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Committee for Medicinal Products for Human Use (CHMP)

Scientific conclusions and grounds recommending the variation to the terms of the marketing authorisation

International non-proprietary name: canagliflozin, metformin

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Scientific conclusions

Taking into account the PRAC Assessment Report on the PSUR for canagliflozin, metformin, the scientific conclusions of CHMP are as follows:

Scientific conclusions and grounds for variation to the terms of the marketing authorisations

Following a request from PRAC, the MAH conducted a search of the clinical trial database identified 69 cases of serious adverse events of renal failure in 65 subjects (17 on canagliflozin 100 mg, 13 on canagliflozin 300 mg, 16 on comparator and 19 still on blinded treatment). Additionally, an analysis of post-marketing data for renal failure and renal impairment events included 382 serious cases involving the use of canagliflozin, and 1 serious case involving the use of canagliflozin/metformin. For completeness, 117 non-serious cases for canagliflozin and 2 non-serious cases for canagliflozin/metformin were also reviewed. Although this events occurred via volume depletion in most of the cases other mechanisms may be involved. Hence, the PRAC agreed with the MAH's proposal to include information on these events in the product information.

Therefore, in view of available data regarding canagliflozin and canaglifozin/metformin, the PRAC considered that changes to the Product Information of medicinal products containing canagliflozin and canaglifozin/metformin were warranted.

The CHMP agrees with the scientific conclusions made by the PRAC.

Grounds for the variation to the terms of the Marketing Authorisation

On the basis of the scientific conclusions for canagliflozin, metformin the CHMP is of the opinion that the benefit-risk balance of the medicinal products containing canagliflozin, metformin is favourable subject to the proposed changes to the product information

The CHMP recommends that the terms of the Marketing Authorisations should be varied.