

## Part VI: Summary of the risk management plan

Summary of risk management plan for [Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki (dimethyl fumarate)

This is a summary of the risk management plan (RMP) for Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki. The RMP details important risks of Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki, how these risks can be minimised, and how more information will be obtained about [Product Name] 120 mg, 240 mg hard gastro-resistant capsules risks and uncertainties (missing information).

Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki's summary of product characteristics (SmPC) and its package leaflet give essential information to healthcare professionals and patients on how Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki.

Important new concerns or changes to the current ones will be included in updates of Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki's RMP.

### I. The medicine and what it is used for

Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki is authorised for the treatment of adult and paediatric patients aged 13 years and older with relapsing remitting multiple sclerosis (RRMS) (see SmPC for the full indication). It contains dimethyl fumarate as the active substance and it is taken orally.

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

Important risks of Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki, together with measures to minimise such risks and the proposed studies for learning more about Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki'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, so that immediate action can be taken as necessary. These measures constitute *routine pharmacovigilance activities*.

### **II.A List of important risks and missing information**

Important risks of Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil

120 mg, 240 mg magasýrupolin hörð hylki 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 Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki. 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).

<b>List of important risks and missing information</b>	
Important identified risks	Progressive Multifocal Leukoencephalopathy (PML)
	Decreases in leukocyte and lymphocyte counts
	Drug-induced liver injury
Important potential risks	Serious and opportunistic infections (other than PML and herpes zoster)
	Malignancies
	Effects on pregnancy outcome
	Interaction with nephrotoxic medications leading to renal toxicity
Missing information	Long-term efficacy and safety
	Safety profile in patients over the age of 55 years
	Safety profile in patients with moderate to severe renal impairment
	Safety profile in patients with hepatic impairment
	Safety profile in patients with severe active gastrointestinal disease
	Increased risk of infection in patients concomitantly taking anti-neoplastic or immunosuppressive therapies

## ***II.B Summary of important risks***

The safety information in the proposed Product 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 Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki

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

There are no studies required for Dimethyl fumarate STADA Nordic 120 mg, 240 mg magasýrupolið hart hylki / Dimethyl fumarate Zentiva Group a.s. 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl fumarate Newbury 120 mg, 240 mg magasýrupolin hörð hylki / Dimethyl fumarate Adalvo / Dimethyl Fumarate G.L. Pharma 120 mg, 240 mg Magasýrupolið hart hylki / Dimethyl Fumarate G.L. Pharma GmbH 120 mg, 240 mg Magasýrupolið hart hylki / Dimetex 120 mg, 240 mg magasýrupolin hörð hylki / Xevil 120 mg, 240 mg magasýrupolin hörð hylki.