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A study to explore the best dose of mitomycin-c inside your eye during retinal detachment surgery

Submission date 12/11/2024	**Recruitment status** Recruiting	[_] Prospectively registered [_] Protocol [_] Statistical analysis plan [_] Results [_] Individual participant data [X] Record updated in last year
Registration date 12/02/2025	**Overall study status** Ongoing	
Last Edited 12/02/2025	**Condition category** Eye Diseases	

Plain English summary of protocol

Background and study aims
The aim of this study is to investigate the potential benefit of a commonly used anti-inflammatory treatment to improve the outcome of surgery in eyes with retinal detachment. The retina is a thin layer which lines the inside of the eye. It is sensitive to light (like the film in a camera) and is necessary for vision. If a hole or tears develop in a retina, it can become detached. Retinal detachment is the most common eye emergency. For 1 in 5 patients the detached retina can develop scar tissue on its surface, a complication termed proliferative vitreoretinopathy (PVR). This PVR scar tissue pulls on the retina preventing the holes or tears from being repaired by standard surgery. The scar tissue increases your risk of the retina detaching again and requires more operations to resolve the problem. Multiple surgeries to remove the scar tissue often result in poor vision outcomes that do not meet the patients' expectations. Efforts to stop and treat PVR scar tissue formation have so far proved unsuccessful.
Mitomycin C (MMC) has been used in ophthalmology for over 30 years, with evidence to show its ability to stop scarring after eye surgery. This low-cost drug is currently used routinely around the globe to treat patients for other eye problems by applying MMC-soaked mini-sponges to the external surface of the eye. This study aims to demonstrate safe and potentially effective MMC treatment by direct application to the internal surface of the patient's eye with retinal detachment, at high risk of PVR. The researchers will define the most appropriate and safest dose of MMC in patients with retinal detachment.

Who can participate?
Patients aged 18 years and over who have a diagnosis of repeated retinal detachment complicated by scar tissue

What does the study involve?
Participants will receive MMC in addition to the standard eye operation

What are the possible benefits and risks of participating?
There are limited previous clinical studies using the IMP for this indication. The safety and effectiveness of intraocular use have been shown in small groups of people (about 30), with no

major safety events occurring. There is a risk that in larger groups of people the safety events will be more common. In this study, the dose will start low at 0.02% MMC and potentially reach 0.05%. In previous studies, higher doses (0.04%) have been used, but not 0.05%.The IMP is commonly used in glaucoma and there have been multiple trials testing the safety of its use. The dosing will be staggered in order to properly monitor each patient in turn. Statistical methods and DSMB judgement will be used to decide on appropriate dose escalation. The maximum dose will be 0.05%, which is only marginally more concentrated than previous studies using 0.04%.
Participants will be observed for the duration of IMP treatment, and have a visit 1 day after surgery and then weekly visits for 3 weeks.
The patients will be attending more frequently during the first 3 months following surgery, with patients attending every 2 weeks for face-to-face clinical visits/assessments until the 3-month visit.
All the study visits and care after surgery are standard practice for retinal detachment. Therefore there is no additional burden introduced by the trial visits.

Where is the study run from?
1. University College London (UK)
2. Moorfields Eye Hospital (UK)

When is the study starting and how long is it expected to run for?
November 2024 to January 2028

Who is funding the study?
Medical Research Council (UK)

Who is the main contact?
1. Jessie Barry, cctu.morph@ucl.ac.uk
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Contact information

Type(s)
Public

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Additional identifiers

EudraCT/CTIS number
Nil known

IRAS number
1010526

ClinicalTrials.gov number
Nil known

Secondary identifying numbers
137155

Study information

Scientific Title
A Phase I/II dose-finding study of intraocular mitomycin-c adjunct in vitrectomy for retinal detachment and proliferative vitreoretinopathy (MORPH-1)

Acronym
MORPH-1

Study objectives
Safety:
To affirm the safety profile of the pharmacological doses of Mitomycin based on the occurrence of Dose Limiting Events (DLEs).

Efficacy:
Successful retinal re-attachment without tamponade at 6 months.

1.+10 ETDRS letters improvement in Visual acuity (ETDRS chart at 4 meters starting distance) at 6 months
2. Presence or absence of PVR grade C as measured/mapped on:
2.1. Widefield fundus photography,
2.2. OCT (Optical coherence tomography)
2.3. Clinical fundus examination
3. Mean 5-10 pc/ms reduction in flare from baseline at 6 months post-surgery

Ethics approval required
Ethics approval required

Ethics approval(s)
Not yet submitted, ref: 24/NE/0211

Study design
Non-randomized study

Primary study design
Interventional

Secondary study design
Non randomised study

Study setting(s)
Hospital

Study type(s)
Safety, Efficacy

Participant information sheet
Not available in web format, please use the contact details to request a participant information sheet

Health condition(s) or problem(s) studied
Proliferative vitreoretinopathy in rhegmatogenous retinal detachment

Interventions
Additional use of mitomycin-C during retinal detachment surgery (vitrectomy) including dose or dose range 0.1, 0.2, 0.3, 0.4, 0.5, and 1 mg/ml, dose frequency one dose per participant, intraocular route of administration and duration of treatment 3 minutes of application.

Intervention Type
Drug

Pharmaceutical study type(s)
Dose response

Phase
Phase I

Drug/device/biological/vaccine name(s)
Mitomycin-C

Primary outcome measure
Safety:
The occurrence of Dose Limiting Events (DLEs) over 7 days post-surgery. Safety events are assessed against DLE criteria from Day 1 to Day 7.

Efficacy:
Successful retinal re-attachment without tamponade at 6-month patient follow-up, as assessed by the slit-lamp examination.

Secondary outcome measures
1. Change in visual acuity (measured using ETDRS chart at 4 metres starting distance) in the MMC treatment group from baseline to the 6-month final follow-up visit
Target: +10 ETDRS letters improvement
Acceptable: +5 ETDRS letters improvement
2. PVR grade, change from baseline to the 6-month final follow-up visit
Target: Complete absence of PVR in 80% of patients in the MMC group
Acceptable: Significant difference in the rate of PVR presence between the MMC group and the wider population
PVR grade C as measured/mapped on widefield fundus photography, OCT and clinical fundus examination
3. Reduction in flare measured by laser flare photometry, change from baseline to the 6-month final follow-up visit
Target: mean 5-10 pc/ms reduction in flare post-surgery/MMC
Acceptable: mean 0-5 pc/ms reduction in flare post-surgery/MMC

Overall study start date
08/11/2024

Completion date
30/01/2028

Eligibility

Key inclusion criteria
1. Individuals aged ≥18 years
2. Recurrent macula-OFF Rhegmatogenous Retinal Detachment (RRD) complicated with Proliferative vitreoretinopathy (PVR) grade C
3. Requiring surgery with silicone oil tamponade
4. Able and agree to provide written informed consent
5. Able and agree to attend all protocol mandated follow-up visits
6. If participants of childbearing potential, must have negative pregnancy test at screening visit and prior to surgery, and agree to at least one form of contraception throughout the duration of the trial

Participant type(s)
Patient

Age group
Adult

Lower age limit
18 Years

Sex
Both

Target number of participants
50

Key exclusion criteria
1. Previous known adverse reaction to mitomycin C (MMC)
2. History of open globe injury
3. A diagnosis of ocular hypertension on two or more pressure-lowering medications
4. Uncontrolled or advanced glaucoma (defined as uncontrolled eye pressure, changes to medication, recent surgery in the last 3 months or being considered for surgical treatment)
5. Uncontrolled uveitis
6. Previous steroid-induced glaucoma
7. Proliferative diabetic retinopathy or vasculopathy
8. Participant who is pregnant or is planning to become pregnant for the duration of the trial
9. Participant who has given birth within the past 6 months or is breastfeeding
10. Suspected ocular/periocular infection (e.g. Herpes Simplex Virus, Varicella Zoster Virus, mycobacterial, fungal disease)
11. Aphakia
12. Participant in whom a lensectomy is planned at the time of surgery
13. Pre-existing anterior chamber intraocular lens
14. Participant enrolled in another interventional clinical trial within 90 days prior to screening
15. Participants receiving any other treatments such as anti-cancer or autoimmune disease

Date of first enrolment
10/01/2025

Date of final enrolment
10/06/2026

Locations

Countries of recruitment
United Kingdom

Study participating centre
-
United Kingdom
-

Sponsor information

Organisation
University College London

Sponsor details
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Sponsor type
University/education

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ROR
https://ror.org/02jx3x895

Funder(s)

Funder type
Research council

Funder Name
Medical Research Council

Alternative Name(s)
Medical Research Council (United Kingdom), UK Medical Research Council, MRC

Funding Body Type
Government organisation

Funding Body Subtype
National government

Location
United Kingdom

Results and Publications

Publication and dissemination plan
1. Peer-reviewed scientific journals
2. Internal report
3. Conference presentation
4. Other publication
5. Submission to regulatory authorities

Intention to publish date
30/01/2029

Individual participant data (IPD) sharing plan

The datasets generated during and/or analysed during the current study will be available upon request from cctu-enquiries@ucl.ac.uk. Participant-level data will be stored securely on UCL CCTU servers. This dataset will not be made public in its raw form to protect the participant's identities and to comply with data protection. A fully anonymised dataset will be made available to researchers on submission and review of a formal request, sent in writing to UCL CCTU and the MORPH-1 Chief investigator.

IPD sharing plan summary

Available on request