REVIEW Open Access

Mental burden and its risk and protective factors during the early phase of the SARS-CoV-2 pandemic: systematic review and meta-analyses



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Abstract

Background: Mental burden due to the SARS-CoV-2 pandemic has been widely reported for the general public and specific risk groups like healthcare workers and different patient populations. We aimed to assess its impact on mental health during the early phase by comparing pandemic with prepandemic data and to identify potential risk and protective factors.

Methods: For this systematic review and meta-analyses, we systematically searched PubMed, PsycINFO, and Web of Science from January 1, 2019 to May 29, 2020, and screened reference lists of included studies. In addition, we searched PubMed and PsycINFO for prepandemic comparative data. Survey studies assessing mental burden by the SARS-CoV-2 pandemic in the general population, healthcare workers, or any patients (eg, COVID-19 patients), with a broad range of eligible mental health outcomes, and matching studies evaluating prepandemic comparative data in the same population (if available) were included. We used multilevel meta-analyses for main, subgroup, and sensitivity analyses, focusing on (perceived) stress, symptoms of anxiety and depression, and sleep-related symptoms as primary outcomes.

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Results: Of 2429 records retrieved, 104 were included in the review (n = 208,261 participants), 43 in the meta-analysis (n = 71,613 participants). While symptoms of anxiety (standardized mean difference [SMD] 0.40; 95% CI 0.15–0.65) and depression (SMD 0.67; 95% CI 0.07–1.27) were increased in the general population during the early phase of the pandemic compared with prepandemic conditions, mental burden was not increased in patients as well as healthcare workers, irrespective of COVID-19 patient contact. Specific outcome measures (eg, Patient Health Questionnaire) and older comparative data (published ≥ 5 years ago) were associated with increased mental burden. Across the three population groups, existing mental disorders, female sex, and concerns about getting infected were repeatedly reported as risk factors, while older age, a good economic situation, and education were protective.

Conclusions: This meta-analysis paints a more differentiated picture of the mental health consequences in pandemic situations than previous reviews. High-quality, representative surveys, high granular longitudinal studies, and more research on protective factors are required to better understand the psychological impacts of the SARS-CoV-2 pandemic and to help design effective preventive measures and interventions that are tailored to the needs of specific population groups.

Keywords: SARS-CoV-2, COVID-19, Early phase, Psychological distress, Pandemic, Health personnel, Systematic review, Meta-analysis, Anxiety, Depression

Introduction

The emergence of novel severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) was described for the first time in Wuhan, China [1, 2] and declared a public health emergency of international concern on 30 January 2020 [3]. The virus spread rapidly and, as of January 14, 2021, led to 90,759,370 confirmed infections and 1,963,169 deaths worldwide [4].

During the early phase of the pandemic, many countries adopted drastic measures, including testing, tracing, self-isolation, and quarantine measures as well as broader population measures ranging from travel bans, school closures, assembly restrictions, curfews, to full lockdowns [5–7]. Besides substantial stressors for individuals and the general public (eg, social isolation, reduced income, restructuring of school, university, and work life) and healthcare systems (eg, disruption of essential health services) [8, 9], the SARS-CoV-2 pandemic has had major socio-economic consequences for the affected countries (eg, global supply chain disruptions) [10, 11]. By drastically changing our way of social interaction (eg, social distancing), it continues to affect many areas of daily life and in line with this social life and participation.

The disease-related threats, containment measures, and associated stressors may have a negative psychological impact on the community at large and potentially even more so on specific risk groups [12–17]. Given the work-related stressors in the context of disease outbreaks (eg, high workload, risk of infection, triage decisions), healthcare workers may suffer from a particularly high burden [18]^{1,2} [20–22]. Patients with pre-existing

physical or mental conditions (eg, chronically ill individuals, psychiatric patients, geriatric patients), people with confirmed COVID-19 diagnosis, those recovering from the infection, or suffering from long COVID-19, and subgroups with special risk exposure (eg, caregivers) may also be at risk of developing stress-related mental symptoms [15, 22–28].

Various systematic reviews have synthesized the evidence on psychiatric symptoms associated with previous highly contagious infectious disease outbreaks (eg, Ebola, SARS-CoV) and the SARS-CoV-2 pandemic [20, 24, 29-35], some of them also narratively summarizing risk and protective factors for mental health [20, 30, 31, 33, 34]. Several meta-analyses have been conducted, either calculating the pooled prevalence of mental symptoms or odds ratios for the risk of mental burden attributable to the SARS-CoV-2 pandemic [20, 24, 29, 32, 33, 36]. Potential moderators of the negative mental health impact were also partly investigated [32]. International evidence indicates an elevated level of mental symptoms in the general public, including symptoms of anxiety, depression, and stress [30-33, 36]. Confirming the risk status of healthcare workers, several reviews also found an increased prevalence of mental symptoms in this group [18]^{1,2} [20, 29, 31, 32]. Finally, a few studies in patient populations (eg, COVID-19 patients, patients with preexisting mental or physical conditions) show increased mental burden [24, 31–33].

There are several shortcomings of reviews published to date. Most either focus on the general population, health-care workers, or patients, with only few publications examining the level of mental burden across all three specified, most relevant population groups [31–33]. Further limitations included a limited search strategy [31], language restrictions [24, 30, 31, 33], or a missing

¹included pandemic studies.

 $^{^2}$ studies already considered in previous systematic review (Gilan, Röthke and colleagues) [19].

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preregistration [20, 24, 29–31, 33, 36]. Most importantly, all but one systematic review failed to compare the mental burden during an ongoing pandemic with the burden before the pandemic [31]. Such comparisons, however, are necessary to quantify the mental burden specifically attributable to the current pandemic. We therefore aimed to assess the mental health impact of the SARS-CoV-2 pandemic by comparing data from the early phase of the current pandemic with prepandemic data in the general population, healthcare workers, and patients. We aimed to identify population-specific risk and protective factors for mental health.

Methods

Review registration

This systematic review [37] was preregistered with PROSPERO (registration no. CRD42020193249) with the title 'Psychological distress, protective factors and resilience during the SARS-CoV-2 pandemic: a systematic review and meta-analysis with comparison to standard data'. Details of the methods are presented in the Additional file 1. The MOOSE Checklist for Meta-analyses of Observational Studies and differences between the protocol and the final review are presented in eTables 1 and 2.

Search strategy and selection criteria

We searched three bibliographic databases from January 1, 2019 to May 29, 2020 (PubMed, PsycINFO, and Web of Science) and inspected the reference lists of included studies. The search strategy comprised terms associated with mental health, pandemics, and the populations of interest (see eMethods 1 in Additional file 1). There were no restrictions concerning language, publication date, or publication format. We did not consider preprint articles. If not reported within a study, we systematically searched for prepandemic comparative data in the same or a similar population (PubMed, PsycINFO; see eMethods 2).

The populations of interest comprised the general population, healthcare workers, and any patients (eg, COVID-19 patients, those with pre-existing physical or mental conditions; eTable 3). Participants were included irrespective of age, health, or employment status. We did not consider infectious disease outbreaks other than due to SARS-CoV-2. To be eligible for the review, studies had to assess at least one mental health outcome, with a broad range of eligible outcomes (ie, anxiety and worrying, depression, posttraumatic stress, sleep, stress, general psychological distress). These outcomes were also considered for a descriptive synthesis of the prevalence (see data analysis). We included original research articles reporting on cross-sectional and longitudinal surveys.

All pandemic studies meeting these criteria were included but were only taken forward to pairwise metaanalyses if using a validated outcome measure and if prepandemic comparative data were available (eTables 4, 5). These were defined as data collected before the exposure to the current pandemic, and in the absence of other disease outbreaks or macrostressors (eg, disasters), in the same country and population group (if available) and using the same outcome measure. In contrast to the review, we only focused on the four most frequently reported mental health outcomes (primary outcomes), including symptoms related to stress, anxiety, depression, or sleep. Posttraumatic stress, although reported more often than sleep, was not considered for pairwise metaanalyses. As this outcome is usually measured in the aftermath of macro-stressors, we were not able to identify adequate comparative data as mentioned above. Comparative data were selected stepwise using four levels to ensure best available comparability between SARS-CoV-2 exposure ('pandemic') studies and prepandemic ('comparative') studies. If representative studies in the same country and population (level 1) were not available, we used prepandemic studies in the same (level 2) or an alternative population (level 3; eg, healthcare workers compared with the general population), before resorting to the best available data in a similar country (level 4).

Study selection, data extraction, and quality assessment

The study selection process for the pandemic studies at the level of titles/abstracts and full-texts was performed in duplicate by two reviewers independently (NR, LG). Any disagreements were resolved by discussion or by consulting a third reviewer (KL). At both title/abstract (κ = 0.90) and full-text level (κ = 0.97), excellent interrater reliability was achieved.

Relevant information for each included study was extracted in duplicate by two reviewers (NR, LG), working independently, using a customized spreadsheet (eTable 6), which was shortened for the extraction of comparative data. Discrepancies were resolved through discussion or by a third reviewer (KL).

Three independent reviewers (NR, JSW, LG) assessed the quality of included studies using the modified National Institutes of Health (NIH) Quality Assessment Tool for Observational Cohort and Cross-Sectional Studies [38] (eTable 7), with disagreements being resolved by discussion or a third reviewer (KL). The level of comparability between pandemic and comparative data was assessed using a self-developed tool with four levels based on the previously mentioned levels for the stepwise selection of comparative data (eTable 8).

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Data analysis

The included studies were synthesized in narrative and tabular form, with a descriptive analysis of prevalence rates for mental health symptoms (ie, proportion of participants beyond a cut-off score reported in the included study) and of risk and protective factors. If adequate comparative data for any of the primary outcomes were available, pairwise meta-analyses were performed for the general population, healthcare workers, and patients, respectively (eMethods 3). Given the multiple uses of comparative studies, we used multilevel meta-analyses [39] for the general population and healthcare workers, with pandemic studies being clustered according to prepandemic comparators. For patients, the multilevel model reduces to the classic random-effects model as different comparative studies were available. Prediction intervals were calculated in meta-analyses with at least four studies to take the large between-study heterogeneity into account [40].

Two sensitivity analyses referred to the quality of pandemic studies and the level of comparability (see Search strategy and selection criteria), by limiting the analyses to very comparable pandemic and prepandemic studies (ie, level 1 and 2 mentioned above).

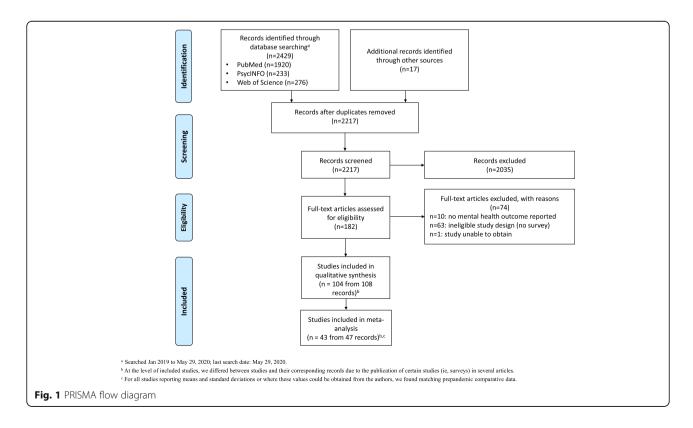
Subgroup analyses for each of the three groups were performed for the surveyed populations (eg, age), characteristics of the pandemic studies (eg, survey start) and of comparative data (eg, publication year), and the relationship of sample sizes in pandemic versus comparative studies, in order to identify potential sources of heterogeneity of the psychological impact of the SARS-CoV-2 pandemic.

Results

Details of the results are presented in the Additional file 2. The systematic search for studies performed during the SARS-CoV-2 pandemic identified 2429 records from database searches and 17 additional records from reference lists, of which 104 studies were included in the review and 43 studies in the metaanalyses (Fig. 1). Of the 104 eligible studies, most studies were performed in the general population (50 studies), followed by 30 studies in healthcare workers, and seven studies in various patient populations. Seventeen studies included mixed samples. Across the three population groups, a total of 208,261 participants ranging from 51 to 52,730 participants [41, 42]¹ from the pandemic studies were included in the review, the number of participants considered in the meta-analyses, in total 71,613, ranged from 127 to 60,213 participants (eTable 9).

The study characteristics of the 104 included pandemic studies (early phase) are presented in Table 1.

Although we imposed no restrictions on the age limits, we identified no studies conducted in children but did find some studies in the general population that included participants below the age of 18 years [47, 58]¹.



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Thus, the mean age of participants in the pandemic studies ranged from 20 (SD not reported) to 56.9 (SD 7.6) years $[49, 99]^1$. The studies covered Asia (67 studies [26, 41–43, 49, 50, 54, 57, 58, 60, 62, 64, 71, 74, 76, 80, 81, 85, 86, 88-91, 93, 98, 101, 102, 104, 109, 111-116, 118-123, 126, 124, 128, 131-133, 136, 137, 140, 141, 143, 145]¹ [18, 48, 59, 73, 84, 87, 92, 100, 105, 106, 108, 117, 134, 138, 139, 142, 144]^{1,2} [127]^{1,3}) thereof from China [42, 49, 50, 57, 58, 60, 62, 64, 71, 80, 81, 85, 86, 88-91, 93, 101, 104, 109, 115, 116, 118-120, 122-124, 126, 128, 131-133, 136, 140, 141, 143, 145¹ [18, 48, 59, 84, 87, 92, 100, 105, 106, 108, 117, 134, 138, 139, 142, 144]^{1,2} [127]^{1,3}, Europe (24 studies) [47, 99, 44–46, 51– 53, 55, 61, 63, 65, 66, 68, 70, 110, 77, 78, 82, 83, 94, 97, 130, 103, 135]¹, North America (six studies) [56, 67, 79, 107, 146, 147]¹, or different continents (seven studies) [69, 72, 75, 95, 96, 125, 129]¹. For 13 studies investigating more than one population, several samples were considered [130–133, 135, 140, 141, 143, 145]¹ [134, 138, 139, 144]^{1,2}. We identified 47 matching prepandemic comparative studies (eTable 10), including one pandemic study reporting adequate comparative data $[127]^{1,3}$ $[148-193]^3$.

Prevalence rates of the six mental health symptoms, that were considered for the review, were available for a varying number of included pandemic studies (Table 2). The proportion of participants beyond a cut-off value in the included studies varied considerably (eg, anxiety in general population: 0.7–64.0%). Based on cut-off values reported in the primary studies (eTable 11), we found increased levels of mental burden during the SARS-CoV-2 pandemic in the general population, healthcare workers, and patients regarding each of the symptoms observed during the current pandemic, that is, without considering the prepandemic situation.

In pairwise meta-analyses comparing pandemic (early phase) with prepandemic data for the four primary outcomes, however, we found only evidence for a small increase of anxiety (standardized mean difference [SMD] 0.40; 95% CI 0.15–0.65; p = .002) and a moderate increase of depressive symptoms (SMD 0.67; 95% CI 0.07–1.27; p = .03) in the general population. No evidence for a change in stress or sleep-related symptoms was identified (Table 3). For healthcare workers compared with healthcare staff before the pandemic, the meta-analyses showed no evidence of any effect on the primary outcomes (Table 3). The same was found for patients (Table 3); however, prepandemic data in patients were only available for four samples. Forest plots are presented in Figs. 2, 3, and eResults 1 in the Additional file 2.

Of the 104 studies, 38 studies were judged to be of fair quality and 57 studies of poor quality, with main

concerns regarding selection bias, the validity of outcome measures, and the description of the sample and the survey period (eTable 12). From nine high-quality studies, four were representative surveys [44, 47, 77, 88]¹. From the 85 pairwise comparisons relevant for meta-analyses, 52 comparisons were of level-1 and 33 of level-2 quality (eTable 13). When excluding low-quality pandemic studies (Table 3), the effects on anxiety and depressive symptoms in the general population increased. The effect on anxiety in the general population was stable in the sensitivity analysis when only best comparable data sets (ie, level-1 and level-2 comparability) were included, while there was no longer evidence for an effect on depressive symptoms (Table 3 and eResults 2 in Additional file 2).

Heterogeneity was considerable in main and sensitivity analyses, with I^2 scores mostly ranging from 90 to 100% and wide prediction intervals (Table 3). We therefore performed subgroup analyses with at least k=5 studies in the main analyses in attempts to explain this heterogeneity (Table 4; eResults 3 in Additional file 2).

Regarding population characteristics (pandemic studies), age was no consistent risk or protective factor. Within the general population, we identified no evidence for a subgroup difference according to stressor exposure except for elevated sleep symptoms in isolated individuals [62]¹. In healthcare workers, there was no evidence for a moderating effect of COVID-19 patient contact on mental health. In different groups of patients, we identified no evidence of differences in anxiety or depression. Compared with COVID-19 patients [131]¹, psychiatric patients reported more stress, with the caveat of few studies [42, 132, 135]¹.

Among general characteristics of the pandemic studies, we found no (consistent) evidence of differences depending on when the surveys started, whether they were conducted in China, or the sample size. We found evidence of an elevated level of depressive symptoms in the general population and patients depending on the specific outcome measure employed (eg, Patient Health Questionnaire [PHQ], Zung Self-Rating Depression Scale [SDS]).

In subgroup analyses for comparative study characteristics, there was no evidence of a consistent moderation of comparison sample sizes.

Across the three populations, we identified a higher level of anxiety and depressive symptoms if included studies were compared to prepandemic data published five or more years before versus a smaller burden in comparison to prepandemic data of less than 2 years ago.

The relationship of sample sizes explained the heterogeneity of the psychological impact of the SARS-CoV-2 pandemic in the general population and patients, with evidence for elevated symptoms of anxiety if similar sample sizes were compared.

³prepandemic comparative studies.

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Table 1 Study characteristics of included main studies

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales
General Popula	tion						
Ahmad et al. (2020) [43] ¹	CS, OBS	Iraq (Kurdistan)	516; 222 (43%); NA (mode: 18–35 years [65.1%])	NA	NA	Anxiety and fear	Binary single item ^a
Bacon et al. (2020) [44] ¹	CS, OBS	United Kingdom	202; 127 (62.9%), 1 diverse; 33.79 (12.48)	NA	March 18– 19, 2020	Anxiety and fear	GAD-7
						Depressive symptoms	BDI-II
Bäuerle et al. (2020) [45] ¹ ,	CS, OBS	Germany	15,037; 10,633 (70.7%), NA (mode: 25–34 years [24.8%])	NA	March 10– May 5, 2020	Anxiety and fear	GAD-7, single item 7-P LS ^a
Teufel et al. (2020) [46] ¹						Depressive symptoms	PHQ-2
						Psychological Distress	DT
Buzzi et al. (2020) [47] ¹	CS, OBS	Italy	2064; NA; NA	100% adolescents	March 2020	Anxiety and fear	4-P LS ^a
Cao et al. (2020) [48] ¹	CS, OBS	China	7143; 4975 (69.7%); NA	NA	NA	Anxiety and fear	GAD-7
Chang et al. (2020) [49] ¹	CS, OBS	China	3881; 2447 (63.1%); 20.00 (NA); P ₂₅₌ 19.00, P ₇₅₌ 22.00]	medical students	January 31, 2019–	Anxiety and fear	GAD-7
				(n = 3359)	February 3, 2020	Depressive symptoms	PHQ-9
Gao J et al. (2020) [50] ¹	CS, OBS	China	4872; 3267 (67.7%); 32.3 (10.0)	NA	January 31– February 02	Anxiety and fear	GAD-7
					2020	Depressive symptoms	WHO-5 ^c
Germani et al. (2020)	CS, OBS	Italy	1011; 720 (71.2%); 24.2 (3.6)	100% age between 18 and 29 years	March 17– 24, 2020	Anxiety and fear	STAI-Y
[51] ¹						Stress	PSS
						Other Outcomes	SDQ
González– Sanguino	CS, OBS	Spain	3480; 2610 (75%); 37–92 (NA)	NA	March 21– 28, 2020	Anxiety and fear	GAD-2
et al. (2020) [52] ¹						Depressive symptoms	PHQ-2
						PTSS	PCL-C-2
						Other outcomes	FACIT-Sp12, MSPSS, SCS
Harper et al. (2020) [53] ¹	CS, OBS	UK	324; 162 (50%); 34–32 (11.71)	NA	March 27– 28, 2020	Anxiety and fear	FCV-19S, PROMIS-SF Anxiety
						Depressive symptoms	PROMIS-SF Depression
						Other outcomes	WHOQOL-BREF
Jahanshahi et al. (2020) [54] ¹	CS, OBS	Iran	1058; 569 (53–8%); NA (mode: 26–35 years)	NA	March 25– 28, 2020	Psychological distress	CPDI
Lauri Korajlija et al. (2020) [55] ¹	CS (repeated), OBS	Croatia	sample 1: 888; 738 ^d (83–1%); 31.3 (10.45) sample 2: 966; 732 ^d (75.8%); 40 (11.94)	NA	1st period: February 24–NA 2nd period: March 19–	Anxiety and fear	11-items 5-P L (based on Swine Flu Anx iety Items, Wheaton et al.

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Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales
					NA		2012) ^a
Lee SA et al. (2020) [56] ¹	CS, OBS	USA	398; 191 (49%); 35.91 (11.73)	NA	March 23– 24, 2020	Anxiety and fear	2 single items 5-P LS ^a
						Other outcomes	Passive suicidal ideation (single item 5-P LS) ^a
Lei et al. (2020) [57] ¹	CS, OBS	China	1593; 976 (61.3%); 32.3 (9.8)	'affected group': quarantined /	February 04–10, 2020	Anxiety and fear	SAS
				relatives quarantined $(n = 420)^b$		Depressive symptoms	SDS
Li Y et al.	CS (part of	China	1442; 891 ^d (61.8%); NA (K-6 <	medical students	February 7–	PTSS	IES-R
(2020) [58] ¹	longitudinal cohort study), OBS		5: 20.0 [1.5]; K-6 ≥ 5: 20.0 [1.6])	(n = 764), nursing students $(n = 211)$, medical technology students $(n = 467)$	13, 2020	Psychological distress	K-6
Liu N et al. (2020) [59] ^{1,2}	CS, OBS	China	285; 155 (54.4%); NA (47.7% < 35)	NA	January 30– February 08, 2020	PTSS	PCL-5
Liu S et al. (2020) [60] ¹	CS, OBS	China	primary school: 209; 116 (56%); NA	primary school students, college	February– March, 2020	Anxiety and fear	3 items, 4-P LS ^a
			college: 198; 130 (62%); NA	students		Other outcomes	SSS
Lopez et al. (2020) [61] ¹	CS, OBS	Spain	878; 544 ^d (62%) or 636 (72% ^d), data in text and Table 1	100% community- dwelling older adults;	NA	Anxiety and fear	a
			inconsistent; NA (mode: 60– 70 years [71% ^d])	age 60–70 (n = 626); age 71–80 (n = 252)		Other outcomes	BRCS, Ryff's PWB (subscales for personal growth and purpose in life)
Ma et al. (2020) [62] ¹	CS, OBS	China	123; 71 ^d (57.7% ^d); 37.4 (10.6)	100% isolated people ^b	January 2020	Anxiety and fear	DASS-21 Anxiety
						Depressive symptoms	DASS-21 Depression
						Stress	DASS-21 Stress
						Sleep-related symptoms	PSQI
						Other outcomes	SF-36
Mazza et al. (2020) [63] ¹	CS, OBS	Italy	2766; 1982 (71.7%); 32.94 (13.2)	NA	March 18– 22, 2020	Anxiety and fear	DASS-21 Anxiety
						Depressive symptoms	DASS-21 Depression
						Stress	DASS-21 Stress
McKay et al. (2020) [64] ¹	CS, OBS	China	908; 752 (82.8%); 40.37 (9.27)	NA	February 24–March 15, 2020	Anxiety and fear	CoVGAD-7, DASS-21 Anxiety
						Depressive symptoms	DASS-21 Depression
Moccia et al. (2020) [65] ¹	CS, OBS	Italy	500; 298 (59.6); NA (mode: 28–37 years, <i>n</i> = 129)	NA	April 10–13, 2020	Psychological distress	K-10
						Other outcomes	TEMPS-A

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Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales
Odriozola- González	CS, OBS	Spain	2530; 1672 (66.1%); 27.9 (12.4)	students ($n = 1944$); administrative staff	March 28– April 3,	Anxiety and fear	DASS-21 Anxiety
et al. (2020) [66] ¹				(n = 247); faculty members and academic staff (n =	2020	Depressive symptoms	DASS-21 Depression
				339) ^b		Stress	DASS-21 Stress
						PTSS	IES
Olagoke et al. (2020) [146] ¹	CS, OBS	USA	501; 277 (55.29%); 32.44 (11.94)	NA	March 25, 2020–NA	Depressive symptoms	PHQ-2
						Other outcomes	Perceived self- efficacy (Ajzen 2002)
Ozamiz- Etxebarria	CS, OBS	Spain	976; 792 (81.1%); NA (mode: 18–25 years [56.5%])	NA	March 11– 15, 2020	Anxiety and fear	DASS-21 Anxiety
et al. (2020) [68] ¹						Depressive symptoms	DASS-21 Depression
						Stress	DASS-21 Stress
Özdin et al. (2020) [69] ¹	CS, OBS	Turkey	343; 169 (49.2%); 37.2 (10.3)	NA	April 14–16, 2020	Anxiety and fear	HAI
						Depressive symptoms	HADS
Perez– Fuentes et al. (2020) [70] ¹	CS, OBS	Spain	1014; 681 (67.2%); 40.87 (12.42)	NA	March 18– 23, 2020	Depressive symptoms	BIP-Q5
Qiu et al. (2020) [41] ¹	CS, OBS	China, Hong Kong, Macao, Taiwan	52,730; 34,131 (64.7%)	NA	January 31– February 2, 2020	Psychological distress	CPDI
Ren et al. (2020) [71] ¹	CS, OBS	China	1172; NA; NA	NA	February 14–March 29, 2020	Anxiety and fear	GAD-7
					29, 2020	Depressive symptoms	PHQ-9
						Stress	PSS-10
						Sleep-related symptoms	ISI
						PTSS	PCL-5
						Other outcomes	MINI suicidality module
Reznik et al. (2020) [72] ¹	CS, OBS	Russia & Belarus	850; 622 (73.2%); 34.8 (13.0)	NA	after March 27, 2020	Anxiety and fear	FCV-19S
Roy et al. (2020) [73] ^{1,2}	CS, OBS	India	662; 339 (51.2%); 29.09 (8.83)	NA	March 22– 24, 2020	Anxiety and fear	18 items 5-P LS ^a
Sakib et al. (2020) [74] ¹	CS, OBS	Bangladesh	8550; 3760 (44%); 26.5 (9.1)	NA	April 1–10, 2020	Anxiety and fear	FCV-19S
						Depressive symptoms	PHQ-9
Satici et al. (2020) [75] ¹	CS, OBS	Turkey	1304; 917 (70.3%); 29.5 (10.5)	NA	NA	Anxiety and fear	DASS-21 Anxiety, FCV- 19S
						Depressive symptoms	DASS-21 Depression
						Stress	DASS-21 Stress
Shammi et al.	CS, OBS	Bangladesh	1066; 405 (38.5%); 27.80	NA	March 28–	Psychological	COVID-19

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Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales	
(2020) [76] ¹			(10.05)		30, 2020	distress	related mental distress (5 items 5-P LS) ^a	
Shevlin et al. (2020) [77] ¹	CS, OBS	UK	2025; 1047 (51.9%); 45.4 (15.9)	NA	March 23– 28, 2020	Anxiety and fear	GAD-7, VAS on COVID-19 anxiety	
						Other outcomes	PHQ-15	
Soraci et al. (2020) [78] ¹	CS, OBS	Italy	249; 229 (92%); 34.50 (12.21)	NA	March 18– 21, 2020	Anxiety and fear	FCV-19S, HADS	
Sutin et al. (2020) [147] ¹	CS, OBS	USA	2094; 1024 (48.9%) ^d ; 51.03 (16.58)	overweight ($n = 706$); obesity ($n = 587$)	mid–March, 2020	Anxiety and fear	13 items 5-P LS ^a	
Tan W et al. (2020) [80] ¹	CS, OBS	China	673; 172 ^d (25.6% ^d); 30.8 (7.4)	NA	February 24–252,020	Anxiety and fear	DASS-21 Anxiety	
						Depressive symptoms	DASS-21 Depression	
						Stress	DASS-21 Stress	
			,	Sleep-related symptoms	ISI			
							PTSS	IES-R
Tian et al. (2020) [81] ¹	CS, OBS	China	1060; 511 (48.2%); 35.01 (12.8)	HCW (n = 42), students (n = 330)	January 31– February 02,	Anxiety and fear	SCL-90 Anxiety	
				20.	2020	Depressive symptoms	SCL-90 Depression	
						Psychological distress	SCL-90 GSI	
						Other outcomes	SCL-90 subscales	
Tsipropoulou et al. (2020)	CS, OBS	Greece	2970; 2153 (72.5%); NA (mode: 18–30 years [52%])	NA	NA	Anxiety and fear	FCV-19S, GAD- 7	
[82] ¹						Depressive symptoms	PHQ-9	
Tull et al. (2020) [79] ¹	CS, OBS	USA	500; 235 ^d (47%); 40 (11.6)	NA	March 27– April 5,	Anxiety and fear	DASS-21 Anxiety, SHAI	
					2020	Depressive symptoms	DASS-21 Depression	
						Stress	DASS-21 Stress	
Voitsidis et al. (2020) [83] ¹	CS, OBS	Greece	2363; 1800 (76.2%); NA (mode: 18–30 years [55%])	NA	April 10-13, 2020	Anxiety and fear	a	
						Depressive symptoms	PHQ-2	
						Sleep-related symptoms	AIS	
						Other outcomes	IUS-12, JGLS	
Wang C et al. (2020a)	(repeated),	China	1738 not counting participants in both surveys;	NA	January 31– February 2,	Anxiety and fear	DASS-21 Anxiety	
[84] ^{1,2} , Wang C et al. (2020b) [85] ¹	OBS		333 in both 1st survey: 1210; 814 ^d or 878 ^d (67.3%); NA (mode: 21.4–30.8	d G	Febru	2020 and February 28–March 1,	Depressive symptoms	DASS-21 Depression
, , , , , , , , , , , , , , , , ,			years [53.1%])		2020	Stress	DASS-21 Stress	
			2nd survey: 861; 646 ^d (75%);			PTSS	IES-R	

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Table 1 Study characteristics of included main studies (*Continued*)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales
			NA (mode: 21.4–30.8 years [46.5%])				
Wang H et al. (2020) [86] ¹	CS, OBS	China	1599; 1068 (66.8%); 33.9 (12.3)	NA	February 1– 4, 2020	Psychological distress	K-6
Wang Y et al.	CS, OBS	China	600; 333 (55.5%); 34 (12)	NA	February 6–	Anxiety	SAS
(2020) [87] ^{1,2}					9, 2020	Depressive symptoms	SDS
Yang H et al. (2020) [88] ¹	CS (repeated), OBS	China	during COVID-19: 3000; 1500 ^d (50%); 34.7 (NA)	NA	end of December 2019 and mid– February, 2020	Other outcomes	Emotional well-being (Kahneman and Deaton, 2010)
Yuan R et al. (2020) [89] ¹	CS, OBS	China	parents of children hospitalised during the	EH $(n = 50)^b$, NEH $(n = 50)^b$	NA	Anxiety	HADS Anxiety, VDAS
			epidemic (EH): 50; 31 (62% ^d); 36.80 (5.20) parents of children hospitalised during			Depressive symptoms	HADS Depression
			the non-epidemic period (NEH): 50; 26 (52% ^d); 37.22 (5.40)			Other Outcomes	SF-36
Zhang SX et al. (2020a)	CS, OBS	China	369; 165 (44.7%); 36.6 (10.5)	NA	February 20–21, 2020	Psychological Distress	K6
[90] ¹ ; Zhang SX et al. (2020b) [91] ¹						Other outcomes	SF12, SWLS
Zhang Y et al. (2020) [92] ^{1,2}	CS, OBS	China	263; 157 (60%); 37.7 (14.0)	NA	January 28– February 05, 2020	PTSS	IES
Zhou SJ et al.	CS, OBS	China	8079; 4326 (53.5%); NA	100% senior high	March 8–15,	Anxiety	GAD-7
(2020) [93] ¹			(median: 16, minimum 12, maximum 18 years)	school students ^b	2020	Depressive symptoms	PHQ-9
Healthcare wor	kers						
Abdessater et al. (2020) [94] ¹	CS, OBS	France	275; 91 ^d (33%) or 83 ^d (30%), ambiguous data; 29.5 (0.47)	100% urologists	March 27– 30, 2020	Stress	a
Ahmed et al. (2020) [95] ¹	CS, OBS	multinational (Pakistan > Saudi Arabia > others)	650; 490 (75%); NA (mode: 20–30 years [54%])	100% dentists	March 10– 17, 2020	Anxiety	8 binary items ^a
Alhaj et al. (2020) [96] ¹	CS, OBS	multinational (Canada, USA, others)	52; 14 (27%); NA (mode: < 30 years [69%])	100% surgeons	April 14–28, 2020	Psychological distress	Affection of mental health (binary single item) ^a
Amerio et al.	CS, OBS	Italy	131; 63 (48.1%); 52.3 (12.2)	100% physicians	March 15-	Anxiety	GAD-7
(2020) [97] ¹				(general practitioners)	April 15, 2020	Depressive symptoms	PHQ-9
						Sleep-related symptoms	ISI
						Other outcomes	SF-12
Badahdah	CS, OBS	Oman	194; 116 ^d (60%); 40.72 (8.53)	100% physicians	early April	Anxiety	GAD-7
et al. (2020) [98] ¹					2020	Stress	PSS-10
						Other	WHO-5 ^c

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Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales	
						outcomes		
Bohlken et al. (2020) [99] ¹	CS, OBS	Germany	396; NA; 165 (42%); 56.9 (7.6)	100% physicians	April 1–6, 2020	Anxiety and fear	Single items 5- P LS ^a	
						Sleep disorders	Single item 5-P LS ^a	
Cai H et al. (2020) [100] ^{1,2}	CS, OBS	China	534; 367 (69%); 36.4 (16.18)	physicians ($n = 233$), nurses ($n = 248$)	January– March, 2020	Anxiety and fear	Single items 4- P LS ^a	
Cai W et al. (2020) [101] ¹	CS, OBS	China	whole sample: 1521; 1149 (75.5% ^d); NA	physicians ($n = 511$), nurses ($n = 546$)	NA	Anxiety and fear	SCL-90 anxiety	
			(mode: 18–30 years, [43.5%])			Depressive symptoms	SCL-90 depression	
						Psychological distress	SCL-90 positive items	
						Other outcomes	SCL-90 subscales, CD- RISC, SSRS	
Chew et al. (2020) [102] ¹	CS, OBS	multinational (Singapore,	906; 583 (64.3%); NA (median [IQR]: 29 [25–35] years)		February 19–April 17,	Anxiety and fear	DASS-21 anxiety	
		India)	professionals ($n = 96$),	professionals ($n = 96$),			Depressive symptoms	DASS-21 depression
				11011 11CVV (17 = 107)		Stress	DASS-21 stress	
						Sleep-related symptoms	Single item 4-P LS ^a	
						PTSS	IES-R	
Consolo et al. (2020) [103] ¹	CS, OBS	Italy	356; 141 (39.6%); NA (mode: 35–55 years [48.6%])	100% dentists	April 2–21, 2020	Anxiety and fear	GAD-7	
Gan et al. (2020) [104] ¹	CS, OBS	China	11,183; 10,811 (96.7%); NA (mode: 20–29 years)	100% nurses	February 4– 10, 2020	Anxiety and fear	VAS on anxiety	
						Stress	VAS on stress	
Huang JZ et al. (2020)	CS, OBS	China	230; 187 (81.3%); NA (mode: 30–39 years [53%])	physicians (n = 70), nurses (n = 160)	February 7– 14, 2020	Anxiety and fear	SAS	
[105] ^{1,2}						PTSS	PTSD-SS	
Kang et al. (2020) [106] ^{1,2}	CS, OBS	China	994; 850 (85.5%); NA (mode: 30–40 years [63.4%])	physicians ($n = 183$), nurses ($n = 811$)	January 29– February 4,	Anxiety and fear	GAD-7 ^e	
					2020	Depressive symptoms	PHQ-9 ^e	
						Sleep-related symptoms	ISI ^e	
						PTSS	IES-R ^e	
Khusid et al. (2020) [107] ¹	CS, OBS	USA	332; 117 (35%); 30.5 (2.6)	100% urologists	April 7-11, 2020	Anxiety and fear	2 items 5-P LS ^a	
						Depressive symptoms	2 items 5-P LS ^a	
Lai et al. (2020) [18] ^{1,2}	CS, OBS	5, OBS China 1257; 964 (76.7%); NA (mode: physicians (n = 493), 26–40 years [64.7%]) nurses (n = 764)		January 29– February 3,	Anxiety and fear	GAD-7		
					2020	Depressive symptoms	PHQ-9	
						Sleep-related symptoms	ISI	
						PTSS	IES	

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Table 1 Study characteristics of included main studies (*Continued*)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales
Mo et al. (2020) [108] ^{1,2}	CS, OBS	China	180; 162 (90%); 32.71 (6.52)	NA	end of February	Anxiety and fear	SAS
					2020	Stress	SOS
Pu et al. (2020) [109] ¹	CS, OBS	China	867: 829 (95.6% ^d); 30.8 (7.1)	100% nurses	NA	Anxiety and fear	SAS
						Other outcomes	TAF
Rossi et al. (2020) [110] ¹	CS, OBS	Italy	1379; 1064 (77.2%); 39.0 (6.0)	physicians ($n = 433$), general practitioners	March 27– 31, 2020	Anxiety and fear	GAD-7
				(n = 86), nurses (n = 472)		Depressive symptoms	PHQ-9
						Stress	PSS
						Sleep-related symptoms	ISI
						PTSS	GPS-PTSD
Sahu et al. (2020) [111] ¹	CS, OBS	India	611; NA; NA (mode: 30–40 years, <i>n</i> = 192 [31-4%])	100% orthopedic surgeons	March 31– April 4, 2020	Stress	Single-item ^a
Shacham et al. (2020) [112] ¹	CS, OBS	Israel	338; 198 (586%); 46.39 (11.2)	dentists ($n = 198$), dental hygienists ($n = 140^{d}$)	March 30– April 10, 2020	Psychological distress	K-6
Suleiman et al. (2020) [113] ¹	CS, OBS	Jordan	308; 113 (36.7%); 30.3 (5.8)	100% physicians	March 23– 27, 2020	Anxiety and fear	Binary single items ^a
Tan B et al. (2020) [114] ¹	CS, OBS	Singapore			Anxiety and fear	DASS-21 anxiety	
				hospital personnel (n = 174)	13, 2020	Depressive symptoms	DASS-21 depression
						Stress	DASS-21 stres
						PTSS	IES-R
Wang S et al. (2020) [115] ¹	CS, OBS	China	123; 111 (90%); 33.75 (8.41)	100% pediatricians; physicians ($n = 48$),	January 30– February 07,	Anxiety and fear	SAS
				nurses (n = 75)	2020	Depressive symptoms	SDS
						Sleep-related symptoms	PSQI
Wu K et al. (2020) [116] ¹	CS, OBS, controlled	China	experimental group: 60; 44 (73%); 33.5 (12.4) comparison group: 60; 45	COVID-19 hospital (n = 60), non-designated hospital =	NA	Anxiety and fear	SAS, SCL-90 anxiety
			(75%) 33.8 (11.9)	comparison group (n = 60)		Depressive symptoms	SCL-90 depression, SDS
						Sleep-related symptoms	PSQI
						PTSS	PCL-C
					Psychological distress	SCL-90 total score	
						Other outcomes	SCL-90 subscales
Xiao et al. (2020a) [117] ^{1,2}	CS, OBS	China	180; 129 (71.7%); 32.31 (4.88)	physicians ($n = 82$), nurses ($n = 98$)	January– February,	Anxiety and fear	SAS
[117] *					2020	Sleep-related	PSQI

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Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales
						symptoms	
						Other outcomes	GSES, SASR, SSRS
Xu J et al. (2020) [118] ¹	CS, OBS, controlled	China	outbreak period: 60; 38 (63.3%); 36.68 (9.67) 'post-epidemic': 60; 32	100% surgeons	January 28– February 29, 2020 and	Anxiety and fear	'Anxiety scale', dream anxiety score
			(53.3%); 35.77 (7.06)		March 2–21, 2020	Depressive symptoms	'Depression score'
						Other outcomes	SF-36
Yin et al. (2020) [119] ¹	CS, OBS	China	371; 228 (61.5%); 35.3 (9.5) physicians: NA	physicians ($n = 67$), nurses ($n = 264$)	February 01–05, 2020	Sleep-related symptoms	PSQI
			nurses: NA			PTSS	PCL-5
Zhang C et al. (2020)	CS, OBS	China	1563; 1293 (83% ^d); NA (mode: 26–40 years, <i>n</i> = 495 [31.7% ^d])	physicians ($n = 454$), nurses ($n = 984$),	January 29– February 03,	Anxiety and fear	GAD-7
[120] ¹			physicians: NA nurses: NA	administrative staff $(n = 30)$, other medical staff $(n = 95)$	2020	Depressive symptoms	PHQ-9
				, ,		Sleep-related symptoms	ISI
						PTSS	IES-R
Zhang SX et al. (2020c)	CS, OBS	Iran		April 5-20, 2020	Anxiety and fear	GAD-2 ^d	
[121] ¹						Depressive symptoms	PHQ-2 ^d
						Psychological distress	K6
						Other outcomes	SF-12
Zhu J et al. (2020) [122] ¹	CS, OBS	China	156; 137 (83%); 34.16 (8.06) physicians: 79; 51 ^d (65% ^d)	physicians ($n = 79$), nurses ($n = 86$)	February 1– 29, 2020	Anxiety and fear	SAS
						Depressive symptoms	SDS
Patients							
Cai X et al. (2020) [123] ¹ ,	CS, OBS	China	126; 66 (52.4%); 45.7 (14.0)	100% cured COVID-19 patients	March 2–12, 2020	,	SAS
Yuan B et al. (2020) [124] ¹						Depressive symptoms	SDS
						PTSS	PTSD-SS
Durankus et al. (2020)	CS, OBS	Turkey	260; 260 (100%); 29.6 (3.8)	100% pregnant women	NA	Anxiety and fear	BAI
[125] ¹						Depressive symptoms	EPDS, BDI
						Psychological distress	Single item 11- P LS ^a
Li X et al. (2020) [126] ¹	CS, OBS	China	76; 35 (46%); 36 (15)	suspected COVID-19 patients	January 31– February 22,	Anxiety and fear	HAMA
					2020	Depressive symptoms	HAMD
Liu X et al. (2020a) [42] ¹	CS, OBS	China	COVID-19 suspected patients: 21; 12 (57.1%); 43.1 (2.6): not COVID-19 suspected	100% schizophrenia patients; COVID-19 suspected patients	January 30– February 21, 2020	Anxiety and fear	HAMA

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 Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales		
			patients: 30; 15 (50%); 45.0 (9.2)	(n = 21), not COVID- 19 suspected patients		Depressive symptoms	HAMD		
				(n = 30)		Stress	PSS		
						Sleep-related symptoms	PSQI		
						Other outcomes	PANSS		
Wu Y et al. (2020) [127] ^{1,3}	CS, OBS, controlled	China	4124; 4124 (100% ^d), NA (median: 30, range = 17–32	100% pregnant women;	January 1– February 9,	Anxiety and fear	EPDS-3A		
			years)	before (group 1: <i>n</i> = 2839)/after (group 2: <i>n</i> = 1284) January 20, 2020	2020	Depressive symptoms	EPDS		
Xu H et al. (2020) [128] ¹	CS, OBS	China	350; 199 (54.1%); NA (mode: 40–60 years [51%])	100% lung cancer patients	March 4–6, 2020	Depressive symptoms	Single item ^a		
						Sleep-related symptoms	Single item ^a		
Yassa et al. (2020) [129] ¹	CS, OBS	Turkey	172; 172 (100%); 27.5 (5.3)	100% pregnant women	ten days after first confirmed COVID-19 death in Turkey	Anxiety and fear	Single ternary item ^a		
Mixed groups									
Büntzel et al. (2020) [130] ¹	CS, OBS	Germany	193; NA; NA (mode: > 60 years)	physicians (n = 47), cancer patients ($n = 146$)	April 16–19, 2020	Anxiety and fear	Single item ^a		
				146)		Stress	Single item ^a		
Guo et al. (2020) [131] ¹	CS, OBS, controlled	China	P:103; 44 (42.7%); 42.5 (12.5); control (GP): 103; 49 (47.6%); 41.5 (13.1)	COVID-19 patients (n = 103), not infected control group (n =	February 10–28, 2020	Anxiety and fear	GAD-7		
			41.5 (15.1)	103)		Depressive symptoms	PHQ-9		
						Stress	PSS-10		
						PTSS	PCL-5		
Hao F et al. (2020) [132] ¹	CS, OBS, controlled	China	P: 76; 51 (37.1%); 32.8 (11.8); control (GP): 109; 68 (62.4%); 33.1 (11.2)	psychiatric patients (n = 76), control group (n = 109)	February 19–22, 2020	Anxiety and fear	DASS-21 anxiety		
			55.1 (11.2)	group (17—109)		Depressive symptoms	DASS-21 depression		
						Stress	DASS-21 stress		
						Sleep-related symptoms	ISI		
						PTSS	IES-R		
Hao X et al. (2020) [133] ¹	CS, OBS, controlled	China	P: 252; 132 ^d (52.4% ^d); 29.3 (11.6); control (GP): 252; 132 ^d (52.4% ^d); 29.4 (11.5)	epilepsy patients (n = 252), control group (n = 252)	February 1– 29, 2020	Psychological distress	K-6		
Huang Y et al. (2020)	CS, OBS	China	7236; 3952 (54.6%); 35.3 (5.6)	GP (n = 4986), HCW (n = 2250)	February 3– 17, 2020	Anxiety and fear	GAD-7		
[134] ^{1,2}								Depressive symptoms	CES-D
						Sleep-related symptoms	PSQI		
lasevoli et al.	CS, OBS,	Italy	461; NA; NA	psychiatric patients	April 13-17,	Anxiety and	GAD-7		

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Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales	
(2020) [135] ¹	controlled		P: 205; NA; NA	(n = 205), caregivers	2020	fear		
			caregivers: 51; NA; NA control (GP): 205; NA; NA	(n = 51), non- psychiatric persons (n = 205)		Depressive symptoms	PHQ-9	
				(===,		Stress	PSS	
						Other outcomes	SPEQ	
Jin YH et al. (2020) [136] ¹	CS, OBS	China	103; 64 (62.1%); NA (median [IQR]: 35 [14.0])	100% infected with SARS-CoV-2; physi- cians, nurses	February 15–29, 2020	Anxiety and fear	Single item multiple choice ^a	
Ko et al. (2020) [137] ¹	CS, OBS	Taiwan	1904; 1282 (67.3%); 38.0 (10.8)	GP (n = NA), HCW (n = NA)	April 10–20, 2020	Other outcomes	Psychological wellbeing (single item 5- P LS) ^a	
Li Z et al. (2020) [138] ^{1,2}	CS, OBS	China	740; 128 (59.8%); 25 (IQR: 22–38.3 years]	GP (n = 214), HCW (n = 526)	February 17–21, 2020	PTSS	Vicarious Traumatization Questionnaire	
Lu W et al. (2020) [139] ^{1,2}	CS, OBS	China	2299; 1785 (77.6%); NA (78% < 40 years)	HCW (n = 2042), GP (n = 257)	February 25.26, 2020	Anxiety and fear	HAMA, NRS on fear	
						Depressive symptoms	HAMD	
Ni et al. (2020) [140] ¹	CS, OBS	GP: 1577; 1218 ^d (60.8%); NA (n = 214)	GP: 1577; 1218 ^d (60.8%); NA (n = 214) 18.24, 202	, ,,,	February 18.24, 2020	Anxiety and fear	GAD-2	
			(mode: 18–34 years [38.6%]) HCW: 214; 147 ^d (68.8%); NA (mode: 18–34 years [58.9%])			Depressive symptoms	PHQ-2	
Sanchez et al. (2020) [67] ¹	CS, OBS	USA	1051; 0 (0%); 35 (15.83)	100% men who have sex with men; HIV-	April 2–13, 2020	Anxiety and fear	Single item ^a	
				patients (<i>n</i> = 122)		Other outcomes	Quality of life (single item) ^a	
Wu W et al. (2020) [141] ¹	CS, OBS	China	4268; 2930 ^d (68.7% ^d); NA HCW: 2110; 1598 ^d (76% ^d); NA	students (n = 2158), HCW (n = 2110)	February 10–21, 2020	Anxiety and fear	Single item ^a	
			Students: 2158; 1332 (62%); NA			Sleep-related symptoms	Single item ^a	
Yuan S et al. (2020) [142] ^{1,2}	L, OBS	China	939; 582 (61.98%); NA (mode: 18–39 years [71.5%])	HCW ($n = 249$), students ($n = 312$)	2 survey periods in	Sleep-related symptoms	PSQI	
					February, 2020	Other outcomes	SRQ	
Zhang J et al. (2020) [143] ¹	CS, OBS	China	205; 115 (56.1% ^d); NA (for infected: 46.9 [15.4]; for guarantined: 36.2 [10.9]; for	P, infected $(n = 57)$, GP, quarantined $(n = 50)$,	February 15–29, 2020	Anxiety and fear	GAD-7	
			general public: 29.6 [12.7])	GP, general public (n = 98)		Depressive symptoms	PHQ-9	
Zhang WR et al. (2020)	CS, OBS	China		February 19–March 6,	Anxiety and fear	GAD-2		
[144] ^{1,2}				,,	202	2020	Depressive symptoms	PHQ-2
							Sleep-related symptoms	ISI
						Other outcomes	SCL-90-R subscales	
Zhu S et al. (2020) [145] ¹	CS, OBS	China	2279 ^d ; 1361 ^d ; NA	HCW (n = 858), GP (n = 1421)	Feb 12–Mar 17, 2020	Anxiety and fear	GAD-7	
						Depressive	PHQ-9	

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Table 1 Study characteristics of included main studies (Continued)

Study	Study design	Country	Sample size; female: No. (%); age: mean (SD) or alternative information on age (eg, mode)	Subgroups	Survey period	Assessed Outcomes	Instruments or scales
						symptoms	
						Psychological distress	SRQ-20

Abbreviations: AIS Athens Insomnia Scale, BAI Beck Anxiety Inventory, BDI Beck Depression Inventory, BDI Call Beck Depression Inventory, BDI Beck Depressio Perception Questionnaire 5. BRCS Brief Resilience Coping Scale, CD-R/SC Connor-Davidson Resilience Scale, CES-D Center for Epidemiologic Studies Depression Scale, CoVGAD-7 Generalized Anxiety Disorder Scale-7 for COVID-19 Anxiety, CPDI CoViD-19 Peritraumatic Distress Index, CS cross-sectional, DASS-21 Depression Anxiety Stress Scale-21, DT Distress Thermometer, EPDS Edinburgh Postnatal Depression Scale, EPDS-3A Edinburgh Postnatal Depression Scale-Anxiety subscale, FACIT-Sp12 Functional Assessment of Chronic Illness Therapy-Spiritual Well-Being Scale, FCV-19S Fear of COVID-19 scale, GAD-2(-7) Generalized Anxiety Disorder Scale-2(/-7), GP general population, GPS-PTSD Global Psychotrauma Scale-posttraumatic stress disorder subscale, GSES General Self-Efficacy Scale, GSI Global Severity Index, HADS Hospital Anxiety and Depression Scale, HAI Health Anxiety Inventory, HAMA Hamilton Anxiety Rating Scale, HAMD Hamilton Depression Rating Scale, HCW healthcare workers, IES Impact of Event Scale, IES-R Impact of Event Scale-Revised, IQR interquartile range, ISI Insomnia Severity Index, IUS-12 Intolerance of Uncertainty Scale-Short Form, JGLS De Jong Gierveld Loneliness Scale, K-6(/- 10) Kessler Psychological Distress Scale-6(/- 10), L longitudinal, MINI Mini International Neuropsychiatric Interview, MSPSS Multidimensional Scale of Perceived Social Support, NA not available, NRS Numeric Rating Scale, OBS observational, P patients, PANSS Positive and Negative Syndrome Scale, PCL-5(-C) Post-traumatic Stress Disorder Checklist-5(/-Civilian Version), PHO-2(/-4/-9/- 15) Patient Health Questionnaire-2(/-4/-9/- 15), PROMIS-SFs Patient Reported Outcomes Measurement Information System short forms, PSQI Pittsburgh Sleep Quality Index, PSS(-10) Perceived Stress Scale(-10), PTSD-SS Post-traumatic Stress Disorder Self-rating Scale, PTSS post-traumatic stress symptoms, Ryff's PWB Ryff's Psychological Wellbeing Scales, SAS Self-Rating Anxiety Scale, SASR Stanford Acute Stress Reaction, SCL-90 Symptom Checklist-90, SCS Self-Compassion Scale, SD standard deviation, SDO Strengths and Difficulties Questionnaire, SDS Self-Rating Depression Scale, SF-12(/-36) Short Form 12 Health Survey, SHAI Short Health Anxiety Inventory, SOS Stress Overload Scale, SPEQ Specific Psychotic Experience Questionnaire, SRQ Stress Response Questionnaire, SRQ-20 20-item Self-Report Questionnaire, SSRS Social Support Rating Scale, SSS Somatic Symptom Scale, STAI-Y State Trait Anxiety Inventory-Y, SWLS Satisfaction With Life Scale, TAF Triage Assessment Form, TEMPS-A Temperament Evaluation of Memphis, Pisa, Paris and San Diego-Anxious, VAS Visual Analogue Scale, VDAS Van Dream Anxiety Scale, WHO-5 World Health Organization- Five Well-Being Index, WHOQOL-BREF abbreviated World Health Organization Quality of Life, 4-/5-/7-/11-P LS

- 4-/5-/6-/11-point Likert-scale a developed by study authors
- ^b included in main analyses for general population but considered separately in subgroup-analyses
- ^c in Gao J et al. WHO-5 is used to assess depressive symptoms, in Badahdah et al. it is used to assess psychological distress
- ^d not directly reported

The risk and protective factors narratively identified for each population are presented in Table 5 and eTables 14 and 15, with most of them being investigated in the general population, and few studies investigating protective factors at all. Most frequently named risk factors across the populations were pre-existing mental disorders, female sex, and concerns about COVID-19 infection, whereas most frequently reported protective factors were older age, good economic situation, and higher education.

Discussion

To our knowledge, this is the first systematic review and meta-analysis to assess the mental health impact of the SARS-CoV-2 pandemic in the general population, health-care workers, and patients, by contrasting data from the early phase of the current pandemic with prepandemic data. We identified 104 independent studies, mainly in the general population, that suggest an increased prevalence of mental burden due to the SARS-CoV-2 pandemic. This finding is in line with previous reviews and meta-analyses that merely pooled the prevalence of or calculated the risk for mental burden in either one or several of these groups [20, 24, 29, 32, 33, 36].

On the other hand, the pairwise meta-analyses for 43 studies across the four primary outcomes revealed different results. Compared with prepandemic data, we only found an elevated level of some mental symptoms

(anxiety, depression) due to the SARS-CoV-2 pandemic in the general population, but not of stress or sleeping problems.

Although healthcare workers were found to be a group at risk for mental health problems during the SARS-CoV-2 pandemic [18]^{1,2} [20, 29, 31, 32], we identified no evidence for an increased mental burden during the early phase when comparing them with healthcare staff prior to the pandemic. Because of a (chronic) work-related risk exposure in daily life [194], as a kind of 'stress inoculation', healthcare professionals might have learned effective strategies (eg, self-efficacy) helping them to cope more professionally with crises than other groups. In contrast to previous findings [20, 195], the level of COVID-19 patient contact did not affect the mental health impact.

Overall, the results of this review paint a more nuanced picture of the mental health consequences of the SARS-CoV-2 pandemic than previous reviews – an observation in line with stress resilience research that identified different trajectories of psychological adaptation after potentially traumatic events, ranging from no mental burden to severe mental illness [196, 197]. Indeed, a recent analysis of 523 healthy subjects from the German LORA study showed a decrease of perceived stress and stressor load while mental health improved during the eight-week measurement after lockdown, indicating that the pandemic and pandemic response may also have

e k-means-clustering method for the 4 tools summarized to 'mental health'

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Table 2 Narrative synthesis of prevalence based on scores above cut-off values for different mental health outcomes

	Number of studies ^a	Lowest reported prevalence (%)	Highest reported prevalence (%)
General population			
Anxiety, worries, fear	24 (18 GP, [45, 47, 49, 50, 52, 57, 63, 66, 68, 69, 71, 77, 93] ¹ [73, 84, 87] ^{1,2} 6 M [132, 140, 145] ¹ [134, 139, 144] ^{1,2})	0.67 (63)	64.0 (46)
Depressive symptoms	18 (13 GP [45, 49, 50, 52, 57, 63, 66, 68, 69, 71, 93] ¹ [84, 87] ^{1,2} , 5 M [132, 140, 145] ¹ [139, 144] ^{1,2})	0.9 (89)	48.3 (48)
PTSS	7 (6 GP [52, 66, 71] ¹ [59, 84, 92] ^{1,2} , 1 M [132] ¹)	7.0 (51)	53.8 (55)
Sleep-related symptoms	6 (3 GP [71, 83] ¹ [84] ^{1,2} , 3 M [132] ¹ [134, 144] ^{1,2})	0.9 (89)	37.6 (131)
Stress	5 (4 GP [66, 68, 71, 83] ¹ , 1 M [132] ¹)	0.9 (89)	67.9 (55)
Psychological distress	7 (5 GP [41, 45, 58, 65, 81] ¹ , 2 M [133, 136] ¹)	1.6 (90)	65.2 (112)
Healthcare workers			
Anxiety, worries, fear	22 (14 HCW [99, 95, 113, 102, 103, 110, 115, 120, 122, 121] ¹ [18, 100, 105, 106] ^{1,2} , 6 M [130, 140, 145] ¹ [134, 139, 144] ^{1,2})	7.0 (108)	92.0 (144)
Depressive symptoms	14 (9 HCW [97, 102, 110, 115, 120, 121, 122] ¹ [18, 106] ^{1,2} , 5 M [140, 145] ¹ [134, 139, 144] ^{1,2})	0.6 (110)	50.4 (18)
PTSS	7 (HCW) [102, 110, 119, 120] ¹ [18, 105, 106] ^{1,2}	3.8 (82)	73.0 (83)
Sleep-related symptoms	9 (7 HCW [99, 102, 110, 115, 120] ¹ [18, 106] ^{1,2} , 2 M [134] ¹ [144] ^{1,2})	8.27 (127)	38.0 (108)
Stress	6 (5 HCW [94, 102, 110, 111] ¹ [108] ^{1,2} , 1 M [130] ¹)	5.2 (102)	56.5 (114)
Psychological distress	5 (4 HCW [96, 101, 112, 121] ¹ , 1 M [145] ¹)	11.1 (101)	90.4 (145)
Patients			
Anxiety, worries, fear	6 (5P [123, 126, 129, 131, 143] ¹ , 1 M [132] ¹)	19.5 (99)	80.2 (143)
Depressive symptoms	8 (7 P [123, 125, 126, 128, 131, 143] ¹ [127] ^{1,3} , 1 M [132] ¹)	27.8 (99)	55.3 (88)
PTSS	2 (1 P [123] ¹ , 1 M [132] ¹)	31.0 (84)	43.4 (89)
Sleep-related symptoms	2 (1 P [128] ¹ , 1 M [132] ¹)	27.6 (89)	66.3 (97)
Stress	1 (M [132] ¹)	17.0 (89)	
Psychological distress	1 (M [133] ¹)	13.1 (90)	

Abbreviations: GP general population, HCW healthcare workers, M mixed samples, P patients, PTSS posttraumatic stress symptoms

positive effects [198]. The number of studies reporting on protective factors in this review was rather limited, especially in healthcare workers and patients. However, these factors might also partly explain the heterogeneity of findings regarding mental health consequences. This is in line with positive aspects (eg, improved social relationships with close social contacts such as families) that were likewise reported for previous infectious disease outbreaks. The importance of taking a 'resilience perspective' in SARS-CoV-2 mental health research and investigating resilience factors has been pointed out previously [19, 22, 197, 199].

Several aspects must be considered when interpreting the results. First, the absence of evidence of effects in healthcare workers and patients in this review does not necessarily mean that there is evidence for the absence of effects of the SARS-CoV-2 pandemic on mental health in these groups. Second, for healthcare workers, the mental burden on individuals probably depends on the location of survey (eg, country, region) and how heavily the respective healthcare systems were burdened in the pandemic timeline (eg, number of hospitalized COVID-19 patients). Among the 13 included studies in meta-analyses for healthcare staff, we could only include

^a reporting prevalence rates for the respective mental health outcome

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Outcome	Studies (samples)	N (pandemic)	N (comp.)	Standardized mean difference (95% CI)		95% prediction interval ^a	
Main analyses							
General population							
Anxiety	23 (26)	49,746	132,145	0.40 (0.15–0.65)	99%	- 0.87-1.67	
Depression	25 (28)	60,213	183,747	0.67 (0.07–1.27)	100%	-2.02-3.36	
Stress	11 (13)	11,600	67,386	0.10 (-0.30-0.50)	100%	-1.39-1.60	
Sleep-related symptoms	4 (4)	3332	7635	0.74 (-1.47-2.96)	100%	-3.68-5.17	
lealthcare workers							
Anxiety	13 (14)	5508	22,204	-0.08 (-0.66-0.49)	99%	-1.75-1.58	
Depression	7 (8)	2226	4605	-0.16 (- 0.59-0.26)	97%	-1.41-1.09	
Stress	3 (3)	1570	2454	0.49 (-0.60-1.57)	99%	/	
Sleep-related symptoms	4 (5)	554	20,024	0.83 (-0.14-1.81)	99%	-1.54-3.21	
atients							
Anxiety	6 (6)	1845	12,458	0.31 (-0.07, 0.69)	93%	-1.08-1.69	
Depression	7 (7)	2138	24,444	0.48 (-0.08-1.04)	98%	-1.58-2.53	
Stress	4 (4)	435	10,061	-0.10 (-0.81-0.61)	98%	-3.54-3.34	
Sleep-related symptoms	2 (2)	127	298	-0.61 (-1.75-0.54)	96%	/	
ensitivity analysis -	- Quality of include	d pandemic studi	es (ie, exclu	usion of poor-quality studies)			
Seneral population							
Anxiety	16 (17)	38,323	81,350	0.53 (0.19–0.86)	100%	-0.90-1.95	
Depression	18 (19)	48,790	136,884	0.83 (0.09–1.57)		-2.17-3.82	
Stress	7 (8)	9110	43,747	0.33 (-0.19-0.84)	100%	-1.20-1.85	
Sleep-related symptoms	3 (3)	2659	6622	0.80 (-1.34-2.94)	100%	/	
lealthcare workers							
Anxiety	4 (4)	1655	4124	-0.18 (-0.78-0.41)		-1.30-0.94	
Depression	4 (4)	1655	2356	0.03 (-0.42-0.47)		-0.73-0.79	
Stress	2 (2)	1376	1872	-0.05 (-0.37-0.26)	95%	/	
Sleep-related symptoms	1 (1)	123	4951	-0.03 (- 0.21-0.15)	/	/	
Patients							
Anxiety	3 (3)	1461	11,116	0.45 (-0.10-1.01)	92%	/	
Depression	3 (3)	1461	21,934	0.21 (-1.08-1.49)	99%	/	
Stress	1 (1)	51	51	0.18 (-0.21-0.57)	/	/	
Sleep-related symptoms	1 (1)	51	207	-0.03 (- 0.33-0.28)	/	/	
Sensitivity analysis - evel-4 studies)	- Level of comparal	oility between inc	luded pand	lemic studies and comparative studies	(ie, excl	usion of level-3 and	
General population							
Anxiety	12 (13)	38,461	32,698	0.40 (0.06–0.74)	99%	-0.77-1.57	
Depression	14 (15)	38,259	78,619	0.77 (-0.23-1.77) 100% -2.72-4		-2.72-4.25	
Stress	7 (8)	8624	12,739	-0.15 (- 0.76-0.46)	99%	-1.84-1.53	
Sleep-related symptoms	2 (2)	2550	5609	1.54 (-1.18-4.27)	100%	/	

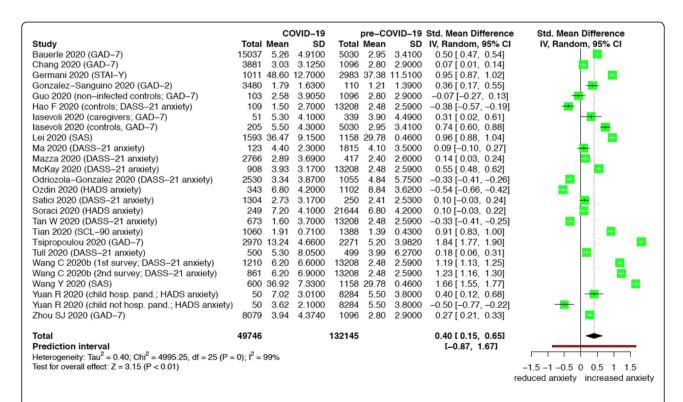
Table 3 Results of main and sensitivity analyses in three populations (Continued)

Outcome	Studies (samples)	N (pandemic)	N (comp.)	Standardized mean difference (95% CI)	l ²	95% prediction interval ^a
Healthcare workers						
Anxiety	7 (8)	3147	9511	-0.54 (-1.23-0.15)	99%	-2.11-1.03
Depression	4 (5)	546	2576	-0.38 (-1.56-0.79)	98%	-2.60-1.84
Stress	/	/	/	/	/	/
Sleep-related symptoms	3 (4)	423	19,804	1.01 (-0.17-2.18)	99%	-1.61-3.63
Patients						
Anxiety	4 (4)	1616	3184	0.23 (-0.33-0.79)	92%	-2.47-2.93
Depression	4 (4)	1704	3205	0 (-0.56-0.56)	93%	-2.69-2.70
Stress	2 (2)	127	217	0.15 (-0.08-0.37)	0%	/
Sleep-related symptoms	2 (2)	127	298	-0.61 (-1.75-0.54)	96%	/

Abbreviations: CI confidence interval, comp. comparative studies, l^2 heterogeneity, N sample size, pandemic included pandemic studies

a few studies from heavily burdened countries (eg, Italy: k = 2; Spain: k = 0; USA: k = 0). However, nine studies in these meta-analyses had been conducted in China, which, compared internationally, was less affected by the

SARS-CoV-2 pandemic [4]. In the subgroup analysis regarding the level of COVID-19 patient contact, we assigned studies to the subgroup 'high level of contact' if at least 50% of the sample had close contact to COVID-



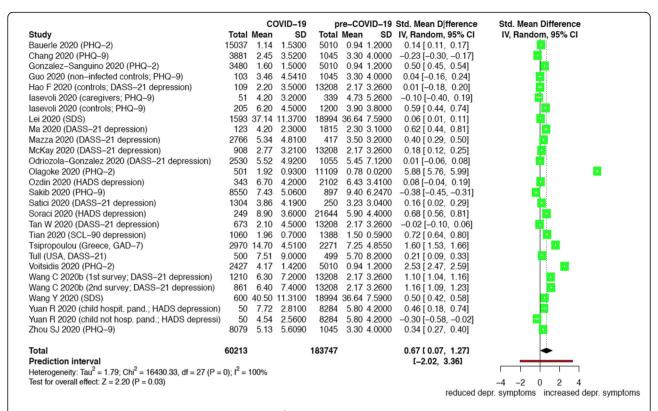
Abbreviations: CI, confidence interval; df, degrees of freedom; I², indicator of statistical heterogeneity; P, p value; SD, standard deviation; Std., standardized; Tau², indicator of statistical heterogeneity; Total, the number of participants; Z, z value; Chi², Chi² test for heterogeneity.

^a Horizontal lines indicate the 95% CI of each study; diamond, the pooled estimate with 95% CI; multilevel meta-analysis.

Fig. 2 Forest plot main analysis, general population, anxiety

^a 95% prediction interval only calculated for meta-analyses with at least k = 4 studies

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Abbreviations: CI, confidence interval; df, degrees of freedom; I², indicator of statistical heterogeneity; P, p value; SD, standard deviation; Std., standardized; Tau², indicator of statistical heterogeneity; Total, the number of participants; Z, z value; Chi², Chi² test for heterogeneity.

a Horizontal lines indicate the 95% CI of each study; diamond, the pooled estimate with 95% CI; multilevel meta-analysis.

Fig. 3 Forest plot main analysis, general population, depression

19 patients (ie, 'frontline healthcare workers'). However, the nature of contact was insufficiently described in the included studies.

Strengths of this review compared with previous publications include the systematic search for comparative prepandemic data for inclusion in pairwise meta-analyses, the stepwise selection of prepandemic studies to ensure best available comparability, and the population-specific analysis of risk and protective factors. One limitation refers to the search methods for pandemic studies (eg, no preprints; no reference lists of reviews) and comparative data (eg, subgroups in general population only partially searched). We had no restrictions regarding the publication format except for the exclusion of preprints which might be viewed as limitation. This restriction might have affected the evidence found in this review compared to others (eg, Cochrane reviews) where preprint articles are included.

The large between-study heterogeneity, a problem shared by previous meta-analyses [20, 24, 32, 33], could not be fully explained by subgroup analyses. This heterogeneity probably resulted from differences between the pandemic studies (eg, countries, sociocultural differences in the perception of mental burden, pandemic outbreak severity, subpopulations, outcome measures) and variability between the comparative studies (eg, study design, outcome measures), respectively. Among the pandemic studies, especially the specific outcome measures used were an important source of heterogeneity. Furthermore, the pandemic and comparative data were heterogeneous (eg, country, population), which could be partially captured by our selfdeveloped tool for the level of comparability and was controlled for by the corresponding sensitivity analysis. We cannot preclude that moderators of effects are present that we, though our best efforts, did not identify and therefore could not control for. Besides, comparative studies with larger sample sizes were preferred, leading to small 95% CIs and a lack of CI overlap with pandemic study findings. Despite the comprehensiveness of this review compared to previous publications, the small number of studies in certain subgroups potentially limited the statistical power (eg, survevs including students).

Apart from specific outcome measures, less recent comparative data, and homogenous sample sizes, the subgroup analyses indicated no consistent determinants of heterogeneity. An elevated level of depression based on the assessment with the PHQ and SDS might – at least for the PHQ-9 – be explained by the high

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Table 4 Results of subgroup analyses for those populations and outcomes with at least k=4 studies in main analysis

subgroup analysis subgroups)	Outcome	Test for subgroup differences ^a	Population	Subgroup difference: elevated effect ^b	Subgroup difference: reduced effect ^b
opulation characteristic	s (main stud	ies)			
age 20 years	Anxiety	$Chi^2 = 9.5$, $df = 5$ ($p = .09$)	GP	/	/
 30 years >30 ≤ 35 years >35 ≤ 40 years >40 ≤ 45 years multiple age groups age not specified 	Depression	Chi ² = 29.3, df = 5 (p < .001)	GP	≤30 years; > 40 ≤ 45 years	/
	Stress	$Chi^2 = 1043.3$, $df = 4$ ($p < .001$)	GP	/	> 40 ≤ 45 years
	Anxiety	$Chi^2 = 8.7$, $df = 4$ ($p = .07$)	HCW	/	/
	Depression	$Chi^2 = 2.2$, $df = 1$ ($p = .14$)	HCW	/	/
	Sleep	$Chi^2 = 0.3$, $df = 1$ ($p = .57$)	HCW	/	/
	Anxiety	Chi ² = 17.14, df = 4 $(p = .002)$	Р	> 40 ≤ 45 years	
	Depression	$Chi^2 = 3.74$, $df = 4$ ($p = .44$)	Р	/	/
tressor exposure	Anxiety	$Chi^2 = 2.8$, $df = 3$ ($p = .42$)	GP	/	/
 General population Students 	Depression	$Chi^2 = 1.9$, $df = 3$ ($p = .60$)	GP	/	/
Others Special exposure	Stress	Chi ² = 0.12, df = 3 (p = .99)	GP	/	/
ovid-19 patient contact	Anxiety	$Chi^2 = 0$, $df = 1$ ($p = .95$)	HCW	/	/
Low contact riskHigh contact risk	Depression	$Chi^2 = 1.0$, $df = 1$ ($p = .31$)	HCW	/	/
riigir comact risk	Sleep	$Chi^2 = 0.2$, $df = 1$ ($p = .69$)	HCW	/	/
ubgroup of patients	Anxiety	$Chi^2 = 0.3$, $df = 2$ ($p = .88$)	Р	/	/
· COVID-19 patients · Pregnant women · Psychiatric patients	Depression	Chi ² = 1.3, df = 2 (p = .51)	Р	/	/
Pandemic study characte	eristics				
urvey start ^c	Anxiety	$Chi^2 = 3.55$, $df = 4$ ($p = .47$)	GP	/	/
y ≤4 weeks y > 4 ≤ 6 weeks	· · · · · ·	Chi ² = 10.15, df = 4 (p = .04)	GP	> 8 weeks	/
> 6 ≤ 8 weeks> 8 weeks	Stress	Chi ² = 0.31, df = 4 (p = .99)	GP	/	/
not specified	Anxiety	$Chi^2 = 7.91$, $df = 4$ ($p = .10$)	HCW	/	/
	Depression	Chi ² = 0.95, df = 2 (p = .62)	HCW	/	/
	Sleep	Chi ² = 4.21, df = 2 (p = .12)	HCW	/	/
	Anxiety	Chi ² = 4.58, df = 2 (p = .10)		/	/
	Depression	Chi ² = 3.08, df = 3 (p = .38)		/	/
tudy conduction China	Anxiety	Chi ² = 0.10, df = 1 (p = .75)	GP	/	/
· China · Non-China	Depression	$Chi^2 = 0.60$, $df = 1$ ($p = .44$)	GP	/	/
NOH-CHINA	Stress	Chi ² = 0.10, df = 1 (p = .76)	GP	/	/
	Anxiety	Chi ² = 2.84, df = 1 (p = .09)	HCW	/	/
	•	$Chi^2 = 0.08$, $df = 1$ ($p = .78$)	HCW	/	/
	Sleep	Chi ² = 0.32, df = 1 (p = .57)	HCW	/	/
	Anxiety	$Chi^2 = 3.35$, $df = 1$ ($p = .07$)	Р	/	/
	Depression	$Chi^2 = 0.62$, $df = 1$ ($p = .43$)	Р	/	/
Outcome measure	Anxiety	Chi ² = 10.7, df = 6 (p = .10)	GP	/	/
AIS BDI DASS-21	Depression	Chi ² = 11.46, df = 5 $(p = .04)$	GP	PHQ-2	/
• EDPS	Stress	Chi ² = 0.16, df = 1 (p = .69)	GP	/	/
• EPDS-3A • GAD-2; GAD-7	Anxiety	Chi ² = 2.80, df = 4 (p = .59)	HCW	/	/
• GAD-2; GAD-7 • HADS	•				

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Table 4 Results of subgroup analyses for those populations and outcomes with at least k = 4 studies in main analysis (Continued)

Subgroup analysis (subgroups)	Outcome	Test for subgroup differences ^a	Population	Subgroup difference: elevated effect ^b	Subgroup difference: reduced effect ^b
• HAMA	Sleep	$Chi^2 = 0.32$, $df = 1$ ($p = .57$)	HCW	/	/
• HAMD • ISI	Anxiety	$Chi^2 = 1.18$, $df = 4$ ($p = .88$)	Р	/	/
PHQ-2; PHQ-9 PSQI PSS SAS SDS SCL-90 STAI-Y	Depression	$Chi^2 = 16.95$, $df = 5$ ($p = .005$)	Р	SDS; PHQ-9	/
Sample size	Anxiety	$Chi^2 = 1.86$, $df = 1$ ($p = .17$)	GP	/	/
• < 1000 • ≥1000	Depression	$Chi^2 = 0.03$, $df = 1$ ($p = .86$)	GP	/	/
_1000	Stress	$Chi^2 = 2.31$, $df = 1$ ($p = .13$)	GP	/	/
	Anxiety	$Chi^2 = 2.83$, $df = 1$ ($p = .09$)	HCW	/	/
	Depression	$Chi^2 = 0$, $df = 1$ ($p = .96$)	HCW	/	/
	Sleep	not possible	HCW	/	/
	Anxiety	$Chi^2 = 3.60$, $df = 1$ ($p = .06$)	Р	/	/
	Depression	$Chi^2 = 0.09$, $df = 1$ ($p = .77$)	Р	/	/
Comparative study cha	aracteristics				
Sample size	Anxiety	$Chi^2 = 0.9$, $df = 3$ ($p = .83$)	GP	/	/
≤500> 1000 ≤ 5000	Depression	$Chi^2 = 3.5$, $df = 4$ ($p = .48$)	GP	/	/
• > 5000 ≤ 10,000	Stress	$Chi^2 = 8.6$, $df = 3$ ($p = .03$)	GP	/	> 5000 ≤ 10,000 participant
• > 10,000	Anxiety	$Chi^2 = 9.93$, $df = 3$ ($p = .02$)	HCW	> 5000 ≤ 10,000 participants	
	Depression	$Chi^2 = 4.3$, $df = 2$ ($p = .12$)	HCW	/	/
	Sleep	$Chi^2 = 0.3$, $df = 1$ ($p = .57$)	HCW	/	/
	Anxiety	$Chi^2 = 0.1$, $df = 2$ ($p = .97$)	Р	/	/
	Depression	$Chi^2 = 3.9$, $df = 2$ ($p = .14$)	Р	/	/
Publication year • ≤1 year ago • ≤2 years ago	Anxiety	$Chi^2 = 8.0$, $df = 5$ ($p = .16$)	GP	/	/
	Depression	$Chi^2 = 12.4$, $df = 5$ ($p = .03$)	GP	> 10 years ago	/
 > 2 ≤ 5 years ago > 5 ≤ 10 years ago 	Stress	$Chi^2 = 11.6$, $df = 4$ ($p = .02$)	GP	/	≤1 year ago
• > 10 years ago	Anxiety	Chi ² = 14.5, df = 3 $(p = .002)$	HCW	> 10 years ago	≤2 years ago
		$Chi^2 = 4.6$, $df = 1$ ($p = .03$)	HCW	/	≤2 years ago
	Sleep	not possible	HCW	/	/
	Anxiety	$Chi^2 = 0.1$, $df = 2$ ($p = .94$)	Р	/	/
	Depression	Chi ² = 17.0, df = 5 (p = .005)	Р	≤1 year ago; > 5 ≤ 10 years ago	/
Pandemic and compar	ative study cha				
Relationship samples	Anxiety	$Chi^2 = 10.0$, $df = 3$ ($p = .02$)	GP	Ratio ≥ 0.5 < 2	/
sizes ^d • Ratio ≥ 2	Depression	$Chi^2 = 4.8$, $df = 3$ ($p = .19$)	GP	/	/
 Ratio ≥ 0.5 < 2 Ratio ≥ 0.1 < 0.5 	Stress	Chi ² = 0.4, df = 2 (p = .84)	GP	/	/
• Ratio < 0.1	Anxiety	$Chi^2 = 4.2$, $df = 2$ ($p = .12$)	HCW	/	/
	Depression	$Chi^2 = 3.8$, $df = 2$ ($p = .15$)	HCW	/	/

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Table 4 Results of subgroup analyses for those populations and outcomes with at least k = 4 studies in main analysis (Continued)

Subgroup analysis (subgroups)	Outcome	Test for subgroup differences ^a	Population	Subgroup difference: elevated effect ^b	Subgroup difference: reduced effect ^b
	Sleep	$Chi^2 = 0.32$, $df = 1$ ($p = .57$)	HCW	/	/
	Anxiety	Chi ² = 17.7, df = 3 $(p < .001)$	Р	Ratio ≥ 0.5 < 2; Ratio < 0.1	/
	Depression	$Chi^2 = 3.0$, $df = 3$ ($p = .39$)	Р	/	/

Abbreviations: AIS Athens Insomnia Scale, BDI Beck Depression Inventory, DASS-21 Depression Anxiety Stress Scale-21, df degrees of freedom, EPDS Edinburgh Postnatal Depression Scale, EPDS-3A Edinburgh Postnatal Depression Scale-Anxiety subscale, GAD Generalized Anxiety Disorder Scale, GP general population, HADS Hospital Anxiety and Depression Scale, HAMA Hamilton Anxiety Rating Scale, HAMD Hamilton Depression Rating Scale, HCW healthcare workers, p. p. value, P. patients, PHQ Patient Health Questionnaire, PSS Perceived Stress Scale, SAS Self-Rating Anxiety Scale, SCL-90 Symptom Checklist-90, SDS Zung Self-Rating Depression Scale, STAI-Y, State Trait Anxiety Inventory-Y

sensitivity to change of this instrument and its usefulness to monitor treatment outcomes [200, 201]. Given the increased mental burden if pandemic studies were compared to older prepandemic data, cohort effects cannot be excluded.

Discrepancies between subgroup analyses and the narrative synthesis of risk and protective factors (eg, COVID-19 patient contact) might be due to methodological differences. Because of the primary use of screening but not diagnostic tools to determine mental burden in the included pandemic studies, this review does not allow any conclusions concerning a putative increase of diagnoses of mental disorders during the early phase of the SARS-CoV-2 pandemic. Consistent with the synthesis of risk factors, the meta-analyses partly showed an increased level of mental symptoms in young and middleaged groups, in line with previous studies [12]. However, more studies including elderly would be needed to clearly investigate age differences, and whether the pandemic works as a 'burning lens' for the already increased mental burden in young people [202]. Finally, given the pandemic timeline, the evidence is substantially based

	Risk factors ^a	Protective factors ^b
General population	 - Mental disorder/or symptoms [44, 49, 51, 52, 58, 64, 69, 74, 78, 82, 83, 116, 132, 135]¹) - Worries about relatives or oneself [51, 57, 64, 66, 74, 75, 80, 89, 82, 83]¹ [48]¹²² - Being female [49, 52, 63, 66, 69, 72, 74, 79, 82, 83, 93]¹ [vs 1x being male] - Previous (chronic) medical disease [52, 55, 63, 64, 69, 85, 135]¹ - Being a student [52, 57, 60, 72, 146]¹ - Personal/social worries about COVID-19 [51, 85, 86, 145]¹ [48]¹² - Physical symptoms [52, 66, 80, 85, 132]¹ - Reduced perceived health [50, 57, 80, 85, 132]¹ - No current relationship [57, 80, 81, 146]¹ - Current local outbreak severity [57, 88, 93, 141]¹ - History of stressful situations [52, 58, 63, 147]¹ - Vulnerability to COVID-19 [53, 85, 146]¹ - Health profession [66, 81, 141]¹ - Own or close person's quarantine [57, 62, 85]¹ 	- Older age [49, 52, 63, 65, 66, 79, 91, 140, 147] ¹ - Good economic situation [52, 79, 88, 140, 146] ¹ [48] ^{1,2} - Satisfaction with/level of information on COVID-19 [45, 49, 52, 85, 88, 93] ¹ - Not being single [66, 80, 88,86] ¹ - Higher education [50, 52, 66, 146] ¹ - Social support [52, 140] ¹ [48] ^{1,2} - Being male [54, 65, 85] ¹
Healthcare workers	 Mental disorder/or symptoms [97, 115, 116, 119, 122]¹ Being female [98, 110, 119, 121]¹ Concern about infection with COVID-19 [103, 109, 120, 121]¹ Exposure to COVID-19 patients [94, 110, 115, 119]¹ Current local COVID-19 severity [94, 118, 141, 107]¹ 	- Older age [98, 110] ¹
Patients	- (Suspected) COVID-19 [42, 131, 143] ¹ - Inflammatory markers in blood [42, 131] ¹ - Physical symptoms [132] ¹	 Higher education [127]^{1,3} Good economic situation [127]^{1,3} Higher lymphocyte ratio in blood [42]¹ Concomitant medical diseases [135]¹

a most frequently reported risk factors: general population: factor was reported as statistically significant risk factor in at least k = 3 studies; healthcare workers: factor reported in at least k=4 studies; patients: factor reported in at least k=2 studies

Chi² = test for subgroup differences

^b ordered by size of effect estimate (SMD)

c since first COVID-19 cases in the respective country or, in case of China, since January 20, 2020

^d ratio of sample size in pandemic study vs comparative study

most frequently reported protective factors: general population: factor was reported as statistically significant protective factor in at least k = 3 studies; healthcare workers: factor reported in at least k = 2 studies (limited number of studies reporting protective factors in this group); patients: factor reported in k = 1study (limited number of studies reporting protective factors in this group)

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on Chinese studies thus potentially limiting the transferability of findings to other contexts.

Further research in other countries (eg, USA), that started later on during the pandemic, could change the findings. The latter is also supported by the wide prediction intervals identified in this review, which indicate uncertainty in our conclusions about whether the pandemic and related stressors do affect mental health [203].

The review has several implications for research and practice. There is an urgent need for representative surveys, in order to allow fair comparisons between the mental burden caused by SARS-CoV-2 in different countries and to examine other risk and protective factors (eg, cultural context). Representative surveys in the general population might also serve to identify specific subgroups at risk for which further studies would be needed. From a public mental health perspective, a stronger focus on (psychosocial) protective factors for mental health would be desirable to derive appropriate contents for preventive measures (eg, pandemic preparedness plans) or health-promoting interventions (eg, resilience training) prior to, during, and after a pandemic [199]. By further investigating the mental health impact of specific stressors - in line with Brooks and colleagues [13] - researchers and practitioners might gain further knowledge about when (eg, in pandemic timeline) and for whom (eg, after exposure to which stressors) interventions should be implemented to buffer negative mental health effects of SARS-CoV-2.

Conclusions

In conclusion, compared with prepandemic data, this review shows different adverse mental health consequences of the early phase of the SARS-CoV-2 pandemic in the examined population groups in contrast to previous research, with healthcare workers being more resilient than expected. The quality of studies varies. High-quality, representative surveys in the general population and specific sub-populations, longitudinal studies, and further research efforts on protective factors are needed to better understand the psychological impacts of the SARS-CoV-2 pandemic and to help design effective preventive measures and interventions that are tailored to the needs of specific population groups.

Supplementary Information

The online version contains supplementary material available at https://doi.org/10.1186/s12992-021-00670-y.

Additional file 1: Methods of the systematic review with meta-analyses. eTable 1. MOOSE Checklist. eTable 2. Differences between protocol

and review. **eMethods 1.** Search strategies for SARS-CoV-2 ('pandemic') studies. **eMethods 2.** Search strategy for prepandemic comparative studies. **eTable 3.** Eligibility criteria for SARS-CoV-2 pandemic studies. **eTable 4.** Eligibility criteria for prepandemic comparative studies. **eTable 5.** Eligibility criteria for pairwise meta-analyses. **eTable 6.** Customized data extraction sheet. **eTable 7.** Modified quality assessment tool. **eTable 8.** Rating of comparability between pandemic and prepandemic comparative studies. **eMethods 3.** Further methodological details of this systematic review and meta-analyses.

Additional file 2: Results of the systematic review with meta-analyses. eTable 9. Details on number of included (pandemic and comparative) studies. eTable 10. Study characteristics of the prepandemic comparative studies. eTable 11. Cut-off values reported in included pandemic studies. eResults 1. Forest plots of main analyses. eTable 12. Quality assessment of included pandemic studies. eTable 13. Assessment of level of comparability between pandemic and prepandemic comparative studies. eResults 2. Forest plots of sensitivity analyses. eResults 3. Detailed results of subgroup analyses. eTable 14. Risk factors in the general population, healthcare workers, and patients. eTable 15. Protective factors in the general population, healthcare workers, and patients.

Abbreviations

COVID-19: Coronavirus disease 2019; LORA: Longitudinal Resilience Assessment; MOOSE: Meta-analyses Of Observational Studies in Epidemiology; NIH: National Institutes of Health; PHO: Patient Health Questionnaire; PRISMA: Preferred Reporting Items for Systematic Reviews and Meta-Analysis; PROSPERO: International Prospective Register of Systematic Reviews; SARS-CoV(– 2): Severe Acute Respiratory Syndrome Coronavirus (– 2); SD: Standard deviation; SDS: Zung Self-Rating Depression Scale; SMD: Standardized mean difference

Acknowledgements

We appreciate the contributions of Prof Raffael Kalisch PhD to the discussion of the findings of our review.

Authors' contributions

AMK, NR, JSW, and KL designed the study. NR and LG, respectively, assessed study eligibility; KL was consulted in case of any disagreements. NR and LG extracted and analyzed data for Table 1, with KL being consulted in case of any disagreements. AMK, NR, and GS designed the statistical analyses. AMK and NR analyzed data for Tables 2, 3, 4 and 5 based on pairwise meta-analyses (including subgroup and sensitivity analyses) and the narrative synthesis of risk/protective factors. GS and HB reviewed the statistical analyses. KL monitored the review process. All authors contributed to the interpretation of the results, with special expertise provided in the field of resilience research (AMK, OT, KL), public health (MC, ER), and evidence-based medicine (CS, JJM). AMK wrote the first draft of the manuscript with input and subsequent edits by all authors. KL is the guarantor. All authors read and approved the final manuscript.

Funding

The CEOsys and the egePan project are funded under a scheme issued by the Network of University Medicine (Nationales Forschungsnetzwerk der Universitätsmedizin (NUM)) by the Federal Ministry of Education and Research of Germany (Bundesministerium für Bildung und Forschung (BMBF); Grant number 01KX2021). The Project RESPOND is funded by the EU RIA-call H2020-SC1-PHE-CORONAVIRUS-2020-2-RTD (Grant number 101016127). The funding body had no role in the design of the study and collection, analysis, and interpretation of data and in writing the manuscript. Open Access funding enabled and organized by Projekt DEAL.

Availability of data and materials

All data generated or analyzed during this study are included in this published article and its supplementary information files. Additional data (eg, detailed extracted data) are available from the corresponding author on request.

Ethics approval and consent to participate

Not applicable.

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Consent for publication

Not applicable.

Competing interests

LG, JSW, and GS have no conflicts of interest to disclose. AMK, NR, OT, MC, ER, HB, CS, JJM, and KL report grants from the Federal Ministry of Education and Research (BMBF), Germany, during the conduct of the study. JJM reports grants from the Federal Ministry of Health (BMG), Germany, outside of the submitted work.

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Received: 2 December 2020 Accepted: 4 February 2021 Published online: 29 March 2021

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