DEPARTMENT OF HEALTH & HUMAN SERVICES

Public Health Service

Food and Drug Administration
Rockville, MD 20857

NDA 21-974

Cipla Limited
ATTN: Ms Vaishali Shridhankar
Regulatory Affairs
289, Bellasis Rd
Mumbai Central
Mumbai 400083

Dear Ms Shridhankar:

Please refer to your new drug application (NDA) 21-974 submitted under rolling review with the final major submission dated, July 20, 2006 received on July 21, 2006, submitted pursuant to section 505(b)(2) of the Federal Food, Drug, and Cosmetic Act for Lamivudine 150mg/Stavudine 30mg Tablets and Lamivudine 150mg/Stavudine 40mg Tablets.

We acknowledge receipt of your submissions dated:
February 16, 2006
March 20, 2006
July 20, 2006
October 12, 2006 (2)
November 17, 2006

This NDA provides for the use of Lamivudine 150mg/Stavudine 30mg Tablets and Lamivudine 150mg/Stavudine 40mg Tablets in combination with other antiretrovirals for the treatment of HIV-1 infection.

We completed our review of this application. It is tentatively approved under 21 CFR 314.105 for use as recommended in the agreed upon labeling **in addition to the changes outlined below** (refer to the enclosed text for the package insert and patient package insert). Also refer to your submission emailed to the Agency on January 18, 2007 and to the agreed upon label dated February 16, 2006 for the immediate container and carton labels**.** The tentative approval is contingent upon information available to the Agency at this time (i.e. information in your application and the status of current good manufacturing practices of the facilities used in manufacturing and testing of the drug product) and is, therefore, subject to change on the basis of any new information that may come to our attention.

Additional changes to package insert and patient package insert:

- **Package insert:**

  -Under the Description and How Supplied section of the Package Insert minor chemistry changes were made. The changes are reflected in the attached label.

- **Patient Package insert:**

  -Under the inactive ingredient section minor chemistry changes were made. The changes are reflected in the attached label.

The listed reference drug products upon which you base your application are subject to a period of patent protection and therefore, final approval of your application under section 505(b) may not be made effective until the period has expired. If you have questions as to when this date will be please contact the Agency at the information provided below.

At least 180 days prior to the expiration of patent protection or when requested, submit a "**MINOR AMENDMENT – FINAL APPROVAL REQUESTED**" as an amendment to this application identifying changes, if any, in the conditions under which your product was tentatively approved. This information should include updated labeling, chemistry, manufacturing and control data, and a safety update. This amendment should include draft final printed labels and labeling which comply with all United States regulations (uniqueness of drug product appearance per 21 CFR 206; child-resistant packaging per 16 CFR 1700. etc). This amendment should be submitted even if none of these changes were made. This amendment should be designated clearly in your cover letter as a "**MINOR AMENDMENT – FINAL APPROVAL REQUESTED**".

Failure to submit this amendment will prompt a review of the application that may result in rescission of the tentative approval status of your application, or may result in a delay in the issuance of the final approval letter.

Any significant change in the conditions outlined in this NDA requires our review before final approval may be granted.

We remind you that you are expected to comply with the reporting requirements provided in 21 CFR 314.80 and 314.81. If the combination 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).

We remind you that, should you intend to market this product in the U.S. after the period of patent protection, you are required to join the antiretroviral pregnancy registry at that time and make the appropriate labeling change that references the existence of the pregnancy registry. In addition, an updated package insert (PI) must be submitted under the Structured Product Labeling requirements (http://www.fda.gov/oc/datacouncil/spl.html) as defined by the Physician's Labeling Rule [21 CFR 201.56, 201.57].

Before we issue a final approval letter, this NDA is not deemed approved. If you believe that there are grounds for issuing the final approval letter before the period of patent protection has expired, you should amend your application accordingly.

This product may be considered misbranded under the Federal Food, Drug, and Cosmetic Act if it is marketed in the U.S. before final approval.

If you have any questions, call Vasavi Reddy, RPh, MPH, Sr. Program Management Officer at (301) 796-0793 or via email at vasavi.reddy@fda.hhs.gov

Sincerely,

{See appended electronic signature page}

Debra Birnkrant, M.D.
Director
Division of Antiviral Drug Products
Office of Drug Evaluation IV
Center for Drug Evaluation and Research

Attachments: PI & PPI, Immediate container & carton label

Emailed CC:
Nicholas Cola
Byron Chemical Company, Inc.
U.S. Agent

This is a representation of an electronic record that was signed electronically and this page is the manifestation of the electronic signature.

/s/

Jeffrey Murray
1/19/2007 10:57:13 AM