



EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

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Pharmacovigilance Risk Assessment Committee (PRAC)

PRAC recommendations on signals

Adopted at the 23-26 October 2017 PRAC meeting

This document provides an overview of the recommendations adopted by the Pharmacovigilance Risk Assessment Committee (PRAC) on the signals discussed during the meeting of 23-26 October 2017 (including the signal European Pharmacovigilance Issues Tracking Tool [EPITT]¹ reference numbers).

PRAC recommendations to provide supplementary information are directly actionable by the concerned marketing authorisation holders (MAHs). PRAC recommendations for regulatory action (e.g. amendment of the product information) are submitted to the Committee for Medicinal Products for Human Use (CHMP) for endorsement when the signal concerns Centrally Authorised Products (CAPs), and to the Co-ordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh) for information in the case of Nationally Authorised Products (NAPs). Thereafter, MAHs are expected to take action according to the PRAC recommendations.

When appropriate, the PRAC may also recommend the conduct of additional analyses by the Agency or Member States.

MAHs are reminded that in line with Article 16(3) of Regulation No (EU) 726/2004 and Article 23(3) of Directive 2001/83/EC, they shall ensure that their product information is kept up to date with the current scientific knowledge including the conclusions of the assessment and recommendations published on the European Medicines Agency (EMA) website (currently acting as the EU medicines webportal).

For CAPs, at the time of publication, PRAC recommendations for update of product information have been agreed by the CHMP at their plenary meeting (6-9 November 2017) and corresponding variations will be assessed by the CHMP.

For nationally authorised medicinal products, it is the responsibility of the National Competent Authorities (NCAs) of the Member States to oversee that PRAC recommendations on signals are adhered to.

Variations for CAPs are handled according to established EMA procedures. MAHs are referred to the available [guidance](#). Variations for NAPs (including via mutual recognition and decentralised procedures) are handled at national level in accordance with the provisions of the Member States.

¹ The relevant EPITT reference number should be used in any communication related to a signal.



The timeline recommended by PRAC for submission of variations following signal assessment is applicable to both innovator and generic medicinal products, unless otherwise specified.

For procedural aspects related to the handling of PRAC recommendations on signals (e.g. submission requirements, contact points, etc.) please refer to the [Questions and Answers on signal management](#).

1. Recommendations for update of the product information

None

2. Recommendations for submission of supplementary information

INN	Signal (EPITT No)	PRAC Rapporteur	Action for MAH	MAH
Efavirenz; tenofovir disoproxil; emtricitabine; efavirenz, emtricitabine, tenofovir disoproxil	Autoimmune hepatitis (18956)	Ana Sofia Diniz Martins (PT)	Supplementary information requested (submission by 3 January 2018)	Gilead Sciences Limited, Bristol-Myers Squibb, Merck Sharp & Dohme Limited
Eltrombopag	Laboratory test interference, interference with bilirubin assay (18955)	Eva Segovia (ES)	Assess in the next PSUR (submission by 9 December 2017)	Novartis Europharm Ltd
Rivaroxaban	Oesophagitis (18954)	Qun-Ying Yue (SE)	Assess in the next PSUR (submission by 24 November 2017)	Bayer AG

3. Other recommendations

INN	Signal (EPITT No)	PRAC Rapporteur	Action for MAH	MAH
Amitriptyline	Drug induced liver injury (DILI) and hepatocellular injury (18890)	Agni Kapou (GR)	Routine pharmacovigilance	MAHs of amitriptyline-containing products
Levonorgestrel intrauterine device (IUD)	Anxiety, panic attacks, mood changes, sleep disorders and restlessness (18849)	Martin Huber (DE)	Routine pharmacovigilance	MAHs of levonorgestrel-releasing intrauterine devices
Levonorgestrel intrauterine device (IUD)	Arthralgia (19109)	Martin Huber (DE)	Routine pharmacovigilance	MAHs of levonorgestrel-releasing intrauterine devices