

A trial of surgery versus no surgery in patients with severe pectus excavatum: looking at improvements in heart and lung function

Submission date 26/04/2024	Recruitment status Recruiting	<input checked="" type="checkbox"/> Prospectively registered <input type="checkbox"/> Protocol
Registration date 30/04/2024	Overall study status Ongoing	<input type="checkbox"/> Statistical analysis plan <input type="checkbox"/> Results
Last Edited 04/06/2024	Condition category Surgery	<input type="checkbox"/> Individual participant data <input type="checkbox"/> Record updated in last year

Plain English summary of protocol

Background and study aims

Pectus Excavatum (PE), also known as funnel chest, is a condition where the ribs and breastbone grow inwards forming a dent in the chest. It can be found in between 1 in 400 and 1 in 1000 people. People with severe PE, where the space between the spine and the chest is limited, can experience symptoms such as breathlessness, dizziness, fainting and pain with exercising. This can be very restrictive in daily life.

Treatment for PE includes surgery, which lifts the sternum up, relieving these symptoms. There are no new or “experimental” procedures being tested in this study: the two types of surgeries called the Nuss and Ravitch procedures are both well-established and regularly performed in patients across the world.

In 2019 the NHS England decommissioned pectus surgery, although it is still fully-funded in the devolved nations. This decision was based on the lack of high-quality comparative data showing an improvement in physical health or heart-lung function. More recently in 2023, NHS England have started funding surgery for only the most severe cases, as assessed by a national expert Multi-Disciplinary Team (MDT), which are then performed at two hospitals in England.

The purpose of this study is to see how surgery to treat PE affects a participant’s ability to be physically active. We will recruit 300 participants overall and compare the risks and benefits of surgery against no surgery as measured by a change in physical health after a year. We will also look at how much it costs the NHS overall.

Who can participate?

Patients with pectus excavatum aged 12 years or older.

What does the study involve?

Participants will have a series of data collected when they join the trial, most data should be available from their medical records, additionally, they will need to complete a number of questionnaires. If they have consented to the trial, then they will be allocated randomly (by a

computer) to either the early or the delayed surgery arms. If they are part of the observational study, they will have the surgery as already planned.

Participants will then be followed initially for up to approximately 3 years post-surgery (at 6 months, 1 year and 3 years after surgery, and for the delayed-surgery group also at 6 months and 1 year after their allocation) and, if further funding is available, then for up to 5 years after surgery. At these follow-ups, participants will be asked about their current status, medications, any hospitalisations, adverse events and other procedures they might have undergone since their last contact point, additionally they will be asked to complete the same questionnaires as at baseline. Participants will also be asked to undergo exercise tests to measure their heart-lung function at 1 and 3 years after surgery, and for the delayed-surgery group also at 1 year after their allocation and before their surgery.

There are two types of surgery, one called Nuss and one called Ravitch. Both types are included in this study, and the surgeon will discuss them with the participant to help decide the best option. Both surgical procedures are well-established practices for the treatment of PE. As such, there are routine care practices to be followed.

What are the possible benefits and risks of participating?

The main benefit for participants in England in groups 1 and 2 is that being part of the study enables access to surgery that may not be available to them as part of NHS care; and it is expected that the surgery will improve their condition. However, there are risks associated with the surgery, including prolonged pain, infection, and complications of the procedure. In the routine surgical follow-ups, there is also the need for a CT scan, which involves a low dose of radiation; with any radiation exposure there is the risk that it may cause cancer many years later. Additional risks and burdens as research participants include the potential loss of confidentiality, as we will collect a lot of personal data, although we will record most of this against a code. Some people will also feel uncomfortable when answering questions about their health or healthcare.

Where is the study run from?

South Tees Hospitals NHS Foundation Trust (UK)

When is the study starting and how long is it expected to run for?

February 2024 to May 2030

Who is funding the study?

National Institute for Health and Care Research (NIHR) (UK).

Who is the main contact?

RESTORE trial team, Dr Lisa Chang
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Contact information

Type(s)

Public

Contact name

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Additional identifiers**EudraCT/CTIS number**

Nil known

IRAS number

331910

ClinicalTrials.gov number

Nil known

Secondary identifying numbers

CPMS 57603, NIHR158749, IRAS 331910

Study information**Scientific Title**

A randomised trial of surgery versus no treatment to RESTORE cardiopulmonary function in severe pectus excavatum

Acronym

RESTORE

Study objectives

Surgery improves physical functioning in patients with pectus excavatum, as measured at one-year.

Ethics approval required

Ethics approval required

Ethics approval(s)

Approved 03/06/2024, East of Scotland Research Ethics Service (Tayside Medical Science Centre, George Pirie Way, Ninewells Hospital, Dundee, DD1 9SY, United Kingdom; +44 1382 383871; tay.eosres@nhs.scot), ref: 24/ES/0034

Study design

Interventional randomized controlled trial with observational cohort

Primary study design

Interventional

Secondary study design

Randomised controlled trial

Study setting(s)

Hospital

Study type(s)

Treatment

Participant information sheet

To follow

Health condition(s) or problem(s) studied

Cardiothoracic surgery, pectus excavatum

Interventions

If a potential participant consents to the randomised trial, a series of screening and baseline assessments will be conducted and recorded. These include capturing data from routine care on medical history, physical exercise and lung function tests, existing scans and images that have been performed to diagnose their condition and the severity of it. In addition to capturing this routine data, participants will be asked to complete a series of questionnaires. Where the patients have had an exercise test some time ago (more than 12 months' ago for people 16 years and over, and more than 6 months' ago for people under 16 years) we will ask them to do a repeat test, as their condition might have changed.

If meeting the eligibility criteria, participants will be allocated (randomised) into one of two groups: to have early surgery (within 3 months of randomisation), or to have surgery after a 12-month delay.

For the late-surgery group, they will be contacted at 6 and 12 months after randomisation to ask about their health and complete the same questionnaires they had at baseline. At 12-months they will also be asked to repeat the exercise test.

For all groups, after surgery, as part of their routine care, it is recommended in routine care that a low-dose CT scan is performed to measure the impact of the surgery, this takes place in a timeframe of up to 6 months after surgery and we will collect this data if it is available. At 6 and 12 months after surgery, all participants will be contacted to ask about their health and complete the same questionnaires as at baseline. At 12-months after surgery they will also be asked to repeat the exercise test.

If patients undergo the Nuss surgical procedure (which involves the insertion of bars into their chest), then at any time from 2.5 - 3 years after surgery they will have the bars removed. These participants will be contacted approximately 6 months after the bars are removed to ask about their health and complete the same questionnaires. They will also be asked to repeat the exercise test.

If patients undergo the Ravitch surgical procedure, they will be contacted 3 years after surgery to ask about their health, complete the same questionnaires and to repeat the exercise test.

Intervention Type

Procedure/Surgery

Primary outcome measure

1. SF-36v2 Physical Function score change between testing before randomisation and 1 year later (control group) or 1 year after surgery (experimental group).
2. Incremental cost per QALY at 1 year. QALY assessed through quality of life questionnaires (EQ-5D-5L, SF-36v2, HADS) completed at baseline and 1 year.

Secondary outcome measures

1. Measures of cardiopulmonary function from cardiopulmonary exercise tests (CPET), at baseline, 1 year (post-surgery for all groups and also post-randomisation for the late surgery group), 3 years post-surgery.
2. Quality of life measures including those that of impact on mental well-being (EQ-5D-5L and SF36v2 mental component scores, and HADS), at baseline, 6 months, 1 year (post-surgery for all groups and also post-randomisation for the late surgery group), 3 years post-surgery.
3. Symptoms measured by Nuss and Phoenix Comprehensive Assessment of Pectus Excavatum Symptoms (PCAPES) Questionnaires, at baseline, 6 months, 1 year (post-surgery for all groups and also post-randomisation for the late surgery group), 3 years post-surgery.
4. Body image measured by the Body Image Disturbance Questionnaire (BIDQ), at baseline, 6 months, 1 year (post-surgery for all groups and also post-randomisation for the late surgery group), 3 years post-surgery.
5. Need for revision surgery (complications including the need for unplanned redo surgery and syncope), data collected across the duration of the patient's participation in the study (up to 3 years).
6. Adverse events of special interest, data collected across the duration of the patient's participation in the study (up to 3 years).
7. Major surgical complications, data collected across the duration of the patient's participation in the study (up to 3 years).
8. Costs to the NHS and patients at 1 year and over the patients' lifetime, measured by Health Resource Usage Questionnaire.

9. QALYs at 1 year and over the patients' lifetime, measured by quality of life questionnaires (EQ-5D-5L, SF-36v2, HADS).
10. Incremental cost per QALY gained over the patients' lifetime, measured by quality of life questionnaires and healthcare resource utilisation.
11. Patient preference measured by a discrete choice experiment, to be conducted at month 16 post-randomisation.
12. Net benefit of the intervention, measured by a combination of outcome measures listed above.

Overall study start date

01/02/2024

Completion date

31/05/2030

Eligibility

Key inclusion criteria

For inclusion in the randomised trial:

1. ≥ 12 years old.
2. A PE deformity with a Haller Index of >3.25 , as measured by the internal width of the chest measured at the widest point divided by the distance from the back of the sternum to the anterior vertebral body at its minimum point on CT scan.
3. The participant must satisfy at least one of the following criteria:
 - 3.1. Significant level of shortness of breath or exercise ability perceived to be below that of their peers (e.g., limited by vigorous activities such as running or lifting heavy objects).
 - 3.2. Presyncope or syncope on exercise.
 - 3.3. Arrhythmias on ECG that may be due to the pectus abnormality.
 - 3.4. Dysphagia or swallowing abnormalities in the absence of any other cause.
4. Provide informed consent/assent and agree to 5 years of follow up.
5. Fit to undergo surgery.

For inclusion in the observational cohort:

1. ≥ 12 years old.
2. Confirmed as eligible and fit for surgery via the national MDT surgical pathway.
3. Providing informed consent to take part in the embedded observational cohort and agreeing to 5 years follow up.

Participant type(s)

Patient

Age group

Mixed

Lower age limit

12 Years

Sex

Both

Target number of participants

Planned Sample Size: 300; UK Sample Size: 300

Key exclusion criteria

For the randomised trial:

1. Patients not fulfilling the inclusion criteria.
2. Symptoms relating to causes other than PE.
3. Unwilling to have surgery for PE.

For the observational cohort:

1. Patients not fulfilling the inclusion criteria.
2. Patients who are willing to join the full randomised trial group of the RESTORE Trial.

Date of first enrolment

01/08/2024

Date of final enrolment

31/08/2026

Locations

Countries of recruitment

England

United Kingdom

Study participating centre

South Tees Hospitals NHS Foundation Trust

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TS4 3BW

Study participating centre

Alder Hey Children's NHS Foundation Trust

Alder Hey Hospital
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Study participating centre

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Study participating centre

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E1 2ES

Study participating centre

University Hospitals Birmingham NHS Foundation Trust

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B15 2GW

Study participating centre

Leeds Teaching Hospitals NHS Trust

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Study participating centre

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Study participating centre

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Study participating centre
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Sponsor information

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Sponsor type
Hospital/treatment centre

Website
<http://southtees.nhs.uk/>

ROR
<https://ror.org/02js17r36>

Funder(s)

Funder type
Government

Funder Name

NIHR Evaluation, Trials and Studies Co-ordinating Centre (NETSCC)

Results and Publications

Publication and dissemination plan

The findings of the study will be disseminated to stakeholders for the RESTORE research, including patients, NHS England and the clinical community nationally and internationally. The Patient Advisory Group and lay co-applicants will work with the trial team to inform the dissemination plans to patients and the public.

Findings will be reported to the funder, Society for Cardiothoracic Surgery (SCTS), the writing committee of the European Society of Thoracic Surgery (ESTS), National Institute for Health and Care Excellence (NICE) and presented at conferences including the European Society of Thoracic Surgeons and the Chest Wall International Group.

The Chief Investigator and co-Chief Investigator will be responsible for ensuring the results of RESTORE are disseminated through peer reviewed journals once the analyses are complete.

Intention to publish date

31/12/2028

Individual participant data (IPD) sharing plan

The datasets generated during and/or analysed during the current study are available from the corresponding author on reasonable request

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Aggregated coded datasets will be available to be shared upon request, following publication of the main findings. The data will be available, as described in the participant information sheet, and according to the consent provided by the participant. Requesters must provide an outline of their analysis plans and agree to adhere to UK data privacy laws as a minimum, access will be granted after consideration of the plan by the study chief investigator, trials unit and sponsor.

IPD sharing plan summary

Data sharing statement to be made available at a later date