Idecabtagene Vicleucel (ide-cel) Shows Similar Efficacy and Toxicity in Patients with Multiple Myeloma Aged 70 and Older Compared to Younger Patients: a Multicenter Cohort Study. Evgenii Shumilov¹, Philipp Berning¹, Markus Maulhardt², Hristo Boyadzhiev^{3,4}, Anca-Maria Albici⁵, Snjezana Janjetovic⁶, Anna Ossami Saidy⁶, Christian Schultze-Florey⁷, Simon Call¹, Natalie Schub⁵, Annamaria Brioli⁷, Michael Daskalakis⁸, Juliane Knust², Justin Hasenkamp², Matthias Stelljes¹, Cyrus Khandanpour⁹, Ulrike Bacher⁸, Christine Hanoun¹⁰, Florian Heidel⁷, Hans Christian Reinhardt¹⁰, Georg Lenz¹, Gerald Wulf², Friedrich Stölzel⁵, Bastian von Tresckow¹⁰, Thomas Pabst³ Department of Hematology and Oncology, University Hospital Muenster, Muenster, Germany. 6. Department of Hematology, Oncology, and Tumor Immunology Helios Klinikum Berlin-Buch Berlin Germany. Department of Hematology, Hemostasis, Oncology and Stem Cell Transplantation, Hannover Medical School, Hannover, Germany. Department of Hematology and Medical Oncology, University Hospital Göttingen, Göttingen, Germany. 8. Department of Hematology and Central Hematology Laboratory, Inselspital, Bern University Hospital, University of Bern, Switzerland. 3. Department of Medical Oncology, University Hospital Bern, University of Bern, Bern, Switzerland. 9. Department of Haematology and Stem Cell Transplantation, West German Cancer Center and German Cancer consortium (DKTK partner 4. Habichtswald Hospital, Kassel, Germany. site Essen), University Hospital Essen, University of Duisburg-Essen, Essen, Germany. 5. University Hospital Schleswig-Holstein, Kiel, Department of Stem Cell Transplantation and Cellular Immunotherapies, Kiel University, Kiel, Germany

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INTRODUCTION

B-cell maturation antigen (BCMA) directed chimeric antigen receptor-T (CAR-T) cell therapies have revolutionized the treatment landscape of relapsed/ refractory (r/r) multiple myeloma (MM). Data on the efficacy and safety in older patients - who often have more co-morbidities and are often underrepresented - remain limited.

OBJECTIVE

To evaluate the efficacy and toxicity profiles of patients aged 70 and older compared to their younger counterparts in a real-world setting.

METHODS

Multicenter analysis retrospective including 132 relapsed/refractory MM pts without CNS manifestation undergoing treatment with ide-cel CAR-T cell between March 2022 and May 2024 at seven tertiary german centers. Patients were grouped by age at CAR-T infusion $(<70 \text{ vs.} \geq 70 \text{ years (yrs)})$. Subsequently, descriptive and survival analyses, including propensity score matching (nearest neighbor 1:1 matching with Charlson combordity index, triple-class refractoriness, and remission status as co-variates), were performed to compare outcomes between both age groups.

	0 years</th <th>≥ 70 years (N=44)</th> <th>P-valu</th>	≥ 70 years (N=44)	P-valu
Adian age at CAR-T infusion Irangel	60.8 [37.0, 69.3]	72.6 [70.0. 82.6]	< 0.00
Female	37 (42.0%)	16 (36.4%)	0.576
ECOG			
0	29 (36.3%)	10 (26.3%)	0.518
1	43 (53.8%)	25 (65.8%)	
2	7 (8.8%)	2 (5.3%)	
3	1 (1.3%)	1 (2.6%)	
Missing	8	6	
Charlson Comorbidity Index (CCI)			
0-1	14 (15.9%)	1 (2.3%)	<0.00
2-3	49 (55.7%)	10 (22.7%)	
≥4	25 (28.4%)	33 (75.0%)	
R-ISS Stage at Diagnosis	40 (04 00()		0.004
	18 (24.3%)	11 (27.5%)	0.304
	31 (41.9%)	11 (27.5%)	
	25 (33.8%)	18 (45.0%)	
Unknown	0 (0%)	0 (0%)	
IVIISSING	14	4	
High risk with 1g gain	26 (20 50/)	6 (13 60/)	0 150
High risk without 1g gain	20 (23.0%) 16 (18.0%)	7 (15.0%)	0.159
No high rick	30(10.270)	25 (56 <u>8</u> %)	
Linknown	7 (Q 00/)	£ (13 6%)	
Plasma cell leukemia	/ (0.070) 1 (1 10/)		1
Extramedullary disease	33 (37 5%)	12 (27,3%)	0.33
MvCARe score	00 (01.070)	12 (21:070)	0.00
low (0-1)	23 (26,1%)	14 (31.8%)	0.714
intermediate (2-3)	53 (60.2%)	26 (59.1%)	
high (4)	12 (13.6%)	4 (9.1%)	
Median previous therapy lines [range]	5.00 [3.00, 15.0]	5.00 [3.00, 12.0]	0.772
_enalidomide refractory	74 (84.1%)	36 (81.8%)	0.806
Friple refractory	69 (78.4%)	28 (63.6%)	0.093
Penta refractory	43 (51.2%)	29 (65.9%)	0.135
Previous BCMA-directed Therapy	12 (13.8%)	7 (15.9%)	0.795
Previous HDCT/ASCT	82 (93.2%)	38 (86.4%)	0.213
Previous alloSCT	6 (6.8%)	2 (4.5%)	0.718
Response status at CAR-I	4 (4 40/)	4 (0.00()	0 45 4
	1 (1.1%)	(2.3%)	0.454
	4 (4.0%) 7 (9.09()	3(7.0%)	
	10 (00 70/)	7 (10.3%)	
	18 (20.7%)	9 (20.9%)	
< PR Missing	57 (05.5%)	23 (53.5%)	
IVIISSING			
Covertional MM regimena	17 (71 00/)	22 (7E 00/)	0.000
Covenuonar wive regimens	$\frac{41}{14} (11.2\%)$	<u>کک (۲</u> ۵.۳%) ۲ (17 ۵۰/۱	0.928
Bispecific antibodios	5 (7 60/)	$\frac{J(17.270)}{2(6.007)}$	
Other	0(1.0%)	∠ (0.9%)	
Missing	0 (0%)	U (U%) 15	
wissing		10	
	18 (20 70/)	15 (21 10/)	0.264
1	62 (71 2%)	27 (61 1%)	0.301
2	6 (6 0%)	21 (01.470) 2 (1 50/)	
3	1 (1 10/_)	<u> </u>	
Missing	1	0 (0 %)	
Median Time Diagnosis to CAP T (vers)		U	
vieurari Time Diagnosis (U CAR-I (years)	6.85 [0, 26.2]	7.90 [0, 19.5]	0.233
rangel			
range] Median			
range] Median Median follow-up of survivors (months)			
range] Median Median follow-up of survivors (months) range]	11.9 [0.400, 24.4]	8.55 [0.400, 26.5]	0.401

Our real-world analysis provides additional support that CAR-T cell therapy is feasible and effective in patients with r/r MM aged 70 yrs or older, demonstrating outcomes and toxicities comparable to those observed in younger patients. Therefore, CAR-T cell therapy should not be withheld for eligible patients above 70 yrs with r/r MM.

RESULTS





RESPONSE RATES Aged <70 Years All patients CR/VGPR: 70.0% CR/VGPR: 13.7% CR/VGPR: 17.7% CR/VGPR: 68.9% 5.7% 40.2% 20.7% 65.5% 61.5% 4.6% 8.5% 17.2% 12.3% Best response Response at CART Best response Response at CART **OUTCOMES AFTER IDE-CEL BEFORE MATCHING** Non-Relapse Mortality Relapse Age Group Age Group 🔶 <70 years + <70 years + ≥70 years + ≥70 years p = 0.18 0.25 p = 0.1710 Time after CAR-T [months] Time after CAR-T [months Progression-free Survival **Overall Survival** Age Group + <70 years Age Group 🕂 <70 years + ≥70 years p = 0.53 p = 0.71 Fime after CAR-T [months No. at ris







index. triple-class refractoriness, and remission status as co-variate

DISCLOSURE

No relationships relevant to the contents of this work to disclose.