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RESEARCH ARTICLE

Psychological distress among healthcare providers during COVID-19 in Asia: Systematic review and meta-analysis

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Abstract

Introduction

COVID-19 pandemic is having a devastating effect on the mental health and wellbeing of healthcare providers (HCPs) globally. This review is aimed at determining the prevalence of depression, anxiety, stress, fear, burnout and resilience and its associated factors among HCPs in Asia during the COVID-19 pandemic.

Material and methods

We performed literature search using 4 databases from Medline, Cinahl, PubMed and Scopus from inception up to March 15, 2021 and selected relevant cross-sectional studies. Publication bias was assessed using funnel plot. Random effects model was used to estimate the pooled prevalence while risk factors were reported in odds ratio (OR) with 95% CI.

Results

We included 148 studies with 159,194 HCPs and the pooled prevalence for depression was 37.5% (95%CI: 33.8–41.3), anxiety 39.7(95%CI: 34.3–45.1), stress 36.4% (95%CI: 23.2–49.7), fear 71.3% (95%CI: 54.6–88.0), burnout 68.3% (95%CI: 54.0–82.5), and low resilience was 16.1% (95%CI: 12.8–19.4), respectively. The heterogeneity was high (I2>99.4%). Meta-analysis reported that both females (OR = 1.48; 95% CI = 1.30–1.68) and nurses (OR = 1.21; 95%CI = 1.02–1.45) were at increased risk of having depression and anxiety [(Female: OR = 1.66; 95% CI = 1.49–1.85), (Nurse: OR = 1.36; 95%CI = 1.16–1.58)]. Females were at increased risk of getting stress (OR = 1.59; 95%CI = 1.28–1.97).

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Abbreviations: Asi-3, Anxiety Sensitivity Index-3; BAS, Beck Anxiety Scales; BDI, Beck Depression Inventory; CAS, Coronavirus Anxiety Scale; CCMD-3, Chinese Classification and the Diagnose Criterion of Mental Disorder; CD-RISC, Connor-Davidson Resilience Scale; CES-D, Center for Epidemiology Studies-Depression; COVID-19, Novel coronavirus 2019; DASS-21, Depression-Anxiety-Stress Scale-21; FCV-19S, Fear of Coronavirus 2019 Scale; FS-HPs, Fear Scale for Healthcare Professionals; GAD, Generalized Anxiety Disorder; HADS, Hospital Anxiety and Depression Scale 14 items; HAMA, Hamilton Anxiety Rating Scale; HAMD, Hamilton Depression Rating Scale; IUS-12, Intolerance of Uncertainty Scale; MBI, Maslach Burnout Inventory; MBI-HSS, Maslach Burnout Inventory-Human Service Survey; MINI, Mini International Neuropsychiatric Interview; NA, Not available; NRS, Numerical rating scale; OLBI, Oldenburg Burnout Inventory; PHQ, Patient Health Questionnaire; PRISMA, Preferred Reporting Items for Systematic Review and Meta-analyses; PSS, Perceived Stress Scale; SAQ, Safety Attitudes Questionnaire; SAS, Zung's Self-Rating Anxiety Scale; SASRQ, Stanford Acute Stress Reaction Questionnaire; SDS, Zung's Self-rating Depression Scale; STAI, State and Trait Anxiety Inventory; STROBE, Strengthening the Reporting of Observational Studies in Epidemiology; WHO-5, WHO Well-being Index.

Conclusion

In conclusion, one third of HCPs suffered from depression, anxiety and stress and more than two third of HCPs suffered from fear and burnout during the COVID-19 pandemic in Asia.

Introduction

The COVID-19 pandemic is both a worldwide healthcare crisis and financial disaster at the biggest scale that has emerged in the last century. With the emergence of new COVID-19 virus variants, COVID-19 is even more infectious and can spread more rapidly through various inter-continentals [1, 2]. To date, about 171 million individuals had contracted the infection caused by this novel coronavirus and more than 3.5 million have succumbed to this virus across 222 countries [3].

There has been increasing concern that COVID-19 has infected nearly 570,000 healthcare providers (HCP) and killed more than 2500 of them in the Americas alone [4]. In the Asia Pacific region, a total of 12,454 HCP have been infected with the novel coronavirus while 171 have succumbed to this virus as of June 11, 2020 [5].

In performing their duties of arresting the spread of COVID-19, the HCP are risking their lives due to a higher risk of virus exposure, high workload demand, irregular or long working hours and increased psychological distress such as depression, anxiety, stress, occupation burnout, fear, low resilience as well as fatigue [1]. In addition, the HCP were barred from taking leaves and separated from their loved ones for up to weeks or even months. Wearing the full personal protective equipment or gear (PPE) that is very uncomfortable for long hours continuously every day while managing patients diagnosed with COVID-19 is extremely exhausting, particularly that this has become a routine task in their daily work. Literature reported that factors associated with personal-, work-, and patient-related burnout among HCPs were those that had direct involvement in COVID-19 management, underlying medical illness, and receiving inadequate psychological support in the workplace [6]. Those with higher total points in the coping score were significantly associated with reduction in anxiety and depression scores [7]. Other significant factors associated with psychological distress inluding but not limited to thought of resignation and reluctant to work, fear of infecting family members, frequent change in infection prevention and control protocol or guideline, and poor social support [8]. All of the aforementioned factors had been determined as factors that are leaving negative psychological impacts on the healthcare workers in Asian countries [8-10].

HCPs experiencing anxiety and depression have impaired physical and mental health and it may affect their wellbeing and work efficacy. In other word, the psychological consequences may contribute to a poorer quality of life among HCPs and suboptimal performance of delivered care. Experiencing these psychological hardships in the long run, can lower the immunity and put HCPs at a higher risk of being infected [11]. Furthermore, HCPs with pre-existing depression may suffer a higher mortality rate if they are hospitalised with COVID-19 [12]. Thus, it is very important to create the awareness on the degree of psychological distress that are encountered by HCPs amidst the COVID-19 pandemic in order to help reduce the incidence of occupation-related burnout and deaths. This can then serve as a platform for the government and policy makers to allocate funding to promote and provide psychosocial support for HCPs during this COVID-19 pandemic which will ultimately led to a better patient care. Recent systematic review involving 32 studies reported on the prevalence of mental health

among healthcare personnel during COVID-19 in Asia but there was no meta-analysis on pooled prevalence [13]. Another recently published systematic review by Md Mahbub Hossain et al involving 35 studies with 41,402 participants reported the prevalence of anxiety and depression during COVID-19 pandemic in South Asia but not in the Asia region specifically [14]. There was no systematic review and meta-analysis from Asia regarding these topics in which the burden of the psychological impacts could be different in view of the varietis of health care system across this region. Therefore, we are performing this systematic review and meta-analysis to determine the prevalence of depression, anxiety, stress, burnout, fear and low resilient among HCPs as well as its associated factors during the COVID-19 pandemic in Asia.

Materials and methods

Protocol

This present study is registered with INPLASY (Number: 202140043). We have also adhered to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guide-line [15] on conducting and reporting this systematic review and meta-analysis result as stated in Table A1 in S1 Table.

Literature search

Two authors (HR and CSM) performed literature search based on four databases (Medline, Cinahl, PubMed and Scopus databases) systematically and independently for potential articles published in 2020 to 13th of March, 2021. A combination of search terms that consists of (depression OR anxiety OR stress OR burnout, professional OR fatigue OR fear OR resilience, psychological OR adjustment) AND (healthcare workers OR medical staff OR healthcare professionals OR medical personnel) AND (coronavirus OR SARS-COV-2 OR COVID-19) AND (Asia) were used and is stated in Table A2 in S2 Table.

Selection criteria

The inclusion criteria for this systematic review were as follow:

- 1. The study design was cross-sectional with a minimum sample size of 100
- 2. The study stated the prevalence of depression, anxiety, stress, burnout, fear and resilience among HCPs during COVID-19 pandemic
- 3. The study evaluated depression, anxiety, stress, burnout, fear and resilience based on validated instrument tools or scales
- 4. The study involved HCPs from Asian countries
- 5. The studies must be published in English peer-reviewed journals.

Studies with the following criteria were excluded:

- 1. Perspective, opinion, review articles, case reports, short communications paper, no full text study and unpublished data
- 2. Data reported in continuous or qualitative format
- 3. Outcomes were not clearly defined by validated tools
- 4. Depression, anxiety, stress, burnout, fear and resilience were reported as independent data
- 5. Technical error was present in the reported data

6. After full-text articles have been assessed for eligibility, those outcomes were grouped into category of severities which were different from our operational definition.

Study selection

We performed the study selection according to the PRISMA guidelines. Studies identified using the search strategies were transferred into Endnote software (version 19) for screening, removing duplication and data extraction. Two authors (CSM and NKY) screened the title and abstract to determine the eligibility of the studies. For those potentially eligible articles, further screening on the full text had been performed to determine the availability of data and whether the articles fulfilled the selection or inclusion criteria (types of studies, participants, setting and outcomes). Any doubt on eligibility was resolved by discussion with a third author (KWL). Finally, those full-text articles which fulfilled all selection criteria were kept for data extraction and subsequent quantitative analysis.

PICO

The participants were HCPs (doctors, dentists, nurses, nurse assistants, midwives, medical assistants, pharmacists and other allied healthcare workers). Exposure was referred to actively providing healthcare-related services in conditions that are high risk for COVID-19 transmission and there is no comparator for the current systematic review. The main outcome for this review was pooled prevalence of depression, anxiety, depression, burnout, fear and resilience among HCPs.

Data extraction

We used Microsoft Excel to perform the data extraction and recording. The following data were extracted independently by two authors (SMC and KYN) and recorded as: Author information, publication year, country of the study, mean age or median of participants, study design, sample size, number of HCPs with and without symptoms of depression, anxiety, stress, burnout, fear and resilience, screening tool for assessment of depression, anxiety, stress, burnout, fear and resilience, sociodemographic and any clinical characteristics of the respondents.

Quality assessment

We used the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) checklist to perform quality assessment on all the included articles [16]. Two authors (CSM and NKY) individually assessed the study quality, and discrepancies were resolved by discussion with third investigator (LKW). STROBE check list consists of 22 items that assessed 6 components in cross-sectional studies. One point would be rewarded for a positive response for each of the items making the total score ranging from 0 to 22. Each article was graded as 'low risk of bias' if STROBE score $\geq 14/22$; or 'high risk of bias' if the score is <14/22 [16]. The STROBE scoring for this systematic review is presented in Appendix A. Studies were included in the analysis regardless of STROBE score and grades. The summary of the quality grade is reported in Table 1.

Operational definition

Psychological distress in this systematic review covers depression, anxiety, stress, fear, burnout, and low resilience symptoms. Presence of depression symptom was applied to those who scored points that belonged to mild to severe depression domain categories. Similarly, this

Table 1. Characteristic of 148 studies.

	No. of article	No. of country	Method of screening	Number of positve case	Total population
All	148	23			159194
Depression [8, 21–25, 27–31, 33, 34, 36–39, 43–50, 54– 58, 60–63, 65–67, 69–74, 76–80, 82, 84, 86, 87, 89, 90, 92, 93, 95, 97, 98, 101, 102, 104–106, 109, 111, 113– 115, 118, 122, 125, 127–131, 135, 136, 138, 139, 141– 144, 147, 149–152, 156–159, 161–163]	98	20	DASS-21, PHQ-2, PHQ-4, PHQ-9, HADS, HAMD, SDS, MINI, CES-D, CCMD-3, SCL-90, STAI, BDI, WHO-5	37630	103628
Anxiety [7–9, 22–25, 28–31, 33–40, 43–54, 56–58, 60– 63, 65–67, 69–74, 76–80, 82, 84, 86–93, 97, 98, 101– 112, 114–116, 118, 120–133, 135–139, 141–143, 147, 149–152, 154–163]	117	21	DASS-21, GAD-2, GAD-7, HAMA, SAS, MINI, PHQ-4, CCMD-3, HADS, ASI-3, SCL-90, COVID-19 Anxiety Scale, CAS, Dispositional cancer worry scale, STAI, IUS- 12 STAI, BAS	38284	99639
Stress [28, 29, 37, 43, 54, 57, 75, 76, 89, 90, 97–100, 102, 104, 105, 111, 118, 121, 122, 125, 127–131, 136, 138, 139, 141, 142, 148, 150, 151, 153, 158, 159, 162, 163]	40	17	DASS-21, PSS, SASRQ, SAQ, STAI.	8599	34010
Burnout [6, <u>32</u> , <u>41</u> , <u>85</u> , <u>140</u> , <u>142</u> , <u>158</u>]	7	5	MBI, ProQOL Scale, MBI-HSS, Adopted Queationnaire, OLBI.	5426	8732
Fear [31, 56, 105, 143, 156]	5	4	FCV-19S, FS-HPS, CCMD-3, NRS, STAI.	3460	4302
Low Resilience [9, 94]	2	2	CD-RISC	78	484

Abbreviation:

ASI-3: Anxiety Sensitivity Index-3;

BAS: Beck Anxiety Scales;

BDI: Beck Depression Inventory;

CAS: Coronavirus Anxiety Scale;

CCMD-3: Chinese Classification and the Diagnose Criterion of Mental Disorder;

CD-RISC: Connor-Davidson Resilience Scale;

CES-D: Center for Epidemiology Studies-Depression;

DASS-21: Depression-Anxiety-Stress Scale-21;

FCV-19S: Fear of Coronavirus 2019 Scale;

FS-HPs: Fear Scale for Healthcare Professionals;

GAD-2: 2-item Generalized Anxiety Disorder;

GAD-7: 7-item Generalized Anxiety Disorder;

HADS: Hospital Anxiety and Depression Scale 14 items;

HAMA: Hamilton Anxiety Rating Scale;

HAMD: Hamilton Depression Rating Scale;

IUS-12: Intolerance of Uncertainty Scale;

MBI: Maslach Burnout Inventory;

MBI-HSS: Maslach Burnout Inventory- Human Service Survey

MINI: Mini International Neuropsychiatric Interview;

NRS: Numerical rating scale

OLBI: Oldenburg Burnout Inventory;

PHQ-2: 2-item Patient Health Questionnaire;

PHQ-4: Patient Health Questionnaire-4;

PHQ-9: 9-item Patient Health Questionnaire;

PSS: Perceived Stress Scale;

SAQ: Safety Attitudes Questionnaire;

SAS: Zung's Self-Rating Anxiety Scale;

SASRQ: Stanford Acute Stress Reaction Questionnaire;

SDS: Zung's Self-rating Depression Scale;

STAI: State and Trait Anxiety Inventory;

WHO-5: WHO Well-being Index;

applied for anxiety, stress and fear symptoms. Burnout is characterized by those who ranked under moderate to severe categories. Low resilience is defined as those classified under a low score category.

Data synthesis

We use the *Open Meta Analyst* and StatsDirect to perform the meta-analyses [17, 18]. We used a random-effects model (DerSimonian and Laird Method) to calculate the pooled prevalence from multiple studies [19, 20]. The numerator would be the total number of cases summed up of HCPs with positive symptoms; denominator would be total number of HCPs in that study. Values of numerator and denominators of all studies were used to derive the pooled prevalence of outcome variables. The random-effects model was used so that the findings were generalisable and more representative presuming that the studies were randomly selected from a bigger population [21]. We used I² statistics index to assess the heterogeneity across the studies. I² index is categorised as low if < 25%, moderate 25–50%, and high > 50%) [20].

Subgroup analyses

Subgroup analyses are useful to examine the between-group differences in terms of the prevalence as a possible cause of heterogeneity across studies. The prevalence of depression, anxiety, stress, burnout, fear and resilience among HCPs in Asia was determined by subgrouping the countries. The data were further examined by subgrouping the gender and occupational subtype of HCPs. The risk factors for depression, anxiety, stress, burnout, fear and resilience were reported in odds ratio (OR) with a 95% confidence interval (CI).

Sensitivity analysis

We performed sensitivity analysis by using leave-one-out meta-analysis to examine how each particular study alters the overall performance of the rest of the studies especially the pooled prevalence estimates and heterogeneity.

Publication bias

We assessed the potential publication bias by visually checking the Funnel plot followed by conducting the Begg's test and Egger's regression test. In Egger's test, a p-value < 0.01 was defined as an indicator for publication bias [22].

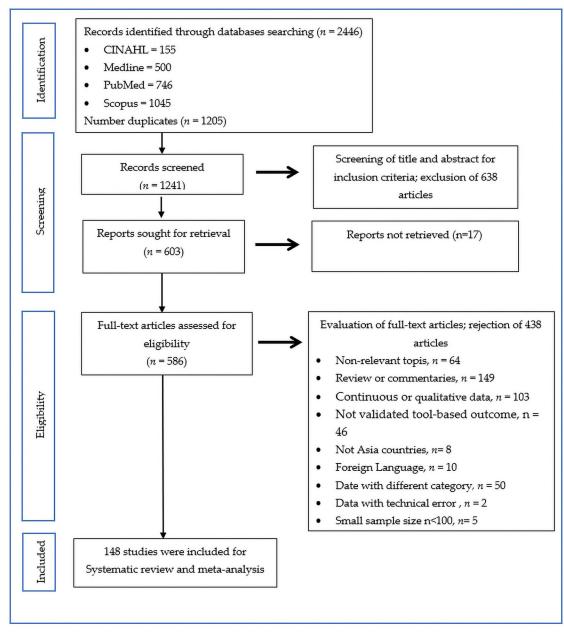
Results

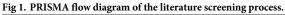
Search result

Fig 1 shows the PRISMA flow diagram used in this review. A total of 2446 articles were identified from four databases: CINAHL (155), Medline (500), PubMed (746) and Scopus (1045). After removing the duplicate records, we performed screening on 1241 articles. Finally, we included 148 studies for systematic review and meta-analysis after further evaluation of eligibility.

Description of included studies

Table 1 summarizes the main characteristics of 148 studies. A total of 159,194 healthcare providers from 23 different Asia countries were involved in this systematic review and meta-analysis. Almost half (n = 70) of the studies were conducted in China [8, 11, 23–90], followed by 15 studies in India [10, 91–104], 11 studies in Turkey [105–115], 10 studies in Saudi Arabia





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[116–125], 6 studies in Pakistan [126–131], 4 studies in Indonesia [132–135] and Nepal [136–139] respectively, 3 studies in Malaysia [140–142], Singapore [143–145], Japan [146–148] and Iran [149–151] respectively with 2 studies in Oman [152, 153], Jordan [154, 155], Philippines [156, 157] and Bangladesh [158, 159] respectively. Besides, Korea [160], Qatar [161], and Iraq [162] which each had one study, there were also some multinational studies that were conducted, with 2 of them involving several countries in Asian Pacific [9, 163] and another one involving both Egypt and Saudi Arabia [164] (Table A4 in S4 Table).

About two thirds (n = 98) of the studies reported the data on depression [8, 23–27, 29–33, 35, 36, 38–41, 45–52, 56–60, 62–65, 67–69, 71–76, 78–82, 84, 86, 88, 89, 91, 92, 94, 95, 97, 99,

100, 103, 104, 106–108, 111, 113, 115–117, 120, 123, 124, 127, 129–133, 137, 138, 140, 141, 143–146, 149, 151–154, 158–161, 163–165]. Different type of tools or scales were used to diagnosed depression in different papers from different Asian countries (Table 1 and S4 Table), such as 2-item Patient Health Questionnaire (PHQ-2), 9-item Patient Health Questionnaire (PHQ-9), Hamilton Depression Rating Scale (HAMD), Hospital Anxiety and Depression Scale 14 items (HADS), Chinese edition of Zung's Self-Rating Depression Scale (SDS), Center for Epidemiology Studies-Depression (CES-D), Chinese Classification and the Diagnose Criterion of Mental Disorder (CCMD-3), Hospital Anxiety and Depression Scale 14 items (HADS), Symptom Checklist 90, and also Beck Depression Inventory (BDI).

A total of 117 out of 148 studies described the anxiety data in terms of frequency or percentage [8, 10, 11, 24–27, 30–33, 35–42, 45–56, 58–60, 62–65, 67–69, 71–76, 78–82, 84, 86, 88–95, 99, 100, 103–114, 116–118, 120, 122–135, 137–141, 143–145, 149, 151–154, 156–165]. Tools that were used to screen for anxiety were Depression-Anxiety-Stress Scale-21 (DASS-21), 2-item and 7-item Generalized Anxiety Disorder (GAD-2& GAD-7), Hospital Anxiety and Depression Scale 14 items (HADS), Self-Rating Anxiety Scale (SAS), Hamilton Anxiety Scale (HAMA), Patient Health Questionnaire-4 (PHQ-4), Coronavirus Anxiety Scale (CAS), Chinese Classification and the Diagnose Criterion of Mental Disorder (CCMD-3), State and Trait Anxiety Inventory (STAI), Dispositional cancer worry scale, Symptom Checklist 90, and Beck Anxiety Scales (BAS).

There were 40 studies which reported on data regarding stress, using five different type of scales which were Depression-Anxiety-Stress Scale-21 (DASS-21), Safety Attitudes Question-naire (SAQ), Perceived Stress Scales (PSS), Stanford Acute Stress Reaction Questionnaire (SASRQ), and State and Trait Anxiety Inventory (STAI) [30, 31, 39, 56, 59, 77, 78, 91, 92, 99–102, 104, 106, 107, 113, 120, 123, 124, 127, 129–133, 138, 140, 141, 143, 144, 150, 152, 153, 155, 160, 161, 164, 165].

Only 7 studies investigated data on burnout in term of its' frequency or percentage. The most common tools used to screen burnout among healthcare providers was Maslach Burnout Inventory (MBI), followed by ProQOL Scale of Chinese version, and Oldenburg Burnout Inventory (OLBI) [9, 34, 43, 87, 142, 144, 160]. A validated questionnaire form was adopted from Michelle Post, Public Welfare, Vol. 39, No. 1, 1981, American Public Welfare Association to examine the prevalence of burnout.

On the other hand, 5 studies reported data on fear, using five different scales which were Fear of Coronavirus 2019 Scale (FCV-19S), Fear Scale for Healthcare Professionals (FS-HPs), Chinese Classification and the Diagnose Criterion of Mental Disorder (CCMD-3), Numerical rating scale (NRS), and State and Trait Anxiety Inventory (STAI) [33, 58, 107, 145, 158]. Besides, two studies that reported on low resilience data were using the Connor-Davidson Resilience Scale (CD-RISC) as screening tool [11, 96].

Pooled prevalence of psychological distress among healthcare providers amidst COVID-19 pandemic

A summary of pooled prevalence of mental illnesses among healthcare providers during COVID-19 pandemic is shown in Table 2 and S1–S5 Figs. The overall pooled prevalence of mild to severe depression among HCPs is 37.5 (95% CI = 33.8–41.3). (Fig 2) Jordan recorded the highest prevalence of depression among HCPs at 78.0% (95% CI = 75.6–80.4), followed by a multicentre study involved Egypt and Saudi Arabia at 69.0% (95% CI = 64.6–73.4), and Iran at 59.6% (95% CI = 37.4–81.9). Pooled prevalence of depression of 52 studies in China was reported as 36.5% (95% CI = 31.7–41.2) while that in Malaysia was 26.6% (95% CI = 17.9–35.3) (S1 Fig).

Domain	Country	N	Total Psychological distress	Total sample size	Prevalence,% (95% CI)	I ² (p-value)	Appendix
Depression	All	98	37630	103628	37.5 (33.8-41.3)	99.49 (<0.001)	Fig 2
	Asian Pacific region	1	51	1146	4.5 (3.3-5.6)	NA	NA
	Bangladesh	2	342	782	43.5 (33.5-53.4)	87.95 (0.004)	Fig A1.1 in (S1 Fig)
	China	52	22772	66052	36.5 (31.7-41.2)	99.51 (<0.001)	Fig A1.2 in (S1 Fig)
	Egypt and Saudi	1	294	426	69.0 (64.6-73.4)	NA	NA
	India	9	1804	5573	33.6 (27.9–39.3)	94.62 (<0.001)	Fig A1.3 in (S1 Fig)
	Indonesia	2	268	1326	20.2 (14.1-26.4)	87.70 (0.004)	Fig A1.4 in (S1 Fig)
	Iran	2	548	928	59.6 (37.4-81.9)	98.10 (<0.001)	Fig A1.5 in (S1 Fig)
	Japan	1	237	848	27.9 (24.9-31.0)	NA	NA
	Jordan	1	907	1163	78.0 (75.6-80.4)	NA	NA
	Korea	1	20	115	17.4 (10.5–24.3)	NA	NA
	Malaysia	2	413	1449	26.6 (17.9-35.3)	91.99 (<0.001)	Fig A1.6 in (S1 Fig)
	Nepal	2	295	879	33.2 (24.9-41.6)	86.17 (0.007)	Fig A1.7 in (S1 Fig)
	Oman	2	550	1541	38.6 (25.9–51.3)	95.19 (<0.001)	Fig A1.8 in (S1 Fig)
	Pakistan	4	4897	10790	44.6 (19.3–69.9)	99.54 (<0.001)	Fig A1.9 in (S1 Fig)
	Qatar	1	54	127	42.5 (33.9–51.1)	NA	NA
	Saudi Arabia	5	1313	2483	44.2 (28.7–59.6)	98.20 (<0.001)	Fig A1.10 in (S1 Fig)
	Singapore	2	1043	3197	41.7 (21.5-61.9)	95.03 (<0.001)	Fig A1.11 in (S1 Fig)
	Singapore and India	1	96	906	10.6 (8.6–12.6)	NA	NA
	Turkey	6	1691	3624	46.8 (23.8-69.8)	99.58 (<0.001)	Fig A1.12 in (S1 Fig)
	Vietnam	1	35	173	20.2 (14.2-26.2)	NA	NA
Anxiety	All	117	38284	99639	39.7(34.3-45.1)	99.78(<0.001)	Fig 3
	Asian Pacific region	1	60	1146	5.2(3.9-6.5)	NA	NA
	Bangladesh	2	414	782	52.1(21.5-82.7)	98.81 (<0.001)	Fig A2.1 in (S2 Fig)
	China	56	16605	54004	31.9 (27.8-36.0)	99.27 (<0.001)	Fig A2.2 in (S2 Fig)
	Egypt and Saudi	1	251	426	58.9(54.2-63.6)	NA	NA
	India	10	1807	4098	44.2(32.6-55.9)	98.41 (<0.001)	Fig A2.3 in (S2 Fig)
	Indonesia	4	1122	2054	56.5(39.0-74.0)	98.56 (<0.001)	Fig A2.4 in (S2 Fig)
	Iran	3	794	1330	59.1(40.8-77.3)	98.07 (<0.001)	Fig A2.5 in (S2 Fig)
	Iraq	2	470	889	51.8(30.2-73.4)	97.78 (<0.001)	Fig A2.6 in (S2 Fig)
	Jordan	1	823	1163	70.8(68.2-73.4)	NA	NA
	Korea	1	23	115	20.0(12.7-27.3)	NA	NA
	Malaysia	2	438	1449	30.2(27.9-32.6)	0 (0.493)	Fig A2.7 in (S2 Fig)
	Nepal	2	343	879	38.8(32.7-44.9)	72.36 (0.06)	Fig A2.8 in (S2 Fig)
	Oman	2	659	1541	50.7(18.0-83.4)	99.33 (<0.001)	Fig A2.9 in (S2 Fig)
	Pakistan	6	4732	11372	52.4(31.9-72.9)	99.45 (<0.001)	Fig A2.10 in (S2 Fig)
	Philippines	2	525	1061	46.3(29.9-62.8)	96.23 (<0.001)	Fig A2.11 in (S2 Fig)
	Qatar	1	53	127	41.7(33.2-50.3)	NA	NA
	Saudi Arabia	8	4988	8426	45.7(31.9-59.5)	99.25 (<0.001)	Fig A2.12 in (S2 Fig)
	Singapore	2	1308	3197	40.9(39.2-42.6)	0 (0.345)	Fig A2.13 in (S2 Fig)
	Singapore and India	1	142	906	15.7(13.3-18.0)	NA	NA
	Turkey	10	3065	5289	48.9(27.0-70.7)	99.8 (<0.001	Fig A2.14 in (S2 Fig)
	Vietnam	1	58	173	33.5(26.5-40.6)	NA	NA

Table 2. Summary of pooled prevalence of psychological distress among healthcare providers during COVID-19 pandemic.

(Continued)

Domain	Country	N	Total Psychological distress	Total sample size	Prevalence,% (95% CI)	I ² (p-value)	Appendix
Stress	All	40	8599	34010	36.4 (23.2–49.7)	99.45 (<0.001)	Fig 4
	China	8	2876	8552	29.3 (12.5-46.1)	99.75 (<0.001)	Fig A3.1 in (S3 Fig)
	Egypt and Saudi	1	238	426	55.9 (51.2-60.6)	NA	NA
	India	7	1078	1845	49.4 (23.8–74.9)	99.51 (<0.001)	Fig A3.2 in (S3 Fig)
	Indonesia	2	570	1326	43.3 (20.5-66.1)	98.71 (<0.001)	Fig A3.3 in (S3 Fig)
	Iran	1	217	217	99.8 (99.1-100.4)	NA	NA
	Jordan	1	287	448	64.1 (59.6-68.5)	NA	NA
	Korea	1	5	115	4.3 (0.6-8.1)	NA	NA
	Malaysia	2	363	1449	26.0 (20.6-31.4)	77.65 (0.034)	Fig A3.4 in (S3 Fig)
	Nepal	1	69	404	17.1 (13.4–20.7)	NA	NA
	Oman	2	486	1541	38.6 (9.5-67.7)	99.12 (<0.001)	Fig A3.5 in (S3 Fig)
	Pakistan	4	635	10790	33.6 (-6.4-73.6)	99.89 (<0.001)	Fig A3.6 in (S3 Fig)
	Qatar	1	39	127	30.7 (22.7–38.7)	NA	NA
	Saudi Arabia	3	128	498	29.9 (0.2-59.6)	99.31 (<0.001)	Fig A3.7 in (S3 Fig)
	Singapore	1	205	3075	6.7 (5.8–7.5)	NA	NA
	Singapore and India	1	47	906	5.2 (3.7-6.6)	NA	NA
	Turkey	3	1334	2118	46.9 (6.2-87.5)	99.76 (<0.001)	Fig A3.8 in (S3 Fig)
	Vietnam	1	22	173	12.7 (7.8–17.7)	NA	NA
Burnout	All	7	5426	8732	68.3 (54.0-82.5)	99.5 (<0.001)	Fig 5
	Asian Pacific region	1	182	301	60.5(54.9-66.0)	NA	NA
	China	3	2859	5025	58.0 (30.5-85.6)	99.78 (<0.001)	Fig A4.1 in (S4 Fig)
	Korea	1	104	115	90.4(85.1-95.8)	NA	NA
	Malaysia	1	184	216	85.2(80.4-89.9)	NA	NA
	Singapore	1	2097	3075	68.2(66.5-69.8)	NA	NA
Fear	All	5	3460	4302	71.3 (54.6-88.0)	99.83 (<0.001)	Fig 6
	Bangladesh	1	370	370	99.9 (99.5-100.2)	NA	NA
	China	2	1890	2353	53.4 (-20.6-127.5)	99.93 (<0.001)	Fig A5.1 in (S5 Fig)
	Singapore	1	89	122	73.0 (65.1-80.8)	NA	NA
	Turkey	1	1111	1457	76.3 (74.1–78.4)	NA	NA
Low resilience	All	2	78	484	16.1(12.8-19.4)	0(0.922)	Fig 7
	China	1	59	364	16.2(12.4-20.0)	NA	NA
	India	1	19	120	15.8(9.3-22.4)	NA	NA

Table 2. (Continued)

https://doi.org/10.1371/journal.pone.0257983.t002

The pooled prevalence of mild to severe anxiety among HCPs in Asian countries was 39.7% (95% CI = 34.3–45.1). (Fig 3) Jordan, again recorded the highest prevalence of anxiety at 70.8% (95% CI = 68.2–73.4) among HCPs while the lowest prevalence of anxiety was found in a study conducted across the Asian Pacific region. Prevalence of anxiety in China was 31.9% (95% CI = 27.8–36.0), which is almost similar with Malaysia at 30.2% (95% CI = 27.9–32.6).

The overall pooled prevalence of mild to severe stress was found to be 36.4% (95% CI = 23.2–49.7) (Fig 4), with the highest prevalence reported in Iran at 99.8% (95% CI = 99.1–100.4) whereas the lowest prevalence was reported in Korea at 4.3% (95% CI = 0.6–8.1).

Pooled prevalence of moderate to severe burnout among HCPs in Asia was 68.3% (95% CI = 54.0–82.5) (Fig 5) which was relatively higher as compared to others mental health issues aforementioned. Korea was ranked number one for the pooled prevalence of burnout at 90.4% (95% CI = 85.1–95.8), followed by Malaysia at 85.2% (95% CI = 80.4–89.9) and Singapore at

Studies	Estim	mate (95	§ C.I.)	Ev/Trt	
Jo et al 2020	0.012	(0.000,	0.025)	3/253	-
		(0.033,		51/1146	- -
		(0.032,		12/170	- - -
Zhang et al A 2021	0.087	(0.061,	0.113)	39/450	
		(0.066, (0.086,		28/276 96/906	
		(0.086,		12/110	
		(0.103,		268/2299	
Zhang et al B 2020	0.122	(0.101,	0.143)	113/927	-
Mi et al 2021	0.133	(0.112,	0.154)	137/1029	+
		(0.113,		117/863	+
		(0.104,		48/339	- -
		(0.133, (0.088,		301/2031 19/126	
Mathur et al 2020	0.170	(0.118,	0.222)	34/200	
Vargaretha et al 2020	0.172	(0.143,	0.200)	117/682	
		(0.105,		20/115	_ - -
		(0.142,		56/302	
Sun et al B 2020 Ni et al 2020	0.188	(0.155, (0.139,	0.222)	101/536 41/214	
		(0.142,		35/173	
		(0.154,		50/245	_
Guo et al 2021	0.207	(0.160,	0.255)	58/280	_ - _
Woon et al 2020	0.221	(0.180,	0.261)	88/399	
Bahadir-Yilmaz et al 2020	0.232	(0.210,	0.254)	338/1457	
		(0.202, (0.216,		151/644 153/612	
		(0.216,			
		(0.175,		31/123	_ -
Kiong et al 2020	0.265	(0.207,	0.322)	59/223	_ - _
Awano et al 2020	0.279	(0.249,	0.310)	237/848	-
		(0.249, (0.245,		211/749 117/404	
		(0.245,		135/456	_
Xing et al A 2020	0.297	(0.259,	0.336)	163/548	
Xiaoming et al 2020	0.302	(0.293,	0.312)	2666/8817	=
		(0.282,		325/1050	-
		(0.273,		164/524	-
		(0.288, (0.271,		354/1124 120/377	
Tan et al 2020	0.318	(0.302,	0.335)	979/3075	
Kumar et al 2021	0.321	(0.260,	0.383)	72/224	
Alshekaili et al 2020	0.323	(0.296,	0.350)	368/1139	
Khanna et al 2020	0.325	(0.306,	0.344)	765/2355	
Wang et al D 2021 Li et al A 2020	0.328	(0.290, (0.299,	0.366)	192/586 299/908	
		(0.327,		1107/3228	-
		(0.273,		53/152	
Wang et al E 2020	0.361	(0.336,	0.385)	546/1514	-
Fang et al 2021	0.364	(0.322,	0.406)	186/511	
		(0.331, (0.308,		178/475 73/194	
		(0.308,		142/370	
Wang et al A 2020	0.393	(0.363,	0.422)	412/1049	
Cao et al 2020	0.400	(0.356,	0.444)	192/480	
		(0.347,		82/197	
		(0.347, (0.339,		82/197 54/127	
		(0.339,		54/12/ 878/2014	-
An et al 2020	0.436	(0.407,	0.465)	481/1103	
Zhu et al A 2020	0.442	(0.367,	0.518)	73/165	
		(0.423,		1014/2285	-
		(0.404,		182/402 46/100	
		(0.362, (0.452,		46/100 4696/10178	
Li et al B 2020	0.467	(0.401,	0.532)	105/225	
Hassannia et al 2020	0.483	(0.438,	0.527)	235/487	
		(0.437,		200/412	
He et al 2021 Chen et al J 2021	0.486	(0.438,	0.535)	196/403 439/902	
Chen et al J 2021 Teng et al 2020	0.487	(0.454, (0.438,	0.537	439/902 194/398	
		(0.438,		137/281	
Al Ammari et al 2021	0.492	(0.455,	0.528)	354/720	
	0.494	(0.442,	0.547)	173/350	
		(0.483,		1730/3460	•
		(0.477, (0.482,		634/1257 792/1563	-
		(0.402,		56/107	
Teo et al 2020	0.525	(0.436,	0.613)	64/122	_ _
Cai et al 2020	0.528	(0.491,	0.564)	374/709	
		(0.446,		76/144	
		(0.528,		1458/2667 173/309	
		(0.542,		549/958	
Zhou et al A 2020	0.576	(0.537,	0.615)	349/606	
Lu et al A 2020	0.579	(0.529,	0.628)	221/382	
		(0.505,		99/171	
		(0.538,		303/522	
AbuSnieneh et al. 2021 Calikan et al.2020	0.596	(0.569, (0.565,	0.677	754/1265	
		(0.613,		274/416	
		(0.646,		294/426	
Arafa et al 2021	0.710	(0.667,	0.752)	313/441	
Pouralizadeh et al 2020	0 735	(0.701,	0.769)	472/642	
Pouralizadeh et al 2020 Zhang et al F 2020	0.755		0.803)	729/939	
Pouralizadeh et al 2020 Zhang et al F 2020 Sahin et al 2020	0.776			003 /	
Pouralizadeh et al 2020 Zhang et al F 2020 Sahin et al 2020 Naser et al 2020	0.776	(0.756,	0.804)	907/1163	<u>-</u>
Pouralizadeh et al 2020 Zhang et al F 2020 Sahin et al 2020 Naser et al 2020 Liang et al 2020	0.776 0.780 0.825	(0.756, (0.801,	0.804) 0.850)	742/899	*
Pouralizadeh et al 2020 Zhang et al F 2020 Sahin et al 2020 Naser et al 2020 Liang et al 2020 Sandesh et al 2020	0.776 0.780 0.825 0.902	(0.756, (0.801, (0.847,	0.804) 0.850) 0.957)	742/899 101/112	-
Pouralizadeh et al 2020 Zhang et al F 2020 Sahin et al 2020 Naser et al 2020 Liang et al 2020	0.776 0.780 0.825 0.902	(0.756, (0.801, (0.847,	0.804) 0.850) 0.957)	742/899 101/112	-
Pouralizadeh et al 2020 Zhang et al F 2020 Sahin et al 2020 Naser et al 2020 Liang et al 2020 Sandesh et al 2020	0.776 0.780 0.825 0.902	(0.756, (0.801, (0.847,	0.804) 0.850) 0.957)	742/899 101/112	*

Fig 2. Forest plot of overall pooled prevalence of depression.

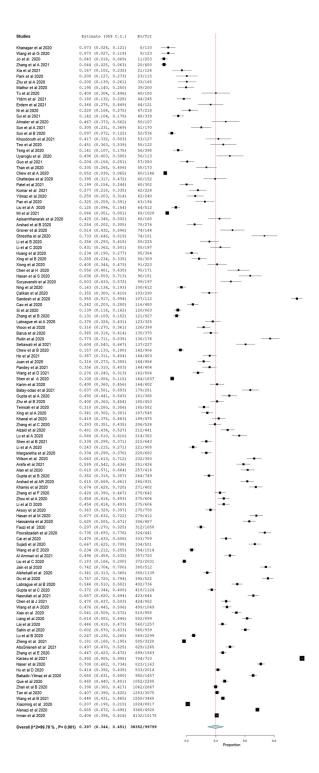


Fig 3. Forest plot of overall pooled prevalence of anxiety.

Studies	Estir	nate (95	% C.I.)	Ev/Trt							
Khanagar et al 2020	0.009	(0.000,	0.027)	1/110							
Park et al 2020	0.043	(0.006,	0.081)	5/115	-	-					
Imran et al 2020	0.044	(0.040,	0.048)	448/10178	-						
Chew et al B 2020	0.052	(0.037,	0.066)	47/906	-	-					
Tan et al 2020	0.067	(0.058,	0.075)	205/3075							
Arshad et al B 2020	0.072	(0.042,	0.103)	20/276	-						
Si et al 2020	0.086	(0.067,	0.104)	74/863							
Mathur et al 2020	0.095	(0.054,	0.136)	19/200							
Guo et al 2021	0.096	(0.062,	0.131)	27/280							
Yldrm et al 2021	0.098	(0.061,	0.135)	24/245							
Liu et al C 2020	0.100	(0.087,	0.113)	203/2031		-					
Than et al 2020	0.127	(0.078,	0.177)	22/173							
Grover et al 2020	0.139	(0.082,	0.195)	20/144			-				
Li et al B 2020	0.160	(0.112,	0.208)	36/225			_				
Sun et al A 2021	0.171	(0.114,	0.227)	29/170							
Pandey et al 2021	0.171	(0.134,	0.207)	69/404			_				
Balay-odao et al 2021	0.178	(0.133,	0.223)	50/281							
Fauzi et al 2020	0.235	(0.210,	0.261)	247/1050			-				
Alshekaili et al 2020	0.238	(0.213,	0.263)	271/1139							
Kumar et al 2021	0.263	(0.206,	0.321)	59/224							
Woon et al 2020	0.291	(0.246,	0.335)	116/399			_ _ _				
Khoodoruth et al 2021	0.307	(0.227,	0.387)	39/127				+			
Margaretha et al 2020	0.317	(0.282,	0.352)	216/682							
Chatterjee et al 2020	0.329	(0.254,	0.404)	50/152				-			
Zhan et al A 2020		(0.417,		789/1794							
Zhan et al B 2020		(0.468,		1298/2667				-			
Khamis et al 2020		(0.486,		215/402							
Sunil et al 2021		(0.485,		169/313					<u> </u>		
Nasrullah et al 2021		(0.511,		354/644					-		
Arafa et al 2021		(0.512,		238/426					•		
Alan et al 2020		(0.525,		238/416					•		
Shahrour et al 2020		(0.596,		287/448							
Patel et al 2021		(0.612,		201/302							
Almater et al 2020		(0.635,		77/107							
Bahadir-Yilmaz et al 2020				1072/1457							
Gu et al 2020		(0.771,		420/522							
Wilson et al 2020		(0.786,		289/350							
Podder et al 2020		(0.825,		330/384							_
Sandesh et al 2020		(0.930,		108/112							
Moayed et al 2021	0.998	(0.991,	1.000)	217/217							-
Overall (I^2=99.95 % , P< 0.001)	0.364	(0.232,	0.497)	8599/34010							
					_			· .			
					0		0.2	0.4 Proportio	0.6 m	0.8	1
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Fig 4. Forest plot of overall pooled prevalence of stress.

https://doi.org/10.1371/journal.pone.0257983.g004

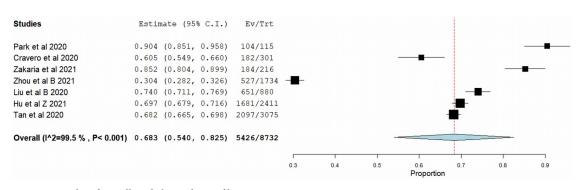


Fig 5. Forest plot of overall pooled prevalence of burnout.

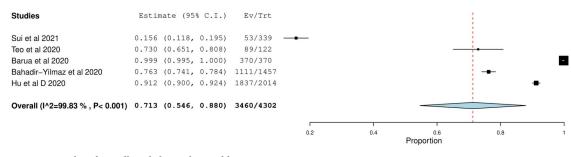


Fig 6. Forest plot of overall pooled prevalence of fear.

https://doi.org/10.1371/journal.pone.0257983.g006

68.2% (95% CI = 66.5-69.8); while China had the lowest prevalence at 58.0% (95% CI = 30.5-85.6) among the 5 countries that were investigated for the prevalence of burnout.

The pooled prevalence of mild to severe fear was 71.3% (95% CI = 54.6–88.0) (Fig.6). The highest prevalence of fear was found in Bangladesh at 99.9% (95% CI = 99.5–100.2) while the lowest prevalence was found in China at 53.4% (95% CI = -20.6-127.5).

Low resilience had the pooled prevalence of 16.1% (95% CI = -12.8-19.4) (Fig 7), with 16.2% (95% CI = 12.4-20.0) and 15.8% (95% CI = 9.3-22.4) being reported in China and India, respectively.

Subgroup analysis for the prevalence of psychological distress among healthcare providers amidst COVID-19 pandemic according to gender and occupation

Table 3 summarized the subgroup analysis of pooled prevalence of mental illness among healthcare providers during COVID-19 pandemic according to gender and occupation. The pooled prevalence of depression was higher in females (40.9%; 95%CI = 33.4-48.4) than males (35.5%; 95%CI = 29.5-41.6). Among all the HCPs, the nurses (39.3%; 95%CI = 33.2-45.3) had the highest prevalence of depression when compared to doctors (36.4%; 95%CI = 30.6-42.3) and allied healthcare personals (34.3%; 95%CI = 23.5-45.1) (\$1 Fig).

Similarly females (50.6%; 95%CI = 43.5–57.6) generally were more anxious than males (41.2%; 95%CI = 32.0–50.4) while analysing the gender subgroup for anxiety. Nurses, assistant nurses and midwives (43.1%; 95%CI = 36.6–49.7) had the highest prevalence for anxiety, followed by doctors and dentists (39.6%; 95%CI = 34.5–44.7), and finally by allied healthcare personals and pharmacists (38.6%; 95%CI = 26.2–51.0).

For the prevalence on stress, the female population (48.1%; 95%CI = 31.6–64.5) was still having higher prevalence than male population (40.4%; 95%CI = 22.8–57.9). Almost half of those who worked as nurses, assistant nurses and midwifes (45.4%; 95%CI = 29.4–61.4)

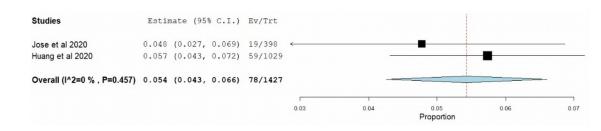


Fig 7. Forest plot of the overall pooled prevalence of low resilience.

Domain	Subgroup	N	Total Psychological distress	Total sample size	Prevalence,% (95% CI)	I ² (p-value)	Appendix
Depression	Female	29	7432	17595	40.9 (33.4-48.4)	99.12 (<0.001)	Fig A1.13 in (<u>S1</u> Fig)
	Male		3883	10556	35.5 (29.5-41.6)	97.49 (<0.001)	Fig A1.14 in (<u>S1</u> Fig)
	Doctor/Dentist 3		9679	23488	36.4 (30.6-42.3)	98.79 (<0.001)	Fig A1.15 in (S1 Fig)
	Nurse/Assistant nurse/midwife	38	8913	21725	39.3 (33.2–45.3)	98.98 (<0.001)	Fig A1.16 in (<u>S1</u> Fig)
	Allied healthcare personal/ Pharmacist	17	966	2578	34.3 (23.5-45.1)	97.96 (<0.001)	Fig A1.17 in (<u>S1</u> Fig)
Anxiety	Female	34	9359	19315	50.6 (43.5–57.6)	99.07 (<0.001)	Fig A2.15 in (<u>S2</u> Fig)
	Male	33	5288	12434	41.2 (32.0–50.4)	99.21 (<0.001)	Fig A2.16 in (<u>S2</u> Fig)
	Doctor/Dentist	41	9942	23846	39.6 (34.5-44.7)	98.37 (<0.001)	Fig A2.17 in (<u>S2</u> Fig)
	Nurse/Assistant nurse/midwife		10445	24679	43.1 (36.6–49.7)	99.25 (<0.001)	Fig A2.18 in (<u>S2</u> Fig)
	Allied healthcare personal / Pharmacist		2279	4302	38.6 (26.2–51.0)	98.70 (<0.001)	Fig A2.19 in (<u>S2</u> Fig)
Stress	Female		1752	9812	48.1 (31.6-64.5)	99.78 (<0.001)	Fig A3.9 in (S3 Fig)
	Male 1		957	6393	40.4 (22.8–57.9)	99.62 (<0.001)	Fig A3.10 in (<u>S3</u> Fig)
	Doctor/Dentist 1		1698	13938	33.5 (22.7-44.2)	99.60 (<0.001)	Fig A3.11 in (<u>S3</u> Fig)
	Nurse/Assistant nurse/midwife		4229	9142	45.4 (29.4–61.4)	99.68 (<0.001)	Fig A3.12 in (<u>S3</u> Fig)
	Allied healthcare personal / Pharmacist	5	149	568	31.4 (15.7–47.1)	95.35 (<0.001)	Fig A3.13 in (<u>S3</u> Fig)
Burnout	Doctor/Dentist	5	1115	1575	74.9 (62.8–87.1)	95.19 (<0.001)	Fig A4.2 in (S4 Fig)
	Nurse/Assistant nurse/midwife 2		1012	1431	80.2 (56.8–103.7)	98.83 (<0.001)	Fig A4.3 in (S4 Fig)
	Allied healthcare personal / Pharmacist	1	24	37	64.9 (49.5-80.2)	NA	NA
Fear	Male	1	233	233	99.8 (99.2–100.4)	NA	NA
	Female	1	147	147	99.7 (98.7–100.6)	NA	NA
	Doctor/Dentist	1	370	370	99.9 (99.5-100.2)	NA	NA
	Nurse/Assistant nurse/midwife	3	3001	3810	61.1 (28.0–94.1)	99.85 (<0.001)	Fig A5.2 in (<u>S5 Fig</u>)
	Allied healthcare personal / Pharmacist	1	89	122	73.0 (65.1–80.8)	NA	NA
Low	Female	1	15	88	17.0 (9.2–24.9)	NA	NA
resilience	Male	1	4	32	12.5 (1.0-24.0)	NA	NA
	Nurse/Assistant nurse/midwife	1	19	120	15.8 (9.3-22.4)	NA	NA

Table 3. Subgroup analysis of pooled prevalence of psychological distress among healthcare providers during COVID-19 pandemic according to gender and occupation.

experience stress, while about one third of doctors or dentists (33.5%; 95%CI = 22.7-44.2), and allied healthcare personals or pharmacists (31.4%; 95%CI = 15.7-47.1), respectively also experienced stress.

For the data on burnout, nurses population (80.2%; 95%CI = 56.8-103.7) remained at the top position in terms of experiencing burnout, followed by the doctors population (74.9%; 95%CI = 62.8-87.1) and finally by allied healthcare personals (64.9%; 95%CI = 49.5-80.2).

For the prevalence of fear, both genders were deemed to have almost similar prevalence with females (99.7%; 95%CI = 98.7–100.6) and males (99.8%; 95%CI = 99.2–100.4). Doctors (99.9%; 95%CI = 99.5–100.2) were having the highest prevalence of fear while nurses had the lowest prevalence (61.1%; 95%CI = 28.0-94.1). Besides, females (17.0%; 95%CI = 9.2-24.9) reported a higher prevalence of having low resilience as compared to males (12.5%; 95% CI = 9.3-22.4).

Subgroup analysis for risk factors associated with psychological distress among healthcare providers amidst COVID-19 pandemic

Table 4 showed the subgroup analysis of the odds ratio of psychological distress according to the risk factors. From this meta-analysis, it was found that odds of depression were increased in females (OR = 1.48; 95%CI = 1.30-1.68) and those who worked as nurses, assistant nurses or midwives (OR = 1.21; 95% CI = 1.02-1.45). Those who worked as doctors, dentist, allied healthcare or pharmacist were found to be the protected against depression (p-value <0.001).

In terms of the risk of getting anxiety, females (OR = 1.66; 95%CI = 1.49–1.85) and nurses, assistant nurse or midwife (OR = 1.36; 95%CI = 1.16–1.58) had a higher risk than doctors, dentist (OR = 0.82; 95%CI = 0.73–0.93), allied healthcare personals and pharmacists (OR = 0.89; 95%CI = 0.74–1.06), with a p-value of <0.001. Besides, females were also a risk factor for the development of stress (OR = 1.59; 95%CI = 1.28–1.97).

Sensitivity analysis and publication bias

We performed sensitivity analysis by omitting every single study step-by step from the metaanalytic model. The result reported no major changes in terms of the pooled prevalence of psychological distress (Fig A6.1-A6.6 in S6 Fig). The visual assessment of the funnel plot for all the psychological distress parameters showed a high publication bias (Fig A7.1-A7.5 in S7 Fig), which was confirmed by Egger's test for depression and anxiety.

Discussion

To date, this is the first systematic review with meta-analysis on psychological distress among HCPs amidst the COVID-19 pandemic in Asian region with country-based estimates. We identified 148 cross-sectional studies from 23 Asia countries and quantitatively determined the subgroup pooled prevalence according to gender and occupations in this region. The pooled prevalence for depression was 37.5% (95%CI: 33.8–41.3), anxiety 39.7(95%CI: 34.3–45.1), stress 36.4% (95%CI: 23.2–49.7), fear 71.3% (95%CI: 54.6–88.0), burnout 68.3% (95%CI: 54.0–82.5), and low resilience was 16.1% (95%CI: 12.8–19.4), respectively.

Among all the psychological distress, fear appeared to be the most common psychological reaction among HCPs whom continued to provide healthcare services during the COVID-19 pandemic, followed by burnout, anxiety, depression and finally by stress. More than two thirds of the HCPs were having fear amidst the COVID-19 pandemic. Our findings is similar to the findings of the previous review in Asia which the prevalence of fear among HCPs ranged between 67% [166] to 77.1% [13]. There are many reason for HCW's fear in this pandemic,

Subgroup analysis	N	Exposure in at risk group	Total at risk group	Exposure in control	Total control	OR (95%CI)	I ² (p-value)	Appendix
Depression								
Gender: Female	29	7432	17595	3789	10556	1.48 (1.30– 1.68)	68.10 (<0.001)	Fig A1.18 in (SI Fig)
Occupation Doctor/ Dentist	22	2889	7329	3334	9109	0.87(0.69– 1.10)	86.47 (<0.001)	Fig A1.19 in (<u>S1</u> Fig)
Nurse/Assistant nurse/midwife	24	2716	7142	3791	9812	1.21 (1.02– 1.45)	79.11 (<0.001)	Fig A1.20 in (<u>S1</u> Fig)
Allied healthcare personal/ pharmacist	16	902	2456	3612	8764	0.93 (0.69– 1.25)	84.75 (<0.001)	Fig A1.21 in (<u>S1</u> Fig)
Anxiety								
Female	33	9319	19215	5288	12434	1.66(1.49– 1.85)	61.68(<0.001)	Fig A2.20 in (<u>S2</u> Fig)
Doctor/ Dentist	23	3546	8474	6135	12404	0.82(0.73- 0.93)	62.85(<0.001)	Fig A2.21 in (<u>S2</u> Fig)
Nurse/Assistant nurse/midwife	24	4002	8561	5690	12456	1.36(1.16– 1.58)	75.58(<0.001)	Fig A2.22 in (<u>S2</u> Fig)
Allied healthcare personal /pharmacist	16	2144	3955	5642	11328	0.89(0.74– 1.06)	68.78(<0.001)	Fig A2.23 in (S2 Fig)
Stress								
Female	14	1752	9812	957	6393	1.59 (1.28– 1.97)	56.14 (0.005)	Fig A3.14 in (S3 Fig)
Doctor/ Dentist	6	481	1571	817	2668	0.80 (0.43– 1.49)	87.67 (<0.001)	Fig A3.15 in (S3 Fig)
Nurse/Assistant nurse/midwife	6	704	2325	594	1914	1.47 (0.80– 2.70)	87.02 (<0.001)	Fig A3.16 in (<u>S3</u> Fig)
Allied healthcare personal /pharmacist	4	113	343	562	1343	0.85 (0.61– 1.17)	20.27 (0.288)	Fig A3.17 in (<u>S3</u> Fig)
Burnout								
Female	0							
Doctor/ Dentist	1	800	1122	881	1289	1.15(0.97– 1.37)	Not sig	
Nurse/Assistant nurse/midwife	2	1012	1431	853	1196	1.93(0.37– 10.12)	93.96(<0.001)	Fig A4.4 in (<u>S4</u> Fig)
Allied healthcare personal /pharmacist	1	24	37	160	179	0.22(0.10- 0.50)	Significant- NA	
Fear								
Female	1	147	147	223	223	0.66 (0.01– 33.44)	Not sig	
Doctor/ Dentist	0							
Nurse/Assistant nurse/midwife	0							
Allied healthcare personal /pharmacist	0							
Low resilience								
Female	1	15	88	4	32	1.44(0.44– 4.71)	NA	
Doctor/ Dentist	0							
Nurse/Assistant nurse/midwife	0		1				1	
Allied healthcare personal /pharmacist	0							

Table 4. Subgroup analysis of odd ratio of psychological distress among HCPs according to gender and occupations.

one of the most common issues face by HCPs are there are fear of failing to provide adequate care for patients, non-limited to only COVID-19 due to limited supply of resources as well as manpower to handle the frequent sudden rise in COVID-19 cases that often strain the health-care facility capacity [167]. Another possible explanation for the high prevalence of fear among HCPs were due to the fear of carrying the virus back home and infecting family and friends as well as fear of stigmatization [168]. It is also could be due to the fear of being infected and need to be quarantined, thereby further exacerbating the pre-existing inadequate numbers of HCWs at the frontline combating COVID-19 [169]. Another reason for fear among HCPs was because they were lacking in Personal Protection Equipment (PPV) rations and had unfamiliarity in using PPV, especially at the beginning of the pandemic. They may have had no training for infection prevention and control protocols especially at the beginning of the pandemic. All these factors may end up with many HCPs succumbing to this virus while providing care to the patients [170, 171].

Burnout was the second most common psychological distress faced by HCPs. The possible explanation could be due to the irregular or long working hours and high workload demand [1]. Furthermore, HCWs need to adapt to the IPC strategies by putting on PPE before starting to work and showering before going home. All this required additional time in preparation and cleaning and can cause fatigue in the long run [172]. This is not surprising on why another systematic review by de Pablo et al., 2020 reported that only 34.4%(95%CI = 19.3-53.5%) of the HCPs suffered from burnout which is lower compared with our finding of 68.3% [172]. The possible explanation for the difference in prevalence of burnout could be due to the fact that the study by de Pablo et al., 2020 examined burnout among HCWs exposed to SARS/ MERS/COVID-19 whereas our systematic focused mainly on COVID-19. Despite SARS/ MERS/COVID-19 are all caused by different strains of coronavirus, however there is an apparent difference in the influence on human imposed by these different coronaviruses' strains. For instance, the mortality rate of COVID-19 is 4.9%, which is higher than SARS (0.96%) and much lower than MERS (34.4%). The duration of the SARS pandemic (cumulative of 8,422 cases) was relative shorter which was brought under control in only 9 months (1 November 2002-31 July 2003) and outbreak of MERS only lasted for two months in both Saudi Arabia (n = 402 cases) and South Korea (n = 150 cases). However, COVID-19 has been around us for more than one year now and the emergence of new variants that are still evolving and its effect largely remains unpredictable, adds to its own pandora box. In view of the influence exerted by COVID-19 is much more devastating than SARS and MERS, therefore HCWs suffer burnout more easily in handing COVID-19 cases than in handling cases of SARS and MERS.

Our review showed that about one third of the HCPs suffered from depression with pooled prevalence of 37.5%, anxiety (39.7%) and stress (36.4%). However, our pooled prevalence of depression was higher comparing with two other systematic reviews by Hossain MM et al and Gonzalo Salazar de Pablo et al where the pooled prevalence of depression ranges from 17.9%-29.9% and anxiety, 22.2% to 43.6% [12, 171]. The pooled prevalence of anxiety in our review fell in between of these two reviews which ranged from 22.2 to 43.6% [12, 171].

Among the 148 papers reviewed, 98 of them examined the prevalence of depression among HCP, with China reporting most number of cases (22,772) of depression since the outbreak of COVID-19. Prevalence of this parameter in China was found to be 36.5%. Prevalence of depression in Malaysian HCP was relatively lower at 26.6% (95%CI: 17.9–35.3; p-value<0.001). The possible reason could be due to the fact that these cross-sectional studies were conducted at the very beginning stage of the outbreak in China before the pandemic was declared, where very little was known about the virus and hence the Chinese healthcare workers were generally experiencing greater mental disturbances.

Jordan, Egypt from the subdivision of Asian continent, Arab Saudi and Iran were the countries with the prevalence of depression and, anxiety of more than 50% when compared with other countries in Asia. The possible explanation could be due to the fact that most of the research were conducted between March to July 2021 at the time that Middle East countries were having the COVID-19crisis. Most of the countries (Jordan, Egypt, Arab Saudi and Iran) had full lockdown or night-time curfew. The reason for the rapid rise in cases were largely due to large religious gatherings, wedding celebrations and other social events where control measures were not sufficiently enforced [173]. The possible reasons were that there was a lack ofhealthcare facilities and equipment to deal with COVID-19 pandemic [174]. Furthermore, the human resources were also insufficient and below the recommendation of WHO as some of HCPs have left the countries following the country's own politic instability [175, 176]. With a weakened healthcare system, the COVID-19outbreak had posed a major challenge on the mental health of HCPs and it explained why the prevalence of depression and anxiety were generally higher as compared with other countries in Asia [177].

On the other hand, we also found that the prevalence of anxiety among healthcare workers in China, India, and Malaysia were 31.9% (95%CI: 27.8–36.0), 44.2% (95%CI: 32.6–55.9) and 30.2% (95%CI: 27.9–32.6), respectively. The lower prevalence of both anxiety and depression in Malaysia as compared with other countries could be due to the studies were conducted during Conditional Movement Control Order (CMCO) period when the condition of outbreak in Malaysia was considered to be relatively under-control, and background of the causative virus had been learnt from China. Besides, the fact that the data was not merely focusing on COVID-19 hospitals but also involving non-COVID centres could have led to the lower anxiety prevalence in Malaysia [140, 142].

With the COVID-19 pandemic hitting many health care facilities that were unprepared to handle it, many healthcare providers who were standing at the frontline were working and pushing themselves to the limit [28]. Pooled prevalence of burnout was found to be 68.3% (95% CI: 54.0–82.5) in our systematic review and meta-analysis, ranging from the lowest prevalence of 58.0% in China to the highest prevalence of 90.4% reported in Korea. Korea has the highest prevalence of burnout as the study was conducted specifically among the the Infectious Disease physicians. This is not unexpected as their work burden is much higher during the COVID-19 pandemic [160].

A cross-sectional study conducted by Dong et al in China revealed that despite the long working hours, healthcare workers were mostly (n = 4,120, 89.2%) motivated and feeling positive towards their task at hand, and remained committed to their professions. During the outbreak period in China around early 2020, Chinese nationals from all over the country, with or without medical background, showed exemplary courage and actively volunteered to assist at Wuhan, the epitome of the COVID-19 pandemic [178, 179]. There was also an increment in healthcare providers' salaries by the Chinese government at the same time. Their firm belief to their professions and strong social support from their nation were believed to be factors contributing to the lowest burnout prevalence among Asian countries [28]. Whereas, in Malaysia, the high prevalence of burnout among nurses could be due to the reduction in their off days as a consequences of more intense shift hours, and being overworked, coupled with a low salary [142].

When analysing the subgroup of associated risk factors of psychological distress, it was found that both females and nurses population were more at risk of getting mental distress such as depression [(Female: OR = 1.48; 95% CI = 1.30–1.68), (Nurse: OR = 1.21; 95% CI = 1.02–1.45)], anxiety [(Female: OR = 1.66; 95% CI = 1.49–1.85), (Nurse: OR = 1.36; 95% CI = 1.16–1.58)], and stress [(Female: OR = 1.59; 95% CI = 1.28–1.97), (Nurse: OR = 1.47; 95% CI = 0.80–2.70)]. Internally, females' nature generally belongs to the sentimental type and they

usually experience hormonal changes which would then affect their mood and emotion [180]. Other than playing a role as a medical professional, most of the time females were also housewives for their family [8]. They tend to be a multitasker where they must take care of their family members' health and well-being as well as going out for marketing in crowded areas to purchase necessities. A lot of time they were lacking support from family and were bombarded with many negative news circulated on social media, which can create negative feelings and make them more tired and prone to psychological illness [181–183]. Besides, it was found that the majority of the nurses were females (95.6%) [184]. Furthermore, nurses generally had longer contact hours with COVID-19 patients than doctors and were working longer hours than usual [36]. Other than taking care of the patients, they had to deal with their families who might be more frustrated, angry, anxious or worried due to lack in family time [142]. These were consistent with the findings from a systematic review by Thatrimontrichai et al which concluded that females, nurses, having direct contact with infected patients, working longer hours and possessing less working experience were the main risk factors leading to mental disturbances among Chinese healthcare workers [13].

High heterogeneity was found in this systematic review, and possible reasons include variation from the participants' characteristics, outcome level and research setting [185]. Firstly, the high heterogeneity for the overall prevalence of psychological distress found in this study could be due to differences in the screening methods and diagnostic criteria in different countries with different ethnicities and research settings (S4 Table). For example, different screening tools were used to determine burnout such as Maslach Burnout Inventory (MBI) ProQOL Scale of Chinese Version and Olenburg Burnout Inventory (OLBI); whereas scales like Patient Health Questionnaaire (PHQ), Depression-Anxiety-Stress Scale-21 (DASS-21), Beck Depression Inventory (BDI), Hamilton depression rating scale (HAMD), etc were used for depression screening. Next, different categories for the severity of disease were reported in different studies. Moreover, even though in the same country, there were different diagnostic criteria applied for psychological distress studies. For example, in China, ten different diagnostic tools were used to detect depression compared with three diagnostic tools used in India, which give rise to a broad range of prevalence of depression in China (31.7%-41.2%) and India (27.9%-39.3%). In our meta-analysis, there was a wide gap in terms of sample size in all 148 studies, ranged from 100 to 14,825 in this review. All these factors would explain why there was a high heterogeneity for the prevalence of psychological distress, for instance, depression in this study.

Multiple factors contributed to publication bias, comprises of study design, sample size, decision of authors, journal editors and reviewers [186]. We had excluded all the unpublished data and studies with a sample size of below 100 in this review. However, we need to weigh the advantage and disadvantages as those published studies have gone through a rigorous review process, which gave a more reliable result compared with those unpublished data. Furthermore, we also excluded studies with outcomes that did not fulfil our operational definition of psychological distress. This systematic only included manuscripts wrote in English due to constraint of resources. Thus, we had to interpret the results of this systematic review carefully within the context of its limitations level and research setting [185].

Strengths and limitations

This review paper highlights the psychological distress of HCPs in the menacing era of the COVID-19 pandemic. Such HCW's mental burden either have not been acknowledged or have been underestimated because most of the healthcare systems have currently focused on coping with the pandemic as its main target. Foremost, psychological distress among HCPs

should be given priority and urgent action is needed to reduce the psychological impacts on HCPs, in order to ensure continuation of effective services to patients amidst the COVID-19 pandemic.

There are some limitations in this review paper. Firstly,our review paper depended on collecting and compiling the published data where the papers that were analysed were mostly periodic, in which only the psychological state of HCPs over a certain period of time were reflected. However, with the progression of time, different conditions of outbreak over the period of one year and shifting to new environments, the targeted population's mental health may have changed. Thus, the psychological impact among healthcare providers amidst the COVID-19 pandemic should ideally be assessed longitudinally. Secondly, high heterogeneity was not an unexpected finding in our review as data were gathered from various studies that were conducted differently in terms of study designs, data collection tools, different study setting and location as well as having varied demographic features of participants. Therefore, our results needs careful interpretation. Thirdly, only English language written articles were recruited in this review, therefore this can result in publication bias. Future studies on interventions to improve the psychological health of HCPs is needed urgently in order to maintain their physical health and productivity in continuing the fight against this pandemic.

Conclusions

In conclusion, the global COVID-19 pandemic has had a devastating impact on the mental health of HCPs. This systematic review synthesizes the quantitative evidence of psychological distress among HCPs in Asian countries, which showed that one third of HCPs suffered from depression, anxiety and stress and more than two third of HCPs suffered from fear and burnout during the COVID-19 pandemic in Asia. Meta-analysis reported both females and nursse were at increased risk of having depression and anxiety. Female HCPs was also at a higher risk of getting stress when compared with the male HCPs. Urgent action are needed to implement a multicultural level interventions to support HCPs in order to reduce the burden of psychological distress during this very challenging COVID-19 pandemic.

Supporting information

S1 Fig. Forest plots of depression. (RAR)
S2 Fig. Forest plots of anxiety. (RAR)
S3 Fig. Forest plots of stress. (RAR)
S4 Fig. Forest plots of burnout. (RAR)
S5 Fig. Forest plots of fear. (RAR)
S6 Fig. Leave-out-one Forest plots. (RAR)
S7 Fig. Funnel plots. (RAR) S1 Table. PRISMA checklist. (DOCX)
S2 Table. Search terms used from 13th of March to 15th of March 2021. (DOCX)
S3 Table. Strobe checklist. (DOCX)
S4 Table. Characteristics of 148 studies. (DOCX)

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