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A study to describe the fertility journey of risdiplam-treated adult male individuals with spinal muscular atrophy

Submission date
22/02/2024

Recruitment status
Recruiting

Registration date
13/03/2024

Overall study status
Ongoing

Last Edited
11/06/2024

Condition category
Nervous System Diseases

[X] Prospectively registered
[ ] Protocol
[ ] Statistical analysis plan
[ ] Results
[ ] Individual participant data
[ ] Record updated in last year

Plain English summary of protocol

Background and study aims
Spinal muscular atrophy (SMA) is an inherited disorder which results in weakness and wasting of muscles used for movement. It is caused by the loss of certain specialized nerve cells in the brain and spinal cord that control muscle movement, known as motor neurons. Risdiplam is approved by the U.S. Food and Drug Administration (FDA) for the treatment of paediatric and adult patients with SMA. The purpose of this study, the MARLIN study, is to gather information about the fertility journey of adult male individuals with SMA who are taking or have taken risdiplam.

Who can participate?
Males with SMA between 18 and 50 years of age, who are taking or have taken risdiplam, and are trying to conceive a child or have previously conceived a child after taking risdiplam, can participate in this study. Additionally, there is an option for the participant's sexual partner /surrogate/gestational carrier to also participate.

What does the study involve?
Participation in this study is fully remote and involves the completion of questionnaires using an electronic platform. It does not include any procedures or doctor visits. The questionnaires include the following topics: demographics, medical history and medications, risdiplam use, sexual history, and fertility journey (including any tests and treatments the participant has undergone in their attempt to conceive a child). Each questionnaire takes approximately 20-45 minutes to complete.
Participants who are actively trying to conceive a child will complete an initial questionnaire and subsequently complete a follow-up questionnaire once a year for up to 3 years. Participants who conceived before study enrollment but after treatment with risdiplam will only complete the initial questionnaire.
Participants who successfully conceive or stop trying to conceive before the end of the 3 years will be considered to have completed the study. If a participant has conceived at the end of 3 years but the outcome of the pregnancy is not yet known, they will complete an additional

follow-up questionnaire to capture this outcome.
If the participant's sexual partner, surrogate, or gestational carrier agrees to take part in the study, they will also be asked to complete similar questionnaires.

What are the possible benefits and risks of participating?
Participants will not receive any direct medical benefit from participating in this study but the information that is learned may help other people with SMA in the future.

Where is the study run from?
Genentech Inc. (USA)

When is the study starting and how long is it expected to run for?
October 2023 to May 2029

Who is funding the study?
Genentech Inc. (USA)

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

Study website
https://www.marlinstudy.com/

Contact information

Type(s)
Public, Scientific, Principal Investigator

Contact name
Dr Clinical Trials

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Additional identifiers

EudraCT/CTIS number
Nil known

IRAS number

ClinicalTrials.gov number
Nil known

Secondary identifying numbers

ML44914

Study information

Scientific Title
Observational study of fertility in risdiplam-treated adult male patients with spinal muscular atrophy (MARLIN)

Acronym
MARLIN

Study objectives
The purpose of the study is to collect and describe fertility-related outcomes in adult males with SMA who have received risdiplam treatment.

Ethics approval required
Ethics approval required

Ethics approval(s)
Approved 26/02/2024, WCG North America (212 Carnegie Center, Suite 301, Princeton, NJ, 08540, United States of America; +1 855-818-2289; Clientcare@wcgclinical.com), ref: 20240680

Study design
Phase IV real-world observational study with prospective and retrospective data collection

Primary study design
Observational

Secondary study design
Longitudinal study

Study setting(s)
Internet/virtual, Other

Study type(s)
Other

Participant information sheet
No participant information sheet available

Health condition(s) or problem(s) studied
Spinal muscular atrophy

Interventions
Adult male participants with SMA, who are currently on treatment with risdiplam or were previously treated with risdiplam, will be enrolled in this study. Participants actively trying to conceive will provide prospective data via questionnaires at baseline and annually for 3 years, with an option for their sexual partner/surrogate/gestational carrier to complete questionnaires. If the partner/surrogate/gestational carrier is pregnant at the end of the 3 years of follow up, an additional annual questionnaire to capture the pregnancy outcome will be

administered. Data will be collected at baseline for those who conceived before study enrollment but after exposure to risdiplam.

If a participant successfully conceives and the outcome of the pregnancy is known, or if a participant stops trying to conceive before the end of the 3 years of follow-up, they will have completed the study early.

Questionnaires will be entered into an electronic platform to be administered remotely following participant consent.

Intervention Type
Other

Primary outcome measure
Whether risdiplam-exposed adult males with SMA conceive, measured using questionnaires completed by the participants at study initiation and annually for up to 3 years

Secondary outcome measures
1. Presence of confounding factors that may impact fertility in enrolled participants, measured using questionnaires completed by the participants at study initiation and annually for up to 3 years
2. Fertility-related healthcare resource utilization, management, and treatment decisions in enrolled participants, measured using questionnaires completed by the participants at study initiation and annually for up to 3 years
3. Outcome of pregnancies, measured using questionnaires completed by the participants at study initiation and annually for up to 3 years

Overall study start date
10/10/2023

Completion date
30/05/2029

Eligibility

Key inclusion criteria
1. Diagnosis of SMA
2. Currently receiving or previously ever received risdiplam treatment
3. Actively trying to conceive or conceived in the past (during or after exposure to risdiplam)
4. Consent to a baseline and annual questionnaire for the duration of the study
5. Able to complete a questionnaire, in English, with or without assistance
6. Able to access a smartphone with internet connection

Participant type(s)
Patient

Age group
Adult

Lower age limit

18 Years

Upper age limit
50 Years

Sex
Male

Target number of participants
30

Key exclusion criteria
1. Participants who are using/used donor sperm for conception
2. Participants who are using/used their own cryopreserved sperm that was not exposed to risdiplam for conception

Date of first enrolment
30/05/2024

Date of final enrolment
30/05/2026

Locations

Countries of recruitment
United States of America

Study participating centre
United BioSource LLC (UBC)
920 Harvest Drive, Blue Bell
Pennsylvania
United States of America
19411

Sponsor information

Organisation
Genentech

Sponsor details
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Sponsor type
Industry

Website
https://www.roche.com/about/

ROR
https://ror.org/04gndp242

Funder(s)

Funder type
Industry

Funder Name
Genentech

Alternative Name(s)
Genentech, Inc., Genentech USA, Inc., Genentech USA

Funding Body Type
Private sector organisation

Funding Body Subtype
For-profit companies (industry)

Location
United States of America

Results and Publications

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

Intention to publish date
30/05/2030

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