



NDA 21-232/S-008

SUPPLEMENT APPROVAL

R & R Registrations
Attention: Ronald Leonardi, Ph.D.
President, R & R Registrations
9915 Caminito Chirimolla
San Diego, CA 92131

Dear Dr. Leonardi:

Please refer to your supplemental new drug application dated October 23, 2008, received October 24, 2008, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act (FDCA) for Orfadin (nitisinone) Capsules.

We acknowledge receipt of your submissions dated February 24, 2009, May 8, 2009, July 23, 2009, and September 30, 2009.

This "Prior Approval" supplemental new drug application provides for conversion of the package insert (PI) to Physician's Labeling Rule (PLR) format.

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 and with the minor editorial revision listed below.

- Highlights: update 'Revised' date to 'January 2010'
- Full Prescribing Information, Section 5: fix numbering (change second 5.1 to 5.2)

As soon as possible, but no later than 14 days from the date of this letter, please submit the content of labeling [21 CFR 314.50(l)(1)(i)] in structured product labeling (SPL) format as described at <http://www.fda.gov/oc/datacouncil/spl.html> that is identical to, except for including the revisions listed above, the enclosed labeling, and labeling submitted September 30, 2009. For administrative purposes, please designate this submission, "SPL for approved NDA 21-232/S-008".

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
Division of Drug Marketing, Advertising, and Communications
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 2253. For instruction on completing the Form FDA 2253, see page 2 of the Form. For more information about submission of promotional materials to the Division of Drug Marketing, Advertising, and Communications (DDMAC), see <http://www.fda.gov/AboutFDA/CentersOffices/CDER/ucm090142.htm>.

Please submit one market package of the drug product when it is available.

LETTERS TO HEALTH CARE PROFESSIONALS

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

MedWatch
Food and Drug Administration
5600 Fishers Lane, Room 12B05
Rockville, MD 20857

REPORTING REQUIREMENTS

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 Heather Buck, Regulatory Project Manager, at (301) 796-1413.

Sincerely,

{See appended electronic signature page}

Donna Griebel, M.D.
Division Director
Division of Gastroenterology Products
Office of Drug Evaluation III
Center for Drug Evaluation and Research

Enclosure: Package Insert

Application
Type/Number

Submission
Type/Number

Submitter Name

Product Name

NDA-21232

SUPPL-8

RARE DISEASE
THERAPEUTICS
INC

ORFADIN (NITISINONE)
2/5/10MG CAPSULES

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

/s/

DONNA J GRIEBEL

01/25/2010