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A study to find out the safety, tolerability and processing of RO7204239 by the body, the effect of RO7204239 on the body in participants with high body weight

Submission date 22/04/2024	**Recruitment status** No longer recruiting	[X] Prospectively registered [_] Protocol [_] Statistical analysis plan [_] Results [_] Individual participant data [_] Record updated in last year
Registration date 22/04/2024	**Overall study status** Ongoing	
Last Edited 30/04/2024	**Condition category** Other	

Plain English summary of protocol

Background and study aims
Obesity is a complex health issue that occurs due to the accumulation of extra (surplus) body fat. This condition significantly increases the risk of developing other diseases and health issues, including heart disease, a group of health conditions that cause a person's blood sugar to become too high (diabetes) and high blood pressure.
This study is testing a medicine called RO7204239. It is being developed as a possible treatment for obesity. RO7204239 is an experimental medicine. This means health authorities (like the U.S. Food and Drug Administration and European Medicines Agency) have not approved RO7204239 for the treatment of obesity. This study aims to test how safe RO7204239 is (at single and multiple doses), what happens once it is in the body, and what RO7204239 does to the body.

Who can participate?
People (males and females) aged 18-80 years with a body weight of at least 90 kg can take part in the study. People who are pregnant, or currently breastfeeding cannot take part in the study.

What does the study involve?
People will be screened to check if they can participate in the study. The screening period will take place about 6 weeks before the start of the treatment.
Everyone who joins this study will be split into two groups to receive RO7204239 given as an injection under the skin. Participants in Group 1 will receive a single dose of RO7204239 on Day 1 and those in Group 2 will receive multiple doses of RO7204239 for about 12 weeks.
Participants will have regular blood tests and will be checked for unwanted effects throughout the study.
This is an open-label study. This means everyone involved, including the participant and the study doctor, will know the study treatment the participant has been given.
During the study, the study doctors will see the participants 11 times (group 1) or 17 times (group 2) during the clinic visits. Study doctors will see how well the treatment is working and any unwanted effects participants may have. Participants will have 12 follow-up visits after

completing the study treatment, during which the study doctor will check on the participant's well-being. The total time of participation in the study will be about 42 weeks for Group 1 and about 54weeks for Group 2. Participants have the right to stop study treatment and leave the study at any time if they wish to do so.

What are the possible benefits and risks of participating?
Taking part in the study may or may not make participants feel better. However, the information collected in the study can help people with obesity in the future.
It may not be fully known at the time of the study how safe the study treatment is in healthy volunteers. The study involves some risks to the participant, but these risks are generally mild and easily monitored. People interested in taking part will be informed about the risks, as well as procedures or tests they may need to undergo. All details of the study will be described in an informed consent document. This includes information about possible side effects.
Risks associated with the study drug
Participants may have unwanted effects of the drug used in this study. These unwanted effects can be mild to severe, even life-threatening, and vary from person to person. During this study, participants will have regular check-ups to see if there are any unwanted effects.
Participants will be told about the known unwanted effects of RO7204239 and possible unwanted effects based on human and laboratory studies or knowledge of similar medicines. Known unwanted effects include local irritation, pain, swelling, hardening of the skin, itching, redness, and rash at the injection site.
The study medicine(s) may be harmful to an unborn baby. Women and men must take precautions to avoid exposing an unborn baby to the study treatment.

Where is the study run from?
F. Hoffmann-La Roche Ltd (Switzerland)

When is the study starting and how long is it expected to run for?
March 2024 to September 2025

Who is funding the study?
F. Hoffmann-La Roche Ltd (Switzerland)

Who is the main contact?
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Contact information

Type(s)
Public, Scientific, Principal Investigator

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Additional identifiers

EudraCT/CTIS number
Nil known

IRAS number

ClinicalTrials.gov number
Nil known

Secondary identifying numbers
BP45369

Study information

Scientific Title
A single-and multiple-dose, open-label, Phase I study to investigate the safety, tolerability, pharmacokinetics, pharmacodynamics, and immunogenicity of subcutaneously administered RO7204239 in adult participants with high body weight

Study objectives
The main purpose of this study is to evaluate the safety and tolerability of single and multiple doses of RO7204239 in participants with high body weight.

Ethics approval required
Ethics approval required

Ethics approval(s)
Approved 18/04/2024, New Zealand Northern B Health and Disability Ethics Committee (Ministry of Health, 133 Molesworth Street, PO Box 5013, Wellington, 6011, New Zealand; +64 (0) 800 400 569; hdecs@health.govt.nz), ref: 2024 FULL 19681

Study design
Phase I open-label non-randomized uncontrolled parallel-group study

Primary study design
Interventional

Secondary study design
Non randomised study

Study setting(s)
Pharmaceutical testing facility

Study type(s)
Safety

Participant information sheet
Not available

Health condition(s) or problem(s) studied
Healthy volunteers

Interventions
Group 1: Participants will receive a single dose of RO7204239, as a subcutaneous (SC) injection on Day 1.

Group 2: Participants will receive multiple doses of RO7204239, as SC injection, every 2 weeks (Q2W) during the first 4 weeks and then every 4 weeks (Q4W) for 8 weeks.

Intervention Type
Drug

Pharmaceutical study type(s)
Pharmacokinetic, Pharmacodynamic

Phase
Phase I

Drug/device/biological/vaccine name(s)
RO7204239

Primary outcome measure
1. Number of participants with treatment-emergent adverse events (AEs) and severity of TEAEs assessed using the National Cancer Institute Common Terminology Criteria for Adverse Events (NCI CTCAE v5.0) from Day 1 up to Day 253 for Group 1 and from Day 1 up to Day 337 for Group 2
2. Number of participants with local and systemic injection reactions assessed using data collected in the electronic case report form (eCRF) from Day 1 up to Day 253 for Group 1 and from Day 1 up to Day 337 for Group 2
3. Number of participants with local injection site pain measured on a 10-point numerical rating scale, completed by participants at multiple timepoints on Day 1 for Group 1 and from Day 1 up to Day 85 for Group 2
4. Number of participants with local injection site pruritus measured on a 10-point numerical rating scale, completed by participants at multiple timepoints on Day 1 for Group 1 and from Day 1 up to 85 for Group 2

Secondary outcome measures
1. Maximum observed concentration (Cmax) of RO7204239 assessed using non-compartmental methods from the plasma samples collected at pre-dose and multiple time points post-dose from Day 1 up to Day 253 for Group 1 and from Day 1 up to Day 337 for Group 2
2. Area under the serum concentration-time curve (AUC) from the time of dosing to the last measurable concentration (AUClast) of RO7204239 assessed using non-compartmental methods from the plasma samples collected at pre-dose and multiple time points post-dose from Day 1 up to Day 253 for Group 1
3. AUC from single dosing time extrapolated to infinity (AUCinf) of RO7204239 assessed using non-compartmental methods from the plasma samples collected at pre-dose and multiple time points post-dose from Day 1 up to Day 253 for Group 1
4. AUC over a dosing interval (AUCtau) of RO7204239 assessed using non-compartmental methods from the plasma samples collected at pre-dose and multiple time points post-dose from Day 1 up to Day 337 for Group 2

5. Absolute levels of total latent myostatin, free latent myostatin and total mature myostatin measured from the serum samples collected at pre-dose and multiple time points post-dose from Day -1 up to Day 253 for Group 1 and from Day -1 up to Day 337 for Group 2
6. Change from baseline in levels of total latent myostatin, free latent myostatin and total mature myostatin measured from the serum samples collected at pre-dose and multiple time points post-dose from Day -1 up to Day 253 for Group 1 and from Day -1 up to Day 337 for Group 2
7. Number of participants with anti-drug antibodies (ADAs) to RO7204239 as measured using plasma samples collected from Day 1 up to Day 253 for Group 1 and Day 1 up to Day 337 for Group 2

Overall study start date
13/03/2024

Completion date
12/09/2025

Eligibility

Key inclusion criteria
1. Healthy participants (other than body weight-related conditions) as determined by medical evaluation including medical history, physical examination, laboratory tests, and cardiac monitoring
2. Body weight ≥90 kilograms (kg) and body mass index (BMI) ≥ 25 kilograms per square meter (kg/m2)
3. Must agree to refrain from starting a significant new exercise routine or major change to a previous exercise/diet routine within 4 weeks before screening and to maintain their level of physical activity consistently through study Day 113

Participant type(s)
Other

Age group
Mixed

Lower age limit
18 Years

Upper age limit
80 Years

Sex
Both

Target number of participants
36

Key exclusion criteria
1. Presence of clinically significant cardiovascular disease (e.g., cardiac insufficiency, unstable angina, cardiomyopathy, congestive heart failure, or valve disorders or defects)

2. Significant allergies to humanized monoclonal antibodies (mAbs)
3. Lymphoma, leukemia, or any malignancy within the past 5 years
4. Breast cancer within the past 10 years
5. Current or chronic history of liver disease or known hepatic or biliary abnormalities
6. Gastric bypass surgery
7. Any abnormal skin conditions or potentially obscuring pigmentation or lesions in the area intended for SC injection that would prevent visualization of potential injection site reactions (ISRs) to RO7204239
8. Use of anti-obesity medications within 6 months prior to enrollment
9. Live vaccine(s) within one month before screening or plans to receive live vaccines during the study or within 28 days of the last study treatment administration
10. Treatment with biologic agents (such as mAbs including marketed drugs) within 3 months or five half-lives (whichever is longer) prior to dosing
11. Presence of hepatitis B surface antigen and/or total hepatitis B core antibody, a positive hepatitis C or human immunodeficiency virus (HIV) antibody test result, and/or evidence of HIV infection at Screening or within 3 months prior to first study treatment administration

Date of first enrolment
10/05/2024

Date of final enrolment
31/07/2024

Locations

Countries of recruitment
New Zealand

Study participating centre
New Zealand Clinical Research Auckland
3 Ferncroft Street
Grafton
Auckland
New Zealand
1010

Sponsor information

Organisation
F. Hoffmann-La Roche Ltd

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Sponsor type
Industry

Website
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Funder(s)

Funder type
Industry

Funder Name
F. Hoffmann-La Roche

Alternative Name(s)
Hoffman-La Roche, F. Hoffmann-La Roche Ltd.

Funding Body Type
Private sector organisation

Funding Body Subtype
For-profit companies (industry)

Location
Switzerland

Results and Publications

Publication and dissemination plan
Planned publication in a high-impact peer-reviewed journal

Intention to publish date
12/09/2026

Individual participant data (IPD) sharing plan
The datasets generated during and/or analysed during the current study are not expected to be made available due to participant-level data not being a regulatory requirement.

IPD sharing plan summary
Not expected to be made available