



NDA 022562/S-012

SUPPLEMENT APPROVAL

Recordati Rare Diseases
Attention: Sandy S. Suh, Pharm.D.
VP, Regulatory Affairs & Chief Compliance Officer
100 Corporate Drive
Lebanon, NJ 08833

Dear Dr. Suh:

Please refer to your supplemental new drug application (sNDA) dated and received March 18, 2020, and your amendments, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act (FDCA) for Carbaglu (carglumic acid) tablets.

This Prior Approval supplemental new drug application provides for additional clinical pharmacology study data from study CARIV-A-001, entitled "Phase I safety and PK study in healthy subjects of a new IV formulation of Carbaglu (carglumic acid) administered at single ascending doses and at different infusion durations and determination of the absolute bioavailability of the already marketed reference oral formulation," and from study Rec 0/0491-IT-PK 0492, entitled "Hepatic in vitro clearance of carglumic acid (REC 0/0491) in 3D human hepatocyte model and in plated human hepatocytes." The application also proposes minor revisions to Sections 2 and 14 of the Prescribing Information.

APPROVAL & LABELING

We have completed our review of this application, as amended. It is approved, effective on the date of this letter, for use as recommended in the enclosed agreed-upon labeling.

CONTENT OF LABELING

As soon as possible, but no later than 14 days from the date of this letter, submit the content of labeling [21 CFR 314.50(l)] in structured product labeling (SPL) format using the FDA automated drug registration and listing system (eLIST), as described at FDA.gov.¹ Content of labeling must be identical to the enclosed labeling (text for the Prescribing Information), with the addition of any labeling changes in pending "Changes Being Effected" (CBE) supplements, as well as annual reportable changes not included in the enclosed labeling.

¹ <http://www.fda.gov/ForIndustry/DataStandards/StructuredProductLabeling/default.htm>

Information on submitting SPL files using eList may be found in the guidance for industry *SPL Standard for Content of Labeling Technical Qs and As*.²

The SPL will be accessible from publicly available labeling repositories.

Also within 14 days, amend all pending supplemental applications that include labeling changes for this NDA, including CBE supplements for which FDA has not yet issued an action letter, with the content of labeling [21 CFR 314.50(l)(1)(i)] in Microsoft Word format, that includes the changes approved in this supplemental application, as well as annual reportable changes. To facilitate review of your submission(s), provide a highlighted or marked-up copy that shows all changes, as well as a clean Microsoft Word version. The marked-up copy should provide appropriate annotations, including supplement number(s) and annual report date(s).

REQUIRED PEDIATRIC ASSESSMENTS

Under the Pediatric Research Equity Act (PREA) (21 U.S.C. 355c), all applications for new active ingredients (which includes new salts and new fixed combinations), new indications, new dosage forms, new dosing regimens, or new routes of administration are required to contain an assessment of the safety and effectiveness of the product for the claimed indication in pediatric patients unless this requirement is waived, deferred, or inapplicable.

Because none of these criteria apply to your application and because this drug product for this indication has an orphan drug designation, you are exempt from this requirement.

REPORTING REQUIREMENTS

We remind you that you must comply with reporting requirements for an approved NDA (21 CFR 314.80 and 314.81).

² We update guidances periodically. For the most recent version of a guidance, check the FDA Guidance Documents Database <https://www.fda.gov/RegulatoryInformation/Guidances/default.htm>.

If you have any questions, contact Nicolas Kong, Regulatory Project Manager at Nicolas.Kong@fda.hhs.gov or 240-402-0269.

Sincerely,

{See appended electronic signature page}

Lisa Soule, M.D.
Associate Director
Division of Rare Diseases and Medical
Genetics (DRDMG)
Office of Rare Diseases, Pediatrics, Urologic
and Reproductive Medicine (ORPUM)
Center for Drug Evaluation and Research

ENCLOSURES:

- Content of Labeling
 - Prescribing Information

This is a representation of an electronic record that was signed electronically. Following this are manifestations of any and all electronic signatures for this electronic record.

/s/

LISA M SOULE
08/26/2020 03:01:06 PM