

BLA 761275/S-008

SUPPLEMENT APPROVAL

Fresenius Kabi USA, LLC
Attention: Ama Berko, PharmD
Director Regulatory Affairs
Three Corporate Drive
Lake Zurich, IL 60047

Dear Ama Berko:

Please refer to your supplemental biologics license application (sBLA), dated and received August 29, 2024, and your amendments, submitted under section 351(k) of the Public Health Service Act for Tynne (tocilizumab-aazg) injection.

This Category D Prior Approval supplemental biologics license application provides for the addition of the following indications:

- **Coronavirus Disease 2019 (COVID-19)**
Hospitalized adult patients with coronavirus disease 2019 (COVID-19) who are receiving systemic corticosteroids and require supplemental oxygen, non-invasive or invasive mechanical ventilation, or extracorporeal membrane oxygenation (ECMO).
- **Cytokine Release Syndrome (CRS)**
Adults and pediatric patients 2 years of age and older with chimeric antigen receptor (CAR) T cell-induced severe or life-threatening cytokine release syndrome.

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, via the FDA automated drug registration and listing system (eLIST), the content of labeling [21 CFR 601.14(b)] in structured product labeling (SPL) format, as described at FDA.gov,¹ that is identical to the enclosed labeling (text for the Prescribing Information

¹ <http://www.fda.gov/ForIndustry/DataStandards/StructuredProductLabeling/default.htm>

and Medication Guide) and include the labeling changes proposed in any pending “Changes Being Effected” (CBE) supplements.

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 via publicly available labeling repositories.

Also within 14 days, amend all pending supplemental applications that include labeling changes for this BLA, including pending “Changes Being Effected” (CBE) supplements, for which FDA has not yet issued an action letter, with the content of labeling [21 CFR 601.12(f)] 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).

REQUIRED PEDIATRIC ASSESSMENTS

Under the Pediatric Research Equity Act (PREA) (21 U.S.C. 355c), all applications for new active ingredients (which includes new salts and new fixed combinations), 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(s) in pediatric patients unless this requirement is waived, deferred, or inapplicable.

- **Cytokine Release Syndrome (CRS)**
At this time, we have determined that, with respect to this indication in pediatric patients 0 to <2 years of age, no pediatric assessment will be required under PREA for your sBLA. You have provided a pediatric assessment for CRS in pediatric patients 2 years of age and older, and nothing is further required at this time.
- **Coronavirus Disease 2019 (COVID-19)**
At this time, we have determined that, with respect to this indication in pediatric patients 0 to <1 year of age, no pediatric assessment will be required under PREA for your sBLA. We are deferring the required pediatric assessment for pediatric patients 1 to <18 years of age. See Deferred Pediatric Assessments below.

Deferred Pediatric Assessment

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>.

Your deferred pediatric study required under section 505B(a) of the Federal Food, Drug, and Cosmetic Act is required a postmarketing study. The status of this postmarketing study must be reported annually according to 21 CFR 601.28 and section 505B(a)(4)(C) of the Federal Food, Drug, and Cosmetic Act. This required study is listed below.

PMR 4806-1 Assessment of Tyenne (tocilizumab-aazg) for the treatment of Coronavirus Disease 2019 (COVID-19) in pediatric patients 1 to <18 years of age.
Final Report Submission: 06/2026

Reports of this required pediatric postmarketing assessment must be submitted as BLA or as a supplement to your approved BLA with the proposed labeling changes you believe are warranted based on the data derived from this assessment. When submitting the reports, please clearly mark your submission **"SUBMISSION OF REQUIRED PEDIATRIC ASSESSMENTS"** in large font, bolded type at the beginning of the cover letter of the submission.

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

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 601.12(f)(4)]. 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-related information [21 CFR 601.12(f)(4)]. The revisions in your promotional materials should include prominent disclosure of the important new safety-related 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 601.12(f)(4).

REPORTING REQUIREMENTS

We remind you that you must comply with reporting requirements for an approved BLA (in 21 CFR 600.80 and in 21 CFR 600.81).

If you have any questions, contact Chau Nguyen, Regulatory Project Manager, at (240)-402-0022 or chau.nguyen@fda.hhs.gov.

Sincerely,

{See appended electronic signature page}

Raj Nair, MD
Division Director
Division of Rheumatology and Transplant Medicine
Office of Immunology and Inflammation
Office of New Drugs
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>

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/

RAJ NAIR
02/28/2025 09:39:54 AM