Cases from the Community: Investigators Discuss Available Research Guiding the Management of Relapsed/Refractory Multiple Myeloma — What Happened at ASH 2025?

A CME/MOC-Accredited Live Webinar

Monday, December 15, 2025 5:00 PM – 6:00 PM ET

Faculty

Sagar Lonial, MD, FACP, FASCO María-Victoria Mateos, MD, PhD

Moderator Neil Love, MD



Faculty



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Chair and Professor
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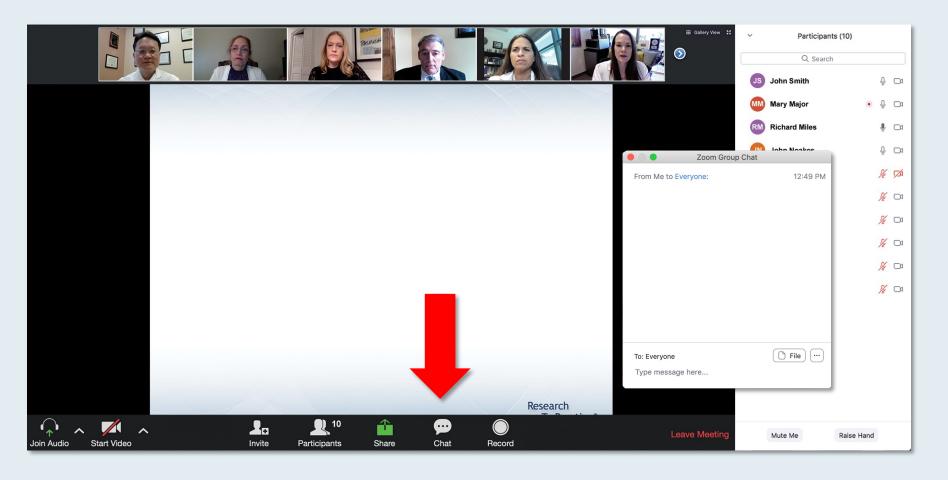
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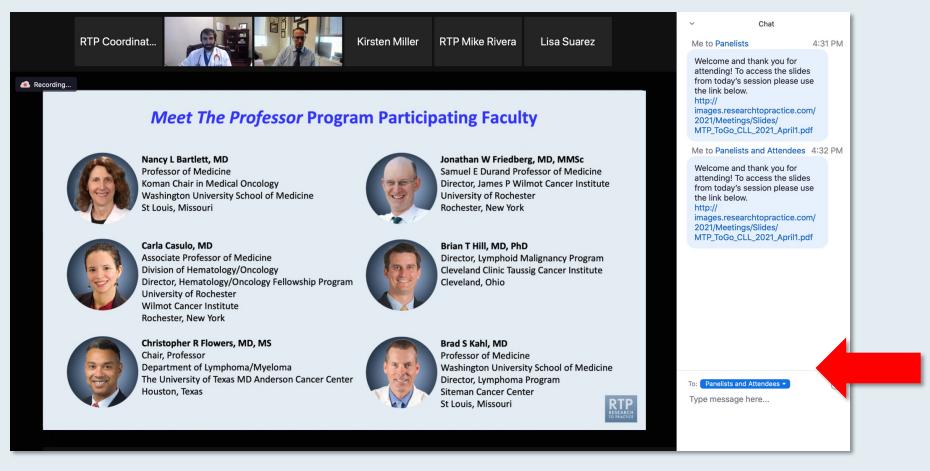


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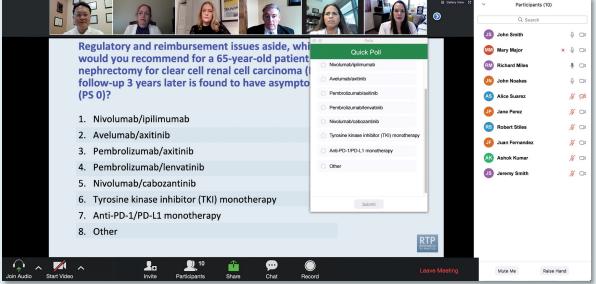


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Relapsed/Refractory Multiple Myeloma — **Proceedings from a Session Held During** the Society of Hematologic Oncology **2025 Annual Meeting**



PROF MELETIOS-ATHANASIOS (THANOS) C DIMOPOULOS, MD ALEXANDRA HOSPITAL



DR NOOPUR RAJE MASSACHUSETTS GENERAL HOSPITAL CANCER CENTER



DR HANS LEE SARAH CANNON RESEARCH INSTITUTE



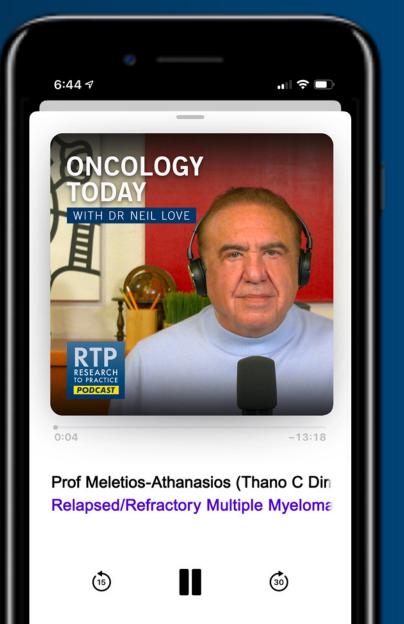


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Practical Perspectives on the Current and Future Management of Immune Thrombocytopenia — What Happened at ASH 2025?

A CME/MOC-Accredited Live Webinar

Tuesday, December 16, 2025 5:00 PM - 6:30 PM ET

Faculty

Hanny Al-Samkari, MD Cindy Neunert, MD, MSCS Professor Francesco Zaja

Moderator Neil Love, MD



Practical Perspectives on the Current Role of Bispecific Antibodies in the Management of Lymphoma — What Happened at ASH 2025?

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Michael Dickinson, MD Laurie H Sehn, MD, MPH

Moderator Neil Love, MD



Expert Second Opinion: Investigators Discuss the Optimal Management of Gastrointestinal Cancers

A CME Symposium Series Held Adjunct to the 2026 ASCO® Gastrointestinal Cancers Symposium

HER2-Positive Gastrointestinal Cancers

Thursday, January 8, 2026

7:15 PM - 8:45 PM PT

(10:15 PM - 11:45 PM ET)

Localized Colorectal Cancer

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7:15 PM - 8:45 PM PT

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Advanced Gastroesophageal Cancers

Friday, January 9, 2026

6:00 PM - 8:00 PM PT

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Grand Rounds

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Optimizing Treatment for Patients with Relapsed/Refractory Chronic Lymphocytic Leukemia

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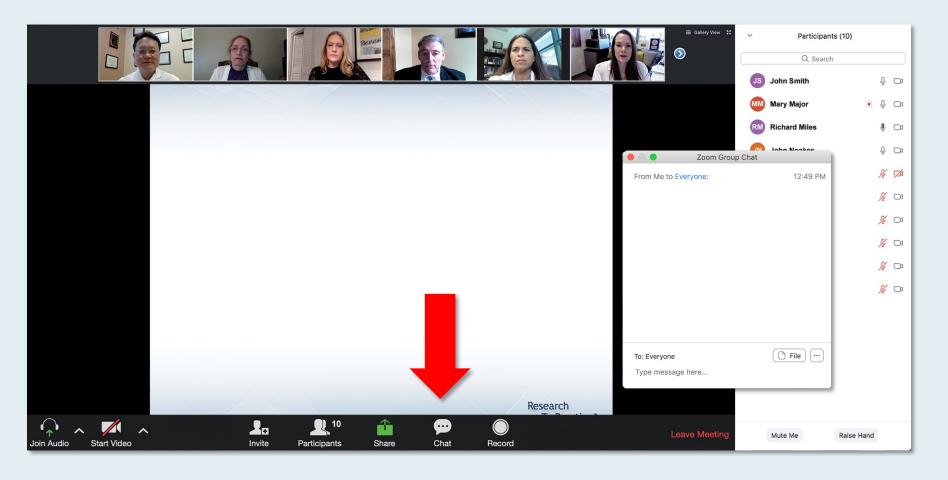
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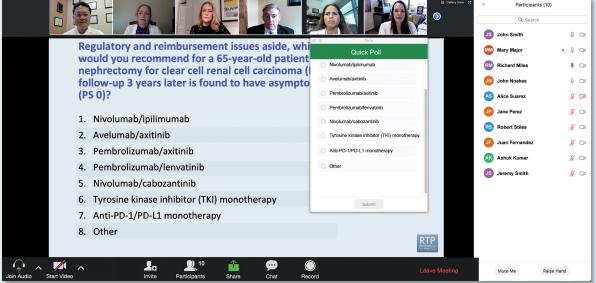


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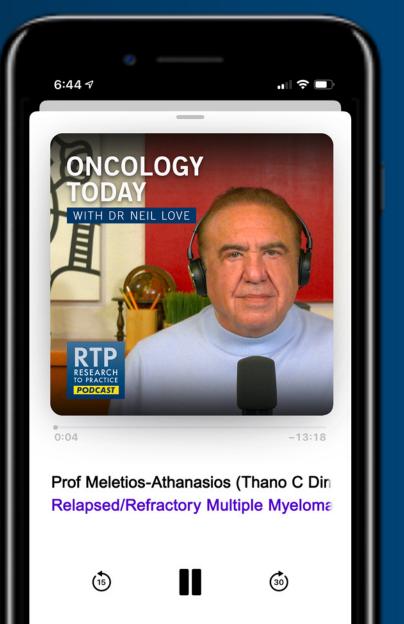


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Sagar Lonial, MD

- Hungria V et al. Long-term responders from the phase 3 DREAMM-7 study of belantamab mafodotin plus bortezomib and dexamethasone vs daratumumab plus bortezomib and dexamethasone in relapsed/refractory multiple myeloma. ASH 2025; Abstract 7236.
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- Lonial S et al. Health-related quality of life with belantamab mafodotin in patients with relapsed or refractory multiple myeloma (RRMM): An exploratory analysis of overall quality of life in DREAMM-7. ASH 2025; Abstract 143.
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Sagar Lonial, MD (continued)

- Landgren O et al. A phase 2 trial of **iberdomide, carfilzomib, daratumumab and dexamethasone** quadruplet therapy for relapsed/refractory multiple myeloma: **The ReKInDLE study**. ASH 2025;Abstract 4108.
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María-Victoria Mateos, MD, PhD

- Costa L et al. Long-term progression-free survival benefit with ciltacabtagene autoleucel in standard-risk relapsed/refractory multiple myeloma. ASH 2025; Abstract 9129.
- Parekh S et al. Earlier use of ciltacabtagene autoleucel (cilta-cel) is associated with better immune fitness
 and stronger immune effects as shown by correlative analysis of peripheral blood and the bone marrow
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- Ailawadhi S et al. KarMMa-3 subgroup analysis in older patients with relapsed/refractory multiple myeloma treated with idecabtagene vicleucel. ASH 2025; Abstract 9044.
- Patel K et al. Phase 2 registrational study of **anitocabtagene autoleucel** for the treatment of patients with relapsed and/or refractory multiple myeloma: **Updated results from iMMagine-1**. ASH 2025;Abstract 4541.
- Ho PJ et al. Minimal residual disease (MRD)-negative outcomes following a novel, in vivo gene therapy
 generating anti—B-cell maturation antigen (BCMA) chimeric antigen receptor (CAR)-T cells in patients with
 relapsed and refractory multiple myeloma (RRMM): Preliminary results from inMMyCAR, the first-in-human
 phase 1 study of KLN-1010. ASH 2025; Abstract LBA-1.



María-Victoria Mateos, MD, PhD (continued)

- Ho PJ et al. Subcutaneous cevostamab demonstrates manageable safety and clinically meaningful activity in relapsed/refractory multiple myeloma (RRMM): First results from the phase Ib CAMMA 3 study. ASH 2025; Abstract 8172.
- Mateos M-V et al. Safety and efficacy of talquetamab + teclistamab in patients with relapsed/refractory multiple myeloma from Phase 1b of RedirecTT-1: Results with an extended median follow-up of 3 years.
 ASH 2025; Abstract 7712.
- Mateos M-V et al. Phase 3 randomized study of teclistamab plus daratumumab versus investigator's choice
 of daratumumab and dexamethasone with either pomalidomide or bortezomib (DPd/DVd) in patients (Pts)
 with relapsed refractory multiple myeloma (RRMM): Results of MajesTEC-3. ASH 2025; Abstract LBA-6.
- Voorhees P et al. Etentamig plus pomalidomide-dexamethasone combination therapy in relapsed or refractory multiple myeloma: A phase 1b dose-escalation and safety expansion study. ASH 2025; Abstract 1875.
- Hamadeh I et al. Low dose tocilizumab for mitigation of cytokine release syndrome with bispecific antibodies in relapsed/refractory multiple myeloma. ASH 2025; Abstract 7258.
- Krishnan A et al. Updated efficacy and safety results of JNJ-5322, a novel, next-generation
 BCMA × GPRC5D × CD3 trispecific antibody, in patients with relapsed/refractory multiple myeloma. ASH 2025; Abstract 7875.



Agenda

Introduction Best of ASH Multiple Myeloma

Case 1 Dr Lorber – 59-year-old man

■ Faculty Presentation: Antibody-Drug Conjugates and Other Emerging Novel
Therapies for Relapsed/Refractory (R/R) Multiple Myeloma (MM) — Dr Lonial

| Case 2 | Dr Morganstein – 84-year-old man |
|--------|----------------------------------|
| Case 3 | Dr Favaro – 64-year-old man |
| Case 4 | Dr Bhatnager – 71-year-old man |
| Case 5 | Dr Lee – 71-year-old man |

■ Faculty Presentation: Integrating CAR T-Cell Therapy and Bispecific Antibodies into the Management of R/R MM — Dr Mateos

Case 6 Dr Rudolph – 56-year-old man



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LBA-1

MRD-negative outcomes following a novel, in vivo gene therapy generating anti-BCMA CAR-T cells in patients with RRMM: Preliminary results from inMMyCAR, the first-in-human Phase 1 study of KLN-1010

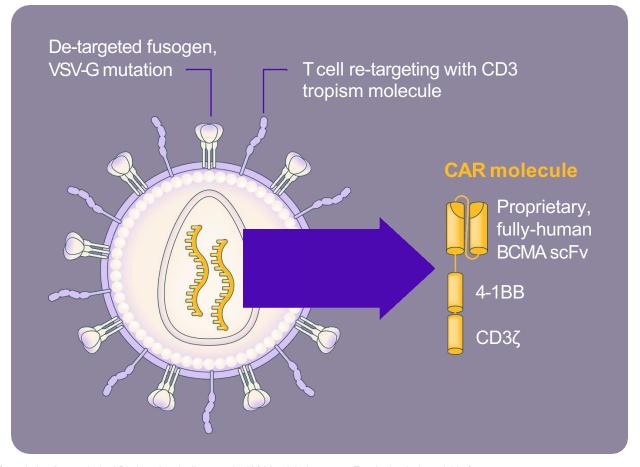
Simon Harrison¹, P. Joy Ho², Sueh-Ii Lim³, Stephanie Talam², Hannah Pahl¹, Dharmesh Dingar⁴, Scott Currence⁴, Travis Quigley⁴, Andrew Spencer³

¹Peter MacCallum Cancer Centre, Melbourne, Victoria, Australia; ²Royal Prince Alfred Hospital, Sydney, New South Wales, Australia; ³The Alfred Hospital, Melbourne, Victoria, Australia; ⁴Kelonia Therapeutics, Inc., Boston, Massachusetts, United States.



KLN-1010: a modified LVV generating anti-BCMA CAR-T cells in vivo

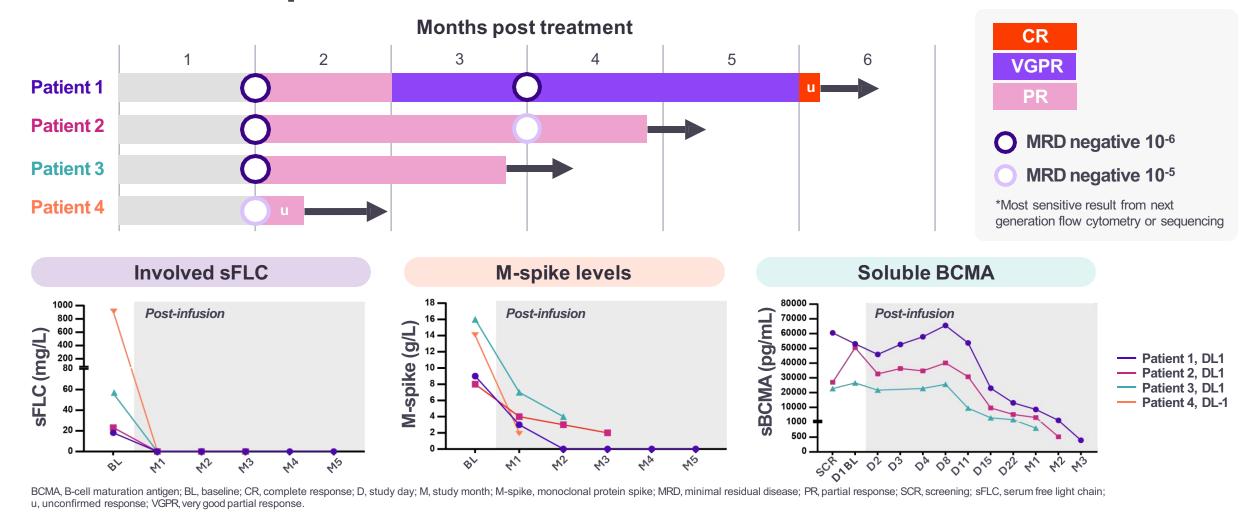
- Envelope-modified, replicationincompetent, self-inactivating lentiviral vector
- De-targeted VSV-G fusogen avoids delivery to LDL-expressing cells while maintaining high transduction efficiency
- Precise re-targeting to T cells with a CD3 scFv; avoids liver uptake and drug sinks
- Anti-BCMA CAR was selected based on high levels of activity to BCMA-positive tumors



BCMA, anti–B-cell maturation antigen; CAR, chimeric antigen receptor; CD3, cluster of differentiation 3; CD3ζ, cluster of differentiation 3 zeta chain; LDL, low-density lipoprotein; LVV, lentiviral vector;scFv, single-chain variable fragment; VSV-G, vesicular stomatitis virus glycoprotein.

Wood JT et al. Toward treatment with gene-modified B cells engineered in vivo using iGPS particles (abstract #1281). Poster presented at: ASGCT 28th Annual Meeting; May 13-17, 2025.

Deep, ongoing MRD-negative responses were observed across first 4 patients



Favorable toxicity profile compared to ex vivo CAR T

- Minimal events of cytopenia; only 1 case of Grade 4 (transient neutropenia related to margination)
- Markedly lower number of events compared to ex vivo CAR-Ttherapies

| TEAEs in >1 patient | Grade 1-2, n | Grade≥3, n |
|---------------------|--------------|------------|
| IRR | 2 | 1 (DLT) |
| Lymphocytosis | 1 | 1 |
| Hypomagnesemia | 2 | 0 |
| Hypokalemia | 2 | 0 |

| TEAEs Grade ≥3 | Patients, n | Study day | Duration, days |
|---------------------|-------------|-----------|----------------|
| Febrile neutropenia | 1 | 1 | 2 |
| IRR | 1 | 1 | 3 |
| Lymphopenia | 1 | 2,8 | 2, 5 |
| Lymphocytosis | 1 | 13 | 3 |
| Anemia | 1 | 15 | 2 |
| Vasovagal syncope | 1 | 27 | 1 |
| Pneumonia | 1 | 86 | 8 |

| Cytopenia | | | |
|------------------|-------------|-----------|----------------|
| | Grade ≥3, n | Study day | Duration, days |
| Anemia | 1 | 15 | 2 |
| Thrombocytopenia | 1 | 16 | 2 |
| Noutropopio | 2 | 1, 15 | 2, 2 |
| Neutropenia | 2 | 14 | 3 |

| Infusion-related reactions | | | |
|----------------------------|--------------|-------------|----------------------|
| | Grade 1-2, n | Grade ≥3, n | Supportive care |
| Dose level 1 | 1 | 1 | Tocilizumab, steroid |
| Dose level -1 | 1 | 0 | Paracetamol |

CAR, chimeric antigen receptor; DLT, dose-limiting toxicity; IRR, infusion-related reaction; TEAE, treatment-emergent adverse event.

Phase 3 Randomized Study of Teclistamab Plus Daratumumab Versus Investigator's Choice of Daratumumab and Dexamethasone With Either Pomalidomide or Bortezomib (DPd/DVd) in Patients With Relapsed Refractory Multiple Myeloma (RRMM): Results of MajesTEC-3

Maria-Victoria Mateos,¹ Nizar J. Bahlis,² Aurore Perrot,³ Ajay K. Nooka,⁴ Jin Lu,⁵ Charlotte Pawlyn,⁶,⁷ Roberto Mina,⁶ Gaston Caeiro,⁶ Alain Kentos,¹⁰ Vania Hungria,¹¹ Donna Reece,¹² Ting Niu,¹³ Anne K. Mylin,¹⁴ Charlotte Toftmann Hansen,¹⁵ Raphael Teipel,¹⁶ Britta Besemer,¹⊓ Meletios A. Dimopoulos,¹ፆ,¹⁰ Elena Zamagni,²⁰,²¹ Satoshi Yoshihara,²² Kihyun Kim,²³ Chang Ki Min,²⁴ Paul Geerts,²⁵ Elena Van Leeuwen-Segarceanu,²⁶ Agata Tyczynska,²⊓ Juan Luis Reguera Ortega,²ፆ Magnus Johansson,²⁰ Markus Hansson,³⁰ Mehmet Turgut,³¹ Mark Grey,³² Surbhi Sidana,³³ Paula Rodriguez-Otero,³⁴ Joaquin Martinez-Lopez,³⁵ Hamza Hashmi,³⁶ Robin Carson,³⊓ Rachel Kobos,³ፆ Weili Sun,³⁰ Kristen Lantz,³⊓ Anne Seifert,⁴⁰ Deborah Briseno-Toomey,⁴¹ Lisa OʻRourke,³⊓ Maria Rubin,³ፆ Diego Vieyra,³⊓ Lijuan Kang,³⁰ Luciano J. Costa⁴²

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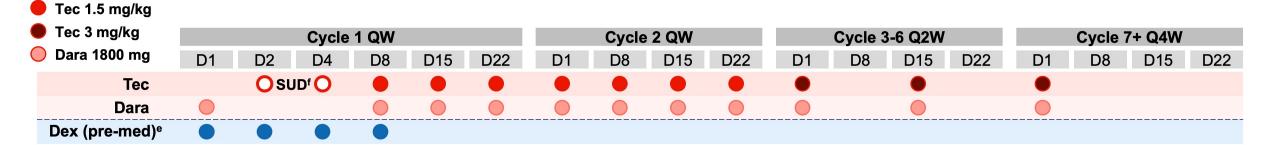
https://www.congresshub.com/ASH2025/Oncology/ Teclistamab/Mateos-LBA

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MajesTEC-3: Phase 3 Study Design

Key inclusion criteria **Primary endpoint** PFS per IRC **Tec-Dara** RRMM • 1-3 prior LOTs including a PI and lenalidomide N = 291**Key secondary endpoints** Patients with only 1 prior LOT must SC dosing following Dara schedule ≥CR^d and ORR^d 1:1 have been lenalidomide refractory per MRD negativity (10⁻⁵) randomization IMWG criteria OS N=587 ECOG PS score of 0-2 MySIm-Q Total Symptom score DPd/DVd 22 Oct 2021 to Key exclusion criteria Other secondary endpoints 29 Sept 2023b N=296 (91% DPd) Prior BCMA-directed therapy Safety by investigator's choice^c Refractory to anti-CD38 mAbsa PK and immunogenicity



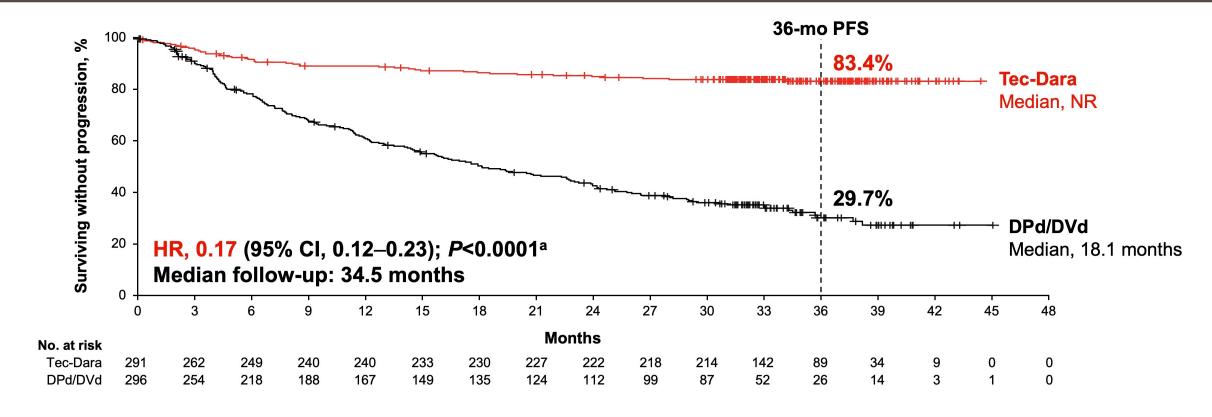
SC dosing aligned with Dara schedule, with monthly dosing after 6 cycles; steroid sparing after Cycle 1 Day 8

^aPrior exposure to anti-CD38 mAbs was permitted. ^bDuring the COVID-19 pandemic. ^cDPd/DVd were administered per the approved schedules. ^dResponse and disease progression were assessed by a blinded IRC per IMWG criteria. ^eDexamethasone, acetaminophen, and diphenhydramine pre-medication was required for the first 2 weeks; subsequent dexamethasone was not required thereafter. ^fPatients received SUD of 0.06 mg/kg and 0.3 mg/kg on Days 2 and 4, respectively.

CR, complete response; Ď, day; Dex, dexamethasone; DPd, daratumumab, pomalidomide, and dexamethasone; DVd, daratumumab, bortezomib, and dexamethasone; ECOG PS, Eastern Cooperative Oncology Group performance status; IMWG, International Myeloma Working Group; IRC, independent review committee; MRD, minimal residual disease; MySIm-Q, Multiple Myeloma Symptom and Impact Questionnaire; ORR, overall response rate; PFS, progression-free survival; PI, proteasome inhibitor; PK, pharmacokinetics; pre-med, pre-medication; QW, weekly; Q2W, every 2 weeks; Q4W, every 4 weeks; SC, subcutaneous; SUD, step-up dosing.



MajesTEC-3: PFS (Primary Endpoint)



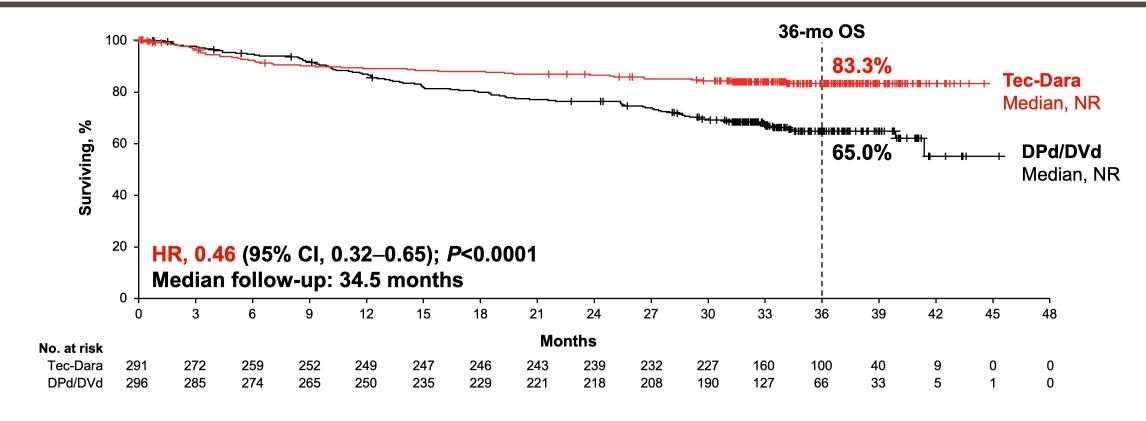
Tec-Dara significantly improved PFS, with a plateauing curve after ~6 months and >90% of patients progression-free at 6 months sustaining such a benefit at 3 years

^aThe *P* value crossed the prespecified stopping boundary for superiority for the first interim analysis (*P*=0.0139). CI, confidence interval; HR, hazard ratio; NR, not reached.





MajesTEC-3: OS



Tec-Dara significantly improved OS versus DPd/DVd, with 83% of patients alive at 3 years



Analysis of RMST demonstrated an OS benefit for Tec-Dara versus DPd/DVd (RMST difference, 2.15 months; *P*=0.0088). RMST, restricted mean survival time.

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Agenda

| Introduction Best of ASH Multiple Myeloma |
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|---|

Case 1 Dr Lorber – 59-year-old man

■ Faculty Presentation: Antibody-Drug Conjugates and Other Emerging Novel
Therapies for Relapsed/Refractory (R/R) Multiple Myeloma (MM) — Dr Lonial

| Case 2 | Dr Morganstein – 84-year-old man |
|--------|----------------------------------|
| Case 3 | Dr Favaro – 64-year-old man |
| Case 4 | Dr Bhatnager – 71-year-old man |
| Case 5 | Dr Lee – 71-year-old man |

■ Faculty Presentation: Integrating CAR T-Cell Therapy and Bispecific Antibodies into the Management of R/R MM — Dr Mateos

Case 6 Dr Rudolph – 56-year-old man





Dr Jeremy Lorber (Beverly Hills, California)

Case Presentation: 59-year-old man with t (11;14) IgA kappa myeloma discovered during workup for new Stage IV kidney disease. Chest wall plasmacytoma. Receives daratumumab with CyBorD and radiation therapy to the plasmacytoma with minimal response



Dr Warren Brenner (Boca Raton, Florida)

Questions for the faculty



Agenda

| Introduction | Best of ASH Multiple Myeloma | |
|---|----------------------------------|--|
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| ■ Faculty Presentation: Antibody-Drug Conjugates and Other Emerging Novel Therapies for Relapsed/Refractory (R/R) Multiple Myeloma (MM) — Dr Lonial | | |
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Case 6 Dr Rudolph – 56-year-old man

Dr Lee - 71-year-old man

Case 5



FDA Approves Belantamab Mafodotin-blmf for Relapsed or Refractory Multiple Myeloma

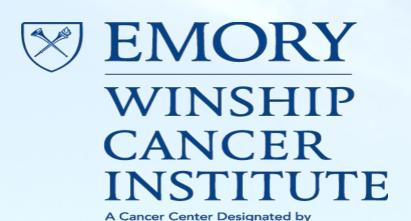
Press Release: October 23, 2025

"The Food and Drug Administration approved belantamab mafodotin-blmf, a B-cell maturation antigen (BCMA)-directed antibody and microtubule inhibitor conjugate, with bortezomib and dexamethasone for adults with relapsed or refractory multiple myeloma who have received at least two prior lines of therapy, including a proteasome inhibitor and an immunomodulatory agent.

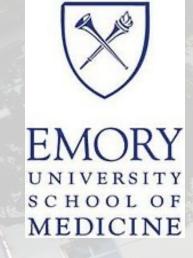
Efficacy was evaluated in DREAMM-7 (NCT04246047), an open-label, randomized, multicenter trial in adults with relapsed or refractory multiple myeloma who had received at least one line of prior therapy. ... Patients were randomized (1:1) to receive either belantamab mafodotin-blmf, bortezomib, and dexamethasone (BVd) or daratumumab, bortezomib, and dexamethasone (DVd).

Efficacy was established based on progression-free survival (PFS) and overall survival (OS). The median PFS was 31.3 months (95% confidence interval [CI]: 23.5, not reached [NR]) in the BVd arm and 10.4 months (95% CI: 7, 13.4) in the DVd arm (hazard ratio [HR] 0.31, 95% CI: 0.21, 0.47). The median OS was NR and 35.7 months (95% CI: 21.1, NR) in respective arms (HR 0.49, 95% CI: 0.32, 0.76)."





the National Cancer Institute



Antibody Drug Conjugates and Other Emerging Novel Therapies for R/R MM

Sagar Lonial, MD
Professor and Chair
Department of Hematology and Medical Oncology
Anne and Bernard Gray Professor in Cancer
Chief Medical Officer, Winship Cancer Institute
Emory University School of Medicine

Long-term Responders From the Phase 3 DREAMM-7 Study of Belantamab Mafodotin Plus Bortezomib and Dexamethasone vs Daratumumab Plus Bortezomib and Dexamethasone in Relapsed/Refractory Multiple Myeloma

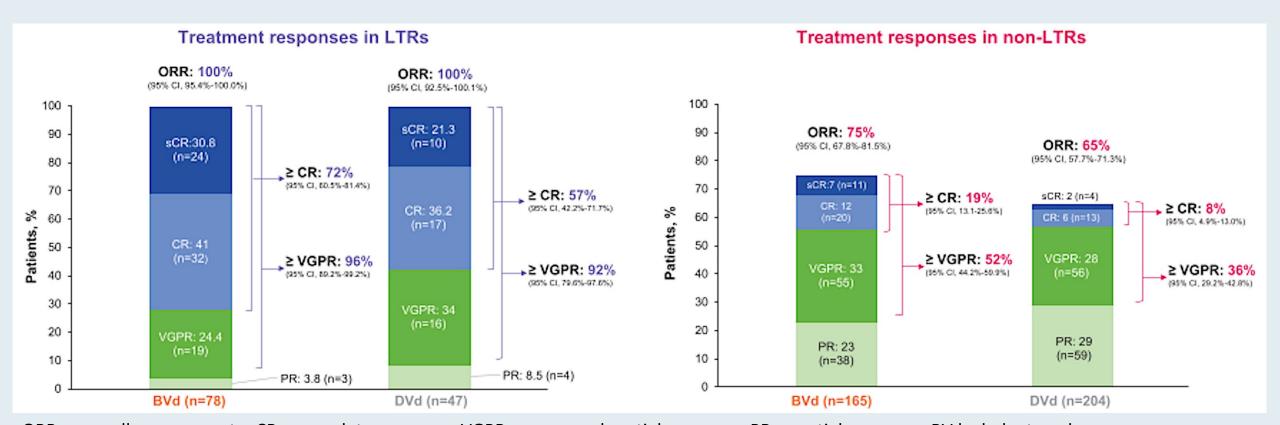
Vania Hungria,¹ Paweł Robak,² Marek Hus,³ Vera Zherebtsova,⁴ Christopher Ward,⁵ P. Joy Ho,⁶ Roman Hájek,⁷ Kihyun Kim,⁸ Sebastian Grosicki,⁹ Hanlon Sia,¹⁰ Adam Bryant,¹¹ Marcelo Pitombeira de Lacerda,¹² Gracia Aparecida Martinez,¹³ Anna Sureda Balarí,¹⁴ Michał Mielnik,³ Maureen Nichols,¹⁵ Jorge Mouro,¹⁶ Zeyad Khalaf,¹⁷ Hena Baig,¹⁸ Margaret Polinkovsky,¹⁹ Nick Pirooz,¹⁹ Sybil Varghese,¹⁹ Joe Lee,¹⁷ Lydia Eccersley,¹⁷ María-Victoria Mateos²⁰

¹Clinica São Germano, São Paulo, Brazil; ²Medical University of Łódź, Poland; ¹Medical University of Lodź, Poland; ¹Medical University of Lodin, Lublin, Poland; ⁴Gorodskaya Klinicheskaya Bol¹nitsa Im. S.p. Botkina, Moscow, Russian Federation; of Sydney, Camperdown, NSW, Australia; ¹Department of Hematooncology, University of Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University, Seoul, Republic of Korea; ¹Medical University of Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University, Seoul, Republic of Korea; ¹Medical University of Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University, Seoul, Republic of Korea; ¹Medical University of Ostrava, Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University, Seoul, Republic of Korea; ¹Medical University of Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University, Seoul, Republic of Korea; ¹Medical University of Ostrava, Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University, Seoul, Republic of Korea; ¹Medical University of Ostrava, Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University of Ostrava, Ostrava, Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University of Ostrava, Ostrava, Ostrava, Ostrava, Czech Republic; ⁵Samsung Medical Center, Sungkyunkwa University of Ostrava, Ostrav

ASH 2025; Abstract 7236.



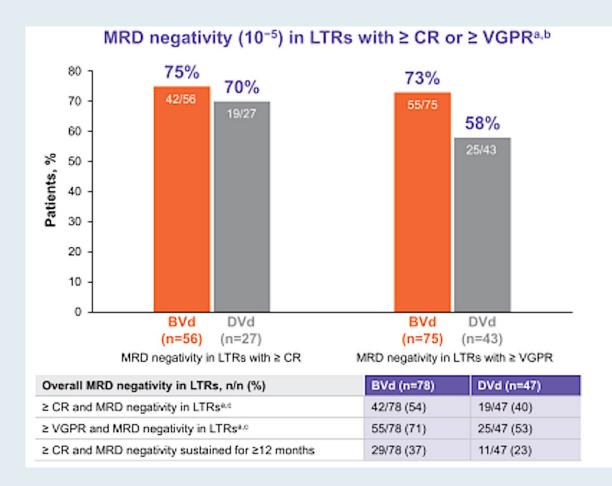
DREAMM-7: Responses in Long-Term Responders (LTRs) and Non-LTRs

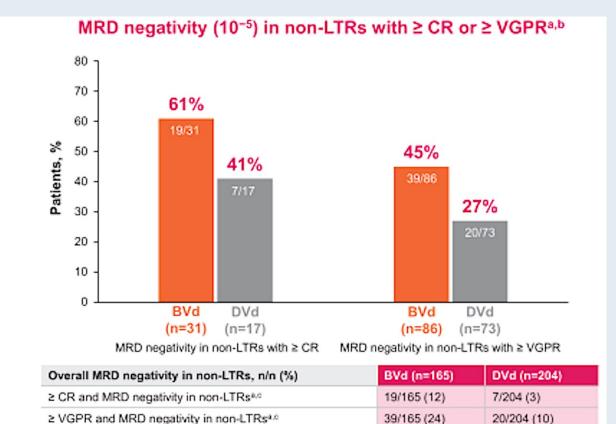


ORR = overall response rate; CR = complete response; VGPR = very good partial response; PR = partial response; BVd = belantamab mafodotin/bortezomib/dexamethasone; DVd = daratumumab/bortezomib/dexamethasone



DREAMM-7: Minimal Residual Disease (MRD) Negativity in Long-Term Responders (LTRs) and Non-LTRs





6/165 (3.6)

0

≥ CR and MRD negativity sustained for ≥12 months



DREAMM-7: Authors' Conclusions



Across both arms, LTRs had slightly different baseline characteristics compared with non-LTRs; baseline characteristics between treatment arms were generally well balanced



Overall, LTRs had fewer prior lines of therapy than non-LTRs, suggesting that earlier treatment with BVd may be associated with long-term clinical benefits



BVd-treated LTRs had deeper and more durable responses compared with DVd-treated LTRs



The presented safety outcomes in LTRs (PFS ≥36 months) were consistent with previous reports in the overall safety population^{3,4}



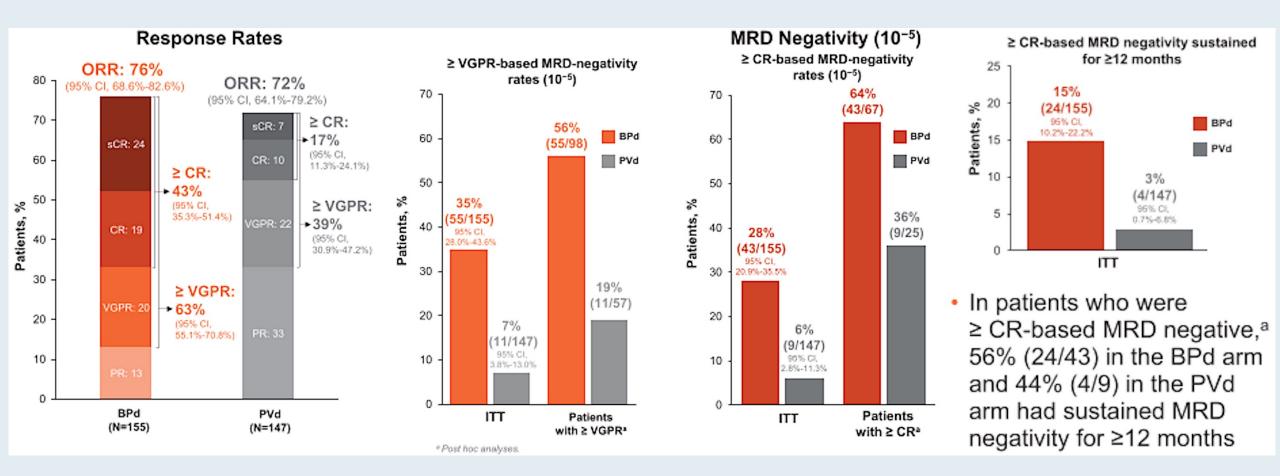
Deep responses and durable outcomes in patients treated with belantamab mafodotin plus pomalidomide and dexamethasone from long-term follow-up of the phase 3 DREAMM-8 study

Suzanne Trudel,¹ Meral Beksaç,² Ludek Pour,³ Sosana Delimpasi,⁴ Vladimir Vorobyev,⁵ Hang Quach,⁶ Ivan Spicka,⁷ Jakub Radocha,⁸ Paweł Robak,⁹ Kihyun Kim,¹⁰ Michele Cavo,¹¹ Kazuhito Suzuki,¹² Syed Zafar,¹³ Ainslee Moore,¹⁴ Marie Duggan,¹⁴ Kristin Morris,¹⁵ Amy Phillips-Jones,¹⁴ Margaret Polinkovsky,¹⁶ Joanna Grams,¹⁷ Ianire Garrobo-Calleja,¹⁸ Elisabet E. Manasanch,¹⁶ Brandon Kremer,¹⁶ Joanna Opalinska,¹⁶ María-Victoria Mateos,¹⁹ Meletios A. Dimopoulos^{20,21}

ASH 2025; Abstract 7677.



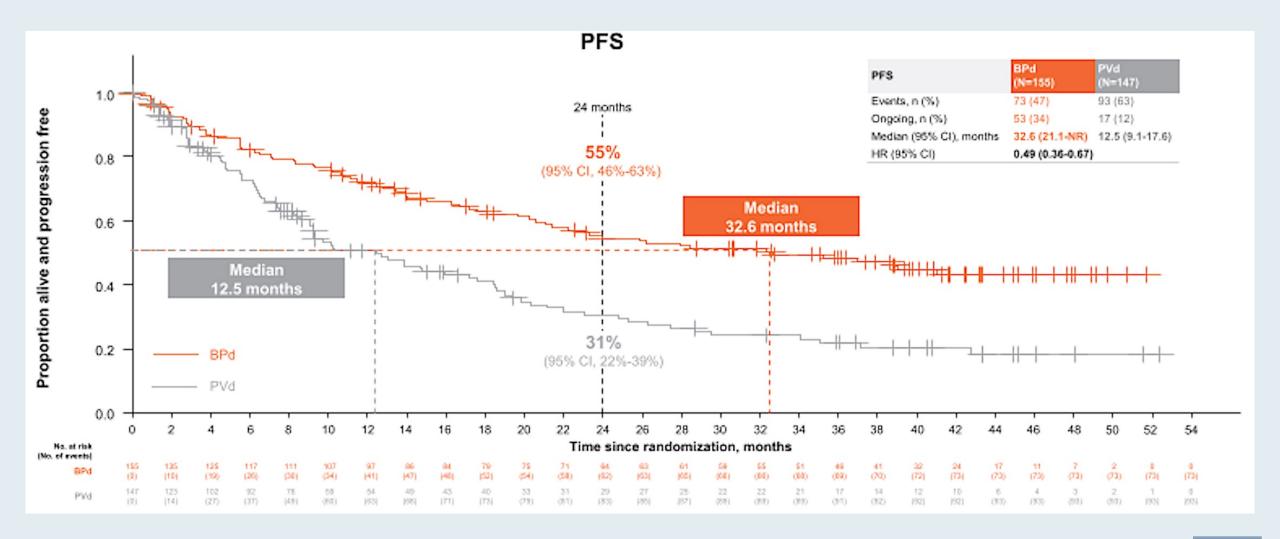
DREAMM-8 Long-Term Follow-Up: Response Rates and MRD Negativity



BPd = belantamab mafodotin/pomalidomide/dexamethasone; PVd = pomalidomide/bortezomib/dexamethasone; ITT = intent to treat



DREAMM-8 Long-Term Follow-Up: Progression-Free Survival (PFS)





DREAMM-8: Authors' Conclusions



BPd led to a 5-fold increase in MRD negativity and sustained MRD negativity vs PVd in patients with RRMM, most of whom had lenalidomide-refractory disease and a quarter of whom had disease refractory to anti-CD38 monoclonal antibodies



3-fold greater median PFS vs PVd (increase of >20 months), with greater than double the benefit following subsequent antimyeloma therapy (median PFS2); follow-up for OS is ongoing



Long-term follow-up from the DREAMM-8 trial demonstrated that BPd maintained superiority over PVd across all efficacy endpoints, including PFS, MRD negativity, sustained MRD negativity, and DOR. Importantly, benefit was maintained following subsequent antimyeloma therapy



Collectively, findings support BPd as a new outpatient and off-the-shelf BCMA treatment as standard of care in MM at first relapse

DOR = duration of response

Trudel S et al. ASH 2025; Abstract 7677.



Health-related quality of life with belantamab mafodotin in patients with relapsed or refractory multiple myeloma: an exploratory analysis of overall quality of life in DREAMM-7

Sagar Lonial,¹ Craig Cole,² Angely Loubert,³ Laurine Bunod,³ Manal Mhari,³ Pralay Mukhopadhyay,⁴ Sandhya Sapra,⁴ Molly Purser,⁴ Jacqueline Nielsen,⁵ Benga Kazeem,⁶ Farrah Pompilus,⁷ Paul G. Richardson⁸

ASH 2025; Abstract 143.

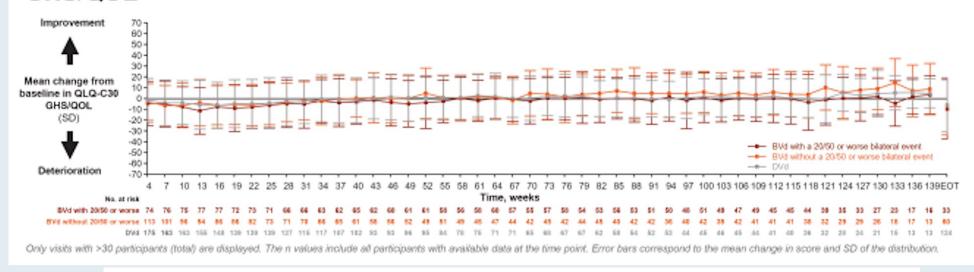


DREAMM-7 Exploratory Analysis: Global Health Status (GHS) and Quality of Life (QoL)

In BVd-treated patients who experienced a bilateral BCVA decline to 20/50 or worse, HRQOL outcomes were stable over time and comparable to those with DVd

 In patients who did not experience a bilateral BCVA decline to 20/50 or worse, a trend toward improvement in GHS/QOL was observed with BVd vs DVd

GHS/QOL



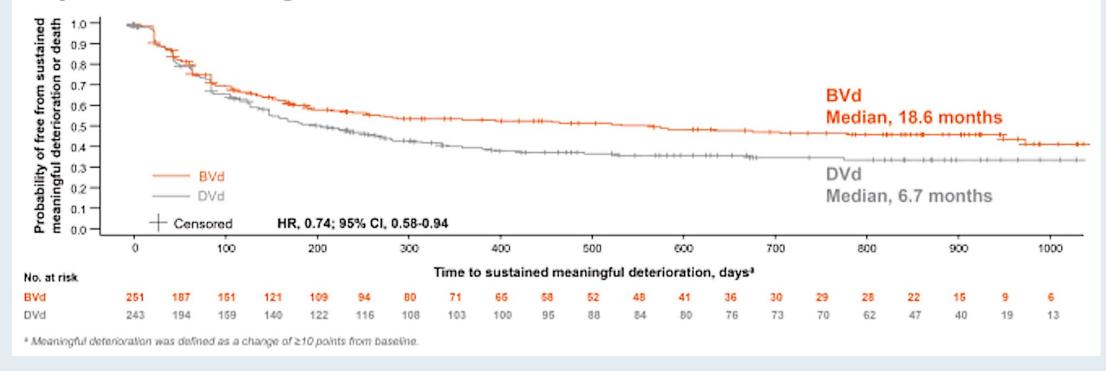
Among patients who experienced a bilateral BCVA decline to 20/50 or worse, the impact on HRQOL subscales was transient and did not meaningfully impact activities of daily life



DREAMM-7 Exploratory Analysis: Treatment Benefit

Treatment with BVd demonstrated a trend toward delayed onset of sustained deterioration in physical functioning

Physical Functioning



A similar trend toward a delay in worsening of disease-specific symptoms was observed.



DREAMM-7 HRQoL: Authors' Conclusions



In BVd-treated patients, HRQOL was not negatively impacted by ocular events, and domain subscales demonstrated comparable outcomes to DVd



In patients with bilateral change of BCVA to 20/50 or worse, HRQOL was maintained, likely due to the transient nature of ocular events and their management with dosage reductions and delays, which have been shown to maintain efficacy^{1,5}



The treatment benefit associated with BVd led to a prolonged time to sustained deterioration in physical functioning, including self-care and walking, with a similar effect seen in disease-specific symptoms



These data provide a comprehensive overview of HRQOL and further support BVd as a potential new therapy for RRMM

NDMM = newly diagnosed multiple myeloma



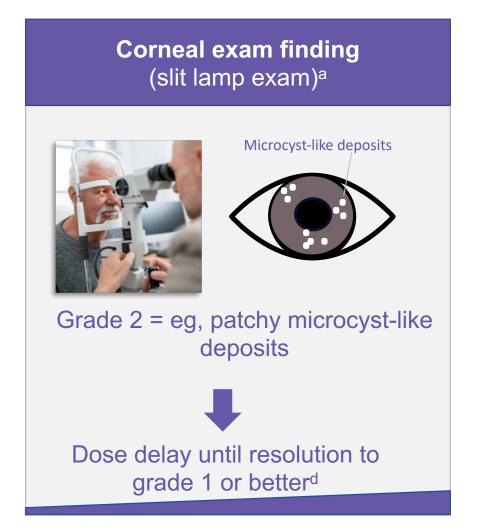
Belantamab Mafodotin (Belamaf) Ocular Events Are Manageable and Reversible With Dose Modifications Guided by Standard Assessments

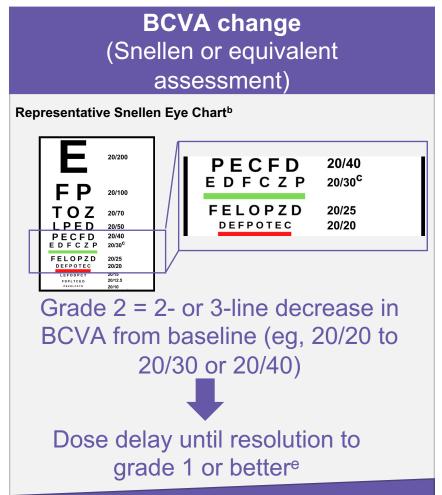
Sagar Lonial,¹ Paul G. Richardson,² Craig E. Cole,³ Asim V. Farooq,⁴ Meghan Berkenstock,⁵ Julie Byrne,⁶ Sumita Roy-Ghanta,⁶ Rachel Rogers,⁶ Amy Phillips-Jones,⁷ Astrid McKeown,⁷ Natalie A. Afshari⁸

¹Winship Cancer Institute, Emory University Hospital, Atlanta, GA, USA; ²Dana-Farber Cancer Institute, Boston, MA, USA; ³Karmanos Cancer Institute, Wayne State University, Detroit, MI, USA, and Michigan State University, Lansing, MI, USA; ⁴University of Chicago Medical Center, Chicago, IL, USA; ⁵Johns Hopkins Wilmer Eye Institute, Baltimore, MD, USA; ⁶GSK, Collegeville, PA, USA; ⁷GSK, Stevenage, UK; ⁸ Shiley Eye Institute, Viterbi Family Department of Ophthalmology, University of California, San Diego, CA, USA

Corresponding author: Sagar Lonial, sloni01@emory.edu

The KVA Scale Used to Grade OEFs Has a Sensitive Criteria for Belamaf Dose Modifications





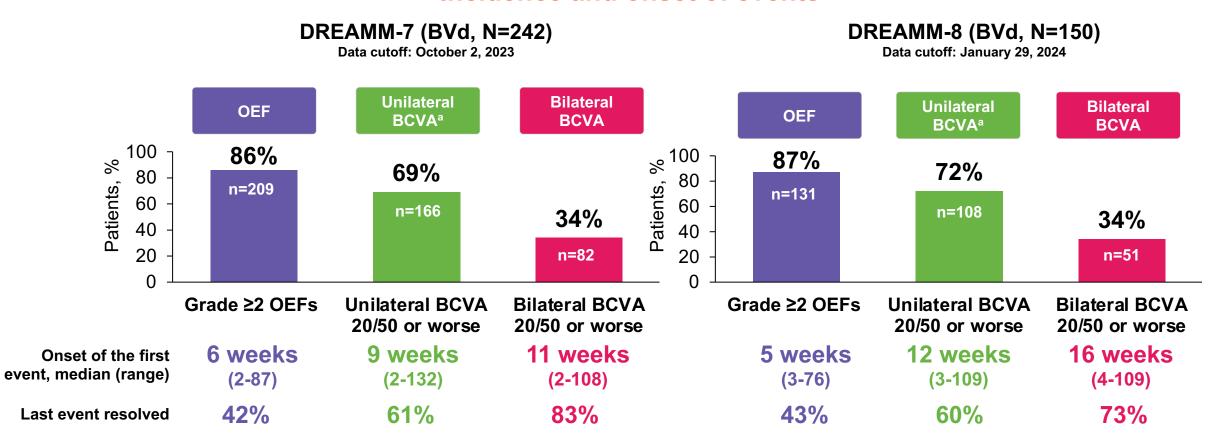
BCVA, best-corrected visual acuity; belamaf, belantamab mafodotin; KVA, Keratopathy and Visual Acuity; OEF, ocular exam finding.

^a The most common corneal exam findings in the trial were superficial punctate keratitis, microcyst-like changes, epithelial haze, and stromal opacity. ^b Not all distances are depicted here.

^c Included 20/30 or 20/32. ^d Mild superficial keratopathy. ^e Decline from baseline of 1 line on Snellen Visual Acuity.

Belamaf Dose Modifications Were Conservatively Guided by OEFs

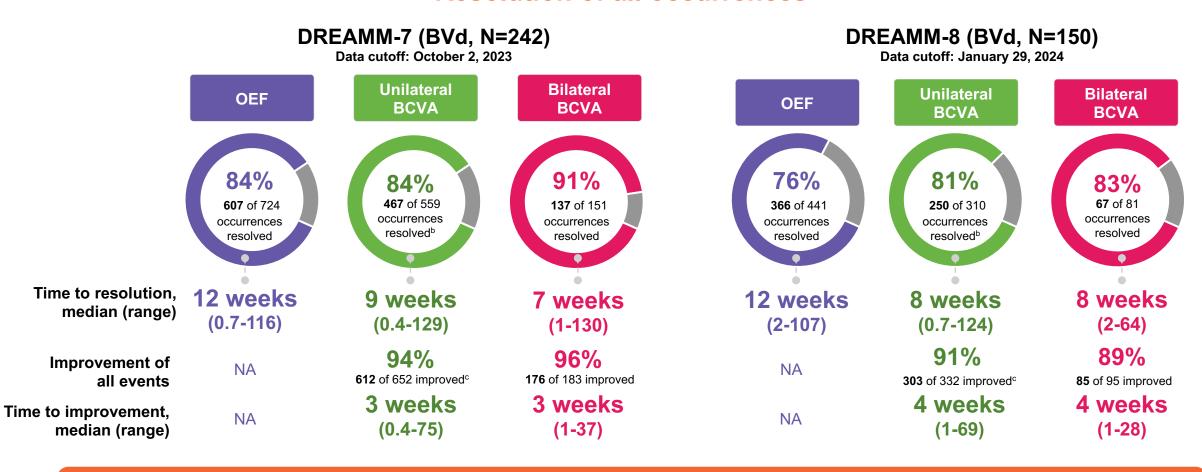
Incidence and onset of events



Grade ≥2 OEFs had a higher incidence and earlier median time to onset than vision changes

Ocular Events Were Reversible in the Majority of Patients

Resolution of all occurrences^a



Patients who experienced a bilateral worsening to 20/50 or worse did so for a median of 7% to 8% of their time on treatment

^a Post hoc analysis. ^b For patients with baseline 20/50 or worse, resolution must additionally be better than 20/50. ^c Only patients with baseline visual acuity better than 20/50 in ≥1 eye are included.

Belantamab Mafodotin (Belamaf) in Combination With Bortezomib, Lenalidomide, and Dexamethasone (VRd) for Patients (Pts) With Transplant-Ineligible (TI) Newly Diagnosed Multiple Myeloma (NDMM): a Focus on Treatment Efficacy and Management/Resolution of Ocular Events in the Phase 1 DREAMM-9 Study

Saad Z. Usmani, MD¹, Michał Mielnik, MD, PhD², Aránzazu Alonso Alonso, MD^{3,4}, Al-Ola Abdallah, MD, PhD⁵, Mamta Garg, MD, FRCP, FRCPath⁶, Wojciech Janowski, MD⁷, Youngil Koh, MD⁸, Chang-Ki Min, MD⁹, Enrique M. Ocio, MD, PhD³, Hang Quach, MD¹⁰, Karthik Ramasamy, MD¹¹, Albert Oriol, MD^{3,12}, Paula Rodriguez-Otero, MD¹³, Ricarda Garcia Sanchez, MD¹⁴, Irwindeep Sandhu, MD, FRCPC¹⁵, Katja Weisel, MD¹⁶, Chris Brawley, MSc¹⁷, Miguel M. Murillo, PhD¹⁸, Fernando Carreño, PhD¹⁹, Jacqueline L. Egger, PhD²⁰, Morrys C. Kaisermann, MD, PhD²¹, Marek Hus, MD²

ASH 2025; Abstract 13646.

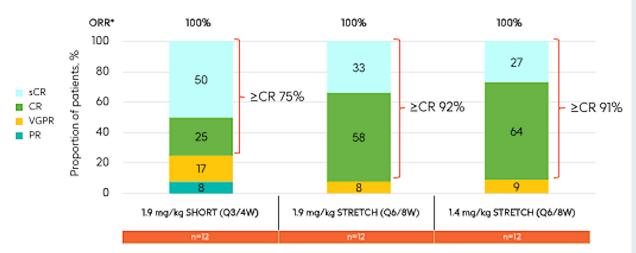


DREAMM-9 Efficacy: Overall Response Rate (ORR) and MRD

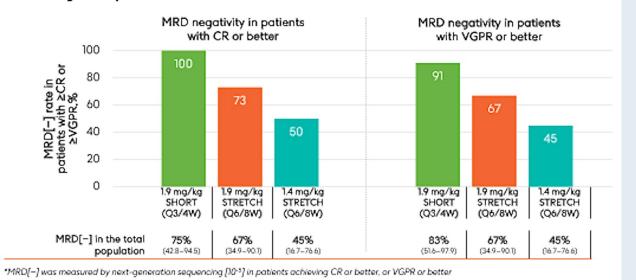
Negativity Rates

• ORR was ≥71% across all dosing cohorts

• ORR was 100% in 1.9 SHORT, 1.9 STRETCH, and 1.4 STRETCH cohorts



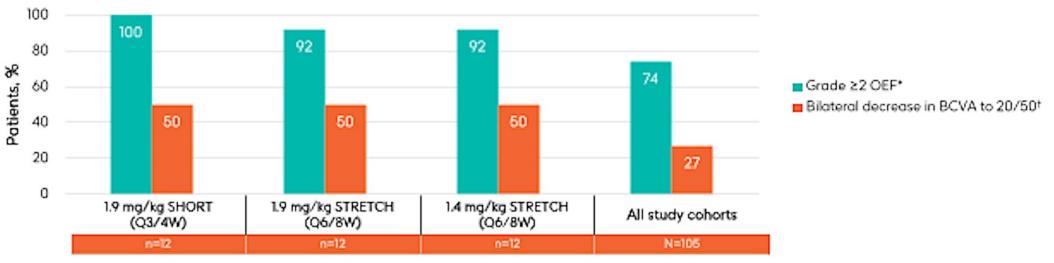
 MRD negativity* in patients with CR or better was higher in cohorts with the highest planned doses, 1.9 SHORT, and 1.9 STRETCH





DREAMM-9 Efficacy: Ocular Event Findings (OEFs)

- In the selected cohorts, occurrence of Gr≥2 OEFs* and bilateral BCVA worsening to 20/50 or worse† were similar
- Ocular events were generally managed with dose modifications and discontinuation rates due to Grade ≥2 OEF events were low at 6% across all 8 cohorts



*OEF were measured on the protocol defined KVA scale comprising corneal findings during a slit lamp examination and/or changes in BCVA. BCVA is visual aculty with the assistance of corrective lenses, assessed using a Snellen chart. BCVA of 20/50 may affect ability to read small print but not activities of daily life or driving. Fin patients in 20/25 BCVA at baseline.

 OEFs were managed by dose delays and schedule extensions and generally reversed when adequate follow up was available



DREAMM-9: Authors' Conclusions

While BVRd demonstrated high ORRs across all cohorts, MRD[-] rates were higher in the 1.9 mg/kg cohorts

Lower-dose-intensity cohorts trended to lower OEF incidence; across cohorts, adverse events including OEFs were manageable via dose reductions/schedule extensions⁸ and rates of discontinuations due to OEFs were low overall

OEFs were **generally reversible and transient**: in the 1.9 mg/kg cohorts, ≥91% of first OEF and 100% of first bilateral BCVA reductions to 20/50 or worse resolved

In line with clinical observations, ER analyses indicated the potential for higher exposure (as expected with a 1.9 mg/kg dose) to improve responses and longer schedules to improve tolerability

Together, in the BVRd regimen, an initial 1.9 mg/kg dose of belamaf with a Q6/8W schedule in patients with NDMM may induce deeper responses, and subsequent schedule extensions (e.g., Q12W) may allow OEF resolution and improve tolerability

These results supported selection of the dose and schedule for belamaf in NDMM including for DREAMM-10 (BRd vs DRd in TI NDMM)^{9,14}

NDMM = newly diagnosed multiple myeloma







A phase 2 trial of iberdomide, carfilzomib, daratumumab and dexamethasone quadruplet therapy for relapsed/refractory multiple myeloma: The **ReKInDLE** study

Ola Landgren, James Hoffman, Abhishek Pandey, Andrew Kowalski, Michael Durante, David Coffey, Marcella Kaddoura, Brian Walker, Leslie Gallardo, Elizabeth Lyubchenko, Massiel Lopez, Fiorela Flores, Liettel Ortega, Rabia Bukhari, Kellye Koubek, Caterine Diaz, Stephanie Mompoint, Sindy Gutierrez, Faika Shah, Stephanie Fernandes, Michelle Armogan, Dickran Kazandjian, Benjamin Diamond

bxd500@miami.edu; @BenDiamondMD

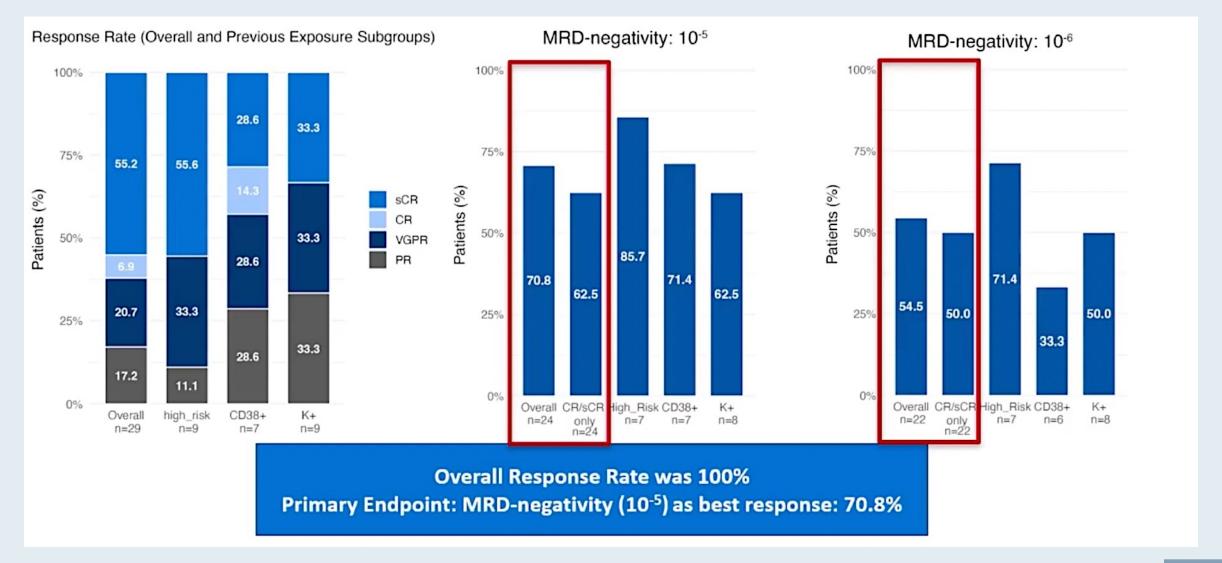


NCT05896228

ASH 2025; Abstract 4108.



ReKInDLE: Response Rates and MRD Analysis





ReKInDLE: Authors' Conclusions

- Iber-DKd appears to be a potent regimen in lenalidomide-refractory multiple myeloma.
- MRD-negativity rate (10⁻⁵) of 70.8% (62.5% in ≥CR only)
 - DKd (CANDOR: MRD-negative CR; 21.8%, Lenalidomide-Refractory; 32%)
 - DPd (APOLLO: MRD-negativity; 9%, Lenalidomide Refractory; 79%)
 - Activity despite prior exposures to K and anti-CD38 therapy.
- Favorable safety profile with predominantly [expected] hematologic toxicity.
 - Further strategies could leverage early growth factor support to further improve dose intensity.
- Deep responses permit time-limited combination therapy and de-escalation to monotherapy.
 - All [n=18] patients in response at time of de-escalation to monotherapy remain in response.
- With the current reality of triple-class exposure and lenalidomide-refractoriness at first relapse, time-limited or response-adapted iberdomide-based combination therapy is worthy of further investigation.
 - EXCALIBER-RRMM (phase III) to report Iber-Dara-Dex (vs Dara-Bor-Dex) in the early relapse setting

DKd = daratumumab/carfilzomib/dexamethasone





Interim analysis of efficacy and safety for Viber-M (ALLG MM25): A Phase Ib/II study of Venetoclax, Iberdomide and Dexamethasone for patients in first or second relapse of Multiple Myeloma with t(11;14)

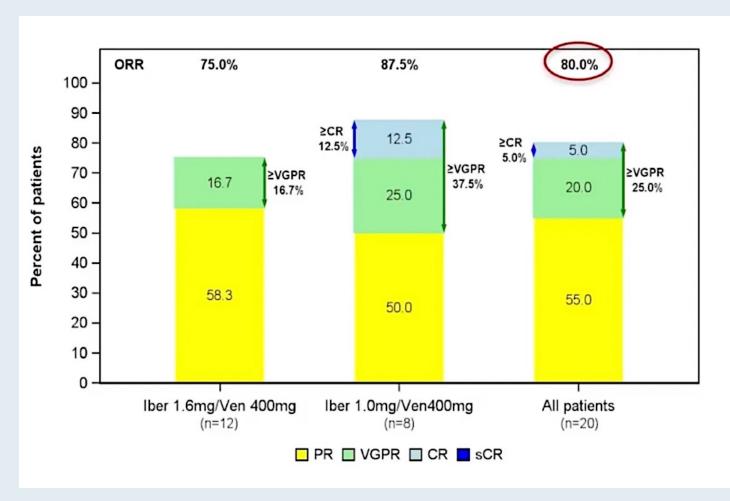
Shirlene Sim^{1,2}, Adam Bryant³, Cecily Forsyth⁴, Olga Motorna⁵, Jennifer Brotchie⁶, Jay Hocking⁷, Angelina Yong⁸, Nicole Chien⁹, Ian Kerridge¹⁰, Hock Choong Lai¹¹, Ann Solterbeck¹², Robert Traficante¹², Maritsa Jocic¹³, Hang Quach^{1,2}

¹St Vincent's Hospital Melbourne, Fitzroy, Victoria, Australia, ²The University of Melbourne, Victoria, Australia, ³Liverpool Hospital, Liverpool, New South Wales, Australia, ⁴Gosford Hospital, Gosford, New South Wales, Australia, ⁵Eastern Health, Box Hill, Victoria, Australia, ⁶South West Healthcare, Warrnambool, Victoria, Australia, ⁷Austin Health, Heidelberg, Victoria, Australia, ⁸Royal Adelaide Hospital, Adelaide, South Australia, Australia, ⁹Auckland City Hospital, Auckland, New Zealand, ¹⁰Royal North Shore Hospital, St Leonards, New South Wales, Australia, ¹¹Townsville University Hospital, Douglas, Queensland, Australia, ¹²Statistical Revelations, Australia, ¹³Australasian Leukaemia and Lymphoma Group, Australia.

Corresponding author: Hang.Quach@svha.org.au



Viber-M (ALLG MM25): Response at the End of Cycle 3



- ORR 80%
 - ≥VGPR 25%
 - ≥CR 5%
- Median time to first response: 41.5 days (range 28 – 62 days)
- Lenalidomide refractory patients (n=13):
 - ORR 85%
 - ≥VGPR 23%
- Daratumumab refractory (n=4):
 - ORR 75%
 - ≥VGPR 25%



Viber-M (ALLG MM25): Authors' Conclusions

- The combination of venetoclax, iberdomide and dexamethasone is deliverable at a dose of <u>venetoclax 400mg</u> and <u>iberdomide 1.0mg</u> in a 28-day cycle
 - Main side effect was neutropenia (50%), but this was manageable
 - Other side effects (e.g. diarrhoea, fatigue, insomnia) were mild, mainly grade 1/2 only
 - Of note, infection rate was low (all grade 35%; grade 3/4 10%)
- This triplet combination demonstrates encouraging early efficacy in t(11;14) MM patients who have had 1-2
 prior lines of therapy, including in Len-refractory patients
 - ➤ ORR of 80% and ≥VGPR rate of 25% at the end of cycle 3 of treatment, with similar responses demonstrated in Len- and Dara-refractory patients
 - At median follow up of 12.2 months, 33/47 patients remain on treatment, with the longest duration on treatment of 26 months and ongoing
- → The Viber-M study is ongoing with 47/50 patients recruited thus far



Safety and Efficacy of Elranatamab in Combination With Iberdomide in Patients With Relapsed or Refractory Multiple Myeloma: Results from the Phase 1b MagnetisMM-30 Trial

Attaya Suvannasankha,¹ Jonathan L. Kaufman,² Ashraf Badros,³ Michel Pavic,⁴ Hock-Choong Lai,⁵ Muhammad S Raza,⁶ Parth S Shah,⁷ Patrick Y. Muller,⁸ Jorge Acosta,⁸ Margaret Hoyle,⁹ Erik R Vandendries,¹⁰ Jay Cheng,¹¹ Alexander Lesokhin¹²

¹Melvin and Bren Simon Comprehensive Cancer Center, Indiana University, Indianapolis, IN, USA; ²Winship Cancer Institute, Emory University, Atlanta, GA, USA; ³Greenebaum Comprehensive Cancer Center, University of Maryland, Baltimore, MD, USA; ⁴Centre Intégré Universitaire de Santé et de Services Sociaux de l'Estrie - Centre Hospitalier Universitaire de Sherbrooke, Quebec, QC, Canada; ⁵Icon Cancer Centre Townsville, Queensland, AU; ⁶Dr. Everett Chalmers Hospital, Halifax, NS, Canada; ⁷Dartmouth Hitchcock Medical Center, Hanover, NH, USA; ⁸Bristol Myers Squibb, Boudry, Switzerland; ⁹Pfizer Inc, Milan, Italy; ¹⁰Pfizer Inc, Cambridge, MA, USA; ¹¹Pfizer Inc, Bothell, WA, USA; ¹²Memorial Sloan Kettering Cancer Center, New York, NY, USA

MagnetisMM-30 Study Design

- MagnetisMM-30 (NCT06215118) is a phase 1b, open-label, multicenter, prospective study
- Part 1 (dose escalation) primary objective was to assess the tolerability and safety of elranatamab in combination with iberdomide to determine the recommended doses of the combination for evaluation in Part 2 (randomized dose optimization)
 - A BOIN approach was used to guide dose escalation/de-escalation in Part 1

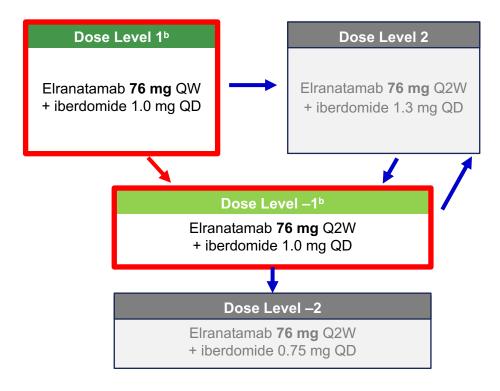
Patients with RRMM

Key inclusion criteria

- Age ≥18 years with MM per IMWG criteria
- ECOG PS 0-1
- 2-4 prior LOTs, including ≥1 IMiD and ≥1 Pla
- Relapsed or refractory to last LOT

Key exclusion criterion

- Stem cell transplant ≤12 weeks prior to enrollment or active GVHD
- Ongoing grade ≥2 peripheral sensory or motor neuropathy; history of grade ≥3 peripheral motor polyneuropathy



Primary endpoint

DLTs during DLT observation period

Secondary endpoints

- AEs and laboratory abnormalities
- ORR°
- CR rate^c
- Time-to-event endpoints^c
- PK
- MRD negativity rate^c
- Immunogenicity

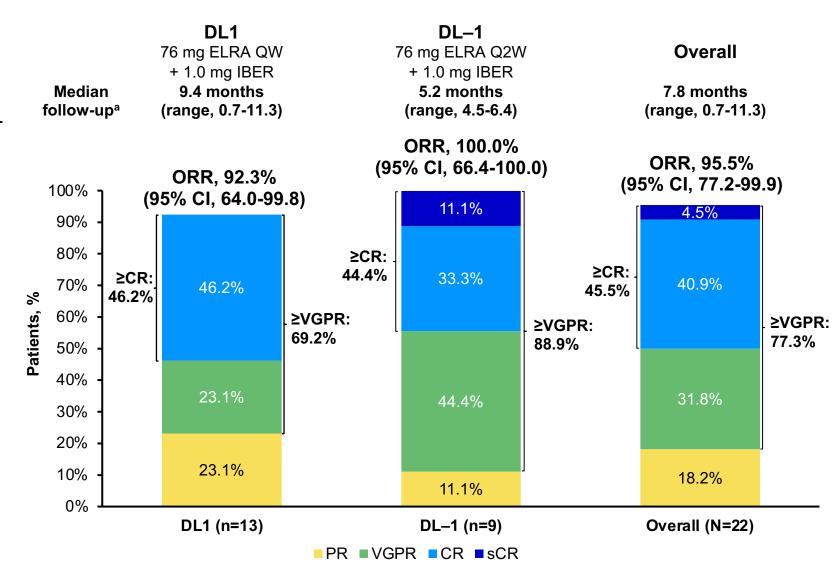
AE=adverse event; BOIN=Bayesian Optimal Interval Design; CR rate=complete response rate; DLT=dose-limiting toxicity; ECOG PS=Eastern Cooperative Oncology Group performance status; GVHD=graft vs host disease; IMiD=immunomodulatory drug; IMWG=International Myeloma Working Group; LOT=line of therapy; MM=multiple myeloma; MRD=minimal residual disease; ORR=objective response rate; PI=proteasome inhibitor; PK=pharmacokinetics; QD=once daily; QW=once weekly; Q2W=once every 2 weeks

a All patients must have received ≥2 consecutive cycles of an IMiD-containing regimen and ≥2 consecutive cycles of a PI or PI-containing regimen; b All patients received an initial 14-day cycle of elranatamab (12 mg on day 1, 32 mg on day 4, 76 mg on day 8) without iberdomide. Iberdomide was dosed at 21 out of 28 days for subsequent cycles; c Per IMWG criteria

ORR

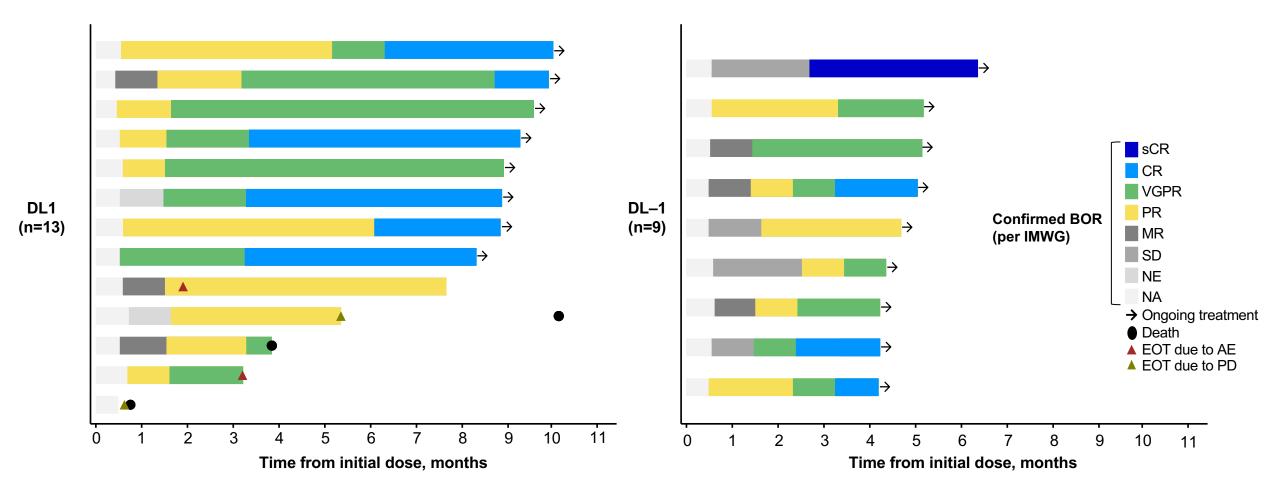
 Overall, the confirmed ORR by investigator was 95.5% (95% CI, 77.2-99.9)

- Responses occurred early
 - Median time to response was1.4 months (range, 0.5-2.7)



^a Simple median of observation times.

Swimmer Plot: Response and PFS per investigator



Grade 5 treatment-emergent adverse events were progressive disease and pancreatic cancer (patient died with VGPR).

AE=adverse event; BOR=best overall response; CR=complete response; DL=dose level; IMWG=International Myeloma Working Group; EOT=end of treatment; MR=minimal response; NA=not assessed; NE=not evaluable; PD=progressive disease; PR=partial response; sCR=stringent complete response; SD=stable disease; VGPR=very good partial response

Agenda

| Introduction | Best of ASH Multiple Myeloma |
|--------------|------------------------------|
| Case 1 | Dr Lorber – 59-year-old man |

■ Faculty Presentation: Antibody-Drug Conjugates and Other Emerging Novel
Therapies for Relapsed/Refractory (R/R) Multiple Myeloma (MM) — Dr Lonial

| Case 2 | Dr Morganstein – 84-year-old man |
|--------|----------------------------------|
| Case 3 | Dr Favaro – 64-year-old man |
| Case 4 | Dr Bhatnager – 71-year-old man |
| Case 5 | Dr Lee – 71-year-old man |

■ Faculty Presentation: Integrating CAR T-Cell Therapy and Bispecific Antibodies into the Management of R/R MM — Dr Mateos

Case 6 Dr Rudolph – 56-year-old man



Case Presentation: 84-year-old morbidly obese man with CAD, CHF and sleep apnea with multiregimen-relapsed myeloma eventually, as part of expanded access, receives belantamab mafodotin with low-dose pomalidomide



Dr Neil Morganstein (Summit, New Jersey)



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Case 6 Dr Rudolph – 56-year-old man





Dr Justin Favaro (Charlotte, North Carolina)

Case Presentation: 64-year-old man s/p CVA with aphasia who has multiregimen-relapsed MM after daratumumab, PIs, IMIDs and selinexor receives teclistamab



Dr Priya Rudolph (Athens, Georgia)

Questions for the faculty



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■ Faculty Presentation: Integrating CAR T-Cell Therapy and Bispecific Antibodies into the Management of R/R MM — Dr Mateos

Case 6 Dr Rudolph – 56-year-old man



Case Presentation: 71-year-old man with kappa light chain myeloma experiences CR on cilta-cel CAR T-cell therapy with hypogammaglobulinemia requiring IVIG, develops melanoma of the abdominal wall



Dr Tina Bhatnagar (Wheeling, West Virginia)



Agenda

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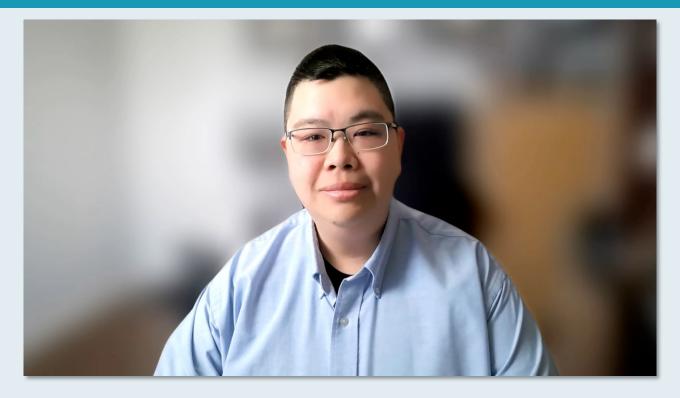
Case 5 Dr Lee – 71-year-old man

■ Faculty Presentation: Integrating CAR T-Cell Therapy and Bispecific Antibodies into the Management of R/R MM — Dr Mateos

Case 6 Dr Rudolph – 56-year-old man



Case Presentation: 71-year-old man with light chain MM receives daratumumab/RVd with VGPR, subsequently completes ASCT → maintenance ixazomib and 2 years later is diagnosed with locally advanced prostate cancer treated with radiation therapy and ADT. Now with myeloma progression



Dr Eric Lee (Fountain Valley, California)



Agenda

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Integrating CAR-T cell therapy and BsAbs into the management of RRMM patients

María-Victoria Mateos Salamanca, Spain





Treatment landscape in Multiple Myeloma today: realistic situation

1st line

ASCT eligible

AntiCD38 + PI + IMiD + Dex

ASCT

Len/Dara-Len

ASCT ineligible

Dara-Len-dex

Dara-VMP/RVd

AntiCD38 + PI + IMiD + Dex

2nd line

Based on sensitivity/refractoriness to Daratumumab and Lenalidomide

Anti-CD38 + Carfilzomib-dex

Anti-CD38 + Pomalidomide-dex

Pomalidomide-bortezomib-dex

Selinexor-bortezomib-dex

Carfilzomib-dex

3rd line

Anti-CD38 + Pomalidomide-dex

Elotuzumab-Pomalidomide-dex

Previous combos if pt eligible

4th line

BCMA-targeted therapy

CAR-T

Ide-cel

Cilta-cel

Teclistamab, Elranatamab

BsAbs

Linvoseltamab

GPRC5D-targeted therapy

BsAbs

Talquetamab

The label is for RRMM after at least 4 PL of therapy

including PI, IMiD and antiCD38 and refractory

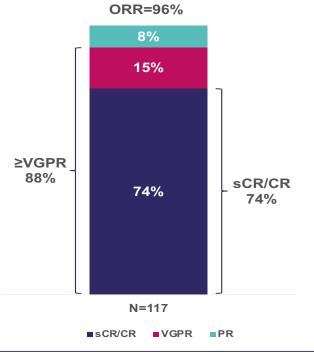
to the last line of therapy





Phase 2 Registrational Study of Anitocabtagene Autoleucel for the Treatment of Patients with (RRMM): Updated Results from iMMagine-1

- Anito-cel is an autologous BCMA-directed CAR T-cell therapy using a novel D-Domain binder
- 117 TCE RRMM pts were dosed: median age 64; 18% EMD; 18% high tumor burden and 40% HRCA
- Median number of PL: 3, TCR 87% and PCR 41%. 76% received bridging therapy. Median f/u: 16 months



| MRD Negativity at 10 ⁻⁵ Sensitivity Level | | |
|--|-----------------|--|
| Overall MRD negativity, % (n/N) | 95% (91/96) | |
| Median time to MRD negativity, months (min - max) | 1.0 (0.9 – 6.4) | |
| MRD negativity sustained for ≥ 6 months, % (n/N) | 83% (54/65) | |
| MRD Negativity at 10 ⁻⁶ Sensitivity Level | | |
| Overall MRD negativity, % (n/N) | 78% (68/87) | |

| N=117 | PFS Rate (%) (95% CI) | OS Rate (%) (95% CI) |
|----------|--------------------------|-------------------------|
| 6-Month | 93.1 (86.7, 96.5) | 95.7 (90.0, 98.2) |
| 12-Month | 82.1 (73.6, 88.1) | 94.0 (87.8, 97.1) |
| 18-Month | 67.4 (55.4, 76.8) | 88.0 (78.8, 93.4) |
| 24-Month | 61.7 (48.0, 72.8) | 83.0 (70.7, 90.5) |

- Safety profile: CRS in 85% (G1 in 68% and G2 in 17); ICANS in 8%
- No delayed NT; no SPM; no enterocolitis
- G3-4 neutropenia: 70% and G3-4 infections: 9%





KarMMa-3: Ide-cel in RRMM after 2-4PL: outcomes in the older population

Methods

 KarMMa-3 (NCT03651128) is an open-label, phase 3 trial in patients with RRMM (Figure 1)^{1,4}

Figure 1. Study design, key eligibility criteria, and study objectives

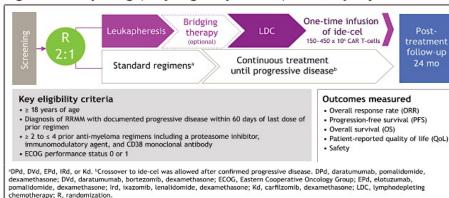


Table 3. Treatment-emergent adverse events of special interest/select adverse events grade ≥3 (safety population)

| | Age < 70 years | | Age ≥ 70 years | |
|------------------------------|----------------------|----------------------------------|---------------------|----------------------------------|
| AESI/Select AE, n (%) | Ide-cel (n = 178) | Standard regimens (n = 99) | Ide-cel (n = 47) | Standard regimens (n = 27) |
| At least one AESI/Select AEa | 70 (39.3) | 29 (29.3) | 14 (29.8) | 13 (48.1) |
| CRS | 8 (4.5) | - | 3 (6.4) | 120 |
| Infections | 48 (27.0) | 20 (20.2) | 12 (25.5) | 6 (22.2) |
| iiNT | 4 (2.2) | - | 3 (6.4) | 1-1 |

Table 2. Response rates

| • | Patients older than 70 years were, overall: with |
|---|---|
| | less HRCA, less TCR and with longer time from the |
| | last line of therapy |

• 20% of pts approximately were older than 70 years

| | Age < 70 years | | Age ≥ 70 years | |
|----------------------|----------------------|-----------------------------------|----------------------|----------------------------------|
| Outcomes | lde-cel (n = 205) | Standard regimens (n = 105) | Ide-cel (n = 49) | Standard regimens (n = 27) |
| ORR, % (95% CI) | 68.8 (62.4, 75.1) | 41.0 (31.5, 50.4) | 81.6 (70.8, 92.5) | 48.1 (29.3, 67.0) |
| P value | < 0.0001 | | 0.0037 | |
| PFS, median (95% CI) | 12.5 (11.2-15.4) | 4.2 (3.5-5.7) | 18.9 (12.1-24.5) | 5.7 (2.2-12.2) |
| P value | < 0.0001 | | 0.0012 | |

Conclusions:

Patients aged ≥ 70 years from the KarMMa-3 trial experienced benefit from ide-cel treatment, as evidenced by a longer median PFS and notable ORR compared with patients treated with standard regimens

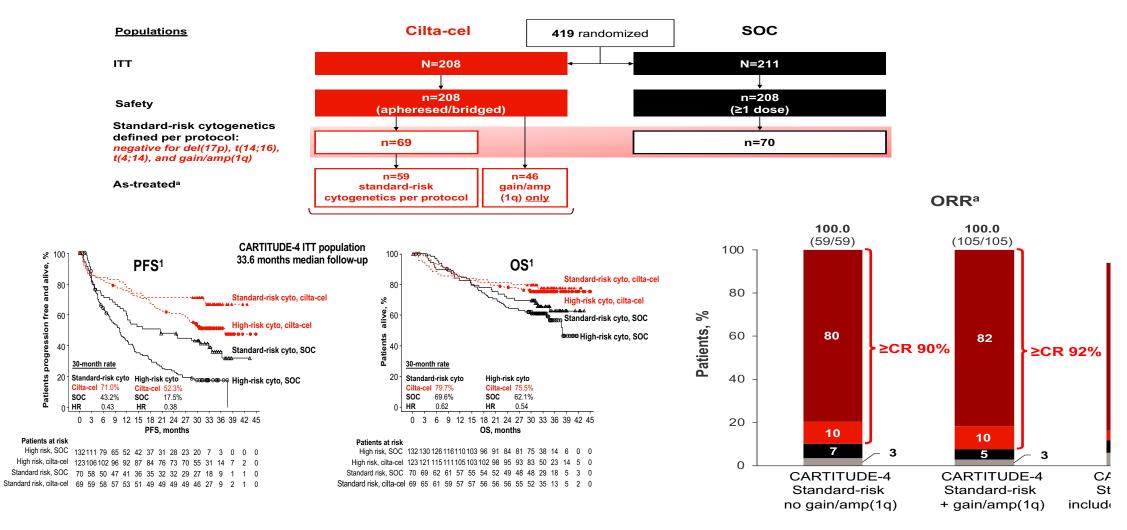
*Includes CRS, neurologic toxicity - Focused 2.0 FDA, and infections.

AE, adverse event; AESI, AE of special interest; CRS, cytokine release syndrome; iiNT, investigator identified neurotoxicity.



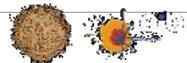


What is the subgroup of patients treated with Cilta-cel in C4 with better outcomes?



Safety profile of cilta-cel in standard-risk population was consistent with overall study population^{1,b}





The better outcomes for standard risk patients in C4 are applicable to C1

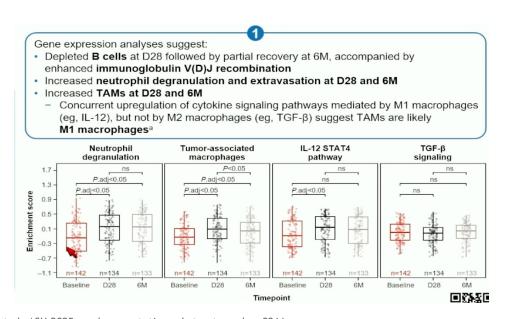
| | 30-month PFS rate, % | 30-month OS rate, % | |
|--|-----------------------------|---------------------|--|
| CARTITUDE-4 (median 2 prior lines) | 59.4% | 76.4% | |
| Cilta-cel Standard-risk cytogenetics per protocol | 80.5 | 87.3 | |
| Cilta-cel Standard-risk cytogenetics per protocol + gain/amp(1q) | 71.7 | 86.1 | |
| CARTITUDE-1 (median 6 prior lines) | Median PFS 36 m and OS 60 m | | |
| Cilta-cel Standard-risk cytogenetics per protocol including gain/amp(1q) | 59.9 | 70.6 | |

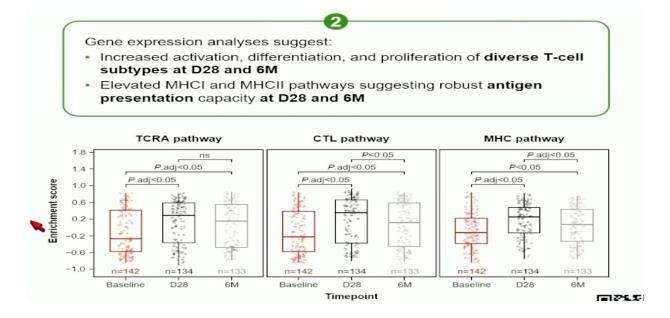


Earlier use of Cilta-cel is associated with better immune fitness and stronger immune effects

- Cilta-cel in both C1 and C4 trials showed better outcomes in earlier lines of therapy (especially after 1 or 2 PL)
- Evaluation included PB samples at D28 and M6:
- Bulk RNA sequencing
- TCR repertoire in C-1
- Immunophenotyping by FCM

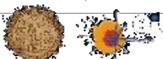
Cartitude-4





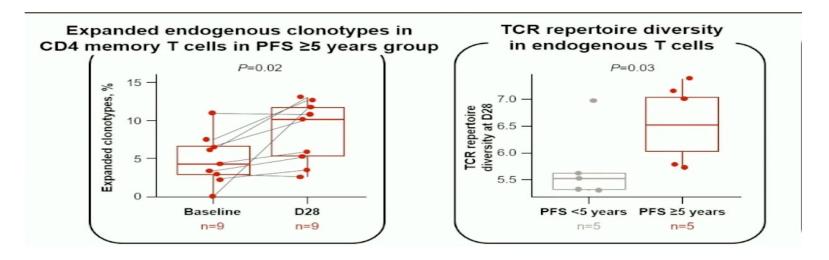






Earlier use of Cilta-cel is associated with better immune fitness and stronger immune effects

Cartitude-1: 33% of pts remain alive and progression-free at 5 years



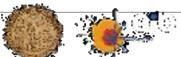
Cartitude-4

Earlier use of cilta-cel may support longer PFS by leveraging a more immunocompetent TME at baseline

CARTITUDE-4 TME: Patients With Shorter PFS (≤18M) and Patients With More pLOT had a More Suppressive TME

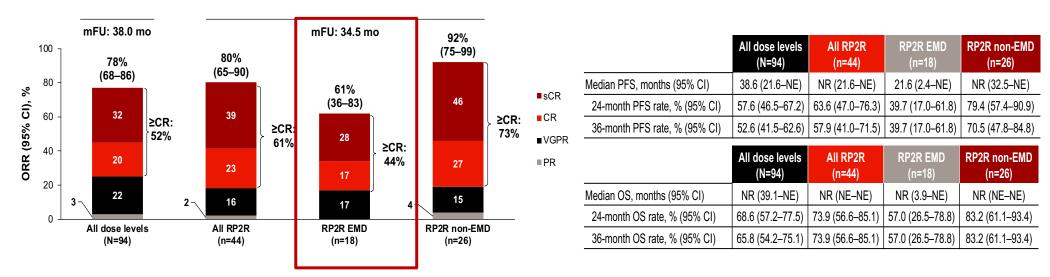






Teclistamab-Talquetamab in RRMM patients: phase 1b of RedirecTT-1 trial

• 94 pts overall and 43 at the RP2R were included. Median number of PL: 4 (86% TCR and 46% with true EMD)

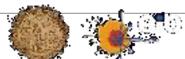


- Safety profile: Infections are the most relevant ones (43% G3-4 at the RP2R); opportunistic infections in 17% of pts; any-grade infection risk peaked early and stabilized by ~6 months; grade ≥3 infection risk was highest in the first 6 months and decreased over time
- On target off tumor toxicity related with GPRC5D are frequent but most are G1-2

Tec-Tal today has a role in patients with true EMD because this population has an unmet need.

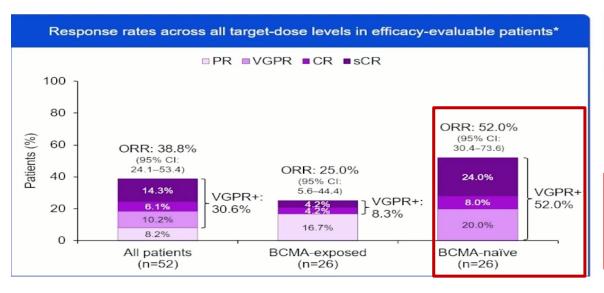
MonumenTAL-9 study will confirm its role in all RRMM patients





Subcutaneous Cevostamab (a novel FcRH5xCD3 bispecific T-cell engager — BiTE) in RRMM patients: Phase Ib CAMMA 3 study

- 58 pts included. Median number of PL: 5 (64% TCR and 48.3% BCMA-TT exposed including CAR-T, ADC and BsAbs)
- Cevostamab was evaluated at different doses and dose > 120 mg is effective. Fixed duration: 13 c

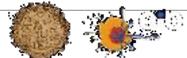


| | All patients (N=52) | BCMA- naïve (n=26) | BCMA- exposed (n=26) |
|---|---------------------------|--------------------------|----------------------------|
| Ongoing response, n/N (%) | 11/19 (57.9%) | 8/13 (61.5%) | 3/6 (50.0%) |
| Median DOR among responders, months (95% CI) | 12.3 (8.3, NE) | NR (11.7, NE) | 8.3 (4.9, NE) |

• Safety profile: Infections (44.8% and G3-4 12%); CRS in 69% and injection-site reaction in 58.6%; neutropenia in 31% (most G3-4)

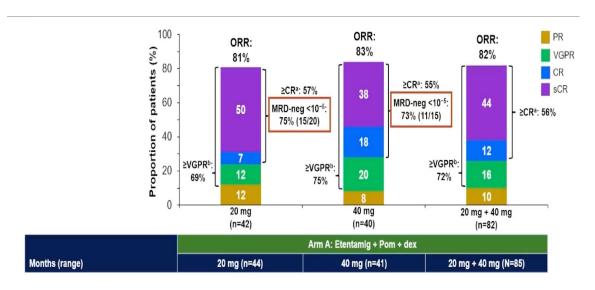
The role of Cevostamab is interesting for the sequencing of T-cell redirecting therapies but.. more data are required and its role is maybe in combination

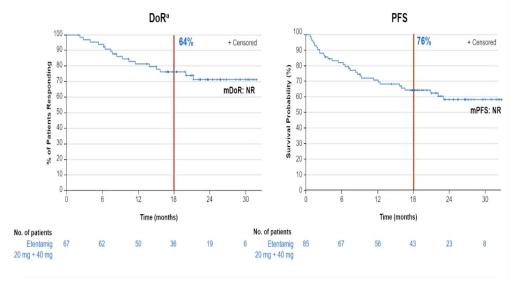




Etentamig (BCMAxCD3 BiTE) plus Pom-dex in RRMM patients: Cohort A of the clinical trial

- 85 pts included. Median number of PL: 4 (73% TCR and 51% refractory to pomalidomide)
- Etentamig was given in this cohort with no SUD, no premedication and Q4W
- Pom and dex at conventional doses from C1D1 and dose for etentamig was 20 or 40 mg





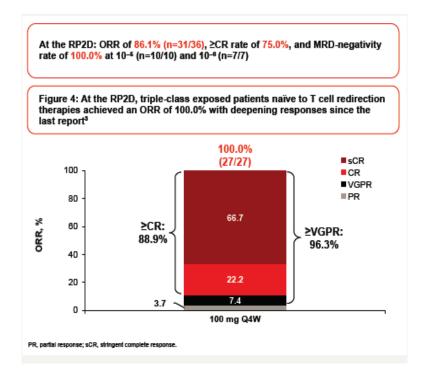
- Safety profile: Infections (89% and G3-4 52%); CRS in 35% (no G3-4); neutropenia G3-4 in 78%. Hepatic toxicity in 25% (G3-4 in 8%)
- Cohort E included some modifications: SUD 1 and Pom included at C2D1. Dose of etentamig 60 mg and less pretreated but the f/u was short

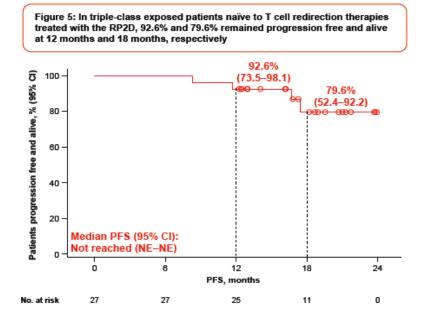


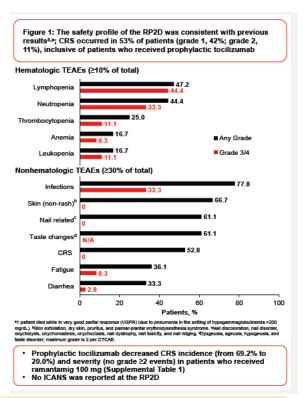


Ramantamig, BCMA-GPRC5D trispecific mAB in RRMM patients

• 36 pts received ramantamig at the RP2D (1 SUD followed by 100 mg flat dose Q4W). Median number of 4 PL (53% TCR and 25% previously exposed to BCMA/GPRc5D)





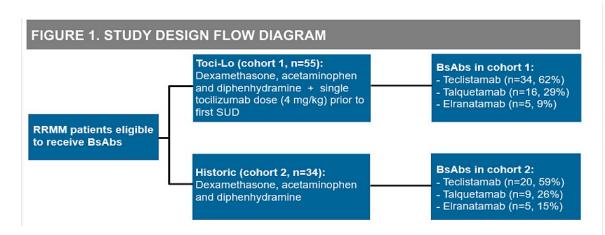


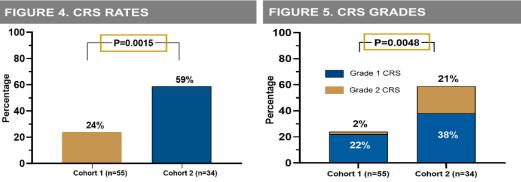
Ramantamig has a great efficacy in patients naïve to T-Cell Redirecting therapy similar to CAR-T It is pending to know the activity in patients previously exposed as well as in specific populations like patients with EMD





Low dose Tocilizumab for mitigation of CRS with BsAbs in RRMM



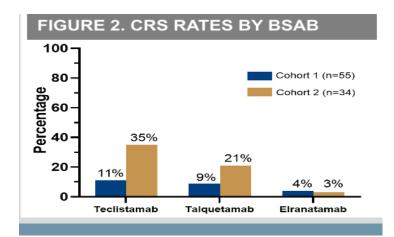


- Of the 13 CRS events in cohort 1, 5 occurred after SUD1 (38%); in cohort 2, 12 occurred after SUD1 (60%).
- Recurrent CRS was noted in 1 patient in cohort 1 compared to 6 patients in cohort 2;
 both received tocilizumab for management.

TABLE 2. RESULTS OF LOGISTIC REGRESSION ANALYSIS

| | Odds ratio | 95% confidence interval | P-value |
|--|--|--|---------|
| Univariate logistic regression | | | |
| Tocilizumab prophylaxis (yes vs. no) | 0.22 | 0.08-0.53 | 0.005 |
| Multivariate logistic regression | | | |
| Tocilizumab prophylaxis (yes vs. no) High risk cytogenetics (yes vs. no) Prior T-cell redirecting therapy (yes vs no) Elevated LDH (yes vs. no) High ALC (yes vs. no) Presence of EMD (yes vs. no) | 0.23 0.64 1.33 0.34 1.94 0.78 | 0.08-0.62 0.21-1.86 0.32-5.50 0.11-1.00 0.60-7.10 0.30-2.25 | 0.0001 |

The rates of neurotoxicity/ICANS were 5% in cohort 1 (n=3) and 6% cohort 2 (n=2). One patient in each cohort developed grade 2 neurotoxicity/ICANS which resolved more than 48 hours following management per institutional guidelines.



Prophylactic single Toci-lo proved to be an effective strategy at reducing CRS incidence and severity during the SUD phase of BsAbs treatment. This low dose instead of the standard (8 mg/kg) may be a cost-effective strategy.





Agenda

Introduction Best of ASH Multiple Myeloma

Case 1 Dr Lorber – 59-year-old man

■ Faculty Presentation: Antibody-Drug Conjugates and Other Emerging Novel
Therapies for Relapsed/Refractory (R/R) Multiple Myeloma (MM) — Dr Lonial

| Case 2 | Dr Morganstein – 84-year-old man |
|--------|----------------------------------|
| Case 3 | Dr Favaro – 64-year-old man |
| Case 4 | Dr Bhatnager – 71-year-old man |
| Case 5 | Dr Lee – 71-year-old man |

■ Faculty Presentation: Integrating CAR T-Cell Therapy and Bispecific Antibodies into the Management of R/R MM — Dr Mateos

Case 6 Dr Rudolph – 56-year-old man



Case Presentation: 56-year-old man with heavily relapsed MM who received multiple prior lines of therapy, including CAR T-cell therapy, is now on talquetamab



Dr Priya Rudolph (Athens, Georgia)



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Practical Perspectives on the Current and Future Management of Immune Thrombocytopenia — What Happened at ASH 2025?

A CME/MOC-Accredited Live Webinar

Tuesday, December 16, 2025 5:00 PM - 6:30 PM ET

Faculty

Hanny Al-Samkari, MD Cindy Neunert, MD, MSCS Professor Francesco Zaja

Moderator Neil Love, MD



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Attendees will also receive an email in 1 to 3 business days with these instructions.

