

To: Ms. Sharon Shapowal
From: Helen S. Gemignani
File: STN 125259/0 CERVARIX
RE: Request for CMC information
Date: October 1, 2007

APPROVED

By Helen S. Gemignani at 7:10 pm, Mar 11, 2008

As communicated by Dr. Robin Levis during inspection of your facility in Rixensart, and as noted in Item #2 of Form 483 issued on September 21, 2007, we request that you address the following items:

1. Regarding the deviation that you reported for filter clogging during filling, we request that you provide your results (including data) from your investigation of the deviation. We also request that you provide all available data to support the [REDACTED] filter following formulation.

2. Please submit validation data to support repeated use of the filters and columns used in the antigen purification process. Please note that the process will be approved only for the number of repeat uses of the filters and columns for which you have validation data. If validation studies have not been completed, we request that you submit interim reports.

3. We request that you submit the most recent stability data for all samples under stability study. In each case, we request that you include the lot number, the date of manufacture and the date of final fill, where applicable.

4. Please submit any updates or changes to the Draft Bulk and Final Product Release Protocols.

5. Since you do not have validation data for [REDACTED] of purified antigen bulks [REDACTED]

(b)(4)

we request that you submit an amendment to the BLA indicating that you are removing these sections of the BLA from review consideration. Please acknowledge.

6. Please submit a translation of GSK Monograph 20092101 which provides a detailed description of final container testing.

7. Please submit corrections to the BLA as detailed below:

a. Provide corrected data on [REDACTED] test for HPV 18 adsorbed monovalent bulks by the [REDACTED] assay – 3.2.P.3.4 – Table 4.

b. Revise the specification for the QC test for General Safety to remove the statement regarding the number of times the test will be conducted post-licensure.

c. Revise the HVAC fresh air specification from [REDACTED] in section 3.2.A.1.

General comment:

8. We are concerned about the number of inaccuracies in the CMC information provided in the BLA that were identified during inspection. We request that you QC the CMC section of the BLA and if any additional inaccuracies are identified, we request that you provide corrections to CBER as soon as possible. Please respond.

In your reply to this memo, we recommend that you restate each item and follow it with your explanation or clarification. Use of this format helps organize the relevant information and provides a self-contained document that facilitates future reference.