

# CENTER FOR DRUG EVALUATION AND RESEARCH

## Approval Package for:

### *APPLICATION NUMBER:*

**203496Orig1s000**

*Trade Name:* Orenitram Extended Release Tablets, 0.125 mg, 0.25 mg, 1mg, and 2.5 mg.

*Generic Name:* Treprostinil

*Sponsor:* United Therapeutics Corporation

*Approval Date:* December 20, 2013

*Indications:* Treatment of pulmonary arterial hypertension (PAH) (WHO Group 1) to improve exercise capacity.

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## 203496Orig1s000

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*APPLICATION NUMBER:*

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**APPROVAL LETTER**



NDA 203496

**NDA APPROVAL**

United Therapeutics Corporation  
Attention: Dean Bunce, RAC  
Executive Vice President,  
Regulatory Affairs and Compliance  
55T. W. Alexander Drive  
P.O. Box 14186  
Research Triangle Park, NC 27709

Dear Mr. Bunce:

Please refer to your New Drug Application (NDA) dated December 23, 2011, received December 27, 2011, resubmitted January 31 and August 16, 2013, under section 505(b)(1) of the Federal Food, Drug, and Cosmetic Act (FDCA), for Orenitram (Treprostinil) Extended Release Tablets, 0.125 mg, 0.25 mg, 1mg, and 2.5 mg.

We acknowledge receipt of your amendments dated August 16, 20, September 13, October 8, 29, November 27, and December 18, 2013.

The August 16, 2013, submission constituted a complete response to our March 22, 2013, action letter.

This new drug application provides for the use of Orenitram (Treprostrinil) Extended Release Tablets, 0.125 mg, 0.25 mg, 1mg, and 2.5 mg for treatment of pulmonary arterial hypertension (PAH) (WHO Group 1) to improve exercise capacity.

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 text.

### **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 <http://www.fda.gov/ForIndustry/DataStandards/StructuredProductLabeling/default.htm>. Content of labeling must be identical to the enclosed labeling (text for the package insert, text for the patient package insert). 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*, available at <http://www.fda.gov/downloads/Drugs/GuidanceComplianceRegulatoryInformation/Guidances/UCM072392.pdf>.

The SPL will be accessible via publicly available labeling repositories.

### **CARTON AND IMMEDIATE CONTAINER LABELS**

We acknowledge your October 29, 2013, submission containing final printed carton and container labels.

Please submit final printed carton and immediate container labels that are identical to the carton and immediate container labels submitted on October 29, 2013, as soon as they are available, but no more than 30 days after they are printed. Please submit these labels electronically according to the guidance for industry *Providing Regulatory Submissions in Electronic Format – Human Pharmaceutical Product Applications and Related Submissions Using the eCTD Specifications (June 2008)*. Alternatively, you may submit 12 paper copies, with 6 of the copies individually mounted on heavy-weight paper or similar material. For administrative purposes, designate this submission “**Final Printed Carton and Container Labels for approved NDA 203496.**” Approval of this submission by FDA is not required before the labeling is used.

Marketing the product with FPL that is not identical to the approved labeling text may render the product misbranded and an unapproved new drug.

### **REQUIRED PEDIATRIC ASSESSMENTS**

Under the Pediatric Research Equity Act (PREA) (21 U.S.C. 355c), all applications for new active ingredients, 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 this drug product for this indication has an orphan drug designation, you are exempt from this requirement.

### **PROMOTIONAL MATERIALS**

You may request advisory comments on proposed introductory advertising and promotional labeling. To do so, submit, in triplicate, a cover letter requesting advisory comments, the proposed materials in draft or mock-up form with annotated references, and the package insert to:

Food and Drug Administration  
Center for Drug Evaluation and Research  
Office of Prescription Drug Promotion  
5901-B Ammendale Road  
Beltsville, MD 20705-1266

As required under 21 CFR 314.81(b)(3)(i), you must submit final promotional materials, and the package insert, at the time of initial dissemination or publication, accompanied by a Form FDA

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