

# Preferences and Priorities for Relapsed Multiple Myeloma Treatments Among Patients and Caregivers in the United States

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**Introduction/Background:** This study aimed to describe patient and caregiver preferences for treatments of relapsed or refractory multiple myeloma (MM).

**Materials and Methods:** A survey including discrete-choice experiment (DCE) and best-worst scaling (BWS) exercises was conducted among US patients with relapsed or refractory MM and their caregivers. The DCE included six attributes with varying levels including progression-free survival (PFS), toxicity, and mode and frequency of administration. In addition, the impact of treatment cost was assessed using a fixed-choice question. The BWS exercise included 18 items (modes and frequency of administration, additional treatment convenience, and toxicity items). The survey was administered online to patients recruited from the Multiple Myeloma Research Foundation CoMMpass study (NCT01454297).

**Results:** The final samples consisted of 94 patients and 32 caregivers. Avoiding severe nerve damage was most important to patients, followed by longer PFS. Caregivers considered PFS to be the most important attribute. We estimate that a third or more of patients were cost-sensitive, meaning their treatment preference was altered based on cost implications. Caregivers were not cost-sensitive. The three most bothersome treatment features in the BWS exercise were risk of kidney failure, lowering white blood cell counts, and weakening the immune system.

**Conclusion:** Patients with relapsed or refractory MM and their caregivers consider many factors including efficacy, toxicity, mode/frequency of administration, and cost in their decisions regarding treatment options. The study provides a basis for future Research on patient and caregiver treatment preferences, which could be incorporated into shared decision-making with physicians.

**Keywords:** discrete choice experiment, preference, patient, caregiver, multiple myeloma

## Introduction

Multiple myeloma (MM), a neoplasm of the plasma cell affecting an estimated 130,000 US individuals, is the second most common hematologic malignancy.<sup>1,2</sup> Despite recent advancements in therapeutic options, MM remains incurable and is associated with considerable symptom burden and impairments in health-related quality of life (HRQoL).<sup>3</sup> In particular, MM and its treatment can cause debilitating complications, including bone pain and fracture, renal insufficiency, cardiovascular complications such as venous thromboembolism and anemia, and peripheral neuropathy.<sup>4-7</sup>

Patients with relapsed or refractory MM are faced with a variety of viable therapeutic options, such that patients and physicians must together consider tradeoffs among the attributes related to efficacy, tolerability, and other aspects of HRQoL of potential therapies in making treatment decisions that reflect patients' preferences, values, and priorities. However, relatively little is known about the preferences of patients and their caregivers in terms of myeloma treatments. Historically, there has been a disconnect between the preferences of physicians and their patients,<sup>8</sup> and better defining patients' preferences can help to bridge that gap.

In addition to the patients themselves, it is important to understand the preferences of caregivers for those with MM, friends or family members who provide physical and/or emotional care, who also bear the burden of MM and often have a key role in treatment decision-making. Due to the level of symptomology related to MM and the relatively high treatment burden,<sup>9</sup> caregivers for patients with MM are even more involved in patient care than other malignancies.<sup>10</sup> But little is known about their preferences for MM treatment.

The objective of this study was to describe the preferences for treatments for relapsed or refractory MM among patients and their caregivers.

## Materials and Methods

A web-based discrete-choice experiment (DCE) and best-worst scaling (BWS) survey was employed to quantitatively assess patients' preferences over a set of MM treatment attributes. Computer tablets were provided to participants.

## Study Population

Convenience samples of patients and caregivers were recruited from sites participating in the Multiple Myeloma Research Foundation CoMMpass study (NCT01454297), a longitudinal study that has enrolled 1150 patients with MM across 91 sites in the United States (US), Canada, and Europe to evaluate clinical parameters, HRQoL, and health care resource utilization over at least 5 years of treatment.<sup>11</sup> For this survey, patients were invited from four CoMMpass sites to complete the online preference survey: Virginia Cancer Specialists, Arlington, Virginia; Washington University School of Medicine, St. Louis, Missouri; Mount Sinai Hospital, New York, New York; and Hackensack University Medical Center, Hackensack, New Jersey. The CoMMpass Patient Support Center also reached out directly to CoMMpass patients to invite them to take the survey. The survey was programmed and administered online by GfK, a market research firm.

Eligible respondents for the patient survey, participating in the CoMMpass study at the time of recruitment, had a physician-confirmed diagnosis of MM, and had been documented as having at least one relapse. Eligible respondents for the caregiver survey were self-reported caregivers, of a person who met the criteria of the patients recruited for the study but not necessarily of patients surveyed within the study. Eligible respondents were adults aged 18 years or older, residents of the US, able to read and understand English, and able to provide informed consent.

## Survey Instrument

The survey instrument was developed following good research practices.<sup>12</sup> Attribute selection and descriptions for both the DCE and BWS exercises were informed by four focus groups conducted with convenience samples of a total of 12 patients with MM and 11 caregivers of patients with MM. Focus group participants were recruited by Snow Companies, a patient-to-patient education, awareness, and word-of-mouth marketing agency. The focus groups identified efficacy, mode and frequency of administration, and several side effects as the most important treatment features. From the list generated in the focus groups, clinicians and treating physicians on the Multiple Myeloma Research Foundation Steering Committee provided input on the attributes and attribute descriptions for the DCE based on their experience and the data available from clinical trials for the therapies of interest. The attributes selected for the DCE reflected clinically relevant endpoints that were identified as important by patients and caregivers. The levels within each attribute were selected based on results from previous clinical trials for comparability with currently available treatment profiles of proteasome inhibitors (bortezomib, carfilzomib, and ixazomib) immunomodulatory drugs (lenalidomide, pomalidomide), and monoclonal antibodies (elotuzumab and daratumumab). However, none of the agents were modeled specifically within the DCE or BWS instrument. Some attributes such as gastrointestinal toxicities nausea, vomiting, diarrhea, and constipation,

and route and frequency of administration were combined as levels within an attribute. This does limit the interpretation of the relative importance of each attribute/level to improve the efficiency of the DCE as the preference weights calculated from the DCE, as discussed further in the statistical analysis section, pertain to the attribute as a whole and not for individual levels of the attribute. However, this was done to improve efficiency and power.

To pretest the survey instrument for comprehension and completeness, face-to-face, semistructured qualitative interviews were conducted with convenience samples of 31 patients and 24 caregivers. The first 20 patients and 9 caregivers were recruited through Hackensack University Medical Center, Virginia Cancer Specialists, and the Washington University School of Medicine. Final pretests with 11 patients and 15 caregivers were recruited through L&E Research. Based on participant feedback, changes were made to the survey text, questions, and attribute and level descriptions. Such changes included clarifying clinical concepts (eg, progression-free survival [PFS], symptoms experienced at MM diagnosis) to be presented in patient-friendly terms, specifying treatment frequency in the descriptions of treatment characteristics, refining the descriptions of the risks of low platelets and low white blood cell counts, and adding questions about patient demographics in the caregiver questions. The data from the focus groups and pretests of the survey instruments were not included in the analysis.

In each DCE question, respondents were asked to choose between two hypothetical, experimentally designed treatments. [Figure 1A](#) presents an example DCE question from the patient survey, [Figure 1B](#) an example of Best-Worst Scaling Question, and [Supplementary Table 1A](#) presents the treatment attributes, attribute levels, and descriptions of the attribute levels provided to respondents evaluated in the DCE. A fractional factorial experimental design was constructed with Sawtooth software using a D-efficient algorithm,<sup>13–17</sup> which was evaluated for level balance and correlation. The full fractional design contained 40 DCE questions, which were used to create five blocks of eight DCE questions each. Respondents were randomly assigned to one of these blocks.

To capture information about the role of cost in treatment choice, a fixed-choice question was shown to respondents after the conclusion of the experimentally determined series of DCE questions. The fixed-choice question was presented in the same format as the DCE questions but asked respondents to choose between two medicine profiles where out-of-pocket cost was added to the attributes in the DCE (see [Supplementary Table 1A](#)).

The focus groups identified more treatment features than could be accommodated in the DCE. To provide an additional source of information on treatment preferences for a larger set of outcomes, a BWS exercise assessing additional treatment features also was included in the survey. Respondents were given three medication attributes and were asked to choose the one that bothered them the least (Best) and the one that bothered them the most (Worst). [Supplementary Table 1B](#) presents an example BWS question from the patient survey. The BWS exercise included 18 items (the modes and frequency of administration included in the DCE, additional treatment convenience items, mild and serious adverse events, and treatment side effects). The experimental design for the BWS questions was also created using Sawtooth software. The full BWS experimental design contained 45 questions, which were divided into five blocks of nine questions. Respondents were randomly assigned to one of the five blocks.

In addition to the DCE and BWS questions, the survey instrument included questions about respondents' demographic characteristics, patient experiences with MM or caregiver experiences caring for a patient with MM. Comprehension questions testing respondents' understanding of the treatment attributes and levels were also included.

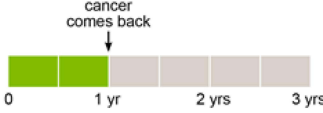
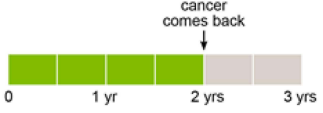




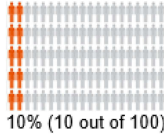
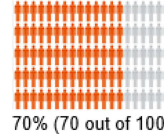
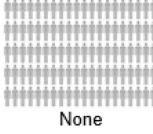
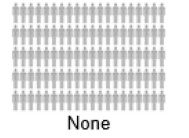

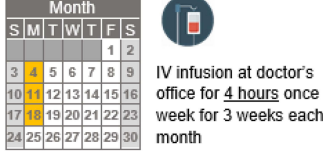
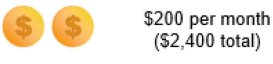
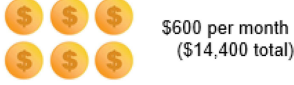
## Statistical Analyses

The analysis sample included all respondents who provided consent and answered at least 1 DCE choice question. Responses to the survey items and respondent characteristics were summarized descriptively.

A random-parameters logit (RPL) regression model was used to analyze the patient choice data collected in this DCE. RPL models avoid potential estimation bias from unobserved preference heterogeneity.<sup>18,19</sup> A conditional multinomial logit regression model was used to analyze the caregiver choice data collected in this due to the small number of caregivers in the sample.

Although the conditional logit model does not control for unobserved preference heterogeneity, the model is recommended and is widely used for DCE analysis.<sup>20</sup>

**A**

Medicine Characteristic	Medicine A	Medicine B
Time until your multiple myeloma comes back or starts advancing again		
Nerve damage		
Stomach problems		
Risk of low blood counts		
Risk of heart failure		
How you take the medicine		
Out-of-pocket cost to you		
Which medicine would you choose?	<input type="checkbox"/>	<input type="checkbox"/>

**B**

I would be bothered by this the MOST	Medicine Characteristics	I would be bothered by this the LEAST
<input type="radio"/>	Treatment weakens your immune system and you get sick more often	<input type="radio"/>
<input type="radio"/>	Having trouble working at your job or at home during treatment	<input type="radio"/>
<input type="radio"/>	Treatment damages kidneys and increases the risk of kidney failure	<input checked="" type="radio"/>

**Figure 1** Example questions from the patient survey. **(A)** Example DCE question with fixed-profile choice question from the patient survey. **(B)** Example best-worst scaling question from the patient survey.

The RPL and conditional logit models yield preference weights that can be interpreted as the relative strength of preference for each attribute level. The scale of these weights is arbitrary and is not directly interpretable, but more-preferred outcomes have higher preference weights. The preference weights can be used to calculate the conditional relative importance estimates for the features associated with MM medicines over the range of levels for each attribute. A conditional relative importance estimate for a feature is the difference between the most-preferred level of that feature and the least-preferred level of that feature. Conditional relative importance scores were scaled so that the most important attribute is equal to 10, and all other scores are relative to that of the most important attribute.

To evaluate the impact of cost on the selected set of treatment profiles, the preference weights from the RPL and conditional logit models were used to estimate the probability that a respondent would select each treatment profile when cost was not included. The predicted probability of selecting each profile in [Figure 1A](#) was compared with the percentage of the sample that selected each profile when cost was included in the fixed-choice question.

A relative score was calculated for each item in the BWS exercise by subtracting the number of times a feature was chosen as least bothersome from the number of times it was chosen as most bothersome, then dividing by the total number of times the feature appeared in the design. The score reflects both the overall ranking of the BWS items and the degree of bothersomeness of each item relative to the other items.

## Results

### Study Population

The final samples consisted of 94 patient respondents with MM and 32 caregivers of individuals with MM (but not necessarily of patients surveyed). Patients were an average age of 65 years old, and 59% were male ([Table 1](#)); caregivers were an average age of 63 years old, and 53% were female ([Table 2](#)).

### Attribute Importance

The mean preference weight estimates for each attribute level are presented in [Supplementary Figure 1](#). Of the attributes and their respective levels presented in the survey, the most important to both patients and caregivers were severe nerve damage and PFS ([Figure 2](#)). Patients considered avoiding severe nerve damage to be most important of the two. Caregivers considered the increase PFS to be most important, although the differences between nerve damage and PFS were not statistically significant for caregivers. The least important attribute to patients was risk of heart failure. The least important attribute to caregivers was gastrointestinal problems. The results also suggest patients and caregivers may prefer oral drugs over other modalities.

### Impact of the Cost on Preference

In the fixed-choice question ([Figure 1A](#)), Medicine A had lower PFS than Medicine B but also had a lower risk of low blood counts, was administered as a weekly pill, and cost \$400 less per month than Medicine B. Using the RPL and conditional logit models to predict the probability that respondents would select Medicine A or Medicine B when cost was not included, the average patient respondent had a 21.8% (95% CI, 4.0–39.7%) chance of selecting Medicine A. Based on this, we would have expected that approximately 4 to 38 patients would select Medicine A if cost data was not included. However, 62 patients (68.1% [95% CI, 58.7–77.6%]) chose Medicine A when cost was included. This suggests that cost influenced the decision of some patients. In the caregiver survey, we would have expected that 12 to 23 (64.9% (95% CI, 25.8–100.0%)) would have chosen Medicine A if cost data was not included. When cost was included, 17 caregivers (54.8% [95% CI, 37.3–72.4%]) chose Medicine A. Therefore, increased cost did not seem to influence the decision of caregivers.

### Most Bothersome Features

Patients and caregivers both considered features that represented treatment-related risks to be among the most bothersome features in the BWS exercise. For patients ([Table 3](#)), the three most bothersome treatment features were kidney damage, reduced white blood cell counts, and a weakened immune system. For caregivers ([Table 4](#)), the three most

**Table I** Respondent Characteristics - Patients

Question	Statistic or Category	Total (N = 94)
Age (years)	Mean (SD)	64.9 (9.8)
	Number over 55 years old	79 (84.0%)
	Minimum, maximum	32, 88
Gender	Female	39 (41.5%)
Race	American Indian or Alaska Native	1 (1.1%)
	Asian Indian	1 (1.1%)
	Black or African American	9 (9.6%)
	Chinese	1 (1.1%)
	Filipino	1 (1.1%)
	White	77 (81.9%)
	Other	1 (1.1%)
	Prefer not to share	5 (5.3%)
Ethnicity	Hispanic or Latino	1 (1.1%)
	Not Hispanic or Latino	88 (93.6%)
	Prefer not to share	5 (5.3%)
Current employment status	Employed full-time	24 (25.5%)
	Employed part-time for reasons not related to MM	4 (4.3%)
	Employed part-time because of MM	5 (5.3%)
Do the following people help take care of you on a daily basis?	My spouse or partner	48 (51.1%)
	My child	8 (8.5%)
Recruitment site	CoMMpass Patient Support Center	25 (26.6%)
	Virginia Cancer Specialists	20 (21.3%)
	Washington University School of Medicine	19 (20.2%)
	Mount Sinai Hospital	20 (21.3%)
	Hackensack University Medical Center	10 (10.6%)
Treatment experience	Currently receiving treatment	82 (87.2%)
Years since diagnosis	Mean (SD)	5.3 (3.38)

bothersome treatment features were kidney damage, weakened bones, and reduced white blood cell counts. Patients and caregivers considered features that were modes of administration to be among the least bothersome features in the BWS given the set of features included.

## Discussion

Recent legislation in the US promotes the inclusion of patient preferences in decision-making at all stages of drug development, the regulatory approval process, and shared decision-making with physicians.<sup>21,22</sup> With this goal in mind,

**Table 2** Respondent Characteristics - Caregivers

Question	Statistic or Category	Total (N = 32)
Age (years)	Mean (SD)	62.5 (12.8)
	Number over 55 years old	24
	Minimum, maximum	29, 83
Gender	Female	17 (53.1%)
Race	American Indian or Alaska Native	1 (3.1%)
	Asian Indian	1 (3.1%)
	Black or African American	3 (9.4%)
	White	26 (81.3%)
	Other	1 (3.1%)
	Prefer not to share	1 (3.1%)
Ethnicity	Hispanic or Latino	2 (6.3%)
	Not Hispanic or Latino	28 (87.5%)
	Prefer not to share	2 (6.3%)
Current employment status <sup>a</sup>	Employed full-time	11 (35.5%)
	Employed part-time for reasons not related to MM	3 (9.7%)
	Employed part-time because of MM	0
How would you describe your relationship with the person you are caring for?	I am caring for my spouse or significant other	25 (78.1%)
	Other	7 (21.9%)
Recruitment site	CoMMpass Patient Support Center	2 (6.3%)
	Virginia Cancer Specialists	8 (25.0%)
	Washington University School of Medicine	9 (28.1%)
	Mount Sinai Hospital	9 (28.1%)
	Hackensack University Medical Center	4 (12.5%)

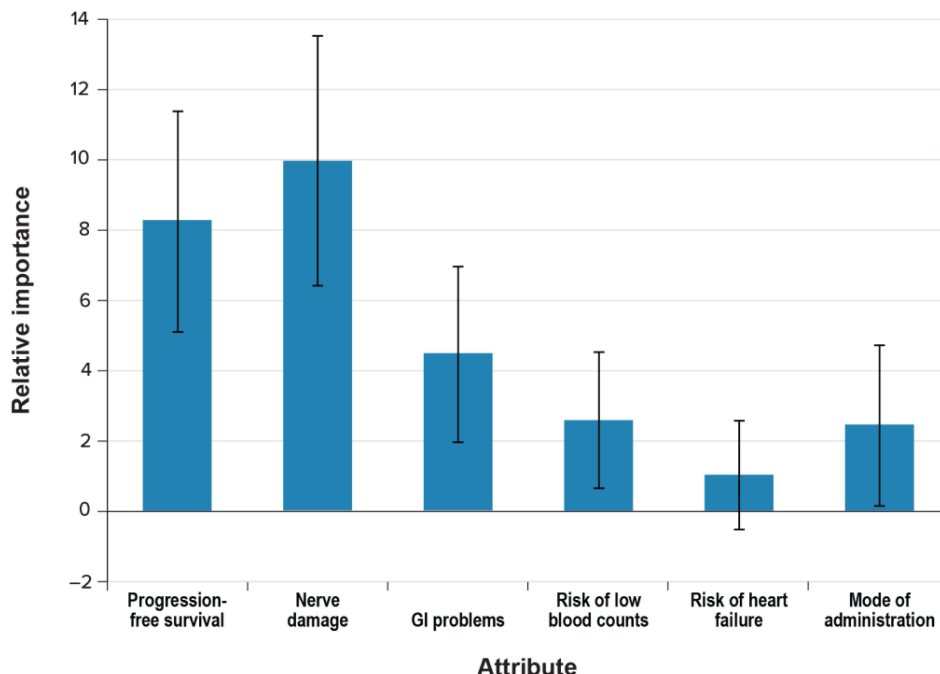
**Note:** <sup>a</sup>One respondent skipped this question; therefore, percentages are calculated out of total non-missing data.

**Abbreviations:** MM, multiple myeloma; SD, standard deviation.

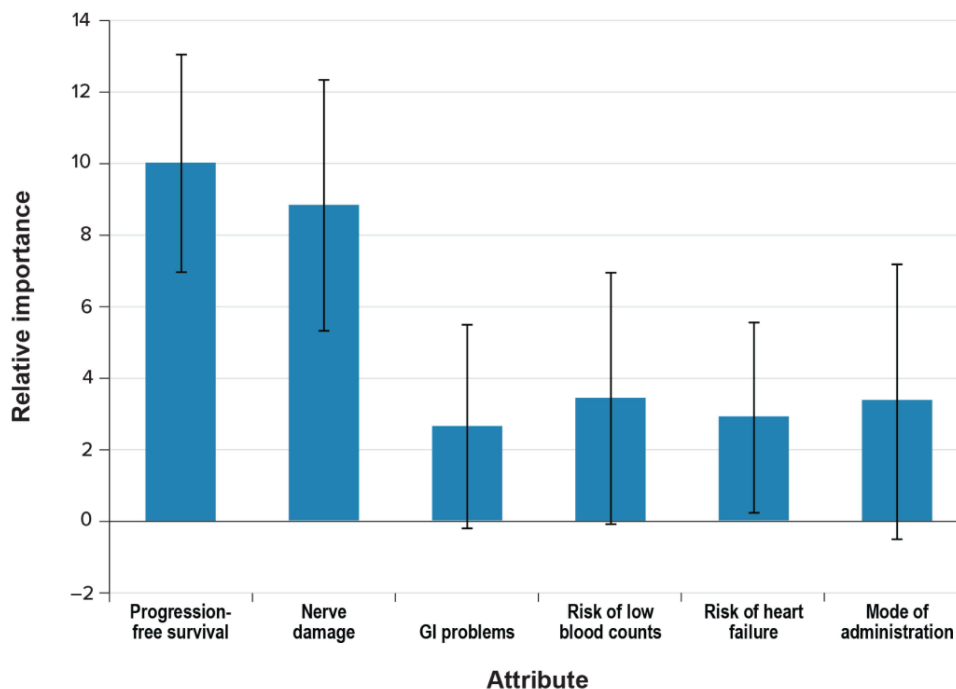
this study aimed to characterize the preferences of patients with relapsed or refractory MM and their caregivers. It found that patients and caregivers placed the most weight on increasing PFS and avoiding severe nerve damage. The relative importance of these two attributes differed between patients and caregivers, with avoidance of severe nerve damage being the most important attribute to patients and prolonged PFS being the most important attribute to caregivers, although the differences in conditional relative importance are not statistically significant. Both patients and caregivers placed more importance on an increase in PFS from 6 months to 12 months than avoiding mild-to-moderate nerve damage. The results indicate that both patients and caregivers make tradeoffs when it comes to treatments to manage relapsed or refractory MM.

Two recent DCE studies examined preferences for MM treatment among patients in the UK and Australia.<sup>23,24</sup> In both studies, patients placed significantly more importance on overall survival (OS) than other attributes. However, the

**A Patients (N = 94)**



**B Caregivers (N = 32)**



**Figure 2** Conditional relative importance of a change in each attribute from the least-preferred level to the most-preferred level. **(A)** Patients. **(B)** Caregivers. **Notes:** The vertical bars surrounding each mean preference weight denote the 95% confidence interval of the point estimate. The levels of each attribute were as follows—progression-free survival: 6 months, 12 months, or 24 months; nerve damage: none, mild to moderate, or severe; GI problems: none, nausea and vomiting, diarrhea, constipation; risk of low blood counts: none, 10%, 25%, 70%; risk of heart failure: none, 3%, 5%; mode of administration (each for 3 weeks each month): daily pill, weekly pill, weekly injection; weekly 4-hour IV infusion, twice-weekly 1-hour IV infusion. **Abbreviation:** GI, gastrointestinal.



**Table 3** Best-Worst Scaling Results - Patients

Feature	Frequency Picked Most Bothersome	Frequency Picked Least Bothersome	Number of Times Item Was Shown	Relative Score
<b>Patients (N = 94)</b>				
Treatment damages kidneys and increases the risk of kidney failure	111	1	126	0.87
Treatment lowers white blood cell counts and increases the risk of serious infection	82	12	153	0.46
Treatment weakens your immune system and you get sick more often	68	9	136	0.43
Needing to go to the hospital to treat a side effect of the treatment	77	26	129	0.40
Treatment weakens your bones and increases the risk of a bone fracture	60	14	126	0.37
Treatment lowers platelet counts and increases the risk of bleeding problems	51	13	126	0.30
Feeling exhausted or tired most days during treatment	66	24	143	0.29
Having thinking or memory problems for a few weeks after receiving treatment	71	27	156	0.28
Changes in mood that may include lack of patience or nonstop talking for 2 or 3 days after each treatment	51	36	132	0.11
Having trouble working at your job or at home during treatment	39	32	150	0.05
4-hour IV, 1 time per week for 3 weeks out of every month	27	44	146	-0.12
1-hour IV, 2 times per week for 3 weeks out of every month	39	59	143	-0.14
You need to travel 3 hours roundtrip once a week to receive your treatment	36	72	160	-0.23
Difficulty sleeping for 2 or 3 nights after each treatment	36	74	146	-0.26
Experiencing hair loss for several months during treatment	20	69	132	-0.37
Injection at doctor's office 1 time per week for 3 weeks out of every month	11	76	139	-0.47
Pill once a week for 3 weeks out of every month	1	121	139	-0.86
Pill daily taken at home with water	0	136	156	-0.87

levels of the attribute for OS ranged from 1 to 5 years in one study and 1 to 9 years in the other, which may have attributed to its high preference weight.

Preference weight is calculated as the difference between utility in the lowest and highest ranked levels of an attribute. The toxicity attribute only included generic “mild or moderate” versus “severe” side effects at various risk

**Table 4** Best-Worst Scaling Results - Caregivers

Feature	Frequency Picked Most Bothersome	Frequency Picked Least Bothersome	Number of Times Item Was Shown	Relative Score
<b>Caregivers (N = 32)</b>				
Treatment damages kidneys and increases the risk of kidney failure	39	1	44	0.86
Treatment weakens your bones and increases the risk of a bone fracture	26	4	45	0.49
Treatment lowers white blood cell counts and increases the risk of serious infection	28	5	49	0.47
Needing to go to the hospital to treat a side effect of the treatment	26	7	47	0.40
Treatment weakens your immune system and you get sick more often	16	2	41	0.34
Treatment lowers platelet counts and increases the risk of bleeding problems	20	6	44	0.32
Having thinking or memory problems for a few weeks after receiving treatment	22	8	52	0.27
Changes in mood that may include lack of patience or nonstop talking for 2 or 3 days after each treatment	16	10	43	0.14
Feeling exhausted or tired most days during treatment	16	15	53	0.02
4-hour IV, 1 time per week for 3 weeks out of every month	13	14	55	-0.02
Experiencing hair loss for several months during treatment	14	19	45	-0.11
You need to travel 3 hours roundtrip once a week to receive your treatment	14	21	48	-0.15
Having trouble working at your job or at home during treatment	9	17	51	-0.16
1-hour IV, 2 times per week for 3 weeks out of every month	13	22	53	-0.17
Difficulty sleeping for 2 or 3 nights after each treatment	9	24	55	-0.27
Injection at doctor's office 1 time per week for 3 weeks out of every month	5	26	43	-0.49
Pill once a week for 3 weeks out of every month	0	37	43	-0.86

levels from 0 to 60% which we may suggest, from a clinical standpoint, is not on the same scale as a difference of 4 or 8 years in OS. In the current study, we did not include OS as an attribute and limited the absolute difference in the PFS attribute to 18 months largely due to this concern.

In the current study, the observed role of severe nerve damage as a driver of treatment preferences warrants further consideration. Both patients and caregivers were particularly averse to severe nerve damage, which was described to patient and caregiver respondents in the survey as “potentially limiting normal daily activities.” Although avoiding

severe nerve damage was important to both patients and caregivers in the study, it was especially so to patients, presumably because they are averse to the limitations that can result from serious nerve damage or because they wish to avoid burdening caregivers. Nerve damage is common with both proteasome inhibitors and immunomodulatory drugs. Bortezomib has the most associated neuropathy with 16% of those receiving intravenous treatment reporting severe neuropathy. Administered predominantly subcutaneously today, it is associated with a 6% incidence of severe neuropathy.<sup>25</sup> With both administration routes, the nerve damage is largely reversible following discontinuation. Given the importance of severe nerve damage to respondents in this study, future research should explore other dimensions of nerve damage as drivers of treatment preferences, including whether such damage is permanent or temporary.

The results also suggest that patients and, potentially, caregivers both seemed to prefer oral drugs over injections, at least for some dosing schedules. Some studies of cancer therapy preferences that included mode of administration have found that patients prefer oral to intravenous administration, including the two aforementioned DCE studies in MM.<sup>23,24,26–30</sup> Future research should explore how mode of administration influences patient preferences, perhaps separately from assessing benefit-risk tradeoffs between efficacy and adverse events. In such a study, a survey to evaluate treatment satisfaction and burden of illness among patients undergoing treatment for MM found that patients treated with oral therapy reported greater effectiveness, satisfaction, and convenience than patients treated with injectable therapy.<sup>31,33</sup>

The role of cost in patient and caregiver preferences is another interesting finding from this study. Although analyses of the impact of cost on treatment preferences is not as robust as the full DCE, there is a clear trend that suggests that cost is important to some patients when evaluating treatment options. Supporting this is evidence from a qualitative study that found financial impacts of treatment to be among patients' highest priorities for MM treatment.<sup>32</sup>

Interestingly, caregivers did not seem to alter treatment selection based on cost. This could suggest that cost is less important to caregivers than patients, but the small sample size in the caregiver cohort limits the potential generalizability of this finding. However, this finding was echoed in the Australian DCE where out-of-pocket costs were also more important to patients than caregivers.<sup>24</sup> However, it must be noted that out-of-pocket costs were not highly important to either group, ranking behind OS, remission duration, mode/frequency of treatment and severe side effects. In the UK study, out-of-pocket costs was the least important of all attributes.<sup>23</sup> While out-of-pocket cost is an important issue in health care worldwide, the differing results may suggest that cost-of-care is a more important issue for patient with MM in the US than in other countries.

Overall our results are very much in line with a recent study using an unbiased artificial intelligence- supported approach that showed that myeloma patients can be divided into two groups: one favoring PFS while the other preferring QoL (through minimizing toxicities).<sup>34</sup> As proposed by the authors of that study, collection of both PFS and QoL data should both be considered to better understand and account for this heterogeneity in patients' preferences and assist the future treatment decision making process.

Some limitations of this study must be noted. The patient and caregiver samples are subject to selection bias, and the results may not be generalizable. In addition, preferences may vary by whether patients have newly diagnosed MM, have experienced a single relapse, or have experienced multiple relapses. Preferences also likely vary based on cultural norms across various countries and cultures. Further, the sample sizes may be insufficient to detect smaller differences in preferences between attribute levels. Lastly, as with all DCE, we must acknowledge that choices based on hypothetical treatments may not reflect real-world decisions that patients, caregivers, and doctors make.

## Conclusions

Patients with relapsed or refractory MM and their caregivers consider many factors including efficacy, toxicity, mode/frequency of administration, and cost in their decisions regarding treatment options. Discussion with patients and their caregivers about their priorities for treatment and the tradeoffs they are willing to make across different treatment attributes could promote shared decision-making and help improve treatment satisfaction. This work supports the need for larger, well-controlled studies of patient preferences in MM care.

## Acknowledgments

This work was funded in part by Millennium Pharmaceuticals, Inc., a wholly owned subsidiary of Takeda Pharmaceutical International Co., Cambridge, MA. Kimberly Moon of RTI Health Solutions provided project management for the analysis. Kate Lothman of RTI Health Solutions provided medical writing services.

## Disclosure

Daniel Auclair reports grants from Takeda, during the conduct of the study performed at MMRF; he is now an AstraZeneca employee. Carol Mansfield and Brennan Mange are salaried employees of RTI Health Solutions which was contracted by Millennium Pharmaceuticals, Inc., a wholly owned subsidiary of Takeda Pharmaceutical International Co. to assist in the analysis and writing. Craig E Cole reports personal fees and/or grants from Oncopeptides, Sanofi, and GlaxoSmithKline, outside the submitted work. Jonathan L Kaufman reports grants and/or personal fees from Janssen, Amgen, AbbVie, Roche, Sutro, TG therapeutics, Incyte, BMS, and Tecnofarma, outside the submitted work. Jeffrey A Zonder reports personal fees and/or grants from Janssen, Takeda, Regeneron, Amgen, Intellia, Eidos, Alnylam, Oncopeptides, BMS, and Caelum, outside the submitted work. Mehul Dalal is a salaried employee of Millennium Pharmaceuticals, Inc., a wholly owned subsidiary of Takeda Pharmaceutical International Co. Joseph R Mikhael has received honoraria from Amgen, BMS, GSK, Karyopharm, Celgene, Janssen, Sanofi, and Takeda Pharmaceuticals. David S Siegel has received honoraria from Amgen, BMS, GSK, Celgene, Janssen, Karyopharm and Takeda Pharmaceuticals. The authors report no other conflicts of interest in this work.

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