



EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

COMIRNATY

Procedural steps taken and scientific information after the authorisation

Application number	Scope	Opinion/ Notification ¹ issued on	Commission Decision Issued ² / amended on	Product Information affected ³	Summary
IB/0142/G	This was an application for a group of variations. B.II.z - Quality change - Finished product - Other variation B.II.f.1.b.5 - Stability of FP - Extension of the shelf life of the finished product - Biological/immunological	01/08/2022	05/08/2022	SmPC	

¹ Notifications are issued for type I variations and Article 61(3) notifications (unless part of a group including a type II variation or extension application or a worksharing application). Opinions are issued for all other procedures.

² A Commission decision (CD) is issued for procedures that affect the terms of the marketing authorisation (e.g. summary of product characteristics, annex II, labelling, package leaflet). The CD is issued within two months of the opinion for variations falling under the scope of Article 23.1a(a) of Regulation (EU) No. 712/2012, or within one year for other procedures.

³ SmPC (Summary of Product Characteristics), Annex II, Labelling, PL (Package Leaflet).



	<p>medicinal product in accordance with an approved stability protocol</p> <p>B.II.f.1.e - Stability of FP - Change to an approved stability protocol</p>				
II/0136	B.II.f.1.z - Stability of FP - Change in the shelf-life or storage conditions of the finished product - Other variation	21/07/2022	05/08/2022	SmPC and PL	The SmPC section 6.1 has been updated as follows: "Within the 1 month shelf life at 2 °C to 8 °C, up to 48 hours may be used for transportation."
II/0135/G	<p>This was an application for a group of variations.</p> <p>B.I.a.3.c - Change in batch size (including batch size ranges) of AS or intermediate - The change requires assessment of the comparability of a biological/immunological AS</p> <p>B.I.a.2.a - Changes in the manufacturing process of the AS - Minor change in the manufacturing process of the AS</p> <p>B.I.z - Quality change - Active substance - Other variation</p> <p>B.I.z - Quality change - Active substance - Other variation</p> <p>B.I.z - Quality change - Active substance - Other variation</p>	21/07/2022	n/a		
II/0134/G	<p>This was an application for a group of variations.</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.7.b - Change in supplier of packaging</p>	14/07/2022	n/a		

	<p>components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.z - Quality change - Finished product - Other variation</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p>				
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PSUSA/10898 /202112	Periodic Safety Update EU Single assessment - covid-19 mRNA vaccine (nucleoside-modified) (COMIRNATY)	07/07/2022	n/a		PRAC Recommendation - maintenance
II/0131	B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method	23/06/2022	n/a		
IAIN/0133/G	This was an application for a group of variations. B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR	15/06/2022	n/a		
II/0124/G	This was an application for a group of variations. B.II.f.z - Stability of FP - Other variation B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing	10/06/2022	n/a		

	processes				
II/0130/G	<p>This was an application for a group of variations.</p> <p>B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP -</p>	02/06/2022	n/a		

	<p>Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.4.c - Change in the batch size (including batch size ranges) of the finished product - The change requires assessment of the comparability of a biological/immunological medicinal product or a new bioequivalence study</p> <p>B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site</p> <p>B.II.b.3.b - Change in the manufacturing process of the finished or intermediate product - Substantial changes to a manufacturing process that may have a significant impact on the quality, safety and efficacy of the medicinal product</p> <p>B.II.z - Quality change - Finished product - Other variation</p> <p>B.II.b.3.b - Change in the manufacturing process of the finished or intermediate product - Substantial changes to a manufacturing process that may have a significant impact on the quality, safety and efficacy of the medicinal product</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP -</p>				
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<p>Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.z - Quality change - Finished product - Other variation</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p>				
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	<p>B.II.d.2.d - Change in test procedure for the finished product - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p>				
IB/0127	B.II.c.z - Change in control of excipients in the Finished Product - Other variation	25/05/2022	n/a		
IA/0128/G	<p>This was an application for a group of variations.</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p>	12/05/2022	n/a		
II/0125/G	<p>This was an application for a group of variations.</p> <p>B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product</p> <p>B.I.z - Quality change - Active substance - Other</p>	12/05/2022	n/a		

	variation				
N/0113	Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification)	11/05/2022	n/a		
II/0104	Update of sections 4.2, 4.8 and 5.1 of the SmPC of Comirnaty 30 microgram/dose and section 4.8 of the SmPC of Comirnaty 10 microgram/dose in order to update immunogenicity, safety and efficacy information regarding heterologous booster vaccination based on published literature data; the Package Leaflet is updated accordingly. C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data	22/04/2022	04/05/2022	SmPC and PL	Please refer to Scientific Discussion 'Comirnaty EMEA/H/C/005735/II/0104'.
II/0126/G	This was an application for a group of variations. B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes	26/04/2022	n/a		
IB/0123	B.I.z - Quality change - Active substance - Other	19/04/2022	n/a		

	variation				
II/0120/G	<p>This was an application for a group of variations.</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p>	07/04/2022	n/a		
IB/0122	B.II.f.1.b.5 - Stability of FP - Extension of the shelf life of the finished product - Biological/immunological medicinal product in accordance with an approved stability protocol	04/04/2022	07/04/2022	SmPC and PL	Update of the product information to extend the shelf life of COMIRNATY 30 µg Concentrate for dispersion for injection (EU/1/20/1528/001) from 9 to 12 months.

II/0116/G	<p>This was an application for a group of variations.</p> <p>B.I.z - Quality change - Active substance - Other variation</p> <p>B.I.z - Quality change - Active substance - Other variation</p> <p>B.I.a.2.a - Changes in the manufacturing process of the AS - Minor change in the manufacturing process of the AS</p> <p>B.I.a.2.a - Changes in the manufacturing process of the AS - Minor change in the manufacturing process of the AS</p> <p>B.I.d.1.c - Stability of AS - Change in the re-test period/storage period or storage conditions - Change to an approved stability protocol</p> <p>B.I.a.1.j - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Replacement or addition of a site where batch control/testing takes place and any of the test method at the site is a biol/immunol method</p> <p>B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product</p>	31/03/2022	07/04/2022	Annex II	
IB/0106/G	<p>This was an application for a group of variations.</p> <p>B.I.b.1.c - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Addition of a new</p>	31/03/2022	n/a		

	<p>specification parameter to the specification with its corresponding test method</p> <p>B.I.b.2.c - Change in test procedure for AS or starting material/reagent/intermediate - Other changes to a test procedure for a reagent, which does not have a significant effect on the overall quality of the AS</p>				
II/0112/G	<p>This was an application for a group of variations.</p> <p>B.II.b.3.b - Change in the manufacturing process of the finished or intermediate product - Substantial changes to a manufacturing process that may have a significant impact on the quality, safety and efficacy of the medicinal product</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.4.c - Change in the batch size (including batch size ranges) of the finished product - The change requires assessment of the comparability of a biological/immunological medicinal product or a new bioequivalence study</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p>	24/03/2022	n/a		
II/0105	B.I.a.1.e - Change in the manufacturer of AS or of a	24/03/2022	n/a		

	starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product				
II/0121/G	<p>This was an application for a group of variations.</p> <p>B.II.f.1.b.5 - Stability of FP - Extension of the shelf life of the finished product - Biological/immunological medicinal product in accordance with an approved stability protocol</p> <p>B.II.f.1.c - Stability of FP - Change in storage conditions for biological medicinal products, when the stability studies have not been performed in accordance with an approved stability protocol</p>	23/03/2022	07/04/2022	SmPC and PL	
IB/0119/G	<p>This was an application for a group of variations.</p> <p>B.II.b.5.z - Change to in-process tests or limits applied during the manufacture of the finished product - Other variation</p> <p>B.II.b.5.z - Change to in-process tests or limits applied during the manufacture of the finished product - Other variation</p> <p>B.II.b.5.z - Change to in-process tests or limits applied during the manufacture of the finished product - Other variation</p>	22/03/2022	n/a		
II/0109/G	<p>This was an application for a group of variations.</p> <p>B.II.b.2.b - Change to importer, batch release</p>	10/03/2022	n/a		

	<p>arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p>				
II/0087	C.I.11.b - Introduction of, or change(s) to, the obligations and conditions of a marketing authorisation, including the RMP - Implementation of change(s) which require to be further substantiated by new additional data to be submitted by the MAH where significant assessment is required	10/03/2022	n/a		
II/0115/G	<p>This was an application for a group of variations.</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p>	03/03/2022	n/a		

B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes

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B.II.b.4.b - Change in the batch size (including batch size ranges) of the finished product - Downscaling down to 10-fold

B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process

B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier

B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change

	<p>in the manufacturing process</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p>				
IB/0114	<p>B.I.b.1.a - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Tightening of specification limits for medicinal products subject to OCABR</p>	03/03/2022	n/a		
IB/0117/G	<p>This was an application for a group of variations.</p> <p>B.III.2.z - Change to comply with Ph. Eur. or with a national pharmacopoeia of a Member State - Other variation</p> <p>A.7 - Administrative change - Deletion of manufacturing sites</p> <p>A.7 - Administrative change - Deletion of manufacturing sites</p>	02/03/2022	n/a		
II/0111	<p>Update of section 4.2 of the SmPC of Comirnaty 30 microgram/dose to lower the age of the booster dose from adults 18 years of age and older to adults and adolescents 12 years of age and older, based on real world evidence collected by the Ministry of Health of Israel. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to make</p>	24/02/2022	28/02/2022	SmPC and PL	<p>SmPC new text</p> <p>A booster dose of Comirnaty may be administered intramuscularly at least 6 months after the second dose in individuals 12 years of age and older.</p> <p>For more information, please refer to the Summary of Product Characteristics.</p>

	<p>minor editorial changes throughout the product information (SmPC and package leaflet).</p> <p>C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data</p>				
II/0093	<p>Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC of Comirnaty 30 microgram/dose and section 4.8 of the SmPC of Comirnaty 10 microgram/dose in order to update efficacy and safety information after booster dose and change posology recommendations for booster use from "individuals 18 years of age and older" to "individuals 16 years of age and older", based on interim results from study C4591031, this is a randomized, placebo-controlled, phase 3 booster efficacy study involving participants ≥ 16 years of age who completed the primary series of BNT162b2 30 µg in Study C4591001, published literature data and post-marketing safety data. The Package Leaflet and Labelling are updated accordingly.</p> <p>C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data</p>	24/02/2022	28/02/2022	SmPC, Labelling and PL	<p>SmPC new text</p> <p>A booster dose of Comirnaty may be administered intramuscularly at least 6 months after the second dose in individuals 16 years of age and older.</p> <p>The relative vaccine efficacy information for participants 16 years of age and older without prior evidence of SARS-CoV-2 infection was 95.3% (95% confidence interval of 89.5% to 98.3%). Relative vaccine efficacy in participants with or without evidence of prior SARS-CoV-2 infection was 94.6% (95% confidence interval of 88.5% to 97.9%), similar to that seen in those participants without evidence of prior infection.</p> <p>The overall safety profile for the booster dose was similar to that seen after 2 doses.</p> <p>For more information, please refer to the Summary of Product Characteristics.</p>
II/0056/G	<p>This was an application for a group of variations.</p> <p>C.I.11.b - Introduction of, or change(s) to, the obligations and conditions of a marketing authorisation, including the RMP - Implementation of</p>	17/02/2022	28/02/2022	Annex II	<p>To address specific obligations on quality, the MA Holder provided relevant data and proposed changes to the dossier, in line with the requirements and due dates set by CHMP.</p> <p>With regards to SO1, the MA Holder has provided additional</p>

<p>change(s) which require to be further substantiated by new additional data to be submitted by the MAH where significant assessment is required</p> <p>C.I.11.b - Introduction of, or change(s) to, the obligations and conditions of a marketing authorisation, including the RMP - Implementation of change(s) which require to be further substantiated by new additional data to be submitted by the MAH where significant assessment is required</p> <p>B.I.b.1.a - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.I.b.1.c - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters</p>				<p>characterisation data as requested.</p> <p>a) The potential for truncated transcripts to produce proteins/peptides was further investigated using a cell-free in vitro expression system. No truncated or other protein species were detected beyond the background bands observed in the negative control sample. The MAH will complement the characterization exercise using the cell-free in vitro translation system with additional tozinameran batches.</p> <p>b) It is sufficiently demonstrated that the major proportion of fragmented species contains the 5'-cap but lacks the poly(A) tail.</p> <p>c) WB results obtained by three different antibodies, specific for the S1 domain, the receptor binding domain and the S2 domain, respectively, were presented and compared to theoretical masses of the S-protein and the subdomains in glycosylated and non-glycosylated forms. It is sufficiently justified that the major band monitored corresponds to the heavily glycosylated S-protein. With regards to SO₂, the MA Holder provided additional information to enhance the control strategy, including the active substance and finished product specifications.</p> <p>a) For active substance the acceptance criterion for 5'-Cap has been tightened. For finished product acceptance criteria for osmolality, LNP size, RNA encapsulation, RNA content and lipids content (ALC-0315, ALC-0159, DSPC, cholesterol) and in vitro expression (IVE) assay have been tightened. The revised limits are sufficiently justified and found acceptable. The acceptance criteria for RNA integrity test in active substance and finished product specifications have been previously tightened during VAR IB-31-G.</p> <p>b) A new test method has been introduced and</p>
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	<p>and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.b - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.I.d.1.c - Stability of AS - Change in the re-test period/storage period or storage conditions - Change to an approved stability protocol</p> <p>B.I.z - Quality change - Active substance - Other variation</p>				<p>acceptably validated.</p> <p>c) The MAH previously presented data supporting the suitability of the ddPCR method, as being capable of detecting changes in the poly(A) tail content. The accuracy of the ddPCR method is sufficiently addressed.</p> <p>d) The acceptance criteria for RNA integrity test in active substance and finished product specifications have been previously tightened during VAR IB-31-G. The justifications for no further tightening of the finished product specifications for RNA integrity and LNP polydispersity in the current variation have been acceptably justified.</p> <p>e) Additional details have been provided to support the suitability of the IVE assay (cell-based flow cytometry) used for potency determination.</p> <p>f) Evaluation of lipid-related impurities in the finished product and discussion on ALC-0315 impact on late migrating species (LMS) and RNA integrity were provided and assessed via parallel VAR II-54-G. Based on the provided information, SO1 and SO2 are considered fulfilled. The provided data also fulfils recommendations on analytical methods (REC 10) and additional stability (REC 20). The Annex IIE has been updated as follows:</p> <ul style="list-style-type: none"> • SO1 (relating to characterisation of the active substance and finished product) is deleted from the list of specific obligations. • SO2 (relating to control strategy, including the active substance and finished product specifications) is deleted from the list of specific obligations.
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IB/0110	C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation	15/02/2022	28/02/2022	SmPC and PL	To update section 4.6 of the SmPC and section 2 of the PL to implement the recommendation on vaccination with Comirnaty in pregnant and breastfeeding women as requested by the CHMP.
II/0108/G	<p>This was an application for a group of variations.</p> <p>B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.2.z - Change in test procedure for an excipient - Other variation</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>	10/02/2022	n/a		

IB/0107/G	<p>This was an application for a group of variations.</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>	07/02/2022	n/a		
II/0102	<p>Update of sections 4.8 and 5.1 of the SmPC in order to update safety and efficacy information 6 months post-dose 2 follow-up for adolescence 12 through 15 years of age based on updated interim results from study C4591001, listed as a specific obligation; this is a Phase 1/2/3, placebo-controlled, observer-blind, interventional, dose-finding, study to evaluate the safety, tolerability, immunogenicity and efficacy of SARS-CoV-2 RNA vaccine candidates against COVID-19 in healthy individuals.</p> <p>C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data</p>	27/01/2022	04/02/2022	SmPC and PL	<p>Updated efficacy analyses were performed with additional confirmed COVID-19 cases accrued during blinded placebo-controlled follow-up, representing up to 6 months after Dose 2 in the efficacy population.</p> <p>In the updated efficacy analysis of Study 2 in adolescents 12 to 15 years of age without evidence of prior infection, there were no cases in 1,057 participants who received the vaccine and 28 cases out of 1,030 who received placebo. The point estimate for efficacy is 100% (95% confidence interval 86.8, 100.0). In participants with or without evidence of prior infection there were 0 cases in the 1,119 who received vaccine and 30 cases in 1,109 participants who received placebo. This also indicates the point estimate for efficacy is 100% (95% confidence interval 87.5, 100.0). For more information, please refer to the Summary of Product Characteristics.</p>
II/0092/G	<p>This was an application for a group of variations.</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a</p>	03/02/2022	n/a		

	<p>biol/immunol method</p> <p>B.I.a.1.j - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Replacement or addition of a site where batch control/testing takes place and any of the test method at the site is a biol/immunol method</p>				
II/0090	<p>B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product</p>	27/01/2022	n/a		
IB/0100	<p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>	24/01/2022	n/a		
II/0080	<p>Update of section 4.4 of the SmPC in order to amend an existing warning on anxiety-related reactions to add hypoaesthesia and paraesthesia and remove tingling sensations following the outcome of the Post-Authorisation Measure MEA-002.8 (EMA/H/C/005735/MEA/002.8, dated 30 September 2021).</p> <p>Update of section 4.8 of the SmPC in order to add hypoaesthesia and paraesthesia to the list of adverse drug reactions (ADRs) with frequency 'not known' following the outcome of the Post-Authorisation Measure MEA-002.8 (EMA/H/C/005735/MEA/002.8, dated 30 September 2021). The Package Leaflet is updated accordingly.</p>	13/01/2022	24/01/2022	SmPC	<p>Following the assessment of clinical trial, literature and post-marketing safety data, a total of 21,793 cases were identified of which approximately 75% occurred within the day after vaccination. When available (~15% of the cases), the duration of the event was reported to be ≤2 days in ~70% of the cases, which is considered a plausible time to onset (TTO).</p> <p>Hypoaesthesia and paraesthesia are not sufficiently covered in the product information by the currently included text regarding stress-related reactions. Although approximately 60% of the cases co-reported symptoms associated with an acute stress response/vasovagal episode, the remaining cases were not observed in the context of stress-related reactions (n = 9446). In these cases, co-reported events included reactogenicity adverse drug reactions (ADRs)</p>

	C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data				listed in the product information. Thus, the existing warning on anxiety-related reactions in the SmPC section 4.4 has been amended to remove tingling sensations and add paraesthesia and hypoaesthesia. These ADRs (paraesthesia and hypoaesthesia) have also been added to the list of adverse drug reactions with frequency 'not known' in the SmPC section 4.8.
II/0101	B.II.f.1.c - Stability of FP - Change in storage conditions for biological medicinal products, when the stability studies have not been performed in accordance with an approved stability protocol	21/01/2022	04/02/2022	SmPC and PL	The SmPC section 6.3 of the 30 micrograms/dose and 10 micrograms/dose Tris-sucrose formulations (EU/1/20/1528/002-005) has been updated to remove the possibility of distribution at -25°C to -15°C. Distribution is performed under frozen conditions at -90°C to -60°C.
II/0075/G	This was an application for a group of variations. B.II.d.2.c - Change in test procedure for the finished product - Substantial change to or replacement of a biol/immunol/immunochemical test method or a method using a biol. reagent or replacement of a biol. reference preparation not covered by an approved protocol B.I.b.2.a - Change in test procedure for AS or starting material/reagent/intermediate - Minor changes to an approved test procedure	13/01/2022	n/a		
PSUSA/10898/202106	Periodic Safety Update EU Single assessment - covid-19 mRNA vaccine (nucleoside-modified) (COMIRNATY)	13/01/2022	n/a		PRAC Recommendation - maintenance
IA/0103	B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the	10/01/2022	n/a		

	dossier) - Replacement or addition of a supplier				
IB/0099/G	This was an application for a group of variations. B.II.b.5.z - Change to in-process tests or limits applied during the manufacture of the finished product - Other variation B.II.b.3.z - Change in the manufacturing process of the finished or intermediate product - Other variation	10/01/2022	n/a		
IB/0098	B.I.c.1.z - Change in immediate packaging of the AS - Other variation	10/01/2022	n/a		
IB/0096/G	This was an application for a group of variations. B.II.c.z - Change in control of excipients in the Finished Product - Other variation B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure	04/01/2022	n/a		
II/0097	B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes	21/12/2021	n/a		
II/0089	B.I.a.3.c - Change in batch size (including batch size	16/12/2021	n/a		

	ranges) of AS or intermediate - The change requires assessment of the comparability of a biological/immunological AS				
II/0054/G	<p>This was an application for a group of variations.</p> <p>B.II.z - Quality change - Finished product - Other variation</p> <p>A.7 - Administrative change - Deletion of manufacturing sites</p> <p>B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure</p> <p>B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure</p> <p>B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure</p> <p>B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.c.1.a - Change in the specification parameters and/or limits of an excipient - Tightening of specification limits</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new</p>	16/12/2021	24/01/2022	Annex II	<p>To address specific obligations SO4 and SO5, a reassessment of the ALC-0315 and ALC-0159 lipid control strategies was performed including impurity understanding and control, specifications for starting materials, intermediates, and the final ALC-0315 and ALC-0159 lipids and supporting data for method validation and stability data were provided. Evaluation of lipid-related impurities in the drug product and discussion on ALC-0315 impact on late migrating species (LMS) and RNA integrity were provided in the response to SO2 (f). Based on the provided information, SO2 (f), SO4 and SO5 are considered fulfilled. The Annex IIE has been updated as follows:</p> <ul style="list-style-type: none"> • SO4 relating to excipient ALC-0315 is deleted from the list of specific obligations. • SO5 relating to excipient ALC-0159 is deleted from the list of specific obligations.

	<p>specification parameter to the specification with its corresponding test method</p> <p>B.II.c.1.a - Change in the specification parameters and/or limits of an excipient - Tightening of specification limits</p> <p>C.I.11.b - Introduction of, or change(s) to, the obligations and conditions of a marketing authorisation, including the RMP - Implementation of change(s) which require to be further substantiated by new additional data to be submitted by the MAH where significant assessment is required</p>				
IB/0094/G	<p>This was an application for a group of variations.</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p>	13/12/2021	n/a		
IB/0091	B.II.f.1.b.5 - Stability of FP - Extension of the shelf life of the finished product - Biological/immunological medicinal product in accordance with an approved stability protocol	09/12/2021	09/12/2021	SmPC and PL	Update of Product Information to extend the shelf life from 6 to 9 months.
IAIN/0095	C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation	07/12/2021	07/12/2021	SmPC and PL	To update sections 4.4 and 4.8 of the SmPC and sections 2 and 4 of the PL to implement the signal recommendations

					on 'Signal assessment report on Myocarditis, pericarditis with Tozinameran (COVID-19 mRNA vaccine (nucleoside-modified) - COMIRNATY)
II/0086	B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product	26/11/2021	n/a		
X/0077	Annex I_2.(c) Change or addition of a new strength/potency	25/11/2021	26/11/2021	SmPC, Labelling and PL	
IB/0088/G	This was an application for a group of variations. B.II.b.4.f - Change in the batch size (including batch size ranges) of the finished product - The scale for a biological/immunological medicinal product is increased/decreased without process change (e.g. duplication of line) B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place	23/11/2021	n/a		
IB/0085/G	This was an application for a group of variations. B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process B.II.b.4.f - Change in the batch size (including batch	19/11/2021	n/a		

	<p>size ranges) of the finished product - The scale for a biological/immunological medicinal product is increased/decreased without process change (e.g. duplication of line)</p> <p>B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p>				
IB/0084/G	<p>This was an application for a group of variations.</p> <p>B.II.z - Quality change - Finished product - Other variation</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.5.b - Change to in-process tests or limits applied during the manufacture of the finished product - Addition of a new test(s) and limits</p>	19/11/2021	n/a		
II/0083/G	<p>This was an application for a group of variations.</p> <p>B.II.e.7.b - Change in supplier of packaging</p>	17/11/2021	n/a		

	components or devices (when mentioned in the dossier) - Replacement or addition of a supplier B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes				
IAIN/0082	C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation	16/11/2021	18/11/2021	SmPC and PL	To update section 4.8 of the SmPC and section 4 of the PL to implement the signal recommendation on erythema multiforme.
IB/0081	B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process	08/11/2021	n/a		
N/0079	Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification)	03/11/2021	18/11/2021	PL	To update section 2 of the PL regarding the wording on immunity of the third dose.
R/0046	Renewal of the marketing authorisation.	14/10/2021	03/11/2021		Please refer to Scientific Discussion 'EMEA/H/C/005735/R/0046`
X/0044/G	This was an application for a group of variations. Extension application to add new pharmaceutical form (dispersion for injection) with a new strength (0.1mg/ml). Update sections 6.4, 6.5 and 6.6 of the SmPC, section 5, 6 and information for healthcare professionals of the PL, section 1 of the Carton Box	14/10/2021	03/11/2021	SmPC, Labelling and PL	Please refer to Scientific Discussion 'EMEA/H/C/005735/X/0044`

	<p>Label as well as section 1 and 5 of the Vial Label to ensure the correct handling by providing dose verification information about strength, age range, colour information of the flip-off plastic cap and greyscale images.</p> <p>Update sections 4.2 of the SmPC ensure the correct handling in accordance to interchangeability of the medicinal product.</p> <p>Update section 8 of Carton Box Label to clarify expiry date "EXP" by adding storage temperature "(at -90°C to -60°C)" to ensure the correct handling of the medicinal product.</p> <p>Change the name of the active substance from COVID-19 mRNA Vaccine (nucleoside modified) to Tozinameran.</p> <p>Annex I_2.(c) Change or addition of a new strength/potency</p> <p>Annex I_2.(d) Change or addition of a new pharmaceutical form</p> <p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p> <p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p> <p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p> <p>A.3 - Administrative change - Change in name of the AS or of an excipient</p>				
II/0078/G	This was an application for a group of variations.	29/10/2021	n/a		

	<p>B.II.z - Quality change - Finished product - Other variation</p> <p>B.II.e.4.b - Change in shape or dimensions of the container or closure (immediate packaging) - The change in shape or dimensions concerns a fundamental part, which may have a significant impact on the delivery, use, safety or stability of the FP</p>				
II/0069/G	<p>This was an application for a group of variations.</p> <p>Grouped variation: Type II, B.II.a.3.b.3, Changes in the composition of the finished product to add sodium hydroxide and hydrochloric acid as pH adjustment excipients (present in the purchased 1500 mM PBS buffer, pH 6.81). Type IB, B.II.b.3.a, Minor changes in the manufacturing process of the finished product to introduce the use of an in-line dilution skid to prepare 50 mM citrate and 150 mM PBS buffers.</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process B.II.a.3.b.3 - Changes in the composition (excipients) of the finished product - Other excipients - Change that relates to a biological/immunological product</p>	28/10/2021	18/11/2021	SmPC, Labelling and PL	The SmPC section 6.1 (EU/1/20/1528/001) has been updated to include the following excipients: Sodium hydroxide (for pH-adjustment) Hydrochloric acid (for pH-adjustment) The Labelling and PL have been updated accordingly.
IB/0076	B.II.g.5.c - Implementation of changes foreseen in	18/10/2021	n/a		

	an approved change management protocol - For a biological/immunological medicinal product				
IB/0074/G	This was an application for a group of variations. B.II.c.z - Change in control of excipients in the Finished Product - Other variation B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)	18/10/2021	n/a		
N/0051	Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification)	14/10/2021	n/a		
II/0072/G	This was an application for a group of variations. B.II.c.z - Change in control of excipients in the Finished Product - Other variation B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change B.II.z - Quality change - Finished product - Other variation	11/10/2021	n/a		
II/0073/G	This was an application for a group of variations. B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing	07/10/2021	n/a		

	<p>processes</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p>				
II/0071	B.II.b.3.c - Change in the manufacturing process of the finished or intermediate product - The product is a biological/immunological medicinal product and the change requires an assessment of comparability	06/10/2021	n/a		
IB/0068/G	This was an application for a group of variations.	04/10/2021	n/a		

	<p>B.II.f.1.e - Stability of FP - Change to an approved stability protocol</p> <p>B.II.b.5.b - Change to in-process tests or limits applied during the manufacture of the finished product - Addition of a new test(s) and limits</p> <p>B.II.b.5.b - Change to in-process tests or limits applied during the manufacture of the finished product - Addition of a new test(s) and limits</p> <p>B.II.b.5.b - Change to in-process tests or limits applied during the manufacture of the finished product - Addition of a new test(s) and limits</p> <p>B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p> <p>B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p>				
II/0067	<p>Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to introduce a booster dose (third dose) of Comirnaty for individuals 18 years of age and older, based on interim safety and immunogenicity data from the interventional study C4591001, " A Phase 1/2/3, placebo-controlled, randomized, observer-blind, dose-finding study to evaluate the safety, tolerability, immunogenicity, and efficacy of SARS-CoV-2 RNA vaccine candidates against COVID-19 in healthy individuals". The package leaflet is updated accordingly.</p> <p>C.I.4 - Change(s) in the SPC, Labelling or PL due to</p>	04/10/2021	05/10/2021	SmPC and PL	Please refer to Scientific Discussion 'EMEA/H/C/005735/II/0067`

	new quality, preclinical, clinical or pharmacovigilance data				
II/0062	Update of sections 4.2 and 4.4 of the SmPC in order to introduce a third dose of Comirnaty for individuals 12 years of age and older who are severely immunocompromised, based on published literature data; the Package Leaflet is updated accordingly. C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data	04/10/2021	05/10/2021	SmPC and PL	Please refer to Scientific Discussion 'EMA/H/C/005735/II/0062'
II/0059	C.I.11.b - Introduction of, or change(s) to, the obligations and conditions of a marketing authorisation, including the RMP - Implementation of change(s) which require to be further substantiated by new additional data to be submitted by the MAH where significant assessment is required	30/09/2021	n/a		
IB/0070	B.I.b.z - Change in control of the AS - Other variation	24/09/2021	n/a		
II/0036	C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data	16/09/2021	23/09/2021	SmPC and PL	This type II variation updates the product information with updated data (March 2021) from the pivotal study C4591001 with the MAH having conducted an interim analysis containing immunogenicity, efficacy and safety data. The presented efficacy data support the previous conclusions in the initial marketing authorisation approval, and all provided subgroup analyses are in agreement with

				<p>previous data. A slightly lower point estimate of efficacy (92%) compared to previous analyses was obtained (95%). In addition, the reported interim data in this interim analysis, including reactogenicity and reported adverse events, supports the safety profile of Comirnaty 30µg in subjects ≥16 years of age obtained at the initial authorisation. Based on the safety data provided in this interim analysis, hyperhidrosis, night sweats, decreased appetite, asthenia and lethargy was added with a frequency uncommon to the list of adverse reactions in section 4.8 of the SmPC.</p> <p>For more information, please refer to the Summary of Product Characteristics.</p>
II/0047/G	<p>This was an application for a group of variations.</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.b.5.d - Change to in-process tests or limits applied during the manufacture of the finished product - Deletion of an in-process test which may have a significant effect on the overall quality of the finished product</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of</p>	16/09/2021	n/a	

	the finished or intermediate product - Minor change in the manufacturing process				
II/0066	B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product	13/09/2021	n/a		
IB/0061	B.II.f.1.b.5 - Stability of FP - Extension of the shelf life of the finished product - Biological/immunological medicinal product in accordance with an approved stability protocol	10/09/2021	23/09/2021	SmPC and PL	<ul style="list-style-type: none"> - Update section 6.3 of the SmPC to change the shelf life from 6 months to 9 months when stored at the intended storage conditions of -90 to -60 °C - implement Annex IV according to revised CHMP Opinion dated 22 July 2021 for procedure EMEA/H/C/005735/II/0030 - amend a mistake in section 4 of the PIL as requested in procedure PAM-MEA-002.6 to indicate that swelling of the face may occur in patients who have had facial dermatological fillers instead of facial cosmetic injections - amend section 4.4 of the SmPC for French and Latvian translations according to the agreed wording with the respective national authorities
II/0065/G	<p>This was an application for a group of variations.</p> <p>B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place</p> <p>B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site</p>	09/09/2021	n/a		

	B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes				
II/0057	B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product	09/09/2021	n/a		
II/0063/G	<p>This was an application for a group of variations.</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place</p> <p>B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site</p> <p>B.II.d.2.a - Change in test procedure for the finished</p>	02/09/2021	n/a		

	<p>product - Minor changes to an approved test procedure</p> <p>B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place</p> <p>B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place</p>				
II/0052/G	<p>This was an application for a group of variations.</p> <p>B.I.b.2.d - Change in test procedure for AS or starting material/reagent/intermediate - Substantial change to or replacement of a biological/immunological/immunochemical test method or a method using a biological reagent for a biological AS</p> <p>B.I.a.1.j - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Replacement or addition of a site where batch control/testing takes place and any of the test method at the site is a biol/immunol method</p>	02/09/2021	n/a		
II/0040/G	<p>This was an application for a group of variations.</p> <p>B.I.b.2.a - Change in test procedure for AS or starting material/reagent/intermediate - Minor changes to an approved test procedure</p> <p>B.II.d.2.c - Change in test procedure for the finished</p>	02/09/2021	n/a		

	<p>product - Substantial change to or replacement of a biol/immunol/immunochemical test method or a method using a biol. reagent or replacement of a biol. reference preparation not covered by an approved protocol</p> <p>B.I.b.2.d - Change in test procedure for AS or starting material/reagent/intermediate - Substantial change to or replacement of a biological/immunological/immunochemical test method or a method using a biological reagent for a biological AS</p>				
II/0053/G	<p>This was an application for a group of variations.</p> <p>A.4 - Administrative change - Change in the name and/or address of a manufacturer or an ASMF holder or supplier of the AS, starting material, reagent or intermediate used in the manufacture of the AS or manufacturer of a novel excipient</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.z - Change in control of excipients in the</p>	25/08/2021	n/a		

	Finished Product - Other variation				
II/0060/G	<p>This was an application for a group of variations.</p> <p>B.II.e.6.b - Change in any part of the (primary) packaging material not in contact with the finished product formulation - Change that does not affect the product information</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.b.3.z - Change in the manufacturing process of the finished or intermediate product - Other variation</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.6.b - Change in any part of the (primary) packaging material not in contact with the finished product formulation - Change that does not affect the product information</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p>	20/08/2021	n/a		

	<p>dossier) - Replacement or addition of a supplier</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.6.b - Change in any part of the (primary) packaging material not in contact with the finished product formulation - Change that does not affect the product information</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p> <p>B.II.b.1.z - Replacement or addition of a manufacturing site for the FP - Other variation</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p> <p>B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier</p>				
IB/0058	B.I.a.3.e - Change in batch size (including batch size ranges) of AS or intermediate - The scale for a biological/immunological AS is increased/decreased without process change (e.g. duplication of line)	17/08/2021	n/a		
IB/0055	B.I.b.1.c - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Addition of a new	17/08/2021	n/a		

	specification parameter to the specification with its corresponding test method				
II/0038/G	<p>This was an application for a group of variations.</p> <p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p> <p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p> <p>C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data</p>	22/07/2021	23/07/2021	SmPC and PL	<p>Based on signal detection and evaluation activity in the post-authorisation setting, the section in 4.4 of the SmPC is updated by adding "vaccine stress-related responses" to the already existing warning on anxiety-related reactions (i.e. including examples on stress-related reactions that may occur in association with the vaccination). The provided data support that these anticipated reactions occur related to vaccination.</p> <p>In addition, as agreed by the PRAC, the product information is also updated to add "extensive swelling of the vaccinated limb" and "facial swelling", both at frequency unknown, to the list of adverse drug reactions in the section 4.8 of the SmPC.</p> <p>The Package Leaflet is updated accordingly.</p> <p>For more information, please refer to the Summary of Product Characteristics.</p>
II/0049/G	<p>This was an application for a group of variations.</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place</p>	20/07/2021	n/a		

II/0041/G	<p>This was an application for a group of variations.</p> <p>B.I.a.1.j - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Replacement or addition of a site where batch control/testing takes place and any of the test method at the site is a biol/immunol method</p> <p>B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product</p> <p>B.I.a.2.a - Changes in the manufacturing process of the AS - Minor change in the manufacturing process of the AS</p> <p>B.I.e.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p>	15/07/2021	n/a		
IB/0048/G	<p>This was an application for a group of variations.</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.4.f - Change in the batch size (including batch size ranges) of the finished product - The scale for a biological/immunological medicinal product is increased/decreased without process change (e.g. duplication of line)</p>	13/07/2021	n/a		

IAIN/0050	<p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p> <p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p>	12/07/2021	13/07/2021	SmPC and PL	To update sections 4.4 and 4.8 of the SmPC and sections 2 and 4 of the PL to implement the signal recommendations from the PRAC Updated Signal assessment report on Myocarditis, Pericarditis with Tozinameran (COVID-19 mRNA vaccine (nucleoside-modified) - COMIRNATY) (EPITT no 19712) adopted by PRAC on 08 July 2021
II/0045/G	<p>This was an application for a group of variations.</p> <p>B.I.b.2.d - Change in test procedure for AS or starting material/reagent/intermediate - Substantial change to or replacement of a biological/immunological/immunochemical test method or a method using a biological reagent for a biological AS</p> <p>B.I.a.1.j - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Replacement or addition of a site where batch control/testing takes place and any of the test method at the site is a biol/immunol method</p> <p>B.I.b.1.b - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Tightening of specification limits</p> <p>B.II.d.2.c - Change in test procedure for the finished product - Substantial change to or replacement of a biol/immunol/immunochemical test method or a method using a biol. reagent or replacement of a biol. reference preparation not covered by an approved protocol</p> <p>B.II.b.2.c.3 - Change to importer, batch release arrangements and quality control testing of the FP -</p>	09/07/2021	n/a		

	<p>Including batch control/testing for a biol/immunol product and any of the test methods is a biol/immunol/immunochemical method</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p>				
II/0043	<p>B.I.a.1.e - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - The change relates to a biological AS or a starting material [-] used in the manufacture of a biological/immunological product</p>	07/07/2021	n/a		
IB/0042/G	<p>This was an application for a group of variations.</p> <p>B.II.e.4.c - Change in shape or dimensions of the container or closure (immediate packaging) - Sterile medicinal products</p> <p>B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p> <p>B.II.e.7.b - Change in supplier of packaging</p>	18/06/2021	n/a		

	<p>components or devices (when mentioned in the dossier) - Replacement or addition of a supplier B.II.e.7.b - Change in supplier of packaging</p> <p>components or devices (when mentioned in the dossier) - Replacement or addition of a supplier B.II.e.7.b - Change in supplier of packaging</p> <p>components or devices (when mentioned in the dossier) - Replacement or addition of a supplier B.II.e.7.b - Change in supplier of packaging</p> <p>components or devices (when mentioned in the dossier) - Replacement or addition of a supplier B.II.e.6.b - Change in any part of the (primary) packaging material not in contact with the finished product formulation - Change that does not affect the product information</p> <p>B.II.e.7.a - Change in supplier of packaging</p> <p>components or devices (when mentioned in the dossier) - Deletion of a supplier</p> <p>B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product</p>				
II/0039/G	<p>This was an application for a group of variations.</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p>	08/06/2021	n/a		

	<p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>				
II/0035/G	<p>This was an application for a group of variations.</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.z - Quality change - Finished product - Other variation</p>	01/06/2021	n/a		

II/0030	<p>Extension of the existing indication from "individuals 16 years of age and older" to "individuals 12 years of age and older" for Comirnaty; as a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 2.0 of the RMP has also been submitted.</p> <p>C.I.6.a - Change(s) to therapeutic indication(s) - Addition of a new therapeutic indication or modification of an approved one</p>	28/05/2021	31/05/2021	SmPC and PL	<p>The application for extension of the indication to include adolescents 12-15 years of age is based on a single pivotal phase 1/2/3 study C4591001. It is an extension of the pivotal efficacy study in adults assessed in the initial approval of Comirnaty. The inferential analysis for the 12-15 year olds which was to demonstrate non-inferior immune responses in this age cohort, compared to subjects 16-25 years from the initial efficacy part of the same study. The effects of Comirnaty in children were investigated in 2,260 children aged 12 to 15 years. The trial showed that the immune response to Comirnaty in this group was comparable to the immune response in the 16 to 25 age group. The efficacy of Comirnaty was calculated in close to 2,000 children from 12 to 15 years of age who had no sign of previous infection. These received either the vaccine or a placebo. Of the 1,005 children receiving the vaccine, none developed COVID-19 compared to 16 children out of the 978 who received the placebo. This means that, in this study, the vaccine was 100% effective at preventing COVID-19 (although the true rate could be between 75% and 100%).</p> <p>The most common side effects in children aged 12 to 15 are similar to those in people aged 16 and above. They include pain at the injection site, tiredness, headache, muscle and joint pain, chills and fever. These effects are usually mild or moderate and improve within a few days from the vaccination.</p> <p>The CHMP concluded that the benefits of Comirnaty in this age group outweigh the risks.</p> <p>The CHMP also noted that due to the limited number of children included in the study, the trial could not have detected rare side effects. The committee also noted that</p>
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					<p>EMA's safety committee PRAC is currently assessing very rare cases of myocarditis and pericarditis that occurred after vaccination with Comirnaty, mainly in people under 30 years of age. Currently there is no indication that these cases are due to the vaccine and EMA is closely monitoring this issue.</p> <p>Despite this uncertainty, the CHMP considered that benefits of Comirnaty in children aged 12 to 15 outweigh the risks, in particular in children with conditions that increase the risk of severe COVID-19.</p>
II/0034	B.II.d.2.c - Change in test procedure for the finished product - Substantial change to or replacement of a biol/immunol/immunochemical test method or a method using a biol. reagent or replacement of a biol. reference preparation not covered by an approved protocol	28/05/2021	n/a		
IB/0037	B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)	27/05/2021	n/a		
II/0033/G	<p>This was an application for a group of variations.</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.c.z - Change in control of excipients in the</p>	25/05/2021	n/a		

	<p>Finished Product - Other variation</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>				
II/0023/G	<p>This was an application for a group of variations.</p> <p>C.I.11.b - Introduction of, or change(s) to, the obligations and conditions of a marketing authorisation, including the RMP - Implementation of change(s) which require to be further substantiated by new additional data to be submitted by the MAH where significant assessment is required</p> <p>B.II.b.5.z - Change to in-process tests or limits applied during the manufacture of the finished product - Other variation</p>	20/05/2021	31/05/2021	Annex II	<p>The MAH has submitted additional validation data, which were a condition to the Marketing Authorisation (Specific Obligation SO3; In order to confirm the consistency of the finished product manufacturing process, the MAH should provide additional validation data. Due date: March 2021). Following assessment of the data, Annex IIE (specific obligation to complete post-authorisation measures for a conditional marketing authorisation) is updated; SO3 is fulfilled and is deleted from the list of specific obligations.</p>
IB/0031/G	<p>This was an application for a group of variations.</p> <p>Sections 6.3 and 6.6 of the SmPC have been updated to change the storage conditions of the unopened thawed vial from '5 days at 2-8°C' to '1 month at 2-8°C'. An additional text field has been added to the outer packaging to insert the expiry date at 2 to 8°C.</p> <p>B.I.z - Quality change - Active substance - Other variation</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.I.b.1.a - Change in the specification parameters</p>	17/05/2021	31/05/2021	SmPC, Labelling and PL	<p>The SmPC, labelling and package leaflet are updated to extend the shelf life of thawed vials from 5 days to 1 months at 2-8°C.</p> <p>In addition, section 6.3 of the SmPC is updated in line with CHMP/QWP/159/96 corr. (Note for Guidance of Maximum Shelf-Life for Sterile Products for Human Use After First Opening or Following Reconstitution).</p>

	<p>and/or limits of an AS, starting material/intermediate/reagent - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.I.a.2.a - Changes in the manufacturing process of the AS - Minor change in the manufacturing process of the AS</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.I.b.1.a - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Tightening of specification limits for medicinal products subject to OCABR</p> <p>B.II.d.1.a - Change in the specification parameters and/or limits of the finished product - Tightening of specification limits</p> <p>B.II.f.1.d - Stability of FP - Change in storage conditions of the finished product or the diluted/reconstituted product</p> <p>B.II.f.1.d - Stability of FP - Change in storage conditions of the finished product or the diluted/reconstituted product</p>				
IA/0032	B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier	12/05/2021	n/a		
II/0028/G	This was an application for a group of variations.	07/05/2021	n/a		

B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method

B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site

B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes

B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place

B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place

B.II.b.1.b - Replacement or addition of a manufacturing site for the FP - Primary packaging site

B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place

B.II.d.2.a - Change in test procedure for the finished

	<p>product - Minor changes to an approved test procedure</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.d.2.d - Change in test procedure for the finished product - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.b.2.a - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p>				
IB/0029	B.II.c.z - Change in control of excipients in the Finished Product - Other variation	04/05/2021	n/a		
II/0027	B.II.g.2 - Introduction of a post approval change management protocol related to the finished product	27/04/2021	n/a		

II/0022/G	<p>This was an application for a group of variations.</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.4.c - Change in the batch size (including batch size ranges) of the finished product - The change requires assessment of the comparability of a biological/immunological medicinal product or a new bioequivalence study</p>	21/04/2021	n/a		
II/0026	B.II.g.2 - Introduction of a post approval change management protocol related to the finished product	20/04/2021	n/a		
IB/0025/G	<p>This was an application for a group of variations.</p> <p>B.II.z - Quality change - Finished product - Other variation</p> <p>B.II.c.1.c - Change in the specification parameters and/or limits of an excipient - Deletion of a non-significant specification parameter (e.g. deletion of an obsolete parameter)</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p>	20/04/2021	n/a		
II/0019	C.I.11.b - Introduction of, or change(s) to, the obligations and conditions of a marketing	15/04/2021	n/a		

	authorisation, including the RMP - Implementation of change(s) which require to be further substantiated by new additional data to be submitted by the MAH where significant assessment is required				
II/0016/G	<p>This was an application for a group of variations.</p> <p>Group of two type II variations C.I.3.b consisting of: C.I.3.b - Change(s) in the SPC, Labelling or PL intended to implement the outcome of a procedure concerning PSUR or PASS or the outcome of the assessment done under A 45/46 - Change(s) with new additional data submitted by the MAH C.I.3.b - Change(s) in the SPC, Labelling or PL intended to implement the outcome of a procedure concerning PSUR or PASS or the outcome of the assessment done under A 45/46 - Change(s) with new additional data submitted by the MAH</p> <p>C.I.3.b - Change(s) in the SPC, Labelling or PL intended to implement the outcome of a procedure concerning PSUR or PASS or the outcome of the assessment done under A 45/46 - Change(s) with new additional data submitted by the MAH C.I.3.b - Change(s) in the SPC, Labelling or PL intended to implement the outcome of a procedure concerning PSUR or PASS or the outcome of the assessment done under A 45/46 - Change(s) with new additional data submitted by the MAH</p>	13/04/2021	14/04/2021	SmPC and PL	This variation pertains to two updates of the section 4.8 of the SmPC to implement the wording agreed by the PRAC following the outcome of the Post Authorisation Measures PAM-MEA-002.1 (EMA/H/C/005735/MEA/002.1) and PAM-LEG-022 (EMA/H/C/005735/LEG/022). In this context, 2 new adverse drug reactions (ADRs) (“diarrhoea”, “vomiting”) were added with very common and common frequencies respectively, and the ADR “pain in extremity” was updated with a footnote. In addition, the ADR concerning “hypersensitivity reactions” was further detailed (e.g. “rash, pruritus, urticaria, angioedema”) with the relevant frequency categories.
IB/0024/G	This was an application for a group of variations.	13/04/2021	n/a		

	<p>B.II.f.1.e - Stability of FP - Change to an approved stability protocol</p> <p>B.II.f.1.e - Stability of FP - Change to an approved stability protocol</p> <p>B.II.f.z - Stability of FP - Other variation</p>				
II/0020/G	<p>This was an application for a group of variations.</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>	31/03/2021	n/a		
II/0017/G	<p>This was an application for a group of variations.</p> <p>B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site</p> <p>B.II.e.4.c - Change in shape or dimensions of the container or closure (immediate packaging) - Sterile medicinal products</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.b.1.c - Replacement or addition of a</p>	29/03/2021	n/a		

	<p>manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p> <p>B.II.b.1.b - Replacement or addition of a manufacturing site for the FP - Primary packaging site</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.d.2.d - Change in test procedure for the finished product - Other changes to a test procedure (including replacement or addition)</p>				
IB/0021	B.II.g.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product	26/03/2021	n/a		
II/0018/G	This was an application for a group of variations.	26/03/2021	n/a		

	B.II.g.2 - Introduction of a post approval change management protocol related to the finished product B.II.g.2 - Introduction of a post approval change management protocol related to the finished product				
IB/0013	B.II.f.1.d - Stability of FP - Change in storage conditions of the finished product or the diluted/reconstituted product	26/03/2021	14/04/2021	SmPC and PL	
IB/0014	B.I.e.5.c - Implementation of changes foreseen in an approved change management protocol - For a biological/immunological medicinal product	24/03/2021	14/04/2021	Annex II	Update of Annex II of the product information to include an additional active substance manufacturing site.
IB/0015	B.II.d.2.d - Change in test procedure for the finished product - Other changes to a test procedure (including replacement or addition)	17/03/2021	n/a		
II/0012	B.II.b.3.c - Change in the manufacturing process of the finished or intermediate product - The product is a biological/immunological medicinal product and the change requires an assessment of comparability	12/03/2021	n/a		
II/0011/G	This was an application for a group of variations. B.I.a.1.j - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Replacement or addition of a site where batch control/testing takes place and any of the test method at the site is a biol/immunol method B.I.b.2.a - Change in test procedure for AS or starting material/reagent/intermediate - Minor	09/03/2021	n/a		

	<p>changes to an approved test procedure</p> <p>B.II.b.2.b - Change to importer, batch release arrangements and quality control testing of the FP - Replacement/addition of a site where batch control/testing takes place for a biol/immunol product and any of the test methods at the site is a biol/immunol method</p> <p>B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure</p>				
II/0009	<p>B.II.f.1.c - Stability of FP - Change in storage conditions for biological medicinal products, when the stability studies have not been performed in accordance with an approved stability protocol</p>	19/02/2021	23/02/2021	SmPC and PL	<p>The SmPC sections 6.3 and 6.6 have been updated to include information related to the transportation of diluted and undiluted product at non-frozen storage conditions and to the handling of temperature excursions once removed from the freezer as follows:</p> <p>Unopened vial;</p> <p>Once removed from the freezer, the unopened vaccine can be stored for up to 5 days at 2 °C to 8 °C. Within the 5 days shelf-life at 2 °C to 8 °C, up to 12 hours may be used for transportation. Prior to use, the unopened vaccine can be stored for up to 2 hours at temperatures up to 30 °C.</p> <p>Diluted medicinal product;</p> <p>Chemical and physical in-use stability, including during transportation, has been demonstrated for 6 hours at 2 °C to 30 °C after dilution in sodium chloride 9 mg/mL (0.9%) solution for injection. From a microbiological point of view, the product should be used immediately. If not used immediately, in-use storage times and conditions are the responsibility of the user.</p>

					<p>Handling of temperature excursions once removed from the freezer</p> <p>Stability data indicate that the unopened vial is stable for up to:</p> <ul style="list-style-type: none"> • 24 hours when stored at temperatures from -3 °C to 2 °C • a total of 4 hours when stored at temperatures from 8 °C to 30 °C; this includes the 2 hours at up to 30 °C detailed above <p>This information is intended to guide healthcare professionals only in case of temporary temperature excursion.</p> <p>The PL has been updated accordingly.</p>
II/0010/G	<p>This was an application for a group of variations.</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.4.a - Change in synthesis or recovery of a non-pharmacopoeial or novel excipient - Minor change</p> <p>B.II.c.2.a - Change in test procedure for an excipient - Minor changes to an approved test procedure</p>	22/02/2021	n/a		
II/0008/G	<p>This was an application for a group of variations.</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for</p>	16/02/2021	n/a		

<p>biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.3.a - Change in the manufacturing process of the finished or intermediate product - Minor change in the manufacturing process</p> <p>B.II.b.1.c - Replacement or addition of a manufacturing site for the FP - Site where any manufacturing operation(s) take place, except batch release/control, and secondary packaging, for biol/immunol medicinal products or pharmaceutical forms manufactured by complex manufacturing processes</p> <p>B.II.b.5.z - Change to in-process tests or limits applied during the manufacture of the finished product - Other variation</p> <p>B.II.b.4.c - Change in the batch size (including batch size ranges) of the finished product - The change requires assessment of the comparability of a biological/immunological medicinal product or a new bioequivalence study</p> <p>B.II.b.4.c - Change in the batch size (including batch size ranges) of the finished product - The change requires assessment of the comparability of a biological/immunological medicinal product or a new</p>				
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	<p>bioequivalence study</p> <p>B.II.b.4.c - Change in the batch size (including batch size ranges) of the finished product - The change requires assessment of the comparability of a biological/immunological medicinal product or a new bioequivalence study</p> <p>B.II.b.5.a - Change to in-process tests or limits applied during the manufacture of the finished product - Tightening of in-process limits</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>				
II/0005	B.II.g.2 - Introduction of a post approval change management protocol related to the finished product	10/02/2021	n/a		
II/0004	B.I.e.2 - Introduction of a post approval change management protocol related to the AS	04/02/2021	n/a		
IB/0007	<p>To update the dose interval for Comirnaty in sections 4.2, 5.1 of the SmPC and section 3 and section "The following information is intended for healthcare professionals only" of the PL to implement the recommendation of the CHMP.</p> <p>The wording of the dose interval in section 4.2 has been updated with a view to reflect better in section 4.2 the evidence from the submitted data and provide more clear guidance as well as to describe further in section 5.1 the available data underlying this revision.</p> <p>The MAH has also taken the opportunity to introduce a few editorial changes and to provide updated</p>	28/01/2021	02/02/2021	SmPC and PL	

	<p>material affected by the change including 'How to prepare and administer poster' and 'Vaccine traceability card'</p> <p>In addition, the company has informed us that they updated the 'Administration video' and the 'Comirnaty website'.</p> <p>C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation</p>				
IB/0006	B.II.c.z - Change in control of excipients in the Finished Product - Other variation	28/01/2021	n/a		
II/0003/G	<p>This was an application for a group of variations.</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.2.d - Change in test procedure for an excipient - Other changes to a test procedure (including replacement or addition)</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.1.b - Change in the specification parameters and/or limits of an excipient - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.c.1.c - Change in the specification parameters and/or limits of an excipient - Deletion of a non-significant specification parameter (e.g. deletion of an obsolete parameter)</p>	27/01/2021	n/a		

	<p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p> <p>B.II.c.z - Change in control of excipients in the Finished Product - Other variation</p>				
IB/0001	B.I.d.1.a.4 - Stability of AS - Change in the re-test period/storage period - Extension or introduction of a re-test period/storage period supported by real time data	26/01/2021	n/a		
II/0002/G	<p>This was an application for a group of variations.</p> <p>Grouped variation:</p> <p>B.II.e.5.c - Change in pack size of the finished product - Change in the fill weight/fill volume of sterile multidose (or single-dose, partial use) parenteral medicinal products, including biological/immunological medicinal products</p> <p>B.II.d.1.d - Change in the specification parameters and/or limits of the finished product - Deletion of a non-significant specification parameter</p> <p>B.II.d.1.c - Change in the specification parameters</p>	08/01/2021	08/01/2021	SmPC, Labelling and PL	<p>The change in pack size and change in specifications were approved.</p> <p>The SmPC sections 2, 4.2, 6.5 6.6 has been updated to reflect the following:</p> <p>One vial (0.45 mL) contains 6 doses of 0.3 mL after dilution. In order to extract six doses from a single vial, low dead-volume syringes and/or needles should be used. The low dead-volume syringe and needle combination should have a dead volume of no more than 35 microliters. If standard syringes and needles are used, there may not be enough of the vaccine to extract a sixth dose from a vial. If the amount of vaccine remaining in the vial after the fifth dose cannot provide a full dose (0.3 ml), the healthcare professional must discard the vial and its contents. There should be no pooling from multiple vials to make up a full</p>

	<p>and/or limits of the finished product - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.d.1.d - Change in the specification parameters and/or limits of the finished product - Deletion of a non-significant specification parameter</p> <p>B.II.d.1.c - Change in the specification parameters and/or limits of the finished product - Addition of a new specification parameter to the specification with its corresponding test method</p> <p>B.II.e.5.c - Change in pack size of the finished product - Change in the fill weight/fill volume of sterile multidose (or single-dose, partial use) parenteral medicinal products, including biological/immunological medicinal products</p>				<p>dose, and any unused vaccine should be discarded 6 hours after dilution.</p> <p>The Labelling and PL have been updated accordingly.</p>
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