



Epidemiological and clinical characteristics of vaccinated COVID-19 patients: A meta-analysis and systematic review

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Abstract

Objective: With the global epidemic of coronavirus disease 2019 (COVID-19), vaccination rates are increasing globally. This study evaluated the relevant clinical manifestations of vaccinated COVID-19 patients. **Methods:** We searched carefully in 11 databases such as PubMed, Embase, Scopus, Cochrane Library, Web of Science, Ovid, China National Knowledge Infrastructure Database, Wan Fang Data, Sinomed, VIP Database, and Reading Showing Database up to 26 March 2022. To search for articles that have described the characteristics of vaccinated patients including epidemiological and clinical symptoms. Statistical analysis of the extracted data using STATA 14.0. **Results:** A total of 58 articles and 263,708 laboratory-confirmed COVID-19 patients were included. Most of the patients in the vaccinated group had more asymptomatic infection and fewer severe illnesses. There were significant differences in ethnicity, and strain infected with COVID-19, and comorbidities (hyperlipidemia, diabetes, obesity, kidney disease, immunocompromised, cardiovascular disease, and tumor) and symptoms (fever, cough, gastrointestinal symptoms, neurological symptoms, and dysgeusia/anosmia) between vaccinated group and unvaccinated group. Oxygen support, use of steroid, days in hospital, hospital treatment, ICU treatment, death, and poor prognosis were also significantly different. **Conclusion:** Compared with the vaccinated group, patients in the unvaccinated group had a more severe clinical manifestations. Vaccines are also protective for infected people.

Keywords

COVID-19, vaccine, epidemiological characteristics, clinical characteristics, meta-analysis

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Introduction

Coronavirus disease 2019 (COVID-19) first appeared in China in 2019 and was defined as a global pandemic in March 2020, which was a major global public health problem. As of 24 August 2022, there have been 595,219,966 confirmed cases of COVID-19, including 6,453,458 deaths.¹ In response to the spread of COVID-19, vaccines are rapidly developed and administered. The World Health Organization also has approved 7 vaccines for emergency use.² Various types of vaccines including mRNA vaccines, viral vector, inactivated vaccine, protein

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subunit, and DNA vaccines are being developed to control the spread of COVID-19.³ In December 2020, the mass vaccination program was launched,⁴ and as of 1 August, 2022, a total of 12,308,330,588 doses of vaccine have been administered worldwide.

Vaccination was significantly effective in reducing viral infections as well as hospitalizations and deaths.⁵⁻⁷ But with the emergence of variants of concern (VOCs), viruses with multiple clusters of mutations in the genome with higher infection rates, vaccines cannot play a 100% preventive effect⁸ and there are many patients infected with COVID-19 after being vaccinated. "Breakthrough infection" was defined as a patient who was fully vaccinated with two doses of the vaccine for more than 14 days.⁹ There are still some people who are hesitant to receive the COVID-19 vaccine, especially those living in developing countries.¹⁰ Residents' misunderstanding and distrust of vaccines may also lead to blind anti-vaccine movements.¹¹

Unfortunately, it is not known whether VOCs can identify antibodies accurately, and whether mutations can evade vaccine-induced immunity and reduce vaccine activity is also of concern. According to some studies, vaccinated patients have lower risk factors for hospitalization, infectious diseases, and death.^{12,13} However, other studies have indicated that VOCs can evade antibody responses¹⁴ and there is a similar Cycle threshold (Ct) value¹⁵ between infected individuals who have not been vaccinated and have been vaccinated.

We conducted this study to assess whether vaccines were protective in COVID-19 patients. We would conduct this research by evaluating the characteristics of vaccinated COVID-19 patients from the epidemiological history, clinical symptoms, treatments, and prognosis of the patients.

Methods

This is a systematic review and meta-analysis following the international prospective register of systematic reviews. We obtained the registration number which was CRD42021293918. Follow the principles of PICOS (Population, Intervention, Comparison, Outcome, Study type). Populations were confirmed COVID-19 patients; intervention means patients had been vaccinated before COVID-19 infection; comparison of general data and clinical data between vaccinated and unvaccinated patients; outcome was to explore whether clinical data and general data were statistically different between two groups; and study types were case-control studies and cohort studies.

Search strategy

We had made a complete search plan and fully searched 11 databases including PubMed, Embase, Scopus, Cochrane

Library, Web of Science, Ovid, China National Knowledge Infrastructure Database, Wan Fang Data, Sinomed, VIP Database, and Reading Showing Database from January 1, 2020 to March 26, 2022. We conducted a comprehensive search for relevant research in relevant fields, not specific countries, regions, or continents. We searched by using "COVID-19," "COVID-19 Virus," "SARS-CoV-2," "coronavirus disease 2019," "2019-nCoV," "severe acute respiratory syndrome coronavirus 2 infection," "Wuhan seafood market pneumonia virus," "vaccine," "vaccines," "vaccination," "breakthrough infections," and "breakthrough." The flow chart of the process and results of literature retrieval was shown in [Figure 1](#).

Study selection

The literature included in this systematic review and meta-analysis included studies with the following characteristics. (1) Patients need to be clearly diagnosed of COVID-19 such as COVID-19 RNA was detected by PCR, isolate the virus or IGM/IGG antibodies from the patient's serum; (2) the study must be a case-control or cohort study; (3) the study must be including the vaccinated and unvaccinated COVID-19 patients; (4) the patient's clinical characteristics including gender, age, symptoms, treatments, prognosis, and other relevant information should be exhibited.

Exclusion criteria were the following items: (1) duplicate literatures, (2) non-Chinese or non-English researches, (3) lack of data or failing to extract data, and (4) meta-analysis and reviews.

Two review authors (Tian and Ren) independently selected articles for initial screening by reading the title and abstract of each article retrieved and then determined whether the full text should be read carefully according to inclusion or exclusion criteria. The final inclusion or exclusion of studies was discussed in consultation between two reviewers after reading the full text. If there is any disagreement, it will be discussed with the third reviewer.

Data extraction

We would extract the following information from the included studies including authors, publication information, sample size, gender, age, comorbidities such as diabetes, obesity, liver and kidney disease, clinical treatment such as antiviral drugs, steroid and supplementary oxygen required, prognosis, and clinical outcomes. It would be discussed with a third author when data extraction was in dispute.

Quality assessment

Each study we included would be assessed for quality by two reviewers using the Newcastle-Ottawa Scale (NOS).¹⁶

Meta-analysis

Statistical analysis was performed using Stata14.0. We assessed the heterogeneity of each study by using Cochran's Q test and I^2 test, if p -value > 0.1 and $I^2 < 50\%$, using a fixed-effects model, otherwise a random-effects model would be used. The possible sources of heterogeneity would be explored through subgroup analysis. A funnel plot was used to evaluate the publication bias of these studies.

Results

Literature search, basic information, and quality assessments

A total of 13,818 literatures were searched. After removing duplicate and did not meet the inclusion criteria literatures and reading 296 full-text articles, 58 literatures were included finally. The detail of the study was shown in Figure 1. The basic characteristics of each study were shown in Table 1. We used the NOS scale to evaluate the

quality of each included study, and all of the studies were rated 6 stars or above.

Gender

A total of 44 articles were included in the study of gender. After the heterogeneity test, we found that $I^2 = 37.4\%$ and $p < 0.1$, indicating that there was low heterogeneity, therefore a random-effect model was used. We found that OR = 0.995 (95% CI: 0.942–1.05, $p = 0.849$, Figure S1(a)), showing no statistical difference in gender between vaccinated and unvaccinated patients. Through subgroup analysis, we found heterogeneity of studies derived from studies published in 2022 (Figure S1(b)). The funnel plot of this study was symmetric by drawing a funnel plot (Figure S1(c)) and Egger's test ($p = 0.47$).

Age

A total of 14 studies were included to analyze the effect of age. We found no statistical difference in the age of the two groups (SMD = 0.085, 95% CI = -0.073–0.189, $p = 0.388$,

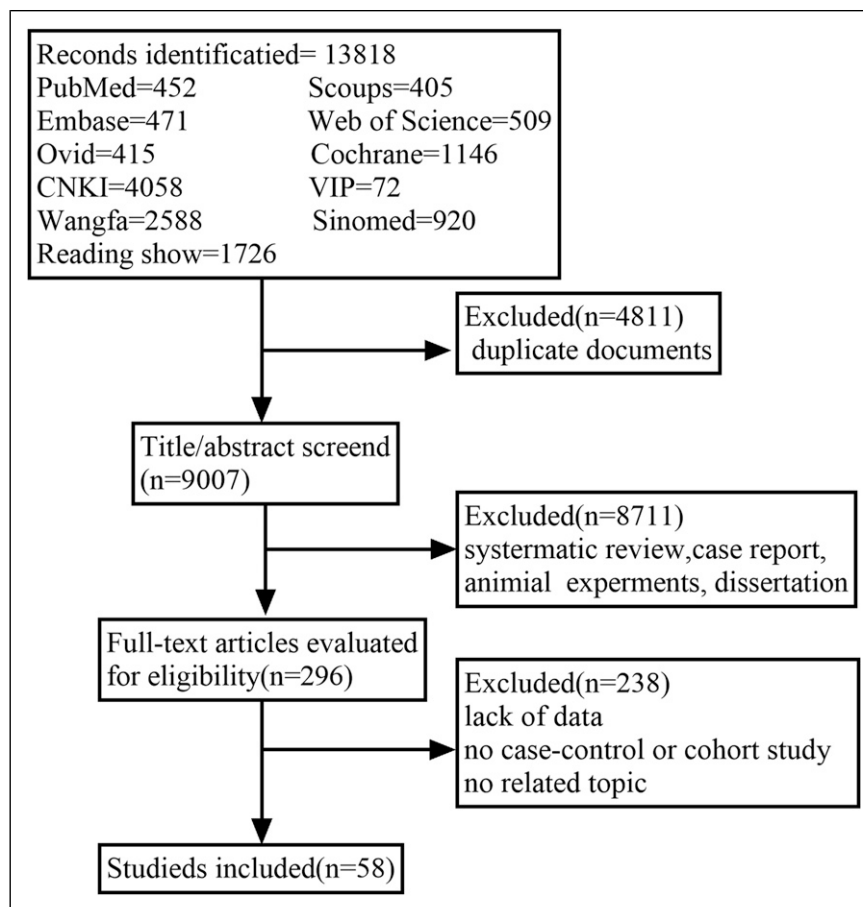


Figure 1. Flowchart of the retrieval process.

Table I. Basic information and quality evaluation of included studies.

Study	Type	Vaccinated	Unvaccinated	NOS			Scores
				Selection	Comparability	Outcome	
Li ¹⁷	Case-control study	44	67	☆☆☆	☆☆	☆☆	7☆
Balachandran ¹²	Case-control study	346	1 100	☆☆☆	☆☆	☆☆	7☆
Wi ¹⁸	Case-control study	142	20	☆☆	☆☆	☆☆	6☆
Tenforde ¹³	Case-control study	314	1669	☆☆☆	☆☆	☆☆	7☆
Hsu ¹⁹	Case-control study	85	85	☆☆	☆☆	☆☆☆	7☆
Blanquart ²⁰	Case-control study	724	5459	☆☆☆	☆☆	☆☆	7☆
Bollineni ²¹	Case-control study	14	56	☆	☆☆	☆☆☆	6☆
Marincu ²²	Case-control study	62	62	☆☆	☆☆	☆☆☆	7☆
Wolff ²³	Case-control study	260	507	☆☆☆	☆☆	☆☆☆	8☆
Kalligeros ²⁴	Case-control study	91	824	☆☆☆	☆☆	☆☆☆	8☆
Fragoulis ²⁵	Cohort study	101	60	☆☆☆	☆☆	☆☆☆	8☆
Trunfio ²⁶	Cohort study	55	110	☆☆☆	☆☆	☆☆	7☆
Safdar ²⁷	Cohort study	45	65	☆☆☆	☆☆	☆☆☆	8☆
Bouton ²⁸	Cohort study	96	329	☆☆	☆☆	☆☆☆	7☆
Bosch ²⁹	Case-control study	126	839	☆☆	☆☆	☆☆☆	7☆
Christensen ³⁰	Case-control study	3346	13,619	☆☆☆	☆☆	☆☆☆	8☆
Yu ³¹	Case-control study	23	50	☆☆☆	☆☆	☆☆☆	8☆
Bayhan ³²	Case-control study	38	190	☆☆☆	☆☆	☆☆☆	8☆
Kustin ³³	Case-control study	396	396	☆☆	☆☆	☆☆	6☆
Toda ³⁴	Case-control study	11	15	☆☆☆	☆☆	☆☆☆	8☆
Tian ⁴	Case-control study	88	41	☆☆☆	☆☆	☆☆☆	8☆
Sayed ³⁵	Case-control study	50	70	☆☆☆	☆☆	☆☆☆	8☆
Lee ³⁶	Case-control study	174	587	☆☆☆	☆☆	☆☆☆	8☆
Luo ³⁷	Case-control study	484	1782	☆☆☆	☆☆	☆☆	7☆
Puhach ³⁸	Case-control study	139	245	☆☆	☆☆	☆☆	6☆
Whittaker ³⁹	Case-control study	716	2487	☆☆☆	☆☆	☆☆☆	8☆
Sanghavi ⁴⁰	Case-control study	149	478	☆☆☆	☆☆	☆☆☆	8☆
Butt (1) ⁴¹	Case-control study	250	250	☆☆☆	☆☆	☆☆☆	8☆
Butt (2) ⁴²	Case-control study	456	456	☆☆☆	☆☆	☆☆☆	8☆
Spiera ⁴³	Case-control study	88	2317	☆☆	☆☆	☆☆☆	7☆
Baltas ⁴⁴	Case-control study	119	476	☆☆☆	☆☆	☆☆	7☆
John ⁴⁵	Case-control study	254	508	☆☆	☆☆	☆☆	6☆
Jacobson ⁴⁶	Case-control study	189	471	☆☆	☆☆	☆☆	6☆
Estofolete ⁴⁷	Case-control study	259	2518	☆☆☆	☆☆	☆☆☆	8☆
Servellita ⁴⁸	Case-control study	39	433	☆☆☆	☆	☆☆☆	7☆
Thangaraj ⁴⁹	Case-control study	376	183	☆☆☆	☆☆	☆☆☆	8☆
Galan-huerta ⁵⁰	Case-control study	53	19	☆☆☆	☆☆	☆☆☆	8☆
Butt (3) ⁵¹	Case-control study	2332	40,540	☆☆	☆☆	☆☆☆	7☆
Romano ⁵²	Case-control study	34	86	☆☆☆	☆☆	☆☆☆	8☆
Basso ⁵³	Case-control study	155	81	☆☆	☆☆	☆☆☆	7☆
Griffin ⁵⁴	Case-control study	12,179	29,989	☆☆☆	☆☆	☆☆☆	8☆
Padovani ⁵⁵	Case-control study	234	50	☆☆☆	☆☆	☆☆	7☆
Anand ⁵⁶	Case-control study	3240	1551	☆☆	☆☆	☆☆☆	7☆
Rovida ⁵⁷	Case-control study	33	20	☆☆	☆☆	☆☆☆	7☆
Taquet ⁵⁸	Case-control study	10,024	83,957	☆☆☆	☆☆	☆☆☆	8☆
Cocchio ⁵⁹	Case-control study	773	12,499	☆☆	☆☆	☆☆☆	7☆
Stupica ⁶⁰	Case-control study	175	354	☆☆☆	☆☆	☆☆☆	8☆
Ioannou ⁶¹	Cohort study	24	31	☆☆	☆☆	☆☆☆	7☆
Bahl ⁶²	Cohort study	954	10,880	☆☆☆	☆☆	☆☆☆	8☆

(continued)

Table 1. (continued)

Study	Type	Vaccinated	Unvaccinated	NOS			Scores
				Selection	Comparability	Outcome	
Muthukrishnan ⁶³	Cohort study	450	718	☆☆☆	☆☆	☆☆☆	8☆
Chia ⁶⁴	Case-control study	84	130	☆☆	☆☆	☆☆	6☆
Zheng ⁶⁵	Case-control study	39	127	☆☆☆	☆☆	☆☆☆	8☆
Chen ⁶⁶	Case-control study	38	38	☆☆☆	☆☆	☆☆☆	8☆
Ma ⁶⁷	Case-control study	46	43	☆☆☆	☆☆	☆☆☆	8☆
Jiang ⁶⁸	Case-control study	85	362	☆☆☆	☆☆	☆☆☆	8☆
Yue ⁶⁹	Case-control study	20	55	☆☆☆	☆☆	☆☆☆	8☆
Papagoras ⁷⁰	Case-control study	48	147	☆☆	☆☆	☆☆☆	7☆
Naik ⁷¹	Case-control study	1010	1003	☆☆	☆☆	☆☆	7☆

Figure S2(a)). Through subgroup analysis, we found that heterogeneity was mainly present in groups with more than 300 people in the population (Figure S2(b)).

Severity of disease

We would assess the severity of the disease, asymptomatic infection, and disease activity. We found that the severe patients in the vaccinated group were 0.247 times lower than those in the unvaccinated group (OR = 0.247, 95% CI: 0.128–0.474, $p < 0.001$, Figure S3(a)), and subgroup analysis found heterogeneity only present in studies with a population of 1000 (Figure S3(b)). The patients with asymptomatic infection were 1.686 times higher than those who were in the unvaccinated group (OR = 1.686, 95% CI: 1.277–2.225, $p < 0.001$, Figure S3(c)), and heterogeneity existed in studies with a vaccinated population of 30 and 1000 (Figure S3(d)). There was no statistical difference between the two groups in terms of disease activity (OR = 1.266, 95% CI: 0.598–2.68, $p = 0.538$, Figure S3(e)).

Epidemiology and personal history

The studies would be conducted from race, lineage, occupation, smoking history, and pregnancy history. We found noted that the vaccinated group being a Caucasian was higher than the unvaccinated group (OR = 1.52, 95% CI: 1.255–1.887, $p < 0.001$, Figure S4(a)); through subgroup analysis, we found that the source of heterogeneity came from studies with a total population more of than 10,000 (Figure S4(b)). For Black or African American (OR = 0.761, 95% CI: 0.517–1.121, $p = 0.167$, Figure S4(c)) and Asian (OR = 1.269, 95% CI: 0.65–2.476, $p = 0.486$, Figure S4(d)), there were no statistical difference between the two groups of patients. In vaccine variants, B.1.1.7 (OR = 1.609, 95% CI: 1.04–2.488, $p = 0.032$, Figure S4(e)) in the vaccinated group was 1.609 times higher than the unvaccinated group. And there were no statistically significant differences in

occupational (doctors or non-doctors) (OR = 1.874, 95% CI: 0.835–4.202, $p = 0.128$, Figure S4(f)), pregnancy history (OR = 0.314, 95% CI: 0.086–1.15, $p = 0.08$, Figure S4(g)), and smoking (OR = 0.824, 95% CI: 0.668–1.071, $p = 0.071$, Figure S4(h)) between the two groups.

Comorbidity

We would analyze whether the presence of comorbidities and the different types of comorbidities. Including 8 literature studies, we found no difference in the presence of comorbidities between the vaccinated and unvaccinated groups (OR = 1.75, 95% CI: 0.936–3.273, $p = 0.08$, Figure S5(a)). Diabetes in the vaccine group was higher than in the unvaccinated group (OR = 1.407, 95% CI: 1.028–1.925, $p = 0.033$, Figure S5(b)). By subgroup analysis, heterogeneity was only present in studies with vaccinated groups of 2000 (Figure S5(c)). More patients with hyperlipidemia (OR = 1.806, 95% CI: 1.316–2.478, $p < 0.001$, Figure S5(d)) in the vaccine group. However, obesity in the vaccinated group was lower than those in the vaccine group (OR = 0.873, 95% CI: 0.843–0.905, $p < 0.001$, Figure S5(e)). By subgroup analysis, heterogeneity exists only in studies with a total population of 10,000 (Figure S5(f)).

There were also more kidney disease (OR = 1.827, 95% CI: 1.182–2.823, $p = 0.007$, Figure S6(a)), immunocompromised patients (OR = 2.615, 95% CI: 1.671–4.092, $p < 0.001$, Figure S6(b)), and organ transplants (OR = 3.727, 95% CI: 2.682–5.18, $p < 0.001$, Figure S6(c)), and cardiovascular (OR = 1.667, 95% CI: 1.076–2.582, $p = 0.022$, Figure S6(d)), and tumors (OR = 1.296, 95% CI: 1.249–1.344, $p < 0.001$, Figure S6(e)) in the vaccinated group than the unvaccinated group.

There were no statistical differentials in liver disease (OR = 0.939, 95% CI: 0.871–1.012, $p = 0.101$, Figure S7(a)), lung disease (OR = 1.054, 95% CI: 0.673–1.65, $p = 0.818$, Figure S7(b)), hypertension (OR = 1.439, 95% CI:

0.878–2.356, $p = 0.149$, [Figure S7\(c\)](#)), and cerebrovascular disease (OR = 1.551, 95% CI: 0.74–3.25, $p = 0.245$, [Figure S7\(d\)](#)).

Symptom

Among the symptom of fever, the morbidity rate of patients in the vaccinated group was lower (OR = 0.583, 95% CI: 0.421–0.808, $p = 0.001$, [Figure S8\(a\)](#)). Patients had lower cough (OR = 0.718, 95% CI: 0.597–0.865, $p < 0.001$, [Figure S8\(b\)](#)), and gastrointestinal symptoms (OR = 0.743, 95% CI: 0.585–0.944, $p = 0.015$, [Figure S8\(c\)](#)) and dysgeusia/anosmia (OR = 0.449, 95% CI: 0.24–0.84, $p = 0.012$, [Figure S8\(d\)](#)) in the vaccinated group. However, among patients with neurological symptoms (OR = 1.950, 95% CI: 1.053–3.61, $p = 0.034$, [Figure S8\(e\)](#)), it was higher.

There was no statistical difference between the two groups of patients in the symptoms of anemia (OR = 2.183, 95% CI: 0.951–5.013, $p = 0.066$, [Figure S9\(a\)](#)), sore throat (OR = 0.785, 95% CI: 0.591–1.041, $p = 0.093$, [Figure S9\(b\)](#)), muscle and joint pain (OR = 0.56, 95% CI: 0.276–1.138, $p = 0.109$, [Figure S9\(c\)](#)), headache (OR = 0.77, 95% CI: 0.467–1.269, $p = 0.305$, [Figure S9\(d\)](#)), fatigue (OR = 0.362, 95% CI: 0.059–2.215, $p = 0.271$, [Figure S9\(e\)](#)), nausea/vomiting (OR = 1.106, 95% CI: 0.729–1.679, $p = 0.636$, [Figure S9\(f\)](#)), low oxygen saturation (OR = 0.79, 95% CI: 0.358–1.744, $p = 0.559$, [Figure S9\(g\)](#)), dyspnea (OR = 0.682, 95% CI: 0.483–1.021, $p = 0.063$, [Figure S9\(h\)](#)) and nasal congestion and runny nose (OR = 1.929, 95% CI: 0.85–4.379, $p = 0.116$, [Figure S9\(i\)](#)).

Treatment

We would analyze two aspects of supportive care and drug treatment. Patients in the vaccinated group required less oxygen support than those in the unvaccinated group (OR = 0.579, 95% CI: 0.418–0.801, $p = 0.001$, [Figure S10\(a\)](#)). Steroid use was lower in the vaccinated group than in the unvaccinated group (OR = 0.679, 95% CI: 0.496–0.929, $p = 0.015$, [Figure S10\(b\)](#)). There was no statistical difference in the use of antiviral drugs (OR = 0.896, 95% CI: 0.595–1.348, $p = 0.597$, [Figure S10\(c\)](#)) between the two groups of patients.

Hospitalization

A total of 4 studies compared the days of hospitalization and we found the duration in days in the vaccinated group was 0.79 times lower than the unvaccinated group (SMD = -0.790 , 95% CI: -1.123 to -0.458 , $p < 0.001$, [Figure S11\(a\)](#)). Fewer patients in the vaccinated group required hospitalization (OR = 0.660, 95% CI: 0.489–0.889, $p = 0.006$, [Figure S11\(b\)](#)), and ICU treatment

(OR = 0.648, 95% CI: 0.479–0.876, $p = 0.005$, [Figure S11\(c\)](#)). There was no statistically significant difference between the two groups of patients who lived in a nursing facility for a long time (OR = 1.773, 95% CI: 0.571–5.5, $p = 0.322$, [Figure S11\(d\)](#)).

Outcome and prognosis

A total of 31 studies were included to investigate the prognosis of patients. And we found that the mortality rate of patients in the vaccinated group was 0.659 times lower than that of patients in the unvaccinated group (OR = 0.659, 95% CI: 0.462–0.94, $p = 0.021$, [Figure S12\(a\)](#)). In the study of patients with poor prognosis, we found that the poor prognosis of patients in the vaccinated group was 0.315 times lower than that of patients in the unvaccinated group (OR = 0.315, 95% CI: 0.148–0.668, $p = 0.003$, [Figure S12\(b\)](#)).

Discussion

We found that vaccination could reduce the severity of disease, which was consistent with the study of Giuseppe,⁷² and more patients with asymptomatic infection were found in the vaccinated group in our study. What plays a central role in vaccines are the antigens.⁷³ Vaccines can stimulate the patient's humoral and cellular immune systems. They play a role similar to live viruses,⁷⁴ and stimulate the body to retain relevant stimulatory T cells and related memory B cells. Studies have shown that T cell responses play an important role in maintaining long-term immunity.⁷⁵ When exposed to the virus, the body can quickly initiate immunity, reduce immune damage and the severity of the disease. Therefore, severe patients in the vaccine group were fewer, mostly asymptomatic infections, in line with global data.^{76–78}

Statistical results show that vaccinated patients were more Caucasian than unvaccinated patients, which may be due to the uneven distribution of vaccines. Studies have shown that 4.2 billion doses of vaccines have been ordered by high-income countries, and 70% of vaccine candidates have been purchased.⁷⁹ The B.1.1.7 strain, which was one of the four major mutants, with more infections in patients in the vaccine group. VOCs have more evasion capabilities for vaccines, changing the efficacy of neutralizing antibodies to affect the efficacy and effectiveness of vaccines. Animal models have demonstrated that the lack of neutralizing antibodies can lead to increased inflammation and poor clinical outcomes.^{80,81} Clinical studies have also shown that neutralizing antibodies are associated with faster virus clearance responses.⁹ Neutralizing antibodies in patients with vaccine breakthrough infection are significantly

lower than those without breakthrough infection of patients.

For COVID-19 patients with pregnancy, there was no difference in infection between the two groups of patients, indicating that pregnancy did not increase the susceptibility of COVID-19's infection. We discovered more comorbidities in the vaccine group than in the unvaccinated group. In a study by Kertes,⁸² it was found that patients with comorbidities had lower levels of antibodies than healthy individuals. Vaccination is more beneficial for patients with comorbidities and immunocompromised patients.⁸³ Studies have shown that vaccinated patients have lower levels of autoimmune IG receptor antibodies and ACE2 receptor antibodies in transplant patients, kidney disease, malignancies, and autoimmune diseases, so they are more susceptible to infection.⁸⁴

In our study, fever and cough, typical symptoms of COVID-19 patients, were lower in the vaccine group. Nevertheless, neurological symptoms and taste/smell disturbance symptoms were higher in the vaccine group, which was consistent with Sagar.⁸⁵ This may be due to the lack of protection of the olfactory cleft and oral cavity by circulating antibodies.

We also found that patients in the vaccine group required less oxygen support and hormone steroids. Due to the characteristics of viral pneumonia, COVID-19 shows symptoms such as abnormal hemorrhagic gas, alveolar exudation, and accumulation of inflammatory markers of inflammatory factors. So oxygen therapy and symptomatic treatment are often used.^{86,87} For antiviral therapeutic drugs, such as Remdesivir, an RNA polymerase (RdRp) blocker, can inhibit the synthesis of viral nucleic acid, and favipiravir inhibits the synthesis of viral genomic RNA to achieve the effect of antiviral therapy.⁸⁸ For immunosuppressants, studies have shown that such drugs can be used to block the effects of inflammatory cytokine storms.⁸⁹

The results showed that patients in the vaccine group required fewer hospitalizations and ICU treatment, and the incidence of death or poor prognosis was lower, indicating that the vaccine was very important for the reduction of hospitalization and mortality. Which was consistent with real-world studies⁹⁰ but inconsistent with research of Lee.⁹¹

Despite the fact that patients are still at risk of contracting COVID-19 after vaccination, our study indicates that vaccination provides a protective effect on us. Vaccines are the pillars of public health, and policy makers should encourage vaccination of people, especially occupational health service workers, whose occupational risk assessment plays an important role in preventing and combating COVID-19.^{92,93} And it is also suggested that for the emergence of different VOCs, the research of vaccines should be increased to maintain the effectiveness of vaccines in the population.

There are, however, limitations to this study as well. Laboratory data and antibodies of the two groups of patients are lacking, imaging characteristics of the different patients are lacking, and comparative research on cytokines has not been conducted. There are few research records on the Ct value and viral load of the two groups of patients, which cannot be included in this study. Therefore, we were unable to assess relative differences in viral load across groups of patients. Due to the different regions and sample sizes included in this study, the heterogeneity of the results of this study is also quite different, which may affect the accuracy of the results. More studies are expected to be added in the future to address these limitations.

Conclusion

Based on our study, the vaccine appears to provide a protective effect for patients with COVID-19, with significant improvements in symptoms, hospitalizations, and patient outcomes. Similarly, we also observed that Caucasians were more likely to be vaccinated, and that those who were vaccinated were more likely to be exposed to VOCs. Our research shows that vaccination is an effective tool for combating COVID-19.

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This manuscript has been approved for publication by all authors. This manuscript represents original research that has not been published elsewhere. A consensus has been reached among the authors regarding the content of this manuscript.

Author contribution

Research design, literature search, and article preparation were conducted by Wen Tian and Xingxiang Ren, statistical methods were contributed by Mei Han, data extraction and quality evaluation were completed by Xu Gao, Yuanyuan Zhang, and Zhihai Chen. Wei Zhang contributed research ideas and reviewed the manuscript. The final manuscript has been approved by all authors.

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Supplemental Material

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