



October 22, 2020

Ashley Rhoades, MBS, RAC
Manager, Regulatory Affairs
Gilead Sciences, Inc.
333 Lakeside Drive
Foster City, CA 94404

Dear Ms. Rhoades:

On February 4, 2020, pursuant to Section 564(b)(1)(C) of the Act, the Secretary of the Department of Health and Human Services (HHS) determined that there is a public health emergency that has a significant potential to affect national security or the health and security of United States citizens living abroad, and that involves the virus that causes COVID-19.¹ On the basis of such determination, the Secretary of HHS on March 27, 2020, declared that circumstances exist justifying the authorization of emergency use of drugs and biological products during the COVID-19 pandemic, pursuant to Section 564 of the Federal Food, Drug, and Cosmetic Act (the Act) (21 U.S.C. 360bbb-3), subject to terms of any authorization issued under that section.²

On May 1, 2020, the Food and Drug Administration (FDA) issued an Emergency Use Authorization (EUA) for emergency use of Veklury® (remdesivir) for the treatment of hospitalized patients with severe 2019 coronavirus disease (COVID-19)³, pursuant to Section 564 of the Act. Veklury is a direct acting antiviral drug that inhibits viral RNA synthesis. At that time, Veklury was an investigational drug and not approved for any indication.

On August 28, 2020, having concluded that revising this EUA was appropriate to protect the public health or safety under Section 564(g)(2) of the Act, FDA reissued the May 1, 2020, letter in its entirety with revisions incorporated to expand the authorized use of Veklury by no longer limiting its use to the treatment of patients with severe disease. In addition, the Fact Sheet for Health Care Providers was revised to provide updated clinical trial results and supporting data.⁴

¹ U.S. Department of Health and Human Services, *Determination of a Public Health Emergency and Declaration that Circumstances Exist Justifying Authorizations Pursuant to Section 564(b) of the Federal Food, Drug, and Cosmetic Act*, 21 U.S.C. § 360bbb-3. February 4, 2020.

² U.S. Department of Health and Human Services, *Declaration that Circumstances Exist Justifying Authorizations Pursuant to Section 564(b) of the Federal Food, Drug, and Cosmetic Act*, 21 U.S.C. § 360bbb-3, 85 FR 18250 (April 1, 2020).

³ For purposes of the May 1, 2020, EUA, patients with severe disease were defined as patients with oxygen saturation (SpO₂) ≤94% on room air or requiring supplemental oxygen or requiring mechanical ventilation or requiring extracorporeal membrane oxygenation (ECMO).

⁴ Prior to the reissuance of the EUA on August 28, 2020, and pursuant to the conditions of authorization, Gilead had requested, and FDA had concurred with, other changes to the Fact Sheets, including but not limited to: (1) clarified dosing and administration recommendations; (2) added sponsor's recommended formula to be used in calculating eGFR (this formula was removed in the August 28, 2020, reissuance); (3) added hypersensitivity reaction and drug interaction information; (4) added safety information from randomized, clinical trials; (5) removed information

On October 1, 2020, having concluded that revising this EUA was appropriate to protect the public health or safety under Section 564(g)(2) of the Act, FDA reissued the August 28, 2020, letter in its entirety with revisions incorporated to the scope and conditions of authorization designating Gilead Sciences, Inc. and its authorized distributors⁵ as the responsible parties for the distribution of Veklury. On October 16, 2020, FDA reissued the October 1, 2020, letter in its entirety with revisions to clarify that an alternate care site (ACS) meeting certain criteria was considered an “inpatient hospital setting” for the purposes of the scope of the EUA, and as such, was within the terms and conditions of FDA’s authorization.

On October 22, 2020, FDA approved NDA 214787 for Veklury (remdesivir), which is indicated for adults and pediatric patients (12 years and older and weighing at least 40 kg) for the treatment of COVID-19 requiring hospitalization. Under its approval, Veklury should only be administered in a hospital or in a healthcare setting capable of providing acute care comparable to inpatient hospital care. Having concluded that revising this EUA is appropriate to protect the public health or safety under Section 564(g)(2) of the Act, FDA is reissuing the October 16, 2020, letter in its entirety with revisions to remove uses previously authorized that are now the subject of the approved NDA 214787 for Veklury, and to continue authorizing Veklury for emergency use to treat suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg.

Veklury has activity in cell culture and animal models against SARS-CoV, MERS-CoV, and SARS-CoV-2. Based on review of the data from the randomized, double-blinded, placebo-controlled trial conducted by NIAID (NCT04280705), from the Gilead-sponsored open-label trial that evaluated different durations of Veklury (NCT04292899), and from the Gilead-sponsored open-label trial that evaluated different durations of Veklury as compared to standard of care (NCT04292730), it is reasonable to believe that Veklury may be effective and the known and potential benefits of Veklury outweigh the known and potential risks of the drug for the treatment of suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg.

Having concluded that the criteria for issuance of this authorization under Section 564(c) of the Act are met, I am authorizing the emergency use of Veklury for treatment of COVID-19, as described in the Scope of Authorization section of this reissued letter (Section II) and subject to the terms of this authorization.

I. Criteria for Issuance of Authorization

I have concluded that the emergency use of Veklury for the treatment of COVID-19 when administered as described in the Scope of Authorization (Section II) meets the criteria for issuance of an authorization under Section 564(c) of the Act, because:

related to the compassionate use program; and (6) added reference to remdesivir’s trade name, Veklury.

⁵ “Authorized Distributor(s)” are identified by Gilead as an entity or entities allowed to distribute authorized Veklury.

1. SARS-CoV-2 can cause a serious or life-threatening disease or condition, including severe respiratory illness, to humans infected by this virus;
2. Based on the totality of scientific evidence available to FDA, it is reasonable to believe that Veklury may be effective in treating suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg and that, when used under the conditions described in this authorization, the known and potential benefits of Veklury when used to treat suspected or laboratory-confirmed COVID-19 outweigh the known and potential risks of such products; and
3. There is no adequate, approved, and available alternative to the emergency use of Veklury to treat suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg.⁶

II. Scope of Authorization

I have concluded, pursuant to Section 564(d)(1) of the Act, that the scope of this authorization is limited as follows:

- The Veklury covered by this authorization will be used only to treat suspected or laboratory-confirmed COVID-19 in hospitalized⁷ pediatric patients weighing 3.5 kg to less than 40 kg *or* hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg administered via intravenous (IV) infusion by a healthcare provider; and
- The use of Veklury covered by this authorization should be in accordance with the dosing regimens as detailed in the authorized Facts Sheets.

Product Description

Veklury is a nucleoside ribonucleic acid (RNA) polymerase inhibitor. Veklury for injection, 100 mg, is a sterile, preservative-free lyophilized solid that is to be reconstituted with 19mL of sterile water for injection and diluted into 0.9% saline prior to intravenous (IV) administration. Following reconstitution, each single-dose, clear glass vial contains a 5 mg/mL Veklury concentrated solution with sufficient volume to allow withdrawal of 20 mL.

The authorized product includes remdesivir for injection⁸ with a vial label and/or carton labeling that is clearly marked for “emergency use authorization”, Veklury for injection with a vial label

⁶ No other criteria of issuance have been prescribed by regulation under Section 564(c)(4) of the Act.

⁷ Individuals determined as being appropriate for acute inpatient hospitalization and who are admitted or transferred to an alternate care site (ACS) that is capable of providing acute care that is comparable to general inpatient hospital care are within the terms and conditions of this Letter of Authorization. An ACS is intended to provide additional hospital surge capacity and capability for communities overwhelmed by patients with COVID-19.

⁸ FDA’s authorization includes remdesivir for injection manufactured and labeled prior to Gilead’s reference to remdesivir’s trade name, “Veklury”, in product labeling.

and/or carton labeling that is clearly marked for “emergency use authorization”, and commercially available⁹ Veklury for injection.

Veklury for injection, 100 mg, vials should be stored below 30 °C until time of use. Following reconstitution with sterile water and dilution with 0.9% saline, the solution can be stored for up to 24 hours at room temperature (20 °C to 25 °C) or 48 hours at refrigerated temperatures (2 °C to 8 °C).

Veklury is authorized for emergency use with the following product-specific information required to be made available to healthcare providers and patients, respectively, through Gilead’s website at www.gilead.com/remdesivir:

- Fact Sheet for Health Care Providers: Emergency Use Authorization (EUA) of Veklury (remdesivir)
- Fact Sheet for Parents and Caregivers: Emergency Use Authorization (EUA) of Veklury (remdesivir) for Coronavirus Disease 2019 (COVID-19)

I have concluded, pursuant to Section 564(d)(2) of the Act, that it is reasonable to believe that the known and potential benefits of Veklury, when used for the treatment of COVID-19 and used in accordance with this Scope of Authorization (Section II), outweigh its known and potential risks.

I have concluded, pursuant to Section 564(d)(3) of the Act, based on the totality of scientific evidence available to FDA, that it is reasonable to believe that Veklury may be effective for the treatment of COVID-19 when used in accordance with this Scope of Authorization (Section II), pursuant to Section 564(c)(2)(A) of the Act.

Having reviewed the scientific information available to FDA, including the information supporting the conclusions described in Section I above, I have concluded that Veklury (as described in this Scope of Authorization (Section II)) meets the criteria set forth in Section 564(c) of the Act concerning safety and potential effectiveness.

The emergency use of your product under an EUA must be consistent with, and may not exceed, the terms of the Authorization, including the Scope of Authorization (Section II) and the Conditions of Authorization (Section III). Subject to the terms of this EUA and under the circumstances set forth in the Secretary of HHS’s determination under Section 564(b)(1)(C) described above and the Secretary of HHS’s corresponding declaration under Section 564(b)(1), Veklury is authorized to treat suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg as described in the Scope of Authorization (Section II) under this EUA, despite the fact that it does not meet certain requirements otherwise required by applicable federal law.

⁹ For the purposes of this Letter of Authorization, commercially available Veklury for injection refers to product in United States distribution under the approved New Drug Application 214787.

III. Conditions of Authorization

Pursuant to Section 564 of the Act, I am establishing the following conditions on this authorization:

Gilead Sciences, Inc. (Gilead) and Authorized Distributors

- A. Gilead and authorized distributor(s) will ensure that the authorized Veklury is distributed and the authorized labeling (i.e., Fact Sheets) will be made available to hospitals and healthcare facilities consistent with the terms of this letter.
- B. Gilead and authorized distributor(s) will ensure that appropriate storage is maintained until the product is delivered to hospitals and healthcare facilities.
- C. Gilead and authorized distributor(s) will ensure that the terms of this EUA are made available to all relevant stakeholders (e.g., U.S. government agencies, state and local government authorities, authorized distributors, healthcare facilities, healthcare providers) involved in distributing or receiving authorized Veklury. Gilead will provide to all relevant stakeholders a copy of this letter of authorization and communicate any subsequent amendments that might be made to this letter of authorization and its authorized accompanying materials (i.e., Fact Sheets).
- D. Gilead may request changes to this authorization, including to the authorized Fact Sheets for Veklury, and FDA may determine that such changes may be permitted without amendment of this EUA, upon concurrence of the Office of Infectious Diseases/Office of New Drugs/Center for Drug Evaluation and Research (CDER), the Counter-Terrorism and Emergency Coordination Staff/Office of the Center Director/CDER, and Office of Counterterrorism and Emerging Threats/Office of the Chief Scientist/Office of the Commissioner.
- E. Gilead will report to FDA serious adverse events and all medication errors associated with the use of the authorized Veklury that are reported to Gilead using either of the following options.

Option 1: Submit reports through the Safety Reporting Portal (SRP) as described on the [FDA SRP](#) web page.

Option 2: Submit reports directly through the Electronic Submissions Gateway (ESG) as described on the [FAERS electronic submissions](#) web page.

Submitted reports under both options should state: “use of Veklury (remdesivir) was under an EUA”. For reports submitted under Option 1, include this language at the beginning of the question “Describe Event” for further analysis. For reports submitted under Option 2, include this language at the beginning of the “Case Narrative” field.

- F. Through a process of inventory control, Gilead and authorized distributor(s) will maintain records regarding distribution of the authorized Veklury (i.e., lot numbers, quantity, receiving site, receipt date).
- G. Gilead and authorized distributor(s) will make available to FDA upon request any records maintained in connection with this EUA.

Hospitals and Other Healthcare Facilities to Whom the Authorized Veklury Is Distributed and Healthcare Providers Administering the Authorized Veklury

- H. Healthcare facilities and healthcare providers will ensure that they are aware of the letter of authorization, and the terms herein, and that the authorized Fact Sheets are made available to healthcare providers and to patients and caregivers, respectively, through appropriate means.
- I. Healthcare facilities and healthcare providers receiving Veklury will track serious adverse events that are considered to be potentially attributable to Veklury use and must report these to FDA in accordance with the Fact Sheet for Healthcare Providers. Complete and submit a MedWatch form (www.fda.gov/medwatch/report.htm), or Complete and submit FDA Form 3500 (health professional) by fax (1-800-FDA-0178) (these forms can be found via link above). Call [1-800-FDA-1088](tel:1-800-FDA-1088) for questions. Submitted reports should state, “use of Veklury (remdesivir) was under an EUA” at the beginning of the question “Describe Event” for further analysis.
- J. Through a process of inventory control, healthcare facilities will maintain records regarding the dispensed authorized Veklury (i.e., lot numbers, quantity, receiving site, receipt date), product storage, and maintain patient information (e.g., patient name, age, disease manifestation, number of doses administered per patient, other drugs administered).
- K. Healthcare facilities will ensure that any records associated with this EUA are maintained until notified by Gilead and/or FDA. Such records will be made available to Gilead, HHS, and FDA for inspection upon request.

Conditions Related to Printed Matter, Advertising and Promotion

- L. All descriptive printed matter, including advertising and promotional material, relating to the use of the Veklury for the treatment of suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg shall be consistent with the authorized labeling, as well as the terms set forth in this EUA and the applicable requirements set forth in the Act and FDA regulations.
- M. No descriptive printed matter, including advertising or promotional material, relating to the use of the Veklury may represent or suggest that such products are safe or effective when used for the treatment of suspected or laboratory-confirmed COVID-19 in hospitalized

pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg.

N. All descriptive printed matter, including advertising and promotional material, relating to the use of the Veklury to treat suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg or hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg shall clearly and conspicuously state that:

- the Veklury has not been approved, but has been authorized for emergency use by FDA to treat suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg *or* hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg;
- the Veklury is authorized under an EUA for the treatment of suspected or laboratory-confirmed COVID-19 in hospitalized pediatric patients weighing 3.5 kg to less than 40 kg *or* hospitalized pediatric patients less than 12 years of age weighing at least 3.5 kg only for the duration of the declaration that circumstances exist justifying the authorization of the emergency use of the Veklury under Section 564(b)(1) of the Act, 21 U.S.C. § 360bbb-3(b)(1), unless the authorization is terminated or revoked sooner.

IV. Duration of Authorization

This EUA will be effective until the declaration that circumstances exist justifying the authorization of the emergency use of drugs and biological products during the COVID-19 pandemic is terminated under Section 564(b)(2) of the Act or the EUA is revoked under Section 564(g) of the Act.

Sincerely,

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RADM Denise M. Hinton
Chief Scientist
Food and Drug Administration