RADIANT 4 - Everolimus Plus Best Supportive Care vs Placebo Plus Best Supportive Care in the Treatment of Patients With Advanced Neuroendocrine Tumors (GI or Lung Origin)

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Main Information

Primary registry identifying number
LBCTR2020011379

Protocol number
CRAD001T2302

MOH registration number
262/ص

Study registered at the country of origin
Yes

Study registered at the country of origin: Specify

Type of registration
Retrospective

Type of registration: Justify
This study was submitted prior to LBCTR initiation

Date of registration in national regulatory agency
13/01/2015

Primary sponsor
Novartis Pharmaceuticals

Primary sponsor: Country of origin
Novartis Pharmaceuticals

Date of registration in primary registry
04/01/2021

Date of registration in national regulatory agency
13/01/2015

Public title
RADIANT 4 - Everolimus Plus Best Supportive Care vs Placebo Plus Best Supportive Care in the Treatment of Patients With Advanced Neuroendocrine Tumors (GI or Lung Origin)

Acronym

Scientific title
A Randomized, Double-blind, Multicenter, Phase III Study of Everolimus (RAD001) Plus Best Supportive Care Versus Placebo Plus Best Supportive Care in the Treatment of Patients With Advanced NET of GI or Lung Origin

Acronym

Brief summary of the study: English
The purpose of this study is to compare the antitumor activity of everolimus plus best supportive care versus placebo plus best supportive care in patients with advanced nonfunctional neuroendocrine tumor of gastrointestinal or lung origin.

Brief summary of the study: Arabic
دراسة عشوائية ومتعددة المراكز في المرحلة الثالثة لدواء Everolimus إيفيروليموس (RAD001) بالإضافة إلى أفضل عناية داعمة مقابل العلاج الإرضائي وأفضل عناية داعمة في علاج المرضى المصابين بحالة متقدمة من أورام الغدد الصمّ العصبية يكون مصدرها معديًا معويًا أو رئويًا - (RADIANT-4) مشع-4

Health conditions/problem studied: Specify
Advanced Nonfunctional NeuroEndocrine Tumor

Interventions: Specify
•Drug: Everolimus
After randomization, patients will receive everolimus once daily until disease progression, intolerable toxicity, or consent withdrawal

Other Name: RAD001

•Drug: Everolimus Placebo
After randomization, patients will receive everolimus placebo once daily until disease progression, intolerable toxicity, or consent withdrawal

Key inclusion and exclusion criteria: Inclusion criteria

•Pathologically confirmed, well differentiated (G1 or G2), advanced (unresectable or metastatic), neuroendocrine tumor of GI or lung origin
•No history of and no active symptoms related to carcinoid syndrome
•In addition to treatment-naive patients, patients previously treated with SSA, Interferon (IFN), one prior line of chemotherapy, and/or PRRT are allowed into the study. Pretreated patients must have progressed on or after the last treatment
•Radiological documented disease progression within 6 months prior to randomization
•Measurable disease
•WHO performance status ≤1
•Adequate bone marrow, liver and renal function

Key inclusion and exclusion criteria: Gender

Both

Key inclusion and exclusion criteria: Specify gender

Key inclusion and exclusion criteria: Age minimum

18

Key inclusion and exclusion criteria: Age maximum

99

Key inclusion and exclusion criteria: Exclusion criteria

•Patients with poorly differentiated neuroendocrine carcinoma, high-grade neuroendocrine carcinoma, adenocarcinoid, pancreatic islet cell carcinoma, insulinoma, glucagonoma, gastrinoma, goblet cell carcinoid, large cell neuroendocrine carcinoma and small cell carcinoma
•Patients with pancreatic NET or NET of origins other than GI or Lung
•Patients with history of or active symptoms of carcinoid syndrome (e.g. flushing, diarrhea)
•Patients with more than one line of prior chemotherapy
•Prior targeted therapy
•Hepatic locoregional therapy within the last 6 months
•Prior therapy with mTOR inhibitors (e.g. sirolimus, temsirolimus, deforolimus)
•Known intolerance or hypersensitivity to everolimus or other rapamycin analogs (e.g. sirolimus, temsirolimus)
•Known impairment of gastrointestinal (GI) function or GI disease that may significantly alter the absorption of oral everolimus
•Uncontrolled diabetes mellitus as defined by HbA1c >8% despite adequate therapy

•Patients who have any severe and/or uncontrolled medical conditions such as:
◦unstable angina pectoris, symptomatic congestive heart failure, myocardial infarction ≤6 months prior to randomization, serious uncontrolled cardiac arrhythmia
◦active or uncontrolled severe infection
◦liver disease such as cirrhosis, decompensated liver disease, and chronic hepatitis (i.e. quantifiable HBV-DNA and/or positive HbsAg, quantifiable HCV-RNA)

•Chronic treatment with corticosteroids or other immunosuppressive agents
•Known history of HIV seropositivity
•Pregnant or nursing (lactating) women

Other protocol-defined inclusion/exclusion criteria may apply.

Type of study

Interventional

Type of intervention

Pharmaceutical

Type of intervention: Specify type

N/A

Trial scope

Therapy

Trial scope: Specify scope

N/A

Study design: Allocation

Randomized controlled trial

Study design: Masking

Open (masking not used)

Study design: Control

Placebo

Study phase

3

Study design: Purpose

Treatment

Study design: Specify purpose

N/A

Study design: Assignment

Study design: Specify assignment

Parallel

N/A

IMP has market authorization

Yes, Lebanon and Worldwide

IMP has market authorization: Specify

Austria, Belgium, Canada, China, Colombia, Czechia, Germany, ...

Name of IMP

everolimus (RAD001)

Year of authorization

2010

Month of authorization

5

Type of IMP

Cell therapy

Pharmaceutical class

proliferation signal inhibitor in the mammalian target of rapamycin (mTOR)

Therapeutic indication

proliferation signal inhibitor in the mammalian target of rapamycin (mTOR)

Therapeutic benefit

Progression Free Survival (PFS)

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

Samples are sent to central quintiles laboratories

Target sample size

5

Actual enrollment target size

5

Date of first enrollment: Type

Actual

Date of first enrollment: Date

25/09/2012

Date of study closure: Type

Date of study closure: Date

Actual

31/12/2021

Recruitment status

Complete

Recruitment status: Specify

Date of completion

17/07/2013

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/NCT01524783

Admin comments

Trial status

Approved

Secondary Identifying Numbers

Full name of issuing authority	Secondary identifying number
Clinical trials.gov	NCT01524783

Sources of Monetary or Material Support

Name
Novartis Pharmaceuticals

Secondary Sponsors

Name
NA

Contact for Public/Scientific Queries

Contact type	Contact full name	Address	Country	Telephone	Email	Affiliation
Public	Ali Shamseddin	Beirut	Lebanon	03344277	as04@aub.edu.lb	American University of beirut Medical Center
Scientific	Hind Khairallah	Sin elfil	Lebanon	01512002#271	Hind.Khairallah@fattal.com.lb	Khalil Fattal et Fils s.a.l.
Public	Joseph Kattan	Beirut	Lebanon	011424942	jkattan62@hotmail.com	Hotel Dieu De France

Centers/Hospitals Involved in the Study

Center/Hospital name	Name of principles investigator	Principles investigator speciality	Ethical approval
American University of Beirut Medical Center	Ali Shamseddin	Hematology	Approved
Hotel Dieu De France	Joseph Kattan	Hematology	Approved

Ethics Review

Ethics approval obtained	Approval date	Contact name	Contact email	Contact phone
American University of Beirut Medical Center	11/03/2013	Fuad Ziyadeh	fz05@aub.edu.lb	961 (0) 1 350 000 ext:5445
Hotel Dieu de France	07/05/2012	Nancy Alam	nancy.alam@usj.edu.lb	961 (0) 1 421000 ext 2335

Countries of Recruitment

Name
Lebanon
Australia
Belgium
Canada
China
Colombia
Greece
Italy
Norway
Saudi Arabia
Turkey
United Arab Emirates
United States of America

Health Conditions or Problems Studied

Condition	Code	Keyword
Neuroendocrine tumor	Endocrine gland, unspecified (C75.9)	Neuroendocrine tumor

Interventions

Intervention	Description	Keyword
ICF, Lab tests , physical exam, radiology	ICF, Lab tests , physical exam, radiology	ICF, Lab tests , physical exam, radiology

Primary Outcomes

Name	Time Points	Measure
Progression Free Survival (PFS) Based on Central Radiology Assessment Per Kaplan-Meier	18 months	18 months

Key Secondary Outcomes

Name	Time Points	Measure
•Overall Survival (OS) Using Kaplan-Meier	18 Months	18 Months
•Overall Safety Evaluation of Everolimus Versus Placebo	5 years	5 years

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