

**FOR FURTHER INFORMATION CONTACT:**

Larry Bauer, Rare Diseases Program, Center for Drug Evaluation and Research, Food and Drug Administration, 10903 New Hampshire Ave., Silver Spring, MD 20993-0002, 301-796-4842, FAX: 301-796-9858, email: [larry.bauer@fda.hhs.gov](mailto:larry.bauer@fda.hhs.gov).

**SUPPLEMENTARY INFORMATION:** FDA is announcing the issuance of a priority review voucher to the sponsor of a rare pediatric disease product application. Under section 529 of the FD&C Act (21 U.S.C. 360ff), added by FDASIA, FDA will award priority review vouchers to sponsors of rare pediatric disease product applications that meet certain criteria. FDA has determined that UNITUXIN (dinutuximab), manufactured by United Therapeutics Corporation, meets the criteria for a priority review voucher. UNITUXIN (dinutuximab) is indicated, in combination with granulocyte-macrophage colony-stimulating factor (GM-CSF), interleukin-2 (IL-2), and 13-cis-retinoic acid (RA), for the treatment of pediatric patients with high-risk neuroblastoma who achieve at least a partial response to prior first-line multiagent, multimodality therapy. Neuroblastoma is the most common pediatric solid tumor occurring outside the brain, and it is the most common cancer in infants.

For further information about the Rare Pediatric Disease Priority Review Voucher Program and for a link to the full text of section 529 of the FD&C Act, go to <http://www.fda.gov/ForIndustry/DevelopingProductsforRareDiseasesConditions/RarePediatricDiseasePriorityVoucherProgram/default.htm>.

For further information about UNITUXIN (dinutuximab), go to the Drugs@FDA Web site at <http://www.accessdata.fda.gov/scripts/cder/drugsatfda/index.cfm>.

Dated: April 2, 2015.

**Leslie Kux,**

*Associate Commissioner for Policy.*

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## DEPARTMENT OF HEALTH AND HUMAN SERVICES

### Food and Drug Administration

[Docket No. FDA-2014-D-1747]

#### Risk Evaluation and Mitigation Strategies: Modifications and Revisions; Guidance for Industry; Availability

**AGENCY:** Food and Drug Administration, HHS.

**ACTION:** Notice.

**SUMMARY:** The Food and Drug Administration (FDA) is announcing the availability of a guidance for industry entitled “Risk Evaluation and Mitigation Strategies: Modifications and Revisions.” This guidance provides information on how FDA will define and process submissions for modifications and revisions to risk evaluation and mitigation strategies (REMS), as well as information on what types of changes to approved REMS will be considered modifications of the REMS and what types of changes will be considered revisions of the REMS. There are different procedures for submission of REMS modifications and revisions to FDA as well as different timeframes for FDA review and action of such changes. In addition, this guidance provides information on how REMS modifications and revisions should be submitted to FDA and how FDA intends to review and act on these submissions. The definitions of REMS modifications and revisions apply to all types of REMS.

**DATES:** Although you can comment on any guidance at any time (see 21 CFR 10.115(g)(5)), to ensure that the Agency considers your comment on this guidance before it begins work on the final version of the guidance, submit either electronic or written comments on the guidance by June 8, 2015. Submit either electronic or written comments concerning the proposed collection of information by June 8, 2015.

**ADDRESSES:** Submit written requests for single copies of the guidance to the Division of Drug Information, Center for Drug Evaluation and Research, Food and Drug Administration, 10001 New Hampshire Ave. Hillandale Building, 4th Floor, Silver Spring, MD 20993-0002; or the Office of Communication, Outreach, and Development, Center for Biologics Evaluation and Research, Food and Drug Administration, 10903 New Hampshire Ave. Bldg. 71, Rm. 3128, Silver Spring, MD 20993-0002. Send one self-addressed adhesive label to assist that office in processing your requests. See the **SUPPLEMENTARY INFORMATION** section for electronic access to the guidance document.

Submit electronic comments on the guidance to <http://www.regulations.gov>. Submit written comments to the Division of Dockets Management (HFA-305), Food and Drug Administration, 5630 Fishers Lane, Rm. 1061, Rockville, MD 20852.

**FOR FURTHER INFORMATION CONTACT:** Kristen Everett, Center for Drug Evaluation and Research, Food and

Drug Administration, 10903 New Hampshire Ave. Bldg. 22, Rm. 6484, Silver Spring, MD 20993-0002, 301-796-0453; or Stephen Ripley, Center for Biologics Evaluation and Research, Food and Drug Administration, 10903 New Hampshire Ave. Bldg. 71, Rm. 7301, Silver Spring, MD 20993-0002, 240-402-7911.

#### **SUPPLEMENTARY INFORMATION:**

##### **I. Background**

FDA is announcing the availability of a guidance for industry entitled “Risk Evaluation and Mitigation Strategies: Modifications and Revisions.” This guidance provides information on what types of changes to approved REMS will be considered modifications and what types of changes will be considered revisions. See section 505-1(h) of the Federal Food, Drug, and Cosmetic Act (the FD&C Act) (21 U.S.C. 355-1(h)). This guidance also provides information on how REMS modifications and revisions should be submitted to FDA and how FDA intends to review and act on these submissions.

If FDA determines that a REMS is necessary to ensure that the benefits of a drug outweigh its risks, FDA is authorized to require a REMS for such drugs under section 505-1 of the FD&C Act,<sup>1</sup> added by section 901 of the Food and Drug Administration Amendments Act of 2007 (Pub. L. 110-85).<sup>2</sup> Section 505-1(g) and (h) of the FD&C Act include provisions for the assessment and modification of an approved REMS.

In 2009, FDA issued draft guidance on the format and content of REMS, REMS assessments, and proposed REMS modifications. In that guidance, based on the language of section 505-1(g) and (h) of the FD&C Act before the amendments made by the Food and Drug Administration Safety and Innovation Act (Pub. L. 112-144) (FDASIA), FDA stated that any proposed modification to an approved REMS, including proposed changes to materials that are appended to the REMS document, must be submitted as a proposed REMS modification in the form of a prior approval supplement and must include a REMS assessment. The guidance stated that the proposed

<sup>1</sup> Section 505-1 of the FD&C Act applies to applications for prescription drugs submitted under subsection 505(b) (*i.e.*, new drug applications) or (j) (*i.e.*, abbreviated new drug applications) of the FD&C Act (21 U.S.C. 355) and applications under section 351 of the Public Health Service Act (*i.e.*, biologics license applications).

<sup>2</sup> See <http://www.fda.gov/RegulatoryInformation/Legislation/FederalFoodDrugandCosmeticAct/FDCAct/SignificantAmendmentsToTheFDCAct/FoodandDrugAdministrationAmendmentsActof2007/default.htm>.

modification(s) may not be implemented until approved by FDA.

FDASIA amended the REMS modification provisions under section 505–1(g) and (h) of the FD&C Act. Section 505–1(h), as amended by FDASIA, requires FDA to review and act on proposed “minor modifications,” as defined in guidance, within 60 days.<sup>3</sup> It also requires FDA to establish, through guidance, that “certain modifications” can be implemented following notification to FDA.<sup>4</sup> In addition, FDASIA requires FDA to review and act on REMS modifications due to approved safety label changes, or to a safety label change that FDA has directed the application holder to make pursuant to section 505(o)(4) of the FD&C Act within 60 days.<sup>5</sup> Finally, FDASIA specifies that proposed REMS modifications no longer require submission of a REMS assessment; instead, proposed modifications must include an adequate rationale for the proposed changes. This guidance is issued pursuant to section 505–1(h)(2)(A)(ii), (h)(2)(A)(iii), and (h)(2)(A)(iv) of the FD&C Act.

This guidance is being issued consistent with FDA’s good guidance practices regulation (21 CFR 10.115). This guidance, except for the portion setting forth the submission procedures for REMS revisions, is being implemented without prior public comment because the Agency has determined that prior public participation is not feasible or appropriate (21 CFR 10.115(g)(2)). The Agency made this determination because, consistent with the requirements of FDASIA, FDA is issuing this guidance to establish a less burdensome policy and process for submitting certain changes to REMS that is consistent with public health. Although the guidance document is immediately in effect, except for the submission procedures for REMS revisions, it remains subject to comment in accordance with the Agency’s good guidance practices. Insofar as this guidance establishes the modifications to an approved REMS that may be implemented following notification to the Secretary under section 505–1(h)(2)(A)(iv)—here referred to as REMS

revisions—it has binding effect, except for the portion of the guidance setting forth the submission procedure for REMS revisions, which will, when final, have binding effect.

## II. Paperwork Reduction Act of 1995

This guidance contains information collection provisions that are subject to review by the Office of Management and Budget (OMB) under the Paperwork Reduction Act of 1995 (44 U.S.C. 3501–3520). The title, description, and respondent description of the information collection are given under this section with an estimate of the annual reporting burden. Included in the estimate is the time for reviewing instructions, searching existing data sources, gathering and maintaining the data needed, and completing and reviewing the collection of information.

We invite comments on these topics: (1) Whether the proposed collection of information is necessary for the proper performance of FDA’s functions, including whether the information will have practical utility; (2) the accuracy of FDA’s estimate of the burden of the proposed collection of information, including the validity of the methodology and assumptions used; (3) ways to enhance the quality, utility, and clarity of the information to be collected; and (4) ways to minimize the burden of the collection of information on respondents, including through the use of automated collection techniques, when appropriate, and other forms of information technology.

*Title:* Guidance for Industry on Risk Evaluation and Mitigation Strategies: Modifications and Revisions

*Description:* The guidance provides information on submitting to FDA modifications and revisions to approved REMS for approved new drug applications (NDAs), abbreviated new drug applications (ANDAs), or biologics license applications (BLAs).

REMS modifications are submitted to FDA as supplements to approved NDAs under 21 CFR 314.70 and for ANDAs under 21 CFR 314.97, and as supplements to approved BLAs under 21 CFR 601.12. The burden hours for preparing and submitting supplements to NDAs and ANDAs is approved by OMB under control number 0910–0001, and for BLAs under control number 0910–0338.

Concerning REMS revisions, application holders should include the following information in each submission: (1) A full description of the changes to the REMS and/or appended materials, the date the changes will be implemented, and a REMS history that outlines all changes made to the REMS since its approval; (2) a clean Word version of the revised REMS and all appended REMS materials; (3) a redlined (tracked changes) Word version of the revised REMS and revised appended REMS materials that shows the changes from the previous versions; (4) an updated REMS supporting document, if needed; and (5) Form FDA 356h indicating that the submission is a REMS revision. (Form FDA 356h is approved by OMB under control number 0910–0338.) Each REMS revision that is submitted to FDA should also be documented in the next annual report for the application under 21 CFR 314.81(b)(2) (the burden hours for preparing and submitting annual reports for NDAs and ANDAs is approved by OMB under control number 0910–0001, and for BLAs under control number 0910–0338). All subsequent REMS submissions (*i.e.*, proposed modifications or additional REMS revisions) should include previously implemented REMS revisions in the REMS document and appended materials, and should be noted in the REMS history.

Currently, there are 117 application holders with approved REMS that include 152 drugs. Based on FDA’s current review of REMS submissions for approved NDAs, ANDAs, and BLAs, and anticipating an average of 1 REMS revision across the entire group of REMS, we estimate that annually a total of approximately 117 application holders (“Number of Respondents” in table 1) will submit to FDA approximately 152 REMS revision submissions (“Total Annual Responses” in table 1) as described in this document and in the guidance. We also estimate that it will take an application holder approximately 30 hours to prepare and submit to FDA each REMS revision (“Average Burden per Response” in table 1).

The total estimated reporting burden for the guidance is as follows:

<sup>3</sup> See section 505–1(h)(2)(A)(ii) of the FD&C Act.

<sup>4</sup> See section 505–1(h)(2)(A)(iv) of the FD&C Act.

<sup>5</sup> See section 505–1(h)(2)(A)(iii) of the FD&C Act.

TABLE 1—ESTIMATED ANNUAL REPORTING BURDEN<sup>1</sup>

Guidance for industry on risk evaluation and mitigation strategies: modifications and revisions	Number of respondents	Number of responses per respondent	Total annual responses	Average burden per response	Total hours
REMS revisions .....	117	1	152	30	4,560

<sup>1</sup> There are no capital costs or operating and maintenance costs associated with this collection of information.

### III. Comments

Interested persons may submit either electronic comments regarding this document to <http://www.regulations.gov> or written comments to the Division of Dockets Management (see **ADDRESSES**). It is only necessary to send one set of comments. Identify comments with the docket number found in brackets in the heading of this document. Received comments may be seen in the Division of Dockets Management between 9 a.m. and 4 p.m., Monday through Friday, and will be posted to the docket at <http://www.regulations.gov>.

### IV. Electronic Access

Persons with access to the Internet may obtain the document at <http://www.fda.gov/Drugs/GuidanceComplianceRegulatoryInformation/Guidances/default.htm>, <http://www.fda.gov/BiologicsBloodVaccines/GuidanceComplianceRegulatoryInformation/default.htm>, or <http://www.regulations.gov>.

Dated: April 2, 2015.

**Leslie Kux,**

*Associate Commissioner for Policy.*

[FR Doc. 2015-08015 Filed 4-6-15; 8:45 am]

**BILLING CODE CODE 4164-01-P**

## DEPARTMENT OF HEALTH AND HUMAN SERVICES

### Health Resources and Services Administration

#### Privacy Act of 1974; Report of an Altered System of Records

**AGENCY:** Health Resources and Services Administration (HRSA), Department of Health and Human Services (HHS or Department).

**ACTION:** Notice of an Altered System of Records.

**SUMMARY:** In accordance with the requirements of the Privacy Act of 1974 (5 U.S.C. 552a), the Health Resources and Services Administration (HRSA) is publishing notice of a proposed alteration of the system of records entitled and numbered “Public Health and National Health Service Corps Scholarship Program (NHSC SP),

National Health Service Corps Loan Repayment Program (NHSC LRP), Students to Service, (S2S), NHSC Student/Resident Experiences and Rotations in Community Health (SEARCH), NURSE Corps Loan Repayment Program (NURSE Corps LRP) formerly the Nursing Education Loan Repayment Program (NELRP), NURSE Corps Scholarship Program (NURSE Corps SP) formerly the Nursing Scholarship Program (NSP), Native Hawaiian Health Scholarship Program (NHHSP), and Faculty Loan Repayment Program (FLRP), Applicants and/or Participants Records System, HHS/HRSA/BHW,” No. 09-15-0037. The proposed alterations affect the system name, system location, categories of records, purposes, routine uses, safeguards, records retention and disposal, system manager title and address, as well as minor editorial corrections and clarifications.

**DATES:** HRSA filed an altered system report with the Chair of the House Committee on Government Reform and Oversight, the Chair of the Senate Committee on Homeland Security and Governmental Affairs, and the Administrator, Office of Information and Regulatory Affairs, Office of Management and Budget (OMB) on March 30, 2015. To ensure all parties have adequate time in which to comment, the altered system, including the routine uses, will become effective 30 days from the publication of the notice or 40 days from the date it was submitted to OMB and Congress, whichever is later, unless HRSA receives comments that require alterations to this notice.

**ADDRESSES:** Please address comments to: Associate Administrator, Bureau of Health Workforce (BHW), Health Resources and Services Administration (HRSA), 5600 Fishers Lane, Room 11W-37, Rockville, MD 20857, telephone (301) 594-4130, or FAX (301) 594-4076. Comments received will be available for inspection at this same address from 9:00 a.m. to 3:00 p.m. (Eastern Standard Time Zone), Monday through Friday.

**FOR FURTHER INFORMATION CONTACT:** BMISS System Manager, Bureau of Health Workforce, Health Resources and Services Administration (HRSA), 5600

Fishers Lane, Room 11W-37, Rockville, MD 20857, Telephone: 301-443-1587.

### SUPPLEMENTARY INFORMATION:

#### I. Explanation of Changes

- The system of records notice name has been shortened to “HHS/HRSA/BHW Scholarship and Loan Repayment Program Records.”

- The Nursing Scholarship Program (NSP) has been renamed the NURSE Corps Scholarship Program (NURSE Corps SP), and the Nursing Education Loan Repayment Program (NELRP) has been renamed the NURSE Corps Loan Repayment Program (NURSE Corps LRP) and all associated records have been merged into a new central database system as noted below.

- A new information system, BHW Management Information System Solution (BMISS), has replaced Bureau of Health Care Delivery and Assistance NET (BHCDANET), and serves as the central database for information concerning the, NHSC SP, NHSC LRP, S2S, NURSE Corps SP, NURSE Corps LRP, FLRP, and NHHSP.

- The system location section has been updated to indicate that electronic records and electronic copies of paper records for applicants and participants under various programs are now stored in BMISS and to include locations of records not stored in BMISS (for example, Ambassador records are electronic but currently maintained in an online Web site directory while a BMISS database is being designed and built, at which point the records will be merged into BMISS).

- The categories of records have been updated to include “information concerning educational loans.”

- The purpose(s) section has been updated to consolidate certain descriptions (*i.e.*, to combine the loan repayment and scholarship program monitoring activities previously described in purpose 4 with the other program selection and monitoring activities previously described in purpose 8), and to include intra-agency transfers of information previously described as routine uses by mistake (*i.e.*, transfers to HHS’ debt and financial management systems).