Participant INFORMATION SHEET (16-17 years): COV006

Investigating a Vaccine Against COVID-19 in children and teenagers

"A phase 2 study to assess the safety and immunogenicity of a recombinant adenovirus-based vaccine against Coronavirus Disease (COVID-19) in children aged 6-17 years of age"

IMPORTANT: If you develop a fever or cough, shortness of breath or become unwell then you must contact the study team on <<contact details>> for advice before attending any visit. If you cannot get hold of the study team, please consult your GP or use 111.

Participation could really make a difference during a public health emergency.

We would like to invite you to take part in our COVID-19 vaccine study. Before you make a decision, it is important you take the time to understand why we are doing this research and what it would involve. Please read the following information carefully and consider discussing it with friends and relatives.

Who is sponsoring, organising and funding the research?

The study is organised and sponsored by the University of Oxford. The study is funded through financial support to the University of Oxford from the National Institute for Health Research (NIHR), which is a UK government funded research agency, and AstraZeneca. Neither your GP nor the researchers are paid for recruiting you into this study.

What is the purpose of this research study?

The purpose of this study is to test how well children and teenagers respond to one of the COVID-19 vaccines currently being used in adults in the UK.

In the first wave of the pandemic, over 600 children were admitted to hospital with COVID-19 disease in the UK. A small number are also more severely affected by Paediatric Inflammatory Multisystem Syndrome Temporally Associated with SARS-CoV-2 (PIMS-TS), associated with a fever and a generalised inflammatory response in the body. Some children have been critically ill and required intensive care.

The WHO declared the COVID-19 epidemic a Public Health Emergency of International Concern on 30th January 2020 and a pandemic on 11th March 2020. This means that the epidemic is expected to spread to all countries of the world and infect 50-80% of people. Vaccines against COVID-19 are currently being used in the UK for adults, but to date there is no approval to use these vaccines in under 18 year olds.

This study will allow us to assess how well the immune systems of children and teenagers respond to immunisation with the Oxford/AstraZeneca Covid-19 vaccine (also known as *ChAdOx1 nCoV-19*). It will also give us valuable information on safety aspects of the vaccine. The data from this study may be used to support further larger scale trials in children, the results of which may be used by AstraZeneca to support approvals of this vaccine for use in children in the future.

Children from Black, Asian and Minority Ethnic groups (BAME) are particularly welcome to take part in this study.

Summary of the study

In total this study will enrol 300 children and young people across the UK.

- Participants will be randomised to two doses of either the *ChAdOx1 nCoV-19* or a licensed vaccine (MenB) that will be used as a 'control' for comparison. We will not be able to tell you which vaccine you are receiving, because this study is investigating the side-effects associated with the vaccine and we do not want reporting to be biased
- Participants and their parents/guardians will not be told which vaccine course they are receiving
- 5 blood tests will be taken over the course of a year to check if there are any problems and to look at immune responses to the vaccine
- For all participants there will be a diary to complete for up to 28 days following vaccination
- Participants will be enrolled in the study for one year

What is the vaccine we are testing?

The vaccine we are testing in this research study is called *ChAdOx1 nCoV-19*. By vaccinating with ChAdOx1 nCoV-19, we are hoping to make the body recognise and develop an immune response that will help stop the SARS-CoV-2 virus from entering human cells and therefore prevent infection.

Over 20,000 adults have now been recruited to clinical trials of ChAdOx1 nCoV-19 worldwide and widespread roll-out of the vaccine in adults in the UK has now begun.

The ChadOx1-nCOV-19 vaccine was approved for emergency use in adults in the UK in December 2020. This approval was based on data from clinical trials enrolling over 20 000 adults, with early analysis showing the vaccine was 60 – 90% effective at preventing COVID-19. Importantly, these early results showed that no study participants receiving the ChadOx1 nCOV-19 vaccine were admitted to hospital with severe COVID-19 disease. These studies also provided important information on the safety of the vaccine, and the expected side -effects; these are described in more detail later in this document. The vaccine is now being rolled out for routine use, and more information about its effectiveness and safety is being gathered all the time.

The ChAdOx1 nCoV-19 vaccine is being given as two doses in adults, between 4-12 weeks apart. Therefore, this study will enrol you to both a 4 and 12 week (1 and 3 month) boosting schedule.

We are interested in evaluating both boost intervals in the trial so that we can provide the data needed to inform policymakers on how to use the vaccine in children and teenagers.

What is the control (comparison) vaccine, MenB?

In this study we will be using a licensed 'MenB' vaccine against group B meningococcus (MenB), one of the most common causes of meningitis and sepsis in children and teenagers. This will be used as an 'active control' vaccine, to help us understand participants' response to ChAdOx1 nCoV-19. We will be using one of the two versions of MenB, Bexsero®, a vaccine which has been routinely given to infants in the UK since 2015, and is licensed for use in the age groups we are studying here.

Both ChadOx1 nCoV-19 and MenB are given as two dose schedules, administered as injections into the muscle around the shoulder region.

Given we don't expect MenB to offer any protection against COVID-19, by comparing immune responses and post-vaccination symptoms between participants receiving ChAdOx1 nCoV-19 and MenB we will get a better understanding of how well children and teenagers respond to ChAdOx1 nCoV-19.

Do I have to take part?

No. It is up to you to decide whether or not to take part. Your decision will not result in any penalty, or changes to your standard medical care. If you do decide that you will take part, you will be given this information sheet to keep (or be sent it electronically) and will be asked to sign a consent form. We would also provide some written information for you to read. You are free to withdraw at any time and without giving a reason, but we may request a follow up appointment for safety reasons.

Can I take part?

In order to be involved in the study you must:

- Be aged between 16 and 17 years (up to 17 years and 8 months)
- Be able and willing (in the Investigator's opinion) to comply with all study requirements, including the follow up visits (and not rely on public transport to attend the trial centre).

You must:

- Allow the Investigators to discuss your medical history with your GP and access all medical records.
- Provide written informed consent. We recommend that parents are present at the first visit, although this is not compulsory.

Study participants cannot participate in this study if any of the following apply:

- are the child of a staff member of the Oxford Vaccine Group on the delegation log
- have a history of laboratory confirmed COVID-19 or a blood test shows that they have had contact with the COVID-19 virus
- have a history of chronic respiratory diseases (resolved childhood asthma is allowed)
- have previously received a vaccine for Meningitis B
- have any vaccine in the 30 days before or after this study vaccine
- have previously had other similar vaccines that might impact on understanding the study results such as adenovirus vectored vaccines or coronavirus vaccines
- have received immunoglobulins or blood products in the 3 months before having the study vaccine
- history of autoimmune conditions
- have immunosuppression or immunodeficiency
- have a history of angioedema
- have a history of severe allergic reaction (anaphylaxis)
- are have a current diagnosis or are having treatment for cancer
- have a bleeding disorder

- have a history of a congenital heart condition
- any other significant disease which might present increased risk to the participant
- are pregnant, or may become pregnant during the study period
- have previously been diagnosed with Kawasaki disease

Mild conditions that are well-controlled would not automatically exclude you from participating. If you are unclear whether you are eligible to be involved in the study you can contact the study team who will be able to advise you.

What will happen if I decide to take part?

You will firstly be allocated to one of the following groups:

Group	No. of participants (N)	Age	Schedule
1	75	12-17 years	ChAdOx1 nCoV-19 (N=60) OR MenB (N=15) with boost at 1 month
2	75		ChAdOx1 nCoV-19 (N=60) OR MenB (N=15) with boost at 3 months

There are 2 plans depending on which group you are in, shown below.

a) If booster given at 1 month

	Screening questionnaire (up to 90 days before vaccination)	Visit 1 **Day 0**	Visit 2 **1 month**	Visit 3 **2 months**	Visit 4 **6 months**	Visit 5 **1 year**
Medical History	X					
Urine pregnancy test (girls 11 and over)		X	X			
Vaccination		X	X			
Blood Tests		X	X	X	X	X
Diary		X	X	X		

b) If booster given at 84 days

	Screening Questionnaire (up to 90 days before vaccination)	V1 0	V2 3 months	V3 4 months	V4 6 months	V5 1 year
Medical History	X					
Urine pregnancy test (girls 11 and over)		X	X			
Vaccination		X	X			
Blood Tests		X	X	X	X	X
Diary		X	X	X		

If you decide to take part in this trial there is a short online questionnaire to complete to check that you are eligible. At the end of the online questionnaire you will be asked if you agree that a researcher can contact you by phone to ask questions about your current health and discuss details of your medical history, if needed.

If you are eligible, you will be invited to a face-to-face visit where you will be asked to sign the consent form in person and you will receive the vaccination on the same day. COVID-19 safety precautions will be taken throughout this trial to maintain infection control. If you are not eligible, your information will not be stored beyond the end of the trial. Online questionnaire

We will ask you to complete an online questionnaire to establish whether you are eligible for the study, which will include questions about their medical history. It should take no more than 15 minutes. In some cases we may require clarification from your General Practitioner before proceeding and we will ask for your consent to contact them if needed. There will be two parts to the questionnaire, one with general questions and a second part which will ask for your permission to contact your GP.

Consent and Vaccination Visit – *up to 1 hour (Signing of consent form, temperature measured, blood test, vaccination and up to 30-60 minutes observation after the vaccination)*

At the first visit, full written informed consent* will be obtained. You will be given at least 24 hours to read and consider the information in the Participant Information Sheet (this document) before attending for the consent and vaccination visit. You will have opportunity to ask any questions of a member of the research team before you sign a consent form and decide to take part.

We will ask you a few questions to check if there have been any problemsfor you since filling in the screening questionnaire. We will check your temperature and we will take blood samples. We would use an anaesthetic cream to numb the skin for the blood test.

A blood sample will be taken to check if you have had contact with the COVID-19 virus prior to vaccination but the result will not stop you from entering the study. Because of the nature of the pandemic, we expect approximately 10% of participants to have had previous contact with COVID-19 and it will be important to include these participants in our results as well. It is important to note that this is a research test that has not been validated for diagnostic purposes, so results cannot be used to provide confirmation of previous infection nor of protection from future infection. Results will not be provided to participants. You will automatically be required to take a urine pregnancy test, we are asking this of all female participants over the age of 11. We ask that all female participants who are sexually active use an effective form of contraception for the duration of the trial.

You will be randomly allocated to receive two doses of ChAdOx1 nCoV-19 or MenB, and you will not know which one you have had. This will be given as an injection at the top of your arm and we will cover the vaccine site with a dressing. We will need to keep an eye on you to make sure they are OK for 60 minutes (+/- 30 minutes) after the vaccine. Overall the visit will take up to one hour.

We may ask to photograph the vaccination site. You will not be identifiable in these photographs, as only the vaccination site and the unique trial number will be visible. These photographs may be shown to other professional staff, used for educational purposes, or included in a scientific publication.

Electronic Symptom Diary "e-diary" – *Completed at home by participant*

We will give you a thermometer, tape measure and an E-diary account to record your symptoms and temperature every day for 7 days after vaccination. After these 7 days we will ask you to record if you are unwell or take any medications over the next 3 weeks. The research staff will monitor the E-Diary and may phone you to ask for more information. You will also be asked to record in the diary any serious medical illnesses or hospital visits you may have over the course of the study.

Booster and Follow up visits – *1 hour booster visit and 3 follow-up appointments of 30 minutes (blood tests and check for side effects or new health problems)*

At the second visit, you will undergo a second blood test, and the diary card will be reviewed. A physical examination will be done if necessary and after a urinary pregnancy test (if you are female) your second vaccination given.

There are then up to a further 3 follow-up visits over the next 9 months (lasting approx. 30 minutes) to ensure everything is fine and to check your symptoms. Blood tests will be performed as outlined in the table above. Wherever possible, visits will take place outside of school hours (including vaccination and follow-up visits). Social distancing will need to be maintained throughout all visits.

Note: due to the high number of planned volunteers in this study, visits may take longer than the estimates given here.

What should I avoid during the trial?

You should not take part in other studies that involve the administration of drugs or vaccines, including trials testing other interventions for COVID-19. If during the trial you are required to receive any vaccinations while enrolled in this study you should inform the research team beforehand and we will discuss with you the most appropriate time to receive them.

Are there any risks from taking part in the trial?

The risks and side effects of the proposed vaccinations and trial procedures are detailed here:

1. *Blood samples*

Drawing blood may cause slight pain, although we will use anaesthetic cream to numb the skin if requested. Occasionally there can be bruising at the site where the needle enters. You may feel light-headed or even faint. Taking blood can sometimes be difficult and we may ask you for a second attempt if needed. Each blood sample will be a maximum volume of 15 mls.

The blood tests we perform will include antibody testing against COVID-19 and looking at other immune responses to the vaccines. We will not be informing you of your levels of immunity against the COVID-19 virus.

2. *Vaccination Side Effects: ChAdOx1 nCoV-19 and MenB*

It is likely that you will experience some symptoms at the vaccination site as well as general symptoms due to vaccination. Like all medicines, this vaccine can cause side effects, although not everybody gets them. In clinical studies with the vaccine, most side effects were mild to moderate in nature and resolved within a few days with some still present a week after vaccination. If side-effects such as pain and/or fever are troublesome, medicines containing paracetamol can be taken.

Side effects that occurred during clinical trials in adults were as follows:

Very Common (may affect more than 1 in 10 people)

- tenderness, pain, warmth, redness, itching, swelling or bruising where the injection is given
- generally feeling unwell
- feeling tired (fatigue)
- chills or feeling feverish
- headache
- feeling sick (nausea)
- joint pain or muscle ache

Common (may affect up to 1 in 10 people)

- a lump at the injection site
- fever
- being sick (vomiting)
- flu-like symptoms, such as high temperature, sore throat, runny nose, cough and chills

Uncommon (may affect up to 1 in 100 people)

- feeling dizzy
- decreased appetite
- abdominal pain
- enlarged lymph nodes
- excessive sweating, itchy skin or rash

In clinical trials there were very rare reports of events associated with inflammation of the nervous system, which may cause numbness, pins and needles, and/or loss of feeling. However, it is not confirmed whether these events were due to the vaccine.

This study will be collecting data on side-effects in children and teenagers.

There have been recent reports of increased Kawasaki- like disease in children during the COVID-19 pandemic. Kawasaki disease is an illness that causes inflammation in the blood vessels in the body. A

fever that lasts for 5 days or more is the first sign. Since the cause of the COVID-19-associated Kawasaki-like disease is unknown, it is not known if the immune response to the vaccine could also be a trigger or if this event will be more likely to occur in those who have been exposed to the virus after receiving the vaccine.

What are the advantages of taking part?

Interim data from adult studies suggest that ChAdOx1 nCoV-19 has 60-90% efficacy against COVID-19 disease. The information gained from this study will support the licensing of this vaccine in children and young adults, if it is shown to be effective in these age groups as well. Participants who receive MenB will reduce their risk of meningitis and sepsis caused by group B meningococcus.

At the end of the study, if you received the MenB vaccine, you will be offered two doses of ChAdOx1 nCoV-19 vaccine if it is approved for use in your age group and you are not eligible under a national immunisation programme. If you received ChAdOx1 nCoV-19, you will be offered 2 doses of the MenB vaccine.

What should you do if you believe you may have developed COVID-19 during the study?

If you are unwell then contact the NHS 111 service or phone 999 if they are severely unwell.

If you have a positive swab performed in the community or are diagnosed as having COVID-19 disease while in the study then you must contact the study team on [insert phone number]

If you are admitted to hospital during the study then you should inform the medical or nursing staff that you are taking part in this trial. We will provide a contact card for you to give to these staff which will have a link to a website for them to fill in details about your admission.

It is important that you understand that if you become seriously unwell and needs to be admitted to hospital, the standard referral routes within the NHS will be used. Participants will be treated the same way as the general population in this context of the COVID-19 pandemic. We are unable to offer extra medical support outside what is available within the NHS for the general public.

Participants who develop COVID-19 symptoms and have a positive PCR test after the first vaccination can only receive a booster dose after a minimum 4 weeks interval from their first PCR positive test, provided their symptoms have significantly improved. The decision to proceed with booster vaccinations in those cases will be at clinical discretion of the investigators. For participants who are asymptomatic and have a positive PCR test, a minimum of 2 weeks from first PCR positivity will be required before boosting.

Will we be compensated for our travel?

Yes, we will reimburse you £10 for each study visit, to cover travel costs incurred whilst participating in this trial.

What if new information becomes available?

Sometimes during the course of a trial, new information relevant to the trial becomes available. If this happens, we will tell you about it and discuss whether you want to, or should, continue in the study. If you decide you will continue to take part, you will be asked to sign an updated consent form. On receiving new information, we may consider it to be in your best interests to withdraw from the study. Your participation in this study may also be stopped at any time by the study doctor or the Sponsor for other reasons.

What will happen if I don't want to carry on with the trial?

If, at any time after agreeing to participate, you change your mind about being involved with this study you are free to withdraw without giving a reason. If you withdraw we would not usually perform any

more research procedures, although occasionally we might need to offer a follow up visit for safety purposes, for example to check the injection site or a blood result. Your decision will not result in any penalty. Unless you state otherwise, any blood taken whilst you have been in the study will continue to be stored and used for research as detailed above. You are free to request that the blood samples are destroyed at any time during or after the study. If you choose to withdraw from the trial, your standard medical care will not be affected.

What if something goes wrong?

The investigators recognise the important contribution that volunteers make to medical research, and make every effort to ensure your safety and well-being. The University of Oxford, as the research Sponsor, has arrangements in place in the unlikely event that you suffer any harm as a direct consequence of participation in this trial.

In the event of harm being suffered, while the Sponsor will cooperate with any claim, you may wish to seek independent legal advice to ensure that you are properly represented in pursuing any complaint. The study doctor can advise you of further action and refer you to a doctor within the NHS for treatment, if necessary. NHS indemnity operates in respect of the clinical treatment which may be provided if you needed to be admitted to hospital.

Complaints statement

If you wish to complain about any aspect of the way in which you have been approached or treated during the course of this study, you should contact the research investigators who will do their best to address your concerns by sending us an email to INSERT SITE EMAIL. Alternatively you may contact the University of Oxford Clinical Trials and Research Governance (CTRG) office on 01865 616480 or the head of CTRG, email ctrg@admin.ox.ac.uk

Would my taking part in this trial be kept confidential?

All information that is collected about you during the course of the research will be coded with a study number and kept confidential. The information is available to the trial team, authorised collaborators, ethical review committees, INSERT LOCALTrust, government regulatory agencies and the Sponsor (University of Oxford), who can ask to access the trial data. Responsible independent monitors may be given access to data for monitoring and/or audit of the trial to ensure we are complying with regulations. They are bound by the same confidentiality rules.

Every effort will be taken to maintain confidentiality. Information about you may be stored electronically on a secure server, and paper notes will be kept in a key-locked filing cabinet at the INSERT SITE NAME or at the Centre for Clinical Vaccinology and Tropical Medicine (CCVTM), University of Oxford. Trial results will be published in a scientific journal but nothing that could identify you will be included in any report or publication.

What will happen to my data?

UK Data protection regulation requires that we state the legal basis for processing information about you. In the case of research, this is 'a task in the public interest.' The University of Oxford is the sponsor for this study, based in the United Kingdom, is the data controller and is responsible for looking after your child's information and using it properly.

We will be using information from your medical records in order to undertake this study and will use the minimum personally-identifiable information possible. We will keep identifiable information about you such as contact details for a minimum of 5 years and until the youngest participant turns 21 years. The need to store this information for longer in relation to licensing of the vaccine will be subject to ongoing review. De-identified research data will be stored indefinitely. If you have agreed that samples can be retained for future research then your personally identifiable information will be

kept with restricted access solely for the purposes of sample management for a minimum of five years after the last sample has been either used or disposed of in order to meet regulatory requirements. We will also store your consent form. Samples will be provided for future research only in a form that does not identify you. We store research data securely at the University of Oxford indefinitely following removal of identifiable information. If you agree to your details being held to be contacted regarding future research, we will retain a copy of the consent form until such time as your details are removed from our database but will keep the consent form and your details separate.

The study team will use your name and contact details, to contact you about the research study, and make sure that relevant information about the study is recorded for your care, in relation to your health during the study and to oversee the quality of the study. At the completion of the study, unless you consent otherwise (e.g. if you request to be informed of other trials), your personal details will not be used to contact you other than exceptional circumstances concerning your safety. If you consent to take part in another study carried out by the INSERT SITE NAME, personal information and medical information including blood test results may be accessed to avoid unnecessary repetition.

Data protection regulation provides you with control over your personal data and how it is used. When you agree to your information being used in research, however, some of those rights may be limited in order for the research to be reliable and accurate. Further information about your rights with respect to your personal data is available at: https://compliance.web.ox.ac.uk/individual-rights

Safety data will be provided to AstraZeneca, in an anonymised format.

Involvement of the General Practitioner (GP)/Family doctor (GP)

In order to enrol into this study, you will be required to tick a box on an online form to say that you consent for us to contact your GP. Your \GPmay be asked to share information about your medical history and give access to any other medical records as required. We will write to your GP to let them know about your enrolment and study completion status, so they can update your medical records accordingly.

What will happen to any samples I give?

If you consent, some of your leftover blood samples can be stored and used for future infectious disease or vaccine related research. This is optional, your participation in this study will not be affected by your decision whether to allow storage and future use of leftover samples. Upon request at any time, your remaining blood samples will be destroyed.

Study blood tests to look at the response of your body to the vaccine will be done both at the University of Oxford and with collaborating laboratories in the UK and in other countries. Any samples or data sent to them would not include information that identifies you.

Will any genetic tests be done?

We may do genetic tests on your blood samples to look at the patterns of genes that regulate your own individual immune response (these are called Human Leukocyte Antigen genes). Doing this helps us to work out which aspects of the immune response to vaccines are due to genetic differences between individuals. We may also look at the expression of certain genes which relate specifically to the immune response to COVID-19, but no genetic tests concerning diseases or conditions other than COVID-19 and other vaccine related responses. Any samples and information recorded will be de-identified, so that we cannot directly identify you. However, DNA is unique, and as such, will never be completely anonymous.

What will happen to the results of the research study?

The results of this research study may be presented at scientific meetings or conferences and published in a scientific medical journal. This may not happen until 1 or 2 years after the study is completed. A copy of the results will be made available to you after the study. You will not be identified in any report or publication.

The de-identified data from this study will be shared with the collaborating partners who are organising and funding this research work. Data from this study may be used to file patents, licence vaccines in the future or make profits in other ways. You will not be paid for any part of this. Data from this study may be used as part of a student post-graduate degree, for example a MD or PhD.

Taking part in future vaccine-related research

With your consent, we would like to keep your contact details after your participation in this study is complete, so we may inform you of opportunities to participate in future vaccine related research. This is entirely optional and your participation in this study will not be affected by your decision to allow or not allow storage of your contact details beyond your participation in this trial. Your details will be stored electronically on a secure server and only authorised individuals at the INSERT SITE NAME will have access to it.

We will not, under any circumstances, share your contact details with any third party institutions without your permission. Being contacted does not oblige you to agree to take part in future research and you can ask us to have your contact details removed from our database at any time.

Who has reviewed the study?

This study has been reviewed by the NHS Research Ethics Service (RES) – Insert name of REC and has been given a favourable ethical opinion. The Medicines and Healthcare products Regulatory Agency (MHRA), which regulates the use of all medicines in the UK, has reviewed the study design and has granted permission to use this unlicensed vaccine in this clinical study.

Further information and contact details

We hope this information sheet has answered all of your questions. If you would like further information about participating in research please visit the following website: http://www.nhs.uk/conditions/Clinical-trials/Pages/Introduction.aspx. For independent advice about participating in this trial you may wish to contact your GP. If you would like to speak to one of our team members to discuss any aspect of this trial or **if you are interested in taking part in the study, please contact us**:

<<Insert Site recruitment contact details (address, email, phone)>>