Real-World Characteristics and Outcomes of Newly Diagnosed Multiple Myeloma Patients Receiving Autologous Stem Cell Transplantation (2018–2024) From the HONEUR Network

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Key Takeaway



Approximately one-third of patients with NDMM undergoing ASCT are ≥65 years of age, indicating real-world practice exceeds traditional trial eligibility. These high-risk profiles associated with reduced survival underscore the importance of prognostic assessments and personalized treatment strategies

Conclusions



Patients received either triplet (52%) or quadruplet (48%) induction therapy followed by ASCT at frontline, over one-quarter of them were ≥65 years, indicating that the real-world transplanted population often exceeds age-based clinical trial criteria



Median TTNT from the start of induction therapy was 60.6 months; while median OS was not reached, 90.7% of the patients were alive at 4 years



Multivariable analysis results indicated that cytogenetics and ISS score were the primary factors influencing TTNT and OS in the transplanted patients independent of age



Disclosures

BGO is an employee of Johnson & Johnson Madrid, Spain

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Introduction

- Eligibility for autologous stem cell transplantation (ASCT) in multiple myeloma clinical trials is primarily determined by age and comorbidities¹⁻³
- Notably, 2025 EHA-EMN guidelines recommend ASCT up to 70 years of age⁴
- Conversely, clinical practice often employs less stringent standards, leading to limited insight into real-world baseline characteristics and outcomes for those with newly diagnosed multiple myeloma (NDMM) receiving ASCT^{2,3,5}
- As new treatment options emerge, real-world data can uncover remaining unmet needs and foster ongoing improvements in the survival of these patients^{2,4,5}
- This study outlines the baseline characteristics, treatment patterns, and outcomes of a cohort of patients with NDMM who underwent ASCT in clinical practice

Methods

- Data were selected from patients starting frontline treatment followed by ASCT from 2018 to 2024 within MyelomaToul - Institut Universitaire du Cancer de Toulouse (IUCT, France)
- Locally collected, patient-level data were analyzed within the HONEUR
- Statistical analyses
- Baseline characteristics (age, sex, International Staging System [ISS] stage, and cytogenetic profiles) and frontline treatment regimens were descriptively analyzed
- Time to next treatment (TTNT), used as a proxy for progression-free survival, and overall survival (OS) were analyzed using the Kaplan-Meier
- A survival subgroup analysis was conducted based on age (<65 and ≥65 years)
- A multivariable analysis was performed to identify the primary drivers of both TTNT and OS

Table 1: Baseline demographics (HONEUR)

		NDMM (n=2945)	Aged <65 years (n=2124)	Aged ≥65 years (n=821)
Median age (range), years		60 (18–79)	57 (18–64)	67 (65–79)
Male, n (%)		1731 (58.8)	1272 (59.9)	459 (55.9)
Age at line of therapy, years, n (%)	<50	371 (12.6)	371 (17.5)	-
	50–64	1753 (59.5)	1753 (82.5)	-
	≥65	821 (27.9)	-	821 (100)
M protein type, n (%)	IgA	469 (15.9)	326 (15.3)	143 (17.4)
	IgG	1246 (42.3)	877 (41.3)	369 (44.9)
	Others	55 (1.9)	45 (2.1)	10 (1.2)
	Unavailable	1175 (39.9)	876 (41.2)	299 (36.4)
	I	789 (26.8)	603 (28.4)	186 (22.7)
ISS stage, ^a n (%)	II	760 (25.8)	534 (25.1)	226 (27.5)
	III	418 (14.2)	299 (14.1)	119 (14.5)
	Unavailable	978 (33.2)	688 (32.4)	290 (35.3)
Cytogenetic profile,ª n (%)	Standard risk	2014 (68.4)	1427 (67.2)	587 (71.5)
	High risk*	369 (12.5)	276 (13)	93 (11.3)
	Indeterminate	562 (19.1)	421 (19.8)	141 (17.2)

Results

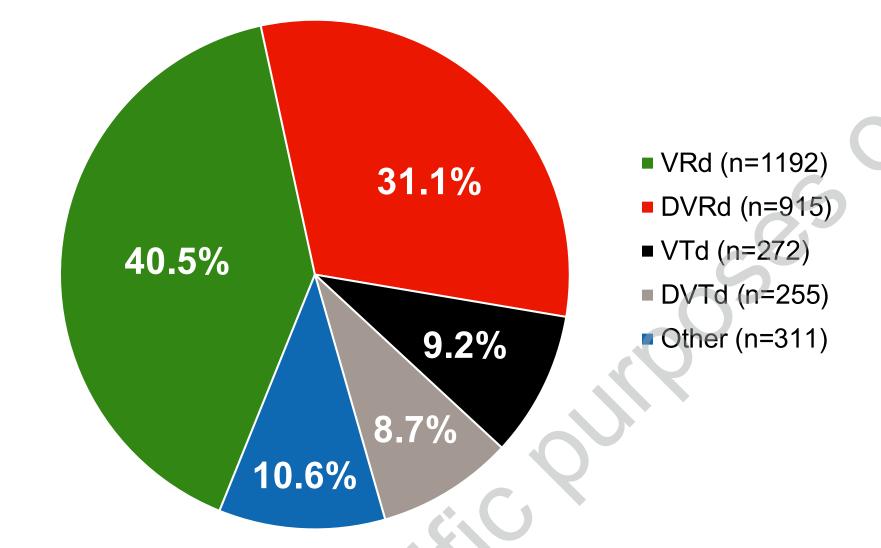
Study population

- 2945 patients underwent ASCT, representing one-third of the patients with NDMM starting treatment between 2018 and 2024
- 72.1% of patients were <65 years (n=2124), 27.9% were ≥65 years (n=821) (**Table 1**)
- Overall, 40% of patients were classified as ISS stage II or III; in the age-stratified analysis, patients aged ≥65 years showed a slightly higher proportion of ISS stage II or III (42%) compared with those aged <65 years (39.2%)
- 12.5% were identified as having high-risk cytogenetics

Treatment regimens at induction (Figure 1)

- The most common induction therapy transplanted patients received was VRd (n=1192; 40.5%); the second most frequent was DVRd (n=915; 31.1%)
- Other induction treatment options included VTd (n=272; 9.2%), DVTd (n=255; 8.7%), and other regimens (n=311; 10.6%)

Figure 1: Induction therapy distribution (n=2945)



), daratumumab: d. dexamethasone: R. lenalidomide: T. thalidomide: V. bortezomib

Efficacy outcomes

- Median follow-up duration from the start of induction therapy was 34.4 months
- Median TTNT from the start of induction was 60.6 months (95% CI, 57.3–64.0) (**Figure 2**)
- While the median OS was not reached, 90.7% (95% CI, 89.1–92.0) of patients were alive at 4 years
- Outcomes stratified by age (Figure 3)
 - Median TTNT was similar in both age subgroups
 - Median OS was not reached for either patient group
- The observed relative mortality risk was 26% higher in patients aged ≥65 years; after adjusting for other prognostic factors, the observed risk increased to 33% but remained nonsignificant (hazard ratio [HR], 1.33; 95% CI, 0.96–1.85)
- This study precedes the widespread uptake and approval of quadruplet anti-CD38 induction, consolidation, and maintenance, which is now considered standard of care (SOC). Therefore, it will be pertinent to repeat these analyses with SOC treatments in the future

Figure 2: Efficacy outcomes (all)

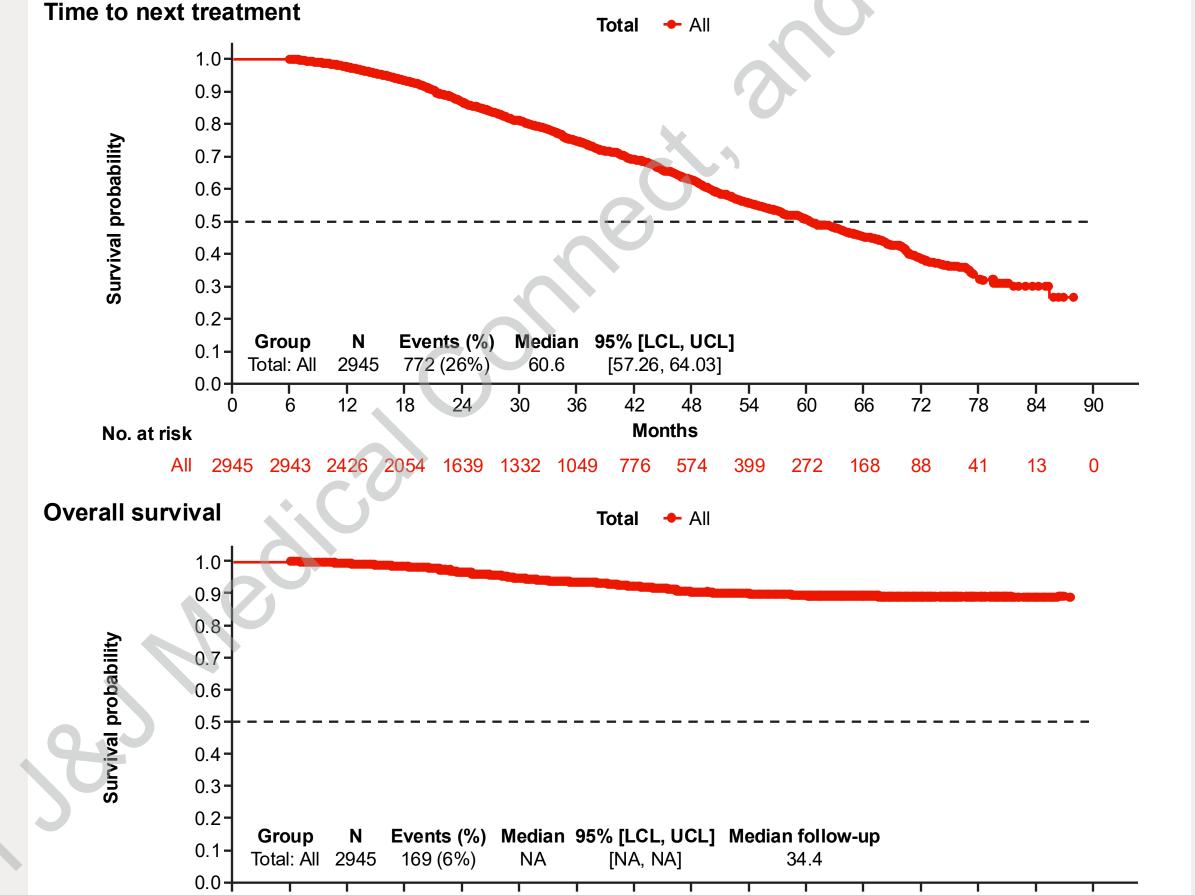
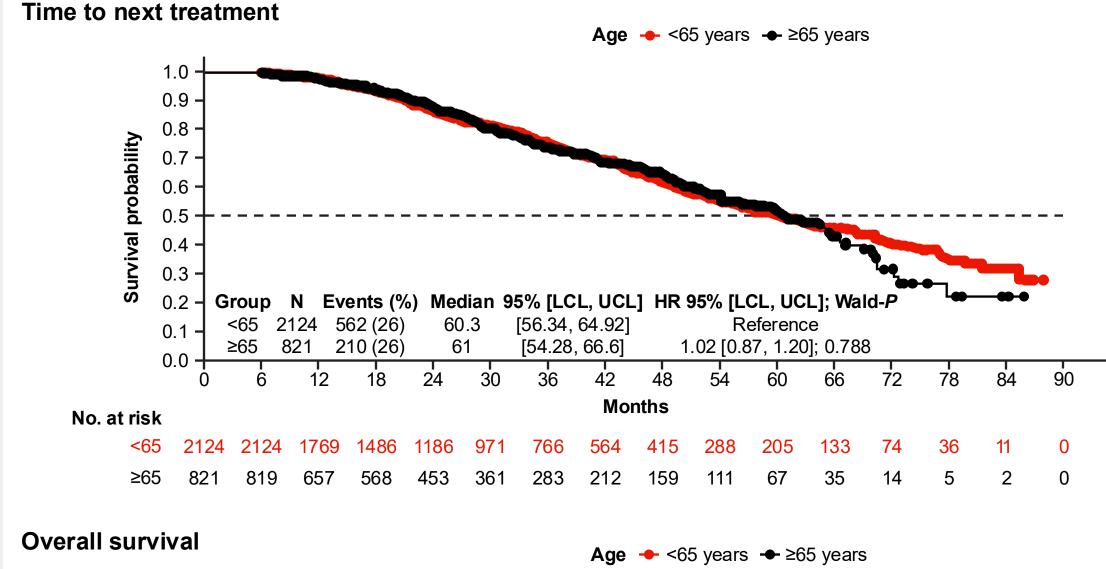
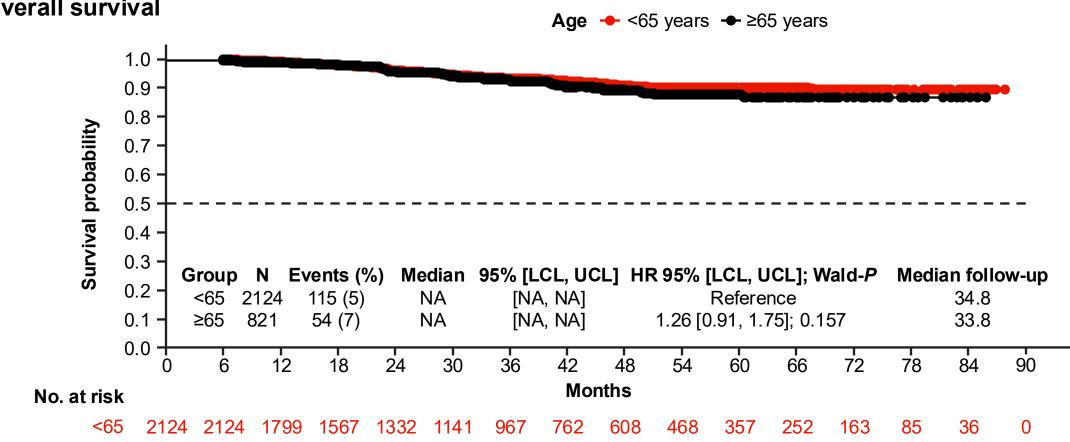


Figure 3: Efficacy outcomes by age group

LCL, lower confidence limit; NA, not applicable; UCL, upper confidence limit.





Multivariable analysis

- Compared with ISS stage I, TTNT was shorter for patients with stage II (HR, 1.5 [95% CI, 1.2–1.9]; *P*<0.001) and stage III disease (HR, 1.9 [95% CI, 1.5–2.5]; *P*<0.001) (**Figure 4**)
- Patients with standard-risk cytogenetics had longer TTNT and OS vs high-risk cytogenetics (TTNT: HR, 1.9 [95% CI, 1.6–2.3]; *P*<0.001; OS: HR, 3.5 [95% CI, 2.4–5.0]; *P*<0.001) (Figure 4)
- Multivariable analysis results indicated that cytogenetics and ISS score were the primary factors influencing TTNT and OS; with these 2 characteristics also driving survival results in the multivariate analysis per age subgroup

Figure 4: Multivariable regression (L1 patients received SCT)

Hazard ratio (multivariable): Time to next treatme	∍nt
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Covariates		Hazard ratio (95% CI)	P value	No. of observations (%)	No. of events (%)
Age, years					
<65	•	Reference		2124 (72.1)	562 (26.5)
≥65		1.06 (0.91, 1.25)	0.456	821 (27.9)	210 25.6)
Sex					
Female		Reference		1214 (41.2)	312 (25.7)
Male	-	1.16 (1.00, 1.34)	0.044	1731 (58.8)	460 (26.6)
ISS					
Stage I	.	Reference		789 (26.8)	136 (17.2)
Stage II		1.48 (1.18, 1.86)	<0.001	760 (25.8)	178 (23.4)
Stage III		1.99 (1.55, 2.54)	<0.001	418 (14.2)	125 (29.9)
Unavailable		1.33 (1.07, 1.66)	0.01	978 (33.2)	333 (34)
Cytogenetic risk					
Standard risk	•	Reference		2014 (68.4)	424 (21.1)
High risk ^a		1.91 (1.58, 2.33)	<0.001	369 (12.5)	140 (37.9)
Unavailable		1.68 (1.36, 2.07)	<0.001	562 (19.1)	208 (37)
M protein					
IgG		Reference		1246 (42.3)	266 (21.3)
IgA	—	0.99 (0.80, 1.24)	0.947	469 (15.9)	117 (24.9)
Others		2.08 (1.28, 3.38)	0.003	55 (1.9)	18 (32.7)
Unavailable		1.37 (1.15, 1.64)	<0.001	1175 (39.9)	371 (31.6)

Hazard ratio (multivariable): Overall survival

Sovariates		(95% CI)	<i>P</i> value	No. of observations (%)	events (%)
Age, years					
<65	•	Reference		2124 (72.1)	115 (5.4)
≥65		1.33 (0.96, 1.85)	0.084	821 (27.9)	54 (6.6)
Sex					
Female	•	Reference		1214 (41.2)	66 (5.4)
Male	+=-	1.21 (0.88, 1.65)	0.239	1731 (58.8)	103 (6)
SS					
Stage I	•	Referenc		789 (26.8)	22 (2.8)
Stage II	-	1.69 (0.99, 2.86)	0.053	760 (25.8)	38 (5)
Stage III		3.23 (1.92, 5.43)	<0.001	418 (14.2)	43 (10.3)
Unavailable	 	1.61 (0.96, 2.71)	0.073	978 (33.2)	66 (6.7)
Cytogenetic risk					
Standard risk		Reference		2014 (68.4)	79 (3.9)
High risk ^a		3.51 (2.45, 5.03)	<0.001	369 (12.5)	52 (14.1)
Unavailable		1.62 (1.00 2.61)	0.049	562 (19.1)	38 (6.8)
M protein					
IgG	•	Reference		1246 (42.3)	56 (4.5)
IgA		1.10 (0.70, 1.71)	0.686	469 (15.9)	31 (6.6)
Others	-	2.49 (0.98, 6.34)	0.056	55 (1.9)	5 (9.1)
Unavailable		1.37 (0.89, 1.92)	0.171	1175 (39.9)	77 (6.6)

^aHigh-risk cytogenetics: presence of del(17p), and/or t(4;14), and/or t(14;16).

1. National Comprehensive Cancer Network (NCCN). Multiple Myeloma. (Version 2.2026). 2. Belotti A, et al. Am J Hematol 2020;95:759-65. 3. Seehaus CM, et al. Hematol Transfus Cell Ther 2024;46 Suppl 6(Suppl 6):S13-S20. 4. Dimopoulos MA, et al. Nat Rev Clin Oncol 2025;22:680-700. 5. Blackburn L, et al. Hemato 2024;5:407-19



Multiple Myeloma

