8 May 2023[1]
EMA/PRAC/164741/2023
Pharmacovigilance Risk Assessment Committee (PRAC)

PRAC recommendations on signals

Adopted at the 11-14 April 2023 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 11-14 April 2023 (including the signal European Pharmacovigilance Issues Tracking Tool [EPITT][2] 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 (24-26 April 2023) 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] Expected publication date. The actual publication date can be checked on the webpage dedicated to PRAC recommendations on safety signals.
[2] The relevant EPITT reference number should be used in any communication related to a signal.

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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

Not applicable

2. Recommendations for submission of supplementary information

INN	Signal (EPITT No)	PRAC Rapporteur	Action for MAH	MAH
Acetazolamide	Choroidal effusion and choroidal detachment (19924)	Ulla Wändel Liminga (SE)	Supplementary information requested (submission by 28 June 2023)	Amdipharm Limited
Apalutamide	Interstitial lung disease (ILD) (19911)	Tiphaine Vaillant (FR)	Assess in the next PSUR (submission by 24 April 2023)	Janssen-Cilag International NV
Elasomeran (COVID-19 mRNA vaccine) - Spikevax	Pemphigus and pemphigoid (19860)	Marie Louise Schougaard Christiansen (DK)	Assess in the next PSUR (submission by 26 August 2023)	Moderna Biotech Spain, S.L.
Enoxaparin	Angiokeratoma (19909)	Menno van der Elst (NL)	Assess in the next PSUR (submission by 2 July 2024)	MAHs for enoxaparin with obligation to submit PSURs
Glucagon-like peptide-1 (GLP-1) receptor agonists: dulaglutide; exenatide; insulin degludec, liraglutide; liraglutide; lixisenatide; insulin glargine, lixisenatide semaglutide	Thyroid cancer (18292)	Mari Thorn (SE)	Supplementary information requested (submission by 26 July 2023)	Novo Nordisk A/S, AstraZeneca AB, Eli Lilly Nederland B.V., Sanofi Winthrop Industrie
Megestrol	Meningioma (19923)	Eamon O'Murchu (IE)	Supplementary information requested (submission by 28 June 2023)	Bausch Health Ireland Limited, Teofarma S.R.L., Pharmaswiss Česká Republika S.R.O.

INN	Signal (EPITT No)	PRAC Rapporteur	Action for MAH	MAH
Pramipexole	Intestinal obstruction (19898)	Anette Kirstine Stark (DK)	Assess in the next PSUR (submission by 15 June 2023)	Boehringer Ingelheim International GmbH
Tozinameran (COVID-19 mRNA vaccine) - Comirnaty	Pemphigus and pemphigoid (19859)	Menno van der Elst (NL)	Assess in the next PSUR (submission by 27 August 2023)	BioNTech Manufacturing GmbH

3. Other recommendations

INN	Signal (EPITT No)	PRAC Rapporteur	Action for MAH	MAH
COVID-19 vaccine (ChAdOx1-S [recombinant]) - Vaxzevria	Pemphigus and pemphigoid (19858)	Jean-Michel Dogné (BE)	Monitor in PSURs	AstraZeneca AB
Evolocumab	Weight increase and abnormal weight gain (19867)	Kimmo Jaakkola (FI)	Routine pharmacovigilance	Amgen Europe B.V.