# Emadine

## 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 |
|---|---|---|---|---|---|
| T/0046 | Transfer of Marketing Authorisation | 20/03/2018 | 23/04/2018 | SmPC, Labelling and PL | |
| PSUSA/1207/ 201705 | Periodic Safety Update EU Single assessment - emedastine | 11/01/2018 | n/a | | PRAC Recommendation - maintenance |
| T/0044 | Transfer of Marketing Authorisation | 06/04/2017 | 16/05/2017 | SmPC, Labelling and PL | |

[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).

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| IB/0043/G | This was an application for a group of variations.<br><br>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<br>A.7 - Administrative change - Deletion of manufacturing sites<br>B.I.a.1.f - Change in the manufacturer of AS or of a starting material/reagent/intermediate for AS - Changes to quality control testing arrangements for the AS -replacement or addition of a site where batch control/testing takes place<br>B.I.b.1.b - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Tightening of specification limits<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation<br>B.I.d.1.a.1 - Stability of AS - Change in the re-test period/storage period - Reduction<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation | 30/01/2017 | n/a | | |

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| | B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation<br>B.I.b.1.z - Change in the specification parameters and/or limits of an AS, starting material/intermediate/reagent - Other variation | | | | |
| IA/0042/G | This was an application for a group of variations.<br>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<br>A.5.b - Administrative change - Change in the name and/or address of a manufacturer/importer of the finished product, including quality control sites (excluding manufacturer for batch release) | 13/05/2016 | n/a | | |
| IA/0041 | 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 | 29/03/2016 | n/a | | |

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| IAIN/0040 | B.II.b.2.c.1 - Change to importer, batch release arrangements and quality control testing of the FP - Replacement or addition of a manufacturer responsible for importation and/or batch release - Not including batch control/testing | 06/01/2016 | 04/01/2017 | Annex II and PL | |
| PSUSA/1207/ 201405 | Periodic Safety Update EU Single assessment - emedastine | 09/01/2015 | n/a | | PRAC Recommendation - maintenance |
| IG/0452 | C.I.8.a - Introduction of or changes to a summary of Pharmacovigilance system - Changes in QPPV (including contact details) and/or changes in the PSMF location | 28/07/2014 | n/a | | |
| II/0037 | Update of sections 2, 4.2, 4.3, 4.6, 4.7, 4.8 and 4.9 of the SmPC following a review of the available clinical data and post-marketing experience with the product. The corresponding sections of the Labelling and Package Leaflet have been updated. The product information has also been updated in accordance with the recommendations of the SmPC Guideline and the latest QRD template version 9. Local representatives were updated for the following countries: Denmark, Norway, Slovakia, Iceland, Finland and Sweden.<br><br>C.I.4 - Change(s) in the SPC, Labelling or PL due to new quality, preclinical, clinical or pharmacovigilance data | 20/03/2014 | 16/03/2015 | SmPC, Annex II, Labelling and PL | The MAH submitted a variation in order to update the sections 2, 4.2, 4.3, 4.6, 4.7, 4.8 and 4.9 of the SmPC following the MAH's review of the available data supporting the safety profile for Emadine eye drops, solution. The review has taken into consideration data from clinical studies and post-marketing experience with the product. As a result of the review the MAH has revised the overall safety profile for Emadine eye drops in section 4.8 of the SmPC the frequency category of five adverse events (headache, eye irritation, dry eye, corneal staining and blurred vision) has been reassigned from Common to Uncommon for consistency with the requirements in the EU SmPC guideline. According to this document ADRs with crude incidence rate between 0.1 to 1% (≥1/1000 to <1/100) should be assigned to the uncommon frequency category; section 4.6 of the SmPC the following sentence was added 'Studies in animals have shown no evidence of impaired fertility (see Section 5.3). No human fertility data are available.' and for the section 5.3 of the |

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| | | | | | SmPC for which the following sentence was reworded to 'There was no evidence of impaired fertility or decreased reproductive capacity in rats administered orally dosages of Emedastine difumarate up to 30 mg/kg/day.' |
| IB/0033 | B.II.e.2.z - Change in the specification parameters and/or limits of the immediate packaging of the finished product - Other variation | 24/07/2013 | n/a | | |
| IG/0324 | C.I.z - Changes (Safety/Efficacy) of Human and Veterinary Medicinal Products - Other variation | 22/07/2013 | n/a | | |
| N/0035 | Update of the local representatives contact details in the Package Leaflet, including addition of a local representative for Croatia.<br><br>Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification) | 18/07/2013 | 05/03/2014 | PL | Update of the local representatives contact details in the Package Leaflet, including addition of a local representative for Croatia. |
| IB/0032 | B.II.b.3.z - Change in the manufacturing process of the finished product - Other variation | 21/06/2013 | n/a | | |
| IG/0274 | A.1 - Administrative change - Change in the name and/or address of the MAH | 19/03/2013 | 05/03/2014 | SmPC, Labelling and PL | |
| WS/0075 | This was an application for a variation following a worksharing procedure according to Article 20 of Commission Regulation (EC) No 1234/2008.<br><br>To replace the current resin which is used for the closures for the drop-trainer packaging system, with two new resins. | 20/01/2011 | 20/01/2011 | | |

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| | B.II.e.1.a.3 - Change in immediate packaging of the finished product - Qualitative and quantitative composition - Sterile medicinal products and biological/immunological medicinal products | | | | |
| N/0027 | Update of the local representatives contact details for Czech Republic, Finland, Iceland, and Slovakia. The MAH also took the opportunity to make minor linguistic amendments in annexes III A and B, and updated the Agency's website address to reflect its new name.<br><br>Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification) | 30/11/2010 | n/a | Labelling and PL | |
| II/0025 | to add a new sterility test method to the release and shelf-life specifications for the Emadine single dose container presentations<br><br>Quality changes | 22/07/2010 | 19/08/2010 | | |
| IB/0026 | To change the name of the finished product.<br><br>IB_02_Change in the name of the medicinal product | 22/01/2010 | n/a | SmPC, Annex II, Labelling and PL | |
| IA/0024 | IA_04_Change in name and/or address of a manuf. of the active substance (no Ph. Eur. cert. avail.) | 05/02/2009 | n/a | | |
| IA/0023 | IA_25_b_02_Change to comply with Ph. - compliance with EU Ph. update - excipient | 05/02/2009 | n/a | | |

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| II/0021 | The MAH applied for an additional site of manufacture for the active ingredient.<br><br>Quality changes | 22/01/2009 | 28/01/2009 | | |
| R/0018 | Renewal of the marketing authorisation. | 20/11/2008 | 13/01/2009 | SmPC, Annex II, Labelling and PL | Based on the review of the available information the CHMP is of the opinion that the quality, the safety and the efficacy of this medicinal product continues to be adequately and sufficiently demonstrated and therefore considers that the benefit/risk profile of Emadine continues to be favourable.<br><br>With this procedure the MAH also updated the product information. This update was based on the requirements in the 2005 Summary of Product Characteristics (SPC) Guidelines and QRD requirements. In detail this meant that the current section 4.8 (Undesirable effects) of the SPC, that was based upon 3 long-term studies, was updated to be based on all short-term, intermediate-term and long-term studies. All identified adverse events were therefore recoded into MedDRA preferred terms and the frequency classification for each adverse drug reaction (ADR) was identified. In addition, based on post-marketing adverse event data, the ADR "tachycardia" was included in section 4.8 of the SPC. Based upon the revision to section 4.8 of the SPC, section 4 of the PIL was subsequently updated. |
| IB/0022 | IB_30_b_Change in supplier of packaging components - replacement/addition | 04/09/2008 | n/a | | |
| IB/0020 | IB_25_a_01_Change to comply with Ph. - compliance with EU Ph. - active substance | 28/08/2008 | n/a | | |

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| IB/0019 | IB_30_b_Change in supplier of packaging components - replacement/addition | 28/08/2008 | n/a | | |
| II/0017 | Change(s) to the pharmaceutical documentation.<br><br>Quality changes | 15/11/2007 | 21/11/2007 | | |
| N/0016 | Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification) | 16/02/2007 | n/a | PL | |
| N/0015 | Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification) | 15/11/2004 | n/a | PL | |
| N/0014 | Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification) | 09/07/2004 | n/a | PL | |
| R/0013 | Renewal of the marketing authorisation. | 17/12/2003 | 05/03/2004 | SmPC, Annex II, Labelling and PL | |
| I/0012 | 20_Extension of shelf-life as foreseen at time of authorisation | 24/07/2003 | 16/09/2003 | SmPC | |
| II/0011 | Update of Summary of Product Characteristics and Package Leaflet | 20/02/2003 | 19/05/2003 | SmPC, Labelling and PL | |
| I/0010 | 01_Change in or addition of manufacturing site(s) for part or all of the manufacturing process | 20/06/2002 | 25/06/2002 | | |
| X/0009 | X-3-iv_Change or addition of a new pharmaceutical form | 21/02/2002 | 30/05/2002 | SmPC, Annex II, Labelling and PL | |

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| I/0008 | 01_Change in or addition of manufacturing site(s) for part or all of the manufacturing process | 04/08/2000 | 04/08/2000 | | |
| I/0007 | 12_Minor change of manufacturing process of the active substance | 04/08/2000 | n/a | | |
| I/0006 | 20a_Extension of shelf-life or retest period of the active substance | 04/08/2000 | n/a | | |
| I/0005 | 24_Change in test procedure of active substance | 04/08/2000 | n/a | | |
| I/0004 | 24_Change in test procedure of active substance | 04/08/2000 | n/a | | |
| I/0003 | 24_Change in test procedure of active substance | 04/08/2000 | n/a | | |
| I/0002 | 01_Change in or addition of manufacturing site(s) for part or all of the manufacturing process | 24/03/2000 | 31/05/2000 | Annex II and PL | |
| N/0001 | Minor change in labelling or package leaflet not connected with the SPC (Art. 61.3 Notification) | 04/02/1999 | 16/03/1999 | Labelling and PL | |