



Tevagrastim

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
IAIN/0065	C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation	31/05/2018		SmPC, Labelling and PL	
II/0064	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	28/09/2017	n/a		

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).



	biological AS				
II/0063/G	<p>This was an application for a group of variations.</p> <p>B.I.a.1.z - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Other variation</p> <p>B.I.a.1.z - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Other variation</p> <p>B.I.a.1.z - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Other variation</p> <p>B.I.a.2.c - Changes in the manufacturing process of the AS - The change refers to a [-] substance in the manufacture of a biological/immunological substance which may have a significant impact on the medicinal product and is not related to a protocol</p>	06/07/2017	n/a		
II/0061	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	10/11/2016	n/a		
IAIN/0062	B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site	12/10/2016	n/a		
IB/0060	C.I.11.z - Introduction of, or change(s) to, the obligations and conditions of a marketing authorisation, including the RMP - Other variation	26/09/2016	n/a		

IB/0059/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.I.b.2.a - Change in test procedure for AS or starting material/reagent/intermediate - Minor changes to an approved test procedure</p>	08/06/2016	n/a		
PSUSA/1391/201509	Periodic Safety Update EU Single assessment - filgrastim	13/05/2016	n/a		PRAC Recommendation - maintenance
IAIN/0058	B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site	12/05/2016	n/a		
II/0055/G	<p>This was an application for a group of variations.</p> <p>B.II.d.1.e - Change in the specification parameters and/or limits of the finished product - Change outside the approved specifications limits range</p> <p>B.II.f.1.e - Stability of FP - Change to an approved stability protocol</p>	07/04/2016	n/a		
IB/0057	B.I.a.1.z - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Other variation	18/03/2016	n/a		
N/0056	Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification)	03/02/2016		PL	

IA/0054	B.II.e.7.b - Change in supplier of packaging components or devices (when mentioned in the dossier) - Replacement or addition of a supplier	15/12/2015	n/a		
II/0052	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	05/11/2015	n/a		
IB/0051/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.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.d - Change in test procedure for the finished product - Other changes to a test procedure (including replacement or addition)</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.f.1.e - Stability of FP - Change to an approved stability protocol</p>	30/01/2015	n/a		

IA/0050	A.7 - Administrative change - Deletion of manufacturing sites	03/10/2014	n/a		
IB/0049	B.I.a.1.k - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - New storage site of MCB and/or WCB	29/08/2014	n/a		
IA/0048/G	This was an application for a group of variations. B.II.b.3.z - Change in the manufacturing process of the finished or intermediate product - Other variation B.II.b.5.c - Change to in-process tests or limits applied during the manufacture of the finished product - Deletion of a non-significant in-process test	15/08/2014	n/a		
IB/0047	B.I.c.1.z - Change in immediate packaging of the AS - Other variation	21/07/2014	n/a		
IB/0046	B.I.c.z - Container closure system of the AS - Other variation	28/05/2014	n/a		
IB/0045	B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation	20/05/2014	n/a		
IA/0044/G	This was an application for a group of variations. B.III.2.a.2 - Change of specification(s) of a former non EU Pharmacopoeial substance to fully comply with the Ph. Eur. or with a national pharmacopoeia of a Member State - Excipient/AS starting material	12/03/2014	n/a		

	B.III.2.a.2 - Change of specification(s) of a former non EU Pharmacopoeial substance to fully comply with the Ph. Eur. or with a national pharmacopoeia of a Member State - Excipient/AS starting material				
IB/0043	B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation	24/01/2014	n/a		
IB/0042/G	This was an application for a group of variations. B.I.b.2.e - Change in test procedure for AS or starting material/reagent/intermediate - Other changes to a test procedure (including replacement or addition) for the AS or a starting material/intermediate B.II.d.2.d - Change in test procedure for the finished product - Other changes to a test procedure (including replacement or addition)	17/12/2013	n/a		
IA/0041/G	This was an application for a group of variations. B.III.2.a.2 - Change of specification(s) of a former non EU Pharmacopoeial substance to fully comply with the Ph. Eur. or with a national pharmacopoeia of a Member State - Excipient/AS starting material B.III.2.a.2 - Change of specification(s) of a former non EU Pharmacopoeial substance to fully comply with the Ph. Eur. or with a national pharmacopoeia of a Member State - Excipient/AS starting material B.III.2.a.2 - Change of specification(s) of a former non EU Pharmacopoeial substance to fully comply with the Ph. Eur. or with a national pharmacopoeia of a Member	22/11/2013	n/a		

	<p>State - Excipient/AS starting material</p> <p>B.III.2.a.2 - Change of specification(s) of a former non EU Pharmacopoeial substance to fully comply with the Ph. Eur. or with a national pharmacopoeia of a Member State - Excipient/AS starting material</p> <p>B.III.2.a.2 - Change of specification(s) of a former non EU Pharmacopoeial substance to fully comply with the Ph. Eur. or with a national pharmacopoeia of a Member State - Excipient/AS starting material</p>				
IB/0039	<p>C.I.2.a - Change in the SPC, Labelling or PL of a generic/hybrid/biosimilar products following assessment of the same change for the reference product - Implementation of change(s) for which NO new additional data is required to be submitted by the MAH.</p> <p>C.I.2.a - Change in the SPC, Labelling or PL of a generic/hybrid/biosimilar products following assessment of the same change for the reference product - Implementation of change(s) for which NO new additional data is required to be submitted by the MAH</p>	30/10/2013	04/12/2014	SmPC and PL	Update of sections 4.4 and 4.8 of the SmPC to include a new adverse reaction, capillary leak syndrome, and a related warning following the same update for the originator. The PL is updated accordingly.
IA/0040	B.II.e.2.a - Change in the specification parameters and/or limits of the immediate packaging of the finished product - Tightening of specification limits	21/10/2013	n/a		
R/0036	Renewal of the marketing authorisation.	30/05/2013	19/07/2013	SmPC, Annex II, Labelling and PL	

IA/0038/G	<p>This was an application for a group of variations.</p> <p>B.II.b.3.z - Change in the manufacturing process of the finished product - Other variation</p> <p>B.II.b.5.c - Change to in-process tests or limits applied during the manufacture of the finished product - Deletion of a non-significant in-process test</p>	11/07/2013	n/a		
IA/0037	<p>B.II.e.2.a - Change in the specification parameters and/or limits of the immediate packaging of the finished product - Tightening of specification limits</p>	11/04/2013	n/a		
IB/0035/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.b - Change to in-process tests or limits applied during the manufacture of the finished product - Addition of a new tests and limits</p>	18/01/2013	n/a		
II/0033/G	<p>This was an application for a group of variations.</p> <p>To introduce a new site for WCB manufacture.</p> <p>To introduce a new WCB lot and some modifications to the manufacturing process of the WCB (change in scale, harvesting time and freezing volume).</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</p>	15/11/2012	n/a		

	biological/immunological product B.I.a.2.c - Changes in the manufacturing process of the AS - The change refers to a [-] substance in the manufacture of a biological/immunological medicinal product and is not related to a protocol				
IB/0034	C.I.3.a - Implementation of change(s) requested following the assessment of an USR, class labelling, a PSUR, RMP, FUM/SO, data submitted under A 45/46, or amendments to reflect a Core SPC - Changes with NO new additional data are submitted by the MAH	24/10/2012	31/10/2012	SmPC, Annex II, Labelling and PL	
IB/0030/G	This was an application for a group of variations. B.I.b.2.a - Change in test procedure for AS or starting material/reagent/intermediate - Minor changes to an approved test procedure B.I.a.4.z - Change to in-process tests or limits applied during the manufacture of the AS - Other variation	21/09/2012	n/a		
IA/0032	A.5.b - Administrative change - Change in the name and/or address of a manufacturer of the finished product, including quality control sites (excluding manufacturer for batch release)	12/09/2012	n/a		
IB/0031	B.II.b.1.z - Replacement or addition of a manufacturing site for the FP - Other variation	10/09/2012	n/a		
II/0029	Replacement of the peptide mapping method. B.I.b.2.d - Change in test procedure for AS or starting material/reagent/intermediate - Change	19/07/2012	19/07/2012		

	(replacement) to a biological/immunological/ immunochemical test method or a method using a biological reagent for a biological AS				
IB/0027	B.1.b.2.e - Change in test procedure for AS or starting material/reagent/intermediate - Other changes to a test procedure (including replacement or addition) for the AS or a starting material/intermediate	10/05/2012	n/a		
IA/0026/G	This was an application for a group of variations. B.1.c.2.b - Change in the specification parameters and/or limits of the immediate packaging of the AS - Addition of a new specification parameter to the specification with its corresponding test method B.1.c.2.c - Change in the specification parameters and/or limits of the immediate packaging of the AS - Deletion of a non-significant specification parameter (e.g. deletion of an obsolete parameter)	14/03/2012	n/a		
N/0024	Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification)	09/12/2011	06/02/2012	PL	
IB/0025	B.1.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)	08/12/2011	n/a		
IA/0022/G	This was an application for a group of variations. B.1.b.1.b - Change in the specification parameters and/or limits of an AS, starting	25/10/2011	n/a		

	<p>material/intermediate/reagent - Tightening of specification limits</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.I.d.1.a.1 - Stability of AS - Change in the re-test period/storage period - Reduction</p>				
II/0021/G	<p>This was an application for a group of variations.</p> <p>Introduction of additional sterile filtration steps in the manufacturing process for the active substance. In addition, new in-process tests and limits are introduced.</p> <p>B.I.a.2.c - Changes in the manufacturing process of the AS - The change refers to a [-] substance in the manufacture of a biological/immunological medicinal product and is not related to a protocol</p> <p>B.I.a.4.b - Change to in-process tests or limits applied during the manufacture of the AS - Addition of a new in-process test and limits</p>	20/10/2011	20/10/2011		
IA/0023	A.1 - Administrative change - Change in the name and/or address of the MAH	20/10/2011	n/a	SmPC, Annex II, Labelling and PL	
IB/0020/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>	20/07/2011	n/a		

	B.II.d.2.a - Change in test procedure for the finished product - Minor changes to an approved test procedure				
IB/0019	B.II.b.3.z - Change in the manufacturing process of the finished product - Other variation	11/05/2011	n/a		
IB/0018/G	<p>This was an application for a group of variations.</p> <p>C.I.3.a - Implementation of change(s) requested following the assessment of an USR, class labelling, a PSUR, RMP, FUM/SO, data submitted under A 45/46, or amendments to reflect a Core SPC - Changes with NO new additional data are submitted by the MAH</p> <p>C.I.2.a - Change in the SPC, Labelling or PL of a generic/hybrid/biosimilar products following assessment of the same change for the reference product - Implementation of change(s) for which NO new additional data are submitted by the MAH</p>	21/02/2011	n/a	SmPC and PL	<p>To implement changes requested in the PSUR 3 Assessment Report (dated 06 August 2010), as well as to bring the Product Information in line with the SmPC of the reference product (Neupogen). Minor linguistic amendments are introduced in the Package Leaflet for the following languages: ES, HU, IT, LT, NL, PL, SL .</p> <p>In addition, the MAH amended the European Medicines Agency web address in section 10 of the SmPC.</p>
IA/0017	A.1 - Administrative change - Change in the name and/or address of the MAH	05/11/2010	n/a	SmPC, Labelling and PL	
IA/0016	<p>To add an alternative site responsible for secondary packaging.</p> <p>B.II.b.1.a - Replacement or addition of a manufacturing site for the FP - Secondary packaging site</p>	29/09/2010	n/a		
IB/0014	B.II.d.2.d - Change in test procedure for the finished	27/08/2010	n/a		

	product - Other changes to a test procedure (including replacement or addition)				
IA/0015	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	13/08/2010	n/a		
II/0011	Addition an alternative site for manufacture (formulation, filling) of drug product. 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, batch control, and secondary packaging, for biological/immunological medicinal products.	24/06/2010	01/07/2010		
IB/0013	To change a test procedure for the active substance B.I.b.2.e - Change in test procedure for AS or starting material/reagent/intermediate - Other changes to a test procedure (including replacement or addition) for the AS or a starting material/intermediate	20/05/2010	n/a		
IB/0012	To change a test procedure for the finished product B.II.d.2.d - Change in test procedure for the finished product - Other changes to a test procedure (including replacement or addition)	20/05/2010	n/a		
II/0004	Extension of the finished product shelf-life from 24 to 30 months.	19/11/2009	21/12/2009	SmPC, Labelling and	

	Change(s) to shelf-life or storage conditions			PL	
IA/0010	IA_28_Change in any part of primary packaging material not in contact with finished product	02/10/2009	02/10/2009	SmPC, Labelling and PL	
IA/0009	IA_28_Change in any part of primary packaging material not in contact with finished product	02/10/2009	02/10/2009	SmPC, Labelling and PL	
IA/0008	IA_28_Change in any part of primary packaging material not in contact with finished product	02/10/2009	02/10/2009	SmPC, Labelling and PL	
IA/0007	IA_28_Change in any part of primary packaging material not in contact with finished product	02/10/2009	02/10/2009	SmPC, Labelling and PL	
IA/0006	IA_28_Change in any part of primary packaging material not in contact with finished product	02/10/2009	02/10/2009	SmPC, Labelling and PL	
IA/0005	IA_28_Change in any part of primary packaging material not in contact with finished product	02/10/2009	02/10/2009	SmPC, Labelling and PL	
IA/0002	IA_01_Change in the name and/or address of the marketing authorisation holder	08/09/2009	n/a	SmPC, Labelling and PL	
N/0001	Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification)	19/02/2009	n/a	PL	