Clinical trial results:
A Phase 2, Open-Label, Single-Arm Study of Pamiparib (BGB-290) for the Treatment of Patients with Metastatic Castration-Resistant Prostate Cancer (mCRPC) with Homologous Recombination Deficiency (HRD)

Summary

EudraCT number	2018-002587-28
Trial protocol	GB ES
Global end of trial date	02 September 2020

Results information

Result version number	v2 (current)
This version publication date	03 December 2021
First version publication date	18 August 2021
Version creation reason	• New data added to full data set Initially since this study was a terminated study and sample availability was low, data were not analyzed.

Trial information

Trial identification

Sponsor protocol code	BGB-290-202

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT03712930
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	BeiGene, Ltd
Sponsor organisation address	BeiGene, Ltd., c/o BeiGene USA, Inc. 2955 Campus Drive, Suite 200 San Mateo, California , United States, 94403
Public contact	BeiGene DDT Call Center, BeiGene, 1 877-828-5568, clinicaltrials@beigene.com
Scientific contact	clinicaltrials@beigene.com, BeiGene, 1 877-828-5568, clinicaltrials@beigene.com

Notes:

Paediatric regulatory details

Is trial part of an agreed paediatric investigation plan (PIP)	No
Does article 45 of REGULATION (EC) No 1901/2006 apply to this trial?	No
Does article 46 of REGULATION (EC) No 1901/2006 apply to this trial?	No

Notes:

Results analysis stage

Analysis stage	Final
Date of interim/final analysis	11 November 2020
Is this the analysis of the primary completion data?	Yes
Primary completion date	06 August 2020
Global end of trial reached?	Yes
Global end of trial date	02 September 2020
Was the trial ended prematurely?	Yes

Notes:

General information about the trial

Main objective of the trial:

To evaluate the efficacy of pamiparib in patients with metastatic castration-resistant prostate cancer (mCRPC) positive for circulating tumor cells (CTC) with homologous recombination deficiency (CTC-HRD-positive; per Epic Sciences assay), as per Prostate Cancer Clinical Trials Working Group [PCWG3] criteria.

Protection of trial subjects:

This trial was designed and monitored in accordance with Sponsor procedures, which comply with the ethical principles of GCP as required by the major regulatory authorities, and in accordance with the Declaration of Helsinki.
The IEC/IRB-approved ICF was signed and dated by the subject or the subject's legally authorized representative before his or her participation in the study. A copy of each signed ICF was provided to the subject or the subject's legally authorized representative. All signed and dated ICFs were retained in each patient's study file or in the site file. For any updated or revised ICFs, written informed consent was obtained using the IEC/IRB-approved updated/revised ICFs for continued participation in the study.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	05 February 2019
Long term follow-up planned	Yes
Long term follow-up rationale	Efficacy
Long term follow-up duration	2 Years
Independent data monitoring committee (IDMC) involvement?	No

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Australia: 6
Country: Number of subjects enrolled	United States: 6
Country: Number of subjects enrolled	Spain: 1
Worldwide total number of subjects	13
EEA total number of subjects	1

Notes:

Subjects enrolled per age group

In utero	0
Preterm newborn - gestational age < 37 wk	0

Newborns (0-27 days)	0
Infants and toddlers (28 days-23 months)	0
Children (2-11 years)	0
Adolescents (12-17 years)	0
Adults (18-64 years)	4
From 65 to 84 years	9
85 years and over	0

Subject disposition

Recruitment

Recruitment details:

Eighteen subjects were screened; 5 subjects failed screening and 13 subjects were dosed.

Pre-assignment

Screening details:

All subjects enrolled had unknown BRCA1/2 status at study entry and were assigned to cohorts 1B or 2B; there were no subjects known to be BRCA1/2 positive at enrollment.

Period 1

Period 1 title	Overall (overall period)
Is this the baseline period?	Yes
Allocation method	Not applicable
Blinding used	Not blinded

Arms

Arm title	Pamiparib

Arm description:

Participants received 60 mg pamiparib orally twice daily

Arm type	Experimental
Investigational medicinal product name	Pamiparib
Investigational medicinal product code	
Other name	BGB-290
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

60 mg pamiparib capsule orally twice daily

Number of subjects in period 1	Pamiparib
Started	13
Completed	0
Not completed	13
Physician decision	1
Death	7
Progressive Disease	2
Study terminated by sponsor	3

Baseline characteristics

Reporting groups	
Reporting group title	Overall

Reporting group description: -

Reporting group values	Overall	Total	
Number of subjects	13	13	
Age categorical Units: Subjects			
Adults (18-64 years)	4	4	
From 65 to 84 years	9	9	
85 years and over	0	0	
Age continuous Units: years			
arithmetic mean	69.3		
standard deviation	± 9.49	-	
Gender categorical Units: Subjects			
Female	0	0	
Male	13	13	

End points

End points reporting groups

Reporting group title	Pamiparib

Reporting group description:

Participants received 60 mg pamiparib orally twice daily

Primary: Objective Response Rate (ORR) Determined by Independent Central Review

End point title	Objective Response Rate (ORR) Determined by Independent Central Review[1]

End point description:

ORR is the percentage of participants with a best objective response of complete response (CR) or partial response (PR) confirmed at a subsequent timepoint ≥ 4 weeks later by an Independent Review Committee (IRC). Efficacy-Evaluable Analysis Set: includes all participants in the Safety Analysis Set who had measurable disease at baseline and at least 1 post-baseline tumor assessment, unless they permanently discontinue pamiparib or the study early due to clinical progression or death before tumor assessment. Participants with available data were included in the analysis.

End point type	Primary

End point timeframe:

Up to 1 year and 6 months

Notes:

[1] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This study stopped prematurely and only 13 subjects enrolled. No summary statistics are available given the limited data from the small number of evaluable subjects.

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	12			
Units: Percentage of participants				
number (not applicable)	0			

Statistical analyses

No statistical analyses for this end point

Primary: Prostate-Specific Antigen (PSA) Response Rate

End point title	Prostate-Specific Antigen (PSA) Response Rate[2]

End point description:

PSA response rate is defined as the percentage of participants with PSA decline ≥ 50% from baseline [confirmed by a second PSA value ≥ 3 weeks later] for CTC-HRD-positive patients with or without measurable disease. PSA-Evaluable Analysis Set: includes all participants in the Safety Analysis Set with ≥3 rising PSA levels with ≥ 1 week between determinations and screening PSA ≥ 2 µg/L) and who had at least 1 post-baseline PSA measurement unless they permanently discontinue pamiparib or the study early due to clinical progression or death before completed PSA assessment.

End point type	Primary

End point timeframe:

Up to 1 year and 6 months

Notes:

[2] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This study stopped prematurely and only 13 subjects enrolled. No summary statistics are available given the limited data from the small number of evaluable subjects.

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13			
Units: Percentage of Participants				
number (not applicable)	0			

Statistical analyses

No statistical analyses for this end point

Secondary: Duration of Response

End point title	Duration of Response

End point description:

DOR is defined as the time from the date of the earliest documented CR or PR (that is subsequently confirmed) to radiographic disease progression or death due to any cause, whichever occurs first. Efficacy-Evaluable Analysis Set: includes all participants in the Safety Analysis Set who had measurable disease at baseline and at least 1 post-baseline tumor assessment, unless they permanently discontinue pamiparib or the study early due to clinical progression or death before tumor assessment. Participants with available data were included in the analysis.

End point type	Secondary

End point timeframe:

Up to 1 year and 7 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	12[3]			
Units: Months	9999			

Notes:

[3] - 9999 = Not applicable; No confirmed CR/PR. Duration of Response is not applicable.

Statistical analyses

No statistical analyses for this end point

Secondary: Objective Response Rate by Investigator

End point title	Objective Response Rate by Investigator

End point description:

ORR is the percentage of participants with a best objective response of complete response (CR) or partial response (PR) confirmed at a subsequent timepoint ≥ 4 weeks later by the investigator. Efficacy-Evaluable Analysis Set: includes all participants in the Safety Analysis Set who had measurable disease

at baseline and at least 1 post-baseline tumor assessment, unless they permanently discontinue pamiparib or the study early due to clinical progression or death before tumor assessment. Participants with available data were included in the analysis.

End point type	Secondary

End point timeframe:

Up to 1 year and 6 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	12			
Units: Percentage of Participants				
number (not applicable)	0			

Statistical analyses

No statistical analyses for this end point

Secondary: Time to Objective Response by Investigator

End point title	Time to Objective Response by Investigator

End point description:

Time to objective response is defined as the time from the date of the first dose of study drug to the first documented confirmed response of CR or PR assessed by the investigator and summarized for participants who have achieved a confirmed objective response. Efficacy-Evaluable Analysis Set: includes all participants in the Safety Analysis Set who had measurable disease at baseline and at least 1 post-baseline tumor assessment, unless they permanently discontinue pamiparib or the study early due to clinical progression or death before tumor assessment. Participants with available data were included in the analysis.

End point type	Secondary

End point timeframe:

Up to 1 year and 6 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	12[4]			
Units: Months	9999			

Notes:

[4] - 9999 = Not applicable; No participants with confirmed CR/PR. Time to response is not applicable.

Statistical analyses

No statistical analyses for this end point

Secondary: Clinical Benefit Rate By Investigator

End point title	Clinical Benefit Rate By Investigator

End point description:

Clinical benefit rate is defined as the percentage of participants with a documented confirmed CR, PR, or stable disease

End point type	Secondary

End point timeframe:

Up to 1 year and 6 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	12			
Units: Percentage of participants				
number (not applicable)	25			

Statistical analyses

No statistical analyses for this end point

Secondary: Time to PSA Response

End point title	Time to PSA Response

End point description:

Time to PSA response is defined as the time from the date of the first dose of study drug to the first PSA decline ≥ 50% that is subsequently confirmed. Assessments are summarized for participants who have achieved a confirmed PSA response. PSA-Evaluable Analysis Set: includes all participants in the Safety Analysis Set with ≥3 rising PSA levels with ≥ 1 week between determinations and screening PSA ≥ 2 µg/L and who had at least 1 post-baseline PSA measurement, unless they permanently discontinue pamiparib or the study early due to clinical progression or death before completed PSA assessment.

End point type	Secondary

End point timeframe:

Up to 1 year and 6 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13[5]			
Units: Months	9999			

Notes:

[5] - 9999 = Not applicable; Participants with PSA response is 0. Time to PSA response is not applicable

Statistical analyses

No statistical analyses for this end point

Secondary: Duration of PSA Response

End point title	Duration of PSA Response

End point description:

Duration of PSA response is defined as the time from the date of the earliest documented PSA response (that is subsequently confirmed) to PSA progression or death due to any cause, whichever occurs first. PSA progression is defined as a ≥ 25% increase in PSA with an absolute increase of ≥ 2 µg/L above the nadir (or above the baseline for patients with no PSA decline after12 weeks), confirmed by a second value ≥ 3 weeks later. The nadir is defined as the lowest value at or after baseline. PSA-Evaluable Analysis Set

End point type	Secondary

End point timeframe:

Up to 1 year and 7 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13[6]			
Units: Months	9999			

Notes:

[6] - 9999 = Not applicable; Participants with PSA Response is 0. Duration of PSA response not applicable

Statistical analyses

No statistical analyses for this end point

Secondary: Time to PSA Progression

End point title	Time to PSA Progression

End point description:

Time to PSA progression is defined as the time from the date of the first dose of study drug to a ≥ 25% increase in PSA with an absolute increase of ≥ 2 ng/mL above the nadir [or above the baseline for participants with no PSA decline after 12 weeks], confirmed by a second value ≥ 3 weeks later. Safety Analysis Set : includes all participants enrolled into the study who received any dose of pamiparib.

End point type	Secondary

End point timeframe:

Up to 1 year and 7 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13			
Units: Months				
arithmetic mean (standard deviation)	3.13 (± 1.533)			

Statistical analyses

No statistical analyses for this end point

Secondary: Time to Symptomatic Skeletal Event

End point title	Time to Symptomatic Skeletal Event

End point description:

Time to symptomatic skeletal event is defined as time from the date of the first dose of study drug to the first symptomatic fracture, radiation or surgery to bone, or spinal cord compression

End point type	Secondary

End point timeframe:

Up to 1 year and 7 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13[7]			
Units: Months	9999			

Notes:

[7] - 9999 = Not applicable; Not participants had SSE

Statistical analyses

No statistical analyses for this end point

Secondary: Radiographic Progression-Free Survival

End point title	Radiographic Progression-Free Survival

End point description:

Radiographic progression-free survival is defined as the time from the date of the first dose of study drug to radiographic disease progression by IRC or death due to any cause, whichever occurs first.

End point type	Secondary

End point timeframe:

Up to 1 year and 7 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13			
Units: Months				
median (confidence interval 95%)	2.6 (1.5 to 3.7)			

Statistical analyses

No statistical analyses for this end point

Secondary: Overall survival (OS)

End point title	Overall survival (OS)

End point description:

Overall survival is defined as the time from the date of the first dose of study drug to death due to any cause. Safety Analysis Set: includes all participants enrolled in the study who received any dose of pamiparib.

End point type	Secondary

End point timeframe:

Up to 1 year and 7 months

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13			
Units: Months				
median (confidence interval 95%)	5.8 (1.6 to 5.9)			

Statistical analyses

No statistical analyses for this end point

Secondary: Number of Participants with Treatment-Emergent Adverse Events graded according to National Cancer Institute-Common Terminology Criteria for Adverse Events (NCI-CTCAE)Version 4.03

End point title	Number of Participants with Treatment-Emergent Adverse Events graded according to National Cancer Institute-Common Terminology Criteria for Adverse Events (NCI-CTCAE)Version 4.03

End point description:

End point type	Secondary

End point timeframe:

From the date of first Pamiparib dose until 30 days after the last dose or initiation of new anti-cancer therapy, whichever occurs first. (Up to 1 year and 7 months)

End point values	Pamiparib			
Subject group type	Reporting group			
Number of subjects analysed	13			
Units: Number				
Participants with at Least One TEAE	13			
Grade 3 or Higher	11			
Serious	6			
Leading to Death	1			
Leading to Treatment Discontinuation	3			
Leading to Dose Modification	11			
Leading to Dose Interruption	9			
Leading to Dose Reduction	5			
Treatment Related TEAEs	11			

Treatment Related Grade 3 or Higher	7			
Treatment Related Serious	1			
Treatment Related Leading to Death	0			
Treatment Related Leading to Discontinuation	3			
Treatment Related Leading to Dose Modification	7			
Treatment Related Leading to Dose Interruption	6			
Treatment Related Leading to Dose Reduction	3			

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information

Timeframe for reporting adverse events:

From the date of first dose of Pamiparib until 30 days after the last dose or initiation of new anti-cancer therapy, whichever occurs first (up to approximately 1 year and 7 months)

Adverse event reporting additional description:

Safety Analysis Set (SAF): included all subjects enrolled in the study who received any dose of pamiparib. The Safety Analysis Set was used for all safety analyses.

Assessment type	Non-systematic

Dictionary used

Dictionary name	MedDRA
Dictionary version	23.0

Reporting groups

Reporting group title	Pamiparib

Reporting group description:

Participants received 60 mg pamiparib orally twice daily

Serious adverse events	Pamiparib		
Total subjects affected by serious adverse events			
subjects affected / exposed	6 / 13 (46.15%)		
number of deaths (all causes)	9		
number of deaths resulting from adverse events	1		
Nervous system disorders			
Hemiparesis			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	2 / 13 (15.38%)		
occurrences causally related to treatment / all	1 / 2		
deaths causally related to treatment / all	0 / 0		
Peripheral swelling			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Blood and lymphatic system disorders			

Anaemia				
subjects affected / exposed	1 / 13 (7.69%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Febrile neutropenia				
subjects affected / exposed	1 / 13 (7.69%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Neutropenia				
subjects affected / exposed	1 / 13 (7.69%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Pancytopenia				
subjects affected / exposed	1 / 13 (7.69%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Respiratory, thoracic and mediastinal disorders				
Respiratory failure				
subjects affected / exposed	1 / 13 (7.69%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Renal and urinary disorders				
Hydronephrosis				
subjects affected / exposed	1 / 13 (7.69%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Ureteric obstruction				
subjects affected / exposed	1 / 13 (7.69%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Musculoskeletal and connective tissue disorders				
Back pain				

subjects affected / exposed	1 / 13 (7.69%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Musculoskeletal chest pain			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Infections and infestations			
Oesophageal candidiasis			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		

Frequency threshold for reporting non-serious adverse events: 0 %

Non-serious adverse events	Pamiparib		
Total subjects affected by non-serious adverse events			
subjects affected / exposed	13 / 13 (100.00%)		
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Cancer pain			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
General disorders and administration site conditions			
Fatigue			
subjects affected / exposed	5 / 13 (38.46%)		
occurrences (all)	5		
Asthenia			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	2		
Reproductive system and breast			

disorders Pelvic pain subjects affected / exposed occurrences (all)	1 / 13 (7.69%) 2		
Respiratory, thoracic and mediastinal disorders Oropharyngeal pain subjects affected / exposed occurrences (all)	1 / 13 (7.69%) 1		
Investigations Weight decreased subjects affected / exposed occurrences (all) Blood alkaline phosphatase increased subjects affected / exposed occurrences (all) Blood creatinine increased subjects affected / exposed occurrences (all) Aspartate aminotransferase increased subjects affected / exposed occurrences (all) Blood lactate dehydrogenase increased subjects affected / exposed occurrences (all) Platelet count decreased subjects affected / exposed occurrences (all)	3 / 13 (23.08%) 4 2 / 13 (15.38%) 2 2 / 13 (15.38%) 3 1 / 13 (7.69%) 1 1 / 13 (7.69%) 1 1 / 13 (7.69%) 1		
Nervous system disorders Dizziness subjects affected / exposed occurrences (all) Dysgeusia subjects affected / exposed occurrences (all) Headache	1 / 13 (7.69%) 1 1 / 13 (7.69%) 1		

subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Taste disorder			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	7 / 13 (53.85%)		
occurrences (all)	11		
Neutropenia			
subjects affected / exposed	2 / 13 (15.38%)		
occurrences (all)	4		
Thrombocytopenia			
subjects affected / exposed	2 / 13 (15.38%)		
occurrences (all)	2		
Leukopenia			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Eye disorders			
Dry eye			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Orbital oedema			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Gastrointestinal disorders			
Nausea			
subjects affected / exposed	7 / 13 (53.85%)		
occurrences (all)	8		
Diarrhoea			
subjects affected / exposed	4 / 13 (30.77%)		
occurrences (all)	4		
Vomiting			

subjects affected / exposed	4 / 13 (30.77%)		
occurrences (all)	4		
Abdominal pain			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Constipation			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Dry mouth			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Dyspepsia			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Gastrooesophageal reflux disease			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Rectal haemorrhage			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Rectal tenesmus			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Skin and subcutaneous tissue disorders			
Decubitus ulcer			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Renal and urinary disorders			
Haematuria			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Musculoskeletal and connective tissue disorders			
Pain in extremity			
subjects affected / exposed	2 / 13 (15.38%)		
occurrences (all)	2		
Back pain			

subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Flank pain			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Musculoskeletal pain			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Myalgia			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Infections and infestations			
Urinary tract infection			
subjects affected / exposed	2 / 13 (15.38%)		
occurrences (all)	2		
Oral candidiasis			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	6 / 13 (46.15%)		
occurrences (all)	6		
Dehydration			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Hypocalcaemia			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	2		
Hypophosphataemia			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		
Malnutrition			
subjects affected / exposed	1 / 13 (7.69%)		
occurrences (all)	1		

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
16 October 2018	To incorporate additional changes based on feedback from the United States Food and Drug Administration (US FDA) To incorporate changes per BeiGene protocol template for better consistency across multiple studies Minor editorial and formatting changes have been made to enhance clarity and readability;

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? Yes

Date	Interruption	Restart date
27 May 2020	Due to changes in standard of care for mCRPC, slow enrollment and lack of responses in this heavily pretreated patient population, the study was terminated.	-

Notes:

Limitations and caveats

None reported