

A trial to determine if a special liquid diet (exclusive enteral nutrition) before Crohn's disease surgery improves recovery

Submission date 25/07/2023	Recruitment status No longer recruiting	<input checked="" type="checkbox"/> Prospectively registered <input type="checkbox"/> Protocol
Registration date 10/08/2023	Overall study status Ongoing	<input type="checkbox"/> Statistical analysis plan <input type="checkbox"/> Results
Last Edited 19/04/2024	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

Crohn's disease is a lifelong inflammatory illness that causes people to have severe stomach pains, chronic diarrhoea, and suffer weight loss. There is no cure. Crohn's disease causes inflammation, ulceration, bleeding and narrowing of the digestive system. People can have periods of good health (remission) and times when symptoms are more active (flare-ups or relapses). Medications can help keep it in remission, however, one-third of people will need surgery to remove/repair part of their diseased gut at some stage.

Previous studies suggest that a special liquid diet might help improve recovery from surgery. This diet is called exclusive enteral nutrition (EEN) because it is the only form of food taken for a period of time. In active Crohn's disease, this special liquid diet can improve symptoms, reduce inflammation and heal the gut better than steroids. EEN is not offered routinely to patients at the moment because we don't have any evidence-based research.

This study aims to find out whether 6 weeks of EEN diet pre-surgery might help participants recover quicker and make the surgery safer with less chance of complications compared to a usual diet pre-surgery.

Who can participate?

Patients aged 16 years and over from across the UK due to have surgery for Crohn's disease

What does the study involve?

Participants are randomly allocated to a minimum of 6 weeks of EEN pre-operatively or the usual diet. All patients will undergo surgery and be followed up for 1 year with information about disease activity, medication use, health-related quality of life and health resource usage is obtained. An embedded study will explore patients' and healthcare professionals' views and experiences of all aspects of the study. There is an optional sub-study which involves a more detailed assessment of outcomes related to early/late surgery and dietary aspects in terms of nutrient intake, dietary habits and food-related quality-of-life. The follow-up schedule is designed to collect data only at time points comparable to common clinical practice to reduce patient burden and the need to attend additional hospital appointments.

What are the possible benefits and risks of participating?

If the liquid diet can be shown to improve symptoms, widespread uptake is anticipated.

Participants in both groups will be undergoing elective surgery as part of their standard medical care. The trial intervention is purely nutritional, with the main risk relating to poor tolerance of the EEN. This could mean that some patients in the EEN arm may not meet their nutritional requirements. However, this risk will be mitigated by having close dietitian follow-up and by not mandating a particular type of EEN to be used within the study. Hospitals have more than one type of EEN available. Participants will be allowed to choose from the EEN available at each site until they can find one that they can tolerate. They can also use a variety of EEN supplements to minimise taste fatigue. Participants will be made aware of the risks and benefits during the consent process and written information will be provided in the patient Information Sheet. The participants' GPs will also be informed that their patients are participating in a clinical trial and of the allocated treatment.

Where is the study run from?

Birmingham Clinical Trials Unit based at the University of Birmingham (UK)

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

May 2022 to April 2026

Who is funding the study?

National Institute for Health and Care Research (NIHR) Health Technology Assessment (HTA) (UK)

Who is the main contact?

ocean@trials.bham.ac.uk

Study website

<https://www.birmingham.ac.uk/ocean>

Contact information

Type(s)

Public

Contact name

Miss Pooja Gaddu

Contact details

Birmingham Clinical Trials Unit (BCTU)

Public Health Building

University of Birmingham

Edgbaston

Birmingham

United Kingdom

B15 2TT

+44 (0)121 415 9120

ocean@trials.bham.ac.uk

Additional identifiers

EudraCT/CTIS number

Nil known

IRAS number

325763

ClinicalTrials.gov number

Nil known

Secondary identifying numbers

2.0, IRAS 325763, CPMS 56722

Study information

Scientific Title

Optimisation before Crohn's surgery using exclusive enteral nutrition

Acronym

OCEaN

Study objectives

To determine if pre-operative exclusive enteral nutrition (EEN) is more clinically and cost-effective compared with usual diet in patients undergoing surgery for Crohn's disease (CD).

Ethics approval required

Ethics approval required

Ethics approval(s)

Approved 11/07/2023, London City & East REC (Research Ethics Committee Centre, 2nd Floor, 2 Redman Place, Stratford, London, E20 1JQ, United Kingdom; +44 (0)207 1048171; cityandeast.rec@hra.nhs.uk), ref: 23/LO/0513

Study design

Multi-centre two-arm parallel-group open-label pragmatic randomized controlled trial with a mixed methods internal pilot

Primary study design

Interventional

Secondary study design

Randomised controlled trial

Study setting(s)

Hospital

Study type(s)

Treatment

Participant information sheet

Not available in web format

Health condition(s) or problem(s) studied

Planned surgery for small bowel and/or colonic Crohn's disease (primary or repeat surgery)

Interventions

The intervention is a minimum of 6 weeks of EEN pre-operatively. The control group is the usual diet.

Following randomisation into the trial, participants will be reviewed by the local dietitian, and their feed will be prescribed. The type of EEN formula prescribed to each participant will be decided by dietitian preference and/or local availability. If participants cannot tolerate the first feed prescribed, an alternate EEN formula can be used, or participants can use a variety of EEN feeds to reduce taste fatigue. Feeds may be concentrated to reduce the volume of feeds consumed. If tolerance remains poor, nasogastric feeding can be offered.

The start date of EEN will be agreed between the dietitian and the participant. An online support tool (FutureLearn Ltd), developed with the patient panel and Crohn's and Colitis UK, will provide information, peer support and assistance to participants randomised to EEN, which will also help facilitate adherence to an EEN diet. The OCEaN trial team have developed some EEN diet patient support sheets to help support patients on the liquid diet, as well as standardise the delivery of the EEN across the sites as much as possible. EEN is often started by the hospital dietitian and then continued by either the hospital or provision is transferred to the community dietitians and/or the GP. This pathway is likely to vary between sites, and so will be directed by local practice and local resources.

Intervention Type

Supplement

Primary outcome measure

Dual primary outcomes at 6 weeks post-surgery:

1. Crohn's Life Impact Questionnaire (CLIQ; a CD-specific patient reported outcome assessing QoL)
2. Post-surgery complications using the Comprehensive Complication Index (CCI)

If we demonstrate benefit for EEN on either of the primary outcomes this establishes effectiveness

Secondary outcome measures

Patient reported:

1. QoL over time using the CLIQ which will be collected post-surgery, fortnightly until 12 weeks post-surgery and then monthly up to 24 weeks post-surgery
2. Post-surgery recovery using the Surgical Quality of Recovery-15 (QoR-15) on day 3 post-surgery (or pre-discharge if discharged before day 3). The QoR-15 is a short form version of the QoR-40 and consists of 15 items each with a numerical rating score of 0-10. The total score therefore ranges from 0 to 150, with higher scores indicating better quality of recovery.

Clinical:

3. Length of post-operative hospital stay (measured in nights in hospital)
4. Length of bowel resected (in centimetres measured along anti-mesenteric border) at time of surgery before being put in formalin
5. Number of anastomoses formed at surgery as documented in the operation notes or from discussion with the operating surgeon

6. Number of participants requiring stoma formation either at index operation or within 30 days of surgery due to re-operation
7. Number of participants who develop an anastomotic leak (either radiological concern or confirmed at reoperation) within 30 days of surgery
8. Number of participants re-admitted within 30 days of date of discharge
9. Number of participants requiring re-operation within 30 days of surgery
10. Number of participants who develop enterocutaneous fistulae within 90 days of surgery. This is defined as any new fistula tract from any point in the gastrointestinal tract opening on to skin at any site from day of surgery to 90 days later. Enterocutaneous fistula diagnosis can be confirmed following clinical assessment. Radiological confirmation is not required.
11. Number of participants who develop clinical recurrence of their CD at 24 and 52 weeks post-surgery as assessed by the Crohn's Disease Activity Index (CDAI)
12. Number of participants who develop endoscopic disease recurrence assessed endoscopically on colonoscopy performed between 24 and 52 weeks post-surgery as part of standard of care. (66, 67) Modified Rutgeert's score should be used to grade the severity of recurrence. (67) If endoscopy is not performed but patient has cross sectional imaging (e.g. Magnetic resonance imaging (MRI), Computerised Tomography (CT) or ultrasound) as part of standard of care (routine assessments), the presence or absence of disease recurrence on imaging can be used as an alternative.
13. Number of participants on steroids at baseline who were able to wean off steroids prior to surgery. Steroid usage and dosage (including prednisolone (oral/rectal), Budesonide (oral /rectal), Hydrocortisone (intravenous/oral)) will be recorded at baseline and on day of surgery to determine change in use or dose. Inhaled or topical steroid use is not relevant to this trial.
14. Safety assessed through adverse event and serious adverse event reporting
15. Number of participants whose planned surgery did not proceed due to clinical improvement. If planned surgery is cancelled by the local team because clinical improvement deems it no longer necessary, these participants will continue to follow the trial protocol (e.g., follow-up assessments etc.)
16. Number of participants who required expedited or emergency surgery. This refers to participants whose elective/planned CD surgery date is brought forward due to clinical deterioration.

Economic outcomes:

17. The EuroQoL-5D-5 Level (EQ-5D-5L) questionnaire and an incremental cost-utility analysis will determine the cost per quality-adjusted life year (QALY) gained over the 52 weeks post-surgery. The EQ-5D-5L will be collected pre-operatively, and then at 6 weeks post-operatively, and then also at 24 and 52 weeks post operatively.

Qualitative outcomes:

18. Interviews will be undertaken with participants randomised to both trial groups, and also with staff involved in the trial. This research aims to provide in-depth qualitative data concerning the acceptability and experience of EEN as a pre-surgical intervention.

Exploratory outcomes:

19. Change in the Harvey-Bradshaw Index (HBI) and CLIQ between baseline and pre-operatively to determine if EEN improves HBI and CLIQ.
20. To determine if baseline microbiome compositional and metabolomic signatures, and changes after 6-weeks of EEN, can predict primary and secondary trial outcomes including post-surgical complications and likelihood of disease recurrence at follow-up. The samples are being collected as part of the main trial in order to measure faecal calprotectin. Further analysis will be performed and funded separately by UoG.

COAST sub-study outcomes:

21. COAST will analyse the subset of participants who are having surgery for terminal ileal CD to determine the impact of surgical timing on QoL and cost.

Microbiome analysis outcomes:

22. As an exploratory trial outcome, we will study if baseline microbiome compositional and metabolomic signatures, and changes after 6-weeks of EEN, can predict primary and secondary trial outcomes including post-surgical complications and likelihood of disease recurrence at follow-up.

Overall study start date

01/05/2022

Completion date

01/04/2026

Eligibility

Key inclusion criteria

1. Any patient undergoing planned surgery for small bowel and/or colonic CD (primary or repeat surgery)
2. Age ≥ 16 years
3. Willingness to go on EEN for the duration of the intervention period (minimum of 6 weeks)
4. Capacity to give consent

Participant type(s)

Patient

Age group

Adult

Lower age limit

16 Years

Upper age limit

99 Years

Sex

Both

Target number of participants

618

Key exclusion criteria

1. Surgery for peri-anal CD, ulcerative colitis, or inflammatory bowel disease unclassified (IBDU)
2. Patients who require parenteral nutrition in the 6 weeks prior to surgery
3. Inability to comply with the trial schedule and follow up

Date of first enrolment

11/04/2024

Date of final enrolment

31/01/2025

Locations**Countries of recruitment**

England

Northern Ireland

Scotland

United Kingdom

Wales

Study participating centre**Dorset County Hospital**

Williams Avenue

DORCHESTER

United Kingdom

DT1 2JY

Study participating centre**Barts Health NHS Trust**

The Royal London Hospital

80 Newark Street

London

United Kingdom

E1 2ES

Study participating centre**County Durham and Darlington NHS Foundation Trust**

Darlington Memorial Hospital

Hollyhurst Road

Darlington

United Kingdom

DL3 6HX

Study participating centre**Imperial College Healthcare NHS Trust**

The Bays

St Marys Hospital

South Wharf Road
London
United Kingdom
W2 1BL

Study participating centre
North Tees and Hartlepool NHS Foundation Trust
University Hospital of Hartlepool
Holdforth Road
Hartlepool
United Kingdom
TS24 9AH

Study participating centre
University College London Hospitals NHS Foundation Trust
250 Euston Road
London
United Kingdom
NW1 2PG

Study participating centre
Liverpool University Hospitals NHS Foundation Trust
Prescot Street
Liverpool
United Kingdom
L7 8XP

Study participating centre
Maidstone and Turnbridge Wells NHS Trust
Hermitage Lane
Maidstone
United Kingdom
ME16 9QQ

Study participating centre
The Newcastle upon Tyne Hospitals NHS Foundation Trust
Freeman Road
Newcastle upon Tyne
United Kingdom
NE7 7DN

Study participating centre
Lewisham and Greenwich NHS Trust
Lewisham High Street
LONDON
United Kingdom
SE13 6LH

Study participating centre
Royal Cornwall Hospitals NHS Trust
Royal Cornwall Hospital
Treliske
Truro
United Kingdom
TR1 3LJ

Study participating centre
Dartford and Gravesham NHS Trust
Darenth Wood Road
DARTFORD
United Kingdom
DA2 8DA

Study participating centre
Croydon Health Services NHS Trust
530 London Road
Thornton Heath
United Kingdom
CR7 7YE

Study participating centre
St Marks Bowel Cancer Screening Centre
Watford Road
Harrow
United Kingdom
HA1 3UJ

Study participating centre
King's College Hospital NHS Foundation Trust
Denmark Hill

LONDON
United Kingdom
SE5 9RS

Study participating centre
Sheffield Teaching Hospitals NHS Foundation Trust
Herries Road
Sheffield
United Kingdom
S5 7AU

Study participating centre
Manchester University NHS Foundation Trust
Cobbett House
Oxford Road
Manchester
United Kingdom
M13 9WL

Study participating centre
Leeds Teaching Hospitals NHS Trust
Beckett Street
LEEDS
United Kingdom
LS9 7TF

Study participating centre
University Hospitals Plymouth NHS Trust
Derriford Road
PLYMOUTH
United Kingdom
PL6 8DH

Study participating centre
The Royal Wolverhampton NHS Trust
New Cross Hospital
Wolverhampton Road
Heath Town
Wolverhampton
United Kingdom
WV10 0QP

Study participating centre
The Dudley Group NHS Foundation Trust
Pensnett Road
Dudley
United Kingdom
DY1 2HQ

Study participating centre
Nottingham University Hospitals NHS Trust
Queens Medical Centre
Derby Road
NOTTINGHAM
United Kingdom
NG7 2UH

Study participating centre
Surrey and Sussex Healthcare NHS Trust
Canada Avenue
Redhill
United Kingdom
RH1 5RH

Study participating centre
Belfast Health and Social Care Trust
A Floor - Belfast City Hospital
Lisburn Road
Belfast
United Kingdom
BT9 7AB

Study participating centre
NHS Greater Glasgow and Clyde
J B Russell House
Gartnavel Royal Hospital
1055 Great Western Road Glasgow
Glasgow
United Kingdom
G12 0XH

Study participating centre
University Hospital Southampton NHS Foundation Trust
Tremona Road
Southampton
United Kingdom
SO16 6YD

Study participating centre
Cambridge University Hospitals NHS Foundation Trust
Hills Road
Cambridge
United Kingdom
CB2 0QQ

Study participating centre
University Hospitals Birmingham NHS Foundation Trust
Queen Elizabeth Hospital
Mindelsohn Way
Edgbaston
Birmingham
United Kingdom
B15 2GW

Study participating centre
University Hospitals of Leicester NHS Trust
Infirmary Square
Leicester
United Kingdom
LE1 5WW

Study participating centre
London North West University Healthcare NHS Trust
Northwick Park Hospital
Watford Road
Harrow
United Kingdom
HA1 3UJ

Study participating centre

Northern Care Alliance NHS Foundation Trust
Stott Lane
Manchester
United Kingdom
M6 8HD

Study participating centre
Medway NHS Foundation Trust
Windmill Road
Gillingham
United Kingdom
ME7 5NY

Sponsor information

Organisation

University of Birmingham

Sponsor details

Research Strategy & Services Division – Research Governance
Birmingham Research Park
97 Vincent Drive
Edgbaston
Birmingham
England
United Kingdom
B15 2SQ
+44 (0)7814 650 003
researchgovernance@contacts.bham.ac.uk

Sponsor type

University/education

Website

<http://www.birmingham.ac.uk/index.aspx>

ROR

<https://ror.org/03angcq70>

Funder(s)

Funder type

Government

Funder Name

Health Technology Assessment Programme

Alternative Name(s)

NIHR Health Technology Assessment Programme, HTA

Funding Body Type

Government organisation

Funding Body Subtype

National government

Location

United Kingdom

Results and Publications

Publication and dissemination plan

On completion of the trial, the data will be analysed, and a Final Study Report prepared. Results of this trial will be submitted for publication in a peer reviewed journal and the findings of the trial will be made public. This manuscript will be prepared by the CI and members of the TMG and submitted to the whole TMG in a timely fashion and in advance of being submitted for publication to allow time for review.

Outputs from this trial will be published under a corporate authorship group. Each publication will include a detailed description of the exact contributions of each person, following accepted guidelines for collaborative authorship models.

Any secondary publications and presentations prepared by investigators must be reviewed and approved the TMG. Manuscripts should be submitted to the TMG in a timely fashion and in advance of being submitted for publication, to allow time for review and resolution of any outstanding issues. In all publications, authors must acknowledge that the trial was performed with the support of NIHR, University Hospitals Birmingham and the University of Birmingham (the Sponsor) and Birmingham Clinical Trials Unit. Intellectual property rights will be addressed in the OCEaN Clinical Trial Site Agreement between Sponsor and site.

Participants can request the published trial results from their PI once available.

Intention to publish date

01/10/2026

Individual participant data (IPD) sharing plan

The datasets generated during and/or analysed during the current study will be available upon request from the BCTU Data Sharing Committee.

ocean@trials.bham.ac.uk

IPD sharing plan summary

Available on request

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
HRA research summary			20/09/2023	No	No