Part VI: Summary of the risk management plan

Summary of risk management plan for Apixaban 2.5 mg and 5 mg film-coated tablets

This is a summary of the risk management plan (RMP) for Apixaban 2.5 mg and 5 mg film-coated tablets. The RMP details important risks of Apixaban 2.5 mg and 5 mg film-coated tablets, how these risks can be minimised, and how more information will be obtained about Apixaban 2.5 mg and 5 mg film-coated tablets's risks and uncertainties (missing information).

Apixaban 2.5 mg and 5 mg film-coated tablets's summary of product characteristics (SmPC) and its package leaflet give essential information to healthcare professionals and patients on how Apixaban 2.5 mg and 5 mg film-coated tablets should be used.

Important new concerns or changes to the current ones will be included in updates of Apixaban 2.5 mg and 5 mg film-coated tablets's RMP.

I. The medicine and what it is used for

Apixaban 2.5 mg / 5 mg film-coated tablet is indicated for:

- Prevention of stroke and systemic embolism in adult patients with non-valvular atrial fibrillation (NVAF), with one or more risk factors, such as prior stroke or transient ischaemic attack (TIA); age $\geq$ 75 years; hypertension; diabetes mellitus; symptomatic heart failure (NYHA Class $\geq$ II).
- Treatment of deep vein thrombosis (DVT) and pulmonary embolism (PE), and prevention of recurrent DVT and PE in adults.

Apixaban 2.5 mg film-coated tablet is indicated for:

- Prevention of venous thromboembolic events (VTE) in adult patients who have undergone elective hip or knee replacement surgery.

It contains Apixaban as the active substance and it is given by oral route of administration.

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

Important risks of Apixaban 2.5 mg and 5 mg film-coated tablets, together with measures to minimise such risks and the proposed studies for learning more about Apixaban 2.5 mg and 5 mg film-coated tablet's 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 *routine risk minimisation* measures.

In the case of Apixaban 2.5 mg and 5 mg film-coated tablets, these measures are supplemented with additional risk minimisation measures mentioned under relevant important risks, below.

In addition to these measures, information about adverse reactions is collected continuously and regularly analysed so that immediate action can be taken as necessary. These measures constitute routine pharmacovigilance activities.

If important information that may affect the safe use of Apixaban 2.5 mg and 5 mg film-coated tablet is not yet available, it is listed under 'missing information' below.

II.A List of important risks and missing information

Important risks of Apixaban 2.5 mg and 5 mg film-coated tablet 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 Apixaban 2.5 mg and 5 mg film-coated tablet. 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);

Summary of safety concerns	
Important Identified Risk	• Bleeding
Important Potential Risk	• Liver Injury • Potential risk of bleeding or thrombosis due to overdose or underdose
Missing Information	• Use in patients with severe renal impairment

II.B Summary of important risks

Important Identified Risk:

Bleeding	
Risk minimisation measures	***Routine risk minimisation measures***: SmPC Section 4.2, Posology and method of administration SmPC Section 4.3, Contraindications SmPC Section 4.4, Special warnings and precautions for use SmPC Section 4.5, Interaction with other medicinal products and other forms of interaction SmPC Section 4.8, Undesirable effects SmPC Section 4.9, Overdose ***Additional risk minimisation measures***: Prescriber Guide Patient Alert Card

Important Potential Risk:

Potential risk of bleeding or thrombosis due to overdose or underdose	
Risk minimisation measures	***Routine risk minimisation measures***: SmPC Section 4.2, Posology and method of administration SmPC Section 4.9, Overdose ***Additional risk minimisation measures***: Prescriber Guide

Missing information

Use in patients with severe Renal Impairment	
Risk minimisation measures	***Routine risk minimisation measures***: SmPC Section 4.2, Posology and method of administration SmPC Section 4.4, Special warnings and precautions for use

	SmPC Section 5.2, Pharmacokinetic properties SmPC provides the dosing recommendation for patients with severe renal impairment for each indication ***<u>Additional risk minimisation measures</u>***: None

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 Apixaban 2.5 mg and 5 mg film-coated tablet.

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

There are no studies required for Apixaban 2.5 mg and 5 mg film-coated tablet.