Clinical trial results:
An Open-Label Extension of BPS-MR-PAH-201 in Pulmonary Arterial Hypertension (PAH) Patients

Summary

EudraCT number	2008-006833-29
Trial protocol	IE BE
Global end of trial date	26 November 2013

Results information

Result version number	v1 (current)
This version publication date	14 June 2020
First version publication date	14 June 2020
Summary attachment (see zip file)	BPS-MR-PAH-202 Synopsis (bps-mr-pah-202-synopsis.pdf)

Trial information

Trial identification

Sponsor protocol code	BPS-MR-PAH-202

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT00792571
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	Lung Biotechnology PBC
Sponsor organisation address	1040 Spring Street, Silver Spring, United States, 20910
Public contact	Lung Biotechnology PBC Study Director, Lung Biotechnology PBC, 1 3016089292,
Scientific contact	Lung Biotechnology PBC Study Director, Lung Biotechnology PBC, 1 3016089292,

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	26 November 2013
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	26 November 2013
Was the trial ended prematurely?	Yes

Notes:

General information about the trial

Main objective of the trial:

This is an open-label extension study for patients who participated in the BPS-MR-PAH-201 study.

Protection of trial subjects:

This study was conducted in accordance with Good Clinical Practices, the ethical principles that have their origin in the Declaration of Helsinki, and Title 21 of the Code of Federal Regulations Sections 50, 56, and 312.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	11 March 2009
Long term follow-up planned	No
Independent data monitoring committee (IDMC) involvement?	No

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Belgium: 4
Country: Number of subjects enrolled	Ireland: 2
Country: Number of subjects enrolled	United States: 12
Worldwide total number of subjects	18
EEA total number of subjects	6

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)	16
From 65 to 84 years	2
85 years and over	0

Subject disposition

Recruitment

Recruitment details:

Participants with pulmonary artery hypertension (PAH) who had completed lead in study BPS-MR-PAH-201 were enrolled in this study

Pre-assignment

Screening details:

A Protocol Amendment was to include an optional arm investigating Beraprost Sodium Modified Release Tablets administered four times daily (QID), however, no participants were enrolled into this arm.

Period 1

Period 1 title	Overall Study (overall period)
Is this the baseline period?	Yes
Allocation method	Non-randomised - controlled
Blinding used	Not blinded

Arms

Arm title	Beraprost Sodium

Arm description:

Beraprost Sodium Modified Release Tablets, 60mcg, b.i.d (twice a day dosing)

Arm type	Experimental
Investigational medicinal product name	beraprost sodium modified release (BPS-MR) tablets 60mcg
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Modified-release tablet
Routes of administration	Oral use

Dosage and administration details:

60 mcg twice a day

Number of subjects in period 1	Beraprost Sodium
Started	18
Completed	8
Not completed	10
Adverse event, serious fatal	1
Consent withdrawn by subject	1
Adverse event, non-fatal	4
Not Specified	4

Baseline characteristics

Reporting groups	
Reporting group title	Beraprost Sodium
Reporting group description:	
Beraprost Sodium Modified Release Tablets, 60mcg, b.i.d (twice a day dosing)	

Reporting group values	Beraprost Sodium	Total	
Number of subjects	18	18	
Age categorical Units: Subjects			
In utero	0	0	
Preterm newborn infants (gestational age < 37 wks)	0	0	
Newborns (0-27 days)	0	0	
Infants and toddlers (28 days-23 months)	0	0	
Children (2-11 years)	0	0	
Adolescents (12-17 years)	0	0	
Adults (18-64 years)	16	16	
From 65-84 years	2	2	
85 years and over	0	0	
Age Continuous Units: years			
arithmetic mean	47.8		
standard deviation	± 13.16	-	
Sex: Female, Male Units:			
Female	14	14	
Male	4	4	
Race (NIH/OMB) Units: Subjects			
American Indian or Alaska Native	0	0	
Asian	3	3	
Native Hawaiian or Other Pacific Islander	0	0	
Black or African American	0	0	
White	15	15	
More than one race	0	0	
Unknown or Not Reported	0	0	
Ethnicity (NIH/OMB) Units: Subjects			
Hispanic or Latino	3	3	
Not Hispanic or Latino	15	15	
Unknown or Not Reported	0	0	
Six-Minute Walk Distance			
A 6 minute walk test (6MWT) was conducted that measured how far a participant could walk in 6 continous minutes. Participants were instructed to walk as far as possible in 6 minutes, and were allowed to slow down and take breaks as needed due to symptoms.			
Units: meters			

arithmetic mean	419.5		
standard deviation	± 73.1	-	
Borg Dyspnea Score			
The Modified Borg scale was an 11 point scale with a score range of 0-10, where 0 indicated no breathlessness at all and 10 indicated maximum breathlessness.			
Units: score on a scale			
arithmetic mean	3.4		
standard deviation	± 2.44	-	

End points

End points reporting groups

Reporting group title	Beraprost Sodium

Reporting group description:

Beraprost Sodium Modified Release Tablets, 60mcg, b.i.d (twice a day dosing)

Primary: Number of Participants Reporting at Least One Treatment-Emergent Adverse Event (TEAE)

End point title	Number of Participants Reporting at Least One Treatment-Emergent Adverse Event (TEAE)[1]

End point description:

A treatment-emergent adverse event (TEAEs) is defined as an event not present prior to the initiation of the treatments or any event already present that worsens in either intensity or frequency following exposure to the treatments. AEs occurring more than 3 days after the last day study drug is taken in the study will not be included in the statistical analyses or summaries (except for subjects with adverse events leading to study drug withdrawn). Only treatment-emergent adverse events occurring during the treatment period of the BPS-MR-PAH-202 study will be summarized. Any adverse event starting prior to the first dose of study drug will be excluded from the summary analyses and only presented in the data listings. A summary of serious and all other non-serious adverse events regardless of causality is located in the Reported Adverse Events module.

End point type	Primary

End point timeframe:

Up to 56 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: The analysis of the primary endpoint was descriptive i.e. no statistical hypothesis test was performed.

End point values	Beraprost Sodium			
Subject group type	Reporting group			
Number of subjects analysed	18			
Units: Participants	18			

Statistical analyses

No statistical analyses for this end point

Primary: Number of Treatment Emergent Adverse Events Reported During The Study

End point title	Number of Treatment Emergent Adverse Events Reported During The Study[2]

End point description:

A treatment-emergent adverse event (TEAE) is defined as an event not present prior to the initiation of the treatment or any event already present that worsens in either intensity or frequency following exposure to the treatment. AEs occurring more than 3 days after the last day study drug was taken in the study was not included in the statistical analyses or summaries (except for participants with adverse events leading to study drug withdrawn). Only TEAEs that occurred during the treatment period of the BPS-MR-PAH-202 study were summarized. Any adverse event starting prior to the first dose of study drug was excluded from the summary analyses and only presented in the data listings. All efficacy results are descriptive; no statistical analysis was conducted. A summary of serious and all other non-

serious adverse events regardless of causality is located in the Reported Adverse Events module.

End point type	Primary

End point timeframe:

Up to 56 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: The analysis of the primary endpoint was descriptive i.e. no statistical hypothesis test was performed.

End point values	Beraprost Sodium			
Subject group type	Reporting group			
Number of subjects analysed	18			
Units: TEAEs	156			

Statistical analyses

No statistical analyses for this end point

Secondary: Mean Change From Baseline in Six Minutes Walk Distance (6MWD) at End of Study

End point title	Mean Change From Baseline in Six Minutes Walk Distance (6MWD) at End of Study

End point description:

The area used for the Six Minute Walk Test (6MWT) was pre-measured at a minimum of 30 meters in length and at least 2 to 3 meters in width. There were no turns or significant curves to the 6-minute walk area. The length was marked with gradations to ensure the accurate measurement of the distance walked. The area was well ventilated with air temperature controlled at 20 to 23°C. Intermittent rest periods were allowed if the participant could no longer continue. If the participant needed to rest briefly, he/she could stand or sit and then begin again when rested but the clock continued to run. At the end of 6 minutes, the tester called "stop" while simultaneously stopping the watch and then measured the distance walked. For the purposes of the 6MWT if a participant was assessed at Baseline using oxygen therapy, then all future 6MWT were conducted in the same manner. All efficacy results are descriptive; no statistical analysis was conducted.

End point type	Secondary

End point timeframe:

Baseline and 56 months

End point values	Beraprost Sodium			
Subject group type	Reporting group			
Number of subjects analysed	11			
Units: meters				
arithmetic mean (standard deviation)	10.55 (± 79.17)			

Statistical analyses

No statistical analyses for this end point

Secondary: Change From Baseline in Borg Dyspnea Score at End of Study

End point title	Change From Baseline in Borg Dyspnea Score at End of Study

End point description:

The modified 0–10 category-ratio Borg scale consists of an 11-point scale rating the maximum level of dyspnea experienced during the 6MWT. Scores range from 0 (for the best condition) and 10 (for the worst condition) with nonlinear spacing of verbal descriptors of severity corresponding to specific numbers. The participant chose the number or the verbal descriptor to reflect presumed ratio properties of sensation or symptom intensity. Baseline was defined as the last non-missing evaluation preceding the first dose of study drug in study BPS-MR-PAH-201. Only participants with both a measurement at baseline and at the given visit are presented. All efficacy results are descriptive; no statistical analysis was conducted.

End point type	Secondary

End point timeframe:

Baseline and 56 months

End point values	Beraprost Sodium			
Subject group type	Reporting group			
Number of subjects analysed	11			
Units: scores on a scale				
arithmetic mean (standard deviation)	-0.09 (± 2.84)			

Statistical analyses

No statistical analyses for this end point

Secondary: Number of Participants That Experienced Clinical Worsening During the Study

End point title	Number of Participants That Experienced Clinical Worsening During the Study

End point description:

Number of Participants that experienced Clinical Worsening in the opinion of the Investigator. Clinical Worsening was defined as any of these events following the Baseline visit: Death, Transplantation or atrial septostomy, Clinical deterioration as defined by: Hospitalization as a result of PAH symptoms or Initiation of any new PAH specific therapy (e.g. ERA, PDE-5 inhibitor, prostanoid). All efficacy results are descriptive; no statistical analysis was conducted.

End point type	Secondary

End point timeframe:

Up to 56 months

End point values	Beraprost Sodium			
Subject group type	Reporting group			
Number of subjects analysed	18			
Units: Participants				
Death	1			
New PAH Therapies	6			
Transplantation or atrial septostomy	0			
Hospitalization	0			

Statistical analyses

No statistical analyses for this end point

Secondary: Number of Participants With a Change in WHO Functional Class

End point title	Number of Participants With a Change in WHO Functional Class

End point description:

Change from Baseline in participant clinical status was recorded according to the World Health Organization (WHO) Functional Class. A change from lower to higher functional class (i.e. 'III to IV' or 'II to III') was considered as a deterioration. A change from higher to lower functional class (i.e. 'III to II' or 'II to I') was considered as an improvement. All efficacy results are descriptive; no statistical analysis was conducted.

End point type	Secondary

End point timeframe:

Baseline and 56 months

End point values	Beraprost Sodium			
Subject group type	Reporting group			
Number of subjects analysed	18			
Units: Participants				
Improved: Change from Class III to Class II	1			
No Change in Class	7			
Deteriorated: Change from Class II to Class III	4			
Not Reported	6			

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information	
Timeframe for reporting adverse events:	
From baseline to 30 days after study treatment discontinuation, up to 56 months.	
Assessment type	Systematic

Dictionary used	
Dictionary name	MedDRA
Dictionary version	16.0

Reporting groups	
Reporting group title	Beraprost Sodium
Reporting group description:	
Beraprost Sodium Modified Release Tablets, 60mcg, b.i.d (twice a day dosing)	

Serious adverse events	Beraprost Sodium		
Total subjects affected by serious adverse events			
subjects affected / exposed	7 / 18 (38.89%)		
number of deaths (all causes)	1		
number of deaths resulting from adverse events			
Injury, poisoning and procedural complications			
Post procedural haemorrhage			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cardiac disorders			
Right ventricular failure			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Atrial fibrillation			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Atrial flutter			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		

Surgical and medical procedures			
Atrial septal defect repair			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Nervous system disorders			
Presyncope			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Syncope			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Respiratory, thoracic and mediastinal disorders			
Worsening Pulmonary arterial hypertension			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Haemoptysis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Infections and infestations			
Lower respiratory tract infection			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Gastroenteritis viral			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Bronchitis			

subjects affected / exposed	1 / 18 (5.56%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		

Frequency threshold for reporting non-serious adverse events: 5 %

Non-serious adverse events	Beraprost Sodium		
Total subjects affected by non-serious adverse events			
subjects affected / exposed	18 / 18 (100.00%)		
Vascular disorders			
Flushing			
subjects affected / exposed	6 / 18 (33.33%)		
occurrences (all)	7		
Peripheral Coldness			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Vasculitis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Surgical and medical procedures			
Tooth Extraction			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Chest Pain			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Fatigue			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences (all)	2		
Generalised oedema			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		

Pain			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Pyrexia			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Immune system disorders			
Seasonal Allergy			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Respiratory, thoracic and mediastinal disorders			
Cough			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	4		
Worsening pulmonary arterial hypertension			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences (all)	3		
Dyspnoea			
subjects affected / exposed	5 / 18 (27.78%)		
occurrences (all)	5		
Nasal congestion			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Sleep apnoea syndrome			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Psychiatric disorders			
Anxiety			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences (all)	2		
Depression			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences (all)	2		
Insomnia			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	3		

Panic Attack			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Investigations			
Liver Function Test Abnormal			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Platelet Count Decreased			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Urine analysis abnormal			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Weight increased			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Cardiac disorders			
Arrhythmia			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	2		
Atrial Fibrillation			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Palpitations			
subjects affected / exposed	4 / 18 (22.22%)		
occurrences (all)	5		
Right ventricular heave			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Nervous system disorders			
Dizziness			
subjects affected / exposed	5 / 18 (27.78%)		
occurrences (all)	5		
Headache			
subjects affected / exposed	6 / 18 (33.33%)		
occurrences (all)	6		
Hypoaesthesia			

subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Presyncope			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Restless Leg Syndrome			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Syncope			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences (all)	2		
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	3		
Ear and labyrinth disorders			
Tinnitus			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Vertigo			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Eye disorders			
Astigmatism			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Cataract			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Eyelid Cyst			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Myopia			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Retinal Detachment			

subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Retinal Tear			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Gastrointestinal disorders			
Abdominal Discomfort			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Abdominal Pain			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences (all)	2		
Abdominal Pain Upper			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Defaecation urgency			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Diarrhoea			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	4		
Gastritis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Irritable Bowel Syndrome			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Nausea			
subjects affected / exposed	5 / 18 (27.78%)		
occurrences (all)	5		
Rectal Haemorrhage			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Vomiting			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	4		

Hepatobiliary disorders			
Biliary Colic			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Cholecystitis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Hepatitis acute			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Skin and subcutaneous tissue disorders			
Rash			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Urticaria			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Skin Irritation			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Renal and urinary disorders			
Haematuria			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Pollakiuria			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Muscle spasms			
subjects affected / exposed	2 / 18 (11.11%)		
occurrences (all)	2		
Muscle Tightness			

subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Muscle twitching			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Musculoskeletal Chest Pain			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Osteoporosis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Pain in extremity			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	3		
Pain in Jaw			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	3		
Infections and infestations			
Bronchiolitis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Bronchitis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Cystitis			
subjects affected / exposed	3 / 18 (16.67%)		
occurrences (all)	3		
Fungal skin infection			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Herpes zoster			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	1		
Nasopharyngitis			
subjects affected / exposed	1 / 18 (5.56%)		
occurrences (all)	2		

Pharyngitis subjects affected / exposed occurrences (all)	1 / 18 (5.56%) 1		
Sinusitis subjects affected / exposed occurrences (all)	1 / 18 (5.56%) 1		
Upper respiratory tract infection subjects affected / exposed occurrences (all)	6 / 18 (33.33%) 8		
Urinary Tract Infections subjects affected / exposed occurrences (all)	1 / 18 (5.56%) 1		
Metabolism and nutrition disorders Fluid overload subjects affected / exposed occurrences (all)	1 / 18 (5.56%) 1		
Hyperuricaemia subjects affected / exposed occurrences (all)	1 / 18 (5.56%) 1		
Hypokalemia subjects affected / exposed occurrences (all)	2 / 18 (11.11%) 2		
Hypomagnesaemia subjects affected / exposed occurrences (all)	1 / 18 (5.56%) 1		
Lactose Intolerance subjects affected / exposed occurrences (all)	1 / 18 (5.56%) 1		

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
29 June 2012	Revised to include an optional substudy investigating the safety, tolerability, and pharmacokinetics of BPS-MR tablets administered four times daily (QID). No subjects were enrolled into the optional substudy before the Sponsor discontinued the study.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported