



NDA 210880

TENTATIVE APPROVAL

Hetero USA, Inc.
Attention: Dr. Soma Raju
U.S. Agent for Hetero Labs Limited Unit III, India
1035 Centennial Ave.
Piscataway, NJ 08854

Dear Dr. Raju:

Please refer to your New Drug Application (NDA) dated February 19, 2018, received February 20, 2018, and your amendments, submitted pursuant to section 505(b)(2) of the Federal Food, Drug, and Cosmetic Act for Dolutegravir, Lamivudine, and Tenofovir Disoproxil Fumarate Tablets, 50 mg/300 mg/300 mg.

This NDA provides for the use of Dolutegravir, Lamivudine, and Tenofovir Disoproxil Fumarate Tablets for the treatment of HIV-1 infection as a complete regimen in adults and pediatric patients weighing 40 kg or greater.

This NDA was reviewed under the President's Emergency Plan for AIDS Relief (PEPFAR).

We have completed our review of this application. It is tentatively approved under 21 CFR 314.105 for use as recommended in the agreed-upon enclosed labeling (text for the prescribing information, text for the patient package insert, carton and immediate container labels submitted on August 8, 2018). Based on the data provided, the expiration dating period is (b) (4) months for Dolutegravir, Lamivudine, and Tenofovir Disoproxil Fumarate Tablets, 50 mg/300 mg/300 mg, in the following packaging configuration when stored (b) (4)

(b) (4)
This determination is based upon information available to the Agency at this time, [i.e., information in your application and the status of current good manufacturing practices (cGMPs) of the facilities used in the manufacture and testing of the drug product]. This determination is subject to change on the basis of any new information that may come to our attention.

The listed drug [Tivicay[®] (dolutegravir)] upon which your application relies is subject to a period of patent and/or exclusivity protection and therefore final approval of your application under section 505(c)(3) of the Act [21 U.S.C. 355(c)(3)] may not be made effective until the period has expired.

To obtain final approval of this application, submit an amendment two or six months prior to the: 1.) expiration of the patent(s) and/or exclusivity protection or 2.) date you believe that your NDA

will be eligible for final approval, as appropriate. In your cover letter, clearly identify your amendment as **“REQUEST FOR FINAL APPROVAL”**. This amendment should provide the legal/regulatory basis for your request for final approval and should include a copy of any relevant court order or judgment settlement, or licensing agreement, as appropriate. In addition to a safety update, the amendment should also identify changes, if any, in the conditions under which your product was tentatively approved, i.e., updated labeling; chemistry, manufacturing, and controls data. This amendment should include labeling which complies with all current U.S. regulations (uniqueness of drug product appearance per 21 CFR 206; [REDACTED] (b) (4) [REDACTED] etc.). If there are no changes, clearly state so in your cover letter. Any changes require our review before final approval and the goal date for our review will be set accordingly.

Until we issue a final approval letter, this NDA is not deemed approved.

Please note that this drug product may not be marketed in the United States without final agency approval under Section 505 of the Act. The introduction or delivery for introduction into interstate commerce of this drug product before the final approval date is prohibited under Section 501 of the Act and 21 U.S.C. 331(d).

We remind you that if you intend to market this product in the United States after the period of patent and exclusivity protection expires, final approval of your application will require that you comply with all applicable U.S. legislation, including the Pediatric Research Equity Act (PREA) (21 U.S.C. 355c), which requires that all applications for new active ingredients, new indications, new dosage forms, new dosing regimens, or new routes of administration contain a pediatric assessment unless this requirement is waived, deferred, or inapplicable. A pediatric assessment contains data gathered from pediatric studies using appropriate formulations for each age group for which the assessment is required, and other data that are adequate to: 1.) assess the safety and effectiveness of the product for the claimed indications in all relevant pediatric subpopulations and 2.) support dosing and administration for each pediatric subpopulation for which the product has been assessed to be safe and effective. You should also join the antiretroviral pregnancy registry at that time and make the appropriate labeling change that references the existence of the pregnancy registry.

We also remind you that you are expected to comply with the reporting requirements provided in 21 CFR 314.80 and 314.81. If the product is to be mass distributed in developing countries, a system of collecting and reporting adverse drug reactions by the distributor would be desirable (e.g., through governmental or nongovernmental agencies distributing the products).

If you have any questions, call David Araujo, Pharm.D., Program Coordinator, at (301) 796-0669.

Sincerely,

{See appended electronic signature page}

Jeffrey S. Murray, M.D., M.P.H.
Deputy Director
Division of Antiviral Products
Office of Antimicrobial Products
Center for Drug Evaluation and Research

ENCLOSURE(S):

Content of Labeling
Carton and Container Labeling

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/

JEFFREY S MURRAY
08/08/2018