Contents lists available at ScienceDirect



International Journal of Infectious Diseases

journal homepage: www.elsevier.com/locate/ijid

# Side effects and perceptions following Sinopharm COVID-19 vaccination





# Balsam Qubais Saeed<sup>a,b,\*</sup>, Rula Al-Shahrabi<sup>c</sup>, Shaikha Salah Alhaj<sup>c</sup>, Zainab Mansour Alkokhardi<sup>c</sup>, Ahmed Omar Adrees<sup>c</sup>

<sup>a</sup> Clinical Sciences Department, College of Medicine, University of Sharjah, Sharjah, UAE

<sup>b</sup> Sharjah Institute for Medical Research, University of Sharjah, Sharjah, UAE

<sup>c</sup> College of Medicine, University of Sharjah, Sharjah, UAE

# ARTICLE INFO

Article history: Received 13 June 2021 Revised 4 August 2021 Accepted 5 August 2021

Keywords: COVID-19 vaccine Sinopharm COVID-19 vaccine Sinopharm BBIBP-CorV Side effects Chronic diseases United Arab Emirates

# ABSTRACT

*Objectives:* Vaccines are one of the best interventions developed for eradicating COVID-19. This study aimed to provide evidence on Sinopharm COVID-19 vaccine side effects.

*Methods:* A cross-sectional survey study was conducted between January and April 2021 to collect data on the effects of the COVID-19 vaccine among individuals in the United Arab Emirates. Demographic data, vaccination and the response of people unwilling to take the COVID-19 vaccine were reported.

*Results:* Side effects post first vaccine dose of normal injection site pain, fatigue and headache were more common in participants aged  $\leq$ 49 years versus >49 years, while pain at the vaccination site, fatigue, lethargy, headache and tenderness were the most common side effects post second dose in both groups. All side effects for both doses were more prevalent among participants aged  $\leq$ 49 years. Side effects were more common in females compared with males for both doses. The most common reason for being unwilling to take the COVID-19 vaccine was that vaccines are not effective.

*Conclusion:* Post-vaccination side effects for the first and second doses were mild and predictable, and there were no hospitalization cases; this data will help reduce vaccine hesitancy.

© 2021 The Author(s). Published by Elsevier Ltd on behalf of International Society for Infectious Diseases.

This is an open access article under the CC BY license (http://creativecommons.org/licenses/by/4.0/)

# Introduction

SARS-CoV-2, causing COVID-19 (Habas et al., 2020), has spread fast worldwide, resulting in various levels of illness. On March 11, 2020, it was announced that SARS-CoV-2 is a worldwide pandemic, and it is with us to this day (Lai et al., 2020). Although numerous therapeutic medications have been presented to resist COVID-19, they remain supportive and require more randomized control studies to determine their efficacy and potency. (Coronavirus disease (COVID-19) 2021, Trivedi et al., 2020).

Vaccines are one of the best interventions developed for eradicating COVID-19, saving millions of lives annually. Moreover, the best option remains an effective, safe vaccine without severe adverse reactions. The lack of effective and approved COVID-19 treatment has triggered a vaccine development race, with 259 COVID-19 vaccine projects underway from November 11, 2020. The rapid creation of vaccinations has increased the risk of vaccine safety issues (Haidere et al., 2021, Petousis-Harris, 2021).

\* Corresponding address. E-mail address: bsaeed@sharjah.ac.ae (B.Q. Saeed). Several candidate COVID-19 vaccines were developed from diverse platforms. One of these was the BBIBP-CorV vaccine (also known as the Sinopharm COVID-19 vaccine) which was made by the Chinese state-owned pharmaceutical business Sinopharm in China and adopted by the United Arab Emirates (UAE) (Zhang et al., 2021). Sinopharm COVID-19 vaccine is an inactivated vaccine that introduces a dead copy of SARS-CoV-2 into the body by a two-dose schedule, with 14 or 21 days between the 2 doses. By inserting the vaccine dose intramuscularly, the dead antigens from the virus are employed to make antibodies that prepare the immune system for future attacks by the virus. (Xia et al., 2021). The traditional inactivated whole-virus vaccines do not lead to clinical disease. In this technology, the inactivated viruses maintain their ability to replicate in vivo with mild or no symptoms (Forni and Mantovani, 2021).

Phases 1 and 2 of the clinical trials for the Sinopharm COVID-19 vaccine were carried out in China over 1 trial for each phase. A total enrolment of 640 participants showed that the vaccine triggered a COVID-19 neutralizing antibody response with a low rate of adverse reactions. The most common side effects were fever and pain at the site of injection and fever; however, these were

https://doi.org/10.1016/j.ijid.2021.08.013

<sup>1201-9712/© 2021</sup> The Author(s). Published by Elsevier Ltd on behalf of International Society for Infectious Diseases. This is an open access article under the CC BY license (http://creativecommons.org/licenses/by/4.0/)

mild and self-limiting and did not require treatment (Sharma et al., 2020). Phase 3 was carried out over 4 trials in the following countries: the UAE, Bahrain, Egypt, Jordan, Peru and Argentina, with a total 69 000 people enrolled. The UAE approved the vaccine on December 9, 2020; the UAE announced that the vaccine was 86% efficacious, according to the interim results of its phase 3 trial (Xia et al., 2021).

Having administered over 2 million doses of the vaccine as of mid-January 2021, the UAE Ministry of Health reported the vaccine to be 100% effective in preventing mild and severe COVID-19 cases (China to run human coronavirus vaccine trial in UAE 2020, Development of an Inactivated Vaccine Candidate, 2021, Cyranoski, 2020, Zahid et al., 2021). Moreover, the World Health Organization (WHO) announced that most side effects of the Sinopharm COVID-19 vaccine in 3 clinical trials on 16 671 participants aged 18–59 years were mild to moderate, with headache, fatigue, and injection site reactions being the most common (World Health Organization (WHO) 2021).

Published data to support adverse reactions to the Sinopharm COVID-19 vaccine are lacking. Only 2 studies focus on this vaccine (Hatmal et al., 2021, Jayadevan et al., 2021), and fear of the new vaccine is a driver of vaccine hesitancy (Riad et al., 2021).

Knowledge on what happens post-vaccination among the general population is still limited. Describing what to expect after the first and second dose of vaccination will help lower apprehension about this type of vaccine, increase public confidence in vaccine safety, and accelerate the vaccination process against COVID-19 (Hatmal et al., 2021, Jayadevan et al., 2021). The results of this study will be reassuring to those who are fearful of the Sinopharm COVID-19 vaccine. Therefore, the goal of this study was to provide evidence on Sinopharm COVID-19 vaccine side effects after receiving the first and second dose, as approved by the UAE.

#### Methodology

#### Study design, setting and participants

This cross-sectional survey-based study was carried out from 10 January to 30 April 2021 to evaluate Sinopharm COVID-19 vaccine adverse reactions among residents in the UAE. The study utilized a self-administered online survey created on Google Forms which had been randomly delivered to individuals (aged  $\geq$ 18 years) using social media sites (Facebook, Email and WhatsApp). Potential participants were directed to a webpage with a brief introduction to the aim and purpose of the study and instructions on how to complete the survey. Informed consent that included statements about voluntary participation and anonymity was sought from all the respondents prior to data collection by sending a standardized general invitation letter with the survey link to accept or decline participation in the study. Participants who declined consent were not permitted to open the survey and participate in the study, and participants could withdraw from the survey at any time in line with the World Medical Association Declaration of Helsinki Ethical principles (Aresté and Salgueira, 2013). Persons who clicked on the link were directed to Google Forms and, to avoid issues of missing data, they had to respond to all the questions or were unable to proceed to the next section of the survey. No incentive or compensation was given to participants (Elm et al., 2007).

Out of 1102 surveys received from respondents, 1080 participants aged  $\geq$ 18 years, from different emirates and nationalities, were included in this study. The study sample included participants who were either vaccinated with the first dose or second dose of Sinopharm COVID-19 vaccine and those who did not receive any COVID-19 vaccine during the early peak of the vaccination campaign in the UAE. Individuals aged <18 years and non-residents in the UAE were excluded from this study as they were

not permitted to receive any vaccine type in the UAE during the conduct of this study. Individuals who had received a COVID-19 vaccine other than Sinopharm were not included in the study, based on the study's aims.

#### Survey and data collection

The survey was designed based on extensive literature searches and guidelines from the WHO, MOHAP, and the Abu Dhabi Department of Health (DOH) in the UAE on the expected adverse reactions post Sinopharm COVID-19 vaccine. The survey questions were multiple choice and were made available in English and Arabic (Xia et al., 2021, World Health Organization (WHO) 2021, COVID-19 Vaccine 2021). The survey was validated by a group of experts who provided feedback on the different items of the survey.

The survey was in three sections. The first section included 7 demographic questions on gender, age, marital status, education level, employment, nationality, and the region of residence. The second section reviewed participants' chronic conditions such as: cancer, autoimmune diseases, chronic respiratory diseases, diabetes, hypertension, obesity, heart disease, allergic reactions to vaccination (any type of immediate allergic reaction to a vaccine or injectable therapy), receiving immunotherapy or inhibitor therapy, severe anemia (level less than 7.0 g/dl), liver diseases, and no chronic condition.

The last section was related to Sinopharm COVID-19 vaccine side effects. In the first section, participants had been asked about COVID-19 vaccination status to determine the scope of the rest of the questions. If the participant had received the vaccine, we asked about the number of doses administered and previous COVID-19 infection; then questions regarding side effects to vaccine first and second dose were asked separately based on each dose side effects, namely, normal/severe pain at the vaccination site, tenderness, redness, pruritus at the vaccination site, fever, headache, fatigue, nausea, diarrhea, cough, allergy, muscle pain, abdominal pain, back pain, and lethargy.

If the participant had not received any COVID-19 vaccine, a question was asked to clarify the reasons behind this. For pilot testing, the survey was passed randomly to 15 participants recently vaccinated who completed the survey after taking the 2 doses and were then excluded from the study. The Cronbach's alpha test of internal consistency was used to evaluate survey reliability. The overall reliability was 0.81, indicating that the survey tool was reliable with good internal consistency (McHugh, 2012, Nunnally and Bernstein, 1994).

#### Sample size

The online Raosoft sample size calculator (Raosoft Inc., Seattle, WA, USA) was used to calculate the sample size of our study. Based on world meter elaboration of the latest United Nations data, the UAE population is approximately 10 million. Around 77.3% of the population had received at least one dose of the vaccine (according to the UAE supreme council for national security). We assumed the confidence level is 95%, the margin of error is 3%, and the response distribution is 50%. With a population of 7.7 million (vaccinated with at least one dose), the sample size recommended was 1067; the study included 1080 respondents meaning a convenience sample size was used (Sample size calculator, 2021, COVID-19 vaccines, 2021).

#### Statistical analysis

The Statistical Package for the Social Sciences version 22.0 (SPSS Inc. Chicago, IL, USA, 2013) was used to carry out descriptive statistics of 1080 participants for the demographic variables (gender,

Variable	Outcomes	Number (n)	Percentage (%)
Gender	Female	760	70.4
	Male	320	29.6
Age	18-29	424	39.2
	30–39	184	17
	40-49	236	22
	50–59	140	13
	60-80	96	8.8
Marital status	Married	600	55.6
	Single	440	40.7
	Widowed	32	3.0
	Divorced	8	0.7
Education level	College level	644	59.7
	High school/diploma	228	21.1
	Postgraduate (Masters/Doctorate)	180	16.6
	Primary	16	1.5
	Illiterate	12	1.1
Employment status	Employed	508	47.1
	Student	304	28.2
	Unemployed	232	21.4
	Retired	36	3.3
Nationality	Non-Emirati	856	79.3
	Emirati	224	20.7
Emirate	Sharjah	572	52.9
	Dubai	288	26.7
	Ajman	116	10.8
	Abu Dhabi	96	8.8
	Umm Al Quwain	8	0.7

Demographic characteristics of participants (n 1080)

#### Table 2

Health status and chronic conditions of participants (n 1080)

Conditions	Number (n)	Percentage (%)
Cancer	8	0.7
Autoimmune disease	4	0.4
Chronic respiratory disease	40	3.7
Diabetes	84	7.8
Hypertension	68	6.3
Obesity	36	3.3
Heart disease	40	3.7
Severe allergic reactions to vaccination	4	0.4
Receiving immunotherapy or inhibitor therapy	4	0.4
Severe anemia	8	0.7
Liver disease	4	0.4
No chronic condition	780	72.2

age, education level, marital status, employment status, nationality and region of residence). In addition, health and chronic conditions, COVID-19-related anamnesis, such as vaccination status, previous infection, and the number of vaccination doses, and the reasons for participants' vaccination hesitancy, were analysed. Data are presented in both frequencies and percentages.

In our study, the mean age was  $37.2\pm13.1$ . Age was classified into 4 groups ranging from 18 to 80 years old. The median age of our sample, 49 years old, was selected to be a cutoff point to evaluate the difference in side effects between 2 groups of participants ( $\leq$ 49 vs >49) using a Chi-squared test. Similarly, a Chi-squared test was performed to assess the correlation between vaccine side effects and gender.

# Results

#### Demographic characteristics

Table 1 shows the demographic data of participants: 760 (70.4%) were female, 320 (29.6%) male. The mean age was  $37.22\pm13.1$  years; 440 (40.7%) were single, 600 (55.6%) married, and 3.7% divorced or widowed. On education level, 644 (59.7%) held a bachelor's degree, 288 (21.1%) a high school degree or be-

low, and 180 (16.6%) a postgraduate degree. The majority (508, 47.1%) were employed, 232 (21.5%) unemployed and 304 (28.2%) were students. Most participants (856, 79.3%) were non-Emirati, 224 (20.7%) were Emirati. Many participants (52.9%) lived in Sharjah, 288 (26.7%) in Dubai and the rest (220, 20.3%) in the other Emirates.

#### Chronic conditions among the participants

Of 1080 participants, 780 (72.2%) were healthy, while 300 (27.8%) had chronic conditions. The most prevalent chronic conditions, as shown in Table 2, were diabetes 7.8% followed by hypertension 6.3%, while, respectively, 3.7%, 3.7%, 3.3%, 0.7%, 0.7%, 0.4%, 0.4%, 0.4%, and 0.4% suffered chronic respiratory disease, heart disease, obesity, cancer, severe anemia, autoimmune disease, severe allergies, were receiving immunotherapy, and had liver disease.

Relationship between the side effects of vaccination and age ( $\leq$ 49 years vs >49 years)

Table 3 compares COVID-19-related anamnesis of vaccinated individuals aged below and above 49. The study shows a significant difference in vaccination status between people who are  $\leq$ 49

COVID-19-related anamnesis of vaccinated individuals (n 1080)

Anamnesis	Outcome	$\leq$ 49 Years Old (844)	>49 Years Old (236)	Total
Vaccinated	Yes	632 (75%)	200 (85%)	832 (77%)
	No	212 (25%)	36 (15%)	248(23%)
Number of doses	One dose	128 (15%)	56 (23%)	184 (17%)
	Two doses	508 (60%)	144 (61%)	652 (60 %)
	None	208 (25%)	36 (15%)	244 (23%)
COVID-19 infection	Yes	140 (16.6%)	36 (15%)	176 (16.1%)
	No	684 (81%)	200 (84%)	884 (81.5%)
	Not sure	20 (2.4%)	0	20 (2.4%)

#### Table 4

Prevalence of general side effects after the first dose of the Sinopharm COVID-19 vaccination among 2 groups of participants ( $\leq$ 49 years old vs >49 years old) (n 1080)

Side effect	${\leq}49$ years (844) n (%)	>49 years (236) n (%)	Total (1080) n (%)	P-value
Normal pain at the vaccination site	356 (42.2)	100 (42.4)	456 (42.2)	0.999
Severe pain at the vaccination site	12 (1.4)	16 (6.8)	28 (2.6)	0.023
Tenderness	44 (5.2)	12 (5)	56 (5.1)	0.963
Redness	8 (0.9)	0	8 (0.7)	0.452
Induration and pruritus at the vaccination site	12 (1.4)	0	12 (1.1)	0.356
Fever	8 (0.9)	4 (1.7)	12 (1.1)	0.631
Headache	92 (10.9)	12 (5)	104 (9.6)	0.178
Fatigue	96 (11.4)	36 (15.3)	132 (12.2)	0.429
Nausea	4 (0.5)	12(5)	16 (1.5)	0.010
Diarrhea	8 (0.9)	0	8 (0.74)	0.452
Cough	8 (0.9)	4 (1.7)	12 (1.1)	0.631
Allergy	8 (0.9)	4 (1.7)	12(1.1)	0.631
Muscle pain	36 (4.3)	32 (13.6)	68 (6.3)	0.010
Abdominal pain	12 (1.4)	8 (3.4)	20 (1.85)	0.324
Back pain	24 (2.8)	20 (8.5)	44 (4.07)	0.054
Lethargy	76 (9)	24 (10)	100 (9.2)	0.631
Other	12 (1.4)	0	8 (0.7)	0.452
None	220 (26.0)	44 (18.6)	264 (24.4)	0.234

years old and >49 years old (*P*=0.000), with participants aged >49 years more likely to be vaccinated (85%) compared with those  $\leq$ 49 (75%). Differences in the number of doses between these 2 groups was not significant (*P*=0.128). However, there was a significant difference between the 2 groups in whether participants had, had COVID-19 infection previously (*P*=0.002), with those aged  $\leq$ 49 more likely to have had a previous COVID-19 infection (16.6%).

Table 4 presents the prevalence of general adverse reactions to the first dose of the Sinopharm COVID-19 vaccine in participants ( $\leq$ 49 years vs >49 years old). The table shows that overall 24.4% (26% vs 18.6%) did not have side effects. Common side effects among both age groups of participants were normal pain at the site of vaccination (42.2%), fatigue (12.2%) and headache (9.6%). The same table indicates a significant difference between the 2 age groups ( $\leq$ 49 years vs >49 years old) for severe pain at the vaccination site (*P*=0.023), nausea (*P*=0.010) and muscle pain (*P*=0.010). Of participants >49 years, 6.8% reported severe pain at the vaccination site compared with 1.42% among those aged  $\leq$ 49. More participants aged >49 years reported nausea and muscle pain compared with participants aged  $\leq$ 49 years, 0.5% vs 5.0% and 4.3% vs 13.6%, respectively.

The tables show no significant difference between the age groups in other side effects, namely normal pain at the vaccination site (P=0.999), tenderness (P=0.963), redness (P=0.452), induration and pruritus at the vaccination site (P=0.356), fever (P=0.631), headache (P=0.178), fatigue (P=0.429), cough and allergy (P=0.631), abdominal pain (P=0.324), back pain (P=0.054), lethargy (P=0.631), and other symptoms (P=0.452).

In the second dose of the Sinopharm COVID-19 vaccine ( $\leq$ 49 years vs >49 years old), there was more prevalence of side effects than for the first dose (Table 5). Among the ( $\leq$ 49 years vs >49 years old) participants, 14% (15% vs 10%) did not have any post-vaccination symptoms. In both age groups, the most preva-

lent side effects of respondents were pain at the vaccination site (32.6%), fatigue (16.3%), lethargy (13.7%), headache (10%), and tenderness (10%). The same table shows that there was a significant difference between both groups ( $\leq$ 49 years vs >49 years old) in fatigue (*P*=0.003), with no significant differences for other side effects.

#### Relationship between the side effects of vaccination and gender

Table 6 shows the prevalence of side effects post the first dose of Sinopharm COVID-19 vaccination among women and men. Our study included 760 females and 320 males. Women had more symptoms from the first dose than males (17% of females had no side effects vs 45% of males). There was a significant relationship between fatigue and gender (P=0.006); however, the results show no significant difference by gender for tenderness (P=0.194), redness (P=0.356), fever (P=0.891) and headache (P= 0.434).

Table 7 presents the prevalence of side effects from the second dose of vaccination among females and males. Women had more side effects from the second dose than males (11.6% of females had no side effects vs 20% of males). Severe pain at the vaccination site (P=0.027) and fatigue (P=0.011) symptoms were significantly higher in females than males (10.5% vs 2.5% and 20.0% vs 7.5%), respectively. In addition, the table shows no significant difference by gender in the symptoms of normal pain at the vaccination site (P=0.482), tenderness (P=0.368), redness (P=0.834), fever (P=0.279) and lethargy (P=0.053).

### Reasons reported by participants for not receiving a COVID 19 vaccine

The three most common reasons reported by participants for unwelling to receive COVID-19 vaccine were: The vaccine is not effective (6.3%), in a category of people not authorized to take the

Prevalence of general side effects after the second dose Sinopharm COVID-19 vaccination among 2 groups of participants ( $\leq$ 49 years old vs >49 years old) (n 1080)

Side effect	${\leq}49$ years (844) n (%)	>49 years (236) n (%)	Total (1080) n (%)	P-value
Normal pain at the vaccination site	272 (32.2)	80 (33.8)	352 (32.6)	0.646
Severe pain at the vaccination site	56 (6.6)	32 (13.6)	88 (8.15)	0.088
Tenderness	72 (8.5)	36 (15.3)	108 (10)	0.131
Redness	16 (2)	0	16 (1.5)	0.285
Induration and pruritus at the vaccination site	8 (0.95)	4 (1.7)	12 (1.1)	0.631
Fever	16 (2)	16 (6.8)	32 (3)	.051
Headache	92 (11)	16 (6.8)	108 (10)	0.346
Fatigue	108 (13)	68 (28.8)	176 (16.3)	0.003*
Nausea	8 (0.95)	4 (1.7)	12 (1.1)	0.631
Diarrhea	8 (0.95)	0	8 (0.7)	0.452
Cough	4 (0.5)	4 (1.7)	8 (0.7)	0.336
Allergy	0	0	0	-
Muscle pain	40 (4.7)	24 (10)	64 (5.9)	0.121
Abdominal pain	12 (1.4)	4 (1.7)	16 (1.5)	0.881
Back pain	20 (2.4)	12 (5.1)	32 (3)	0.28
Lethargy	104 (12.3)	44 (18.6)	148 (13.7)	0.217
Others	8 (0.95)	0	8 (0.7)	0.595
None	128 (15)	24 (10)	152 (14)	0.323

Table 6

Prevalence of the general side effects after 1st dose Sinopharm COVID-19 Sinopharm COVID-19 vaccination among 2 group of participants (females vs males), n (1080)

Side Effect	Female (760)	Percent	Male (320)	Percent	Total (1080)	Percent	p-value
Normal Pain at the vaccination site	312	41%	144	45.0%	456	42.2%	0.571
Sever pain at the vaccination site	28	3.7%	0	0	28	2.6%	0.081
Tenderness	48	6.3%	8	2.5%	56	5.2%	0.194
Redness	8	1.0%	0	0	8	0.7%	0.356
Induration and pruritus at the vaccination site	12	1.6%	0	0	12	1.0%	0.257
Fever	8	1.1%	4	1.25%	12	1.0%	0.891
Headache	80	10.5%	24	7.5%	104	9.6%	0.434
Fatigue	120	15.8%	12	3.75%	132	12.2%	0.006
Nausea	16	2.1%	0	0	16	1.5%	0.190
Diarrhea	8	1.1%	0	0	8	0.7%	0.356
Cough	12	1.6%	0	0	12	1.0%	0.257
Allergy	8	1.0%	4	1.25%	12	1.0%	0.891
Muscle pain	60	7.9%	8	2.5%	68	6.3%	0.094
Abdominal Pain	20	2.6%	0	0	20	1.8%	0.142
Back pain	44	5.8%	0	0	44	4.1%	0.028
Lethargy	96	12.6%	4	1.25%	100	9.3%	0.003
I did not get any side effect	132	17.0%	144	45.0%	264	24.0%	0.000

Bolded numbers are significant

vaccine (5.2%), and believing that the vaccine has many side effects (4.4%). Participants also reported not having enough time to take the vaccine (3.7%), unavailability of the vaccine (3%), the vaccine not being approved by the WHO (1.5%), being afraid of needles (1.1%), and, the least common reason, waiting for another vaccine (0.4%) (Table 8).

#### Discussion

In August 2020, Trials 1 and 2 of the Sinopharm vaccine were completed and showed that the vaccine triggered a COVID-19 neutralizing antibody response with a low rate of adverse reactions. The most common adverse reactions were pain at the injection site and fever, but all were mild and self-limiting. Moreover, no treatment was required for any side effect (Xia et al., 2021). The UAE were among the first to conduct phase 3 clinical trials of the vaccine, which found the vaccine to have an 86% efficacy rate, according to interim results (Xia et al., 2021).

Most studies have assessed post-vaccination adverse reactions of the Pfizer–BioNTech, Moderna, and AstraZeneca vaccines (El-Shitany et al., 2021, Chapin-Bardales et al., 2021, Kadali et al., 2021, Menni et al., 2021, Riad et al., 2021), while only 2 studies focus on the Sinopharm COVID-19 vaccine (Jayadevan et al., 2021, Hatmal et al., 2021). No published studies focus on Sinopharm COVID-19 vaccine side effects in the UAE to the author's knowledge.

The finding of our study shows that the side effects of this vaccine appear to be mild. A quarter of participants reported they did not have any symptoms post the first vaccination shot while had mild symptoms following vaccination. For the second dose, 14% of participants did not report symptoms; however, the majority had mild and predictable side effects. None of the side effects wre severe or required hospitalization. Our results were in line with a study in India where the frequency of experiencing symptoms following each dose of the vaccine was 24.4% (Sinopharm) (Jayadevan et al., 2021).

For the first dose of vaccination, the findings showed statistically significant differences in the prevalence of severe pain at the injection site, nausea and muscle pain, between the participants aged  $\leq$ 49 vs >49. For the second dose, there was a significant difference between the 2 age groups in the prevalence of fatigue. Moreover, for both groups, the side effects for the second dose were normal pain at the vaccination site (42.2%), fatigue (12.2%), headache (9.6%), lethargy (9.2%), and muscle pain (6.3%). The reported symptoms in our study were similar to symptoms reported in the Phase 1/2 trial of the Sinopharm vaccine; WHO, DOH and MOH in the UAE reported that the most adverse reactions were injection site reactions, headaches and fatigue. They indicated that the most common systematic adverse reaction was pain at the in-

Prevalence of general side effects after the second dose of Sinopharm COVID-19 vaccination among 2 groups of participants (females vs males) (n 1080)

Side effect	Female (760)	Percent	Male (320)	Percent	Total (1080)	Percent	<i>p</i> -value
Normal pain at the vaccination site	264	34.7%	100	31.3%	352	32.6%	0.482
Severe pain at the vaccination site	80	10.5%	8	2.5%	88	8.1%	0.027
Tenderness	84	11%	24	7.5%	108	10%	0.368
Redness	12	1.6%	4	1.25%	16	1.5%	0.834
Induration and pruritus at the vaccination site	12	1.6%	0	0	12	1%	0.257
Fever	28	3.7%	4	1.25%	32	3%	0.279
Headache	80	10%	28	8.75%	108	10%	0.648
Fatigue	152	20%	24	7.5%	176	16.3%	0.011
Nausea	12	1.6%	0	0	12	1.1%	0.257
Diarrhea	8	1.1%	0	0	8	0.7%	0.356
Cough	8	1.1%	0	0	8	0.7%	0.356
Allergy	0	0	0	0	0	0	_
Muscle pain	52	6.8%	12	3.75%	64	6%	0.321
Abdominal pain	12	1.6%	4	1.25%	16	1.5%	0.834
Back pain	28	3.7%	4	1.25%	32	3%	0.279
Lethargy	124	16%	24	7.5%	148	13.7%	0.053
None	88	11.6%	64	20%	152	14.07%	0.072

#### Table 8

Reasons for participants not receiving the COVID 19 vaccine

Reasons	No. (1080)	Percentage
The vaccine is not available	32	3%
The vaccine has many side effects	48	4.4%
I don't believe that the vaccine is effective	68	6.3%
I am afraid of needles	12	1.1%
My body can fight the virus without a vaccine	4	0.4%
Vaccine is not approved by the WHO	16	1.5%
I am from the categories who are not authorized to take the vaccine	56	5.2%
I am waiting for another vaccine	4	0.4%
I don't have time	40	3.7%
Agree to take the vaccine	800	74%

jection site and fever, which were self-limiting and patients recovered; none of the symptoms was severe or required hospitalization [(COVID-19 Vaccine 2021, World Health Organization (WHO) 2021, Xia et al., 2021)].

Injection site pain is reported in several reports on vaccine side effects (El-Shitany et al., 2021, Hatmal et al., 2021 Jayadevan et al., 2021, Riad et al., 2021). Injection into a relaxed muscle leads to less pain compared with a tensed one; therefore, researchers recommend lowering the patient's arm to be injected to reduce pain. In addition, vaccines in situ should be kept at a low temperature; the Sinopharm COVID-19 vaccine should be stored at normal refrigeration temperature. Injection without optimal warming may increase the probability of pain at the injection site (China State-Backed Covid Vaccine Has 86% Efficacy 2021, Hatmal et al., 2021, Riad et al., 2021).

The Food and Health Bureau of Hong Kong evaluation report on CoronaVac (another inactivated virus vaccine) reported that common adverse reactions ( $\geq$ 10%) were injection site pain, headache, and fatigue (Food and Health Bureau (FHB) 2021).

The study of Riad and others among healthcare workers in Turkey found that injection site pain (41.5%), fatigue (23.6%) and headache (18.7%) were reported by more than 10% of the participants; this result was similar to our result for the Sinopharm COVID-19 vaccine (Riad et al., 2021). In our study, the younger adults ( $\leq$ 49 years) were more frequently affected than older age groups (>49 years). Vaccine reactogenicity is known to correlate with transient elevation of inflammatory cytokines, suggesting that the vaccine reactogenicity declines with age, but it is not considered a reliable sign of a desirable immune response (Hervé et al., 2019). Similar results to ours are reported in various COVID-19 vaccine side effects studies (Jayadevan et al., 2021, Polack et al., 2020, Riad et al., 2021). Moreover, according to the Centers for Disease

Control and Prevention, the side effects tend to be more noticeable after the first dose (Possible Side Effects After Getting a COVID-19 Vaccine 2021).

Similar to the findings of recently published studies (El-Shitany et al., 2021, Hatmal et al., 2021), we observed that the frequency of adverse effects to the second shot of vaccine was slightly higher than to the first dose except for nausea (1.5% vs 1.1% [first vs second dose]), allergy (1.1% vs 0.0%), cough (1.1% vs 0.7%), abdominal pain (1.85% vs 1.5%), and back pain (4.1% vs 3.0%). This finding could be interpreted on the basis of immune system response. The immune system could produce cytokines with an inflammatory effect on the blood vessels, muscles and other tissues; it may also produce flu-like symptoms that last for days after vaccination (Zhang et al., 2021).

Women were generally more likely to have side effects from vaccination than men; 83% of females reported side effects compared with 55% of males post first dose of vaccination, 98.5% females versus 80% males post second dose. Previous studies on COVID-19 vaccines have reported more side effects within females compared with males for both doses (El-Shitany et al., 2021, Jayadevan et al., 2021, Riad et al., 2021). According to the findings of the study on CoronaVac, local and systemic side effects were more prevalent after the second dose in females than males, which demonstrates the similar efficacy of this vaccine to Sinopharm (Riad et al., 2021).

The difference of side effects between genders reported for inactivated virus vaccines such as influenza, measles-mumps-rubella combination vaccine, attenuated Japanese encephalitis, and attenuated Dengue vaccines suggest that females have stronger immune responses and side effects are more frequent and more intense (Klein et al., 2010, Klein and Pekosz, 2014). In our study, 1.1% of participants had allergic symptoms post first vaccination dose, while no cases for allergy were reported post second dose. This finding confirms that participants in this study with an allergic reaction to the first dose did not receive the second vaccination dose. There are no published data on the safety of the second dose of COVID-19 vaccine after an allergic reaction to the first dose, al-though one study indicated that anyone with an immediate allergic reaction history of any severity to any component of mRNA COVID-19 vaccines or to polyethylene glycol or polysorbate should not be vaccinated with the Pfizer-BioNTech or Moderna COVID-19 vaccine (Kounis et al., 2021).

Allergic reactions to vaccines not attributed to the active vaccine itself might be caused by inactive ingredients such as egg protein, gelatin, formaldehyde, thimerosal, or neomycin which contribute to specific immunoglobulin E-mediated immediate reactions. According to the European Medicines Agency, excipients are constituents of a medicinal form apart from the active substance; they are inert substances added to vaccines to improve stability, increase solubility, improve absorption, influence palatability, or create a distinctive appearance. Excipients can cause various clinical allergic reactions ranging from skin disorders to life-threatening systemic reactions (Caballero and Quirce, 2020).

Our study noted that the most prevalent chronic conditions among the UAE participants were diabetes (7.80%) and hypertension (6.30%). Our results are consistent with the Dubai Statistical Center study, which shows that UAE nationals have a high prevalence of diabetes and hypertension diseases. Furthermore, another study among patients with chronic disease in the UAE reported that 74.1% of individuals were diabetic (CDC 2020, Osama et al., 2011).

Recent studies have shown that COVID-19 vaccine hesitancy levels vary. In New York, approximately 29% of residents claimed that they will refuse a vaccine, compared with 20% of Canadian residents, and 6% of residents in the United Kingdom. (Latimer, 2020, Henley, 2020).

In our study, a few respondents reported that they did not want to receive the COVID-19 vaccine. the common reasons for indecision and rejection of COVID-19 vaccines were the individuals that the vaccine is ineffective, being of a category of persons not authorized to take the vaccine, and being afraid of vaccine side effects (4.4%). Knowing what to expect post-vaccination will help in public education to dispel myths and lower apprehension about the Sinopharm COVID-19 vaccine. Similar reasons were reported in several studies. For example, in a study among individuals aged >18, the most common reasons for rejection of vaccine were that the participants did not think that the vaccine can be reliable as it is new and that COVID-19 infection is a biological weapon and the vaccine will serve those who produce this virus (Akarsu et al., 2021).

# Conclusion

Fear of the unknown is a driver of vaccine hesitancy. This study showed that first and second dose post-vaccination adverse reactions of Sinopharm COVID-19 vaccine were common side effects that were mild, predictable, non-serious and non-lifethreatening. To our knowledge, this is the first study dealing with the Sinopharm vaccine and evaluating the side effects among a UAE population, the results may help reduce public vaccine hesitancy.

# **Conflicts of interest**

The authors declare no conflict of interest.

# Funding

This research was funded by the University of Sharjah, UAE.

#### **Ethical Approval**

The study was approved by the Research Ethics Committee of the University of Sharjah in the UAE on 22/3/2021, with reference no. REC-21-02-09-07.

#### References

- Akarsu B, Özdemir DC, Baser DA, Aksoy H, Fidancı I, Cankurtaran M. While studies on COVID-19 vaccine is ongoing, the public's thoughts and attitudes to the future COVID-19 vaccine. Int J Clin Pract 2021;75:e13891.
- Aresté N, Salgueira M. World Medical Association Declaration of Helsinki: ethical principles for medical research involving human subjects. JAMA 2013;310(20):2191–4.
- Caballero ML, Quirce S. Delayed Hypersensitivity Reactions Caused by Drug Excipients: A Literature Review. J. Investig. Allergol. Clin. Immunol. 2020;30:400–8.
- CDC. 2020 https://www.cdc.gov/coronavirus/2019-ncov/need-extra-precautions/ people-with-medical-conditions.html
- Chapin-Bardales J, Gee J, Myers T. Reactogenicity Following Receipt of mRNA-Based COVID-19 Vaccines. JAMA 2021;325(21):2201–2. doi:10.1001/jama.2021.5374.
- China State-Backed Covid Vaccine Has 86% Efficacy, UAE Says". Bloomberg.com. 2020-12-09.2021 Retrieved 2020-12-09. https://www. bloomberg.com/news/articles/2020-12-09/uae-says-sinopharm-vaccinehas-86-efficacy-against-covid-19
- China to run human coronavirus vaccine trial in UAE. Reuters. 2020; Jun https://www.reuters.com/article/us-health-coronavirus-china-vaccine/ china-to-run-human-coronavirus-vaccine-trial-in-uae-idUSKBN23U2H8.
- Coronavirus disease (COVID-19): How is it transmitted?. 2021, https://www.who. int/emergencies/diseases/novel-coronavirus-2019/question-and-answers-hub/ q-a-detail/coronavirus-disease-covid-19-how-is-it-transmitted
- COVID-19 Vaccine, Awareness Guide, April, 2021, https://www.dha.gov.ae/Asset% 20Library/COVID19/Covid19\_Vaccine\_EN.pdf
- Cyranoski D. Arab nations first to approve Chinese COVID vaccine despite lack of public data. Nature 2020;588(7839):548.
- Development of an Inactivated Vaccine Candidate, BBIBP-CorV, with Potent Protection against SARS-CoV-2. 2021. Retrieved 1 June 2021, https://dx.doi.org/10.1016%2Fj.cell.2020.06.008
- Elm E, Altman DG, Egger M, Pocock SJ, Gøtzsche C, Vandenbroucke JP. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement: guidelines for reporting observational studies. The Lancet 2007;370(9596):1453–7.
- El-Shitany NA, Harakeh S, Badr-Eldin SM, et al. Minor to Moderate Side Effects of Pfizer-BioNTech COVID-19 Vaccine Among Saudi Residents: A Retrospective Cross-Sectional Stud. International Journal of General Medicine 2021;14:1389–401.
- Food and Health Bureau (FHB). Report on Evaluation of Safety, Efficacy and Quality of CoronaVac COVID-19 Vaccine (Vero Cell) Inactivated. 2021. Available online: https://www.fhb.gov.hk/download/our\_work/health/201200/e\_ evaluation\_report\_CoronaVac.pdf
- Forni G, Mantovani A. COVID-19 Commission of Accademia Nazionale deiLincei, Rome. COVID-19 vaccines: Where we stand and challenges ahead. Cell Death Differ 2021;28:626–39.
- Habas K, Nganwuchu C, Shahzad F, Gopalan R, Haque M, Rahman S, et al. Resolution of coronavirus disease 2019 (COVID-19). ERATCK 2020;18(12):1201–11. doi:10.1080/14787210.2020.1797487.
- Haidere M, Ratan Z, Nowroz S, Zaman S, Jung Y, Hosseinzadeh H, Cho J. COVID-19 Vaccine: Critical Questions with Complicated Answers. Biomol Ther 2021;29(1):1–10. doi:10.4062/biomolther.2020.178.
- Hatmal MM, Al-Hatamleh MAI, Olaimat AN, Hatmal M, Alhaj-Qasem DM, Olaimat TM, Mohamud R. Side Effects and Perceptions Following COVID-19 Vaccination in Jordan: A Randomized, Cross-Sectional Study Implementing Machine Learning for Predicting Severity of Side Effects. Vaccines 2021;9:556.
- Henley J; Guardian correspondents. Coronavirus causing some anti-vaxxers to waver, experts say; 2020. https://www.theguardian.com/world/2020/apr/21/ anti-vaccination-community-divided-how-respond-to-coronavirus-pandemic. Accessed July 27, 2020.
- Hervé C, Laupèze B, Del Giudice G, Giudice GD, et al. The how's and what's of vaccine reactogenicity. npj Vaccines 2019;4(39). doi:<u>10.1038/s41541-019-0132-6</u>.
  Jayadevan R, Shenoy R, Anithadevi TS. Survey of symptoms follow-
- Jayadevan R, Shenoy R, Anithadevi TS. Survey of symptoms following COVID-19 vaccination in India.2021 medRxiv preprint doi: https://doi.org/10.1101/2021.02.08.21251366.
- Kadali RAK, Janagama R, Peruru S, Malayala SV. Side effects of BNT162b2 mRNA COVID-19 vaccine: A randomized, cross-sectional study with detailed self-reported symptoms from healthcare workers. Int. J. Infect. Dis. 2021;106:376–81.
- Klein SL, Jedlicka A, Pekosz A. The Xs and Y of immune responses to viral vaccines. Lancet Infect. Dis. 2010;10:338–49.
- Klein SL, Pekosz A. Sex-based biology and the rational design of influenza vaccination strategies. J. Infect. Dis. 2014;209(Suppl. 3):S114.

- Kounis NG, Koniari I, Gregorio CD, Velissaris D, Petalas K, Brinia A, Assimakopoulos SF, Gogos C, Kouni SN, Kounis GN, Calogiuri G, Ming-Yow Hung M. Allergic Reactions to Current Available COVID-19 Vaccinations: Pathophysiology, Causality, and Therapeutic Considerations. Vaccines 2021;9(3):221.Lai C, Shih T, Ko W, Tang H, Hsueh P. Severe acute respiratory syndrome coron-
- Lai C, Shih T, Ko W, Tang H, Hsueh P. Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and coronavirus disease-2019 (COVID-19): The epidemic and the challenges. Agents 2020;55(3).
- Latimer K. About 20% of people in recent survey said they wouldn't take COVID-19 vaccine; 2020. https://www.cbc.ca/news/canada/saskatchewan/ covid-survey-first-round-results-1.5541053. Accessed July 27, 2020.
- Chugh ML. Interater reliability: the kappa statistic. Biochem Med (Zagreb) 2012;22(3):276–82 PMID: 23092060; PMCID: PMC3900052.
- Menni C, Klaser K, May A, Polidori L, et al. Vaccine side-effects and SARS-CoV-2 infection after vaccination in users of the COVID Symptom Study app in the UK: A prospective observational study. Lancet Infect. Dis. 2021. doi:10.1016/S1473-3099(21)00224-3.
- Nunnally JC, Bernstein IH. The Assessment of Reliability. Psychometric Theory 1994;3:248–92.
- Osama H, Ibrahim M, Jirjees FJ, Mahdi HM. Barriers affecting compliance of patients with chronic diseases: A preliminary study in United Arab Emirates (UAE) population. AJPCR 2011;4(suppl 2).
- Petousis-Harris H. Correction to: Assessing the Safety of COVID-19 Vaccines: A Primer. Drug Saf 2021;44(4):507 -507. doi:10.1007/s40264-020-01023-1.
- Polack F, Thomas S, Kitchin N, et al. Safety and efficacy of the BNT162b2 mRNA covid-19 vaccine. N Engl J Med 2020;383:2603–15. doi:10.1056/NEJMoa2034577.
- Possible Side Effects After Getting a COVID-19 Vaccine 2021. Available online: https: //www.cdc.gov/coronavirus/2019-ncov/vaccines/expect/after.html (accessed on 25 April 2021).
- Riad A, Abdulqader H, Morgado M, Domnori S, Koščík M, Mendes JJ, Klugar M, Kateeb E. Global Prevalence and Drivers of Dental Students' COVID-19 Vaccine Hesitancy. Vaccines 2021;9(6):566.

- Riad A, Pokorná A, Attia S, Klugarová J, Koš<sup>\*</sup>cík M, Klugar M. Prevalence of COVID-19 Vaccine Side Effects among Healthcare Workers in the Czech Republic. 2021. J. Clin. Med. 2021;10:1428.
- Riad A, Sağıroğlu D, Batuhan Üstün B, Pokorná A, Attia S, Klugar M, et al. Prevalence and Risk Factors of CoronaVac Side Effects: An Independent Cross-Sectional Study among Healthcare Workers in Turkey. J. Clin. Med. 2021;10(12):2629.
- Riad A, Sağıroğlu D, Batuhan Üstün B, Pokorná A, Klugarová J, Attia S, Klugar M. Prevalence and Risk Factors of CoronaVac Side Effects: An Independent Cross-Sectional Study among Healthcare Workers in Turkey. J. Clin. Med. 2021;10(12):2629.
- Sharma O, Sultan AA, Ding H, Chris R. A Review of the Progress and Challenges of Developing a Vaccine for COVID-19. Front. Immunol 2020;11.
- Trivedi A, Sharma S, Ashtey B. Investigational treatments for COVID-19. Pharm. J. 2020. doi:<u>10.1211/pj.2020.20208051</u>.
- World Health Organization (WHO): Evidence Assessment: Sinopharm/BBIBP COVID-19 vaccine, for recommendation by the strategic advisory group of experts (sage) on immunization prepared by the sage working group on covid-19 vaccines. 2021
- Xia S, Duan K, Zhang Y, Zhao D, Zhang H, Xie Z, Yang X. Effect of an inactivated vaccine against SARS-CoV-2 on safety and immunogenicity outcomes: Interim analysis of 2 randomized clinical trials. JAMA 2021;324(10):951–60. doi:10.1001/jama.2020.15543.
   Zahid MN, Moosa MS, Perna S, Ebtisam EB. A review on COVID-19 vaccines:
- Zahid MN, Moosa MS, Perna S, Ebtisam EB. A review on COVID-19 vaccines: stages of clinical trials, mode of actions and efficacy. Arab. J. Basic Appl. Sci. 2021;28(1):225–33.
- Zhang Y, Zeng G, Pan H, Li C, Hu Y, Chu K, et al. Safety, tolerability, and immunogenicity of an inactivated SARS-CoV-2 vaccine in healthy adults aged 18–59 years: a randomised, double-blind, placebo-controlled, phase 1/2 clinical trial. Lancet Infect Dis 2021;21(2):181–92. doi:10.1016/s1473-3099(20)30843-4.