PUBLIC SUMMARY OF THE RISK MANAGEMENT PLAN

COMIRNATY AND COMIRNATY ORIGINAL/OMICRON BA.1 AND COMIRNATY ORIGINAL/OMICRON BA.4-5, COMIRNATY OMICRON XBB.1.5 (COVID-19 mRNA VACCINE)

Marketing Authorization Numbers:

68225 / 68710 / 69047/69127

Concentrate for dispersion for injection 30 micrograms/dose

Dispersion for Injection 30 micrograms/dose

Concentrate for dispersion for injection 10 micrograms/dose

Dispersion for Injection 15/15 micrograms/dose

Dispersion for injection 30 micrograms/dose

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LIST OF ABBREVIATIONS

Abbreviation	Definition of Term
COPD	chronic obstructive pulmonary disease
CoV	coronavirus
COVID-19	coronavirus disease 2019
EPAR	European public assessment report
EU	European Union
MERS	middle Est respiratory syndrome
mRNA	messenger ribonucleic acid
PSUR	Periodic safety update report
RMP	risk management plan
RNA	ribonucleic acid
SARS	severe acute respiratory syndrome
SmPC	summary of product characteristics
VAC4EU	Vaccine monitoring Collaboration for Europe
VAED	vaccine-associated enhanced disease
VAERD	vaccine-associated enhanced respiratory disease

OVERVIEW

The Risk Management Plan (RMP) is a comprehensive document submitted as part of the application dossier for market approval of a medicine. The RMP summary contains information on the medicine's safety profile and explains the measures that are taken in order to further investigate and follow the risks as well as to prevent or minimise them. The RMP summary for Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms), and of Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) is a concise document and does not claim to be exhaustive.

As the RMP is an international document, the summary might differ from the "Arzneimittelinformation / Information sur le médicament" approved and published in Switzerland, e.g., by mentioning risks occurring in populations or indications not included in the Swiss marketing authorization.

Please note that the reference document which is valid and relevant for the effective and safe use of Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms) and of Comirnaty Original/Omicron BA.4-5 (15/15 micrograms), Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) in Switzerland is the "Arzneimittelinformation / Information sur le médicament" (see www.swissmedic.ch) approved and authorised by Swissmedic. Pfizer is fully responsible for the accuracy and correctness of the content of the published RMP summary of Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and of Comirnaty Omicron XBB.1.5 (30 micrograms)/dose).

Summary of risk management plan for Comirnaty, Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5 (30 micrograms)/dose)**.**

This is a summary of the risk management plan (RMP) for Comirnaty, for Comirnaty Original/Omicron BA.1 (15/15 micrograms) and for Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5. The RMP details important risks of Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms) and of Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5 (30 micrograms)/dose), how these risks can be minimised, and how more information will be obtained about Comirnaty's, Comirnaty Original/Omicron BA.1 (15/15 micrograms) and Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) risks and uncertainties (missing information).

Comirnaty, Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) summary of product characteristics (SmPC) and its package leaflet give essential information to healthcare professionals and patients on how Comirnaty, Comirnaty Original/Omicron BA.1 (15/15 micrograms) and Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) should be used.

This summary of the RMP for Comirnaty, for Comirnaty Original/Omicron BA.1 (15/15 micrograms), Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) should be read in the context of all this information including the assessment report of the evaluation and its plain-language summary, all which is part of the European Public Assessment Report (EPAR).

Important new concerns or changes to the current ones will be included in updates of Comirnaty's, Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms), Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) RMP.

I. The Medicine and What It Is Used For

Comirnaty is a vaccine for active immunisation to prevent COVID-19 caused by SARS-CoV-2 virus, in individuals 5 years of age and older. Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5 (30 micrograms/dose dispersion for injection are indicated for active immunisation to prevent COVID-19 caused by SARS-CoV-2 virus, in individuals 12 years of age and older who have previously received at least a primary vaccination course against COVID-19 (see SmPC for the full indication). Both contain nucleoside-modified messenger RNA encapsulated in lipid nanoparticles as the active substance and are given intramuscularly.

Further information about the evaluation of Comirnaty's, of Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms), and of Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) benefits can be found in Comirnaty's, Comirnaty Original/Omicron BA.1 (15/15 micrograms), Comirnaty Original/Omicron BA.4-5 (15/15 micrograms), and Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) EPAR, including in its plain-language summary, available on the EMA website, under the medicine's webpage www.ema.europa.eu/en/medicines/human/EPAR/comirnaty.

II. Risks Associated with the Medicine and Activities to Minimise or Further Characterise the Risks

Important risks of Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms), and of Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) together with measures to minimise such risks and the proposed studies for learning more about Comirnaty's, Comirnaty Original/Omicron BA.1 (15/15 micrograms), Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) risks, are outlined below.

Measures to minimise the risks identified for medicinal products can be:

- Specific Information, such as warnings, precautions, and advice on correct use, in the package leaflet and SmPC addressed to patients and healthcare professionals;
- Important advice on the medicine's packaging;
- The authorised pack size — the amount of medicine in a pack is chosen so to ensure that the medicine is used correctly;
- The medicine's legal status — the way a medicine is supplied to the patient (e.g. with or without prescription) can help to minimise its risks.

Together, these measures constitute *routine risk minimisation* measures.

In addition to these measures, information about adverse events is collected continuously and regularly analysed, including PSUR assessment so that immediate action can be taken as necessary. These measures constitute *routine pharmacovigilance activities*.

If important information that may affect the safe use of Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and of Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) is not yet available, it is listed under 'missing information' below.

II.A. List of Important Risks and Missing Information

Important risks of Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and of Comirnaty Omicron XBB.1.5 (30 micrograms)/dose) are risks that need special risk management activities to further investigate or minimise the risk, so that the medicinal product can be safely administered. Important risks can be regarded as identified or potential. Identified risks are concerns for which there is sufficient proof of a link with the use of Comirnaty, of Comirnaty Original/Omicron BA.1 (15/15 micrograms),Comirnaty Original/Omicron BA.4-5 (15/15 micrograms) and Comirnaty Omicron XBB.1.5 (30 micrograms)/dose). Potential risks are concerns for which an association with the use of this medicine is possible based on available data, but this association has not been established yet and needs further evaluation. Missing information refers to information on the safety of the medicinal product that is currently missing and needs to be collected (e.g., on the long-term use of the medicine).

Table 1. List of Important Risks and Missing Information

Important identified risks	Myocarditis and Pericarditis
Important potential risks	None
Missing information	Use in pregnancy and while breast feeding
	Use in immunocompromised patients
	Use in frail patients with co-morbidities (e.g. chronic obstructive pulmonary disease [COPD], diabetes, chronic neurological disease, cardiovascular disorders)
	Use in patients with autoimmune or inflammatory disorders
	Interaction with other vaccines
	Long term safety data

II.B. Summary of Important Risks

The safety information in the Product Information is aligned to the reference.

Table 2. Important Identified Risk: Myocarditis and Pericarditis

Evidence for linking the risk to the medicine	Events of Myocarditis and Pericarditis have been reported.
Risk factors and risk groups	Post-authorization reports have been reported more frequently in adolescent and young adult male patients following the second dose of vaccine; however, reports have been received for adult males and females of broader age range and following the first vaccination also.
Risk minimisation measures	Routine risk minimisation measures SPC sections Warnings and Precautions and Undesirable Effects Additional risk minimisation measures: Direct Healthcare Professional Communication (DHPC) letter and communication plan
Additional pharmacovigilance activities	• C4591009 • C4591011 • C4591012 • C4591021 (former ACCESS/VAC4EU) • C4591038 (former C4591021 sub-study) • C4591036 (former Pediatric Heart Network study) • C4591051 • C4591052 See section II.C of this summary for an overview of the post-authorisation development plan.

Table 3. Missing Information: Use in Pregnancy and while Breast Feeding

Risk minimisation measures	Routine risk minimisation measures: SPC section Pregnancy, Lactation Additional risk minimisation measures: Noadditional risk minimisation measures.
Additional pharmacovigilance activities	• C4591009[a] • C4591011[a] • C4591015 • C4591021 (former ACCESS/VAC4EU)[a] • C4591022[a] • C4591051[a] • C4591052[a] See section II.C of this summary for an overview of the post-authorisation development plan.

a. Please note that studies C4591009, C4591011, C4591021 (former ACCESS/VAC4EU) C4591022, C4591051 and C4591052 address only "Use in pregnancy" and not "Breast feeding".

Table 4. Missing Information: Use in Immunocompromised Patients

Risk minimisation measures	Routine risk minimisation measures: SPC sections Warnings and Precautions Additional risk minimisation measures: No additional risk minimisation measures.
Additional pharmacovigilance activities	• C4591009[a] • C4591011 • C4591012 • C4591021 (former ACCESS/VAC4EU) • C4591024 (former Safety and Immunogenicity in high risk adults) • C4591051 • C4591052 See section II.C of this summary for an overview of the post-authorisation development plan.

a. Addresses AESI-based safety events of interest

Table 5. Missing Information: Use in Frail Patients with Co-morbidities (e.g. chronic obstructive pulmonary disease (COPD), diabetes, chronic neurological disease, cardiovascular disorders)

Risk minimisation measures	Routine risk minimisation measures: SPC section Properties/Effects Additional risk minimisation measures: No additional risk minimisation measures.
Additional pharmacovigilance activities	• C4591011 • C4591012 • C4591021 (former ACCESS/VAC4EU) C4591024 (former Safety and immunogenicity in high risk adults) • C4591052 See section II.C of this summary for an overview of the post-authorisation development plan.

Table 6. Missing Information: Use in Patients with Autoimmune or Inflammatory Disorders

Risk minimisation measures	Routine risk minimisation measures: None. Additional risk minimisation measures: No additional risk minimisation measures.
Additional pharmacovigilance activities	• C4591011 • C4591012 • C4591021 (former ACCESS/VAC4EU) • C4591024 (former Safety and immunogenicity in high risk adults) • C4591052 See section II.C of this summary for an overview of the post-authorisation development plan.

Table 7. Missing Information: Interaction with other Vaccines

Risk minimisation measures	Routine risk minimisation measures: SPC section Interactions Additional risk minimisation measures: No additional risk minimisation measures.
Additional pharmacovigilance activities	• C4591030 (Co-administration study with seasonal influenza vaccine) See section II.C of this summary for an overview of the post-authorisation development plan.

Table 8. Missing Information: Long Term Safety Data

Risk minimisation measures	Routine risk minimisation measures: None. Additional risk minimisation measures: No additional risk minimisation measures.
Additional pharmacovigilance activities	• C4591007 • C4591009 • C4591011 • C4591012 • C4591021 (former ACCESS/VAC4EU) • C4591038 (former C4591021 substudy) • C4591036 (former PHN) • C4591051 • C4591052 See section II.C of this summary for an overview of the post-authorisation development plan.

II.C. Post-Authorisation Development Plan

II.C.1. Studies which are Conditions of the Marketing Authorisation

None

II.C.2. Other Studies in Post-Authorisation Development Plan

Study	Purpose of the study
C4591007	To assess the safety, tolerability, immunogenicity, and efficacy of the BNT162b2 RNA-based COVID-19 vaccine candidate against COVID-19 in healthy paediatric subjects.
C4591009	To assess the occurrence of safety events of interest, including myocarditis and pericarditis, in the general US population (all ages), pregnant women, the immunocompromised and persons with a prior history of COVID-19 within selected data sources participating in the US Sentinel System.
C4591011	To assess whether individuals (all ages) in the US DoD MHS experience increased risk of safety events of interest, following receipt of the COVID-19 mRNA vaccine.
C4591012	To assess whether individuals in the US Veteran's Affairs Health System experience increased risk of safety events of interest, following receipt of the COVID-19 mRNA vaccine including the bivalent Omicron modified vaccine.
C4591015	To assess safety and immunogenicity in pregnant women In addition, exploratory objectives include: (a) To describe the immune response in infants born to breastfeeding maternal participants vaccinated with prophylactic COVID-19 mRNA vaccine during pregnancy. (b) To describe the safety of maternal immunisation in infants born to breastfeeding maternal participants vaccinated with prophylactic COVID-19 mRNA vaccine during pregnancy.
C4591014	To estimate the effectiveness of COVID-19 mRNA vaccine against hospitalisation and emergency department admission for acute respiratory illness due to SARS-CoV-2 infection and to assess the effectiveness of bivalent Omicron-modified vaccines following their introduction in all authorized age groups.
WI255886	To estimate the effectiveness of COVID-19 mRNA vaccine against hospitalisation for acute respiratory illness due to SARS-CoV-2 infection and to assess the effectiveness of bivalent Omicron-modified vaccines following their introduction in individuals 18 years of age and older.
C4591024 (former Safety and immunogenicity in high-risk adults)	Safety, tolerability and immunogenicity based on representative medical conditions (≥18 years: NSCLC, CLL, in hemodialysis for end-stage renal disease).
C4591021 (former ACCESS/ VAC4EU)	Assessment of potential increased risk of adverse events of special interest (AESI) among individuals (all ages) after being vaccinated with COVID-19 mRNA vaccine, including individuals less than 12 years of age. Estimating the time trend, in relation to DHPC letter dissemination, of the proportion of individuals who received real-world clinical assessments for myocarditis/pericarditis following Comirnaty vaccination.
C4591038 (former C4591021 substudy)	To describe clinical course (treatment, survival, hospitalisations, long-term cardiac outcomes) of myocarditis and pericarditis among individuals diagnosed with myocarditis and/or pericarditis after receiving at least 1 dose of the Pfizer-BioNTech COVID-19 vaccine and among individuals diagnosed with myocarditis and/or pericarditis who had no prior COVID-19 vaccination, using a cohort. study.
C4591022	To assess whether pregnant women receiving BNT162b2 experience increased risk of pregnancy and infant safety outcomes, including major congenital malformations, spontaneous abortion, stillbirth, preterm delivery, small for gestational age, and small for age postnatal growth to one year of age relative to pregnant women who received no COVID-19 vaccines during pregnancy.
C4591036 (former Pediatric Heart Network study)	To characterize the clinical course, risk factors, long-term sequelae, and quality of life in children and young adults <21 years with acute post-vaccine myocarditis including myocarditis after the bivalent Omicron modified vaccine.

Study	Purpose of the study
C4591030 (Co-administration study with seasonal influenza vaccine)	Safety and immunogenicity of COVID-19 mRNA vaccine and quadrivalent seasonal influenza vaccine when administered separately or concomitantly.
C4591031 Substudy E	To describe the safety and tolerability profile of BNT162b2 (30 µg or 60 µg), BNT162b2 OMI (30 µg or 60 µg), and bivalent BNT162b2 and BNT162b2 OMI (30 µg or 60 µg) given as a fourth dose to BNT162b2 experienced participants >55 years of age and experienced participants 18-to 55 years of age
C4591044	To describe the safety/tolerability and immune response to BNT162b5 Bivalent and BNT162b2 Bivalents given as a 2nd booster dose to COVID-19-vaccine-experienced participants ≥12 years of age.
C4591048	To investigate the safety, tolerability, and immunogenicity of bivalent BNT162b2 RNA-based vaccine candidate(s) in healthy children.
C4591051	To ensure comprehensive understanding of real-world safety of the Pfizer-BioNTech COVID-19 bivalent Omicron-modified vaccine in large samples of general US populations.
C4591052	To ensure comprehensive understanding of real-world safety of the Pfizer-BioNTech COVID-19 bivalent Omicron-modified vaccine in large samples of general EU populations.