Durvalumab and Olaparib in Metastatic or recurrent Endometrial Cancer

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The combination of PARP and PD-L1 inhibition has a synergistic effect and will result in progression free survival improvement.

Ethische beoordeling	Positief advies
Status	Werving gestopt
Type aandoening	-
Onderzoekstype	Interventie onderzoek

Samenvatting

ID

NL-OMON20492

Bron
NTR

Verkorte titel
DOMEC

Aandoening

Advanced (recurrent, refractory or metastatic) Endometrial cancer or carcinosarcoma of the uterus

Ondersteuning

Primaire sponsor: Leiden University Medical Center
Overige ondersteuning: Sponsor LUMC, partial funding by LUMC and co-founding by AstraZeneca

Onderzoeksproduct en/of interventie

Uitkomstmaten

Primaire uitkomstmaten

Progression free survival

Toelichting onderzoek

Achtergrond van het onderzoek

The DOMEC trial is designed as a DGOG, prospective, multi-center, phase II study for 55 patients with advanced (recurrent, refractory or metastatic) endometrial cancer, including carcinosarcoma of the uterus. Patients must have had one prior platinum-based chemotherapeutic regimen or not be able/willing to get chemotherapy. The aim is to investigate the efficacy of the combination therapy of olaparib tablets 300mg twice daily and durvalumab 1500mg i.v. every 4 weeks in terms of progression free survival. Secondary objectives are to investigate objective response rate, overall survival, safety and predictive biomarkers.

Doel van het onderzoek

The combination of PARP and PD-L1 inhibition has a synergistic effect and will result in progression free survival improvement.

Onderzoeksopzet

Baseline assessment exists of medical history including toxicity assessment, blood chemistry, hematological screening, dipstick urinalysis, a pregnancy test (in women of child-bearing potential), ECG, imaging (e.g. CT thorax/abdomen or MRI) and complete physical examination (incl. height, weight, WHO performance status and vital signs). Optionally, extra tumor biopsy will be performed for RAD51 testing (only at baseline) and at baseline, 6 and 12 weeks extra bloodsamples for immunomonitoring (50cc). Every 4 weeks during and at the end of therapy physical examination, blood chemistry and hematology will be performed by the subject’s own oncologist. Imaging will be repeated every three months. Every three months after last dose WHO, hematology and chemistry and tumor assessment will be reported.

Onderzoeksproduct en/of interventie

olaparib tablets 300mg twice daily orally and durvalumab 1500mg by IV infusion every 4 weeks

Contactpersonen

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Deelname eisen

Belangrijkste voorwaarden om deel te mogen nemen (Inclusiecriteria)

To be included into the DOMEC-trial, patients must be (1) at least 18 years old, (2) have a WHO performance score of 0-1, (3) weight more than 30kg and (4) have histologically confirmed diagnosis of EC or carcinosarcoma of the uterus. There must be (5) a documented progressive disease (metastatic or locally advanced) according to RECIST 1.1 criteria. (6) Disease must be not amendable to local therapy, chemotherapy and hormonal therapy (or patient is not able/willing to get chemotherapy). (7) Organ system function should be adequate, defined as adequate bone marrow function (Haemoglobin ≥ 10.0 g/dL, Absolute neutrophil count (ANC) ≥ 1.5 x 10^9 /L, Platelet count ≥ 100 x 10^9 /L), liver function (Total bilirubin ≤ 1.5 x institutional upper limit of normal (ULN), Aspartate aminotransferase (AST) and Alanine aminotransferase (ALT) ≤ 2.5 x ULN (in case of lever metastases ≤ 5x ULN) and kidney function (creatinine clearance ≥51 mL/min calculated according to Cockcroft-Gault or 24 hour urine clearance). (8) Life expectancy must be at least 16 weeks.

Belangrijkste redenen om niet deel te kunnen nemen (Exclusiecriteria)

Patients with (1) history of leptomeningeal carcinomatosis, symptomatic brain metastases (uncontrolled despite of corticosteroids) or spinal cord compression are not eligible. Other exclusion criteria are (2) severe concomitant diseases; (3) active or prior documented autoimmune or inflammatory disorders; (4) active primary immunodeficiency; (5) active infections including tuberculosis, HIV, hepatitis B or C or (6) other malignant disease (except adequately treated non-melanoma skin cancer, lentigo maligna or carcinoma in situ without evidence of disease). (7) Prior treatment with PARP, PD1 or PD-L1 inhibitor; (8) QTc interval or family history of long QT syndrome; (9) severe psychiatric illness; (10) irreversible grade ≥2 toxicity from previous anti-cancer therapy; (11) major surgery in the last 2 weeks; (12) prior allogeneic bone marrow transplantation or double umbilical cord blood transplantation; (13) inability to swallow oral medication; (14) concurrent treatment with another investigational agent during the conduct of the trial or (15) expected or known intolerance to olaparib or

durvalumab will prohibit inclusion; as well as (16) pregnancy or breast feeding.

Onderzoeksopzet

Opzet

Type:	Interventie onderzoek
Onderzoeksmodel:	Anders
Toewijzing:	N.v.t. / één studie arm
Blindering:	Open / niet geblindeerd
Controle:	N.v.t. / onbekend

Deelname

Nederland

Status:	Werving gestopt
(Verwachte) startdatum:	01-05-2019
Aantal proefpersonen:	55
Type:	Werkelijke startdatum

Voornemen beschikbaar stellen Individuele Patiënten Data (IPD)

Wordt de data na het onderzoek gedeeld: Nog niet bepaald

Ethische beoordeling

Positief advies

Datum:	07-03-2019
Soort:	Eerste indiening

Registraties

Opgevolgd door onderstaande (mogelijk meer actuele) registratie

Geen registraties gevonden.

Andere (mogelijk minder actuele) registraties in dit register

Geen registraties gevonden.

In overige registers

Register	ID
NTR-new	NL7569
Ander register	METC LUMC : P19.019

Resultaten