

Real-world Experience With Talquetamab Clinical Management in Relapsed Refractory Multiple Myeloma (RRMM): A Qualitative Study of US Healthcare Providers

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Key Takeaways



Talquetamab SUD was administered in various clinical settings, including inpatient, outpatient, and hybrid models

The 0.8 mg/kg Q2W dosing schedule was most common; however, changing to Q4W (0.8 mg/kg for all but one center, which used 0.4 mg/kg) was reported as a real-world strategy for symptom management among patients who responded to therapy

HCPs adopted a variety of strategies to manage oral, skin, and nail symptoms; prophylactic use of dexamethasone and nystatin mouthwash or zinc and vitamin B complex may be effective strategies to alleviate oral symptoms

Conclusions



Current clinical practices indicated that there is variation in the SUD care setting for talquetamab treatment (half as inpatient and half as outpatient/hybrid models)



Most HCPs used a 0.8 mg/kg Q2W dosing schedule in SUD and treatment phases; for some patients with response to therapy, switching to a Q4W schedule (0.8 mg/kg for all but one center, which used 0.4 mg/kg) is a real-world adverse event management strategy



HCPs recommended symptom management and prophylactic strategies, such as reducing dosing frequency. This is supported by two recent post hoc analyses of MonumentAL-1 that indicated dose modification may help manage GPRC5D-related symptoms after patients achieve response^{4,6}; however, there remain limited real-world data and further evidence is needed to inform the optimal dosing schedule of talquetamab and long-term efficacy and safety



While HCPs provided valuable insights into real-world practices with talquetamab, the small sample size of HCPs and the limited generalizability of qualitative assessments are key limitations of this study

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<https://www.congresshub.com/Oncology/IMS2024/Talquetamab/Schinke-Real-World>

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Introduction

- Talquetamab is a first-in-class, G protein-coupled receptor family C group 5 member D (GPRC5D) × CD3 bispecific monoclonal antibody that showed strong clinical efficacy in heavily pretreated patients with relapsed/refractory multiple myeloma (RRMM) in the phase 1/2 MonumentAL-1 study (ClinicalTrials.gov Identifier: NCT03399799)^{1,2}
- Based on the MonumentAL-1 clinical study data, talquetamab was approved in 2023 in the United States at 2 dosing schedules for patients with RRMM who received ≥4 prior lines of therapy, including a proteasome inhibitor, an immunomodulatory agent, and an anti-CD38 monoclonal antibody: (1) a weekly (QW) dosing schedule that includes 3 step-up doses followed by 0.4 mg/kg QW thereafter; and (2) a biweekly (Q2W) dosing schedule that includes 4 step-up doses followed by 0.8 mg/kg Q2W thereafter³
- Because GPRC5D is a novel target, GPRC5D-related, on-target, off-tumor side effects may be unfamiliar to physicians and patients^{1,4,5}
- Given talquetamab's unique target and recent approval, there are limited real-world data on talquetamab and an unmet need for information on real-world dosing and symptom management

Results

Patient characteristics

- Of the 21 invited HCPs, 11 participated in the study, representing healthcare centers across 9 states (Table 1)
- 10 HCPs took part in the IDIs, and the subsequent expert panel included 6 HCPs (5 from the IDIs and 1 additional expert who was unavailable for participation in the IDIs; Table 1)
 - On average, participants had 10.5 years of experience, collectively treating an average of 100 patients with RRMM per month

Table 1: Interviewee and panelist demographics

Characteristic	Interviewees, n (%) (N=10)	Panelists, n (%) (N=6)
Principal practice setting		
University/academic medical center	10 (100)	6 (100)
HCP role		
Oncologist/hematologist	5 (50)	3 (50)
Nurse practitioner	2 (20)	2 (33)
Clinical oncology pharmacist	2 (20)	1 (17)
Myeloma program director	1 (10)	0
Number of patients treated for RRMM each month		
1-10	0	0
11-50	3 (30)	1 (17)
51-100	3 (30)	3 (50)
101-200	3 (30)	1 (17)
>200	1 (10)	1 (17)
Years of experience		
0-5	3 (30)	2 (33)
6-10	1 (10)	1 (17)
11-15	3 (30)	1 (17)
16-20	3 (30)	2 (33)
Practice's census region		
South	3 (30)	2 (33)
Midwest	3 (30)	2 (33)
West	3 (30)	1 (17)
Northeast	1 (10)	1 (17)

HCP, healthcare provider; RRMM, relapsed/refractory multiple myeloma.

Clinical utility and sequencing

- IDI participants considered various patient factors, such as age, caregiver support, comorbidities, performance status, and disease history, when initiating talquetamab
- Most HCPs (n=7) explicitly stated that they mainly used B-cell maturation antigen (BCMA)-targeted therapies first, followed by talquetamab, but 2 of these HCPs also reported using talquetamab earlier to reserve the BCMA target for chimeric antigen receptor T-cell (CAR-T) therapy
- Most interviewed HCPs (n=8) used talquetamab as a bridging therapy while awaiting CAR-T therapy manufacture, especially for very sick or frail patients or those with aggressive disease; panelists indicated that talquetamab was given for 1 or 2 cycles after cell collection or apheresis and stopped 2 weeks before CAR-T therapy
 - Panelists also reported they would consider giving talquetamab again after CAR-T therapy if patients had previously used talquetamab as bridging therapy to CAR-T therapy
- Talquetamab combined with pomalidomide was noted as a potential option for some patients (especially for those not achieving the expected response of very good partial response); however, there is not yet a uniform practice on this and additional data are needed to inform this treatment approach

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Objective

- To capture real-world practices of US healthcare providers (HCPs) regarding talquetamab clinical utilization patterns, step-up dosing (SUD), treatment dosing schedules, dose modifications, and symptom (oral, skin, and nail) management

Methods

Study design and population

- This study was conducted between February and March of 2024 and employed a protocol-driven, mixed-methods qualitative design with 2 unblinded phases
 - Phase 1: in-depth interviews (IDIs) with HCPs who had early experience administering talquetamab
 - Phase 2: an expert panel of HCPs with similar qualifications to those in the IDIs, with the majority being the same HCPs
- A pool of US healthcare institutions was identified with early experience using talquetamab through preapproval access programs or clinical trials

- From these institutions, 55 HCPs with board certification or specialization in oncology/hematology were selected who had treated patients with RRMM and who had treated patients with talquetamab (via clinical trial or preapproval patient access program)
 - Among these, 21 HCPs from 11 states were contacted via email to participate in 1-hour individual IDIs
 - Convenience sampling was employed to select HCPs who were subsequently recruited based on their interest in participation, and efforts were made to ensure geographic diversity

Data collection

- IDIs were conducted virtually using a standardized discussion guide with open-ended questions about HCPs' experiences treating triple-class-exposed RRMM with talquetamab, alongside a pre-IDI survey for baseline data; recordings were transcribed and analyzed by 2 independent reviewers to identify recurring themes and insights
- After the IDIs, HCPs joined a 2.5-hour virtual expert panel to discuss talquetamab clinical management, using a summary of the insights gained from the IDIs to guide and deepen the conversation

- 5 panelists noted that their institutions were developing talquetamab "tool kits" and symptom management protocols

Table 2: Summary of SUD setting and dosing practices among IDI participants

Dosing practices*	Interviewees, n (%) (N=10)
SUD site of care	
Inpatient	5 (50)
Outpatient	3 (30)
Hybrid	2 (20)
SUD schedule	
Q2W	7 (70)
QW	3 (30)
SUD day schedule	
1-3-5-7	4 (40)
1-4-7	2 (20)
1-3-5-7/8	2 (20)
1-3-5	1 (10)
1-4-7-10/11	1 (10)
Treatment dosing schedule	
Q2W ^b	8 (80)
QW ^c → Q2W ^b	1 (10)
Use of both QW ^c and Q2W ^b	1 (10)
Modifications for patients responding well to treatment	
Has used a Q4W ^d schedule	6 (60)
Has not used a Q4W ^d schedule	4 (40)

HCP, healthcare provider; IDI, in-depth interview; Q2W, once every 2 weeks (biweekly); Q4W, once every 4 weeks; QW, once per week (weekly); SUD, step-up dosing.
*These values represent the number of HCPs who mentioned a given practice during their IDI. More HCPs may have used a strategy in practice but did not mention it during the IDI.
^bQ2W dosing at 0.8 mg/kg.
^cQW dosing at 0.4 mg/kg.
^dQ4W dosing at 0.8 mg/kg for all but 1 center, which used 0.4 mg/kg.

Table 3: Summary of GPRC5D-related symptom management among IDI participants

Symptom management strategies*	Interviewees, n (%) (N=10)
Oral symptom management	
Dietitian	5 (50)
Consider Q4W ^b	5 (50)
Steroid (eg, dexamethasone) mouth rinse	4 (40)
Dietary changes	4 (40)
Biotène	3 (30)
Saliva substitutes	3 (30)
Nystatin (for those with oral thrush from steroid)	1 (10)
Ice packs	1 (10)
Salt solution	1 (10)
Gum/lozenges	1 (10)
AE protocol	1 (10)
Skin symptom management	
Creams (including urea-based creams, moisturizers, and ammonium lactate cream [Amlactin])	8 (80)
Topical steroids	7 (70)
Hydration	3 (30)
Abrasion to help healing	2 (20)
Consider Q4W ^b	2 (20)
Dermatologist referral	1 (10)
Nail symptom management	
No plan	4 (40)
Cosmetics	3 (30)
Nail polish	1 (10)
Gloves	1 (10)
Dermatologist referral	1 (10)

AE, adverse event; GPRC5D, G protein-coupled receptor family C group 5 member D; HCP, healthcare provider; IDI, in-depth interview; Q4W, once every 4 weeks.
*These values represent the number of HCPs who mentioned a given strategy for symptom management during their IDI. There may be additional HCPs who made use of a strategy in practice but did not mention it during the IDI.
^bQ4W dosing at 0.8 mg/kg for all but 1 center, which used 0.4 mg/kg.

Figure 1: Management strategies for symptoms (oral, skin, and nail) from the expert panel

Oral symptoms:

- Dysphagia
 - Prophylactic use of dexamethasone and nystatin mouth rinses 3x/day, starting at SUD and before symptom onset
 - Prophylactic use of zinc and vitamin B complex to delay oral toxicities
 - Ice pack/wrap around cheeks 3x/day to reduce circulation
 - Altering diet to address taste alterations and nutrition consultation if weight loss occurs
- Dry mouth
 - Biotène, saliva alternatives, lozenges, sour citrus, sour candy, and sugar-free candy or gum
- Other considerations
 - Decreasing dose frequency from Q2W to Q4W^b (dose reduction was not found to help)

Skin symptoms:

- Moisturizers and antihistamines for dryness and itchiness, respectively
- Creams, including topical steroids, ammonium lactate cream (Amlactin), and urea-based creams (2x/day) for hands and feet
- Moisturizer brands such as CeraVe, Vanicream, Eucerin, and Cetaphil
- Avoiding contact with alcohols (ie, perfumes/fragrances) that dry out skin
- A thick emollient used on the full body at least once a day, as well as topical steroids
- Avoiding hot showers and applying emollients after drying off from cool showers

Nail symptoms:

- Vitamin E cuticle oil and ammonium lactate (Amlactin) around the nails/cuticles
- Good emollients and low-dose topical steroids for peeling around the nails
- Urea-based treatments on hands and feet (2x/day)
- Wearing cotton gloves with Aquaphor in them overnight to prevent nails falling off
- Wearing soft socks and avoiding trauma to the nails
- Keeping good nail hygiene, including keeping nails short