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Research Article

Bone-targeted agent use for bone metastases from breast cancer and prostate cancer: A patient survey



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ABSTRACT

Background: In order to design studies assessing the optimal use of bone-targeted agents (BTAs) patient input is clearly desirable.

Methods: Patients who were receiving a BTA for metastatic prostate or breast cancer were surveyed at two Canadian cancer centres. Statistical analysis of respondent data was performed to establish relevant proportions of patient responses.

Results: Responses were received from 141 patients, 76 (53.9%) with prostate cancer and 65 (46.1%) with breast cancer. Duration of BTA use was < 3 months (15.9%) to > 24 months (35.2%). Patients were uncertain how long they would remain on a BTA. While most felt their BTA was given to reduce the chance of bone fractures (77%), 52% thought it would slow tumour growth. Prostate patients were more likely to receive denosumab and breast cancer patients, pamidronate. There was more variability in the dosing interval for breast cancer patients. Given a choice, most patients (49–57%) would prefer injection therapy to oral therapy (21–23%). Most patients (58–64%) were interested in enrolling in clinical trials of de-escalated therapy.

Conclusion: While there were clear differences in the types of BTAs patients received, our survey showed similarity for both prostate and breast cancer patients with respect to their perceptions of the goals of therapy. Patients were interested in participating in trials of de-escalated therapy. However, given that patients receive a range of agents for varying periods of time and in different locations (e.g. hospital vs. home), the design of future trials will need to be pragmatic to reflect this.

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1. Introduction

Despite the widespread use of bone-targeted agents in the care of patients with bone metastases from prostate and breast cancer, questions regarding the optimal; choice of agent, dose, dosing frequency, and duration of therapy remain unanswered [1–6]. Our research group is interested in designing and performing pragmatic de-escalation trials in both breast and prostate cancer patients. However, before undertaking this research, it is important to survey potential patients regarding this type of study with

respect to their understanding of their disease and use of their bone-targeted therapies (i.e. denosumab, zoledronate, pamidronate or clodronate). The information obtained will facilitate appropriate pragmatic trial designs, which will positively impact patients' potential willingness to participate.

This report summarizes the conduct and findings of a survey designed to collect information from Canadian patients with breast or prostate cancer regarding their views of their current treatment regimen. In addition, their opinions regarding the possibility of less frequent treatment administrations and participation in potential research studies to assess the effectiveness of this treatment option were also sought. While we are aware of similar patient surveys around the use of bone-targeted agents for osteoporosis [7] and multiple myeloma [8], we are not aware of published work in patients with metastatic bone disease from solid tumors.

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2. Materials and methods

2.1. Survey development

The survey was designed to collect specific information about the use of current bone-targeted therapy and patients' perception of the goals of this treatment and alternative treatment schedules. Second, the survey gathered information regarding their perspectives on the concept of de-escalated bisphosphonate therapy, designed to help the study team develop an understanding of how patients would value a change in their therapy. Our survey was conceived and developed using an iterative approach by the research team with backgrounds in oncology and epidemiology, including expertise in survey design. The survey is provided in a supplemental appendix.

2.2. Study population

Patients attending outpatient clinics at two Canadian cancer centres (The Ottawa Hospital Cancer Centre in Ottawa and the Sunnybrook Odette Cancer Centre in Toronto, Canada) who were receiving or had recently been receiving bone-targeted therapy for bone metastases from either prostate or breast cancer were approached. The survey was available in both paper and online formats (implemented via www.fluidsurveys.com), whichever the patient preferred to use, and took about 10 min to complete. The survey was open to patients seen between August 1st and October 31st, 2012. Local institutional research ethics board (REB) approval for the study was granted at both participating hospitals. However, given the REB requirement that no personal patient identifiers could be collected, we were unable to collect information on the number of patients who took the survey away from clinic but then chose not to complete it. As a result, a response rate cannot be calculated.

2.3. Statistical analysis

The survey consisted of close-ended multiple-choice and hybrid questions (i.e. choose-one-and specify) which were analyzed using a descriptive summary of findings in the form of frequencies and percentages using Microsoft Excel 2010 (Microsoft Corporation, Seattle, Washington) and SAS software (version 9.2, SAS, Cary, North Carolina). There were no formal hypotheses tested using the data. This data was used to qualitatively judge the respondents' views of their current treatments and to determine their willingness to be treated with de-escalated bone-targeted therapy to allow the study team to clarify the value and feasibility of a future trial around this clinical question. Analyses were performed to present the overall response profile, as well as responses within the breast cancer and prostate cancer subgroups. Bar plots were generated to help present categorical data findings.

3. Results

3.1. Respondent population characteristics

A total of 141 patients responded, and included totals of 65 (46.1%) and 76 (53.9%) patients diagnosed with breast cancer and prostate cancer, respectively (Table 1). All patients had bone metastasis. A total of 100% of breast cancer patients and 94.7% of prostate cancer patients were aware that they had received treatment with a bone-targeted agent at some point. Respondents had received bone-targeted agents for: < 3 months (17.0%), between 3 and 12 months (37.6%), between 12 and 24 months (22.0%), and > 24 months (19.1%); 4.3% of patients did not

Table 1
Overview of respondent population.

Respondent characteristics	Distribution of responses		
	Breast cancer	Prostate cancer	Overall
Number of respondents	65 (46.1%)	76 (53.9%)	141
Cancer spread to bones?			
Yes	65 (100%)	76 (100%)	141 (100%)
No	0	0	0
Previous bone targeted agent therapy?			
Yes	65 (100%)	72 (94.7%)	137 (97.2%)
No	0 (0%)	3 (3.9%)	3 (2.1%)
No response	0	1 (1.3%)	1 (0.7%)
Duration of bone targeted agent therapy?			
< 3 months	8 (12.3%)	16 (21.1%)	24 (17.0%)
3–12 months	21 (32.3%)	32 (22.7%)	53 (37.6%)
12–24 months	14 (21.5%)	17 (22.4%)	31 (22.0%)
> 24 months	22 (33.8%)	5 (6.6%)	27 (19.1%)
No response	0 (0%)	6 (7.9%)	6 (4.3%)
Expectations, bone targeted agent treatment duration?			
2 years	0 (0%)	0 (0%)	0 (0%)
As long as I am well enough	5 (7.7%)	2 (2.6%)	7 (5.0%)
As long as my doctor prescribes	30 (46.2%)	31 (40.8%)	61 (43.2%)
Unknown	29 (44.6%)	34 (44.7%)	63 (44.7%)
Other	0 (0%)	1 (1.3%)	1 (0.1%)
No response	1 (1.5%)	8 (10.5%)	9 (6.4%)

respond. Patients varied in the amount of time for which they expected to remain on therapy: 44.7% did not know for how long, 43.2% indicated as long as their doctor keeps prescribing it, 5.0% believed it to be as long as they remain well enough to attend their centre to receive it, and 6.4% did not respond. These response patterns were similar for breast cancer and prostate cancer patients with a few small differences noted (Table 1).

3.2. Current treatment

Among prostate cancer patients, the most commonly administered agent was denosumab (63.1%, as well as an additional 5.3% who in the past had received denosumab), followed by zoledronic acid (19.7%) (Fig. 1). Overall 44.7% of patients received their treatment at the hospital chemotherapy unit, while 27% received treatment via injections administered at home by a nurse or another individual (Fig. 2). A total of 31 patients (40.8% of all prostate cancer patients, or 45.6% of those responding to the related question) reported being aware of potential side effects of their treatment. According to patients' descriptions, the potential side effects of bone targeted agents were; diarrhoea, constipation, vomiting, fatigue, flu-like symptoms, osteonecrosis of the jaw, hypocalcaemia, bone pain and water retention.

Within the breast cancer cohort, the most commonly used agent was pamidronate (87.7%), followed by zoledronic acid (9.2%; Fig. 1). Overall, 93.8% of patients reported receiving their treatments in the chemotherapy unit at their hospital (Fig. 2). Overall, 34 patients (52.3%) were receiving their treatment every 3 months (Fig. 3). A total of 28 patients (43.1%) reported awareness of potential side effects, similar to those described by prostate cancer patients.

3.3. Patient's viewpoints regarding the indication for current bone-targeted treatment

The most commonly reported reasons were: to reduce the chance of bone fractures due to cancer (78.0%), to slow the growth of cancer (53.2%), to reduce the chance of spinal damage (49.0%), to prolong survival (46.8%), and to reduce the chance of cancer pain (46.1%) (Fig. 4). Fewer patients indicated that it was related

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