Summary of the risk management plan

Summary of risk management plan for Fulvestrant 250 mg solution for injection in pre-filled syringe. The RMP details important risks of Fulvestrant 250 mg solution for injection in pre-filled syringe, how these risks can be minimised, and how more information will be obtained about Fulvestrant 250 mg solution for injection in pre-filled syringe, risks and uncertainties (missing information). Fulvestrant 250 mg solution for injection in pre-filled syringe´s summary of product characteristics (SmPC) and its package leaflet give essential information to healthcare professionals and patients on Fulvestrant 250 mg solution for injection in pre-filled syringe should be used.

I. The medicine and what it is used for

Fulvestrant 250 mg solution for injection in pre-filled syringe is authorised for:

- as monotherapy for the treatment of estrogen receptor positive, locally advanced or metastatic breast cancer in postmenopausal women:

  - not previously treated with endocrine therapy, or
  - with disease relapse on or after adjuvant antiestrogen therapy, or disease progression on antiestrogen therapy.

- in combination with palbociclib for the treatment of HR-positive, HER2-negative locally advanced or metastatic breast cancer in women who have received prior endocrine therapy.

It contains Fulvestrant as active substance and it should be administered as two consecutive 5 ml injections by slow intramuscular injection (1-2 minutes/injection), one in each buttock (gluteal area).

II. Risks associated with the medicine and activities to minimise or further characterise the risks

Important risks of Fulvestrant 250 mg solution for injection in pre-filled syringe together with measures to minimise such risks are outlined below.

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

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

Together, these measures constitute a routine risk minimisation measures.

If important information that may affect the safe use of Fulvestrant is not yet available, it is listed under "missing information" below.

II.A List of important risks and missing information

Important risks of Fulvestrant 250 mg solution for injection in pre-filled syringe are risks that need special risk management activities to further investigate or minimise the risk, so that the medicinal product can be safely administered. Important risks can be regarded as identified or potential. Identified risks are concerns for which there is sufficient proof of a link with the use of Fulvestrant 250 mg solution for injection in pre-filled syringe.
Potential risks are concerns for which an association with the use of this medicine is possible based on available data, but this association has not been established yet and needs further evaluation. Missing information refers to information on the safety of the medicinal product that is currently missing and needs to be collected (e.g. on the long-term use of the medicine).

Table 1. Summary of Safety Concerns: list of important risk and missing information.

Important Identified Risk	• Injection site reactions • Increased risk of bleeding at the injection site • Venous thromboembolic events • Hypersensitivity reactions • Hepatobiliary disorders
Important Potential Risk	• Reduced bone mineral density (osteopenia) and osteoporosis • Ischaemic cardiovascular events • Endometrial dysplasia • Interstitial lung disease • Vasculitis • Pulmonary microembolism of oily solutions • Reprotoxicity (fertility, pregancy and lactation)
Missing information	• Paediatric use • Use with severe hepatic impairment • Use with renal impairment

II.B Summary of important risks

The safety information is aligned to the reference medicinal product.

II.C Post-authorisation development plan

II.C.1 Studies which are conditions of the marketing authorisation

There are no studies which are conditions of the marketing authorisation or specific obligation of Fulvestrant 250 mg solution for injection in pre-filled syringe.

II.C.2 Other studies in post-authorisation development plan

There are no studies required for Fulvestrant 250 mg solution for injection in pre-filled syringe.