

Appropriate use of bone protective agents in hormone sensitive prostate cancer

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INTRODUCTION:

- Bone protective agents (BPAs) treat metastatic castration resistant prostate cancer (mCRPC)
- **Zoledronic acid** and **denosumab** improve progression free survival (PFS), overall survival (OS), and rates of skeletal related adverse events (SREs) in men with mCRPC¹
- Patients with metastatic hormone sensitive prostate cancer (mHSPC) may not have similar SRE or OS benefit, as observed in CALGB 90202/Alliance and STAMPEDE trials^{2,3}
- Men with mHSPC and concomitant osteopenia or osteoporosis can receive BPA at reduced dosing intervals:
 - Zoledronic acid 5 mg every 12 months
 - Denosumab 60 mg every 6 months
- During COVID-19, infusion capacity was limited to reduce viral transmission and for COVID-19 treatment with monoclonal antibodies

OBJECTIVE:

- To identify the rate of inappropriate BPA use in mHSPC and quantify its effects in terms of unnecessary infusion visits and added cost

METHODS:

- Pharmacists reviewed the treatment indication for every patient receiving **zoledronic acid** or **denosumab** over a 1-year period
 - For men with mHSPC and osteopenia/osteoporosis:
 - **Zoledronic acid** 5 mg annually or **denosumab** 60 mg every 6 months was considered appropriate
 - For men with mCRPC or hypercalcemia:
 - **Zoledronic acid** 4 mg every 12 weeks was considered appropriate
- Medication costs were calculated using institutional data
- Infusion visits were calculated based on the drug administration schedule



Inappropriate bone protective agent use leads to increases in unnecessary infusion clinic visits and medication-related costs

BPA As Prescribed	mHSPC and Low Bone Density	mHSPC and Normal Bone Density	Annual Reduction in Infusion Visits*
Zoledronic acid 4 mg every 12 weeks	4 patients	1 patient	16 visits (20→4)
Denosumab 120 mg every 4 weeks	8 patients	10 patients	226 visits (234→8)

*with BPA conversion to zoledronic acid 5 mg annually



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RESULTS:

- Data was collected from September 2020 to September 2021
- For men with mHSPC:
 - 5 patients received **zoledronic acid** 4 mg every 12 weeks
 - 4 patients met indication for **zoledronic acid** 5 mg annually
 - 1 patient did not meet an indication for BPA use
 - 18 patients received **denosumab** 120 mg every 4 weeks
 - 8 patients eligible for conversion to **zoledronic acid** 5 mg annually
 - 10 patients did not meet an indication for BPA use
- Switching patients to a more appropriate BPA dosing schedule is estimated to:
 - Reduce total infusion visits from 254 to 12
 - Increase cost savings of > **\$165,000**

CONCLUSION:

- A total of 23 men at our single healthcare institution received off-guideline BPA medications
- Implementation of pharmacist oversight is associated with a reduction of 242 infusion visits per year and significant cost savings

DISCUSSION:

- Future directions include providing education for systemwide pharmacists and oncologists in the Intermountain Healthcare System on proper BPA use and dosing schedules and a systemwide analysis on impact of infusion visits and medication cost savings

REFERENCES:

1. Fizazi K, Carducci M, Smith M, et al. Denosumab versus zoledronic acid for treatment of bone metastases in men with castration-resistant prostate cancer: a randomised, double-blind study. *Lancet*. 2011 Mar 5;377(9768):813-22. doi: 10.1016/S0140-6736(10)62344-6. Epub 2011 Feb 25. PMID: 21353695; PMCID: PMC3090685.
2. Smith MR, Halabi S, Ryan CJ, Hussain A, Vogelzang N, Stadler W, Hauke RJ, Monk JP, Saylor P, Bhoopalam N, Saad F, Sanford B, Kelly WK, Morris M, Small EJ. Randomized controlled trial of early zoledronic acid in men with castration-sensitive prostate cancer and bone metastases: results of CALGB 90202 (alliance). *J Clin Oncol*. 2014 Apr 10;32(11):1143-50. doi: 10.1200/JCO.2013.51.6500. Epub 2014 Mar 3. PMID: 24590644; PMCID: PMC3970172.
3. James ND, Sydes MR, Clarke NW, Mason MD, Dearnaley DP, Anderson J, Popert RJ, Sanders K, Morgan RC, Stansfeld J, Dwyer J, Masters J, Parmar MK. Systemic therapy for advancing or metastatic prostate cancer (STAMPEDE): a multi-arm, multistage randomized controlled trial. *BJU Int*. 2009 Feb;103(4):464-9. doi: 10.1111/j.1464-410X.2008.08034.x. Epub 2008 Oct 8. PMID: 18990168.

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