Health Canada | Santé Canada

Health Products and Food Branch
Direction générale des produits de santé et des aliments

TERMS AND CONDITIONS

Company: Moderna Biopharma Canada Corporation

Product: Spikevax Bivalent (Original / Omicron BA. 4/5) (elasomeran/devasomeran) mRNA vaccine

Dossier ID: HC6-024-E267589

Background:

The Food and Drug Regulations allows the Minister to impose or amend terms and conditions, and request additional information, in relation to a COVID-19 drug submission, Drug Identification Number (DIN), or establishment licence, at any time while it is in effect. In light of the severity of the COVID-19 pandemic, this allows the Minister to act quickly to gather important safety information or mitigate risk in a timely manner.

Depending on the requirement, terms and conditions can be ongoing (e.g., require a monthly report), have a defined time (e.g., a report is due to Health Canada on a specific day) or are to be completed once the data is available (e.g., a clinical trial is completed).

The status of the Terms and Conditions will be updated on a regular basis.

Status as of October 4, 2023:

Table: Terms and Conditions

	Terms and Conditions	Issued	Status
1	Provide the Day 29 immunogenicity, safety, and reactogenicity data from Study mRNA-1273-P205 Part H, as soon as the data become available.	November 3, 2022	Ongoing
2	Provide the Day 181 immunogenicity, safety, and reactogenicity data from Study mRNA-1273-P205 Part H, as soon as the final study report becomes available.	November 3, 2022	Ongoing
3	For the Drug Substance (DS) Provide stability updates in a timely manner.	November 3, 2022	Closed
4	Provide Stability updates in a timely manner, for the DP.	November 3, 2022	Closed
5	Moderna Biopharma Canada Corporation is required to treat adverse reactions associated with Spikevax Bivalent (Original / Omicron BA.4/5) as priority and submit the	November 3, 2022	Closed

	corresponding reports to Health Canada without delay.		
6	Moderna Biopharma Canada Corporation is required to submit Monthly Safety Summary Reports (MSSRs) for Spikevax Bivalent (Original / Omicron BA.4/5), unless otherwise determined by Health Canada. The MSSR should be submitted within 15 days after the last day of a month, beginning after the first full calendar month after authorization. These reports should contain the following: a. Interval and cumulative number of reports (serious and non-serious), overall and by age groups and in special populations (e.g. pregnant women); b. Reporting interval and cumulative number of reports per High Level Term (HLT) and System Organ Class (SOC); c. Reporting interval and cumulative number of adverse event reports in Canada and globally stratified by seriousness, age groups, gender, dose and in special populations (e.g. pregnant women). Should specific demographic data not be available, Moderna Biopharma Canada Corporation shall document this and provide an explanation in the MSSR; d. Estimated exposure and use patterns including doses distributed/administered in the reporting interval/cumulative period, stratified by country, including any available data on age groups, race, ethnicity, indigenous populations and remote communities. Should specific demographic data not be available, Moderna Biopharma Canada Corporation shall document this and provide an explanation in the MSSR; e. Actions taken in the reporting interval for safety reasons and changes to reference safety information; f. Overview of new/ongoing/closed signals during the reporting interval, and discussion of proposed risk minimization measures (if applicable). Reviews of safety topics identified by Health Canada and/or foreign regulators; g. Updated list of Adverse Events of Special Interest (AESIs) from regulatory authorities, internationally recognized collaborations, and scientific literature, including the Safety Platform for Emergency vaccines (SPEAC) list. Summaries of reported cases of all AESIs and Risk Management Plan (RMP) safety concerns (including the additional missing information): numbers and relevant cases for the reporting interval/cumulative period including complete evaluation, time to-onset, observed/expected analyses, and causality assessment (if applicable);	November 3, 2022	Closed

	h. Fatal reports – reporting interval/cumulative number of reports, including observed/expected analyses and discussion of relevant cases; i. Vaccination errors should be included when a pattern of errors leading to safety issue and/or risk minimization activities are considered warranted (e.g. changes of the product labelling, communication to healthcare professional and the public). Otherwise, data can be presented and discussed in the Periodic Safety Update Reports (PSURs)/Periodic Benefit Risk Evaluation Reports (PBRERs); j. Summary outcomes of some of the routine pharmacovigilance activities (as presented in the EU RMP Part III and applied in the Canadian context) should be included for the purpose of rapid signal detection and communication activities. Summary of all ongoing registries and studies should be included in the six-month scheduled PSURs/PBRERs, unless a safety signal is identified that requires immediate regulatory action; k. Overall risk/benefit assessment.		
7	Moderna Biopharma Canada Corporation is required to submit Periodic Safety Update Reports (PSURs)/Periodic Benefit Risk Evaluation Reports (PBRERs) every 6 months for Spikevax Bivalent (Original / Omicron BA.4/5), unless otherwise determined by Health Canada. The core PSUR/PBRER format should follow international guidance for COVID-19 vaccines.	November 3, 2022	Closed
8	Moderna Biopharma Canada Corporation is required to submit an updated core RMP with the Canadian Addendum in a timely manner if a safety issue is identified that requires immediate regulatory action or as requested by Health Canada. The RMP format should follow the guidance (Guidance Document Submission of Risk Management Plans and Follow-Up Commitments) and should include the following: a. A safety specification that details the identified risks, potential risks, and missing information for Spikevax Bivalent (Original / Omicron BA.4/5); b. A pharmacovigilance plan that details specific measures to be taken to identify and report safety issues in COVID-19 patients, including adverse reaction reporting, periodic reporting, and ongoing/planned studies; and c. A risk minimization plan, if applicable, to manage risks that may require additional measures beyond those considered standard (for instance, labelling).	November 3, 2022	Ongoing
9	Moderna Biopharma Canada Corporation to submit final snapshots of all components of the electronic platform (linked to any foreign or Canadian specific labels),	November 3, 2022	Ongoing

	containing the approved Canadian-specific information for Spikevax Bivalent (original/omicron BA.4/5) in French and English for Health Canada's review and records, prior to launch of the electronic platform, and for each subsequent update.		
10	Moderna Biopharma Canada Corporation is requested to develop and distribute a Health Product Risk Communication (HPRC), in French and English, should a decision be made to import, for Canadian sites, non-Canadian labelled supplies. In this case, the HPRC would need to be developed with Health Canada approval and endorsement, to inform healthcare professionals that Moderna Biopharma Canada Corporation will delay implementation of Canadian-specific inner/outer labels following NDS-CV approval for Spikevax Bivalent (original/omicron BA.4/5), and that interim inner/outer labels will be used for the short term. Please note the following: a. Moderna Biopharma Canada Corporation should include images and texts of these labels in the HPRC and clearly outline all deviations from Canadian requirements; b. The HPRC should direct healthcare professionals to the electronic platform where they can find information about the approved Canadian-specific labelling in both official languages c. The HPRC should include an alternative method for the health care professionals to obtain a paper copy of the HPRC and/or Product Monograph (PM) by mail or fax from Moderna Biopharma Canada Corporation, if they cannot access the internet	November 3, 2022	Closed
11	Moderna Biopharma Canada Corporation to commit to developing Canadian specific bilingual labelling for Spikevax Bivalent (original/omicron BA.4/5) and implementing such labelling once supplies are transitioned to Canadian dedicated supplies. Health Canada should be kept informed of estimated timelines and proposed strategies concerning the development and implementation of Canadian-specific bilingual labels. a. During the period prior to implementation of the Canadian-specific bilingual labeling, Canadian reference labels should be made available to healthcare professionals.	November 3, 2022	Ongoing

12	Moderna Biopharma Canada Corporation is requested to develop and distribute a company-led customer communication, in English and French, should a decision be made to import non-Canadian labelled supplies for Canadian sites. The customer communication should aim to inform healthcare professionals that Moderna Biopharma Canada Corporation will delay the implementation of Canadian-specific inner/outer labels following NDS-CV approval for Spikevax Bivalent (Original / Omicron BA. 4/5) (elasomeran/devasomeran) mRNA vaccine, and that interim inner/outer labels will be used for the short term. Please note the following: a) Moderna Biopharma Canada Corporation should consider including images and texts of these labels in the customer communication and clearly outline all deviations from Canadian requirements; b) The customer communication should direct healthcare professionals to the electronic platform where they can find information about the approved Canadian-specific labelling in both official languages c) The customer communication should not resemble a Health Canada-endorsed Health Product Risk Communication and should not reflect a red banner at the top of the document. A company-led risk communication may take the form of a letter carrying the company letterhead, for example. The company-led customer communication does not require review by Health Canada; however, Moderna Biopharma Canada Corporation is welcome to seek courtesy feedback from BRDD. Moderna Biopharma Canada Corporation is expected to exercise due discretion to ensure prompt finalization and dissemination of the customer communication. Please provide a copy of the final, signed customer communication in English and French via eCTD once available.	April 4, 2023	Ongoing
13	Provide immunogenicity and safety data from Study mRNA-1273- P203 as soon as the data become available, including: a) Open label arm evaluating safety, reactogenicity and effectiveness of the BA.4/BA.5- containing bivalent vaccine in vaccine naïve aged 12 to 17 years, as part of a 2-dose primary series, with the doses administered 6 months apart.	May 18, 2023	Ongoing
14	Provide the final Clinical Study Report for parts of Study P203 mentioned above, when available.	May 18, 2023	Ongoing

15	Provide safety data from Study mRNA-1273- P204, as soon as the data become available, including: a) Booster Phase of Study P204 (ongoing SPIKEVAX monovalent booster study in participants aged 6 to 11 years). b) Open label arm evaluating the safety of the SPIKEVAX Bivalent (elasomeran/imelasomeran)/mRNA-1273.214 in participants aged 6 to 11 years who have not yet received a booster dose after their primary series with mRNA-1273.	May 18, 2023	Ongoing
16	Provide the final Clinical Study Report for parts of Study P204 mentioned above, when available.	May 18, 2023	Ongoing
17	Provide post-marketing safety data derived from Study P904, and Study P920 for pediatric age groups, as soon as the data become available.	May 18, 2023	Ongoing
18	Provide post-marketing effectiveness data derived from Study P901 for pediatric age groups, as soon as the data become available.	May 18, 2023	Ongoing
19	Provide immunogenicity data from Study P306 assessing mRNA-1273.214 booster dose in participants 6 months to < 6 years of age, as soon as the data become available.	May 18, 2023	Ongoing
20	Provide reactogenicity (solicited local and systemic ARs through 7 days after booster dose or third dose, and unsolicited AEs through 28 days after booster dose or third dose injection) and safety data from Study P306 assessing mRNA-1273.214 booster dose in participants 6 months to < 6 years of age, as soon as the data become available.	May 18, 2023	Ongoing
21	Moderna Biopharma Canada Corporation is required to submit Periodic Safety Update Reports (PSURs)/Periodic Benefit Risk Evaluation Reports (PBRERs) every 6 months, unless otherwise determined by Health Canada.	June 7, 2023	Ongoing