Clinical trial results:
Double-blind, randomised clinical study comparing efficacy and safety of Clindamycin/Benzoyl Peroxide Gel (10 mg/g + 30 mg/g) (Test) vs. DUAC 10 mg/g + 30 mg/g Gel (Reference) vs. Vehicle in patients with papulopustular acne

Summary

EudraCT number	2017-000521-13
Trial protocol	CZ
Global end of trial date	01 March 2021

Results information

Result version number	v1 (current)
This version publication date	09 April 2022
First version publication date	09 April 2022

Trial information

Trial identification

Sponsor protocol code	17-01/ClinBPO-30

Additional study identifiers

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

Notes:

Sponsors

Sponsor organisation name	Dermapharm AG
Sponsor organisation address	Lil-Dagover-Ring 7, Gruenwald, Germany, 82031
Public contact	Clinical Research Department, Dermapharm AG, +49 89641860, Clinicaltrials.Dermapharm@dermapharm.com
Scientific contact	Clinical Research Department, Dermapharm AG, +49 89641860, Clinicaltrials.Dermapharm@dermapharm.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	21 October 2021
Is this the analysis of the primary completion data?	Yes
Primary completion date	01 March 2021
Global end of trial reached?	Yes
Global end of trial date	01 March 2021
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

Evaluation of the efficacy and safety of a new gel containing 10 mg/g Clindamycin and 30 mg/g Benzoyl peroxide vs. DUAC 10 mg/g + 30 mg/g Gel (Reference) vs. vehicle in patients with papulopustular acne

Protection of trial subjects:

In the current clinical trial patients below the age of 18 have been included. In such a case, an age-appropriate written subject information sheet was handed over to adolescent patients and an appropriate information session was to be performed by the investigator. The legal guardian(s) received a comparable document and an information session for adults. Before the start of screening and randomisation for the current clinical trial, the legal guardian(s) had to sign the informed consent form(s) and the adolescent patient the informed assent form. In case any of the parties (legal guardian(s) or adolescent patient) refused their consent, a participation of the adolescent patient in the trial was not possible.

Background therapy:

There was no background therapy.

Evidence for comparator:

The comparator contains the same ingredients in the same concentration as the test product and has a marketing license for the study indication.

Actual start date of recruitment	03 April 2019
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	Czechia: 676
Worldwide total number of subjects	676
EEA total number of subjects	676

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)	378
Adults (18-64 years)	298
From 65 to 84 years	0
85 years and over	0

Subject disposition

Recruitment

Recruitment details:

16 study centers in Czechia; first patient first visit: 25 April 2019; last patient last visit: 01 March 2021

Pre-assignment

Screening details:

Main criteria for inclusion: Women, men and adolescents of ≥ 12 years of age; Diagnosis of "papulopustular acne" according to generally accepted criteria; On the face, ≥ 25 non-inflammatory lesions and ≥ 20 inflammatory lesions, thereof ≤ 2 nodular lesions; Investigator`s Global Assessment (IGA) of acne severity grade 2, 3 or 4

Period 1

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

Blinding implementation details:

The tubes containing the study medications were neutral white. The attached labels were identical for all three preparations. All three study medications were indistinguishable with respect to visual or odorous characteristics. The random code was transferred to the data base not before the following actions were completed: data base closure, finalisation of the statistical analysis plan, a Blind Data Review and a subsequent Blind Data Report.

Arms

Are arms mutually exclusive?	Yes
Arm title	ClinBPO 30

Arm description:

Test product

Arm type	Experimental
Investigational medicinal product name	Clindamycin/Benzoyl Peroxide Gel (10 mg/g + 30 mg/g)
Investigational medicinal product code	D10AF51
Other name	
Pharmaceutical forms	Gel
Routes of administration	Cutaneous use

Dosage and administration details:

A thin layer of gel should be applied to the affected area once a day. The treatment area was defined as the acne affected areas on the face whereas face was considered as the area bounded by ears, hairline and lower margin of the mandibles. Contact with the mouth, eyes, lips, other mucous membranes or areas of irritated or broken skin should be avoided.

Arm title	Duac (10 mg/g + 30 mg/g)

Arm description:

Reference Product

Arm type	Active comparator
Investigational medicinal product name	Duac (10 mg/g + 30 mg/g)
Investigational medicinal product code	D10AF51
Other name	
Pharmaceutical forms	Gel
Routes of administration	Cutaneous use

Dosage and administration details:

A thin layer of gel should be applied to the affected area once a day. The treatment area was defined as the acne affected areas on the face whereas face was considered as the area bounded by ears, hairline and lower margin of the mandibles. Contact with the mouth, eyes, lips, other mucous membranes or

areas of irritated or broken skin should be avoided.

Arm title	Vehicle
Arm description: -	
Arm type	Placebo
Investigational medicinal product name	Placebo
Investigational medicinal product code	D10AF51
Other name	
Pharmaceutical forms	Gel
Routes of administration	Cutaneous use

Dosage and administration details:

A thin layer of gel should be applied to the affected area once a day. The treatment area was defined as the acne affected areas on the face whereas face was considered as the area bounded by ears, hairline and lower margin of the mandibles. Contact with the mouth, eyes, lips, other mucous membranes or areas of irritated or broken skin should be avoided.

Number of subjects in period 1	ClinBPO 30	Duac (10 mg/g + 30 mg/g)	Vehicle
Started	223	225	228
Completed	211	213	199
Not completed	12	12	29
Consent withdrawn by subject	1	3	1
Poor tolerability (patient)	3	1	1
Adverse event, non-fatal	2	2	2
Technical-logistic reasons	2	-	-
COVID-19 related	1	-	2
Lost to follow-up	1	-	1
Lack of efficacy	2	6	22

Baseline characteristics

Reporting groups

Reporting group title	Treatment Period

Reporting group description: -

Reporting group values	Treatment Period	Total	
Number of subjects	676	676	
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)	378	378	
Adults (18-64 years)	298	298	
From 65-84 years	0	0	
85 years and over	0	0	
Gender categorical Units: Subjects			
Female	437	437	
Male	239	239	

Subject analysis sets

Subject analysis set title	Safety data set
Subject analysis set type	Safety analysis

Subject analysis set description:

Comprises all randomised patients who had administered the study medication at least once and who provided at least one safety related outcome.

Subject analysis set title	FAS
Subject analysis set type	Full analysis

Subject analysis set description:

Consists of all patients as randomised who received study medication at least once and have a baseline assessment and at least one post-baseline assessment of the number of papulopustular acne lesions.

Subject analysis set title	PP
Subject analysis set type	Per protocol

Subject analysis set description:

Comprises all patients of the FAS who do not exhibit any major protocol violations.

Reporting group values	Safety data set	FAS	PP
Number of subjects	676	673	628
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0

Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	378	377	364
Adults (18-64 years)	298	296	264
From 65-84 years	0	0	0
85 years and over	0	0	0
Gender categorical Units: Subjects			
Female	437	435	394
Male	239	238	234

End points

End points reporting groups

Reporting group title	ClinBPO 30
Reporting group description: Test product	
Reporting group title	Duac (10 mg/g + 30 mg/g)
Reporting group description: Reference Product	
Reporting group title	Vehicle
Reporting group description: -	
Subject analysis set title	Safety data set
Subject analysis set type	Safety analysis
Subject analysis set description: Comprises all randomised patients who had administered the study medication at least once and who provided at least one safety related outcome.	
Subject analysis set title	FAS
Subject analysis set type	Full analysis
Subject analysis set description: Consists of all patients as randomised who received study medication at least once and have a baseline assessment and at least one post-baseline assessment of the number of papulopustular acne lesions.	
Subject analysis set title	PP
Subject analysis set type	Per protocol
Subject analysis set description: Comprises all patients of the FAS who do not exhibit any major protocol violations.	

Primary: Treatment effect (inflammatory lesions)

End point title	Treatment effect (inflammatory lesions)
End point description:	
End point type	Primary
End point timeframe: Treatment start (Visit V1) to end-of-treatment (EOT) examination at Visit V8 (week 12).	

End point values	ClinBPO 30	Duac (10 mg/g + 30 mg/g)	Vehicle	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	207	213	227	
Units: percent change				
arithmetic mean (standard deviation)	82.7 (± 24.21)	83.7 (± 20.86)	51.5 (± 41.57)	

Statistical analyses

Statistical analysis title	Analysis of efficacy

Statistical analysis description:

The first part of the primary objective of this study was to show therapeutic equivalence of the test

preparation ClinBPO 30 as compared to the reference DUAC. Therapeutic equivalence was statistically proven if the two-sided 95% confidence interval (CI) for μ INFL-ClinBPO - μ INFL-DUAC was completely contained within [-10.0, 10.0].

Comparison groups	ClinBPO 30 v Duac (10 mg/g + 30 mg/g)
Number of subjects included in analysis	420
Analysis specification	Pre-specified
Analysis type	equivalence
Parameter estimate	Mean difference (final values)
Point estimate	-1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-5.38
upper limit	3.3

Statistical analysis title	Superiority of Test over Vehicle

Statistical analysis description:

In order to verify assay sensitivity of the study design, superiority of the two active preparations over vehicle was tested by means of two-sided significance tests with α =0.05. The primary test of superiority was carried out for the ITT data set.

Comparison groups	ClinBPO 30 v Vehicle
Number of subjects included in analysis	434
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.0001
Method	t-test, 2-sided

Statistical analysis title	Superiority of Reference over Vehicle

Statistical analysis description:

In order to verify assay sensitivity of the study design, superiority of the two active preparations over vehicle was tested by means of two-sided significance tests with α =0.05. The primary test of superiority was carried out for the ITT data set.

Comparison groups	Vehicle v Duac (10 mg/g + 30 mg/g)
Number of subjects included in analysis	440
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.0001
Method	t-test, 2-sided

Primary: Treatment effect (total number of lesions)

End point title	Treatment effect (total number of lesions)

End point description:

End point type	Primary

End point timeframe:

Treatment start (Visit V1) to end-of-treatment (EOT) examination at Visit V8 (week 12).

End point values	ClinBPO 30	Duac (10 mg/g + 30 mg/g)	Vehicle	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	207	213	227	
Units: percentage change				
arithmetic mean (standard deviation)	75.1 (± 23.27)	75.9 (± 22.55)	43.7 (± 36.84)	

Statistical analyses

Statistical analysis title	Analysis of efficacy

Statistical analysis description:

The first part of the primary objective of this study was to show therapeutic equivalence of the test preparation ClinBPO 30 as compared to the reference DUAC. Therapeutic equivalence was statistically proven if the two-sided 95% confidence interval (CI) for μ TOTAL-ClinBPO - μ TOTAL-DUAC was completely contained within [-10.0, 10.0].

Comparison groups	ClinBPO 30 v Duac (10 mg/g + 30 mg/g)
Number of subjects included in analysis	420
Analysis specification	Pre-specified
Analysis type	equivalence
Parameter estimate	Mean difference (final values)
Point estimate	-0.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-5.23
upper limit	3.56

Statistical analysis title	Superiority of Test over Vehicle

Statistical analysis description:

In order to verify assay sensitivity of the study design, superiority of the two active preparations over vehicle was tested by means of two-sided significance tests with α =0.05. The primary test of superiority was carried out for the ITT data set.

Comparison groups	ClinBPO 30 v Vehicle
Number of subjects included in analysis	434
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.0001
Method	t-test, 2-sided

Statistical analysis title	Superiority of Reference over Vehicle

Statistical analysis description:

In order to verify assay sensitivity of the study design, superiority of the two active preparations over vehicle was tested by means of two-sided significance tests with α =0.05. The primary test of superiority was carried out for the ITT data set.

Comparison groups	Vehicle v Duac (10 mg/g + 30 mg/g)
Number of subjects included in analysis	440
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.0001
Method	t-test, 2-sided

Adverse events

Adverse events information	
Timeframe for reporting adverse events: From inclusion visit (day 0, Visit V1) to end-of-treatment (EOT) examination at Visit V8 (week 12).	
Assessment type	Non-systematic

Dictionary used	
Dictionary name	MedDRA
Dictionary version	24.0

Reporting groups	
Reporting group title	ClinBPO 30
Reporting group description: Test product	
Reporting group title	Duac (10 mg/g + 30 mg/g)
Reporting group description: Reference Product	
Reporting group title	Vehicle
Reporting group description: -	

Serious adverse events	ClinBPO 30	Duac (10 mg/g + 30 mg/g)	Vehicle
Total subjects affected by serious adverse events			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
number of deaths (all causes)	0	0	0
number of deaths resulting from adverse events	0	0	0
Infections and infestations			
Complicated appendicitis			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Frequency threshold for reporting non-serious adverse events: 0.05 %

Non-serious adverse events	ClinBPO 30	Duac (10 mg/g + 30 mg/g)	Vehicle
Total subjects affected by non-serious adverse events			
subjects affected / exposed	55 / 223 (24.66%)	62 / 224 (27.68%)	46 / 228 (20.18%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Gastroenteropancreatic neuroendocrine tumour disease			

subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Melanocytic naevus			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Skin papilloma			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Pregnancy, puerperium and perinatal conditions			
Pregnancy			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
General disorders and administration site conditions			
Application site dermatitis			
subjects affected / exposed	2 / 223 (0.90%)	1 / 224 (0.45%)	1 / 228 (0.44%)
occurrences (all)	2	1	1
Application site dryness			
subjects affected / exposed	12 / 223 (5.38%)	15 / 224 (6.70%)	5 / 228 (2.19%)
occurrences (all)	12	15	5
Application site erythema			
subjects affected / exposed	9 / 223 (4.04%)	9 / 224 (4.02%)	4 / 228 (1.75%)
occurrences (all)	9	9	4
Application site exfoliation			
subjects affected / exposed	3 / 223 (1.35%)	2 / 224 (0.89%)	0 / 228 (0.00%)
occurrences (all)	3	2	0
Application site hypersensitivity			
subjects affected / exposed	2 / 223 (0.90%)	3 / 224 (1.34%)	1 / 228 (0.44%)
occurrences (all)	2	3	1
Application site irritation			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Application site pain			
subjects affected / exposed	6 / 223 (2.69%)	7 / 224 (3.13%)	6 / 228 (2.63%)
occurrences (all)	6	7	6
Application site plaque			

subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Application site pruritus			
subjects affected / exposed	2 / 223 (0.90%)	3 / 224 (1.34%)	1 / 228 (0.44%)
occurrences (all)	3	3	1
Application site scab			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Pyrexia			
subjects affected / exposed	1 / 223 (0.45%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	1	1	0
Immune system disorders			
Milk allergy			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Reproductive system and breast disorders			
Dysmenorrhoea			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	2 / 228 (0.88%)
occurrences (all)	1	0	2
Polycystic ovaries			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Respiratory, thoracic and mediastinal disorders			
Cough			
subjects affected / exposed	5 / 223 (2.24%)	2 / 224 (0.89%)	1 / 228 (0.44%)
occurrences (all)	5	2	1
Rhinorrhoea			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Psychiatric disorders			
Sleep disorder			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Investigations			
Arthroscopy			

subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Injury, poisoning and procedural complications			
Joint injury			
subjects affected / exposed	1 / 223 (0.45%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	1	1	0
Nervous system disorders			
Headache			
subjects affected / exposed	3 / 223 (1.35%)	3 / 224 (1.34%)	2 / 228 (0.88%)
occurrences (all)	4	4	2
Migraine			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Ear and labyrinth disorders			
Motion sickness			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	1	0	1
Diarrhoea			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Dyspepsia			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Gastrooesophageal reflux disease			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Lip dry			

subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Tooth malformation			
subjects affected / exposed	1 / 223 (0.45%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	1	1	0
Toothache			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Skin and subcutaneous tissue disorders			
Acne			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Dermatitis allergic			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Dermatitis contact			
subjects affected / exposed	1 / 223 (0.45%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	1	1	0
Dry skin			
subjects affected / exposed	1 / 223 (0.45%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	1	1	0
Erythema			
subjects affected / exposed	3 / 223 (1.35%)	1 / 224 (0.45%)	1 / 228 (0.44%)
occurrences (all)	3	1	1
Hand dermatitis			
subjects affected / exposed	2 / 223 (0.90%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	2	0	0
Perioral dermatitis			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Photosensitivity reaction			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Pruritus			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0

Skin exfoliation			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Solar dermatitis			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Renal and urinary disorders			
Urinary tract discomfort			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Myalgia			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Infections and infestations			
Body tinea			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Bronchitis			
subjects affected / exposed	0 / 223 (0.00%)	3 / 224 (1.34%)	3 / 228 (1.32%)
occurrences (all)	0	3	3
COVID-19			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	2 / 228 (0.88%)
occurrences (all)	0	1	2
Cystitis			
subjects affected / exposed	1 / 223 (0.45%)	0 / 224 (0.00%)	0 / 228 (0.00%)
occurrences (all)	1	0	0
Furuncle			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Gastrointestinal viral infection			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1

Genital herpes			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	2	0
Hordeolum			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Impetigo			
subjects affected / exposed	0 / 223 (0.00%)	1 / 224 (0.45%)	0 / 228 (0.00%)
occurrences (all)	0	1	0
Nasopharyngitis			
subjects affected / exposed	4 / 223 (1.79%)	3 / 224 (1.34%)	1 / 228 (0.44%)
occurrences (all)	4	3	1
Oral herpes			
subjects affected / exposed	2 / 223 (0.90%)	1 / 224 (0.45%)	1 / 228 (0.44%)
occurrences (all)	3	1	1
Pharyngitis			
subjects affected / exposed	1 / 223 (0.45%)	1 / 224 (0.45%)	1 / 228 (0.44%)
occurrences (all)	1	1	1
Pulpitis dental			
subjects affected / exposed	0 / 223 (0.00%)	0 / 224 (0.00%)	1 / 228 (0.44%)
occurrences (all)	0	0	1
Rhinitis			
subjects affected / exposed	2 / 223 (0.90%)	1 / 224 (0.45%)	2 / 228 (0.88%)
occurrences (all)	2	1	2
Upper respiratory tract infection			
subjects affected / exposed	3 / 223 (1.35%)	5 / 224 (2.23%)	6 / 228 (2.63%)
occurrences (all)	5	5	7
Urinary tract infection			
subjects affected / exposed	0 / 223 (0.00%)	2 / 224 (0.89%)	0 / 228 (0.00%)
occurrences (all)	0	2	0
Viral infection			
subjects affected / exposed	2 / 223 (0.90%)	1 / 224 (0.45%)	1 / 228 (0.44%)
occurrences (all)	3	1	1

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? No

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

Limitations of the trial such as small numbers of subjects analysed or technical problems leading to unreliable data.

None reported

Notes: