

# Durvalumab and Olaparib in Metastatic or recurrent Endometrial Cancer

Gepubliceerd: 07-03-2019 Laatste bijgewerkt: 13-12-2022

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

|                             |                       |
|-----------------------------|-----------------------|
| <b>Ethische beoordeling</b> | Positief advies       |
| <b>Status</b>               | Werving gestopt       |
| <b>Type aandoening</b>      | -                     |
| <b>Onderzoekstype</b>       | Interventie onderzoek |

## Samenvatting

### ID

NL-OMON20492

### Bron

Nationaal Trial Register

### 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 DOMEK 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

## Publiek

LUMC  
Judith Kroep

+31-71-5263464

## Wetenschappelijk

LUMC

Judith Kroep

+31-71-5263464

## Deelname eisen

### Belangrijkste voorwaarden om deel te mogen nemen (Inclusiecriteria)

To be included into the DOME 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  $\geq 10.0$  g/dL, Absolute neutrophil count (ANC)  $\geq 1.5 \times 10^9$  /L, Platelet count  $\geq 100 \times 10^9$  /L), liver function (Total bilirubin  $\leq 1.5 \times$  institutional upper limit of normal (ULN), Aspartate aminotransferase (AST) and Alanine aminotransferase (ALT)  $\leq 2.5 \times$  ULN (in case of liver metastases  $\leq 5 \times$  ULN) and kidney function (creatinine clearance  $\geq 51$  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  $\geq 2$  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 |
| Blinding:        | 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