



A phase II open label, randomized, three-arm, multicenter study of LAG525 given in combination with spartalizumab (PDR001), or with spartalizumab and carboplatin, or with carboplatin, as first or second line therapy in patients with advanced triple-negative breast cancer

22/11/2024 12:08:40

Main Information

Primary registry identifying number

LBCTR2019020196

Protocol number

CLAG525B2101

MOH registration number

33223/2018

Study registered at the country of origin

Yes

Study registered at the country of origin: Specify**Type of registration**

Retrospective

Type of registration: Justify

LCTR was recently initiated, original file was previously submitted by Paper

Date of registration in national regulatory agency

06/08/2018

Primary sponsor

Novartis Pharma Services Inc.

Primary sponsor: Country of origin

Novartis Pharmaceuticals

Date of registration in primary registry

07/01/2020

Date of registration in national regulatory agency

06/08/2018

Public title

A phase II open label, randomized, three-arm, multicenter study of LAG525 given in combination with spartalizumab (PDR001), or with spartalizumab and carboplatin, or with carboplatin, as first or second line therapy in patients with advanced triple-negative breast cancer

Acronym**Scientific title**

A phase II open label, randomized, three-arm, multicenter study of LAG525 given in combination with spartalizumab (PDR001), or with spartalizumab and carboplatin, or with carboplatin, as first or second line therapy in patients with advanced triple-negative breast cancer

Acronym**Brief summary of the study: English**

The purpose of this study is to assess the efficacy, safety, and PK characteristics of the following three combinations: i) LAG525 + spartalizumab; ii) LAG525 + spartalizumab + carboplatin, and iii) LAG525 + carboplatin in subjects with advanced TNBC and up to one prior line of systemic treatment for metastatic disease. A thorough biomarker strategy to address key aspects of tumor immunogenicity will be implemented in the study.

Brief summary of the study: Arabic



دراسة مرحلة ثانية مفتوحة اللصاقة وعشوائية التوزيع ومتعددة المراكز من ثلاث مجموعات حول دواء LAG525 المعطى بالاشتراك مع دواء سبارتاليزوماب (PDR001) أو مع سبارتاليزوماب وكاربوبلاتين، أو مع كاربوبلاتين، كعلاج أساسي أو كعلاج خيار ثان لدى المرضى المصابين بسرطان الثدي الثلاثي، السلبي المتقدم

Health conditions/problem studied: Specify

Triple Negative Breast Cancer

Interventions: Specify

LAG525/ PDR001/ Carboplatin

Key inclusion and exclusion criteria: Inclusion criteria

- 1-Patient has advanced (loco-regionally recurrent not amenable to curative therapy or metastatic) breast cancer.
- 2-Patient must have measurable disease, i.e., at least one measurable lesion as per RECIST 1.1 criteria (Tumor lesions previously irradiated or subjected to other loco-regional therapy will only be considered measurable if disease progression at the treated site after completion of therapy is clearly documented)
- 3-Patient progressed after adjuvant or 1 prior systemic treatment in the metastatic setting. Patients with de novo metastatic disease are eligible if they received 1 prior line of therapy
- 4-Patient must have received prior systemic treatment that included taxane-based chemotherapy for adjuvant or metastatic disease
- 5-Patient must have a site of disease amenable to biopsy, and must be willing to undergo a new tumor biopsy at screening and during therapy on this study, the latter if medically feasible. Patients with an available archival tumor tissue do not need to perform a tumor biopsy at screening if patient has not received anti-cancer therapy since the biopsy was taken.
- 6-Patient has histologically and/or cytologically confirmed diagnosis of advanced TNBC (based on most recently analyzed biopsy, local lab) meeting the following criteria: HER2 negative in situ hybridization test or an IHC status of 0 or 1+, and ER and PR expression is <1 percent as determined by immunohistochemistry (IHC)

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

90

Key inclusion and exclusion criteria: Exclusion criteria

- 1-Patient has received prior immunotherapy as anticancer treatment such as anti-LAG-3, anti-PD-1, anti-PD-L1, or anti-PD-L2 antibody (any line of therapy).
- 2-Patient received prior neoadjuvant or adjuvant therapy with a platinum agent or mitomycin and experienced recurrence within 12 months after the end of the platinum-based or mitomycin containing therapy or received Platinum or mitomycin for metastatic disease
- 3-Patient has had major surgery within 14 days prior to starting study treatment or has not recovered to grade 1 or less from major side effects.
- 4-Patient with presence of CTCAE grade 2 toxicity or higher due to prior cancer therapy. Exception to this criterion; patients with any grade of alopecia are allowed to enter the study..
- 5-Patient has received radiotherapy \leq 4 weeks prior to randomization (\leq 2 weeks for limited field radiation for palliation), and has not recovered to grade 1 or better from related side effects of such therapy (with the exception of alopecia).
- 6-Patient has a known hypersensitivity to other monoclonal antibodies, platinum-containing compounds, or to any of the excipients of LAG525, spartalizumab, or carboplatin.
- 7-Patient has symptomatic central nervous system (CNS) metastases or CNS metastases that require local CNS-directed therapy (such as radiotherapy or surgery), or increasing doses of corticosteroids within the 2 weeks prior to first dose of study treatment. Patients with treated brain metastases should be neurologically stable and without CNS progression for at least 12 weeks prior to randomization and have discontinued corticosteroid treatment (with the exception of < 10 mg/day of prednisone or equivalent for an indication other than CNS metastases) for at least 4 weeks before first dose of any study treatment.

Type of study

Interventional

Type of intervention

Pharmaceutical

Type of intervention: Specify type

N/A

Trial scope

Other

Trial scope: Specify scope

Study design: Allocation

Randomized controlled trial

Study design: Masking

Open (masking not used)

Study design: Control

Active

Study phase

2

**Study design: Purpose**

Treatment

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**

LAG525

Year of authorization**Month of authorization****Type of IMP**

Immunological

Pharmaceutical class

LAG525 is a high-affinity, ligand-blocking humanized IgG4 antibody (stabilized hinge, S228P) against LAG-3 that blocks the binding of MHC Class II to LAG-3.

Therapeutic indication

Patients with triple negative breast cancer

Therapeutic benefit

'Overall response rate (ORR) per RECIST v1.1 per investigators' assessment up to 8 cycles

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 with DNA**

Biospecimen description



Central Laboratory Q2 Solutions, The Alba Campus, Rosebank, Livingston, EH54 7EG, United Kingdom

Lab Tests to be done:

Hematology Hematocrit, Hemoglobin, Platelets, White blood cells, Differential (Basophils, Eosinophils, Lymphocytes, Monocytes, Neutrophils, Bands

Chemistry Albumin, Alkaline phosphatase, ALT , AST , Gamma-glutamyl-transferase (GGT), Lactate dehydrogenase (LDH), Calcium, Magnesium, Phosphorus, Chloride, Sodium, Potassium, Creatinine, Creatinine clearance, Creatine kinase, Direct Bilirubin, Indirect Bilirubin, Total Bilirubin, Total Cholesterol, Blood Urea Nitrogen (BUN) or Urea, Uric Acid, Amylase, Lipase, Glucose Coagulation International normalized ratio [INR]), Activated partial thromboplastin time (APTT)

Thyroid TSH, Free T3 and Free T4

Hepatitis markers HBV-DNA, HBsAg, HBsAb, HBcAb, HCV RNA-PCR

Cytokines IFN- γ , IL-6, IL-1, TNF- α

Pregnancy Test serum pregnancy hCG test

Target sample size

6

Date of first enrollment: Type

Actual

Date of study closure: Type

Actual

Recruitment status

Complete

Date of completion

13/09/2019

IPD sharing statement plan

No

Additional data URL

<https://clinicaltrials.gov/ct2/show/record/NCT03499899?term=CLAG525B2101&rank=1>

Admin comments

Trial status

Approved

Actual enrollment target size

6

Date of first enrollment: Date

31/10/2018

Date of study closure: Date

23/07/2020

Recruitment status: Specify

On Hold

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.

Secondary Identifying Numbers

| Full name of issuing authority | Secondary identifying number |
|--------------------------------|------------------------------|
| Clinical Trials. gov | NCT03499899 |



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 | Joseph Kattan | Beirut | Lebanon | 009613635913 | jkattan62@hotmail.com | Hotel Dieu De France |
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| Public | Fadi Farhat | Saida | Lebanon | +9613753155 | drfadi.trials@gmail.com | Hammoud Hospital |

Centers/Hospitals Involved in the Study

| Center/Hospital name | Name of principles investigator | Principles investigator speciality | Ethical approval |
|--|---------------------------------|------------------------------------|------------------|
| Hotel Dieu De France | Dr Joseph Kattan | Hematology Oncology | Approved |
| Middle East Institute of Health | Dr Dany ABi Gerges | Hematology Oncology | Approved |
| Hammoud Hospital University Medical Center | Dr Fadi Farhat | Hematology Oncology | Approved |

Ethics Review

| Ethics approval obtained | Approval date | Contact name | Contact email | Contact phone |
|--|---------------|---------------|-----------------------------|----------------------------|
| Hotel Dieu de France | 03/07/2018 | Nancy Alam | nancy.alam@usj.edu.lb | +961 1421000 ext 2335 |
| Middle East Institute of Health | 16/08/2018 | Ahmad Ibrahim | ahmad_O_Ibrahim@hotmail.com | +961 (0) 3 233 560 |
| Hammoud Hospital University Medical Center | 16/07/2018 | Ahmad Zaatari | zaatari@hammoudhospital.com | +961 (0) 7 723111 ext 1160 |



Countries of Recruitment

| Name |
|--------------------------|
| Australia |
| Belgium |
| Canada |
| France |
| Germany |
| Hungary |
| Italy |
| Japan |
| Lebanon |
| Singapore |
| Spain |
| Thailand |
| United States of America |

Health Conditions or Problems Studied

| Condition | Code | Keyword |
|---------------|-----------------------------|----------------------|
| breast cancer | Breast, unspecified (C50.9) | Tripple negative ABC |

Interventions

| Intervention | Description | Keyword |
|---|---|---|
| Physical examination, height, weight, Hematology, Chemistry, Ferritin, Creatinine, Cleatinine Clearance, Hepatitis, Pregnancy Test, Urine Dipstick, Microscopic Urinalysis, Proteinuria, Urine Pregnancy Test, Liver function test, Ocular exam, audiometry, ECG, Electrocardiogram, PK sampling, vital signs, Growth and development | Physical examination, height, weight, Hematology, Chemistry, Ferritin, Creatinine, Cleatinine Clearance, Hepatitis, Pregnancy Test, Urine Dipstick, Microscopic Urinalysis, Proteinuria, Urine Pregnancy Test, Liver function test, Ocular exam, audiometry, ECG, Electrocardiogram, PK sampling, vital signs, Growth and development | Physical examination, height, weight, Hematology, Chemistry, Ferritin, Creatinine, Cleatinine Clearance, Hepatitis, Pregnancy Test, Urine Dipstick, Microscopic Urinalysis, Proteinuria, Urine Pregnancy Test, Liver function test, Ocular exam, audiometry, ECG, Electrocardiogram, PK sampling, vital signs, Growth and development |



Primary Outcomes

| Name | Time Points | Measure |
|---|-------------|-----------|
| Overall response rate (ORR) per RECIST v1.1 per investigators' assessment | 24 months | 24 Months |

Key Secondary Outcomes

| Name | Time Points | Measure |
|-----------------------------|-------------|-----------|
| Duration of response (DOR) | 3 years | 3 years |
| Overall Survival (OS) | 3 years | 3 years |
| Clinical Benefit Rate (CBR) | 24 months | 24 months |

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