European Directorate for the Quality of Medicines & HealthCare
Direction européenne de la qualité du médicament & soins de santé
COUNCIL OF EUROPE
CONSEIL DE L'EUROPE

Certification of Substances Department

KREATIVE ORGANICS PRIVATE LIMITED
Dr Krishnamohan SEELAMSETTY
D-123, Phase-III, I.D.A. Jeedimetla Village
Quthbullapur Mandal, Medchal-Malkajgiri District
India-500 055 Hyderabad, Telangana

CEP_RZ_PH_2014-020-1318702

Strasbourg, 23 September 2020

MFE / mr

Re: R1-CEP 2014-020-Rev 00/ Sodium picosulfate

Dear Dr SEELAMSETTY,

Please find enclosed the renewed certificate granted for **Sodium picosulfate** following the evaluation of the dossier.

If you find a mistake on the CEP, you should notify EDQM within 3 months. After this deadline, any complaint will no longer be considered valid.

Please note that a copy of this certificate has to be sent to your customers and/or any relevant competent authority to allow them to update their documentation and in particular the relevant marketing authorisations for medicinal products.

Yours faithfully,

Hélène BRUGUERA
Head of Department

Address: 7 Allée Kastner, CS 30026
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edqm
European Directorate for the Quality of Medicines & HealthCare | Direction européenne de la qualité du médicament & soins de santé
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Certification of Substances Department

Certificate of suitability
No. R1-CEP 2014-020-Rev 00

Name of the substance:
SODIUM PICOSULFATE

Name of holder:
KREATIVE ORGANICS PRIVATE LIMITED
D-123, Phase-III, I.D.A. Jeedimetla Village
Quthbullapur Mandal, Medchal-Malkajgiri District
India-500 055 Hyderabad, Telangana

Site(s) of production:
SEE ANNEX 1

THIS CERTIFICATE SUPERSEDES THE PREVIOUS CERTIFICATE
R0-CEP 2014-020-REV 00

After examination of the information provided on the manufacturing method and subsequent processes (including purification) for this substance on the site(s) of production listed in annex, we certify that the quality of the substance is suitably controlled by the current version of the monograph **SODIUM PICOSULFATE** no. 1031 of the European Pharmacopoeia, current edition including supplements, only if it is supplemented by the test(s) mentioned below, based on the analytical procedure(s) given in annex.

- Test for residual solvents by gas chromatography (Annex 2)
  Acetone not more than 5000 ppm
  Ethanol not more than 5000 ppm

A risk management summary for elemental impurities has been provided. (Annex 3)

The re-test period of the substance is 4 years if stored in a polyethylene bag, in an aluminium bag placed in a polyethylene drum.

The holder of the certificate has declared the absence of use of material of human or animal origin in the manufacture of the substance.

The submitted dossier must be updated after any significant change that may alter the quality, safety or efficacy of the substance.

Manufacture of the substance shall take place in accordance with the Good Manufacturing Practice and in accordance with the dossier submitted.

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Failure to comply with these provisions will render this certificate void.

This certificate is renewed from **16 October 2020** according to the provisions of Resolution AP-CSP (07) 1, and of Directive 2001/83/EC and Directive 2001/82/EC and any subsequent amendment, and the related guidelines.

This certificate has three annexes of 1 page each.

This certificate has:

lines.

On behalf of the
Director of EDQM

Strasbourg, 23 September 2020

DECLARATION OF ACCESS *(to be filled in by the certificate holder under their own responsibility)*

KREATIVE ORGANICS PRIVATE LIMITED, as holder of the certificate of suitability

R1-CEP 2014-020-Rev 00 for Sodium picosulfate

hereby authorises .....
(*name of the pharmaceutical company*)

to use the above-mentioned certificate of suitability in support of their application(s) for the following Marketing Authorisation(s): (*name of product(s) and marketing number(s), if known*)

The holder also certifies that no significant changes to the operations as described in the CEP dossier have been made since the granting of this version of the certificate.

Date and Signature (*of the CEP holder*):

Address: 7 Allée Kastner, CS 30026
F-67081 Strasbourg (France)
Tel: +33 (0) 3 88 41 30 30 – Fax: +33 (0) 3 88 41 27 71 - e-mail: cep@edqm.eu
Internet: http://www.edqm.eu

Annex 1: Site(s) of production for R1-CEP 2014-020-Rev 00

Production of Sodium picosulfate :
KREATIVE ORGANICS PRIVATE LIMITED
D-123, Phase-III, I.D.A. Jeedimetla Village
Quthbullapur Mandal, Medchal-Malkajgiri District
India-500 055 Hyderabad, Telangana

Residual Solvents (GC

Chromatographic conditions:

Column type	:	30 M x 0.53mm ID, 3µm fused silica analytical column
Carrier gas	:	Nitrogen
Flow rate	:	4.0 mL
Detector	:	Flame ionization detector (FID)
Detection range	:	4
Injector port temperature	:	150^0C
Detector port temperature	:	260^0C
Column oven temperature	:	

45^0C (hold for 4.0 minutes) then raise to 120^0C @ 25^0C / min (hold for 2.0 minutes) then raise to 230^0C @ 35^0C/min (hold for 2.0 minutes).

Equilibration time : 1.0 minute

Diluent: Dimethylformamide (DMF)

Blank: Dimethylformamide

Standard Solution: 10ppm solution of acetone and ethanol:

Take 12.7µL of acetone and 12.7µL of ethanol into a 10mL volumetric flask and dilute with DMF upto the mark. Take 1mL of this solution into a 100mL volumetric flask and dilute with DMF upto the mark.

Test Solution: Dissolve 0.1g of test sample to 10mL with DMF.

Procedure:

1. Condition the column for at least 30 minutes and ensure no peaks are eluting from the column. Allow to equilibrate the column.
2. Take 5mL of standard solution and transfer into vial and calculate % RSD for the peak area responses for acetone and ethanol peaks (six injections). The RSD for the peak areas less than 15% and for RT 2%.
3. Take 5mL of test solution and transfer into vial and inject in duplicate.

Calculation:

Determine the amount in ppm using following formula:

$$ppm = \frac{\text{Sample Area}}{\text{Standard Area}} \times \frac{\text{Standard concentration µg/mL}}{\text{Sample concentration µg/mL}} \times 10^6$$

Annexure-1: Elemental Impurities Risk Assessment Summary:

Intended route of administration/Use of the Substance: Sodium Picosulfate Oral dosage form				
Element	**Class**	**Intentionally added?**	**Considered in risk management?**	**Conclusion**
Cd	1	No	Yes	Absent
Pb	1	No	Yes	Absent
As	1	No	Yes	Absent
Hg	1	No	Yes	Absent
Co	2A	No	Yes	Absent
V	2A	No	Yes	Absent
Ni	2A	Yes	Yes	Absent
Tl	2B	No	No	No risk identified
Au	2B	No	No	No risk identified
Pd	2B	No	No	No risk identified
Ir	2B	No	No	No risk identified
Os	2B	No	No	No risk identified
Rh	2B	No	No	No risk identified
Ru	2B	No	No	No risk identified
Se	2B	No	No	No risk identified
Ag	2B	No	No	No risk identified
Pt	2B	No	No	No risk identified
Li	3	No	No	No risk identified
Sb	3	No	No	No risk identified
Ba	3	No	No	No risk identified
Mo	3	No	No	No risk identified
Cu	3	No	No	No risk identified
Sn	3	No	No	No risk identified
Cr	3	No	No	No risk identified

Note: "Absent" (meaning less than 30% of ICH Q3D option 1 limit, as defined under 3.1.1 of "Implementation of ICH Q3D in the Certification Procedure" PA/PH/CEP (16) 23, 1R guidance.