Annex I

Scientific conclusions and grounds for the variation to the terms of the Marketing Authorisation(s)

Scientific conclusions

Taking into account the PRAC Assessment Report on the PSUR(s) for domperidone, the scientific conclusions are as follows:

In January 2016, the PRAC recommended in the context of the apomorphine PSUSA procedure (PSUSA/00000227/201505) an update of the product information of all domperidone-containing products in order to facilitate the safe use of domperidone together with apomorphine in the specific population of patients with Parkinson's disease.

In line with this recommendation, sections 4.3, 4.4 and 4.5 of the SmPC for all domperidone containing products should be updated to reflect, that domperidone, as an exception, may be co-administered with apomorphine, as far as all the recommended precautions as detailed in the apomorphine SmPC are fulfilled.

The CMDh agrees with the scientific conclusions made by the PRAC.

Grounds for the variation to the terms of the Marketing Authorisation(s)

On the basis of the scientific conclusions for domperidone the CMDh is of the opinion that the benefit-risk balance of the medicinal product(s) containing domperidone is unchanged subject to the proposed changes to the product information.

The CMDh reaches the position that the marketing authorisation(s) of products in the scope of this single PSUR assessment should be varied. To the extent that additional medicinal products containing domperidone are currently authorised in the EU or are subject to future authorisation procedures in the EU, the CMDh recommends that such marketing authorisations are varied accordingly.

Annex II

Amendments to the product information of the nationally authorised medicinal product(s)

Amendments to be included in the relevant sections of the Product Information (new text **underlined and in bold**, deleted text ~~strike through~~)

Summary of Product Characteristics

Section 4.3 Contraindications

Domperidone is contraindicated in the following situations:

• …

• co-administration with QT-prolonging drugs**, at the exception of apomorphine (see sections 4.4 and 4.5)**

• …

Section 4.4 Special warnings and precautions for use

Cardiovascular effects

(…)

Use with apomorphine :

Domperidone is contra-indicated with QT prolonging drugs including apomorphine, unless the benefit of the co-administration with apomorphine outweighs the risks, and only if the recommended precautions for co-administration mentioned in the apomorphine SmPC are strictly fulfilled. Please refer to the apomorphine SmPC.

Section 4.5 Interaction with other medicinal products and other forms of interaction

Increased risk of occurrence of QT-interval prolongation, due to pharmacodynamic and/or pharmacokinetic interactions.

Concomitant use of the following substances is contraindicated

QTc-prolonging medicinal products

(…)

- **apomorphine, unless the benefit of the co-administration outweighs the risks, and only if the recommended precautions for co-administration are strictly fulfilled. Please refer to the apomorphine SmPC.**

Package Leaflet

- Other drugs and <Product Name>

  <Product Name> and apomorphine

  Before you use <Product Name> and apomorphine, your doctor will ensure that you tolerate both medicines when used simultaneously. Ask your doctor or specialist for a personalised advice. Please refer to the apomorphine leaflet.

Annex III

Timetable for the implementation of this position

Timetable for the implementation of this position

Adoption of CMDh position:	July CMDh meeting
Transmission to National Competent Authorities of the translations of the annexes to the position:	2 September 2017
Implementation of the position by the Member States (submission of the variation by the Marketing Authorisation Holder):	1 November 2017