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A study to evaluate safety and pharmacokinetics (processing by the body) of single ascending doses of GDC-5780 in healthy participants

Submission date 02/02/2022	**Recruitment status** No longer recruiting	[X] Prospectively registered [_] Protocol [_] Statistical analysis plan [_] Results [_] Individual participant data [_] Record updated in last year
Registration date 09/02/2022	**Overall study status** Completed	
Last Edited 07/02/2022	**Condition category** Other	

Plain English summary of protocol

Background and studyaims
This study will evaluate the safety (side effects) and how the body processes the treatment (pharmacokinetics) of the drug GDC-5780 in healthy volunteers.
The purpose of this first-in-human study is:
1. To find out how safe GDC-5780 is when given as a single dose
2. To find out how GDC-5780 will be distributed and eliminated from the body

GDC-5780 is an experimental drug, which means health authorities have not approved GDC-5780 for the treatment of any disease, and it has not been tested in people before this study.

Who can participate?
Healthy participants aged between 18 to 65 years.

What does the study involve?
Participants may be asked to be in the study for up to 40 days. This includes:
- A Screening Period of up to 28 days before the start of the study where tests will be done to check if the participants are eligible to take part in the study.
- Treatment Period where participants willhave to check in to the clinic 2 days before receiving the study treatment and will have to stay in the clinic for 5 nights and receive a singledoseof GDC-5780or placebo (drug without an active substance).
- Follow-up Period where participantswill have toreport to the clinicfor a check-up2 times, with the last visit taking place about 14 days after the dose of study drug.

The first 8 participants will get a single dose of GDC-5780 or placebo as intravenous (IV) infusion (through the vein) over 2 hours. The dose for new participants joining the study will be adjusted according to the test results of previous participants. Participants joining the study at later stages will get higher doses of the study drug.
All participants will be closely monitored throughout the study to ensure that the study drug is

safe and tolerable. After the dose of study treatment, the study doctor will follow-up participants for 14 days.

What are the possible benefits and risks of participating?
Participants will not receive any benefit from participating in this study, but the information that is learned may help people withUTIs in the future.
No clinical information is available for GDC-5780 to date, as this is a first-in-humanstudy. The expected risks for GDC-5780, determined according to the mechanism of action and results from nonclinical studies (laboratory studies on animals) are listed below:

- Reaction during or following the drug infusion that may mimic an allergic reaction and could include symptoms such as fever, chills, rash, low blood pressure, and difficulty breathing
- Sudden decrease in kidney function
- Transient loss of muscle coordination; awkward, uncoordinated walking; or unsteadiness when walking

There may be a risk in exposing an unborn child to the study drug, and all risks are not known at this time. Participants must take precautions to avoid exposing an unborn child to the study drug. Participantswhoare pregnant, become pregnant, or are currently breastfeeding, cannot take part in this study.

Where is the study run from?
Roche (USA)

When is the study starting and how long is it expected to run for?
October 2021 to August 2022

Who is funding the study?
Roche (USA)

Who is the main contact?
global-roche-genentech-trials@gene.com

Contact information

Type(s)
Public

Contact name
Dr Clinical Trials

Contact details
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Additional identifiers

EudraCT/CTIS number

Nil known

IRAS number

ClinicalTrials.gov number
Nil known

Secondary identifying numbers
GV43221

Study information

Scientific Title
A Phase I, randomized, double-blind, single ascending dose study to evaluate the safety and pharmacokinetics ofGDC-5780 in healthy subjects

Study objectives
The purpose of this study is to evaluate the safety and pharmacokineticsof single doses of GDC-5780 in healthy participants.

Ethics approval required
Old ethics approval format

Ethics approval(s)
Approved 09/12/2021, WCG IRB (1019 39th Ave SE, Suite 120, Puyallup WA98374, USA; +1 800-562-4789; no email provided) ref: 20216447

Study design
Phase I first-in-human participant- and investigator-blinded randomized placebo-controlled dose-escalation study in healthy participants

Primary study design
Interventional

Secondary study design
Randomised controlled trial

Study setting(s)
Other

Study type(s)
Treatment

Participant information sheet
No participant information sheet available

Health condition(s) or problem(s) studied
Safety and pharmacokineticsof single doses of GDC-5780 in healthy participants

Interventions

1. GDC-5780 – Single Ascending Dose: Participants will receive a single dose of GDC-5780 as intravenous (IV) infusion over 2 hours on Day 1 in the first cohort. The dose will be escalated in subsequent cohorts, as per the Safety Monitoring Committee decision in consultation with the investigator.
2. Placebo - Participants will receive a single dose of placebo as an IV infusion over 2 hours on Day 1.

Intervention Type
Drug

Phase
Phase I

Drug/device/biological/vaccine name(s)
GDC-5780

Primary outcome measure
1. Percentage of participants withadverse events (AEs) measured from Day 1until 14 days after the dose of study drug
2. Percentage of participants withadverse events based onseverity per Division of AIDS (DAIDS) Toxicity Grading Scale from Day 1until14 days after thedose of study drug
3. Percentage of participants with clinically significant change in vital signs measured using body temperature, respiratory rate, pulse rate, and blood pressure fromDay 1 until 14 days after thedose of study drug
4. Percentage of participants with clinically significant laboratory test abnormalities measured using blood and urine samples fromDay 1until14 days after the dose of study drug
5. Percentage of participants with clinically significantelectrocardiogram (ECG) abnormalities measured using single 12-lead ECG or Holter ECG recordings from Day 1until14 days after thedose of study drug

Secondary outcome measures
1. Maximum observed plasma concentration (Cmax) of GDC-5780 measured from blood samples taken at multiple timepoints over Days 1, 2, 3, and 4
2. Time to reach maximum plasma concentration (Tmax) of GDC-5780, measured from blood samples taken at multiple timepoints over Days 1, 2, 3, and 4
3. Terminal half-life (t1/2) of GDC-5780 measured from blood samples taken at multiple timepoints over Days 1, 2, 3, and 4
4. Area under the plasma concentration-time curve from time zero to 24 hours after dosing (AUC (0-24)) with GDC-5780 measured from blood samples taken at multiple timepoints over Days 1, 2, 3, and 4
5. Area under the plasma concentration-time curve from time zero to the time of last quantifiable concentration (AUC (0-last)) of GDC-5780 measured from blood samples taken at multiple timepoints over Days 1, 2, 3, and 4
6. Area under the plasma concentration-time curve from time zero to infinity (AUC (0-infinity)) of GDC-5780 measured from blood samples taken at multiple timepoints over Days 1, 2, 3, and 4
7. Renal clearance (CLr) measured from multiple pooled urine samples taken at multiple timepoints from Day 1 to Day 4
8. Fractional excretion of GDC-5780 measured from multiple pooled urine samples taken at multiple timepoints from Day 1 to Day 4

Overall study start date

26/10/2021

Completion date
05/08/2022

Eligibility

Key inclusion criteria
1. Body mass index ≥18.5 and <30 kg/m²
2. Body temperature of 35°C–37.6°C at screening
3. Systolic blood pressure of 90–139 mmHg and diastolic blood pressure of 45–89 mmHg at screening
4. Agree to abstain from consumption of grapefruit, grapefruit hybrids, oranges, orange hybrids, pomelos, exotic citrus fruits, and all orange and grapefruit-type fruit juices from 48 hours prior to check-in until clinic check-out (i.e., end of residential stay)
5. Agree to abstain from consumption of alcohol from 48 hours prior to cliniccheck-in until clinic check-out
6. Agreement to abstain from consumption of caffeine-containing foods and beverages(e.g., coffee, tea, chocolate, energy drinks, soda) from 48 hours prior to clinic check-in until clinic check-out

Participant type(s)
Healthy volunteer

Age group
Adult

Lower age limit
18 Years

Sex
Both

Target number of participants
56

Key exclusion criteria
1. Pregnancy or breastfeeding, or intention of becoming pregnant during the study or within 14 days after the dose of study drug
2. Planned procedure or surgery during the study
3. Clinical laboratory values outside the normal reference range for the test laboratory at screening, Day−2, or Day−1. Participants with an estimated glomerular filtration rate <90 mL/min /1.73 m² at screening, Day−2, or Day−1, as calculated using the Chronic Kidney Disease Epidemiology Collaboration equation, must be excluded from the study
4. Positive HIV test at screening
5. Positive hepatitis B surface antigen (HBsAg), hepatitis B core antibody (HBcAb), hepatitis B virus (HBV) DNA test at screening, hepatitis C virus (HCV)
6. Urine sample positive for drugs of abuse at screening or Day−2
7. Treatment with investigational biologic therapy within 90 daysor 5 drug eliminationhalf-lives (if known), whichever is longer, prior to initiation of study drug

8. Acute illness within 14 days prior to screening
9. Vaccination within 14 days prior to initiation of study drug

Date of first enrolment
02/03/2022

Date of final enrolment
22/07/2022

Locations

Countries of recruitment
United States of America

Study participating centre
PRA International Clinical Pharmacology Centre
9755 Ridge Dr.
Lenexa, KS
United States of America
66219

Sponsor information

Organisation
Roche (United States)

Sponsor details
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Sponsor type
Industry

Website
http://www.roche.com/about_roche/roche_worldwide.htm

ROR
https://ror.org/011qkaj49

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
05/08/2023

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