Study of Two Doses of Crizanlizumab Versus Placebo in Adolescent and Adult Sickle Cell Disease Patients (STAND)

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Main Information

Primary registry identifying number
LBCTR2019060244

Protocol number
CSEG101A2301

MOH registration number

Study registered at the country of origin
Yes

Study registered at the country of origin: Specify

Type of registration
Prospective

Type of registration: Justify
N/A

Date of registration in national regulatory agency

Primary sponsor
Novartis Pharma Services Inc.

Primary sponsor: Country of origin
Novartis Pharmaceuticals

Date of registration in primary registry
15/10/2019

Date of registration in national regulatory agency

Public title
Study of Two Doses of Crizanlizumab Versus Placebo in Adolescent and Adult Sickle Cell Disease Patients (STAND)

Acronym

Scientific title
A Phase III, Multicenter, Randomized, Double-blind Study to Assess Efficacy and Safety of Two Doses of Crizanlizumab Versus Placebo, With or Without Hydroxyurea/ Hydroxycarbamide Therapy, in Adolescent and Adult Sickle Cell Disease Patients With Vaso-Occlusive Crises (STAND)

Acronym

Brief summary of the study: English
The purpose of this study is to compare the efficacy and safety of 2 doses of crizanlizumab (5.0 mg/kg and 7.5 mg/kg) versus placebo in adolescent and adult sickle cell disease (SCD) patients with history of vaso-occlusive crisis (VOC) leading to healthcare visit.

Brief summary of the study: Arabic
دراسة مرحلة ثالثة، متعددة المراكز، عشوائيّة التوزيع ومزدوجة التعمية لتقييم فعاليّة وسلامة جرعتين من دواء كريزانليزوماب مقابل الدواء الوهمي، مع أو بدون علاج هيدروكسي يوريا / هيدروكسي كاربامايد، لدى المرضى المراهقين والبالغين المصابين بداء الكريات المنجليّة مع نوبات انسداد وعائيّ (STAND)

Health conditions/problem studied: Specify
Sickle Cell Disease

Interventions: Specify
•Drug: Crizanlizumab (SEG101)
Crizanlizumab will be supplied in single use 10 mL glass vials at a concentration of 10 mg/mL. One vial contains 100 mg of crizanlizumab. This is a concentrate for solution for infusion IV.

Other Name: SEG101

•Drug: Placebo
Placebo will be supplied in single use 10 mL glass vials at a concentration of 10 mg/mL. One vial contains 100 mg of placebo. This is a concentrate for solution for infusion IV.

Key inclusion and exclusion criteria: Inclusion criteria

1.Written informed consent must be obtained prior to any screening procedures
2.Male or female patients aged 12 years and older on the day of signing informed consent. Adolescent include patients aged 12 to 17 years old and adults ≥ 18 years and older
3.Confirmed diagnosis of SCD by hemoglobin electrophoresis or high performance liquid chromatography (HPLC) [performed locally]. All SCD genotypes are eligible, genotyping is not required for study entry

4.Experienced at least 2 VOCs leading to healthcare visit within the 12 months prior to screening visit as determined by medical history. Prior VOC leading to healthcare visit must include:
a.Pain crisis defined as an acute onset of pain for which there is no other medically determined explanation other than vaso- occlusion -
b.a visit to a medical facility and/or healthcare professional,
c.and receipt of oral/parenteral opioids or parenteral nonsteroidal anti-inflammatory drug (NSAID) analgesia As well as other complicated crises, such as acute chest syndrome, priapism, and hepatic or splenic sequestration

5.If receiving HU/HC or erythropoietin stimulating agent or L-glutamine, must have been receiving the drug for at least 6 months prior to Screening visit and plan to continue taking at the same dose and schedule until the subject has reached one year of study treatment

6.Patients must meet the following central laboratory values at the screening visit:
◦Absolute Neutrophil Count ≥1.0 x 109/L
◦Platelet count ≥75 x 109/L
◦Hemoglobin: for adults (Hb) ≥4.0 g/dL and for adolescents (Hb) ≥5.5 g/dL
◦Glomerular filtration rate ≥ 45 mL/min/1.73 m2 using CKD-EPI formula in adults, and Shwartz formula in adolescents
◦Direct (conjugated) bilirubin < 2.0 x ULN
◦Alanine transaminase (ALT) < 3.0 x ULN

7.ECOG performance status ≤2.0 for adults and Karnofsky ≥ 50% for adolescents

Key inclusion and exclusion criteria: Gender

Both

Key inclusion and exclusion criteria: Specify gender

Key inclusion and exclusion criteria: Age minimum

12

Key inclusion and exclusion criteria: Age maximum

99

Key inclusion and exclusion criteria: Exclusion criteria

1.History of stem cell transplant.
2.Participating in a chronic transfusion program (pre-planned series of transfusions for prophylactic purposes) and/or planning on undergoing an exchange transfusion during the duration of the study; episodic transfusion in response to worsened anemia or VOC is permitted.
3.Contraindication or hypersensitivity to any drug or metabolites from similar class as study drug or to any excipients of the study drug formulation. History of severe hypersensitivity reaction to other monoclonal antibodies, which in the opinion of the investigator may pose an increased risk of serious infusion reaction.
4.Received active treatment on another investigational trial within 30 days (or 5 half-lives of that agent, whichever is greater) prior to Screening visit or plans to participate in another investigational drug trial.
5.Women of child-bearing potential, defined as all women physiologically capable of becoming pregnant unless they are using highly effective methods of contraception during dosing and for 15 weeks after stopping treatment.
6.Concurrent severe and/or uncontrolled medical conditions which, in the opinion of the Investigator, could cause unacceptable safety risks or compromise participation in the study.

7.History or current diagnosis of ECG abnormalities indicating significant risk of safety such as:
◦Resting QTcF ≥470 msec at pretreatment (baseline) for both male and female or inestability to determine QTc
◦Concomitant clinically significant cardiac arrhythmias (e.g ventricular tachycardia), and clinically significant second or third degree AV block without a pacemaker
◦History of familial long QT syndrome or know family history of Torsades de Pointes

8.Not able to understand and to comply with study intructions and requirements.

Type of study

Interventional

Type of intervention

Pharmaceutical

Type of intervention: Specify type

N/A

Trial scope

Safety

Trial scope: Specify scope

N/A

Study design: Allocation

Randomized controlled trial

Study design: Masking

Blinded (masking used)

Study design: Control

Placebo

Study phase

3

Study design: Purpose

Prevention

Study design: Specify purpose

N/A

Study design: Assignment

Parallel

Study design: Specify assignment

N/A

IMP has market authorization

No

IMP has market authorization: Specify

Name of IMP

SEG101 - Crizanlizumab

Year of authorization

Month of authorization

Type of IMP

Immunological

Pharmaceutical class

anti-human P-selectin antibody G1

Therapeutic indication

prevention of vaso-occlusive crises (VOCs) in patients of all genotypes with sickle cell disease (SCD)

Therapeutic benefit

To compare the efficacy of 5.0 mg/kg versus placebo and 7.5 mg/kg of crizanlizumab versus placebo in addition to standard of care.
To compare the efficacy of 7.5 mg/kg versus placebo on the annualized rate of all VOCs (managed at home + leading to healthcare visit), based on documentation by health care provider following contact with participant.
To compare the efficacy of 5.0 mg/kg versus placebo on the annualized rate of all VOCs (managed at home + leading to healthcare visit)

Study model

N/A

Study model: Explain model

N/A

Study model: Specify model

N/A

Time perspective

N/A

Time perspective: Explain time perspective

N/A

Time perspective: Specify perspective

N/A

Target follow-up duration

Target follow-up duration: Unit

Number of groups/cohorts

Biospecimen retention

Samples without DNA

Biospecimen description

All Blood samples and Urine Samples will be shipped to Covance Geneva Central Lab

Target sample size

10

Actual enrollment target size

2

Date of first enrollment: Type

Actual

Date of first enrollment: Date

07/08/2019

Date of study closure: Type

Actual

Date of study closure: Date

26/02/2021

Recruitment status

Recruiting

Recruitment status: Specify

Date of completion

30/06/2020

IPD sharing statement plan

No

IPD sharing statement description

Novartis is committed to sharing with qualified external researchers, access to patient-level data and supporting clinical documents from eligible studies. These requests are reviewed and approved by an independent review panel on the basis of scientific merit. All data provided is anonymized to respect the privacy of patients who have participated in the trial in line with applicable laws and regulations.

This trial data availability is according to the criteria and process described on www.clinicalstudydatarequest.com

Additional data URL

https://clinicaltrials.gov/ct2/show/record/NCT03814746?term=cseg101a2301&rank=1

Admin comments

Trial status

Approved

Secondary Identifying Numbers

Full name of issuing authority	Secondary identifying number
Clinicaltrials.gov	NCT03814746

Sources of Monetary or Material Support

Name
Novartis Pharma Services Inc.

Secondary Sponsors

Name
NA

Contact for Public/Scientific Queries

Contact type	Contact full name	Address	Country	Telephone	Email	Affiliation
Public	Adlette Inati	Tripoli	Lebanon	03228033	adlette.inati@lau.edu.lb	Nini Hospital
Scientific	Hind Khairallah	Sin El Fil	Lebanon	+961 1 512002 Ext. 271	Hind.Khairallah@fattal.com.lb	Khalil Fattal et Fils s.a.l.

Centers/Hospitals Involved in the Study

Center/Hospital name	Name of principles investigator	Principles investigator speciality	Ethical approval
Nini Hospital	Adlette Inati	Hematology	Approved

Ethics Review

Ethics approval obtained	Approval date	Contact name	Contact email	Contact phone
Nini Hospital	20/05/2019	Nabil Kabbara	Nabil.kabbara@hopitalnini.com	+961 (0) 6 431 400 ext 1062

Countries of Recruitment

Name
Lebanon
Belgium
Netherlands
United Kingdom
United States of America
Jordan

Health Conditions or Problems Studied

Condition	Code	Keyword
Sickle Cell Disease	Sickle-cell disorders (D57)	SCD

Interventions

Intervention	Description	Keyword
ICF, Lab tests, IMP , Questionnaires	ICF, Lab tests, IMP , Questionnaires	ICF, Lab tests, IMP , Questionnaires

Primary Outcomes

Name	Time Points	Measure
Rate of vaso-occlusive crisis (VOC) events leading to	1 year	1 year
To compare the efficacy of 5.0 mg/kg versus placebo and 7.5 mg/kg of crizanlizumab versus placebo in addition to standard of care	1 year	1 year

Key Secondary Outcomes

Name	Time Points	Measure
•Rate of all VOCs leading to healthcare visit and treated at home	1 year, 5 years	1 year, 5 years
•Number of days with VOC leading to healthcare visit	1 year	1 year

Trial Results

Summary results

Study results globally

Date of posting of results summaries

Date of first journal publication of results

Results URL link

Baseline characteristics

Participant flow

Adverse events

Outcome measures

URL to protocol files