Summary of risk management plan for Emtricitabine/Tenofovir disoproxil Krka

This is a summary of the risk management plan (RMP) for Emtricitabine/Tenofovir disoproxil Krka. The RMP details important risks of Emtricitabine/Tenofovir disoproxil Krka, how these risks can be minimised, and how more information will be obtained about Emtricitabine/Tenofovir disoproxil Krka 's risks and uncertainties (missing information).

Emtricitabine/Tenofovir disoproxil Krka 's summary of product characteristics (SmPC) and its package leaflet give essential information to healthcare professionals and patients on how Emtricitabine/Tenofovir disoproxil Krka should be used.

This summary of the RMP for Emtricitabine/Tenofovir disoproxil Krka should be read in the context of all this information including the assessment report of the evaluation and its plain-language summary, all which is part of the European Public Assessment Report (EPAR).

Important new concerns or changes to the current ones will be included in updates of Emtricitabine/Tenofovir disoproxil Krka 's RMP.

I. The medicine and what it is used for

Emtricitabine/Tenofovir disoproxil Krka is indicated in antiretroviral combination therapy for the treatment of HIV-1 infected adults, for the treatment of HIV-1 infected adolescents, with NRTI resistance or toxicities precluding the use of first line agents, aged 12 to < 18 years and indicated in combination with safer sex practices for pre-exposure prophylaxis to reduce the risk of sexually acquired HIV-1 infection in adults at high risk.

It contains Emtricitabine/Tenofovir disoproxil as the active substance and it is given orally.

Further information about the evaluation Emtricitabine/Tenofovir disoproxil Krka’s benefits can be found in Emtricitabine/Tenofovir disoproxil Krka’s EPAR, including in its plain-language summary, available on the EMA website, under the medicine’s webpage https://www.ema.europa.eu/en/medicines/human/EPAR/efavirenzemtricitabinetenofovir-disoproxil-krka

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

Important risks of Emtricitabine/Tenofovir disoproxil Krka, together with measures to minimise such risks and the proposed studies for learning more about Emtricitabine/Tenofovir disoproxil Krka'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 Emtricitabine/Tenofovir disoproxil Krka, these measures are supplemented with additional risk minimisatio measures mentioned under relevant important risks, below.

In addition to these measures, information about adverse reactions is collected continuously and regularly analysed , including PSUR assessment 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 Emtricitabine/Tenofovir disoproxil Krka is not yet available, it is listed under 'missing information' below.

II.A List of important risks and missing information

Important risks of Emtricitabine/Tenofovir disoproxil Krka 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 Emtricitabine/Tenofovir disoproxil Krka. 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 risks:	Getting HIV-1 infection (HIV-1 acquisition, including infection resulting from non-adherence) (PrEP indication) (emtricitabine/tenofovir)
	Development of resistance in patients with unrecognized or acute HIV-1 infection (PrEP indication) (emtricitabine/tenofovir)
	Kidney problems (Renal toxicity) (tenofovir)
	Bone problems (Bone events due to proximal renal tubulopathy/loss of BMD) (tenofovir)
Important potential risks:	None
Missing information:	Limited information on use in pregnant and breastfeeding women (Safety in pregnancy and lactation) (tenofovir)

II.B Summary of important risks

Summary of important risk that have corresponding additional risk minimisation activities is:

Important identified risk **< HIV-1 acquisition, including infection resulting from non-adherence (PrEP indication*)(emtricitabine/tenofovir)>***	
Risk minimisation measures	Routine risk minimisation measure: SmPC section 4.4. where also rcommendation for frequent reconfirmation of HIV negativity and stressing of importance of adherence to therapy are included. PL section 2 and 3., also with the information how the patients must take the medicine, recommendation about telling the doctor about recent flu-like illness, and recommendations for additional measures for lowering of risk of getting HIV. Additional risk minimisation measures: Educational materials: – PrEP educational brochure for prescribers entitled 'Important Safety Information for Prescribers About Emtricitabine/Tenofovir disoproxil Krka for a Pre-exposure Prophylaxis (PrEP) Indication' – PrEP Checklist for prescribers – PrEP educational brochure for the individual at risk entitled 'Important Information About Emtricitabine/Tenofovir disoproxil Krka to Reduce the Risk of getting Human Immunodeficiency Virus (HIV) Infection' – PrEP reminder card

Important identified risk **< Development of resistance in patients with unrecognized or acute HIV-1 infection (PrEP indication) *)(emtricitabine/tenofovir)>***	
Risk minimisation measures	Routine risk minimisation measure: SmPC sections 4.3 and 4.4. where also recommendation for frequent reconfirmation of HIV negativity is included. PL section 2 also with the information on how to detect early signs and symptoms of HIV infection, recommendation about telling the doctor about recent flu-like illness, and recommendations for additional measures for lowering of risk of getting HIV. Additional risk minimisation measures: Educational materials: – PrEP educational brochure for prescribers entitled 'Important Safety Information for Prescribers About Emtricitabine/Tenofovir disoproxil Krka for a Pre-exposure Prophylaxis (PrEP) Indication' – PrEP Checklist for prescribers – PrEP educational brochure for the individual at risk entitled 'Important Information About Emtricitabine/Tenofovir disoproxil Krka to Reduce the Risk of getting Human Immunodeficiency Virus (HIV) Infection' – PrEP reminder card

Important identified risk < Renal toxicity *(tenofovir)>*

Risk minimisation measures	Routine risk minimisation measure: SmPC sections 4.2, 4.4, 4.5 and 4.8,. SmPC section 4.4 where also advice is given on monitoring the renal function, including additional renal management for HIV-1 infected patients. PL section 2 and 4 where the information on how to detect early signs and symptoms of renal toxicity is also included. Additional risk minimisation measure: Educational materials: HIV pediatric renal educational brochure

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 Emtricitabine/Tenofovir disoproxil Krka.

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

There are no studies required for Emtricitabine/Tenofovir disoproxil Krka.