Effect of smartphone app on post-traumatic stress disorder in COVID-19 convalescent patients
A protocol for systematic review and meta-analysis
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Abstract
Background: The outbreak of Coronavirus Disease 2019 (COVID-19) seriously affects humans’ health worldwide physically and mentally. Studies revealed that the prevalence of post-traumatic stress disorder (PTSD) increased under this condition. PTSD can change the structure of patients’ central nervous system, and increase the risk of anxiety or depression, thus greatly affecting the quality of patients’ life and their families. PTSD is preventable, and the effects of early prevention are better. Non-drug intervention can prevent or reduce the psychological sequelae after hospitalization, help patients understand the experience during hospitalization, and be beneficial to their psychological rehabilitation. Whether smartphone app based intervention can be an alternative therapy for PTSD in terms of COVID-19 convalescent patients is still controversial. Therefore, we conducted a meta-analysis and systematic review to evaluate the effects of smartphone app based intervention on PTSD in COVID-19 convalescent patients, so as to provide some guidance for clinical application.

Methods: The literatures that are related to the smartphone app based intervention and PTSD in COVID-19 convalescent patients from inception to February 2021 will be searched. The following databases are our focused areas: ClinicalTrials.gov, Cochrane Central Register of Controlled Trials repositories, PubMed, Embase, and Web of Science databases. According to the inclusion and exclusion criteria, 2 investigators would independently screen the literature extract data and evaluate the risk of bias in the included studies. Meta-analysis was performed with RevMan5.3 software.

Results: The results of this meta-analysis will be submitted to a peer-reviewed journal for publication.

Conclusion: The conclusion of our study could provide evidence for the judgment of whether smartphone app based intervention is an effective intervention on PTSD in COVID-19 convalescent patients.

PROSPERO registration number: CRD42021240340.

Abbreviations: COVID-19 = Coronavirus Disease 2019, PTSD = post-traumatic stress disorder, RCTs = randomized controlled trials, SARS-CoV-2 = Severe Acute Respiratory Syndrome Coronavirus-2.

Keywords: Coronavirus Disease 2019, post-traumatic stress disorder, smartphone app, systematic review

1. Introduction
Coronavirus Disease 2019 (COVID-19) is a global pandemic caused by the Severe Acute Respiratory Syndrome Coronavirus-2 (SARS-CoV-2). [1-3] It has become a key issue and has seriously threatened the public health around the world. [4] The outbreak of this infectious disease has affected people’s physical health and mental health, including the infected and not infected. Post-traumatic stress disorder (PTSD) and PTSD-related symptoms are prevalent and disabling conditions occur as a consequence of traumatic events. [5] PTSD refers to a psychological disorder and it occurs late and persists irreversible once it occurs. [6] The lifetime prevalence rate of PTSD is about 4% that is often irreversible once it occurs. [7] People with PTSD may be accompanied by persistent negative emotions, including anger, guilt, fear, or shame. [9] PTSD can result in structural changes in patients’ central nervous system, and increase the risk of anxiety or depression, thus seriously affecting the quality of patients’ life. [10] If it is not intervened, it will seriously disrupt patients’ daily life. [11]
A number of studies have pointed out that PTSD is preventable, and the effects of early prevention are satisfac-
tory.[12–14] The latest PTSD treatment guidelines recommend that patients should receive psychotherapy in the outpatient
first.[15,16] However, many factors limit patients’ access to
treatment, including the limited number of qualified therapists
and the high cost of psychotherapy.[16] In addition, patients are
often afraid of discrimination, so they are reluctant to take the
initiative to visit a doctor. The use of smartphones is more
popular in both developed and developing countries.[17]
Smartphone app can provide patients with flexible and
convenient treatment, and may also make these patients who
have no intention of treatment be willing to receive timely
treatment.[18]

By 2017, more than 300,000 health-related apps were
available to consumers, of which 490 were for mental health
and behavioral disorders.[19] Therefore, mobile app may have
potential advantages in the improvement of PTSD.

At present, there is no consistent conclusion on whether
smartphone app based intervention can reduce the incidence
of PTSD in the convalescent stage of COVID-19. Considering
the low-cost and easy-to-accept intervention method, smartphone
app based intervention has application value in the
field of psychological reconstruction during the rehabilitation
of COVID-19. Therefore, this study conducted a meta-analysis
of the literature on the intervention effects of smartphone app based
intervention on PTSD in convalescent patients with COVID-19 to
provide objective basis for the intervention of PTSD.

2. Methods

2.1. Study registration

Our protocol has been registered on the PROSPERO. The
registration number is CRD42021240340. We strictly abide by
the Preferred Reporting Items for Systematic Review and Meta-
Analysis Protocols (PRISMA-P) guidelines.

2.2. Inclusion and exclusion criteria for study selection

2.2.1. Inclusion criteria. The inclusion criteria were randomized
controlled trials (RCTs), with smartphone app based intervention
as the main form of intervention.

2.2.2. Exclusion criteria.

1. Repeated publication;
2. Incomplete literature;
3. Nonrandomized controlled trials.

2.3. Types of participants

COVID-19 patients who have been clearly diagnosed and now in
recovery period, regardless of sex, age, race, or educational and
economic status, will be enrolled in the review. Participants must
be between 18 and 80 years old.

2.4. Interventions and controls

The intervention in the experimental group was based on mobile
phone app, and the control was multiple control measures,
including blank, placebo, usual or standard care, health
education, psychosocial therapy, and drug therapy.

2.5. Types of outcome measures

PTSD score after intervention.

2.6. Search strategy

The following electronic databases will be searched from
inception to February 2021: ClinicalTrials.gov, Cochrane
Central Register of Controlled Trials repositories, PubMed,
EmBase, and Web of Science databases. As for other sources, we
also plan to manually search for the unpublished conference
articles and the bibliography of established publications. The
search terms on PubMed are as follows: “smartphone∗,” or
“app,” and “computer∗” or “phone∗”; COVID-19 (e.g.,
“Corona Virus Disease 2019” or “Corona Virus”); RCT
(“randomized” or “randomly” or “clinical trial”). The combi-
nation of Medical Subject Headings (MeSH) and text words will
be applied. These search terms are summarized in Table 1.

| Number | Search terms |
|--------|--------------|
| #1 | Web [Title/Abstract] |
| #2 | Online [Title/Abstract] |
| #3 | Internet [Title/Abstract] |
| #4 | App [Title/Abstract] |
| #5 | Application [Title/Abstract] |
| #6 | Computer [Title/Abstract] |
| #7 | E-mail [Title/Abstract] |
| #8 | Phone [Title/Abstract] |
| #9 | Smartphone [Title/Abstract] |
| #10 | OR/1–9 |
| #11 | Stress Disorders, Post-Traumatic [MeSH] |
| #12 | Neuroses, Post-Traumatic [Title/Abstract] |
| #13 | PTSD [Title/Abstract] |
| #14 | Post-Traumatic Stress Disorders [Title/Abstract] |
| #15 | Acute Post-Traumatic Stress Disorder [Title/Abstract] |
| #16 | Chronic Post-Traumatic Stress Disorder [Title/Abstract] |
| #17 | Delayed Onset Post-Traumatic Stress Disorder [Title/Abstract] |
| #18 | Neurones, Posttraumatic [Title/Abstract] |
| #19 | Posttraumatic Stress Disorders [Title/Abstract] |
| #20 | Stress Disorder, Post Traumatic [Title/Abstract] |
| #21 | Stress Disorders, Posttraumatic [Title/Abstract] |
| #22 | Acute Post Traumatic Stress Disorder [Title/Abstract] |
| #23 | Chronic Post Traumatic Stress Disorder [Title/Abstract] |
| #24 | Delayed Onset Post Traumatic Stress Disorder [Title/Abstract] |
| #25 | Neurones, Post Traumatic [Title/Abstract] |
| #26 | Post Traumatic Stress Disorders [Title/Abstract] |
| #27 | Post-Traumatic Neuroses [Title/Abstract] |
| #28 | Post-Traumatic Stress Disorder [Title/Abstract] |
| #29 | Posttraumatic Neuroses [Title/Abstract] |
| #30 | Posttraumatic Stress Disorder [Title/Abstract] |
| #31 | Stress Disorder, Post-Traumatic [Title/Abstract] |
| #32 | Stress Disorder, Posttraumatic [Title/Abstract] |
| #33 | Stress Disorders, Post Traumatic [Title/Abstract] |
| #34 | OR/11–33 |
| #35 | Corona Virus [Title/Abstract] |
| #36 | Corona Virus Disease 2019 [Title/Abstract] |
| #37 | COVID-19 [Title/Abstract] |
| #38 | Novel coronavirus [Title/Abstract] |
| #39 | Novel coronavirus pneumonia [Title/Abstract] |
| #40 | OR/35–39 |
| #41 | Random* [Title/Abstract] |
| #42 | Clinical trial [Title/Abstract] |
| #43 | OR/41–42 |
| #44 | #10 AND #34 AND #40 AND #43 |
2.7. Data collection and analysis

2.7.1. Studies selection and data extraction. The details of the selection process will be displayed in the PRISMA flow chart (Fig. 1). First of all, the 2 reviewers will independently check the titles and abstracts of the search results, and initially sift through the articles. After reading the full text of the preliminary selected articles, the 2 independent reviewers will select qualified studies on the basis of our predetermined inclusion criteria. Finally, the

![Flow diagram of study selection process.](Figure 1)
articles that are selected by the 2 independent commentators will be integrated after putting forward repetitive parts. When the 2 independent reviewers disagree with each other, a third reviewer will make the decision. Extracted contents include author, publication date, country, sample size, age, type, content and the duration of smartphone app based intervention, PTSD scale, result measurement data, intervention time, and other details.

2.7.2. Assessment of risk of bias in included studies. The risk of bias assessment method set out in the Cochrane Handbook for Systematic Reviews of Interventions will be adopted on the included RCTs.

2.7.3. Measure of treatment effect. Standardized mean difference is applied to measure the efficacy of 95% confidence interval.

2.7.4. Dealing with missing data. The accuracy of the data may affect the results of the study, and even lead to different conclusions. Therefore, we will contact the corresponding authors to supplement the research results with insufficient or missing data. If the corresponding author cannot be contacted, the data will be processed separately, and the potential impacts of incomplete data on the results will be analyzed and reported as well.

2.7.5. Assessment of heterogeneity. The heterogeneity assessment will be conducted by the Review Manager 5.3. According to the Cochrane Handbook, Chi-Squared test and I^2 value could be used to evaluate the heterogeneity. If the I^2 value is less than 50%, it represents that the relative heterogeneity is small, and the fixed effect model should be adopted. Otherwise, the random effect model will be used.

2.7.6. Assessment of reporting biases. If enough studies are available, we will evaluate the funnel plot to report bias.\[^{20}\]

2.7.7. Data synthesis. The Review Manager 5.3 software will be used for the quantitative data analysis, including mapping overall forest plot, heterogeneity analysis, and subgroup analysis. If the I^2 value is less than 50%, it represents that the relative heterogeneity is small, and the fixed effect model should be adopted. Otherwise, the random effect model will be used.

2.7.8. Subgroup analysis. Considering the possibility of a high degree of heterogeneity, we will perform a subgroup analysis, if necessary, to explain the underlying causes of heterogeneity. According to the course of intervention and the content of app intervention, subgroup analysis was carried out.

2.7.9. Sensitivity analysis. Sensitivity analysis will be conducted to assess the robustness of the results. After the data are synthesized, we will exclude the merged studies one by one for sensitivity analysis to see if there exist any significant changes in the comprehensive results.

2.7.10. Quality of evidence. The evidence evaluation of all results will be summarized by the suggested assessment, development and assessment (GRADE) method.\[^{21}\] The level of evidence will be divided into high, moderate, low, and very low quality.

2.7.11. Ethical review and informed consent of patients. The content of this article does not involve moral approval or ethical review and will be presented in print or at relevant conferences.

3. Discussion

The occurrence of PTSD will seriously reduce the life quality of COVID-19 survivors and influence their physical and mental recovery.\[^{22–24}\] Therefore, it is of great significance to prevent the occurrence of PTSD in survivors of COVID-19. Recently, there has been the emergence of smartphone app based intervention that is increasingly used to treat PTSD. More and more evidence shows that, smartphone app based intervention has positive effects on the life quality, stress relief, and health improvement of patients with PTSD.\[^{25–29}\] Previous studies have suggested that the intervention of mental illness based on smartphone app is effective.\[^{30,31}\] Therefore, mobile app may have potential advantages in terms of improving PTSD.

Due to the differences in sample, time, frequency, method, and duration of app intervention, the specific effects of PTSD symptoms may be different. However, it is still necessary to carry out studies with large sample size, and well-designed and long-term follow-up in the future. In addition, future researches should set up a monitoring module in app to objectively record the app login frequency and the duration of the research object, so as to make the research results more accurate, and further explore the impacts of intervention measures based on mobile phone app on PTSD.

4. Conclusion

As far as we know, this is the first systematic review and meta-analysis. The findings of this study will prove the effectiveness of smartphone applications in the intervention of PTSD in convalescent COVID-19 patients.

Author contributions

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Software operating: Yunyan Wang.
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Validation: Xia Yang, Haiyan Chen.
Visualization: Haiyan Chen.
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