



NDA 21-017/S-013

Eli Lilly and Company  
Attention: Jeffrey L. Winn, D.D.S., R.Ph.  
Senior Regulatory Research Scientist  
U.S. Regulatory Affairs  
Lilly Corporate Center  
Indianapolis, IN 46285

Dear Dr. Winn:

Please refer to your supplemental new drug application dated July 17, 2003, received July 18, 2003, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act for Humalog<sup>®</sup> Mix 75/25 (75% insulin lispro protamine suspension and 25% insulin lispro injection, [rDNA origin]).

We acknowledge receipt of your submissions dated October 14 (2), and November 18, 2003.

This supplemental new drug application provides for an additional manufacturing site for the assembly of the Humalog Mix 75/25 Pen in (b)(4)----- at the Lilly facility in Indianapolis, Indiana.

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

The final printed labeling (FPL) must be identical to the enclosed labeling (text for the package insert, text for the patient package insert, Humalog Mix 75/25 Pen User Manual, immediate container, and carton labels).

Please submit the FPL electronically according to the guidance for industry titled Providing Regulatory Submissions in Electronic Format – NDA. Alternatively, you may submit 20 paper copies of the FPL as soon as it is available, in no case more than 30 days after it is printed. Please individually mount 15 of the copies on heavy-weight paper or similar material. For administrative purposes, this submission should be designated "FPL for approved supplement NDA 21-017/S-013." Approval of this submission by FDA is not required before the labeling is used.

In addition, submit three copies of the introductory promotional materials that you propose to use for this product. Submit all proposed materials in draft or mock-up form, not final print. Send one copy to this division and two copies of both the promotional materials and the package insert directly to:

Division of Drug Marketing, Advertising, and Communications, HFD-42  
Food and Drug Administration  
5600 Fishers Lane  
Rockville, MD 20857

If you issue a letter communicating important information about this drug product (i.e., a “Dear Health Care Professional” letter), we request that you submit a copy of the letter to this NDA and a copy to the following address:

MEDWATCH, HFD-410  
FDA  
5600 Fishers Lane  
Rockville, MD 20857

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

If you have any questions, call Julie Rhee, Regulatory Project Manager, at (301) 827-6424.

Sincerely,

*{See appended electronic signature page}*

David G. Orloff, M.D.  
Director  
Division of Metabolic  
and Endocrine Drug Products, HFD-510  
Office of Drug Evaluation II  
Center for Drug Evaluation and Research

- Enclosures:
1. Package insert (A3.0 NL 3710 AMP)
  2. INFORMATION FOR THE PATIENT (A3.0 NL 3720 AMP)
  3. Disposable Insulin Delivery Device (Humalog Mix 75/25 Pen) User Manual (A3.0 NL 3730 AMP)
  4. Immediate container label for Humalog Mix 75/25 Pen (NL 9230 AMX)
  5. Carton label for Humalog Mix 75/25 Pen (NL 2530 AMS)



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**This is a representation of an electronic record that was signed electronically and  
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/s/

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David Orloff  
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