Trial record **1 of 1** for: CPDR001F2301

Previous Study | Return to List | Next Study

A Study of the Anti-PD1 Antibody PDR001, in Combination With Dabrafenib and Trametinib in Advanced Melanoma

ClinicalTrials.gov Identifier: NCT02967692

Recruitment Status : Recruiting
First Posted : November 18, 2016
Last Update Posted : March 16, 2018

See **Contacts and Locations**

Sponsor:
Novartis Pharmaceuticals

Information provided by (Responsible Party):
Novartis (Novartis Pharmaceuticals)

Study Details | **Tabular View** | **No Results Posted** | Disclaimer | How to Read a Study Record

Study Description

Go to

Brief Summary:
To evaluate the safety and efficacy of the combination of an anti-PD-1 antibody (PDR001), a BRAF inhibitor (dabrafenib) and a MEK inhibitor (trametinib) in unresectable or metastatic BRAF V600 mutant melanoma

Condition or disease	Intervention/treatment	Phase
Melanoma	Biological: PDR001 Other: Placebo Drug: Dabrafenib Drug: Trametinib	Phase 3

Study Design

Go to

Study Type : Interventional (Clinical Trial)
Estimated Enrollment : 538 participants

Allocation:	Randomized
Intervention Model:	Parallel Assignment
Masking:	Quadruple (Participant, Care Provider, Investigator, Outcomes Assessor)
Primary Purpose:	Treatment
Official Title:	A Randomized, Double-blind, Placebo-controlled, Phase III Study Comparing the Combination of PDR001, Dabrafenib and Trametinib Versus Placebo, Dabrafenib and Trametinib in Previously Untreated Patients With Unresectable or Metastatic BRAF V600 Mutant Melanoma
Actual Study Start Date :	February 17, 2017
Estimated Primary Completion Date :	July 19, 2019
Estimated Study Completion Date :	February 13, 2020

Resource links provided by the National Library of Medicine

MedlinePlus related topics: Melanoma

Drug Information available for: Trametinib Dabrafenib

Genetic and Rare Diseases Information Center resources: Neuroendocrine Tumor Neuroepithelioma

U.S. FDA Resources

Arms and Interventions

Go to

Arm	Intervention/treatment
Experimental: Investigational treatment arm Part 1: Safety run-in Up to 18 evaluable patients with previously untreated unresectable or metastatic BRAF V600 mutated melanoma will be enrolled and treated at different dose levels to determine the recommended Phase 3 regimen of PDR001 in combination with dabrafenib and trametinib. Part 2: Biomarker cohort Approximately 20 patients with previously unresectable or metastatic BRAF V600 mutated melanoma will be enrolled to describe changes in the immune microenvironment and biomarker modulations Part 3: Randomized double blind Approximately 500 patients with previously untreated unresectable and metastatic BRAF V600 mutated melanoma will be enrolled to compare the anti-tumor activity of PDR001 in combination with dabrafenib and trametinib versus placebo plus dabrafenib and trametinib.	Biological: PDR001 PDR001 will be supplied in vial in liquid or lyophilized pharmaceutical form. PDR001 will be administered via intravenous infusion over 30 minutes (up to 2 hours) once every 4 or 8 weeks. Drug: Dabrafenib Dabrafenib will be provided by the sponsor to the investigative site or supplied locally as commercially available. Dabrafenib will be administered orally twice daily (150 mg BID) for Days 1-28 of a 28-day cycle. Other Name: Tafinlar® Drug: Trametinib Trametinib will be provided by the sponsor to the investigative site or supplied locally as commercially available. Trametinib will be administered orally once daily (2 mg QD) for Days 1-28 of a 28-day cycle. Other Name: Mekinist®
Placebo Comparator: Placebo comparator arm Matching placebo in combination with dabrafenib and trametinib	Other: Placebo Placebo will be a Dextrose 5% in water (D5W) infusion supplied by the site.

	Other Name: Placebo control Drug: Dabrafenib Dabrafenib will be provided by the sponsor to the investigative site or supplied locally as commercially available. Dabrafenib will be administered orally twice daily (150 mg BID) for Days 1-28 of a 28-day cycle. Other Name: Tafinlar® Drug: Trametinib Trametinib will be provided by the sponsor to the investigative site or supplied locally as commercially available. Trametinib will be administered orally once daily (2 mg QD) for Days 1-28 of a 28-day cycle. Other Name: Mekinist®

Outcome Measures

Go to

Primary Outcome Measures :

1. Safety Run-In (Part 1): Incidence of dose limiting toxicities (DLTs) [Time Frame: 8 weeks]

   Incidence of DLTs during the first 8 weeks of treatment with PDR001 in combination of dabrafenib and trametinib

2. Biomarker cohort (Part 2): Immune microenvironment and biomarker modulation [Time Frame: 2 years]

   Changes in PD-L1 levels and CD8+ cells upon treatment with PDR001 in combination with dabrafenib and trametinib

3. Randomized (Part 3): Progression-Free Survival (PFS), investigator assessed by RECIST 1.1 [Time Frame: Up to disease progression or death due to any cause, whichever occurs first (5 years)]

   Progression-free survival is defined as the time from the date of first dose to the date of the first documented radiological progression using RECIST 1.1 response criteria

Secondary Outcome Measures :

1. Overall survival [Time Frame: Up to death due to any cause (5 years)]

   Overall survival is defined as the time from date of randomization to date of death due to any cause

2. Overall response rate [Time Frame: Up to disease progression or death due to any cause, whichever occurs first (5 years)]

   ORR is defined as the proportion of subjects with confirmed best overall response of complete response (CR) or partial response (PR), as per investigator's assessment by RECIST 1.1

3. Duration of response [Time Frame: Up to disease progression or death due to any cause, whichever occurs first (5 years)]

   Duration of response is defined as the time from first documented response of CR or PR to date of first

documented progression or death, according to RECIST 1.1 criteria

4. Disease control rate [Time Frame: Up to disease progression or death due to any cause, whichever occurs first (5 years)]

   Disease control rate is defined as the proportion of patients with CR or PR or subjects with SD lasting for a duration of at least 24 weeks as per local review according to RECIST 1.1 criteria

5. Global health status/quality of life score of the EORTC QLQ-C30 [Time Frame: Up to 60 days post progression (5 years)]

   Patient's health-related quality of life

6. Global health status/quality of life score of the FACT-M subscale [Time Frame: Up to 60 days post progression (5 years)]

   Patient's health-related quality of life

7. Global health status/quality of life score of the EQ-5D-5L [Time Frame: Up to 60 days post progression (5 years)]

   Patient's health-related quality of life

8. Time to 10 point definitive deterioration in overall quality of life score from EORTC QLQ-C30 [Time Frame: Up to 60 days post progression (5 years)]

   Patient's health-related quality of life

9. PFS by PD-L1 expression [Time Frame: Up to disease progression or death due to any cause, whichever occurs first (5 years)]

   PFS analysis will be performed by PD-L1 subgroup (positive, negative) where a positive status is defined as having ≥ 1% expression and a negative status is defined as having < 1% expression.

   Additionally PD-L1 subgroups will also be assessed using defined by a PD-L1 expression level cut-off of 10%, where a positive status is defined as having ≥ 10% expression and a negative status is defined as having < 10% expression.

10. OS by PD-L1 expression [Time Frame: Up to disease progression or death due to any cause, whichever occurs first (5 years)]

    OS analysis will be performed by PD-L1 subgroup (positive, negative) where a positive status is defined as having ≥ 1% expression and a negative status is defined as having < 1% expression.

    Additionally PD-L1 subgroups will also be assessed using defined by a PD-L1 expression level cut-off of 10%, where a positive status is defined as having ≥ 10% expression and a negative status is defined as having < 10% expression.

Eligibility Criteria Go to

Choosing to participate in a study is an important personal decision. Talk with your doctor and family members or friends about deciding to join a study. To learn more about this study, you or your doctor may contact the study research staff using the contacts provided below. For general information, Learn About Clinical Studies.

Ages Eligible for Study:	18 Years and older (Adult, Senior)
Sexes Eligible for Study:	All
Accepts Healthy Volunteers:	No

Criteria

Inclusion criteria Part 1: Safety run-in

- Histologically confirmed, unresectable or metastatic melanoma with BRAF V600 mutation
- Aspartate transaminase (AST) < 2.5× ULN and Alanine transaminase (ALT) < 2.5× ULN
- ECOG performance status ≤ 1

Part 2: Biomarker cohort

- Histologically confirmed, unresectable or metastatic melanoma with BRAF V600 mutation
- At least two cutaneous or subcutaneous or nodal lesions for tumor sample collection
- ECOG performance status ≤ 2

Part 3: Double-blind, randomized, placebo-controlled part

- Histologically confirmed, unresectable or metastatic melanoma with BRAF V600 mutation
- ECOG performance status ≤ 2

Exclusion Criteria:

Part 1: Safety run-in

- Subjects with uveal or mucosal melanoma
- Any history of CNS metastases
- Prior systemic anti-cancer treatment for unresectable or metastatic melanoma
- Prior loco-regional treatment for unresectable or metastatic melanoma in the last 6 month
- Prior neoadjuvant and/or adjuvant therapy for melanoma completed less than 6 months
- Radiation therapy within 4 weeks prior to start of study treatment
- Active, known, suspected or a documented history of autoimmune disease

Parts 2 & 3: Biomarker cohort & double-blind, randomized, placebo-controlled part

- Subjects with uveal or mucosal melanoma
- Clinically active cerebral melanoma metastasis
- Prior systemic anti-cancer treatment for unresectable or metastatic melanoma
- Prior loco-regional treatment for unresectable or metastatic melanoma in the last 6 month
- Prior neoadjuvant and/or adjuvant therapy for melanoma completed less than 6 months
- Radiation therapy within 4 weeks prior to start of study treatment
- Active, known, suspected or a documented history of autoimmune disease

Other protocol-defined Inclusion/Exclusion may apply.

Contacts and Locations

Go to

Information from the National Library of Medicine

To learn more about this study, you or your doctor may contact the study research staff using the contact information provided by the sponsor.

Please refer to this study by its ClinicalTrials.gov identifier (NCT number): ***NCT02967692***

Contacts

Contact: Novartis Pharmaceuticals 1-888-669-6682 novartis.email@novartis.com

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Show 193 Study Locations

Sponsors and Collaborators

Novartis Pharmaceuticals

Investigators

Study Director: Novartis Pharmaceuticals Novartis Pharmaceuticals

More Information

Go to

Responsible Party: Novartis Pharmaceuticals
ClinicalTrials.gov Identifier: NCT02967692 History of Changes
Other Study ID Numbers: **CPDR001F2301**
2016-002794-35 (EudraCT Number)
First Posted: November 18, 2016 Key Record Dates
Last Update Posted: March 16, 2018
Last Verified: March 2018

Individual Participant Data (IPD) Sharing Statement:

Plan to Share IPD: Undecided

Plan 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

Studies a U.S. FDA-regulated Drug Product: Yes
Studies a U.S. FDA-regulated Device Product: No

Keywords provided by Novartis (Novartis Pharmaceuticals):

PDR001
dabrafenib
trametinib
melanoma
immunotherapy
anti-PD-1
combination treatment
malignant skin cancer
skin cancer
BRAF V600

PD 1 inhibitor
anti PD1
PD-1

unresectable BRAF V600 mutated melanoma
metastatic BRAF V600 mutated melanoma

Additional relevant MeSH terms:

Melanoma
Neuroendocrine Tumors
Neuroectodermal Tumors
Neoplasms, Germ Cell and Embryonal
Neoplasms by Histologic Type
Neoplasms
Neoplasms, Nerve Tissue
Nevi and Melanomas
Trametinib
Dabrafenib
Antineoplastic Agents
Protein Kinase Inhibitors
Enzyme Inhibitors
Molecular Mechanisms of Pharmacological Action