Clinical trial results:
A RANDOMIZED, DOUBLE-BLIND, VEHICLE-CONTROLLED, PARALLEL-GROUP TRIAL TO ASSESS THE EFFICACY, SAFETY AND TOLERABILITY OF P-3073 FOR TOPICAL TREATMENT OF NAIL PSORIASIS

Summary

EudraCT number	2015-002365-34
Trial protocol	LV DE CZ PL BG GR
Global end of trial date	08 February 2017

Results information

Result version number	v1 (current)
This version publication date	20 February 2018
First version publication date	20 February 2018

Trial information

Trial identification

Sponsor protocol code	PM1434

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT02606760
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	Polichem S.A.
Sponsor organisation address	50, Val Fleuri Luxembourg, legally represented by branch in Lugano-Pazzallo, Switzerland, CH-6912
Public contact	Director, Clinical Development, Polichem S.A., +41 0919864024, maurizio.caserini@polichem.com
Scientific contact	Director, Clinical Development, Polichem S.A., +41 0919864024, maurizio.caserini@polichem.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	08 February 2017
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	08 February 2017
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

The primary objective of this Phase III study was to evaluate the efficacy of P-3073 in the treatment of nail psoriasis.

Protection of trial subjects:

The clinical trial was conducted in compliance with globally accepted standards of good clinical practice (as defined in the ICH E6 guideline for good clinical practice, January 1997), in agreement with the Declaration of Helsinki for biomedical research in humans, revised version of Edinburgh (Scotland, 2000) including the Note of Clarification in paragraph 29, Washington (2002) and the announcements for the 'Principles for Correct Implementation of Clinical Trials'.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	23 November 2015
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	Poland: 124
Country: Number of subjects enrolled	Russian Federation: 27
Country: Number of subjects enrolled	Bulgaria: 64
Country: Number of subjects enrolled	Czech Republic: 67
Country: Number of subjects enrolled	Germany: 20
Country: Number of subjects enrolled	Latvia: 56
Worldwide total number of subjects	358
EEA total number of subjects	331

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)	315
From 65 to 84 years	43
85 years and over	0

Subject disposition

Recruitment

Recruitment details:

The study was conducted in Bulgaria, Czech Republic, Germany, Latvia, Poland and Russia between 23 November 2015 (first subject first visit) and 08 February 2017 (last subject last visit).

Pre-assignment

Screening details:

A total of 378 subjects were enrolled and 358 subjects were randomised in the study and of them, 352 subjects were treated.

Period 1

Period 1 title	Overall Trial (overall period)
Is this the baseline period?	Yes
Allocation method	Randomised - controlled
Blinding used	Double blind
Roles blinded	Subject, Investigator

Arms

Are arms mutually exclusive?	Yes
Arm title	P-3073

Arm description:

Subjects with mild to moderate psoriatic fingernail/s, were treated with P-3073 nail solution as a topical application once daily for 24 weeks.

Arm type	Experimental
Investigational medicinal product name	P-3073
Investigational medicinal product code	
Other name	Calcipotriol
Pharmaceutical forms	Medicated nail lacquer
Routes of administration	Topical use

Dosage and administration details:

Subjects with mild to moderate psoriatic fingernail/s, were treated with P-3073 nail solution as a topical application once daily for 24 weeks.

Arm title	Vehicle

Arm description:

Subjects with mild to moderate psoriatic fingernail/s, were treated with vehicle solution matched to P-3073 nail solution as a topical application once daily for 24 weeks.

Arm type	Placebo
Investigational medicinal product name	Placebo
Investigational medicinal product code	
Other name	Vehicle
Pharmaceutical forms	Medicated nail lacquer
Routes of administration	Topical use

Dosage and administration details:

Subjects with mild to moderate psoriatic fingernail/s, were treated with vehicle solution matched to P-3073 nail solution as a topical application once daily for 24 weeks.

Number of subjects in period 1	P-3073	Vehicle
Started	181	177
Treated	176	176
Completed	158	155
Not completed	23	22
Consent withdrawn by subject	10	13
Adverse event, non-fatal	1	1
Pregnancy	1	1
Follow Up Visit Wrongly Not Performed	1	-
Lost to follow-up	7	3
Lack of efficacy	1	1
Protocol deviation	2	3

Baseline characteristics

Reporting groups	
Reporting group title	P-3073
Reporting group description:	
Subjects with mild to moderate psoriatic fingernail/s, were treated with P-3073 nail solution as a topical application once daily for 24 weeks.	
Reporting group title	Vehicle
Reporting group description:	
Subjects with mild to moderate psoriatic fingernail/s, were treated with vehicle solution matched to P-3073 nail solution as a topical application once daily for 24 weeks.	

Reporting group values	P-3073	Vehicle	Total
Number of subjects	181	177	358
Age categorical Units: Subjects			
Age continuous Units: years arithmetic mean standard deviation	 48.54 ± 13.32	 47.94 ± 13.45	 -
Gender categorical Units: Subjects			
Female	72	63	135
Male	109	114	223
Nail Psoriasis Severity Index (NAPSI) Matrix Score			
Nail matrix psoriasis was assessed by the presence of any features including nail pitting, leukonychia, red spots in the lunula, and crumbling in each quadrant of the nail. The score was 0 if the findings were not present, 1 if they were present in 1 quadrant of the nail, 2 if present in 2 quadrants of a nail, 3 if present in 3 quadrants of a nail, and 4 if present in 4 quadrants of a nail. Thus each nail had a matrix score (0-4). The number of mild to moderate psoriasis fingernails analysed at the baseline visit are 1165 and 1166 for the reporting groups P-3073 and Vehicle respectively.			
Units: Score on a scale arithmetic mean standard deviation	 1.35 ± 1.04	 1.27 ± 1.06	 -
Nail Psoriasis Severity Index (NAPSI) Bed Score			
Nail bed psoriasis was assessed by the presence of any features including onycholysis, oil drop (salmon patch) dyschromia, splinter hemorrhages, and nail bed hyperkeratosis in each quadrant of nail. The score was 0 if the findings were not present, 1 if they were present in 1 quadrant of the nail, 2 if present in 2 quadrants of a nail, 3 if present in 3 quadrants of a nail, and 4 if present in 4 quadrants of a nail. Thus each nail had a nail bed score(0-4). The number of mild to moderate psoriasis fingernails analysed at the baseline are 1165 and 1166 for P-3073 and Vehicle group, respectively			
Units: Score on a scale arithmetic mean standard deviation	 1.55 ± 0.98	 1.54 ± 0.94	 -

End points

End points reporting groups

Reporting group title	P-3073
Reporting group description:	
Subjects with mild to moderate psoriatic fingernail/s, were treated with P-3073 nail solution as a topical application once daily for 24 weeks.	
Reporting group title	Vehicle
Reporting group description:	
Subjects with mild to moderate psoriatic fingernail/s, were treated with vehicle solution matched to P-3073 nail solution as a topical application once daily for 24 weeks.	
Subject analysis set title	Full Analysis Set (FAS)
Subject analysis set type	Full analysis
Subject analysis set description:	
FAS consisted of all randomized subjects to whom the investigational drug was dispensed.	
Subject analysis set title	Safety analysis set (SAF)
Subject analysis set type	Safety analysis
Subject analysis set description:	
SAF consisted of all randomized subjects with at least one documented application of any study drug.	

Primary: Change From Baseline in Total Nail Psoriasis Severity Index (NAPSI) Score at Week 24

End point title	Change From Baseline in Total Nail Psoriasis Severity Index (NAPSI) Score at Week 24
End point description:	
Nail psoriasis severity index (NAPSI) was used to asses nail psoriasis. The nail was assessed for nail matrix and bed psoriasis in each quadrant of the nail. The score was 0 if the findings were not present, 1 if they were present in 1 quadrant of the nail, 2 if present in 2 quadrants of a nail, 3 if present in 3 quadrants of a nail, and 4 if present in 4 quadrants of a nail. Thus each nail had a matrix score (0-4) and a nail bed score (0-4), and the total nail score was the sum of those two (0-8). The sum of the scores from all involved fingernails was 0-80 (total NAPSI score for that subject at that time). "n" signifies those subjects who were evaluable for this measure at given time point for each group.	
End point type	Primary
End point timeframe:	
Baseline, Week 24	

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[1]	177[2]		
Units: Score on the scale				
arithmetic mean (standard deviation)				
Change at week 24 (n=162, 157)	-3.16 (± 7.72)	-2.87 (± 8.39)		

Notes:

[1] - FAS

[2] - FAS

Statistical analyses

Statistical analysis title	P-3073 vs Vehicle difference

Statistical analysis description:

The linear effects model included the treatment group and the pooled site as dummy effects and the total NAPSI Score at screening as continuous covariate.

Comparison groups	P-3073 v Vehicle
Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.7333
Method	Linear effects model
Parameter estimate	Adjusted mean difference
Point estimate	-0.29
Confidence interval	
level	95 %
sides	2-sided
lower limit	-1.99
upper limit	1.4
Variability estimate	Standard error of the mean
Dispersion value	0.86

Secondary: Nail Physician’s Global Assessment (PGA) Response Rate at Week 24

End point title	Nail Physician’s Global Assessment (PGA) Response Rate at Week 24

End point description:

Nail PGA score was used to assess the fingernail. For each affected fingernail, the proximal nail matrix, the distal nail matrix, and the nail bed were scored as 0-clear, 1-almost clear, 2-mild, 3-moderate or 4-severe. A responder fingernail was defined as a score of “clear” (0) or “almost clear” (1) and at least 2 points improvement from the baseline score. Nail PGA response rate is the proportion of subjects mild to moderate fingernails with nail PGA response which is the ratio of “number of subjects mild to moderate responder fingernails” by ”number of affected mild to moderate fingernails at baseline” for each subject. "n" signifies subjects who were evaluable for this measure at given time point for each group.

End point type	Secondary

End point timeframe:

Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[3]	177[4]		
Units: Ratio of subject’s responder fingernails				
arithmetic mean (standard deviation)				
Response rate at Week 24 (n=162, 157)	0.26 (± 0.31)	0.29 (± 0.32)		

Notes:

[3] - FAS

[4] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1

Statistical analysis description:

A generalized linear mixed effects model for repeated measures (MMRM) was modelled to obtain an estimate of the probability to achieve a PGA response after 24 weeks of treatment. Only subjects with at least one mild to moderate fingernail were considered.

Comparison groups	P-3073 v Vehicle
Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.7913
Method	MMRM Model
Parameter estimate	Median difference (net)
Point estimate	-0.01
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.07
upper limit	0.06

Secondary: Change From Baseline in Nail Psoriasis Severity Index (NAPSI) Matrix Score at Week 24

End point title	Change From Baseline in Nail Psoriasis Severity Index (NAPSI) Matrix Score at Week 24

End point description:

NAPSI was used to asses nail psoriasis. The nail was assessed for nail matrix psoriasis in each quadrant of the nail. Nail matrix psoriasis was assessed by the presence of any features including nail pitting, leukonychia, red spots in the lunula, and crumbling in each quadrant of the nail. The score was 0 if the findings were not present, 1 if they were present in 1 quadrant of the nail, 2 if present in 2 quadrants of a nail, 3 if present in 3 quadrants of a nail, and 4 if present in 4 quadrants of a nail. Thus each nail had a matrix score (0-4). The number of mild to moderate psoriatic fingernails analysed at the baseline visit were 1165 and 1166 for the reporting groups P-3073 and Vehicle respectively. "n" signifies nails of subjects which were evaluable for this measure at given time point for each group.

End point type	Secondary

End point timeframe:

Baseline, Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[5]	177[6]		
Units: Score on the scale				
arithmetic mean (standard deviation)				
Change at Week 24 (n= 1038, 1040)	-0.24 (± 1.21)	-0.26 (± 1.22)		

Notes:

[5] - FAS

[6] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1
Comparison groups	P-3073 v Vehicle

Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.382
Method	MMRM Model
Parameter estimate	Adjusted mean difference
Point estimate	0.04
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.05
upper limit	0.14

Secondary: Change From Baseline in Nail Psoriasis Severity Index (NAPSI) Bed Score at Week 24

End point title	Change From Baseline in Nail Psoriasis Severity Index (NAPSI) Bed Score at Week 24

End point description:

NAPSI was used to asses nail psoriasis. The nail was assessed for nail bed psoriasis in each quadrant of the nail. Nail bed psoriasis was assessed by the presence of any features including onycholysis, oil drop (salmon patch) dyschromia, splinter hemorrhages, and nail bed hyperkeratosis in each quadrant of the nail. The score was 0 if the findings were not present, 1 if they were present in 1 quadrant of the nail, 2 if present in 2 quadrants of a nail, 3 if present in 3 quadrants of a nail, and 4 if present in 4 quadrants of a nail. Thus each nail had a nail bed score (0-4). The improvement in NAPSI Bed Score was defined as the achievement of a value of NAPSI Bed Score equal to zero (i.e. completely clearance of psoriatic signs). The number of mild to moderate psoriatic fingernails analysed at the baseline visit were 1165 and 1166 for the reporting groups P-3073 and Vehicle respectively. "n" signifies nails of subjects which were evaluable for this measure at given time for each group

End point type	Secondary

End point timeframe:

Baseline, Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[7]	177[8]		
Units: Score on the scale				
arithmetic mean (standard deviation)				
Change at Week 24 (n= 1038, 1040)	-0.24 (± 1.04)	-0.17 (± 1.10)		

Notes:

[7] - FAS

[8] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1
Comparison groups	P-3073 v Vehicle

Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.0668
Method	MMRM Model
Parameter estimate	Adjusted mean difference
Point estimate	-0.08
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.16
upper limit	0.01

Secondary: Change from Baseline in Subject's Quality-Of-Life as Assessed by Dermatology Life Quality Index (DLQI) at Week 24

End point title	Change from Baseline in Subject's Quality-Of-Life as Assessed by Dermatology Life Quality Index (DLQI) at Week 24

End point description:

Dermatology Life Quality Index (DLQI) score was used to evaluate the impact of nail psoriasis on subject's quality-of-life. The DLQI Total score range from 0 (no effect) to 30 (extremely affected). The higher the score, the more quality of life is impaired. "n" signifies those subjects who were evaluable for this measure at given time point for each group.

End point type	Secondary

End point timeframe:

Baseline, Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[9]	177[10]		
Units: Score on the scale				
arithmetic mean (standard deviation)				
Change at week 24, n=162, 157	-2.52 (± 5.54)	-2.64 (± 5.41)		

Notes:

[9] - FAS

[10] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1

Statistical analysis description:

The linear mixed effects model for repeated measures included treatment, pooled site, visit and treatment-by-visit interaction as fixed effects and baseline value as covariate. The unstructured variance-covariance matrix was used to take into account correlation among repeated measures within subject.

Comparison groups	P-3073 v Vehicle

Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.7885
Method	MMRM Model
Parameter estimate	Adjusted mean difference
Point estimate	0.14
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.88
upper limit	1.16

Secondary: Proportion of Subject's Mild to Moderate Fingernails With Improvement in Nail Psoriasis Severity Index (NAPSI) Score at Week 24

End point title	Proportion of Subject's Mild to Moderate Fingernails With Improvement in Nail Psoriasis Severity Index (NAPSI) Score at Week 24

End point description:

NAPSI was used to asses nail psoriasis. The nail was assessed for nail matrix and bed psoriasis in each quadrant of the nail. Each nail was given a score for nail bed psoriasis (0-4) and nail matrix psoriasis (0-4) depending on the presence of any features of nail psoriasis in that quadrant. The score was 0 if no findings, and 4 if findings in 4 quadrants of a nail. The total nail score was the sum of those two (0-8). The sum of the scores from all involved fingernails was 0-80 (total NAPSI score for that subject at that time). The improvement in NAPSI score was defined as the achievement of a value of NAPSI score equal to zero. Proportion of subjects’ mild to moderate fingernails with improvement in NAPSI Score is the ratio of “number of improved mild to moderate fingernails in NAPSI Score” by ”number of affected mild to moderate fingernails at baseline” for each subject. "n" signifies the subjects who were evaluable for this measure at given time point for each group.

End point type	Secondary

End point timeframe:

Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[11]	177[12]		
Units: Ratio of subject’s improved fingernails				
arithmetic mean (standard deviation)				
Week 24 (n=162, 158)	0.20 (± 0.28)	0.22 (± 0.30)		

Notes:

[11] - FAS

[12] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1

Statistical analysis description:

A generalized linear mixed effects model was modelled to obtain estimate of the probability to achieve an improvement after 24 weeks of treatment. Only subjects with at least one mild to moderate

fingernail were considered.

Comparison groups	P-3073 v Vehicle
Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.7434
Method	MMRM Model
Parameter estimate	Adjusted mean difference
Point estimate	-0.01
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.07
upper limit	0.05

Secondary: Proportion of Subject's Mild to Moderate Fingernails With Improvement in Nail Psoriasis Severity Index (NAPSI) Matrix Score at Week 24

End point title	Proportion of Subject's Mild to Moderate Fingernails With Improvement in Nail Psoriasis Severity Index (NAPSI) Matrix Score at Week 24

End point description:

NAPSI was used to asses each nail psoriasis. The nail was assessed for nail matrix psoriasis in each quadrant of the nail. Nail matrix psoriasis was assessed by the presence of any features including nail pitting, leukonychia, red spots in the lunula, and crumbling in each quadrant of the nail. Each nail had a nail matrix score of 0-4 where was 0 if no findings, and 4 if findings in 4 quadrants of a nail. The improvement in NAPSI matrix score was defined as the achievement of a value of NAPSI matrix score equal to zero (i.e. completely clearance of psoriatic signs). Proportion of subjects' mild to moderate fingernails with improvement in NAPSI matrix Score is the ratio of "number of improved mild to moderate fingernails in NAPSI matrix Score" by "number of affected mild to moderate fingernails at baseline" for each subject. "n" signifies the subjects who were evaluable for this measure at given time point for each group.

End point type	Secondary

End point timeframe:

Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[13]	177[14]		
Units: Ratio of subject's improved fingernails				
arithmetic mean (standard deviation)				
Week 24 (n=162, 158)	0.28 (± 0.32)	0.28 (± 0.32)		

Notes:

[13] - FAS

[14] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1

Statistical analysis description:

A generalized linear mixed effects model was modelled to obtain estimate of the probability to achieve an improvement after 24 weeks of treatment. Only subjects with at least one mild to moderate fingernail were considered.

Comparison groups	P-3073 v Vehicle
Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.9753
Method	Linear mixed effects model
Parameter estimate	adjusted mean difference
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.06
upper limit	0.07

Secondary: Proportion of Subject's Mild to Moderate Fingernails With Improvement in Nail Psoriasis Severity Index (NAPSI) Bed Score at Week 24

End point title	Proportion of Subject's Mild to Moderate Fingernails With Improvement in Nail Psoriasis Severity Index (NAPSI) Bed Score at Week 24

End point description:

NAPSI was used to asses each nail psoriasis. The nail was assessed for nail bed psoriasis in each quadrant of the nail. Nail bed psoriasis was assessed by the presence of any features including onycholysis, oil drop (salmon patch) dyschromia, splinter hemorrhages, and nail bed hyperkeratosis in each quadrant of the nail. Each nail had a nail bed score of 0-4 where was 0 if no findings, and 4 if findings in 4 quadrants of a nail. The improvement in NAPSI Bed Score was defined as the achievement of a value of NAPSI Bed Score equal to zero (i.e. completely clearance of psoriatic signs). Proportion of subjects’ mild to moderate fingernails with improvement in NAPSI Bed Score is the ratio of “number of improved mild to moderate fingernails in NAPSI Bed Score” by ”number of affected mild to moderate fingernails at baseline” for each subject. "n" signifies the subjects who were evaluable for this measure at given time point for each group.

End point type	Secondary

End point timeframe:

Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[15]	177[16]		
Units: Ratio of subject’s improved fingernails				
arithmetic mean (standard deviation)				
Week 24 (n=162, 158)	0.16 (± 0.24)	0.18 (± 0.26)		

Notes:

[15] - FAS

[16] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1

Statistical analysis description:

A generalized linear mixed effects model was modelled to obtain estimate of the probability to achieve an improvement after 24 weeks of treatment. Only subjects with at least one mild to moderate fingernail were considered.

Comparison groups	P-3073 v Vehicle
Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.9988
Method	Linear mixed effects model
Parameter estimate	Adjusted mean difference
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.05
upper limit	0.05

Secondary: Subject Acceptance of Study Therapy at Week 24

End point title	Subject Acceptance of Study Therapy at Week 24

End point description:

The subjects' acceptability of the study therapy was evaluated on the 4-point scale ranged from 1 = poor (very unpleasant and unsatisfactory), 2 = moderate (not fully satisfactory), 3 = good (satisfactory) to 4 = very good (fully satisfactory). "N" signifies those subjects who were evaluable for this measure for each group.

End point type	Secondary

End point timeframe:

Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	171[17]	167[18]		
Units: Subject				
Poor	26	20		
Moderate	34	41		
Good	64	63		
Very Good	39	39		
Not Evaluated	8	4		

Notes:

[17] - FAS subjects evaluable for this end point.

[18] - FAS subjects evaluable for this end point.

Statistical analyses

No statistical analyses for this end point

Secondary: Change in Discomfort Due to Fingernail Psoriasis as Measured on a Visual Analogue Scale (VAS) at Week 24

End point title	Change in Discomfort Due to Fingernail Psoriasis as Measured on a Visual Analogue Scale (VAS) at Week 24

End point description:

The VAS was used to measure the amount of discomfort felt by the subject in fingernail; it ranges from no discomfort (0) to worst possible discomfort (100). "n" signifies those subjects who were evaluable for this measure at given time point for each group.

End point type	Secondary

End point timeframe:

Baseline, Week 24

End point values	P-3073	Vehicle		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	181[19]	177[20]		
Units: Score on a scale				
arithmetic mean (standard deviation)				
Change at Week 24, n=156, 157	-12.63 (± 24.95)	-11.25 (± 27.45)		

Notes:

[19] - FAS

[20] - FAS

Statistical analyses

Statistical analysis title	Statistical Analysis 1

Statistical analysis description:

The linear mixed effects model for repeated measures included treatment, pooled site, visit and treatment-by-visit interaction as fixed effects and baseline value as covariate. The unstructured variance-covariance matrix was used to take into account correlation among repeated measures within subjects.

Comparison groups	P-3073 v Vehicle
Number of subjects included in analysis	358
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.78
Method	MMRM Model
Parameter estimate	Adjusted mean difference
Point estimate	-0.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-5.63
upper limit	4.23

Adverse events

Adverse events information	
Timeframe for reporting adverse events:	
From start of study drug application until follow up (Week 28)	
Assessment type	Systematic

Dictionary used	
Dictionary name	MedDRA
Dictionary version	19.0

Reporting groups	
Reporting group title	P-3073
Reporting group description:	
Subjects with mild to moderate psoriatic fingernail/s, were treated with P-3073 nail solution as a topical application once daily for 24 weeks.	
Reporting group title	Vehicle
Reporting group description:	
Subjects with mild to moderate psoriatic fingernail/s, were treated with vehicle solution matched to P-3073 nail solution as a topical application once daily for 24 weeks.	

Serious adverse events	P-3073	Vehicle	
Total subjects affected by serious adverse events			
subjects affected / exposed	10 / 176 (5.68%)	4 / 176 (2.27%)	
number of deaths (all causes)	0	0	
number of deaths resulting from adverse events	0	0	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Breast cancer			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Injury, poisoning and procedural complications			
Limb traumatic amputation			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nail injury			
alternative assessment type: Non-systematic			

subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Muscle rupture			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thermal burn			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Syncope			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pregnancy, puerperium and perinatal conditions			
Pregnancy			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	2 / 176 (1.14%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory, thoracic and mediastinal disorders			
Chronic obstructive pulmonary disease			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin and subcutaneous tissue disorders			

Psoriasis			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal and urinary disorders			
Renal colic			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal and connective tissue disorders			
Muscular weakness			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Psoriatic arthropathy			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

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

Non-serious adverse events	P-3073	Vehicle	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	105 / 176 (59.66%)	91 / 176 (51.70%)	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Anogenital warts			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Skin papilloma			

alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Vascular disorders			
Hypertension			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	2 / 176 (1.14%)	
occurrences (all)	2	2	
Arteriosclerosis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Hypotension			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Surgical and medical procedures			
Atherosclerosis prophylaxis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Tooth extraction			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	0 / 176 (0.00%)	
occurrences (all)	4	0	
Cataract operation			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Wisdom teeth removal			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Sebaceous cyst excision			
alternative assessment type: Non-systematic			

subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
General disorders and administration site conditions			
Peripheral swelling			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	5	
Fatigue			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Pyrexia			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	3 / 176 (1.70%)	
occurrences (all)	1	9	
Immune system disorders			
Allergy to arthropod bite			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Allergy to arthropod sting			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Sarcoidosis			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Hypersensitivity			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Reproductive system and breast disorders			

Dysmenorrhoea			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	2	
Menstrual disorder			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Oligomenorrhoea			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Respiratory, thoracic and mediastinal disorders			
Rhinorrhoea			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Oropharyngeal pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	2 / 176 (1.14%)	
occurrences (all)	4	2	
Cough			
alternative assessment type: Non-systematic			
subjects affected / exposed	6 / 176 (3.41%)	1 / 176 (0.57%)	
occurrences (all)	8	1	
Psychiatric disorders			
Anxiety			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Depression			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Insomnia			

alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Stress			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Sleep disorder			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Investigations			
Blood cholesterol increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Aspartate aminotransferase increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	3 / 176 (1.70%)	
occurrences (all)	2	3	
Blood bilirubin increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Alanine aminotransferase increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	2 / 176 (1.14%)	
occurrences (all)	2	2	
Blood glucose increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	1 / 176 (0.57%)	
occurrences (all)	4	1	
Blood glucose abnormal			
alternative assessment type: Non-systematic			

subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Blood pressure increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Blood triglycerides increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	2 / 176 (1.14%)	
occurrences (all)	3	3	
Blood potassium increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	1 / 176 (0.57%)	
occurrences (all)	2	1	
Gamma-glutamyltransferase increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	0 / 176 (0.00%)	
occurrences (all)	4	0	
Glucose urine present			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Transaminases increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Liver function test abnormal			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	2 / 176 (1.14%)	
occurrences (all)	2	2	
Liver function test increased			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	

Injury, poisoning and procedural complications			
Arthropod bite			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Contusion			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	4 / 176 (2.27%)	
occurrences (all)	3	6	
Injury corneal			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Fall			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Eye injury			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Limb injury			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	1 / 176 (0.57%)	
occurrences (all)	2	1	
Nail injury			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Laceration			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Road traffic accident			
alternative assessment type: Non-			

systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Skin injury			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Traumatic haematoma			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Wound			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Congenital, familial and genetic disorders			
Type V hyperlipidaemia			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Cardiac disorders			
Angina pectoris			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Nervous system disorders			
Dizziness			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Headache			
alternative assessment type: Non-systematic			
subjects affected / exposed	31 / 176 (17.61%)	27 / 176 (15.34%)	
occurrences (all)	89	56	
Cervical radiculopathy			

alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Hypoaesthesia			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Intercostal neuralgia			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Tremor			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Syncope			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Migraine			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Ear and labyrinth disorders			
Ear pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Middle ear inflammation			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Eye disorders			

Cataract			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Chalazion			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Conjunctival hyperaemia			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Visual acuity reduced			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Gastrointestinal disorders			
Abdominal distension			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Abdominal pain upper			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	3 / 176 (1.70%)	
occurrences (all)	13	3	
Abdominal pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Aphthous ulcer			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Diarrhoea			
alternative assessment type: Non-systematic			

subjects affected / exposed	1 / 176 (0.57%)	4 / 176 (2.27%)	
occurrences (all)	1	4	
Dyspepsia			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	1 / 176 (0.57%)	
occurrences (all)	3	1	
Gastrointestinal disorder			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Gastrooesophageal reflux disease			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Nausea			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Toothache			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	3 / 176 (1.70%)	
occurrences (all)	4	6	
Skin and subcutaneous tissue disorders			
Dermatitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Dermatitis allergic			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Eczema			
alternative assessment type: Non-systematic			

subjects affected / exposed	1 / 176 (0.57%)	2 / 176 (1.14%)	
occurrences (all)	1	2	
Hyperkeratosis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Erythema			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Onychalgia			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Nail disorder			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	2 / 176 (1.14%)	
occurrences (all)	2	3	
Pruritus			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Onychoclasis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	3	0	
Psoriasis			
alternative assessment type: Non-systematic			
subjects affected / exposed	8 / 176 (4.55%)	6 / 176 (3.41%)	
occurrences (all)	10	8	
Skin disorder			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	2 / 176 (1.14%)	
occurrences (all)	0	2	

Rebound psoriasis			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Skin ulcer			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Skin irritation			
alternative assessment type: Non-systematic			
subjects affected / exposed	18 / 176 (10.23%)	13 / 176 (7.39%)	
occurrences (all)	51	32	
Solar dermatitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Swelling face			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Renal and urinary disorders			
Cystitis noninfective			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Renal colic			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	1 / 176 (0.57%)	
occurrences (all)	3	1	
Haematuria			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Renal cyst			
alternative assessment type: Non-systematic			

subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Proteinuria			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Urinary tract inflammation			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Endocrine disorders			
Hypothyroidism			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Musculoskeletal and connective tissue disorders			
Arthralgia			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	4 / 176 (2.27%)	
occurrences (all)	6	4	
Arthritis			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Back pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	7 / 176 (3.98%)	9 / 176 (5.11%)	
occurrences (all)	9	19	
Muscle spasms			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Joint swelling			
alternative assessment type: Non-systematic			

subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Musculoskeletal pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	2 / 176 (1.14%)	
occurrences (all)	1	2	
Intervertebral disc protrusion			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Pain in extremity			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Musculoskeletal chest pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Myalgia			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	2 / 176 (1.14%)	
occurrences (all)	2	2	
Psoriatic arthropathy			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Neck pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	3	
Spinal pain			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	1 / 176 (0.57%)	
occurrences (all)	4	1	

Infections and infestations			
Conjunctivitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Bronchitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	2 / 176 (1.14%)	
occurrences (all)	4	2	
Bronchitis bacterial			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Cystitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Ear infection			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Folliculitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	3	1	
Gastroenteritis			
alternative assessment type: Non-systematic			
subjects affected / exposed	3 / 176 (1.70%)	0 / 176 (0.00%)	
occurrences (all)	4	0	
Gingivitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Gastroenteritis viral			
alternative assessment type: Non-systematic			

subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Hordeolum			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Gastrointestinal infection			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	2	1	
Infection			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Herpes zoster			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Influenza			
alternative assessment type: Non-systematic			
subjects affected / exposed	6 / 176 (3.41%)	5 / 176 (2.84%)	
occurrences (all)	6	5	
Paronychia			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Nasopharyngitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	19 / 176 (10.80%)	11 / 176 (6.25%)	
occurrences (all)	22	14	
Pharyngitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	0 / 176 (0.00%)	
occurrences (all)	4	0	

Pulpitis dental			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Sinusitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Rhinitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	4 / 176 (2.27%)	
occurrences (all)	4	4	
Skin infection			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	2	
Tinea versicolour			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Urinary tract infection			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	2 / 176 (1.14%)	
occurrences (all)	2	3	
Upper respiratory tract infection			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Tonsillitis			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Vaginal infection			
alternative assessment type: Non-systematic			

subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Viral infection			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	1 / 176 (0.57%)	
occurrences (all)	2	1	
Metabolism and nutrition disorders			
Diabetes mellitus inadequate control			
alternative assessment type: Non-systematic			
subjects affected / exposed	1 / 176 (0.57%)	1 / 176 (0.57%)	
occurrences (all)	1	1	
Hypercholesterolaemia			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	3 / 176 (1.70%)	
occurrences (all)	4	3	
Diabetes mellitus			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Gout			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	1	
Hyperglycaemia			
alternative assessment type: Non-systematic			
subjects affected / exposed	0 / 176 (0.00%)	1 / 176 (0.57%)	
occurrences (all)	0	2	
Hyperuricaemia			
alternative assessment type: Non-systematic			
subjects affected / exposed	2 / 176 (1.14%)	0 / 176 (0.00%)	
occurrences (all)	2	0	
Hyperlipidaemia			
alternative assessment type: Non-systematic			

subjects affected / exposed	1 / 176 (0.57%)	0 / 176 (0.00%)	
occurrences (all)	1	0	
Hypertriglyceridaemia			
alternative assessment type: Non-systematic			
subjects affected / exposed	4 / 176 (2.27%)	4 / 176 (2.27%)	
occurrences (all)	4	4	

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
14 March 2016	- 'Subjects acceptance of study therapy at week 24' was added as secondary endpoint. - Definition of the secondary endpoint related to Nail PGA response was updated. - Rewording of the exclusion criteria related to the use of concomitant topical and systemic psoriatic treatments.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported