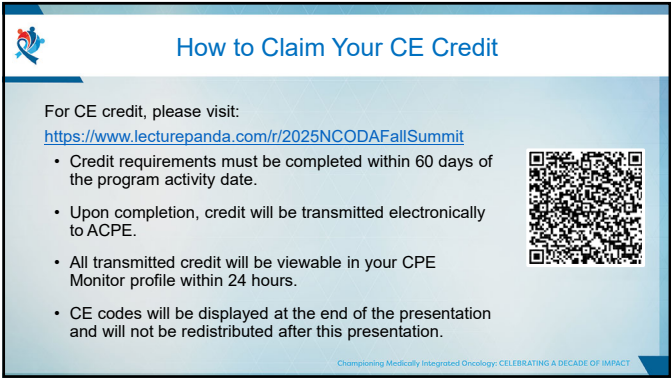




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OBJECTIVES

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1. Discuss the role of quadruplet therapy in transplant-eligible and transplant-ineligible multiple myeloma and identify appropriate use in the frontline setting

2. Review chimeric antigen receptor (CAR) T-cell therapies' use in relapsed/refractory multiple myeloma

3. Summarize the role of BCMA-targeting bispecific antibodies in the management of relapsed/refractory multiple myeloma

4. Determine an evidence-based treatment plan for relapsed/refractory multiple myeloma with available sequencing data

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DISCLOSURES

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There are no relevant conflicts of interest to disclose for this presentation for the following speakers and planners:

Jordan Snyder, PharmD, BCOP

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Multiple Myeloma

Multiple myeloma is characterized by the abnormal proliferation of malignant plasma cells

Accounts for 1% of all cancer diagnosis and 10% of all hematologic malignancies

Median age at diagnosis is 65 years

Occurs more commonly in men and African Americans

Cause of nearly 13,000 deaths in the United States each year

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Plasma Cell Disorder Spectrum

Monoclonal Gammopathy of Undetermined Significance (MGUS)

- Asymptomatic
- Increases risk of developing multiple myeloma at ~1% per year

Smoldering Myeloma

- Asymptomatic
- Increases risk of developing multiple myeloma at ~10% per year

Multiple Myeloma

- Rate of progression depends on cytogenetics and development of new mutations



Holm MJ, et al. J Clin Hematol. 2023;4(1):35-42.

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Quad Therapy in Newly Diagnosed Myeloma

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Doublets to Triplets

Trial	Population	Intervention	Outcomes	Impact
VISTA (2010)	NDMM, transplant ineligible (N=682)	VMP vs MP	•35% reduced risk of death at 3 years (HR 0.653, P<0.001) •3-year OS: 68.5% vs 54%	Addition of bortezomib significantly prolongs survival
GIMEMA-MMY-3006 (2012)	NDMM, transplant eligible (N=474)	VTD vs TD	•Median PFS: 60 months vs 41 months •10-year survival estimate: 34% vs 17%	Addition of bortezomib into double HSCT improved PFS and OS
SWOG S0777 (2017)	NDMM, without immediate HSCT (N=525)	VRD vs RD	•Median PFS: 43 mo vs 30 mo (HR 0.712, 95% CI 0.56-0.906; P=0.0018) •Median OS: 75 mo vs 64 mo (HR 0.709; 95% CI 0.524-0.959; P=0.025)	Triplet therapy became standard of care in NDMM

V: bortezomib; M: melphalan; P: prednisone; T: thalidomide; D: dexamethasone; R: lenalidomide; IMiD: immunomodulator; mo: months; HR: hazard ratio; CI: confidence interval; PFS: progression free survival; OS: overall survival

Holm MJ, et al. J Clin Oncol. 2010;28(13):2259-2265; Trebban P, et al. Lancet Haematol. 2020;7(10):e81-e873; Durie BG, et al. The Lancet. 2017;389(10066):519-527

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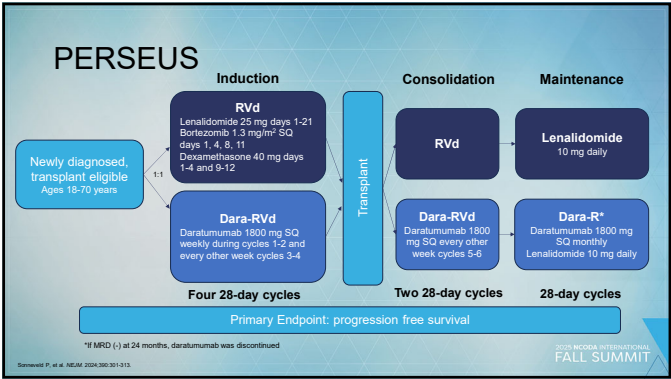
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Quad Therapies in Transplant Eligible				
Trial	Population	Intervention	Outcomes	Impact
CASSIOPEIA (2019)	NDMM, transplant eligible (N=1085)	DaraVTD vs VTD with post HSCT consolidation and maintenance	•sCR at day 100: 29% vs 20% •Median PFS: NR in either group	Clinical benefits seen in induction, consolidation, and maintenance Less impact in US as RVD considered superior to VTD
GRIFFIN (2020)	NDMM, transplant eligible (N=207)	DaraRVD vs RVD with post HSCT consolidation and maintenance with DaraR or R	•ORR post-consolidation: 99% vs 91.8% (P=0.016) •4-year PFS: 87.2% vs 70% (HR 0.45; 95% CI 0.21-0.95; P=0.032) •Median OS: NR for both groups	Addition of dara to RVD resulted in higher response rates with deepening responses over time

Mousse P, et al. Lancet. 2019;394:29-38. Mousse P, et al. Lancet Oncol. 2020;21(5):636-645. Voorhes PM, et al. Blood. 2020;135(5):636-645. Voorhes PM, et al. Lancet Oncol. 2023;10110:6329-6337.

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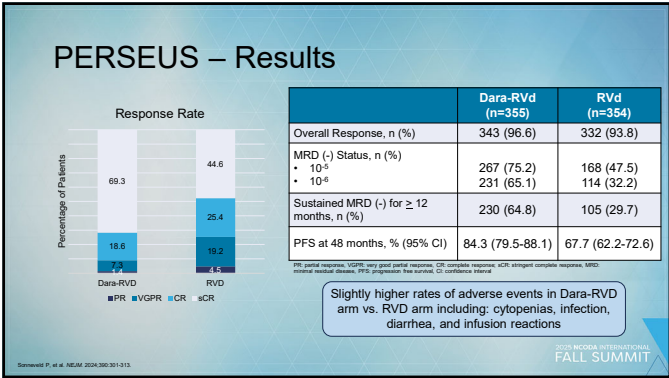
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PERSEUS – Patient Demographics		
Baseline Characteristics	DaraRVd (n=355)	RVd (n=354)
Median Age, years (range)	61 (32-70)	59 (31-70)
Male, n (%)	211 (59.4)	205 (57.9)
Race, n (%)		
• White	330 (93)	323 (91.2)
• Black	5 (1.4)	4 (1.1)
ECOG PS, n (%)		
• 0-1	335 (94.3)	338 (95.4)
Cytogenetic Risk, n (%)		
• Standard	264 (74.4)	266 (75.1)
• High	76 (21.4)	78 (22)
Median time since diagnosis, months	1.2	1.1

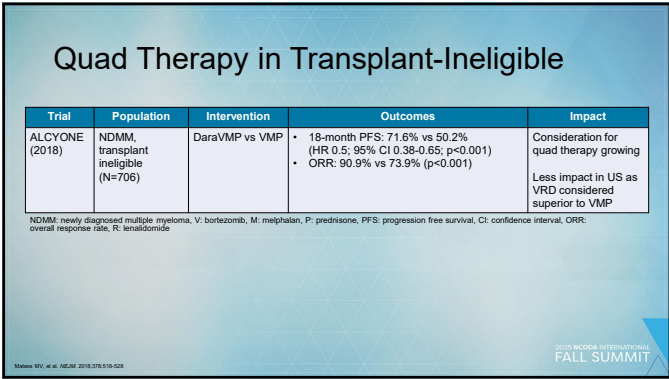
PS: performance status

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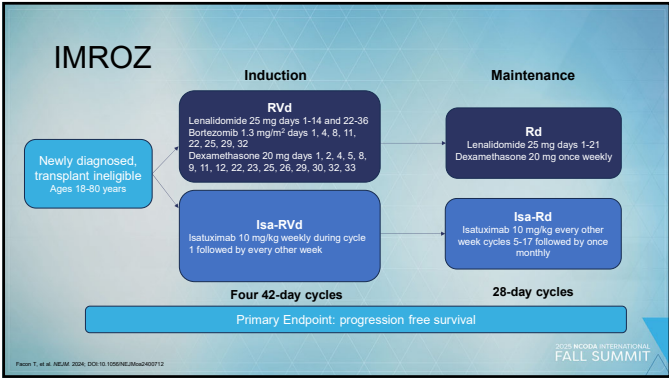
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IMROZ – Patient Demographics

Baseline Characteristic	IsaRVd (n=265)	RVd (n=181)
Median Age, years (range)	72 (60-80)	72 (55-80)
Male, n (%)	143 (54)	94 (51.9)
Race • White • Black	192 (72.5) 2 (0.8)	131 (72.4) 2 (1.1)
ECOG PS • 0 - 1	235 (88.7)	162 (89.5)
Cytogenetic Risk • Standard • High	207 (78.1) 40 (15.1)	140 (77.3) 34 (18.8)
Median time since diagnosis, months	1.2	1.2

PS: performance status

Patel T, et al. NEJM. 2024. DOI:10.1056/NEJMoa2402712

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IMROZ – Results

Overall Response

	IsaRVd (n=265)	RVd (n=181)
ORR, %	91.3	92.3
CR or better, %	74.7	64.1
60-month PFS, %	63.2	45.2
MRD- 10 ⁻⁴ , %	58.1	43.6
Estimated OS at 60 months, %	72.3	66.3

PR: partial response; VGPR: very good partial response; CR: complete response; sCR: stringent complete response; ORR: overall response rate; MRD: minimal residual disease; PFS: progression free survival; OS: overall survival

Slightly higher rates of adverse events with IsaRVd despite higher rates of discontinuation with RVd

Patel T, et al. NEJM. 2024. DOI:10.1056/NEJMoa2402712

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BENEFIT

Induction
IsaRd
Isatuximab 10 mg/kg weekly during cycle 1 followed by every other week cycles 2-12
Lenalidomide 25 mg days 1-21
Dexamethasone 20 mg once weekly
Isa-RVd
Bortezomib 1.3 mg/m² days 1, 8, 15 of each cycle
Twelve 28-day cycles

Consolidation
IsaR
Isatuximab 10 mg/once monthly
Isa-RV
Bortezomib 1.3 mg/m² days 1 and 15
Six 28-day cycles

Maintenance
Isa-R
28-day cycles

Primary Endpoint: Rate of MRD

Patel T, et al. Nature Med. 2024;30:2320-2331

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BENEFIT – Patient Demographics

Baseline Characteristic	IsaRVd (n=135)	IsaRd (n=135)
Median Age, years (IQR)	73.2 (71-76)	73.6 (71-76)
Male, n (%)	74 (55)	71 (53)
ECOG PS, n (%) • 0 or 1	125 (93)	119 (88)
Cytogenetic Risk, n (%) • Standard • High	68 (53) 13 (10)	75 (60) 10 (8)
Median time from diagnosis, months	1	0.9

PS: performance status

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BENEFIT – Results

Overall Response

	IsaRVd (n=135)	IsaRd (n=135)
ORR, %	85	78
CR or better, %	58	31
24-month PFS, % (95% CI)	85.2 (79.2-91.7)	91.5 (86.5-96.8)
24-month OS, % (95% CI)	91.1 (86.1-96.4)	80.0 (73.3-87.4)

PR: partial response, VGPR: very good partial response, CR: complete response, ORR: overall response rate, PFS: progression-free survival, CI: confidence interval, OS: overall survival

Slightly higher rates of adverse events with IsaRVd versus IsaRd including high rates of peripheral neuropathy

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CEPHEUS

Induction

Maintenance

Newly diagnosed, transplant ineligible
Ages 18-70 years

RVd

Lenalidomide 25 mg days 1-14
Bortezomib 1.3 mg/m² days 1, 4, 8, 11
Dexamethasone 20 mg days 1, 2, 4, 5, 8, 9, 11, 12

Dara-RVd

Daratumumab 1800 mg weekly cycles 1-2 followed by every 3 weeks cycles 3-8

Rd

Lenalidomide 25 mg days 1-21
Dexamethasone 40 mg once weekly

Dara-Rd

Daratumumab 1800 mg once monthly

Eight 21-day cycles

28-day cycles

Primary Endpoint: Overall MRD (-)

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CEPHEUS – Patient Demographics

Baseline Characteristic	DaraRVd (N=197)	RVd (N=198)
Median Age, years (range)	70 (42-79)	70 (31-80)
Male, n (%)	87 (44)	111 (56.1)
Race, n (%)		
• White	162 (82.2)	156 (78.8)
• Black	10 (5.1)	9 (4.5)
ECOG PS, n (%)		
• 0 -1	174 (88.3)	184 (92.9)
Cytogenetic Risk, n (%)		
• Standard	149 (75.6)	149 (75.3)
• High	25 (12.7)	27 (13.6)
Median time since diagnosis, months	1.2	1.3

PS: performance status

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CEPHEUS – Results

Overall Response

	DaraRVd (n=135)	RVd (n=135)
≥ CR, %	81.2	61.6
MRD (-) rate, %		
• 10 ⁻⁵	60.9	39.4
• 10 ⁻⁶	46.2	27.3
Median PFS, months	NR	52.6

PR: partial response, VGPR: very good partial response, CR: complete response, MRD: minimal residual disease, PFS: progression free survival

Slightly higher rates of adverse events with DaraRVd compared to RVd including: cytopenias, infection, pneumonia, injection site reactions

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Quad Therapy in Newly Diagnosed Myeloma

- Quad therapy has changed the landscape of newly diagnosed multiple myeloma
- Quad therapy showed higher rates of MRD-negativity and response rates in transplant-eligible and –ineligible myeloma
- Higher rates of adverse events were seen in patients receiving quad therapy, but did not impact outcomes

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QUESTION 1

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PL is a 52-year-old male with a past medical history of hypertension and GERD. He presents in clinic with newly diagnosed multiple myeloma. ECOG PS is 0 and labs are unremarkable. He is deemed a transplant candidate. What is an appropriate 1st line regimen for PL?

a. IsaRVd

b. DaraRVd

c. RVd

d. DRd

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CAR-T in Early Relapse

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T-Cell Approvals in Multiple Myeloma

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2021

Idecabtagene vicleucel

2022

Ciltacabtagene autoleucel

Teclistamab

2023

Elranatamab

Talquetamab

2024

Ciltacabtagene autoleucel: 2nd line

Idecabtagene vicleucel: 3rd line

2025

Linvoseltamab

FDA Oncology (Cancer) Hematology Multiple Myeloma Approval Notifications. Updated March 20, 2024. Accessed September 19, 2025. www.fda.gov/drugs/development-resources/updates-approved-drugs/hematology-cancer-hematology-multiple-myeloma-approval-notifications

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CAR-T in Late Relapse

Trial	Population	Intervention	Outcomes	Conclusions
KarMMa	Relapsed myeloma • Following ≥ 3 lines of therapy	Idecabtagene vicleucel	• ORR: 73% (95% CI: 66-81) • ≥ CR: 33% • mPFS: 8.8 months (95% CI: 5.6-11.6)	Ide-cel induced responses in heavily pre-treated patients with multiple myeloma
CARTITUDE-1	Relapsed myeloma • Following ≥ 3 lines of therapy	Ciltacabtagene autoleucel	• ORR: 97% (95 CI: 91.2-99.4) • sCR: 67% • 5-year PFS: 33% • mOS: 60.7 months (95% CI: 41.9-NE)	Early, deep, and durable responses were achieved in heavily pre-treated patients with cilta-cel

mPFS: median progression free survival, Ide-cel: idecabtagene vicleucel, CR: complete response, CI: confidence interval, mPFS: median progression free survival, mOS: median overall survival, NR: not reached, Cilta-cel: ciltacabtagene autoleucel, sCR: stringent complete response

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KarMMa-3

Relapsed/refractory multiple myeloma

2:1

Idecabtagene vicleucel (n=254)

Standard of care* (n=132)

	Ide-cel (n=254)	SOC (n=132)
Median age (range), years	63 (30-81)	63 (42-83)
Extramedullary disease, n (%)	61 (24)	32 (24)
High tumor burden, n (%)	71 (28)	34 (26)
High risk cytogenetics, n (%)	107 (42)	61 (46)
Median prior lines (range)	3 (2-4)	3 (2-4)
Previous autologous HSCT, n (%)	214 (84)	114 (86)
Triple-class refractory, n (%)	164 (65)	89 (67)

*Standard of care (SOC) regimens include: daratumumab, pomalidomide, dexamethasone; daratumumab, bortezomib, dexamethasone; lenalidomide, dexamethasone; carfilzomib, dexamethasone; elotuzumab, pomalidomide, dexamethasone

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KarMMa-3: Results

Overall Response Rates

Response	Ide-cel	SOC
PR	35	5
VGPR	22	10
sCR	11	27

	Ide-cel (n=254)	SOC (n=132)
Median PFS, mo (95% CI)*	13.3 (11.8-16.1)	4.4 (3.4-5.9)
6-month PFS, %	73	30
Median time to response, mo (range)	2.9 (0.5-13)	2.1 (0.9-9.4)
Median duration of response, mo (95% CI)	14.8 (12-18.6)	9.7 (5.4-16.3)

*p < 0.001, VGPR: very good partial response, sCR: stringent complete response, mo: months

Progression-free survival benefit with Ide-cel was seen across subgroups, including presence or absence of high-risk cytogenetics, high tumor burden, or triple-class refractory status

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KarMMa-3: Toxicity

	Ide-cel (n=250)	SOC (n=126)
CRS, n (%)	197 (88)	0 (0)
Neutropenia, n (%)	195 (78)	55 (44)
Anemia, n (%)	165 (66)	45 (36)
Infection, n (%)	146 (58)	68 (54)
Thrombocytopenia, n (%)	136 (54)	36 (29)
Nausea, n (%)	112 (45)	34 (27)
Diarrhea, n (%)	85 (34)	30 (24)
Neurotoxic event, n (%)	34 (15)	0 (0)

CRS: cytokine release syndrome

Ide-cel resulted in significantly longer PFS versus SOC regimens. These results were observed across subgroups and may benefit difficult to treat patients

Median time to onset of CRS: 1 day (1-14 days)

Median duration of CRS: 3.5 days (1-51 days)

72% of patients receiving Ide-cel required tocilizumab for CRS

28% of patients receiving Ide-cel required corticosteroids for CRS

Median time to onset of neurotoxic event: 3 days (1-317 days)

Median duration of neurotoxic event: 2 days (1-37 days)

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CARTITUDE-4

Relapsed/refractory multiple myeloma

1:1

Cilta-cel (n=208)

Standard of care (SOC)* (n=211)

	Cilta-cel (n=208)	SOC (n=211)
Median age (range), years	61.5 (27-78)	61 (35-80)
Soft tissue plasmacytomas, n (%)	44 (21.2)	35 (16.6)
Bone marrow plasma cells > 60%, n (%)	42 (20.4)	43 (20.7)
High-risk cytogenetics, n (%)	123 (59.4)	132 (51)
Previous lines of therapy, n (%)		
1	68 (32.7)	68 (32.2)
2	83 (39.9)	87 (41.2)
3	57 (27.4)	56 (26.2)
Triple-class refractory, n (%)	30 (14.4)	33 (15.6)

*SOC regimens include: pomalidomide, bortezomib, dexamethasone; daratumumab, pomalidomide, dexamethasone

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CARTITUDE-4: Results

Overall Response Rates

Percentage of Patients

58.2

14.9

8.2

3.4

Cilta-cel (n=208)

15.2

6.8

23.7

21.8

SOC (n=211)

■ PR

■ VGPR

■ CR

■ sCR

	Cilta-cel (n=208)	SOC (n=211)
Median time to response, mo (range)	2.1 (0.9-11.1)	1.2 (0.6-10.7)
Median time to best response, mo (range)	6.4 (1.1-18.6)	3.1 (0.8-20.6)
Minimal residual disease negative, n (%)	126 (60.6)	33 (15.6)
30-month PFS, %*	59.4	25.7
30-month OS, %	76.4	63.8

*p < 0.0001; VGPR: very good partial response; sCR: stringent complete response; mo: months; NR: not reached

PFS benefit with cilta-cel was seen across subgroups, including presence of high-risk cytogenetics and prior lines of therapy

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CARTITUDE-4: Toxicity

	Cilta-cel (n=208)	SOC (n=126)
Neutropenia, n (%)	187 (89.9)	177 (85.1)
CRS, n (%)	134 (76.1)	0 (0)
Infection, n (%)	129 (62)	148 (71.2)
Anemia, n (%)	113 (54.3)	54 (26)
Thrombocytopenia, n (%)	113 (54.3)	65 (31.2)
Nausea, n (%)	101 (48.6)	38 (18.3)
Hypogammaglobulinemia, n (%)	88 (42.3)	13 (6.2)

Median time to CRS onset: 8 days (1-23 days)

Median duration of CRS: 3 days (1-17 days)

CRS: cytokine release syndrome

Cilta-cel had a favorable risk-benefit profile compared to standard of care. Due to the progression free survival benefit, cilta-cel is an option for patients on first relapse

San Miguel J, et al. N Engl J Med. 2023;389:335-47

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CAR-T in Earlier Lines

	KarMMa-3	CARTITUDE-4
Median age, years (range)	63 (30-81)	61.5 (27-78)
High risk cytogenetics, n (%)	107 (42)	123 (59.4)
ORR, n (%)	181 (71)	176 (84.6)
PFS, months, (95% CI)	13.3 (11.8-16.1) (18.6-month follow-up)	NR (15.9-month follow-up)
Any grade adverse events, n (%)		
• Cytokine release syndrome	197 (88)	134 (76.1)
• Neutropenia	146 (58)	187 (89.9)
• Thrombocytopenia	136 (54)	113 (54.3)
• Infection	146 (58)	129 (62)
• Neurotoxic events	34 (15)	36 (20.5)

ORR: overall response rate; NR: not reached

Rodriguez-Otero P, et al. N Engl J Med. 2023;389:350-61; San Miguel J, et al. N Engl J Med. 2023;389:335-47

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CAR-T in Relapsed/Refractory Myeloma

- CAR-T in earlier lines of therapy provides deep and durable remissions compared to standard of care chemotherapy
- May be an appropriate option in certain patients depending on performance and disease status
- No new safety signals identified when used in early relapse

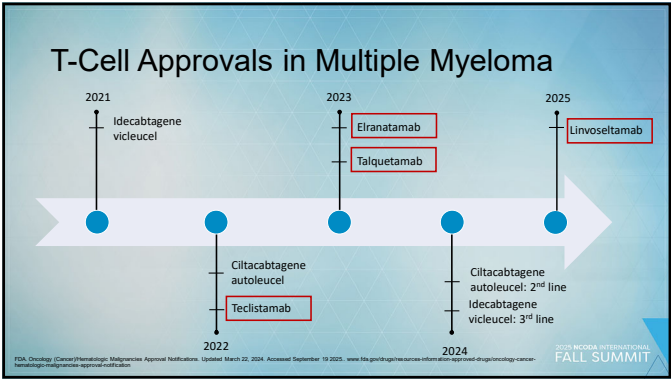
San Miguel J, et al. N Engl J Med. 2023;389:335-47; San Miguel J, et al. N Engl J Med. 2023;389:335-47

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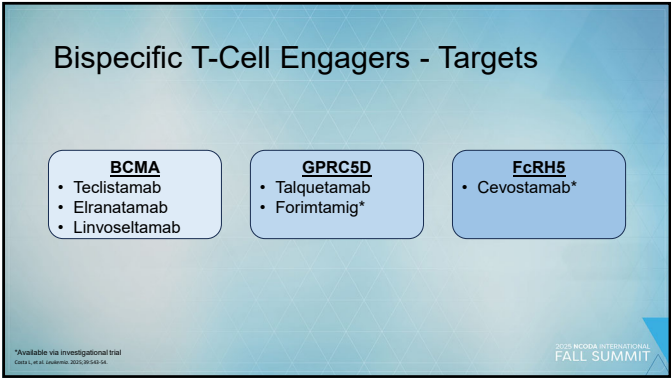
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BCMA Bispecific T-Cell Engagers

Bispecific T-Cell Engager	Indication	Route	Maintenance Dosing	Hospitalization?	REMS
Teclistamab	Relapsed MM after 4 or more lines of therapy	SQ	1.5 mg/kg SQ once weekly x 6 months* 1.5 mg/kg every 2 weeks	Yes – 48 hours after each step-up dose and first treatment dose	Yes
Elranatamab		SQ	76 mg SQ once weekly x 6 months 76 mg SQ every 2 weeks x 6 months* 76 mg SQ once monthly*	Yes – 48 hours after step-up dose 1 and 24 hours after step-up dose 2	Yes
Linvoseltamab		IV	200 mg IV once weekly weeks 4-13 200 mg IV every 2 weeks week 14-24 200 mg IV once monthly*	Yes – 24 hours after step-up dose 1 and 2	Yes

*In responders only. MM, multiple myeloma; SQ, subcutaneous; IV, intravenous.

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MAJESTEC-1

Relapsed Multiple Myeloma

- ≥ 18 years
- 3 prior lines of therapy
- No prior BCMA exposure

Teclistamab

- Target dose: 1.5 mg/kg SQ once weekly

Patient Characteristics	n=165
Median age (range), years	64 (33-84)
Soft tissue plasmacytomas, n (%)	28 (17)
High-risk cytogenetics, n / total n (%)	38/148 (15.5)
Median number of prior lines, n (range)	5 (2-14)
Refractory Status, n (%)	128 (77.6)
• Triple-class	50 (30.3)
• Penta-drug	

Moffatt, P, et al. N Engl J Med. 2022; 387:690-905.

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MAJESTEC-1 - Results

Response Rate

Response	Percentage of Patients
PR	19.4
VGPR	6.7
CR	32.7
sCR	4.2

	n=165
VGPR or better, n (%)	97 (58.8)
Median PFS, months (95% CI)	11.3 (8.8-17.1)
Median OS, months (95% CI)	18.3 (15.1-NE)

Adverse Events

	n=165
Any Adverse Event, n (%)	165 (100)
Cytokine Release Syndrome, n (%)	119 (72.1)
Neutropenia, n (%)	117 (70.9)
Thrombocytopenia, n (%)	66 (40)
Pneumonia, n (%)	30 (18.2)
Neurotoxic Events, n (%)	24 (14.5)

Teclistamab provided deep and durable responses in a heavily pre-treated patient population

Moffatt, P, et al. N Engl J Med. 2022; 387:690-905.

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MagnetisMM-3

Relapsed Multiple Myeloma

- ≥ 18 years
- 3 prior lines of therapy
- No prior BCMA exposure

Eirnatamab

- Target dose: 76 mg SQ once weekly x6 cycles followed by 76 mg SQ every 2 weeks

Patient Characteristics		n=123
Median age (range), years		68 (36-89)
Soft tissue plasmacytomas, n (%)		28 (17)
High-risk cytogenetics, n (%)		31 (25.2)
Median number of prior lines, n (range)		5 (2-22)
Refractory Status, n (%)		119 (96.7)
• Triple-class		
• Penta-drug		52 (42.3)

Leadwinn AS, et al. Nature Med 2023;29:2259-67

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MagnetisMM-3

Response Rate

■ PR ■ VGPR ■ CR ■ sCR

	n=123
ORR, % (95% CI)	61 (51.8-69.6)
VGPR or better, n (%)	69 (56.1)
Median PFS, months (95% CI)	NE (9.9-NE)
Median OS, months (95% CI)	NE (13.9-NE)

Adverse Events	
Any Adverse Event, n (%)	123 (100)
Cytokine Release Syndrome, n (%)	71 (57.7)
Neutropenia, n (%)	60 (48.8)
Thrombocytopenia, n (%)	38 (30.9)
Infection, n (%)	86 (69.9)
ICANS, n (%)	4 (3.4)

Eirnatamab demonstrated durable responses that continued to deepen after the transition to biweekly dosing

Leadwinn AS, et al. Nature Med 2023;29:2259-67

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LINKER MM-1

Relapsed Multiple Myeloma

- ≥ 18 years
- 3 prior lines of therapy
- No prior BCMA immunotherapies

Linvoseltamab

- Target dose: 50 mg or 200 mg IV once weekly x 12 weeks followed by 200 mg IV every other week x 12 weeks followed by 200 mg IV once monthly

Patient Characteristics		200 mg (n=117)
Median age (range), years		70 (37-91)
Soft tissue plasmacytomas, n (%)		19 (16.2)
High-risk cytogenetics, n (%)		46 (39.3)
Median number of prior lines, n (range)		5 (2-16)
Refractory Status, n (%)		96 (82.1)
• Triple-class		
• Penta-drug		33 (28.2)

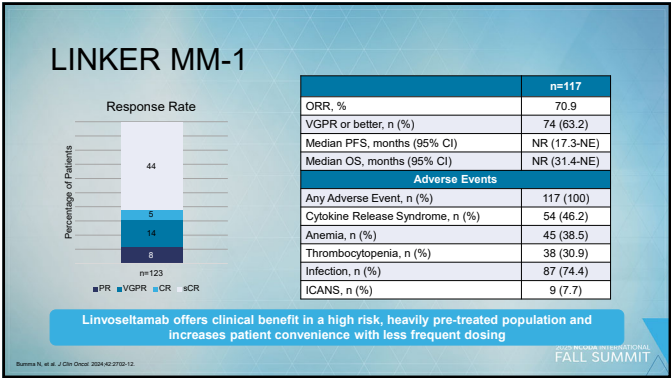
Burns N, et al. J Clin Oncol 2024;42:2703-12

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BCMA Bispecific T-Cell Engagers

Bispecific T-cell Engager	ORR	≥ VGPR	PFS	CRS Rate		ICANS Rate	
				All Grades	≥ Grade 3	All Grades	≥ Grade 3
Teclistamab	63%	58.8%	11.3 mo (14.1 mo follow up)	72.1%	0.6%	14.5%	0.6%
Eltanetamab	61%	56.1%	NR (14.7 mo follow up)	71%	0	3.4%	0
Linvoseltamab	70.9%	63.2%	NR (14.3 mo follow up)	46.2%	0.9%	7.7%	2.6%

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QUESTION 2

PL completed induction with Dara-Rvd and completed as autologous stem cell transplant. He is currently on daratumumab + lenalidomide maintenance. During his 1-year work up, his bone marrow biopsy showed 30% monoclonal plasma cells.

What is an appropriate 2nd line therapy?

- a. Teclistamab
- b. Ide-cel
- c. Cilta-cel
- d. DRd

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Sequencing BCMA Agents

- BCMA-targeting agents have changed the management of relapsed/refractory multiple myeloma
- Several BCMA-targeting agents available including: cilta-cel, ide-cel, teclistamab, elranatamab, linvoseltamab, belantamab (via clinical trial or compassionate use)
- Concerns with T-cell exhaustion and development of resistance are growing concerns with repeated BCMA-targeting agents
- With access to several agents, optimal sequencing while important, is still yet to be determined

Chenmaitre M, et al. ASCTO Daily News. Sequencing Anti-BCMA Therapies in Multiple Myeloma: What Is Their Optimal Use with Other Available Treatment Options? 12/2024. Accessed 9/18/2025. <https://myeloma.asctodo.org/sequencing-and-treatment-agent-multiple-myeloma-from-optimal-use-after-relapse>. CSM 11. Blood. 2024;145(25):2502-27.



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CARTITUDE-2 Cohort C

Evaluation of cilta-cel in BCMA-exposed relapsed/refractory myeloma

	All (n=20)	Prior ADC (n=13)	Prior BsAb (n=7)
Median age (range), years	62.5 (44-81)	66 (44-81)	60 (49-71)
High-risk cytogenetics, n (%)	3 (15)	2 (15.4)	1 (14.3)
Median prior lines, n (range)	8 (4-13)	8 (4-13)	8 (6-12)
Penta drug refractory, n (%)	11 (55)	7 (53.8)	4 (57.1)
Anti-BCMA refractory, n (%)	16 (80)	11 (84.6)	5 (71.4)

Time from last anti-BCMA to cilta-cel infusion: 62-944 days

Duration of prior anti-BCMA: 1-527 days

Cohen AD, et al. Blood. 2022;141(12):1919-203.

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CARTITUDE-2 Cohort C - Results

	All (n=20)	Prior ADC (n=13)	Prior BsAb (n=7)
ORR, % (95% CI)	60 (36.1-80.9)	61.5 (43.1-86.1)	57.1 (18.4-90.1)
MRD (-) 10-5, %	35	39	28
Median DOR, months (95% CI)	11.5 (7.9-NE)	11.5 (7.9-NE)	8.2 (4.4-NE)
Median PFS, months (95% CI)	9.1 (1.5-NE)	9.5 (0.99-NE)	5.3 (0.6-NE)

Results suggest response after prior BCMA studies, larger prospective trials are needed for a better understanding of sequencing

No additional safety concerns identified

Cohen AD, et al. Blood. 2022;141(12):1919-203.

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Patients more likely to respond if:

- Shorter duration of prior BCMA agent
- Longer time between cilta-cel and prior BCMA agent

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QUESTION 3

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RT is a 71-year-old female with relapsed/refractory multiple myeloma. She has a past medical history of treatment-related peripheral neuropathy, hypothyroidism, and hypertension. She has previously received Dara-RVd, HSCT, Daratumumab/lenalidomide maintenance, KPd, DPd, and most recently cilta-cel (currently day +180). She is now progressing.

What would be an appropriate next line of therapy?

a. Elranatamab

b. Talquetamab

c. CyBorD

d. Belantamab

e. a or b

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SUMMARY

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- Landscape of multiple myeloma treatment is rapidly evolving
- Quad therapies have become the main stay in treatment of transplant-eligible and –ineligible newly diagnosed multiple myeloma
- The use of t-cell redirecting therapies are highly active in multiple myeloma and are continuing to be investigation in combination and in earlier lines of therapy
- BCMA-targeting agents are effective in a heavily pre-treated patient population, but optimal sequencing is still under investigation

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QUESTION & ANSWER

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Myeloma:
From Current to Cutting Edge

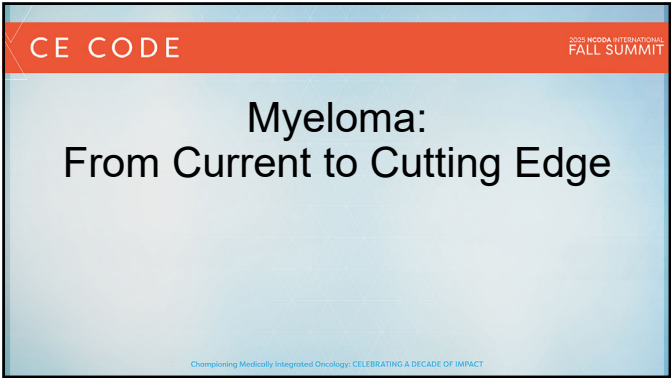
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