NDA 212157/S-002

GENERAL ADVICE

Dr. Reddy's Laboratories Limited.
c/o Dr. Reddy's Laboratories Inc.
Attention: Harpreet Kaur
Regulatory Affairs Associate
107 College Road East
Princeton, NJ 08540

Dear Mr. Harpreet:

Please refer to your supplemental new drug application (sNDA) dated and received November 6, 2020, and your amendments, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act for Elyxyb (celecoxib) oral solution.

We also refer to the supplement approval letter dated April 28, 2021. That letter inadvertently omitted the Medication Guide. The Medication Guide that was omitted (and that is included with the Supplement 2 approval) is appended to this letter.

If you have any questions, call Daniel Ngembus, Regulatory Project Manager, at (301) 837-7345.

Sincerely,

{See appended electronic signature page}

Alice T.D. Hughes, M.D.
Deputy Director for Safety
Division of Neurology 2
Office of Neuroscience
Center for Drug Evaluation and Research

Enclosure:
- Content of Labeling
  - Medication Guide

Medication Guide
ELYXYB (ee-lix'-ib))
(celecoxib) oral solution

What is the most important information I should know about ELYXYB?

ELYXYB contains celecoxib (a non-steroidal anti-inflammatory drug or NSAID). NSAIDs, including ELYXYB, can cause serious side effects, including:

- **Increased risk of a heart attack or stroke that can lead to death.** This risk may happen early in treatment and may increase:
  - with increasing doses of NSAIDs
  - with longer use of NSAIDs

Do not take ELYXYB right before or after a heart surgery called a "coronary artery bypass graft (CABG)."

Avoid taking NSAIDs, including ELYXYB, after a recent heart attack, unless your healthcare provider tells you to. You may have an increased risk of another heart attack if you take NSAIDs after a recent heart attack.

- **Increased risk of bleeding, ulcers, and tears (perforation) of the esophagus (tube leading from the mouth to the stomach), stomach and intestines:**
  - anytime during use
  - without warning symptoms
  - that may cause death

The risk of getting an ulcer or bleeding increases with:

- past history of stomach ulcers, or stomach or intestinal bleeding with use of NSAIDs
- taking medicines called "corticosteroids", "antiplatelet drugs", "anticoagulants", "SSRIs" or "SNRIs"
- increasing doses of NSAIDs
- longer use of NSAIDs
- smoking
- drinking alcohol
- older age
- poor health
- advanced liver disease
- bleeding problems

ELYXYB should only be used:

- exactly as prescribed
- for the shortest time needed

What is ELYXYB?

ELYXYB is a prescription medicine used for the acute treatment of migraine attacks with or without aura in adults.

- ELYXYB is not used as a preventive treatment of migraine.
- It is not known if ELYXYB is safe and effective in children.

Who should not take ELYXYB?

Do not take ELYXYB:

- if you are allergic to celecoxib or any of the ingredients in ELYXYB. See the end of this Medication Guide for a complete list of ingredients in ELYXYB.
- If you are allergic to sulfonamides.
- if you have had an asthma attack, hives, or other allergic reaction with aspirin or any other NSAIDs.
- right before or after heart bypass surgery.

Before taking ELYXYB, tell your healthcare provider about all of your medical conditions, including if you:

- have liver or kidney problems.
- have a history of stomach ulcer or bleeding in your stomach or intestines.
- have heart disease or risk factors that increase your chance of getting heart disease.
- have high blood pressure.
- have asthma.
- are pregnant or plan to become pregnant. Taking NSAIDs, including ELYXYB, at about 20 weeks of pregnancy or later may harm your unborn baby. If you need to take NSAIDs for more than 2 days when you are between 20 and 30 weeks of pregnancy, your healthcare provider may need to monitor the

amount of fluid in your womb around your baby. **You should not take NSAIDs after about 30 weeks of pregnancy.**

- are breastfeeding or plan to breast feed. ELYXYB may pass into your breast milk. Talk with your healthcare provider about the best way to feed your baby if you take ELYXYB.

Tell your healthcare provider about all of the medicines you take, including prescription or over-the-counter medicines, vitamins or herbal supplements. NSAIDs, including ELYXYB, and some other medicines can interact with each other and cause serious side effects. **Do not start taking any new medicine without talking to your healthcare provider first.**

How should I take ELYXYB?

See the detailed "Instructions for Use" on how to take ELYXYB solution.

- Take ELYXYB exactly as your healthcare provider tells you to take it.
- Take ELYXYB by mouth with or without food.
- Do not take more than one dose in a 24-hour period.
- Use ELYXYB for the fewest number of days a month, as needed.

What are the possible side effects of ELYXYB?

ELYXYB can cause serious side effects, including:

See "What is the most important information I should know about ELYXYB?

- liver problems including liver failure
- new or worse high blood pressure
- heart failure
- kidney problems including kidney failure
- life-threatening allergic reactions
- asthma attacks in people who have asthma
- life-threatening skin reactions
- medication overuse headaches. Some people who use too much ELYXYB may have worse headaches (medication overuse headache). If your headaches get worse, your healthcare provider may decide to stop your treatment with ELYXYB.
- low red blood cells (anemia)
- **Other side effects of NSAIDs include:** stomach pain, constipation, diarrhea, gas, heartburn, nausea, vomiting, and dizziness.

Get emergency help right away if you get any of the following symptoms:

- shortness of breath or trouble breathing
- chest pain
- weakness in one part or side of your body
- slurred speech
- swelling of the face or throat

Stop taking ELYXYB and call your healthcare provider right away if you get any of the following symptoms:

- nausea
- more tired or weaker than usual
- diarrhea
- itching
- your skin or eyes look yellow
- indigestion or stomach pain
- flu-like symptoms
- vomit blood
- there is blood in your bowel movement or it is black and sticky like tar
- unusual weight gain
- skin rash or blisters with fever
- swelling of the arms, legs, hands and feet

If you take too much ELYXYB, call your healthcare provider or get medical help right away.

These are not all the possible side effects of NSAIDs. For more information, ask your healthcare provider or pharmacist about NSAIDs. Call your doctor for medical advice about side effects. You may report side effects to FDA at 1-800-FDA-1088.

Other information about NSAIDs

- Aspirin is an NSAID but it does not increase the chance of a heart attack. Aspirin can cause bleeding in the brain, stomach, and intestines. Aspirin can also cause ulcers in the stomach and intestines.
- Some NSAIDs are sold in lower doses without a prescription (over-the counter). Talk to your healthcare provider before using over- the-counter NSAIDs for more than 10 days.

General information about the safe and effective use of ELYXYB

Medicines are sometimes prescribed for purposes other than those listed in a Medication Guide. Do not use ELYXYB for a condition for which it was not prescribed. Do not give ELYXYB to other people, even if they have the same symptoms that you have. It may harm them.

If you would like more information about ELYXYB, talk with your healthcare provider. You can ask your pharmacist or healthcare provider for information about ELYXYB that is written for health professionals.

Manufactured for: Dr. Reddy's Laboratories Limited

This Medication Guide has been approved by the U.S. Food and Drug Administration. Issued: 11/2020

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/

ALICE HUGHES
05/03/2021 10:24:53 AM

NDA 212157/S-002

SUPPLEMENT APPROVAL

Dr. Reddy's Laboratories Limited.
c/o Dr. Reddy's Laboratories Inc.
Attention: Harpreet Kaur
Regulatory Affairs Associate
107 College Road East
Princeton, NJ 08540

Dear Mr. Harpreet:

Please refer to your supplemental new drug application (sNDA) dated and received November 6, 2020, and your amendments, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act (FDCA) for Elyxyb (celecoxib) oral solution.

We also refer to our letter dated October 15, 2020, notifying you, under Section 505(o)(4) of the FDCA, of new safety information that we believe should be included in the labeling for nonsteroidal anti-inflammatory drug (NSAID) products. This information pertains to the serious risks of fetal renal dysfunction, oligohydramnios, and neonatal renal impairment, and Drug Reaction with Eosinophilia and Systemic Symptoms (DRESS).

This supplemental new drug application provides for revisions to the labeling for Elyxyb (celecoxib) oral solution, consistent with our October 15, 2020, letter.

APPROVAL & LABELING

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.

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 FDA.gov.[1] Content of labeling must be identical to the enclosed labeling (text for the Prescribing Information and Medication Guide), with the addition of any labeling changes in pending "Changes Being Effected" (CBE) supplements, as well as annual reportable changes not included in the enclosed labeling.

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*.[2]

The SPL will be accessible from publicly available labeling repositories.

Also within 14 days, amend all pending supplemental applications that include labeling changes for this NDA, including CBE supplements for which FDA has not yet issued an action letter, with the content of labeling [21 CFR 314.50(l)(1)(i)] in Microsoft Word format, that includes the changes approved in this supplemental application, as well as annual reportable changes. To facilitate review of your submission(s), provide a highlighted or marked-up copy that shows all changes, as well as a clean Microsoft Word version. The marked-up copy should provide appropriate annotations, including supplement number(s) and annual report date(s).

PROMOTIONAL MATERIALS

You may request advisory comments on proposed introductory advertising and promotional labeling. For information about submitting promotional materials, see the final guidance for industry *Providing Regulatory Submissions in Electronic and Non-Electronic Format-Promotional Labeling and Advertising Materials for Human Prescription Drugs.*[3]

[1] http://www.fda.gov/ForIndustry/DataStandards/StructuredProductLabeling/default.htm
[2] We update guidances periodically. For the most recent version of a guidance, check the FDA Guidance Documents Database https://www.fda.gov/RegulatoryInformation/Guidances/default.htm.

[3] For the most recent version of a guidance, check the FDA guidance web page at https://www.fda.gov/media/128163/download.

You must submit final promotional materials and Prescribing Information, accompanied by a Form FDA 2253, at the time of initial dissemination or publication [21 CFR 314.81(b)(3)(i)]. Form FDA 2253 is available at FDA.gov.[4] Information and Instructions for completing the form can be found at FDA.gov.[5]

All promotional materials that include representations about your drug product must be promptly revised to be consistent with the labeling changes approved in this supplement, including any new safety information [21 CFR 314.70(a)(4)]. The revisions in your promotional materials should include prominent disclosure of the important new safety information that appears in the revised labeling. Within 7 days of receipt of this letter, submit your statement of intent to comply with 21 CFR 314.70(a)(4).

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 Daniel Ngembus, Regulatory Project Manager, at (301) 837-7345.

Sincerely,

{See appended electronic signature page}

Alice T.D. Hughes, M.D.
Deputy Director for Safety
Division of Neurology 2
Office of Neuroscience
Center for Drug Evaluation and Research

ENCLOSURE(S):

- Content of Labeling
  - Prescribing Information
  - Medication Guide

[4] http://www.fda.gov/downloads/AboutFDA/ReportsManualsForms/Forms/UCM083570.pdf
[5] http://www.fda.gov/downloads/AboutFDA/ReportsManualsForms/Forms/UCM375154.pdf

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/s/

ALICE HUGHES
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