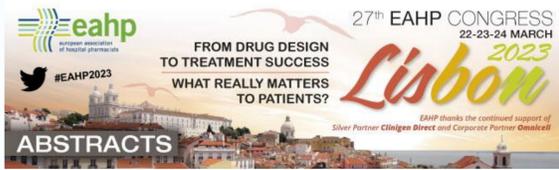


# MEDICATION-RELATED OSTEONECROSIS OF THE JAWS AND CDK4/6 INHIBITORS

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## BACKGROUND AND RELEVANCE

5PSQ-026

Medication-related osteonecrosis of the jaw (MRONJ) is a relatively uncommon but serious complication of osteoclast inhibitors therapy with intravenous bisphosphonates and denosumab. Dose, schedule, and duration of inhibition are associated with MRONJ risk. Marcianò et al.<sup>1</sup> launched an alert about a possible association between MRONJ and cyclin-dependent kinase (CDK)4/6 inhibitors in breast cancer patients with osteoclast inhibitors therapy.

## AIM AND OBJETIVES

Evaluate the use of CDK4/6 inhibitors as a risk factor for MRONJ in our cohort of patients with metastatic cancer and denosumab

## MATERIALS AND METHODS

Retrospective observational study. All patients with denosumab (January 2011-February 2022) were included. Cases of MRONJ found were described. Relationship between CDK4/6 inhibitors and MRONJ was analysed with a Chi-square analysis.

## RESULTS

363 patients with denosumab were included. 21 cases of MRONJ were detected: 62.5% women, 57.1%(12/21) with breast cancer, 19%(4/21) prostate cancer, and 9.5%(2/21) lung cancer. 42.9% with extraosseous metastases. Median treatment duration for denosumab was 19 months (1-52). 7 with a CDK4/6 inhibitors (3 palbociclib, 2 abemaciclib and 2 ribociclib). Median treatment duration with CDK4/6 inhibitors was 27 months (10-35). The mean time from the start of denosumab to the appearance of the event was 23 months (16-29). Incidence of this complication in patients treated with denosumab but without CDK4/6 inhibitors was 5.24% (14/267) and 7.29% (7/96) in patients with denosumab and a CDK4/6 inhibitor. Although the group with CDK4/6 inhibitors had a higher incidence of MRONJ cases, the difference was not significant (0.461).

## CONCLUSION AND RELEVANCE

The incidence of MRONJ in our cohort of patients with metastatic cancer and denosumab was higher in the group of patients with CDK4/6 inhibitors. However, this difference was not significant. Our data are somewhat higher than those reported in the literature according to which the risk of MRONJ with denosumab is 1.1% during the first year, 3.7% the second year and 4.6% per year thereafter. Studies with more patients would be necessary to confirm the relationship between the use of CDK4/6 inhibitors and MRONJ.

## REFERENCES

1. Marcianò, A.; Guzzo, G.M.; Peditto, M.; Picone, A.; Oteri, G. Medication-Related Osteonecrosis of the Jaws and CDK4/6 Inhibitors: A Recent Association. *Int. J. Environ. Res. Public Health* 2020, 17, 9509.

