Part VI: Summary of the risk management plan

Mycophenolate mofetil 500 mg powder for concentrate for solution for infusion

This is a summary of the risk management plan (RMP) for mycophenolate mofetil. The RMP details important risks of mycophenolate mofetil, how these risks can be minimised, and how more information will be obtained about mycophenolate mofetil risks and uncertainties (missing information).

Mycophenolate mofetil's summary of product characteristics (SmPC) and its package leaflet give essential information to healthcare professionals and patients on how mycophenolate mofetil should be used.

Important new concerns or changes to the current ones will be included in updates of mycophenolate mofetil's RMP.

I. The medicine and what it is used for

Mycophenolate mofetil is authorised for following indication.

500 mg powder for concentrate for solution for infusion:

Mycophenolate mofetil is indicated in combination with ciclosporin and corticosteroids for the prophylaxis of acute transplant rejection in patients receiving allogeneic renal or hepatic transplants.

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

Important risks of mycophenolate mofetil, together with measures to minimise such risks and the proposed studies for learning more about mycophenolate mofetil'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 addition to these measures, information about adverse reactions is collected continuously and regularly analysed, including PSUR assessment and signal management activity, 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 mycophenolate mofetil is not yet available, it is listed under 'missing information' below.

II.A List of important risks and missing information

Important risks of mycophenolate mofetil are risks that need special risk management activities to further investigate or minimise the risk, so that the medicinal product can be safely taken. 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 mycophenolate mofetil. 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)

Important identified risks	• Hypersensitivity • Spontaneous abortion and congenital malformations in women • Lymphomas and other malignancies, particularly of the skin • Opportunistic infections, fatal infections and sepsis • Bone marrow depression • Gastrointestinal adverse events particularly ulceration, haemorrhage and perforation
Important potential risks	• Exacerbations of conditions due to hereditary deficiency of hypoxanthine-guanine phosphoribosyl-transferase (HGPRT) • Risk of infection with live vaccines • Reduced effectiveness of vaccines • Interstitial lung disease and pulmonary fibrosis • Increased risk of certain infections, possibly gastrointestinal haemorrhage and pulmonary oedema in elderly population • Spontaneous abortion and congenital malformations in men

Missing Information	• Use in cardiac or hepatic transplant patients with severe chronic renal impairment • Use in cardiac transplant patients with severe hepatic parenchymal disease • Use in paediatric cardiac or hepatic transplant patients • Use in paediatric population < 2 years in renal transplant • Treatment with mycophenolate mofetil during hepatic transplant rejection

II.B Summary of important risks

Important Identified Risks: **Spontaneous abortion and congenital malformations in women**	
Risk minimisation measures	Routine risk minimisation measures: Section 4.4, 4.6, and 4.8 of proposed Accord Mycophenolate SmPC and corresponding sections of PIL have information on this safety concern. Routine risk minimisation measures recommending specific clinical measures to address the risk: In order to assist patients in avoiding foetal exposure to mycophenolate and to provide additional important safety information, the Marketing Authorisation holder will provide educational materials to healthcare professionals. The educational materials will reinforce the warnings about the teratogenicity of mycophenolate; provide advice on contraception before therapy is started and guidance on the need for pregnancy testing. Full patient information about the teratogenic risk and the pregnancy prevention measures should be given by the physician to women of childbearing potential and, as appropriate, to male patients. This information was included in Section 4.4. Other routine risk minimisation measures include; the labelling; and the prescription only status of the product. Additional risk minimisation measures: DHPC communication letter, Patient Guide (Patient Alert Card) and Mycophenolate Mofetil Guide for Healthcare Providers.

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 mycophenolate mofetil.

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

There are no studies required for mycophenolate mofetil.