

Mental health symptoms during the COVID-19 pandemic in developing countries: A systematic review and meta-analysis

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Background This systematic review aims to 1) summarize the prevalence of anxiety, depression, distress, insomnia, and PTSD in the adult population during the first year of the COVID pandemic in developing countries and 2) uncover and highlight the uneven distribution of research on mental health in all developing countries across regions.

Methods Several literature databases were systemically searched for meta-analyses published by September 22, 2021, on the prevalence rates of mental health symptoms in developing countries worldwide. We meta-analysed the raw data of the individual empirical results from the previous meta-analysis papers in developing countries in different regions.

Results The prevalence rates of mental health symptoms were summarized based on 341 empirical studies with a total of 1 704 072 participants from 40 out of 167 developing countries in Africa, Asia (East, Southeast, South, and West), Europe, and Latin America. Comparatively, Africa (39%) and West Asia (35%) had the worse overall mental health symptoms, followed by Latin America (32%). The prevalence rates of overall mental health symptoms of medical students (38%), general adult students (30%), and front-line health care workers (HCWs) (27%) were higher than those of general HCWs (25%) and general populations (23%). Among five mental health symptoms, distress (29%) and depression (27%) were the most prevalent. Interestingly, people in the least developing countries suffered less than those in emergent and other developing countries. The various instruments employed lead to result heterogeneity, demonstrating the importance of using the well-established instruments with the standard cut-off points (eg, GAD-7, GAD-2, and DASS-21 for anxiety, PHQ-9 and DASS-21 for depression, and ISI for insomnia).

Conclusions The research effort on mental health in developing countries during COVID-19 has been highly uneven in the scope of countries and mental health outcomes. This meta-analysis, the largest on this topic to date, shows that the mental health symptoms are highly prevalent yet differ across regions. The accumulated systematic evidence from this study can help enable the prioritization of mental health assistance efforts to allocate attention and resources across countries and regions.

The COVID-19 pandemic has truly been a global pandemic, particularly impactful in developing countries due to the lack of resources and initiatives for tackling mental health issues in the already overburdened and fragile health care systems [1,2]. For instance, the developing country of Bangladesh has a psychiatrist-population ratio of 0.13 per 100 000, far below the recommended range of 3.8 to 15.8 per 100 000 [3]. The overburdened and fragile health care systems and slower or non-existent access to vaccination impose psychological stress on ordinary citizens, who suffer under restrictions and lockdowns for multifaceted reasons

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including, but not limited to, family separations, reduced social contact, less unemployment, work stress, increased loneliness, the overabundance of (mis)information on social media, and various other related factors [4-6]. Global assessments of mental health symptoms under the COVID-19 pandemic across developing countries remain a challenging yet vital endeavour [7].

However, developing countries lack large-scale monitoring of the prevalence of mental health symptoms at the national level, as implemented in developed countries such as the USA and the UK. Instead, developing countries can only rely on individual primary studies from independent researchers (eg, [8], and as a result, there is an uneven geographical distribution of such studies under the COVID-19 pandemic [9]. For instance, a review revealed only 12 out of the 48 countries in Africa have been researched on mental health during COVID-19 [10]. To map the status of mental health in developing countries worldwide, this study aims to summarize the prevalence of anxiety, depression, distress, insomnia, and PTSD in their adult populations under the COVID pandemic. The second aim of this study is to uncover and highlight the uneven distribution of research on mental health across all developing countries across regions.

METHODS

To achieve such aims, we systematically searched and leveraged the existing meta-analyses on the topic to identify individual studies on the prevalence rates of mental health symptoms in developing countries worldwide. We aimed to reveal the inadequacy, if not the absence of, studies in certain countries, to highlight the need to initiate relevant research on the neglected countries. We also conducted a meta-regression to assess the prevalence rates of anxiety, depression, and insomnia in developing countries across regions, based on the two categories of developing countries in terms of the level of development (least developing countries vs emergent and other developing countries). We wanted to reveal heterogeneities to enable evidence-based health care assistance and resource allocation across developing countries, which continue to struggle under the COVID-19 pandemic.

This systematic review and meta-analysis were conducted in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) statement 2019 and registered in the International Prospective Register of Systematic Reviews (PROSPERO: CRD42020220592).

Data sources and search strategy

This study is built upon the existing meta-analyses; we contacted their authors to ask for their coding data for aggregation, since relevant meta-analyses already exist at the scale of individual countries or regions. We searched PubMed, Embase, PsycINFO, Web of Science, medRxiv, and Google Scholar in English for meta-analyses on mental health symptoms of the key adult populations during COVID-19 from January 1, 2020 to September 22, 2021. For example, the following Boolean operators on three sets of keywords were used in Web of Science:

- (ALL= ((2019-nCoV OR 2019nCoV OR COVID-19 OR SARS-CoV-2 OR (Wuhan AND coronavirus)) AND (“depressi*” OR “anxi*” OR “insomnia” OR “sleep” OR “distress” OR “PTSD” OR “post-traumatic stress disorder” OR “mental health” OR “psychiatric”))) AND (TS= “meta-analysis”).

The search targeted meta-analyses that focused on the prevalence of anxiety, depression, distress, insomnia, and PTSD in specific regions or countries during COVID-19. Meta-analyses that did not specify regions or countries were excluded. When multiple meta-analyses existed on the same region, the most comprehensive analysis was chosen. We contacted the authors of these meta-analyses to request their original coding data. Figure S1 in the [Online Supplementary Document](#) details the flowchart of the overall search process.

Selection criteria

To be included in this review, the evidence must have studied the prevalence of at least one mental symptom outcome (eg, anxiety, depression, distress, insomnia, and PTSD) of adult populations such as frontline health care workers (HCW), general HCWs, general adult population, medical students, and general adult students in any developing countries based on the definition of [11] during the COVID-19 pandemic, published in English.

We excluded empirical studies using the following criteria:

1. Population: children, adolescents, or specific niche adult populations such as COVID-19 patients, inpatients, or other patients, adults under quarantine, or pregnant/postpartum women in developing countries

2. Methodological approaches: non-primary studies such as reviews, qualitative or case studies, interventional studies, interviews, or news reports
3. Measurements: non-validated mental health instruments (i.e., self-made questionnaires) or instruments without a validated cut-off score to calculate a prevalence rate (i.e., STAI, SCL-90 for anxiety and depression).

Selection process and data extraction

The coding data from the original meta-analyses were reviewed and recoded based on a single pre-developed coding protocol to ensure the consistency and comparability of the results [10]. If the authors of the existing meta-analyses did not share the coding data, we identified their original empirical studies, independently extracted the relevant data into a coding book based on the same coding protocol [10] and cross-checked their coding. We assessed the eligibility of each study by reading its full text to remove redundant empirical studies and code relevant information such as the authors and year of the study, title, publication status, sample locations, date of data collection, sample size, response rate, population, age (mean, SD, min, and max), gender proportion, instruments, cut-off scores used, the prevalence/mean/SD of the mental health outcome, and other notes or comments. In cases where the two coders disagreed, a lead researcher checked the study independently to determine its coding. The lead researcher integrated all the coding information and reviewed the key information such as mental outcomes, instruments, outcome levels, prevalence, population, sample, and regions.

To consistently analyse the data, we verified the independence of mental health symptoms and samples. For instance, if a study used more than one instrument to measure a mental health outcome, we reported the results based on the most popular instrument. We used the three typical cut-off levels of mental health symptoms (above mild, above moderate, and severe) as standards for reporting the prevalence above mild, moderate, and severe levels. If an empirical study reported the prevalence rates differently from the three-level norm with cut-off points, such as at overall level, we converted the prevalence rates into above mild, above moderate, or severe based on the typical cut-off points of the instruments used.

Assessment of bias risk

The Mixed Methods Appraisal Tool (MMAT) was used as a quality assessment tool [12-14]. Two reviewers independently assessed scores (ranging from 0 [low] to 7 [high]) for the studies using the tool dictionary and guidelines, crosschecked their coding, and resolved disagreements. Studies were categorized as high, medium, or low quality based on the score of >6, 5-6, or <5, respectively.

Statistical analysis

A random-effects model was used (the *metaprop* package in version 16.1 of Stata) to compute the pooled estimates of outcome prevalence between populations by assuming that these studies were randomly selected from their targeted populations [15].

Given the high degree of heterogeneity of the true differences in the effect sizes [16], we ran a meta-regression to regress the prevalence upon outcomes (five types of mental health symptoms), severity of outcome (above mild/above moderate/severe), five major population groups (frontline HCWs, general HCWs, general population, adult students, medical students), and on continents or regions, sample size, research design, and study quality. Given the size of Asia, which contains 60% of the world population, we used the sub-continental regions of Asia (Central, East, Southeast, West, and South). The other continents were not subdivided due to being smaller both in terms of populations as well as the number of conducted studies, so that the regions do not contain too few samples.

The meta-analytical results of our study enable the prediction of prevalence rates while accounting for multiple factors at the same time, thus offering a superior model over prior meta-analyses, which accounted for predictors separately [17,18]. Hence, based on the results of meta-regression, we predicted the prevalence rates of anxiety, depression, and insomnia symptoms at mild above, moderate above, and severe for frontline HCWs, general HWCs, general population, and general students in the seven regions. Due to small sample size, we did not predict the prevalence rates on distress and PTSD or medical students. The statistical significance is taken at the 95% confidence interval level.

The DOI plot and the Luis Furuya-Kanamori index [19] were constructed to assess publication bias [20,21]. We used event ratio as the primary effect measure for the pooled estimates.

RESULTS

Study screening

The search generated smaller elementary meta-analyses on mental health symptoms during COVID-19 [9,10,18,22-29], and we were able to obtain the original coding results from seven of them. The aggregation resulted in a total of 461 studies, 341 of which were unique studies that fit the criteria for this meta-analysis of developing countries (Figure S1 in the **Online Supplementary Document**).

Table 1. Characteristics of the studies on mental health in developing countries during the COVID-19 pandemic

CHARACTERISTICS	TOTAL NUMBER OF STUDIES/SAMPLES*	PERCENTAGE (%)	LEVEL OF ANALYSIS
Overall	341/404	100.00	
Population	404	100.00	Sample
Frontline HCWs	73	18.07	
General HCWs	126	31.19	
General population	145	35.85	
Adult students	43	10.64	
Medical students	17	4.21	
Outcome †:	1433	100.00	Prevalence
Anxiety	650	45.36	
Depression	551	38.45	
Distress	38	2.65	
Insomnia	158	11.03	
PTSD	36	2.51	
Severity:	1433	100.00	Prevalence
Mild above	557	38.87	
Moderate above	542	37.82	
Overall	21	1.47	
Severe	313	21.84	
Region:	343†	100.00	Study
Africa	29	8.50	
Asia:			
East Asia	147	43.11	
South Asia	48	14.08	
Southeast Asia	20	5.87	
West Asia	18	5.28	
Europe	18	5.28	
Latin America	61	17.89	
Oceania	0	0.00	
Design:	341	100.00	Study
Cross-sectional	329	96.48	
Cohort	12	3.52	
Publication status:	341	100.00	Study
Preprint	15	4.40	
Published	326	95.60	
Quality:	341	100	Study
High	48	14.80	
Medium	235	68.91	
Low	58	17.10	
	Medium (Mean)	Range	
Number of participants	535 (4218)	257-1252	Sample
Female portion	65.55% (64.57%)	12.10%-76.10%	Study
Response rate	83.30% (75.04%)	63%-95.12%	Study

PTSD – Posttraumatic stress disorder

*A study may include multiple independent samples. An independent sample in a study may report anxiety, depression, and insomnia at the levels of mild above, moderate above, and severe. Hence, the total number of prevalence rates is larger than the total number of independent samples.

†Two studies included studies from different regions.

Study characteristics

Table 1 summarizes the study characteristics of the 341 empirical studies with a total of 1 704 072 participants in 404 samples. Out of the 404 samples, 73 (18.07%) were from frontline HCWs, 126 (31.19%) from general HCWs, 145 (35.85%) from the general population, 43 (10.84%) from adult students, and 17 (4.21%) from medical students. Table S1 in the **Online Supplementary Document** reported the study characteristics of each of the 341 studies. Among the 404 samples, about one-third investigated the general population (35.85%) or general HCWs (31.19%), almost one-fifth studied frontline HCWs (18.07%), one-tenth focused on general adult students (10.64%), and only 4.21% studied medical students.

More than 80% of the studies covered anxiety and depression symptoms (45.36% and 38.87%, respectively). Just over one-tenth investigated insomnia symptoms (11.03%); few studies investigated PTSD (2.65%) and distress (2.51%). The studies reported the prevalence rates using cut-offs at the “mild above” (38.87%), “moderate above” (37.82%), and “severe above” (21.84%) level of symptom severity.

Almost all studies, 329 in total, employed cross-sectional surveys, with only 12 cohort studies (3.52%). The MMAT indicated 48 (14.08%) studies were of good quality (score >6/7), 235 (68.91%) studies were of medium quality (score = 5-6), and 58 (17.01%) studies were of low quality (score <5). The median number of individuals per sample was 535 (range = 19-746 217).

We next break down the studies by country and region. **Table 2** and **Figure 1** show that, in total, there were 50 countries from four continents with at least one mental health prevalence study under COVID-19: Africa (11 countries studied out of 56 developing countries), Asia (17 countries studied out of 49 developing countries), Europe (nine countries studied out of 15 developing countries), and Latin America (13 countries studied out of 37 developing countries). There have been no studies in the continent of Oceania (zero countries out of 10 developing countries). Within Asia, there have been studies in East Asia, Southeast Asia, South Asia, and West Asia, but not in Central Asia (five developing countries). However, few studies collected samples from multiple countries or regions. For example, one study [30] collected data from both Africa and Latin America, while another [31] collected data from five Southeast

Table 2. Country distribution of mental health prevalence studies in developing countries during COVID-19 pandemic

CONTINENT*	REGION	COUNTRY‡	N OF STUDY	PERCENTAGE	
Africa	Northern Africa	11 countries studied out of 56 developing countries	29	8.45	
			13	3.78	
		Egypt	6	1.75	
		Libya	3	0.87	
		Morocco	2	0.58	
		Tunisia	2	0.58	
	Sub-Saharan Africa			16	4.67
			Cameroon	2	0.58
			Democratic Republic of the Congo	1	0.29
			Ethiopia	7	2.04
			Mali	2	0.58
			Nigeria	2	0.58
			South Africa	1	0.29
			Togo	2	0.58
Asia		17 countries studied out of 49 developing countries	236	68.22	
	Central Asia	0 countries studied out of 5 developing countries	0	0.00	
	East Asia	1 country studied out of 6 developing countries			
		China	147	42.86	
	Southeast Asia		5 countries studied out of 9 developing countries	20	5.83
			Indonesia	2	0.58
			Malaysia	10	2.92
			Philippines	1	0.29
			Thailand	2	0.58
			Vietnam	5	1.46
	South Asia		5 countries studied out of 9 developing countries	49	14.29
			Bangladesh	10	2.92
			India	24	7.00
			Nepal	6	1.75
			Pakistan	8	2.33
			Sri Lanka	1	0.29
		West Asia		6 countries studied out of 18 developing countries	20
			Saudi Arabia	1	0.29
			Jordan	2	0.58
			Kuwait	1	0.29
	Oman		1	0.29	
	Saudi Arabia		6	1.75	
	Turkey	7	2.04		
Europe	Eastern Europe	9 countries studied out of 15 developing countries	18	5.25	
		Albania	2	0.58	
		Bosnia and Herzegovina	1	0.29	
		Bulgaria	1	0.29	
		Kosovo	1	0.29	
		Poland	4	1.17	
		Romania	1	0.29	
		Russia	3	0.87	
		Serbia	4	1.17	
		Ukraine	1	0.29	
Latin America (including the Caribbean)	South America	13 countries studied out of 37 developing countries	62	18.08	
			53	15.46	
		Argentina	7	2.04	
		Bolivia	1	0.29	
		Brazil	32	9.33	
		Chile	1	0.29	
		Colombia	1	0.29	
		Ecuador	3	0.87	
		Paraguay	1	0.29	
		Peru	6	1.75	

Table 2. continued

CONTINENT*	REGION	COUNTRY‡	N OF STUDY	PERCENTAGE
Latin America (including the Caribbean)	Caribbean		2	0.58
		Haiti	1	0.29
		Trinidad and Tobago	1	0.29
	Central America		7	2.04
		Mexico	6	1.75
		Panama	1	0.29
		Did not report country name	1	0.29
Oceania		0 countries studied out of 10 developing countries	0	0.00

*Two continents, Antarctica and North America, are not listed as they do not have developing countries.

‡The developing countries are defined based on the IMF definition.

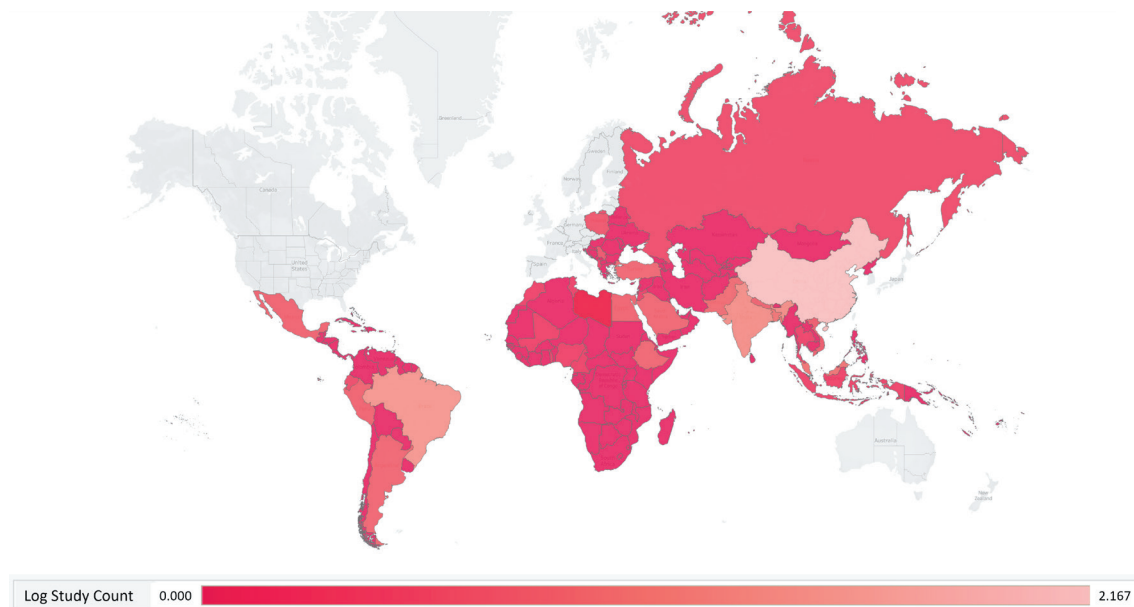


Figure 1. Country distribution of mental health prevalence studies in developing countries during COVID-19 pandemic.

and South Asia countries. Overall, our analysis contains seven regions: Africa, East Asia, Europe, Latin America, South Asia, Southeast Asia, and West Asia.

The 341 papers employed a wide arrange of instruments to assess mental health (Table 3). The popular measures are GAD (50.18%), DASS (18.64%), and SAS (11.83%) for anxiety, PHQ (49.38%) and DASS (21.99%) for depression, K6 (29.93%); for distress, ISI (68.75%) and PSQI (20.31%) for insomnia, and PCL (54.55%), and IES (36.36%) for PTSD.

Major issues from findings of the key study characteristics

Our analysis reveals several widespread issues in mental health research during COVID-19, such as a wide array of used instruments, inconsistent reporting of prevalence rates, inconsistent use and reporting of cut-off points, varied cut-off values for determining the overall prevalence as well as the severity, and other issues on reporting standards and terminologies. Table 3 summarizes popular instruments used for measuring the five mental health symptoms with their primary cut-off point and different variants. Table S2 in the **Online Supplementary Document** summarizes the full list of instruments used by the individual studies included by this meta-analysis. All these issues may contribute to the heterogeneity and confusion in accumulating evidence.

A myriad of instruments: The individual studies on mental health research during COVID-19 employed a wide variety of instruments with varying degrees of popularity and validity, making it challenging to compare or accumulate evidence.

Admixed outcome severity level: The individual studies reported the prevalence rates at a range of symptom severity. First, the studies use different terminologies when reporting the overall prevalence rates. The overall

Table 3. Popular instruments measuring mental health symptoms and their cut-off points

Outcome	Instrument	Outcome level	PRIMARY CUT-OFF		Reference	CUT-OFF VARIANTS		Reference
			Lower bound	Upper bound		Lower bound	Upper bound	
Anxiety	GAD-7	Mild	5	9	[32]	4, 7	10	[33,34]
		Moderate	10	14		11	16	
		Severe	15	21		17	22	
	SAS	Mild	50	59	[35]	45	N/A	[36]
		Moderate	60	69		N/A	59	
		Severe	70	N/A		60	74	
	DASS-21	Mild	7	9	[37]	8	9	[38,39]
		Moderate	10	14		7	N/A	
		Severe	15	42		15	19	
	BAI	Mild	8	15	[40]	22	35	[41]
		Moderate	16	25		36	63	
		Severe	26	63				
	HADS	Mild	8	10	[42]	7	10	[43]
		Moderate	11	14		10	21	
		Severe	15	21				
Depression	PHQ-9	Mild	5	9	[44]	6	8	[45]
		Moderate	10	14		9	14	
		Severe	15	27		15	27	
	DASS-21	Mild	10	12	[46]	1, 9, 10	13	[47-49]
		Moderate	13	20		14	20	
		Severe	21	42		11	N/A	
	HADS	Mild	8	10	[42]	7	10	[43]
		Moderate	11	14		10	21	
		Severe	15	21				
	CES-D-10	Mild	9	13	[50]	10	N/A	[33,51]
		Moderate	14	N/A		11	16	
	Insomnia	ISI	Mild	8	14	[52]	9	
Moderate			15	21	10		N/A	
Severe			22	28				
AIS	Mild above	6	N/A	[54]	5	N/A	[55]	
	PSQI	Overall	10	N/A	[56]	8	N/A	[57]
PTSD	IES-R	Mild	20	N/A	[58]	9, 18, 24	N/A	[59-61]

PHQ – Generalized Anxiety Disorder scale, DASS – Depression Anxiety Stress Scale, SAS – Self-rating Anxiety Scale, HADS – Hospital Anxiety Depression Scale, BAI – Beck Anxiety Inventory, HAMA – Hamilton Anxiety Rating Scale, PHQ – Patient Health Questionnaire depression scale, CES – Center for Epidemiologic Studies Depression Scale, ISI – Insomnia Severity Index, PSQI – Pittsburgh Sleep Quality Index, AIS – Athens Insomnia Scale, PSQI – Pittsburgh Sleep Quality Index, IES – Impact of Event Scale – Revised, PTSD – Posttraumatic stress disorder

prevalence rate could indicate the percentage with moderate symptoms or above, or mild symptoms or above (eg, [62]). Even worse, many studies did not specify if the overall prevalence rate used cut-off at the level of above mild or above moderate. Second, some studies use other terminologies, such as “extremely severe” [63], “very severe” [64], or “very high” [65], “moderate-severe” [66], “moderate to severe” [67,68], “moderately severe” [69], and “poor” (40), making it even more challenging to categorize symptoms. We manually recoded all the studies that indicated their cut-off scores.

Clarity on the cut-off points used to determine severity: Some studies employed non-standard or unusual cut-off scores [70], at times without referencing validation studies that supported the use of those special cut-off scores (eg, [50,71]). Some studies did not report the cut-off score used or did not provide any references [72,73], making the comparison and accumulation difficult. All cut-off variants of the 5 mental health symptoms are listed in **Table 3**.

Pooled prevalence rates of mental health symptoms

Table 4 reports the pooled prevalence rates of mental health symptoms by subgroups of population, outcome, severity, and region. The meta-analyses generally found mental health symptoms to be highly prevalent yet different across regions. Comparatively, Africa had the worst overall mental health symptoms (39%), followed

Table 4. The pooled prevalence rates of mental health disorders by subgroups of population, outcome, severity, and region

FIRST-LEVEL SUBGROUP	SECOND-LEVEL SUBGROUP	SAMPLE SIZE (N)*	PREVALENCE (%)	95% CI
Aggregated			26%	25, 27
Region	Africa	15 391	39%	35, 44
	Asia:			
	East Asia	1 402 610	18%	16, 19
	South Asia	44 184	32%	28, 35
	Southeast Asia	20 914	20%	16, 24
	West Asia	13 289	36%	30, 42
	Europe	15 023	32%	26, 38
	Latin America	196 411	35%	32, 37
Population	Frontline HCWs	71 539	27%	24, 29
	General HCWs	123 698	25%	23, 27
	General population	697 481	23%	21, 25
	Adult students	796 214	30%	27, 32
	Medical students	18 890	38%	32, 44
Outcome†	Anxiety	1 257 838	25%	24, 27
	Depression	321 495	27%	26, 29
	Distress	76 074	29%	21, 37
	Insomnia	41 440	24%	21, 27
	PTSD	10 975	20%	13, 27
Severity‡	Above mild	1 213 070	43%	41, 44
	Above moderate	385 010	21%	20, 23
	Above severe	158 396	8%	7, 9
	Overall	61 346		

CI – confidence interval, PTSD – posttraumatic stress disorder, HCW – health care worker

*The total independent samples are larger than the number of studies because some studies included multiple samples.

†The total sample sizes are larger than the total sample of the 404 independent samples because one sample can assess multiple mental health outcomes.

by West Asia (36%) and Latin America (35%). Among different populations, medical students had the worst overall mental health symptoms (38%), followed by general students (30%) and frontline HCWs (27%). Adults suffered most from distress symptoms (29%), followed by depression (27%) and anxiety (25%). Overall, a whopping 43% of adults in developing countries suffered from mild above mental health symptoms, 21% suffered moderate above, and 8% severe mental health symptoms.

The results of subgroup analyses of popular instruments of mental health symptoms show the various instruments lead to different results (Table 5). While the prevalence rates of anxiety measured by GAD (27%) and DASS (29%) are relatively close, they are significantly different from those measured by SAS (7%), HADS (39%), and BAI (17%). The prevalence rates of depression differ significantly among studies with different measurements, specifically, PHQ (30%), DASS (26%), HADS (33%), SDS (12%), and CES (36%). The prevalence rates of distress measured by DASS (30%) and CPDI (27%) are significantly higher than those measured by K6 (18%). The prevalence rates of insomnia measured by ISI are 22%, which was significantly different from those measured by PSQI (29%) and AIS (44%). While the prevalence rates of PTSD are 15% when measured by PCL by 36% measured by IES. At least partially due to the popularity and standardized usage, the anxiety symptoms measured by GAD-7, GAD-2, and DASS-21 appear more comparable than those measured using other measurements. Similarly, the depression symptoms measured by PHQ-9 or DASS-21 and the insomnia symptoms by ISI appear to be more comparable.

Meta-regression on the prevalence of mental health symptoms

To better explain the heterogeneity of the prevalence of mental health symptoms, Table 6 reports the results of a meta-regression analysis. The meta-analytical model explained over 51% of the variance of mental health

Table 5. Instruments measuring mental health symptoms and the results of subgroup analyses on instruments*

INSTRUMENT	FREQUENCY	PERCENTAGE	PREVALENCE RATE (%)†
Anxiety	279		
GAD (GAD-7/GAD-2)	140	50.18%	27 (26, 29)
DASS-21	52	18.64%	29 (26, 33)
SAS	33	11.83%	7 (6, 9)
HADS	23	8.24%	39 (30, 48)
BAI	6	2.15%	17 (5, 34)
HAMA	4	1.43%	28 (14, 44)
Depression	241		
PHQ (PHQ-9/PHQ-2)	119	49.38%	30 (28, 32)
DASS-21	53	21.99%	26 (22, 30)
HADS	21	8.71%	33 (24, 42)
SDS	18	7.47%	12 (08, 16)
CES (CES-D-20/9/10)	12	4.98%	36 (26, 47)
Distress	27		
K-6	8	29.63%	18 (12, 26)
DASS	4	14.81%	30 (19, 43)
CPDI	4	14.81%	27 (9, 91)
Insomnia	64	100%	
ISI	44	68.75%	22 (19, 25)
PSQI	13	20.31%	29 (21, 38)
AIS	4	6.25%	44 (31, 58)
PTSD	22		
PCL	12	54.55%	15 (6, 25)
IES	8	36.36%	36 (22, 51)

PHQ – Generalized Anxiety Disorder scale, DASS – Depression Anxiety Stress Scale, SAS – Self-rating Anxiety Scale, HADS – Hospital Anxiety Depression Scale, BAI – Beck Anxiety Inventory, HAMA – Hamilton Anxiety Rating Scale, PHQ – Patient Health Questionnaire depression scale, SDS – Self-rating Depression Scale, CES – Center for Epidemiologic Studies Depression Scale, K-6 – the six items Kessler mental distress scale, CPDI – COVID-19 Peritraumatic Distress Index, ISI – Insomnia Severity Index, PSQI – Pittsburgh Sleep Quality Index, AIS – Athens Insomnia Index, PCL – Posttraumatic Stress Disorder Checklist – Civilian Version, IES – Impact of Event Scale

*Only instruments used in at least four studies were included.

†Values are expressed with a 95% confidence interval

Table 6. Meta-regression results on mental health symptoms during COVID-19

VARIABLES	COEFFICIENT*	SE	P VALUE
Outcome			
Anxiety (reference):			
Depression	0.06 (0.01, 0.10)	0.02	0.014
Distress	0.09 (-0.04, 0.23)	0.07	0.17
Insomnia	-0.02 (-0.09, 0.05)	0.04	0.57
Posttraumatic stress disorder	0.03 (-0.10, 0.17)	0.07	0.64
Outcome level			
Mild above	0.46 (0.41, 0.51)	0.02	<0.001
Moderate above (reference)			
Severe	-0.42 (-0.48, -0.37)	0.03	<0.001
Population:			
General HCWs	-0.12 (-0.18, -0.06)	0.03	<0.001
General population	-0.11 (-0.17, 0.05)	0.03	<0.001
Frontline HCWs (reference)			
General students	0.01 (-0.07, 0.10)	0.04	0.80
Medical students	0.09 (-0.01, 0.20)	0.05	0.086
Region:			
Africa	0.16 (0.07, 0.25)	0.05	0.001
East Asia	-0.38 (-0.45, -0.30)	0.04	<0.001

VARIABLES	COEFFICIENT*	SE	P VALUE
South Asia (reference)			
Southeast Asia	-0.31 (-0.42, -0.19)	0.06	<0.001
West Asia	0.09 (-0.01, 0.19)	0.05	0.083
Europe	-0.06 (-0.17, 0.06)	0.06	0.34
Latin America	0.08 (-0.01, 0.17)	0.05	0.070
Sample size/1000000	-0.82 (-1.51, -0.13)	0.00	0.018
Quality	0.03 (0.00, 0.05)	0.01	0.064
Published vs Preprint (reference)			
Least developing vs Emerging developing (reference)	-0.12 (-0.22, -0.03)	0.05	0.012
Cross-sectional vs Cohort (reference)			
Constant	1.02 (0.79, 1.25)	0.12	<0.001
R ²	0.51		
Wald X ² (16)	1488		<0.001

HCWs – Healthcare workers; SE – standardized error
*Values are expressed with a 95% confidence interval.

symptoms among these studies ($R\text{-squared}=51.8\%$, $\tau^2=0.15$). The prevalence rates of depression ($P=0.012$) are significantly higher than those of anxiety (reference). The prevalence of severe mental health symptoms is significantly lower than that of moderate mental illness (reference) ($P<0.001$), which is in turn significantly lower than that of mild mental illness ($P<0.001$). The prevalence rates of general HCWs and the general population are significantly lower ($P<0.001$) than those of frontline HCWs (reference) ($P<0.001$).

The prevalence of mental health symptoms of African adults was significantly higher than in South Asia (reference) ($P=0.001$), which in turn was significantly higher than in East Asia ($P<0.001$) and Southeast Asia ($P<0.001$) yet not significantly different from West Asia, Europe, and Latin America ($P>0.05$). The prevalence rates reported by studies with larger sample size are significantly lower than those of studies with smaller sample size ($P=0.018$). The prevalence rates of mental health symptoms in emergent and other developing countries were significantly higher than those in the least developing countries (reference) ($P=0.012$). Analyses of studies with a higher quality rating ($P=0.06$), publication status ($P=0.72$), and research design ($P=0.31$) did not predict significant prevalence rates.

Table 7 shows the predicted prevalence rates of mental health symptoms by populations, outcomes, severity, and regions by the meta-analytical regression model. **Figure 2** and **Figure 3** show the predicted prevalence rates of depression and anxiety symptoms in different countries or regions, respectively.

Sensitivity analysis

Our meta-analytical model considered several factors, such as publication status (insignificant), sample size (significant), and article quality score (insignificant). Furthermore, excluding each study one by one from the meta-analytical model did not significantly alter the findings. A visual inspection of the sensitivity plot, however, revealed that there is significant asymmetry. Figure S2 in the **Online Supplementary Document** reports the DOI plot in combination with the Luis-Kanamori (LFK) index, which has higher sensitivity and power than a funnel plot [74,75]. LFK index scores of ± 1 , between ± 1 and ± 2 , or ± 2 indicate “no asymmetry”, “minor asymmetry”, and “major asymmetry” respectively, and hence the LFK index of 5.61 represents major asymmetry. Therefore, the presence of publication bias is likely for mental health prevalence studies under COVID-19.

DISCUSSION

By systematically reviewing studies on the prevalence rates of mental health symptoms in developing countries worldwide, we were able to identify and reveal the uneven number of studies on mental health symptoms across countries to highlight the need to initiate relevant studies on the neglected countries. The meta-regression results provide better evidence on the prevalence rates of anxiety, depression, and insomnia in developing countries across regions and based on the level of development.

Table 7. The predicted prevalence rates of mental health disorders by populations, outcomes, severity, and region based on the meta-analytical regression model*

	PREVALENCE RATE (%)†						
	Africa	East Asia	Europe	Latin America	South Asia	Southeast Asia	West Asia
Frontline HCWs:							
Sample	K=4, n=1259	K=47, n=66208	K=5, n=1717	K=2, n=1477	K=8, n=2349	K=5, n=1081	K=2, n=819
Aggregated anxiety	43 (39, 47)	19 (16, 21)	32 (28, 37)	39 (35, 43)	35 (31, 39)	21 (18, 26)	39 (35, 44)
Mild above anxiety	61 (57, 65)	35 (32, 38)	51 (45, 56)	58 (53, 62)	53 (49, 58)	38 (33, 43)	58 (53, 63)
Moderate above anxiety	38 (34, 43)	15 (13, 18)	28 (24, 33)	35 (31, 39)	31 (27, 35)	18 (14, 22)	35 (31, 40)
Severe anxiety	20 (16, 23)	4 (2, 5)	12 (9, 16)	17 (14, 20)	14 (11, 17)	5 (3, 8)	17 (13, 21)
Aggregated depression	46 (41, 50)	21 (19, 23)	35 (30, 40)	42 (38, 46)	38 (34, 46)	24 (20, 28)	42 (38, 47)
Mild above depression	64 (60, 68)	38 (35, 41)	53 (48, 59)	60 (56, 64)	56 (52, 60)	41 (36, 46)	61 (56, 65)
Moderate above depression	41 (37, 46)	17 (15, 20)	31 (26, 36)	38 (34, 42)	34 (30, 38)	20 (16, 24)	38 (33, 43)
Severe depression	22 (18, 26)	5 (3, 6)	14 (10, 18)	19 (15, 22)	16 (12, 19)	6 (4, 9)	19 (15, 23)
Aggregated insomnia	42 (37, 47)	18 (15, 21)	32 (26, 37)	38 (34, 43)	34 (30, 39)	21 (16, 26)	38 (33, 44)
Mild above insomnia	60 (55, 65)	34 (30, 38)	50 (44, 56)	57 (52, 61)	52 (47, 58)	37 (32, 43)	57 (51, 62)
Moderate above insomnia	37 (33, 42)	15 (12, 17)	27 (22, 33)	34 (29, 39)	30 (25, 35)	17 (13, 22)	34 (29, 40)
Severe insomnia	19 (15, 23)	3 (2, 5)	11 (8, 15)	16 (12, 20)	13 (10, 17)	5 (2, 8)	16 (12, 21)
General HCWs							
Sample	K=12, n=3928	K=49, n=77532	K=6, n=3445	K=19, n=12821	K=20, n=9393	K=9, n=2747	K=11, n=6066
Aggregated anxiety	37 (33, 41)	14 (12, 16)	27 (23, 31)	33 (30, 36)	29 (26, 33)	17 (13, 20)	34 (30, 38)
Mild above anxiety	55 (51, 59)	29 (27, 32)	45 (40, 50)	52 (48, 55)	47 (43, 51)	33 (28, 37)	52 (48, 56)
Moderate above anxiety	33 (29, 37)	11 (9, 13)	23 (19, 27)	29 (26, 33)	26 (22, 29)	14 (10, 17)	30 (26, 34)
Severe anxiety	15 (12, 18)	2 (1, 3)	8 (6, 11)	12 (10, 15)	10 (7, 12)	3 (1, 5)	13 (10, 16)
Aggregated depression	4 (36, 43)	16 (14, 18)	29 (25, 34)	36 (33, 39)	32 (28, 36)	19 (15, 23)	36 (32, 40)
Mild above depression	58 (54, 62)	32 (29, 34)	47 (42, 52)	54 (51, 58)	50 (46, 54)	35 (31, 40)	55 (50, 59)
Moderate above depression	35 (31, 39)	13 (11, 15)	25 (21, 30)	32 (28, 35)	28 (24, 32)	15 (12, 19)	32 (28, 36)
Severe depression	17 (14, 20)	2 (2, 4)	10 (7, 13)	14 (12, 17)	12 (9, 14)	4 (2, 6)	15 (11, 18)
Aggregated insomnia	36 (32, 40)	13 (11, 16)	26 (21, 31)	32 (29, 36)	29 (24, 33)	16 (12, 20)	33 (28, 37)
Mild above insomnia	54 (50, 59)	28 (25, 32)	44 (38, 49)	51 (46, 55)	46 (42, 51)	32 (27, 37)	51 (46, 56)
Moderate above insomnia	32 (27, 36)	11 (8, 13)	22 (18, 27)	28 (24, 32)	25 (21, 29)	13 (9, 17)	29 (24, 33)
Severe insomnia	14 (11, 18)	1 (1, 3)	8 (5, 11)	12 (9, 15)	9 (6, 12)	2 (1, 4)	12 (9, 16)
General population:							
Sample	K=15, n=7245	K=70, n=484401	K=7, n=6249	K=35, n=170137	K=9, n=14950	K=8, n=14348	K=1, n=1798
Aggregated anxiety	37 (34, 41)	15 (13, 16)	28 (23, 32)	34 (31, 37)	30 (26, 34)	17 (14, 21)	34 (30, 39)
Mild above anxiety	56 (52, 60)	30 (27, 32)	45 (40, 50)	52 (49, 56)	48 (44, 52)	33 (29, 38)	53 (48, 57)
Moderate above anxiety	33 (29, 37)	12 (10, 13)	24 (20, 28)	30 (27, 33)	26 (23, 30)	14 (11, 18)	30 (26, 34)
Severe anxiety	15 (12, 19)	2 (1, 3)	9 (6, 12)	13 (11, 15)	10 (8, 13)	3 (1, 5)	13 (10, 16)
Aggregated depression	4 (36, 44)	17 (15, 19)	3 (26, 35)	37 (34, 40)	33 (29, 36)	9 (16, 23)	37 (33, 41)
Mild above depression	59 (55, 63)	32 (30, 35)	48 (43, 53)	55 (52, 58)	51 (47, 55)	36 (31, 41)	55 (51, 60)
Moderate above depression	36 (32, 40)	13 (12, 15)	26 (22, 31)	32 (29, 36)	29 (25, 32)	16 (13, 20)	33 (28, 37)
Severe depression	17 (14, 21)	3 (2, 4)	10 (7, 14)	15 (12, 17)	12 (9, 15)	4 (2, 6)	15 (12, 19)
Aggregated insomnia	37 (32, 41)	14 (12, 16)	27 (22, 32)	33 (29, 37)	29 (25, 34)	16 (12, 21)	33 (28, 38)
Mild above insomnia	55 (50, 60)	29 (26, 32)	44 (39, 50)	51 (47, 56)	47 (42, 52)	32 (27, 38)	52 (46, 57)
Moderate above insomnia	32 (28, 37)	11 (9, 13)	23 (18, 28)	29 (25, 33)	25 (21, 30)	13 (10, 17)	29 (24, 34)
Severe insomnia	15 (11, 18)	2 (1, 3)	8 (5, 12)	12 (9, 15)	10 (7, 13)	3 (1, 5)	12 (9, 16)
General students:							
Sample	K=0, n=0	K=15, n=769771	K=4, n=3612	K=9, n=12523	K=7, n=4598	K=3, n=2738	K=5, n=2637
Aggregated anxiety		19 (16, 22)	33 (28, 38)	4 (36, 44)	36 (31, 40)	22 (18, 26)	4 (35, 45)
Mild above anxiety		35 (32, 39)	51 (46, 57)	58 (54, 62)	54 (49, 59)	39 (34, 44)	58 (53, 63)
Moderate above anxiety		16 (13, 19)	29 (24, 34)	35 (31, 40)	31 (27, 36)	18 (14, 23)	36 (31, 41)
Severe anxiety		4 (2, 6)	12 (9, 16)	17 (14, 21)	14 (11, 18)	5 (3, 8)	17 (14, 21)
Aggregated depression		21 (18, 25)	36 (30, 41)	42 (38, 47)	38 (34, 43)	24 (20, 29)	43 (38, 48)
Mild above depression		38 (34, 42)	54 (48, 60)	61 (56, 65)	57 (52, 62)	42 (36, 47)	61 (56, 66)
Moderate above depression		18 (15, 21)	31 (26, 37)	38 (34, 43)	34 (29, 39)	21 (16, 25)	38 (33, 43)
Severe depression		5 (3, 7)	14 (10, 18)	19 (16, 23)	16 (12, 20)	7 (4, 10)	19 (15, 24)
Aggregated insomnia		18 (15, 22)	32 (26, 38)	39 (34, 44)	35 (29, 40)	21 (16, 26)	39 (33, 45)
Mild above insomnia		34 (30, 39)	50 (44, 57)	57 (52, 62)	53 (47, 59)	38 (32, 44)	57 (52, 63)
Moderate above insomnia		15 (12, 19)	28 (22, 34)	34 (29, 40)	31 (25, 36)	18 (13, 23)	35 (29, 40)
Severe insomnia		3 (2, 6)	12 (8, 16)	16 (12, 21)	13 (10, 18)	5 (2, 8)	17 (12, 21)

K – number of samples, n – number of participants, HCW – health care worker

*Due to the small number of samples on distress, posttraumatic stress disorder (PTSD), and medical students, it is not meaningful to report the prevalence rates by different regions.

†All values are expressed with a 95% confidence interval.

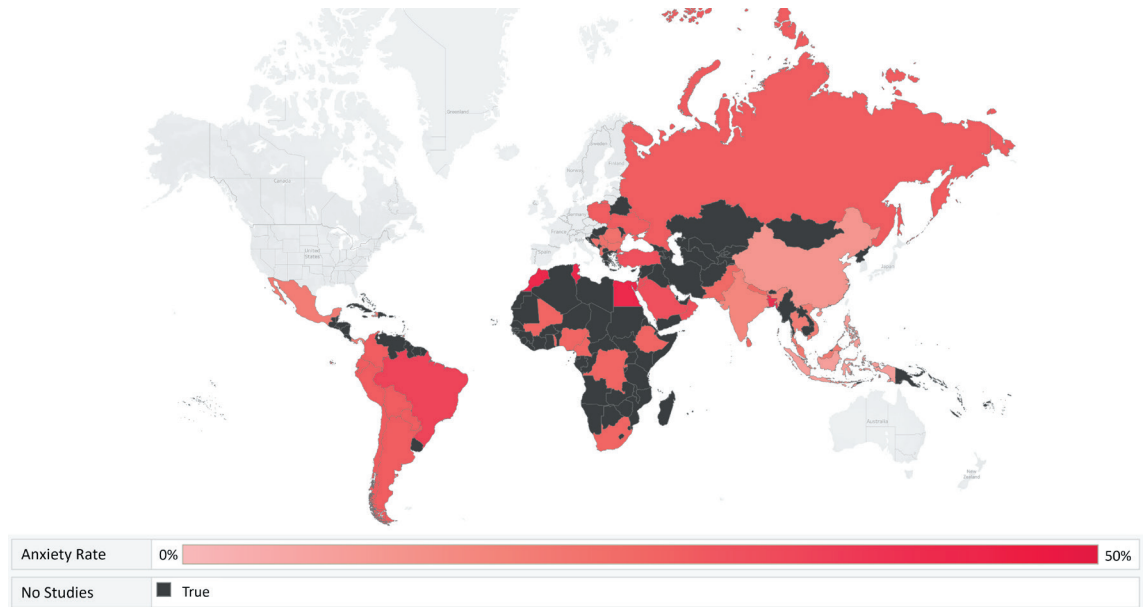


Figure 2. The predicted prevalence rates of depression symptoms by countries or regions, based on the meta-analytical regression model.

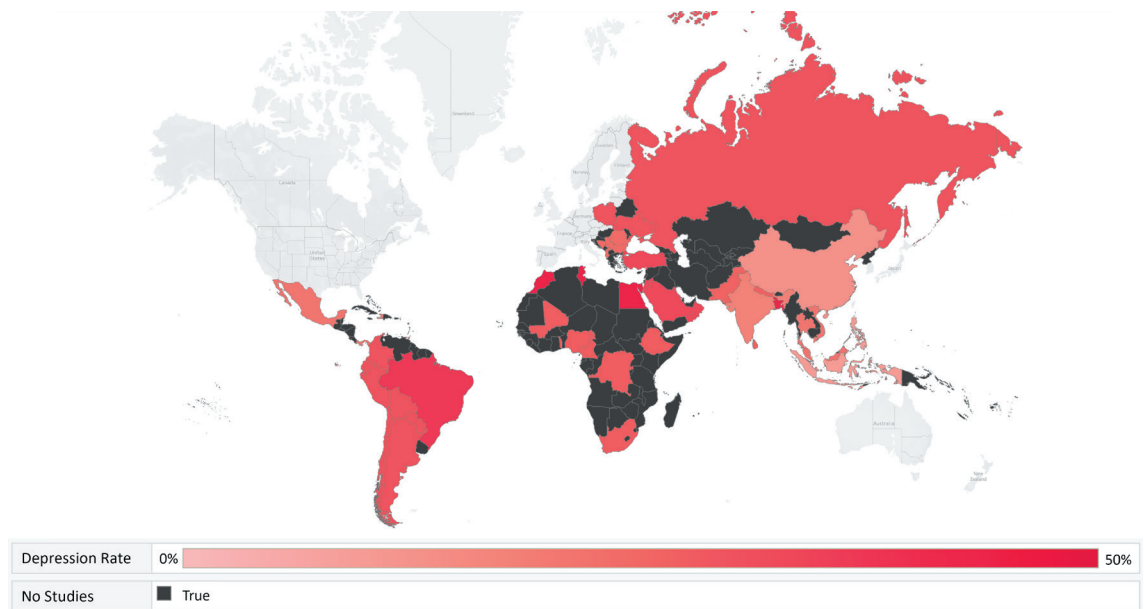


Figure 3. The predicted prevalence rates of anxiety symptoms by countries or regions, based on the meta-analytical regression model.

Uneven research efforts across regions

This study found that the research effort on mental health during COVID-19 has been highly uneven across countries, regions, and mental health outcomes. The studies are far from evenly distributed across countries: there have been 147 studies on China (42.9%), 32 on Brazil (9.3%), 24 on India (7.0%), 10 on Bangladesh (2.9%) and Malaysia (2.9%), yet only 40 out of the 167 developing countries have been studied, leaving 127 developing countries without a single study. Granted, China, India, and Brazil are huge countries, and China was the first to experience the COVID-19 crisis, so a high number of studies in those countries is commendable. Still, there have been no studies in 45 out of 56 countries in Africa (80.4%), 32 out of 49 countries in Asia (65.3%), 24 out of 37 countries in Latin America (64.9%), and 6 out of 15 countries in Europe (40%). The highly uneven distribution of research efforts calls for research attention on the unstudied countries. Given the lack of studies in many countries at the country level, the evidence from our meta-analysis may help by giv-

ing at least the mental health prevalence from nearby countries or regions, to enable health care organizations who need data for evidence-based decision making.

Meta-regression findings

Thanks to a large number of samples in developing countries overall, we were able to conduct meta-regression to account for multiple predictors at the same time to enable better prediction of the prevalence of each mental health symptom. The accumulated evidence shows that several predictors are significantly associated with prevalence rates of mental symptoms during COVID-19, including the severity and type of mental symptoms, population, region, sample size, and study characteristics.

The severity of mental symptoms, largely unaccounted for in prior meta-analyses, was found to contribute greatly to the heterogeneity of prevalence rates; hence, future research on mental health needs to break down and pay special attention to the severity and specify its level. The significant differences revealed by this study call for more meta-analyses on varying levels of severity to provide evidence for practitioners relevant to their concerns.

Among the mental health symptoms examined, distress and depression generally had the highest prevalence rates. Our findings suggest that practitioners need to pay more attention to distress and depression of various populations under the COVID-19 pandemic [76-78].

While not significantly higher than frontline HCWs, general adult students and medical students suffered more than general HCWs and the general population. More than a two-third of studies investigated general HCWs and the general population to generate more meta-analytical evidence, which suggests that policymakers and health care organizations need to further prioritize frontline HCWs and students in this ongoing pandemic. Medical (including nursing) students [79] are worthy of special attention.

A mental health research agenda during COVID-19

Our systemic review and meta-analysis uncovers several widespread problems in the individual papers that impede evidence accumulation. We offer a few concrete suggestions for focusing research and reporting future mental health studies for authors, editors, and reviewers (Table 8), to improve the quality of mental health studies and to facilitate evidence accumulation in future meta-analyses. To make results consistent and comparable, we strongly suggest researchers to use standardized scales with well-established cut-off points (see Table 3 for the popular instruments to measure mental health symptoms and their cut-off points and Table 5 for the results of subgroup analyses on instruments).

Table 8. A list of recommendations for conducting and reporting future mental health research studies

Outcome and instrument	1) Study health outcomes that have higher prevalence rates, eg, distress
	2) Use the well-established instruments with the standard cut-off points listed in Table 3.
Severity of the symptoms	3) Report more levels of severity of symptoms and the cut-off points used
	4) Specify the meaning of overall prevalence, whether above mild or above moderate
	5) Specify the cut-off values used with the reasons/references
Characteristics of the samples	6) Report the sampling dates
	7) Report the age/gender of the participants
	8) Report participant rate
Population	9) Separate and focus on frontline HCWs from general HCWs
	10) Separate and focus on general adult students and medical students
Study design	11) More future research using cohort designs

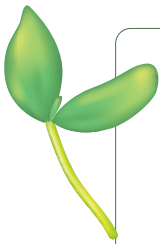
Study limitations and future research

This meta-analysis has a few limitations. First, there may be some bias because all studies were English. Second, 96.48% of studies included in this meta-analysis were cross-sectional surveys, and we call for more cohort studies to examine the effect over time [80]. Third, the validity of our findings rests upon the quality and reporting of the original studies. As discussed before, individual mental health studies varied in their usage of instruments, cut-off scores, the use of cut-off scores to define mental symptoms, and the reporting standards. For example, the overall prevalence refers to “above the cut-off of mild” in some studies, yet “above the cut-off of moderate” in others. Worse, many studies report the overall prevalence without specifying which/how cut-off scores are used. While we focused on the severity, the cut-off points, and the ways in which individual

studies used this information, various approaches contribute to additional noise and variance in the analysis. It is also possible that the diagnostic systems might need to be adjusted across contexts, but such adjustments need to be carefully validated and reported. Fourth, we are limited in examining linear effect, and future research may examine nonlinear effect, as past research has shown age and distance to epicentre may have nonlinear effect on mental health [8,81,82]. Lastly, various classification schemes and terminologies exist on “developing countries” exist, such as low- and middle-income countries (LMIC), newly industrialized countries, emerging markets, third world countries, etc. and future research may use our data to analyse based on other classification schemes.

CONCLUSION

Since the COVID-19 pandemic started in November 2019, hundreds of studies have documented the mental health of major populations by the key mental outcomes and varying levels of severity across the world. This systematic review and meta-analysis synthesized the evidence on the prevalence rates of mental health symptoms in developing countries under the COVID-19 pandemic. We hope this meta-analysis reveals and synthesizes not only the accumulative evidence on mental health research but also reveals key directions for this important research stream.



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Authorship contributions: JC: conceptualization, methodology, validation, formal analysis, investigation, resources, data curation, visualization, writing the original draft, writing and editing the revisions, supervision. SZ: conceptualization, methodology, validation, formal analysis, investigation, data curation, visualization, writing the original draft, writing and editing the revisions, Supervision. AY: investigation (Data). JAY: investigation, resources. All authors were involved in approving the manuscript. The corresponding author attests that all listed authors meet authorship criteria and that no others meeting the criteria have been omitted.

Competing interests: All authors have completed the ICMJE Unified Competing Interest Form (available upon request from the corresponding author) and declare no conflicts of interests.

Additional material

Online Supplementary Document

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