health disorders appears to exacerbate their effect on non-communicable disease onset. Older age, higher body mass index, female sex, smoking status and number of cigarettes smoked, low educational attainment and ethnicity were found to exert a significant effect on the association between mental health disorders and non-communicable disease onset.

**Conclusion:** Findings from this review provide evidence to guide health practice and policy to reduce the burden of non-communicable diseases and mental health disorders. Interventions targeted towards identifying and treating mental health illnesses and the factors linking mental and physical health lower the burden of mental disorders and prevent the subsequent development of non-communicable diseases.

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### Introduction

Worldwide, about 74% of all deaths are attributed to non-communicable diseases (NCDs), of which more than three-quarters occur in low- and middleincome countries (LMICs).<sup>1</sup> NCD prevention and control has mainly focused on the four conditions responsible for the greatest number of deaths. Known as the 'Big Four', cardiovascular diseases (CVDs), followed by cancers, chronic respiratory diseases and diabetes, comprise the highest rates of mortality as they contribute to 33.3 million deaths globally. These NCDs are all driven by

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modifiable behaviours, namely, unhealthy diets, physical inactivity, tobacco smoking and harmful use of alcohol.<sup>2,3</sup> These behavioural determinants are widely recognised as the main entry points for the prevention and control of NCDs.<sup>4</sup>

According to the American Psychiatric Association, mental disorders, also called mental illnesses, are classified as any condition characterised by cognitive and emotional disturbances, abnormal behaviours, impaired functioning or any combination of these.<sup>5</sup> The World Health Organization (WHO) identifies depressive disorders and anxiety disorders, including post-traumatic stress disorder (PTSD), as the two main diagnostic categories comprising common mental health disorders.<sup>6</sup> In 2012, the WHO identified three different categories of risks affecting an individual's mental health, namely, individual attributes and behaviours, social and economic circumstances and environmental factors.<sup>7</sup> Although the WHO called these factors 'risks to mental health', they are similar to the social determinants of mental health that were proposed later by Marmot et al. in 2014.8 At the individual and behavioural level, poor dietary habits<sup>9</sup> and smoking<sup>10</sup> were found to be associated with depression. In addition, socioeconomic conditions, such as unemployment and financial hardships, predispose individuals to mental illnesses.<sup>7</sup> According to the WHO, the poorest segment of society is often the most susceptible to poor mental health.<sup>11</sup> The stress that accompanies absolute poverty is a powerful risk factor for mental disorders such as depression and substance dependence.<sup>12</sup>

Common mental illnesses and NCDs often coexist. Systematic reviews linked cancer with anxiety disorders<sup>13,14</sup> and PTSD.<sup>15</sup> Meta-analyses linked diabetes with a range of mental disorders, including depression and PTSD.<sup>16</sup> Across countries, the odds ratios for the association of heart disease with mood disorders, anxiety disorders and alcohol dependence were found to be 2.1, 2.2 and 1.4, respectively.<sup>17</sup>

The exact mechanism linking common mental disorders (CMDs) to NCDs remains to be clearly understood. It has been suggested that they influence each other directly through physiological systems (e.g. neuroendocrine system). For instance, depression is shown to provoke a cascade of events in the hypothalamic-pituitary-adrenal axis, which is normally stimulated in response to stress, leading to insulin resistance and eventually to type 2 diabetes.<sup>18</sup> On the other hand, there is evidence associating individual, social and environmental factors, known to be determinants of mental health, with NCDs. Individual factors that were mentioned earlier to be associated with

depression, namely, smoking and poor dietary habits, are known to be the main entry points for the prevention and control of NCDs.<sup>4</sup> Poverty, a social determinant of mental health,<sup>19</sup> is shown to exert a similar effect on NCD incidence. In a large-scale population cohort in the UK, it was found that the segment of the population who were more socioeconomically deprived had an increased risk for developing heart disease (incidence rate ratio 1.61, 95% CI 1.58 to 1.64) and did so earlier in life than those from the most affluent group (adjusted difference -3.51 years, 95% CI -3.77 to -3.25). Factors such as health inequity, poor access to basic services such as clean water and electricity, political instability, discrimination and armed conflicts are just examples of environmental determinants of mental health<sup>7</sup> that were also found to be associated with higher rates of diabetes and CVDs<sup>20</sup> and with an increased incidence of cancer.<sup>21</sup>

Physical health outcomes for people living with a chronic physical condition are worse in the presence of a mental illness, including a reduced quality of life and a shortened lifespan. Studies show that mental illness contributes to an increased risk of mortality from CVDs and cancer.<sup>22-24</sup> According to the World Mental Health Surveys, an 8–12% increase in mortality, due to smoking, diabetes, history of myocardial infarction and hypertension, was found among people with common mental health disorders.<sup>17</sup> Figures of myocardial infarctions, a NCD complication, are higher among individuals subjected to stressful situations such as armed conflicts or natural disasters compared to pre-emergency figures.<sup>25</sup>

Despite the supportive evidence that links CMDs and their determinants to NCDs, and despite the fact that physical health outcomes get worse for people living with a NCD in the presence of a mental illness, the connection between them is usually overlooked in clinical settings and, to a great extent, in country-level policy-making. This has been attributed to factors such as limited government capacities for policy development, weak health systems and insufficient finances, among other factors.<sup>26,27</sup> In 2005, member states in the WHO European Region endorsed the 'no health without mental health' statement owing to mental health's contribution to years of disability.<sup>28</sup> The WHO's Global Action Plan (2013–20), to reduce the global burden of non-communicable illnesses and preventable mortality, includes mental disorder prevention and control objectives.<sup>29</sup> However, mental and physical disease prevention initiatives remain essentially separate and independent from one another.<sup>30</sup> The current systematic review aims to address the link between mental illness and NCDs by exploring the aforementioned individual, social and environmental

determinants associated with both conditions. It aims to answer the research question: 'What are the individual, social and environmental determinants that lead to the development of NCDs {CVD, cancers, respiratory diseases [including chronic obstructive pulmonary disease (COPD)] and diabetes mellitus [DM]} among patients with common mental health disorders (depression and anxiety disorders including PTSD)?' Considering these overlapping determinants may be a practical and effective way of providing integrated prevention programmes to reduce the burden of these conditions as well as the associated disability. This review is part of an entire PhD research work on the social determinants of mental health and how they affect population health.

### **Methods**

This review was conducted in accordance with a published protocol prepared by the authors and published on PROSPERO with a registration number CRD42019157800.<sup>31</sup>

The review follows the latest Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) 2020 reporting guidelines.<sup>32</sup>

### Inclusion and exclusion criteria

The current review utilised the population, intervention/ exposure, control, outcome and study (PICOS) design format as a systematic and rigorous method for delineating inclusion and exclusion criteria (Table 1). This approach ensures coherence and uniformity in the item selection process.<sup>33</sup> This review included observational studies (case-control, retrospective and prospective cohort studies) only as they can provide valuable evidence for establishing a causal relationship between common mental health disorders and NCDs. Cross-sectional studies were excluded because, unlike cohort studies, they did not measure the incidence of disease and, unlike casecontrol studies, could not measure the effect of multiple exposures on disease outcome.<sup>34</sup> The exclusion of interventional studies was deemed necessary due to their limited suitability as the primary approach for evaluating the incidence of NCDs. Experimental studies can offer valuable insights into the efficacy of interventions in preventing and managing NCDs. However, in the current review, we did not prioritise examining such aspects.

In the included studies, CMDs were identified via an established clinician-administered instrument [e.g. Structured Clinical Interview for *Diagnostic and Statistical Manual of Mental Disorders* (SCID) and the Composite

International Diagnostic Interview (CIDI)], clinical records and/or using a validated self-report instrument [e.g. Patient Health Questionnaire-9 item (PHQ-9) and Generalised Anxiety Disorder-7, Hospital Anxiety and Depression Scale]. There are two main references to classify mental disorders: the International Classification of Disease (ICD) published by the WHO and the Diagnostic and Statistical Manual of Mental Disorders (DSM) published by the American Psychiatric Association, the most recent versions of which are ICD-11 and DSM-5, respectively. DSM is primarily used by clinicians for diagnosis, while ICD is primarily used for the coding of diseases in clinical settings and for epidemiological purposes. The listing of conditions differs between the two. For example, the ICD-11 lists depressive disorders under mood disorders, whereas the DSM-5 lists depressive disorders as a separate category.<sup>35,36</sup> The classification of conditions is routinely revised. For example, PTSD was previously classified under anxiety disorders in DSM-4 and is currently listed in DSM-5 under 'Trauma- and Stressor-Related Disorders'.<sup>37</sup>

The incidence of the NCD conditions, on the other hand, was based on either self-report, clinical or biochemical assessments.

We excluded studies that involved participants with comorbid NCDs, given that NCDs were the outcome and each could independently be linked to the incidence of the other. For example, studies have shown that there are biological mechanisms associated with DM that increase the risk for CVD among diabetic patients.<sup>38</sup> Also, we excluded studies that did not involve participants who have CMDs, namely, depression and/or anxiety disorders. Studies reporting on the incidence of NCDs without reporting the exposure (individual, social and/or environmental determinants) leading to the development of NCDs were excluded.

### Search criteria

Our searches were restricted to English. There were no geographical restrictions and no time frame limits. One author (ZJ) searched MEDLINE, PsycINFO, EMBASE and CINAHL from database inception to August 2019 using a search strategy developed and validated for the right use of Boolean phrases and medical subject heading (MeSH) terms with the support of a specialist librarian. We searched each database by combining four different concepts related to NCDs along with three different concepts of mental health disorders, namely, 'cardiovascular diseases', 'diabetes', 'cancers', 'chronic obstructive pulmonary disease' with 'anxiety', 'depression' and 'post-traumatic stress disorder'. The search encompassed the combination of various terms for each

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#### TABLE 1 Title and abstract screening tool following the PICOS format

	Inclusion criteria	Exclusion criteria
Participants	Participants ≥ 18 years old With common mental health problem	Participants < 18 years old Participants with comorbid mental health and NCD conditions Participants with pre-existing NCD Participants with severe mental health illness
Exposure	<ul> <li>Reports at least one outcome of the following:</li> <li>Individual determinants such as BMI and/or dietary intake, physical inactivity, drug abuse, harmful use of alcohol, low self-esteem, criminal/anti-social behaviour, treatment abiding/ seeking behaviour</li> <li>Social circumstances such as bereavement/loneliness, socio-economic status/position, family/neighbourhood cohesion or support, marital status, education level, exposure to violence/ abuse, low income/debt/poverty, unemployment, work stress</li> <li>Environmental factors such as poor civic amenities, injustice, discrimination/social inequalities, gender inequalities, exposure to war or disaster</li> </ul>	Reports on any of these: quality of life, satisfaction with life, happiness
Control	Participants who do not develop one of the four NCDs	Not applicable
Outcome	Incidence of common NCDs (diabetes, cancers, COPD and CVD)	Incidence of illnesses other than common NCDs Comorbidity of NCDs with mental health problem
Study design	Observational studies (cohort studies and case-control studies)	Cross-sectional studies Experimental studies Interviews/focus groups/case studies Review articles and meta-analysis Reports (conference abstracts, study registrations, study protocol papers, result reports, policy briefs, book chapters, dissertations and theses, editorials, commentaries and website content)

concept and included both free text words and MeSH terms, utilising relevant search options corresponding to each resource. A sample of the electronic search strategy is available in *Appendix* 1.

#### **Study selection**

Zeina Jamal imported the retrieved studies into EndNote to remove duplicates. After removing all duplicates, references were then imported to Excel for title and abstract screening. Another reviewer (MR) screened 10% of the titles and abstracts of all potentially eligible articles using a checklist for inclusion and exclusion criteria (*Table 1*). A list of full-text articles was identified. A third reviewer (AN) was available for mediation throughout the full-text screening and the data extraction phase. Disagreements were resolved via group consensus.

### **Data extraction**

For each article, the first author (ZJ) extracted all data according to a pre-specified template (see *Appendix 2*). The main variables covered relate to study identifiers, context of the study and aims, data collection details, measurement of main outcome and important findings.

#### **Synthesis**

The process of the synthesis was guided by the synthesis without meta-analysis guidelines.<sup>39</sup> For initial syntheses of the studies, they were grouped based on the NCD outcome (Tables 2-5). Based on individual study appraisals of quality, studies were further grouped into categories of high, medium and low quality. All available evidence, regardless of quality of evidence, was initially collated, after which information from high-quality studies only was synthesised. An overview of all relevant exposures (the determinants) across all outcomes was then collated, with subsequent focused synthesis on studies of high quality only. A narrative synthesis, rather than meta-analysis, was used to consider the association between the different determinants in relation to the incidence of the NCD outcomes, given the diverse nature of the latter and the numerous determinants that were studied. Other sources of heterogeneity are described below.

### Heterogeneity

Several sources of heterogeneity are present within this systematic review. Studies included utilised different tools for assessing mental disorders, and the results themselves are presented differently [i.e. hazard ratio (HR), relative

et al. (2010)**       Epidemiologic Catchment       pro- spec- spec- tudinal       61.2% (MDE and dysphoric episode)       MDE dysphoric episode)       MDE Altaus       status cancer tudinal       Colon, of cancer status       MDE diagnosis tudinal       MDE HR 1.87 (1.16 certificates       45 yeas); HR 1.03 Cl (1.02 to 1.04) for overall cancer Smoking status         VSA       longi- tudinal       tudinal       Alcohol Marital status       Gender Ling, tudinal       longi- tudinal       HR 3.43 Cl 4.30 to 273.74) for lung cancer         VSA       longi- tudinal       HR 4.4 (1.08 to to 273.74) for lung cancer       IR 4.4 (1.08 to to 273.74) for lung cancer         VSA       HR 4.8 (1.98 to 23.90) for prostate cancer* Dysphoric episode       HR 4.88 (1.98 to 23.90) for prostate cancer* Dysphoric episode       HR 1.69 (1.3 to 2.19) for overall cancer         HR 1.42 (1.17 to 4.98) for       SES       SES       SES       SES	Study	Country/name of the study	Design	Duration of follow- up	Type of assessed MH	Measurement of MH condition	Covariates	Outcome	Measurement of outcome	Association of depression/ anxiety with overall incident cancer and with cancer subtypes (95% CI)	Important results/ highest risk group	CASP quality assessment
	Gross <i>et al.</i> (2010) <sup>40</sup>	Epidemiologic Catchment Area Study,	pro- spec- tive longi-	24 years	(MDE and dysphoric		status Age Gender Ethnicit Alcohol depen- dence Marital status	(colon, prostate, lung, skin and	of cancer diagnosis plus death	MDE HR 1.87 (1.16 to 3.01) for overall cancer DIS/DSM-III MDE HR 4.4 (1.08 to 17.6) for breast cancer <sup>a</sup> DIS/DSM-III MDE (single episode) HR 6.88 (1.98 to 23.90) for prostate cancer <sup>b</sup> Dysphoric episode HR 1.69 (1.3 to 2.19) for overall cancer Dysphoric episode HR 2.42 (1.17 to 4.98) for breast cancer <sup>a</sup> Dysphoric episode HR 2.42 (1.17	45 years): HR 1.03 CI (1.02 to 1.04) for overall cancer Smoking status HR 34.3 CI (4.30 to 273.74) for lung	Medium

### TABLE 2 Characteristics of selected studies linking depression with incident cancer

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Study	Country/name of the study	Design	Duration of follow- up		Type of assessed MH	Measurement of MH condition	Covariates	Outcome	Measurement of outcome	Association of depression/ anxiety with overall incident cancer and with cancer subtypes (95% CI)	Important results/ highest risk group	CASP quali
Penninx et al. (1998) <sup>41</sup>	Established Populations for Epidemiologic Studies of the Elderly, USA	Cohort pro- spec- tive longi- tudinal	10 years	4825; F 64.6%	Depression (chronic depression)	CES-D scale	<ul> <li>Age</li> <li>Gender</li> <li>Alcohol intake</li> <li>Smoking</li> <li>Ethnicity</li> <li>Physical disability</li> <li>Hospital admis- sions</li> </ul>	Cancer	Medicare hospitalisa- tion records plus death certificates	Chronic depression HR 1.88 (1.3 to 3.14) for overall cancer Chronic depression HR 4.8 (1.01 to 22.79) for uterus, adnexa cancers <sup>a</sup>	Not chronically depressed mood X smoking: Ex-smoker HR 1.65 CI (1.29 to 2.1)	High
											Current smoker HR 1.96 Cl (1.16 to 3.34) Chronically depressed mood X smoking: Ex-smoker HR 2.17 Cl (0.68 to 6.88) Current smoker HR 2.8 Cl (0.65 to 11.94)	
Linkins et al. (1990) <sup>52</sup>	Household survey Washington County, USA;	Cohort pro- spec- tive longi- tudinal	12 years	2264; F 58%	Depression	CES-D scale	<ul><li>Gender</li><li>Smoking</li></ul>	Cancer	Examination of records of the Washington County Cancer Register	Depressed mood at initial survey No (reference) Yes RR 1.09 (0.69 to 1.71)	Depressed mood present Current smokers RR 2.6 Cl (1.41 to 4.80) Current smoker 1–14 cigarettes/day RR 0.75 Cl (0.10 to 5.46)	Low

**TABLE 2** Characteristics of selected studies linking depression with incident cancer (continued)

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CASP, Critical Appraisal Skills Programme; CES-D, Centre for Epidemiological Studies Depression Scale; DIS, Diagnostic Interview Schedule; HR, hazard ratio; MDE, major depressive episode; MMPI, Minnesota Multiphasic Personality Inventory; RR, Relative Risk; SES, Socio-economic status.

a Analysis for women only.

b Analysis for men only.

c Cancer sites associated with smoking are buccal cavity and pharynx, pancreas, larynx, bronchus and lung, cervix.

Study	Country/name of the study	Design	Duration of follow- up		assessed	Measurement of MH condition	Covariates	Outcome	Measurement of outcome	Association of depression/ anxiety with overall incident cancer and with cancer subtypes (95% CI)	Important results/ highest risk group	CASP qualit
											15-24 cigarettes/ day RR 2.6 Cl (1.11 to 6.07) ≥ 25 cigarettes/day RR 4.47 Cl (1.88 to 10.60) Depressed mood present ≥ 25 cigarettes/day RR 18.47 Cl (4.58 to 74.41) for cancer sites associated with smoking <sup>c</sup>	
Persky <i>et al.</i> (1987) <sup>53</sup>	Western Electric Study, USA	Cohort pro- spec- tive longi- tudinal	20 years	2017; F 0%	Depression	MMPI	<ul> <li>Age</li> <li>Smoking (# of cig- arettes/ day)</li> <li>Alcohol (ml/day)</li> <li>Occu- pational status</li> <li>Family history of cancer</li> <li>BMI</li> <li>Serum choles- terol</li> </ul>	Cancer	Self-report of cancer diagnosis plus death certificates	High D profile proportional Hazards coefficient 0.3188 p = 0.055	Age coefficient 0.0885 <i>p</i> < 0.001 Cigarettes (#/day) 0.0129 <i>p</i> = 0.035	Low

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of	selected stud	ies linking d	lepression	and/or PTSE	) with incident as	sthr	na				
9	Design	Duration of follow- up		Type of assessed MH	Measurement of MH condition	Ca	ovariates	Outcome	Measurement of outcome	Adjusted association of depression/ anxiety with overall incident asthma (95% CI)	Important results/ highest risk group
	Prospective cohort design	20 years	3614; F 64.1%	Depression	CES-D scale	• • •	Age Gender Race Education level PA Smoking BMI	Asthma	A new report of asthma medication use and/or self-reported provider diagnosis of asthma	Baseline depression RH 1.26 (1.02 to 1.56) Cumulative exposure to depression <sup>a</sup> RH 1.15 (1.02 to 1.29)	Gender Female RH 1.93 (1.55 to 2.41) BMI (< 25 reference) BMI = 25-29 HR 1.19 (0.95 to 1.50) BMI = 30+ RH 1.41 (1.10 to 1.81)
											Education ( <hs is<="" td=""></hs>

### TABLE 3 Characteristics of selected stud

Country/name

of the study

The Coronary

Development design

Artery Risk

in Young Adults (CARDIA) Study, USA

Study

et al. (2014)<sup>42</sup>

Brunner

ω

CASP

High

reference) HS

RH 0.74 (0.51 to 1.09) Some college RH 0.64 (0.43 to 0.95) College or more RH 0.66 (0.44 to 0.99)

quality

assessment

Study	Country/name of the study	Design	Duration of follow- up		assessed	Measurement of MH condition	Covariates	Outcome	Measurement of outcome	Adjusted association of depression/ anxiety with overall incident asthma (95% CI)	Important results/ highest risk group	CASP quality assessment
De La Hoz <i>et al.</i> (2016) <sup>43</sup>	World Trade Centre Worker and Volunteer Medical Screening Program, USA	Prospective cohort design	Mean = 4.95 years	3757; F 16.15%	PTSD	PTSD checklist questionnaire	<ul> <li>Gender</li> <li>Age</li> <li>Race/ethnicity</li> <li>BMI</li> <li>Weight gain</li> <li>Education level</li> <li>Occupational exposure</li> </ul>	Asthma	Self-report of new physician- diagnosed asthma	OR 2.41 (1.85 to 3.13) Recovering PTSD OR 1.34 (0.81 to 2.23) Late-onset PTSD OR 3.07	Gender (male as reference) Female OR 1.38 (1.03 to 1.85) Race/ethnicity (white is reference) Latino OR 0.89 (0.69 to 1.16) African American OR 0.62 (0.42 to 0.91) Asian OR 1.32 (0.63 to 2.74) Other OR 3.58 (1.52 to 8.46) BMI (normal is reference) Overweight OR 1.1 (0.79 to 1.54) Obese OR 1.42 (1.01 to 2.00)	Medium

CASP, Critical Appraisal Skills Programme; CES-D, Centre for Epidemiological Studies Depression Scale; HS, High school; OR, Odds ratio; PA, Physical activity; RH, Relative Hazard. a Cumulative exposure model of depression is derived by allowing depressive symptom status to vary by time taking into account the number of instances of elevated symptoms before asthma onset.

b Definitions of PTSD categories: baseline PTSD (PTSD at baseline study visit), recovering PTSD (PTSD at baseline visit but not at the follow-up visit), late-onset PTSD (PTSD not present only at the follow-up visit), chronic PTSD (PTSD present at both visits).

Study	Country/name of the study	Design	Duration of follow- up		assessed	Measurement of MH condition		ovariates	Outcome	Measurement of outcome	Adjusted association of depression/PTSD with overall incident asthma (95% CI)	results/ highest risk	CASP quality assessment
Boyko <i>et al.</i> (2010) <sup>44</sup>	Seattle Epidemiologic Research and Information Centre, Department of Veterans Affairs Puget, Washington, USA	Prospective cohort longitudinal	3 years	44,754; F 25.8%	Depression and PTSD	Self- administered PHQ and PTSD check- list civilian version	• • • • •	Ethnicity Education	DM	Self-report of physician diagnosis	PTSD OR 2.07 (1.31 to 3.29)	Age OR 1.06 (1.05 to 1.08) BMI OR 1.2 (1.17 to 1.23) Race/ ethnicity (Caucasian is reference) African American OR 1.46 (1.10 to 1.94) Hispanic OR 1.35 (0.89 to 2.05) Asian OR 1.67 (1.11 to 2.52) Other OR 1.07 (0.52 to 2.19) Separated from military as of follow-up (No is reference) Yes OR 2.18 (1.61 to 2.93) Component (Reserve/ (Guard is reference)	

This article should Jamal Z, Hom R, A online ahead of pi				
This article should be referenced as follows: Jamal Z, Hom R, Ager A. Effect of contributir online ahead of print May 22 2024). Global	Study	Country/name of the study	Design	Duration of follow- up
This article should be referenced as follows: Jamal Z. Hom R. Ager A. Effect of contributing factors on the incidence of non-communicable diseases among adults with common mental health disorders: a systematic review [published online ahead of print May 22 2024]. Global Health Res 2024. https://doi.org/10.3310/CTDH4677	Mezuk et al. (2008) <sup>45</sup>	Sound Health Care System, Seattle, Washington	Prospective cohort longitudinal	23 years
<u>a</u>				

<ul> <li>2.3 years</li> <li>1071; Depression DIS</li> <li>Age</li> <li>Age&lt;</li></ul>	:ASP uality ssessment	highest risk	association of depression/PTSD with overall incident asthma (95% CI)	Measurement of outcome	Outcome	Covariates	Measurement of MH condition	Type of assessed MH		Duration of follow- up
79.3%• Gendereach wave;(1981-2005)*stratified byIwith• EducationexcludingHR 1.62 (1.03 toeducationalMDD• Racediabetes cases2.55)attainment*61.7%• BMIthat began(1993-2005)Collegeno• Smokingwithin 1 yearHR 2.04 (1.09 todegree or		OR 0.74 (0.56 to 0.98) Depression × gender (Non- depressed men OR 1.14 (0.6 to 2.15) Non- depressed women OR 1.63 (1.26 to 2.13) Depressed women OR 0.51								
onset and HR 0.45 before the age (0.11 to 1.80) of 30 High school or less HR 2.26 (1.11 to 4.6)	1edium	stratified by educational attainment <sup>c</sup> College degree or more HR 0.45 (0.11 to 1.80) High school or less HR 2.26	(1981-2005) <sup>a</sup> HR 1.62 (1.03 to 2.55) (1993-2005) HR 2.04 (1.09 to	each wave; excluding diabetes cases that began within 1 year of depression onset and before the age	DM	<ul><li>Gender</li><li>Education</li><li>Race</li><li>BMI</li></ul>	DIS	Depression	79.3% with MDD 61.7% no	e 23 years
cont	continued									

Adjusted

		Selected S		uepressio				continueu/					
Study	Country/name of the study	Design	Duration of follow- up		assessed	Measurement of MH condition		ariates	Outcome	Measurement of outcome	Adjusted association of depression/PTSD with overall incident asthma (95% CI)	results/ highest risk	CASP quality assessment
							• • • •	Alcohol use Stairs climbed per day Frequency of eating bal- anced meals Social contact with relatives				MDD x college degree or more (reference) <sup>d</sup> MDD × high school or less HR 4.1 (1.84 to 9.16) No MDD × college degree or more HR 0.92 (0.27 to 3.15) No MDD × high school or less HR 1.44 (0.82 to 2.55)	

TABLE 4 Characteristics of selected studies linking depression and/or PTSD with incident DM (continued)

CASP, Critical Appraisal Skills Programme; DIS, Diagnostic Interview Schedule; MDD, major depressive disorder.

a Adjusted for age, race, gender, education, smoking status, alcohol use, social network size and antidepressant use.

b Adjusted for age, race, gender, education, smoking status, alcohol use, social network size, antidepressant use, BMI, family history of diabetes, stairs climbed per day and frequency of contact with relatives.

c Adjusted for age, race, gender, smoking status, alcohol use, BMI and family history of diabetes.

d Adjusted for age, race, gender, smoking status, alcohol use, BMI, family history of diabetes and social network characteristics.

Study	Country/name of the study	Design	Duration of follow- up		Type of assessed MH	Measurement of MH condition	Covariates	Outcome	Measurement of outcome	Adjusted association of depression/PTSD with overall incident asthma (95% CI)	Important results/ highest risk group	CASP quality assessme
O'Neil <i>et al.</i> (2016) <sup>46</sup>	South-Eastern Australia/ Geelong Osteoporosis Study	Prospective longitudinal study		860 women	Depression and anxiety	SCID-I/NP	<ul> <li>Age</li> <li>Education</li> <li>BMI</li> <li>Alcohol frequency</li> <li># years smoker</li> </ul>	Occurrence of a CHD	Hospital medical records	Primary outcome (incidence yes/no): Baseline depres- sion (OR 3.28, Cl 1.36 to 7.9) Baseline anxiety (OR 0.62, Cl 0.18 to 2.14) Secondary outcome (all cardiac events): Depression OR 2.74 Cl (1.42 to 5.29) Anxiety OR 0.34 Cl (0.12 to 0.99) Number of CHD events Depression IRR 1.03 Cl (0.40 to 2.63) Anxiety IRR 1.64 Cl (0.35 to 7.76)	Primary outcome: Age OR 1.07 (1.02 to 1.11) Education OR 0.49 (0.27 to 0.87) Secondary outcome: Age OR 1.04 (1.01 to 1.07) Years of smoking OR 1.03 (1.00 to 1.05) Number of CHD events Age IRR 1.07 (1.01 to 1.13) BMI IRR 17.21 (1.26 to 235.46) Alcohol frequency IRR 3.29 (1.40 to 7.73)	High
Balog et al. (2017) <sup>47</sup>	Hungarian Epidemiological Panel Survey/ Hungary	Prospective longitudinal study	4 years	2725; 56.44%	Depression	BDI-S	<ul> <li>Age</li> <li>Education al attain- ment (6 categories</li> <li>Alcohol abuse</li> </ul>	CVD - morbidity including hypertension ) treatment/ CVD event incidence	Self-reported	Depressive symptomatology factor OR 1.15 Cl (0.97 to 1.36)	Age OR 1.05 (1.04 to 1.07) BMI OR 1.05 (1.01 to 1.10)	Low

ent

Study	Country/name of the study	Design	Duration of follow- up	Type of assessed MH	Measurement of MH condition	Co	ovariates	Outcome	Measurement of outcome	Adjusted association of depression/PTSD with overall incident asthma (95% CI)	Important results/ highest risk group	CASP quality assessment
						•	Smoking status (smoker vs. non- smoker) Physical activity BMI					
Anda et al. (1993) <sup>48</sup>	USA (NHANES follow-up study of 1987)	longitudinal	length of follow-up		General Well-being Questionnaire	• • • • • • • • • • • • • • • • • • • •	Age Sex Race Education Marital status Smoking status Physical activity	Ischaemic heart disease incidence and fatality	Hospital discharge records Cause of death was determined via death records coded against ICD-9	Fatal IHD Depressed affect No (Ref) Yes RR 1.5 (1.0 to 2.3) Non-fatal IHD Depressed affect No (Ref) Yes RR 1.6 (1.1 to 2.4)	Additive interaction Non-smoker/ Non- depressed (Ref) Depressed affect Adjusted RR 1.31 Smoker only Adjusted RR 1.65 Smoker and depressed Adjusted RR 2.97	Medium

#### **TABLE 5** Characteristics of selected studies linking depression and/or PTSD with CVD (continued)

BDI-S, Beck's Depression Inventory; CASP, Critical Appraisal Skills Programme; CHD, coronary heart disease; IHD, ischaemic heart disease; IRR, Incidence rate ratio; SCID-I/NP, Structured Clinical Interview for DSM-IV, Nonpatient edition.

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risk, coefficients etc.). Also, there is some between-study variance in terms of assessing the impact of certain determinants. For example, the impact of smoking was assessed in some studies based on the duration of smoking, while in others on the number of cigarettes smoked per day.

### **Quality assessment**

The Critical Appraisal Skills Programme (CASP) checklists for assessing the quality of cohort<sup>49</sup> and case-control studies<sup>50</sup> were used. The CASP checklists, devised for use in health-related research, were used for quality assessment of papers as 'high', 'medium' or 'low' quality. CASP is commonly used in qualitative evidence syntheses and is endorsed by the Cochrane Collaboration.<sup>51</sup> All articles were assessed for quality by a reviewer (ZJ), and another reviewer (AN) was available for mediation to arrive at a consensus through discussion using the CASP checklists.

### Results

The study selection process is summarised in Figure 1, following PRISMA reporting guidelines.<sup>32</sup> More than 19,000 studies were found in the initial search. A total of 11 studies from 3 countries were included in the final review, with the majority (n = 9) conducted in the USA. All studies were longitudinal prospective studies, presenting outcomes for a total of 71,896 people. Four studies determined the association between depression and cancer, two studies depression with CVD and one study depression with diabetes. One study determined the association between depression and anxiety together with diabetes and another with CVD. No studies were found examining association between CMDs and COPD per se, but two studies addressing asthma were included (one considering depression and the other PTSD). A summary of the included studies is presented in Table 6.

### Cancer

Evidence linking social determinants to cancer incidence: *Table 7* summarises evidence grouped per determinant. It suggests that the determinants that showed a significant association were all associated with an increase in cancer incidence; however, one high-quality study and one medium-quality study were available. The latter suggested that age may be associated with cancer incidence; however, smoking status probably is.

# Summary of findings from retrieved articles

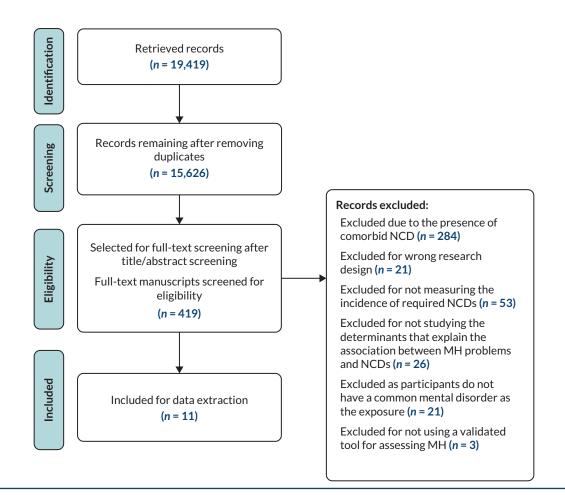
Four papers studied the effects of individual (smoking, age, gender, alcohol intake, body mass index and

ethnicity) and social (marital status, occupational status and socioeconomic status) mental health determinants on cancer incidence (Table 2). None of the included studies adjusted for environmental factors. Follow-up period ranged between 10 and 24 years. Three of the four papers reported a significant association between depression and incident cancer.<sup>40,41,53</sup> Two population-based studies assessed depression at three different time intervals. Both studies revealed a specificity to the association between depression and hormonally mediated cancers, thus supporting the hypotheses about a common biological pathway between depression and cancer.40,41 Penninx et al. measured chronic depression at three different times (1982, 1985 and 1988) in order to consider if depression is related to temporary stressful life circumstances that may remit shortly after.<sup>41</sup> Unlike baseline depression which was not significantly associated with incident cancer, chronically depressed mood, persistent for 6 years among individuals aged 71 years and above, was associated with an increased HR of 1.88 (95% CI = 1.3 to 3.14) for developing cancer after adjusting for age, sex, alcohol intake, smoking, race, physical disability and hospital admissions. A similar HR of 1.87 (CI 1.16 to 3.01) and 1.69 (CI 1.3 to 2.19) for the development of cancer among persons diagnosed with major depression and dysphoria, respectively, was observed in the study by Gross et al.<sup>40</sup> The association between depression and incident cancer in the study by Persky et al. among male employees was significant (p < 0.05); yet it was only observed when cancer was diagnosed during the first 10 years of follow-up.

The analysis of interaction between smoking, depression and incident cancer showed mixed results in all four studies. Smoking habits were unlikely to be related to the increased cancer risk among chronically depressed individuals. In the study where incident cancer was examined among persons with chronic depression, the HR for non-smokers exceeded that of smokers who are not chronically depressed. Compared with the non-chronically depressed participants, chronically depressed participants were more often females and of older age and less often smokers or excessive drinkers. Also, a more strict cut-off point for CES-D scores was adopted, 20 instead of 16, to yield a higher accuracy for the identification of participants with severe depression among older individuals.<sup>41</sup> In contrast, three studies found a significant association between smoking status and the incidence of cancer among depressed individuals.40,52,53 The study by Persky et al. found an association between number of cigarettes smoked and cancer risk among employees with depression, with no specification to the type of cancer. In the other two studies, the interaction was observed at cancer sites related to smoking, such as

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### FIGURE 1 PRISMA flow diagram.

#### TABLE 6 Description of study characteristics

Study characteristic	Number of studies (N = 11)	Study characteristic	Number of studies (N = 11)	
Countries		Diagnostic Interview Schedule (DIS/ DSM-III)	2	
USA Australia	9 1	Centre for Epidemiological Studies Depression Scale (CES-D)	3	
Hungary	1	PTSD checklist	2	
Study design Prospective longitudinal cohort	11	Minnesota Multiphasic Personality Inventory (MMPI)	1	
Sample size		SCID-I/NP	1	
< 1000	1	General Well-Being Schedule	1	
1000-3000	5	NCD		
≥ 3000	5	DM	2	
CMD		CVD	3	
Depression	8	Cancer	4	
Anxiety	1	COPD	0	
Both	2	(Asthma = 2) CASP quality assessment		
CMD screening tool		High	3	
PHQ-9	1	Medium	4	
Beck's Depression Inventory (BDI-S)	1	Low	4	

### TABLE 6 Description of study characteristics (continued)

**TABLE 7** Summary of the contribution of mental health determinants to cancer incidence, categorised per quality of evidence using the CASP tool [high quality (H), medium quality (M) and low quality (L)]

Quality of included studies (results of CASP tool)					
Social determinants	н	М	L		
Age		Gross et al. (2010) <sup>40</sup> +	Linkins <i>et al.</i> (1990) <sup>52</sup> + Persky V. <i>et al.</i> (1987) +		
Smoking status	Penninx <i>et al</i> . (1998) <sup>41</sup> +	Gross et al. (2010) <sup>40</sup> +	Linkins <i>et al.</i> (1990) <sup>52</sup> +		
Number of cigarettes/day			Linkins <i>et al.</i> (1990) <sup>52</sup> + Persky <i>et al.</i> (1987) <sup>53</sup> +		

+ sign indicates the direction of the association.

**TABLE 8** Summary of the contribution of mental health determinants to asthma incidence, categorised per quality of evidence using the CASP tool [high quality (H), medium quality (M) and low quality (L)]

Quality of included studies (results of CASP tool)					
Social determinants	н	М			
Female gender	Brunner <i>et al</i> . (2014) <sup>42</sup> +	De La Hoz <i>et al.</i> (2016) <sup>43</sup> +			
BMI	Brunner <i>et al.</i> (2014) <sup>42</sup> +	De La Hoz <i>et al.</i> (2016) <sup>43</sup> +			
Latino and African ethnicities		De La Hoz et al. (2016) <sup>43</sup> –			
Educational attainment	Brunner <i>et al</i> . (2014) <sup>42</sup> –				
Land signs indicate the direction of the acception					

+ and – signs indicate the direction of the association.

buccal cavity and pharynx cancer<sup>52</sup> and lung cancer.<sup>40</sup> As indicated in *Table 7*, age and smoking status were the only two covariates adjusted for in the study by Linkins *et al.* which might have resulted in an overestimation of the impact of smoking.

The two studies adjusting for older age found it to be significantly associated with an increased risk for developing cancer.<sup>40,53</sup> Except for 'never having been married' which had a protective effect against the development of cancer [HR 0.37 (CI 0.18 to 0.75)],<sup>40</sup> none of the remaining covariates in all studies had any significant association with incident cancer among depressed individuals.

# Chronic obstructive pulmonary disease and asthma

### Evidence linking social determinants to asthma incidence

Available evidence from a medium-quality article suggests that female gender and BMI may be associated with an increased risk of developing asthma, and evidence from the high-quality article confirms that this is probably the case. Evidence from a medium-quality article suggests that being Latino or African, compared to being white, may be associated with decreased likelihood of developing asthma. Finally, based on one high-quality study, educational attainment is probably negatively associated with developing the condition (*Table 8*).

### Summary of findings from retrieved articles

As noted previously, no studies linking depression and/ or anxiety with incident COPD per se were identified in the literature. However, asthma is a form of lung disease that shares similar pathophysiological characteristics with COPD.<sup>54</sup> Also, there is evidence linking COPD to incident asthma. A recent longitudinal study of women, followed prospectively for 13 years, found that more than one in three women with asthma developed COPD. Sociodemographic, lifestyle and environmental risk factors were shown to have a significant association with the progression from asthma to COPD. Older age, obesity, cigarette smoking, unemployment, low education attainment and rural residence were all found to play a significant role in the progression to COPD.<sup>55</sup> Some risks associated with the progression from asthma to COPD were also found to be significantly associated with the progression from depression and anxiety to asthma.<sup>42,43</sup> Since there was evidence linking asthma to COPD incidence and since there was also evidence that mental health disorders could play a role in this, it was logical for the research team to include studies that looked at asthma as a disease outcome.

Two longitudinal studies identified in this review investigated the link between mental ill-health and incident asthma. As shown in *Table 3*, both studies reported significant associations with factors such as gender and obesity as independent predictors of incident asthma. In the multicentre Coronay Artery Risk Development in young adults (CARDIA) study, baseline and chronic depression were significantly associated with incident asthma.<sup>42</sup> The chronicity of PTSD in the study by de la Hoz also had a greater cumulative impact on the incidence of asthma. In a multivariable analysis, lifetime non-smokers with probable PTSD were found to be 2.41 times more likely to develop asthma than non-smokers without PTSD at baseline (OR

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**TABLE 9** Summary of the contribution of mental health determinants to diabetes incidence categorised per quality of evidence using the CASP tool [high quality (H), medium quality (M) and low quality (L)]

Quality of included studies (results of CASP tool)					
Social determinants	м	L			
Age		Boyko <i>et al</i> . (2010) <sup>44</sup> +			
BMI		Boyko <i>et al</i> . (2010) <sup>44</sup> +			
Latino and African ethnicities		Boyko et al. (2010) <sup>44</sup> +			
Educational attainment	Mezuk <i>et al.</i> (2008) <sup>45</sup> –				
+ and – signs indicate the direction of the association.					

= 2.41, 95% CI 1.85 to 3.13). The risk increased when participants had probable PTSD at baseline and follow-up visits. Participants with chronic PTSD were more than four times more likely to develop asthma (OR = 4.18, 95% CI 3.06 to 5.72). That the magnitude of association was greater between late-onset or chronic PTSD and incident asthma than with recovering probable PTSD and incident asthma suggests that treating PTSD could potentially reduce risk for asthma.<sup>43</sup>

In both studies, incident asthma was defined either by a new report of asthma medication use and/or by self-reported physician diagnosis. Self-report of asthma has been found to be acceptable in epidemiologic studies with respect to physician diagnosis and is not linked to misclassification bias.<sup>56</sup> Misclassification bias is a pervasive systematic error that leads to the inaccurate classification of individuals participating in a study.<sup>57</sup> In other words, individuals who provide self-reports of asthma are prone to exhibit comparable asthma states to those who have received a medical diagnosis, hence suggesting a reasonable degree of accuracy in self-reporting.<sup>58</sup>

### **Diabetes mellitus**

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### Evidence linking social determinants to diabetes mellitus incidence

*Table 9* suggests that there is inconclusive evidence linking age, BMI and ethnicity to increased incidence of diabetes, due to low quality of evidence. However, educational attainment may be negatively associated with increased risk for diabetes incidence, as per one medium-quality study.

### Summary of findings from retrieved articles

Two studies reporting the incidence of DM as an outcome were included. Mezuk *et al.* tested prospectively the

relationship between depression and type 2 diabetes among residents of East Baltimore.<sup>45</sup> Depression assessed via Diagnostic Interview Schedule (DIS) was collected in 1982, 1993 and 2005. The DIS is a standardised interview protocol conducted by individuals without professional clinical training. Its purpose is to determine diagnoses of major depressive disorder (MDD) and depressive syndrome, as well as ascertain the age at which these disorders first appeared and their recent occurrence, using diagnostic algorithms.<sup>59</sup> The onset of depression was determined as the earliest age when symptoms of depression began. The average age of depression onset was 30.9 years, and that of type 2 diabetes was 55 years. Respondents self-reported diabetes incidence, which may also have underdetected cases of diabetes. On average, it took 16 years for diabetes to develop after the onset of depression. Depression was found to be associated with diabetes onset [HR 1.62 (1.03 to 2.55)] after controlling for age, gender, race, education, social network size and antidepressant use. This association was more evident after controlling for additional sociodemographic characteristics and health behaviours [HR 2.04 (1.09 to 3.81)]. In a stratified analysis, the risk was elevated among those with 12 years of education or less (Table 4). No effect was seen for behavioural factors such as smoking or physical activity; however, that could be due to the actual measures not being sensitive enough to capture an effect. For instance, information on current smoking status was collected and not on number of cigarettes smoked or duration of smoking. For alcohol intake, information was collected on number of days where participants drank in the past month. Survival bias is also another limitation in such a study where the duration of follow-up was 23 years. Survival bias is observed when examining the relationship between a certain exposure during early life and a future health outcome during later stages of life. The presence of bias in the study may introduce a potential distortion in the results, as the examination exclusively focuses on the participants who have survived while neglecting to consider the characteristics of those who did not survive.<sup>60</sup>

Diabetes risk was measured in 44,754 members of the US military service followed up for 3 years after assessing depression and anxiety at baseline.<sup>44</sup> Diabetes onset was self-reported, and the survey instrument included sociodemographics, weight, height and military service information. In a univariate comparison of baseline characteristics by diabetes status at follow-up, those with MDD and PTSD, screened using the self-administered PHQ-9 and PTSD checklist, respectively, had higher odds for developing diabetes: unadjusted OR 1.95 (1.23 to 3.11) and unadjusted OR 2.56 (1.78 to 3.67). In the multivariable model, only baseline PTSD, and not depression, was significantly associated with diabetes onset after adjusting

for age, sex, BMI, education, race/ethnicity, military service characteristics and mental health conditions OR 2.07 (1.31 to 3.29). The authors suggested that the high occurrence of both conditions at baseline suggests that depression may serve as a surrogate marker for PTSD in other studies not measuring both conditions. Also, participants might have under-reported mental health symptoms due to fear of adverse consequences on their military records. Increased odds for diabetes were seen among participants of older age, higher BMI, non-Caucasians and those separated from the military. Being on active duty seems to have a protective effect against developing diabetes, which could be due to the fact that those selected for deployment pass a medical screening and are usually healthier and more fit. In this study, the interaction between gender and depression with DM onset suggests that gender plays a role in the development of DM. Non-depressed males were found to be at a significantly increased risk for developing DM. There is inconclusive evidence however whether the same association remains when depression ensues.

### Cardiovascular diseases

## Evidence linking social determinants to cardiovascular disease incidence

High-quality evidence suggests that age and educational attainment are probably associated with a positive and a negative association, respectively, with the incidence of CVD. Smoking status may be associated with an increased risk of CVD, and there is inconclusive evidence linking BMI to an increased incidence of CVD due to low-quality evidence only being available (*Table 10*).

### Summary of findings from retrieved articles

Three studies examining the effect of depression on CVD incidence were identified, with follow-up periods ranging between 4 and 23 years. A prospective study of 860 women randomly selected from Australian electoral rolls was followed up for 18 years to study the impact of depression and anxiety on events requiring hospitalisation admission confirming coronary heart disease (CHD) incidence (primary outcome) and additional diagnosis of stable/unstable angina and 'not otherwise defined' CHD events (secondary outcome).<sup>46</sup> Depression and anxiety diagnosis was conducted by mental health professionals, at baseline and 10 years later, utilising the Structured Clinical Interview for Diagnostic and Statistical Manual of Mental Disorders, fourth edition, non-patient edition, which is the gold standard for assessing such disorders. Depression (OR 3.28, Cl 1.36 to 7.9), and not anxiety (OR 0.62, CI 0.18 to 2.14), was found to be an independent risk factor for CHD after adjusting for demographic, anthropometric, health and clinical factors.

**TABLE 10** Summary of the contribution of mental health determinants to CVD incidence, categorised per quality of evidence using the CASP tool [high quality (H), medium quality (M) and low quality (L)]

Quality of included studies (results of CASP tool)					
Social determinants	н	м	L		
Age	O'Neil <i>et al.</i> (2016) <sup>46</sup> +		Balog et al. (2017) <sup>47</sup> +		
BMI			Balog <i>et al.</i> (2017) <sup>47</sup> +		
Smoking status		Anda et al. (1993) <sup>48</sup> +			
Educational attainment	O'Neil <i>et al.</i> (2016) <sup>46</sup> –				

+ and – signs indicate the direction of the association.

Neither depression nor anxiety was significantly associated with recurrent cardiac episodes. The association shown between depression and CHD incidence was of greater magnitude than the association seen with other risk factors (*Table 5*). This suggests a need to set depression as a target in primary prevention of CHD as well as its influence on risk factor behaviours, an approach that could be useful for helping alleviate CVD burden globally.

Another study by Balog *et al.* examined the concurrent effects of depression and vital exhaustion (described by the authors as loss of energy, increased irritability and general demoralisation) on CVD morbidity in a random sample of 2317 participants from the Hungarian Epidemiological Panel Survey.<sup>47</sup> Greater age and BMI were positive predictors of CVD incidence after 4 years of follow-up, according to the logistic regression model, while depression did not have a significant predictive role. In addition to the relatively short follow-up duration, all information, including that of mental distress, was collected via self-administered questionnaires. Therefore, information pertaining to CVD incidence and/or diagnosis is highly unreliable. Also, survival bias is another limitation in such a case.

Using a subscale from the General Well-Being Questionnaire, depressed affect was assessed in a population sample of 2832 US citizens to examine the association of ischaemic heart disease (IHD).<sup>48</sup> After a mean follow-up period of 12.4 years, depression was found to be associated with an increased risk of fatal and non-fatal incidence of IHD (*Table 5*) after adjusting for age, sex, race, education, marital status, smoking, total cholesterol, systolic blood pressure,

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BMI, alcohol use and physical activity. No analysis was run to measure the individual effect of confounding variables on the incidence of IHD. The interaction between depression and smoking indicated a synergistic effect, increasing the relative risk for developing IHD.

### Discussion

Findings from the systematic review suggest that the chronicity of depression and PTSD could be associated with increased risk for cancer and asthma onset, respectively, and therefore treating depression and PTSD could prevent progression into the development of the latter NCDs. Moreover, the high association between depression and CHD suggests the need to target depression as primary prevention to prevent the progression into CHD. On the other hand, factors such as educational attainment, number of cigarettes smoked, female gender, older age, elevated BMI and ethnicity may contribute to an elevated risk for developing NCDs among adults with depression and/or PTSD. Therefore, in order to reduce the incidence of NCDs, mental disorders should be identified and treated, and health and prevention strategies focusing primarily on high-risk groups, where factors listed above are present, can and should be adopted.

Our findings are consistent with cohort studies where older age,<sup>61</sup> low educational attainment,<sup>62</sup> higher BMI<sup>63</sup> and smoking status<sup>61</sup> were found to be associated with CVD incidence. In the current review, the study by O'Neil has identified depression as an independent risk factor for CVD onset,<sup>46</sup> which is also suggested by other longitudinal studies.<sup>64-66</sup> We, therefore, recommend to include depression as a target for primary prevention of CVD.

Ethnicity was found to be associated with the onset of diabetes. Factors identified earlier as being related to CVD incidence, except smoking, were also reported in this review to be associated with diabetes onset. This is in agreement with findings from the literature that linked older age, higher BMI<sup>67,68</sup> and low educational attainment<sup>69</sup> with incident diabetes. Our results regarding the type of mental disorder associated with type 2 diabetes incidence contradict findings of the literature, however. While numerous research has demonstrated a bidirectional association between depression and type 2 diabetes,<sup>70,71</sup> our review indicates that PTSD, and not depression, was found to increase incident diabetes when the effects of depression and PTSD were studied together. A study by Scherrer et al. found that reducing PTSD symptoms lowered the risk of type 2 diabetes<sup>72</sup> among participants with PTSD. In the same study, depression remission contributed to lower diabetes onset among participants diagnosed with

comorbid PTSD and depression, and not with depression alone. Therefore, there may be further benefit from reducing PTSD symptoms to lower type 2 diabetes risk in populations that are exposed to potentially traumatic events, such as veterans. For other populations exposed to such events, for example, refugees and the general population affected by adverse events, studies are lacking. Therefore, it is difficult at this point to generalise a recommendation as to target PTSD for the primary prevention of type 2 diabetes.

In relation to cancer incidence, three out of four studies demonstrated an association between depression and incident cancer. This is in line with two previous metaanalyses.<sup>73,74</sup> Although not all studies included in this review showed a significant impact of depression on cancer initiation, some trends emerged regarding the extended duration of depression having an important role in a slowly developing health condition such as cancer. This view is shared by Garssen<sup>75</sup> who suggested in 2004 specific risk factors among this population, namely, smoking, BMI, age and alcohol intake. This research also found that behavioural factors, namely smoking<sup>76</sup> and alcohol intake,<sup>77</sup> were associated with increased cancer onset.

Our review showed a significant association between depression and PTSD with incident asthma. In 2005, a review that explored the interaction between depression and asthma as comorbidities suggested that the respiratory tract (mainly the airways) is greatly reactive to the psychological changes of individuals.<sup>78</sup> The findings of our study suggest that depression could lead to asthma onset via this mechanism. Regarding PTSD, our findings are consistent with those in the literature whereby PTSD was found to be positively associated with asthma onset.<sup>79,80</sup> Our study identified risk factors such as female gender, increased BMI and low educational attainment as being related to asthma onset. Therefore, special attention should be given to such population groups in clinical practice and public health interventions that aim to reduce the incidence of asthma.

In addition to the complexity in conducting longitudinal studies to test the causal association between common mental health disorders and NCDs, there appears to be a challenge in relation to the reporting of health conditions. Epidemiological investigations and population-based studies mainly rely on self-reports to obtain information on health outcomes, such as diabetes, given the high cost and low efficiency of clinical diagnosis.<sup>81,82</sup> A number of studies assessing the accuracy of self-reported data, using test characteristics such as sensitivity (the fraction of people who fit the medical criteria for the outcome and self-report a diagnosis, i.e. true positive) and specificity (the fraction of people who have normal measures and do not report a diagnosis, i.e. true negative), gave mixed

results.<sup>83-85</sup> Compared to biochemical measurements, selfreported data on diabetes among Chinese adults ≥ 45 years showed low sensitivity and a moderate overall agreement. Factors associated with accurate self-reported data were female sex, older age, high socioeconomic status, better educational attainment and recent utilisation of medical services.<sup>85</sup> Goldman et al. attributed the lack of accuracy in self-reports to several factors, such as undiagnosed diseases, inability to recall, unwillingness to disclose medical information and lack of accessibility to health services.<sup>84</sup> This could further be exacerbated among individuals facing psychological stressors. Depression, for instance, was found to be associated with barriers deterring individuals from utilising medical services. Factors such as mistrust in health providers and the health system,<sup>86</sup> loss of income<sup>87</sup> and low motivation to seek medical care<sup>88</sup> may all result in underdiagnoses leading to under-reporting. Therefore, the above-mentioned issues that feed into under-reporting should be taken into account when conducting future research on this topic.

Another common challenge pertains to the identification of cases of common mental health disorders. There is no single diagnostic tool for such conditions. For instance, several valid and reliable instruments are used by primary care clinicians and researchers to identify cases of depression and anxiety.<sup>89-91</sup> The selection of the appropriate tool depends on factors such as feasibility, administration time and personnel and interest in measuring the severity of the condition or monitoring the response to therapy. These tools tend to under-report cases compared with the clinical psychiatric examination and hinder the ability to compare results from different studies.

### **Strengths and limitations**

This systematic review provides additional evidence on how common mental health disorders are linked to the onset of the four most common NCDs. The particularity of the evidence that emerged has to do with the type of individual studies: the included longitudinal studies provide valuable evidence for establishing a causal relationship between common mental health disorders and NCDs in this specific direction. However, it should be noted that the utilisation of a longitudinal research design does not completely preclude the possibility of other explanations, such as the influence of third variables. Consequently, it does not definitively demonstrate a causal relationship or enable the formulation of causal claims.<sup>92</sup> Other limitations of the current review should be acknowledged. First, our review did not include studies from the grey literature and was limited to literature published in English language only. Second, measures of depression and anxiety varied among studies, with the majority relying on self-report using different screening tools with various cut-off points. This might have caused a misclassification of exposure and, hence, lessened the association between mental disorders and the incidence of NCDs. Third, measures of covariates were different across studies which makes it difficult to compare their effects on the onset of diseases. And finally, the majority of included studies limited their assessment of mental conditions to one point in time which might have been a transient episode. The association found between chronic depression/PTSD and incident NCDs emphasised the importance of assessing the chronicity of mental illnesses and their impact on health.

### Conclusion

This study contributed to the existing global body of evidence regarding the factors that are directly linked to the onset of NCDs. As a result, policy-makers and practitioners can utilise the findings to customise interventions that target these specific factors, with the aim of interrupting the cycle between mental health disorders and other NCDs. Interventions need to target those with common mental health disorders and take into account: age (older age groups are more at risk), sex (females are more at risk), educational level (individuals with <12 years of education are more at risk) and smoking status (smokers are more at risk). Furthermore, there is an urgent need to conduct studies to understand the linkages between mental health and NCDs in LMICs, given the absence of evidence in those settings and the increasing burden of NCDs and mental health conditions they are experiencing. Finally, research to evaluate public health interventions and to assess the responsiveness of healthcare systems (mainly at the primary care level) is essential to alleviate the burden of those conditions at population level. In summary, this review adds to the body of evidence that prioritising the social determinants of mental health is crucial for enhancing public health outcomes through the implementation of a preventative health promotion strategy. Through this approach, it becomes possible to address the systemic determinants of health that are often overlooked, which can contribute to the development of disease.93

**Reporting community engagement and involvement:** Since this was a review article, there was no patient and public involvement in this study.

**Equality, diversity and inclusion**: Research around mental and physical health multimorbidity is limited, despite the

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fact that the global burden of mental illnesses and NCDs is high. This review contributes to the existing global body of evidence regarding the factors that are directly linked to the onset of NCDs. Findings can be utilised to inform interventions for high-risk populations worldwide to prevent development of NCDs.

### **Additional information**

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### **CRediT** contribution statement

Zeina Jamal (https://orcid.org/0000-0002-5078-3416): Conceptualisation (equal), Data curation (lead), Formal analysis (lead), Investigation (lead), Project administration (lead), Visualisation (lead), Writing – original draft (lead), Writing – editing and reviewing (equal). Alastair Ager (https://orcid.org/0000-0002-9869-544X): Conceptualisation (equal), Investigation (supporting), Supervision (equal), Visualisation (supporting), Writing – editing and reviewing (equal). Rebecca Horn (https://orcid.org/0000-0002-9474-3563): Conceptualisation (equal), Investigation (supporting), Supervision (equal), Visualisation (supporting), Writing – editing and reviewing (equal), Visualisation (supporting), Writing – editing and reviewing (equal), Visualisation (supporting), Writing – editing and reviewing (equal).

### **Disclosure of interests**

**Full disclosure of interests:** Completed ICMJE forms for all authors, including all related interests, are available in the toolkit on the NIHR Journals Library report publication page at https://doi.org/10.3310/CTDH4677.

**Primary conflicts of interest:** The authors declare that they have no competing interests.

### **Data-sharing statement**

Requests for access to data should be addressed to the corresponding author.

#### **Ethics statement**

This study is classified as a systematic review, and consequently, ethical approval was not required.

#### Information governance statement

This is a systematic review, and therefore, the current research did not handle any personal information.

### Official Development Assistance (ODA) compliance statement

The research project holds considerable importance in relation to mental health and NCDs due to its direct applicability

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to the developmental challenges encountered by LMICs. These countries face an immense burden in addressing the challenges posed by mental illness and NCDs, as well as grappling with inadequacies in properly addressing and treating these conditions. Although the main focus of this study was not specifically directed towards LMICs, the findings of this research will have substantial implications for policy formulation and practical implementation in LMIC settings. By gaining a comprehensive understanding of the complex interplay between mental health, NCDs and the determinants influencing them, policy-makers and healthcare professionals can develop more efficacious approaches to prevent, promptly detect and manage these conditions within LMICs.

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The contractual start date for this research was in June 2017. This article began editorial review in March 2023 and was accepted for publication in December 2023. The authors have been wholly responsible for all data collection, analysis and interpretation, and for writing up their work. The Global Health Research editors and publisher have tried to ensure the accuracy of the authors' article and would like to thank the reviewers for their constructive comments on the draft document. However, they do not accept liability for damages or losses arising from material published in this article. This article was published based on current knowledge at the time and date of publication. NIHR is committed to being inclusive and will continually monitor best practice and guidance in relation to terminology and language to ensure that we remain relevant to our stakeholders.

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### List of abbreviations

BMI	body mass index
CASP	Critical Appraisal Skills Programme
CHD	coronary heart disease
CIDI	Composite International Diagnostic Interview
CMD	common mental disorder
COPD	chronic obstructive pulmonary disease
CVD	cardiovascular disease
DM	diabetes mellitus
ICD	International Classification of Disease
IHD	ischaemic heart disease
MeSH	medical subject heading
NCD	non-communicable disease
PHQ-9	Patient Health Questionnaire-9 item
PICOS	population, intervention/exposure, control, outcome and study design
PRISMA	Preferred Reporting Items for Systematic Reviews and Meta- Analyses
PTSD	post-traumatic stress disorder
SCID	Structured Clinical Interview for Diagnostic and Statistical Manual of Mental Disorders
WHO	World Health Organization

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### Appendix 1 MEDLINE search strategy

#	Searches	Results
1	TI, AB cardiovascular disease* OR CVD OR heart disease* OR vascular disease* OR cerebrovascular diseases	415,338
2	(MM 'Cardiovascular Diseases') OR (MM 'Pathological Conditions, Signs and Symptoms') OR (MM 'Disorders of Environmental Origin') OR (MH 'Endocrine System Diseases+') OR (MH 'Nutritional and Metabolic Diseases+') OR (MH 'Vascular Diseases+') OR (MH 'Heart Diseases+')	3,753,932
3	TI, AB type 2 diabetes OR type ii diabetes or noninsulin dependent or non insulin depen- dent OR non-insulin dependent OR NIDDM OR adult onset diabetes OR maturity onset OR diabetes mellitus	415,051
4	(MM 'Diabetes Mellitus, Type 2') OR (MM 'Diabetes Complications/DI/ET/PX/EP/CO')	102,692
5	TI, AB neoplas* OR cancer* OR carcinoma* OR tumor* OR tumour* OR malignan* OR leukaemia OR leukemia OR lymphoma	3,805,137
6	(MM 'Neoplasms/PX/ET/EP/CO/DI') OR (MM 'Carcinogenesis')	77,036
7	TI, AB COPD OR chronic obstructive pulmo- nary disease OR lung disease OR pulmonary disease	238,793
8	(MM 'Pulmonary Disease, Chronic Obstructive/CO/DI/EP/ET/PX')	10,078
9	(MM 'Non-communicable Diseases/PX/EP') OR (MM 'Communicable Diseases, Emerging/ DI/EP/PX/ET/CO')	2133

#	Searches	Results
10	TI, AB stress disorder OR PTSD OR Post- traumatic OR Posttraumatic OR psychological distress OR emotional distress	96,438
11	(MM 'Stress Disorders, Post-Traumatic/CO/ DI/EP/ET/PX')	16,533
12	TI, AB anxiety OR phobi* OR panic OR anxio*	60,099
13	(MM 'Anxiety Disorders+/DI/CO')	10,301
14	(TI depress*) OR (AB depress*)	429,598
15	(MM 'Depressive Disorder/CO/EP/PX')	15,364
16	(MM 'Mental Health')	20,351
17	TI ('follow up' OR 'follow-up') OR AB ('follow-up' OR 'follow up')	895,827
18	TI longitudinal OR AB longitudinal	224,732
19	S1 OR S2 OR S3 OR S4 OR S5 OR S6 OR S7 OR S8 OR S9	7,384,756
20	S10 OR S11 OR S12 OR S13 OR S14 OR S15 OR S16	569,216
21	S19 AND S20 (publication type: observational study)	886
22	S19 AND S20 (no limiters applied)	100,223
23	S22 AND S18	3439
24	S22 AND S17	9104
25	S23 OR S24 OR S21	<b>9153</b> (full text)

### Appendix 2 Data extraction sheet template

Duration Study	# of cohort/ case-control Gender population (%F)	Age of MH		Determinants assessed	assessing	incident	· ·
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