



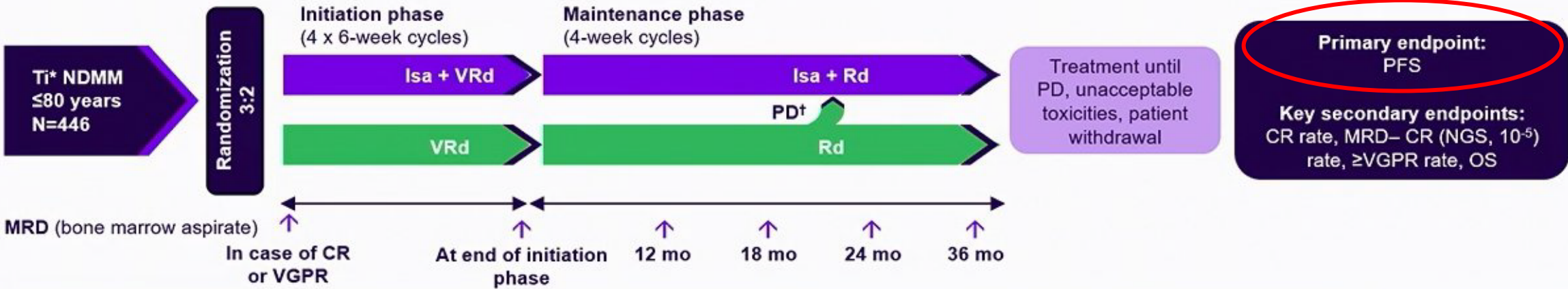
# Post IMS 2024 : actualités en 1<sup>ère</sup> ligne chez les patients non éligibles à la greffe

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# IMROZ: IsaVRd vs VRd



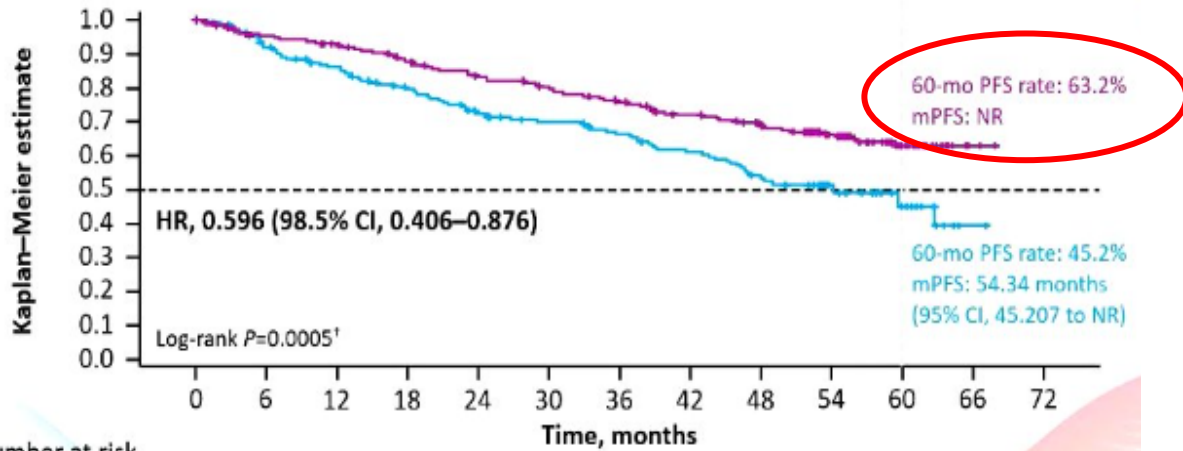
Day		1	8	15	22	29	36	43
Initiation phase	Isa IV (C1 only)	10 mg/kg						
	Isa IV (C2-4)	10 mg/kg						
	V SC	1.3 mg/m <sup>2</sup>						
	R PO†	25 mg						
	d IV/PO§	20 mg						



# IMROZ: IsaVRd vs VRd

## PFS

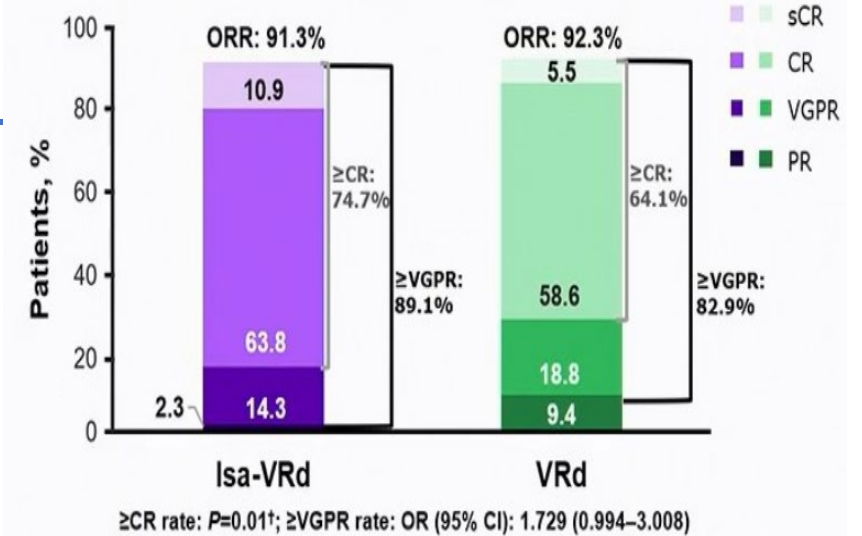
162 PFS events: 84 (31.7%) in Isa-VRd; 78 (43.1%) in VRd\*



Number at risk	0	6	12	18	24	30	36	42	48	54	60	66	72
Isa-VRd	265	243	234	217	201	190	177	164	153	104	43	2	0
VRd	181	155	141	121	104	96	89	81	70	51	20	2	0

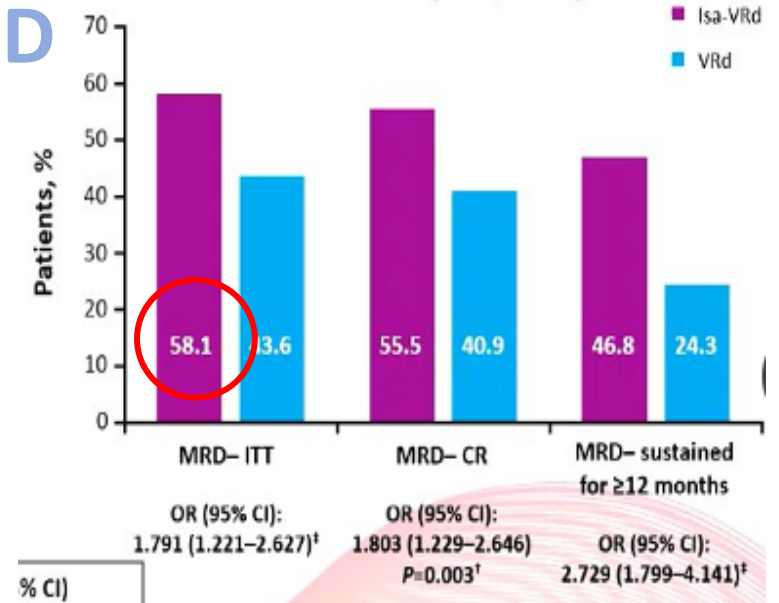
Preferred term, n (%)	Isa-VRd (n=263)		VRd (n=181)	
	Any grade	Grade ≥3	Any grade	Grade ≥3
<b>Infections</b>	240 (91.3)	118 (44.9)	157 (86.7)	69 (38.1)
Pneumonia	79 (30.0)	53 (20.2)	35 (19.3)	23 (12.7)
Upper respiratory tract infection	90 (34.2)	2 (0.8)	61 (33.7)	2 (1.1)
<b>Peripheral sensory neuropathy</b>	143 (54.4)	19 (7.2)	110 (60.8)	11 (6.1)

## Best Overall Response



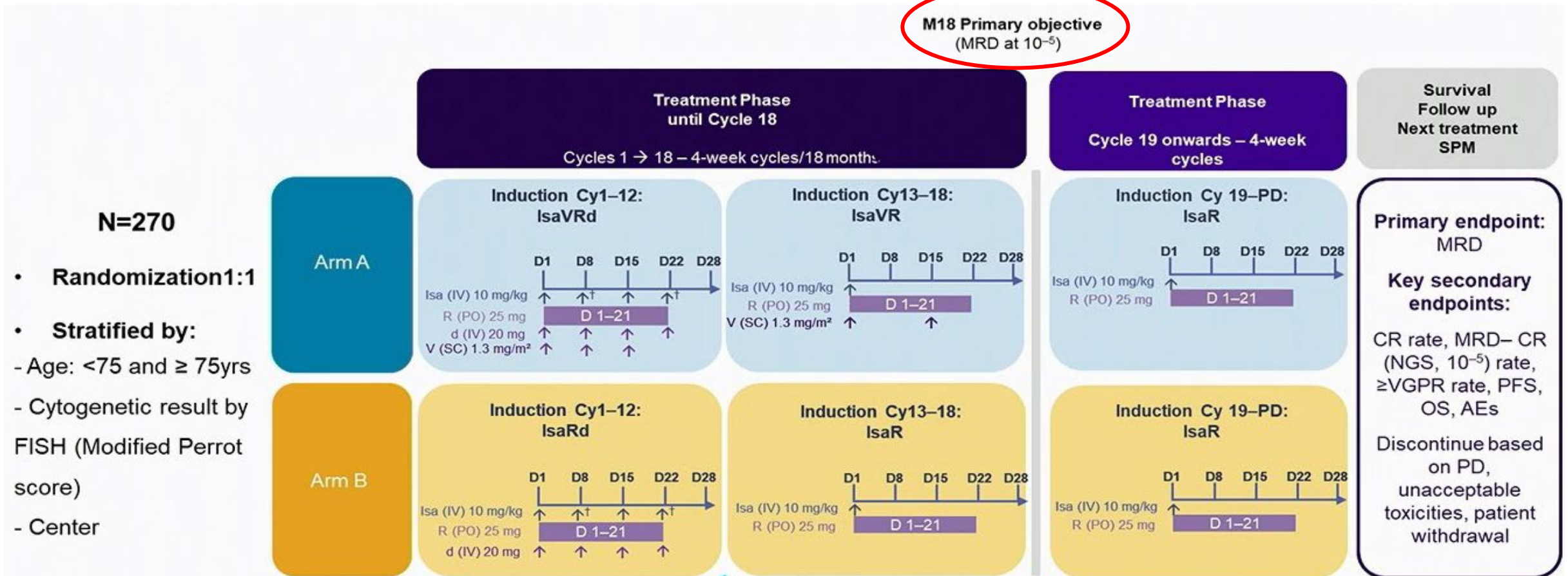
## MRD

### MRD Rate (NGS, \* 10<sup>-5</sup>)



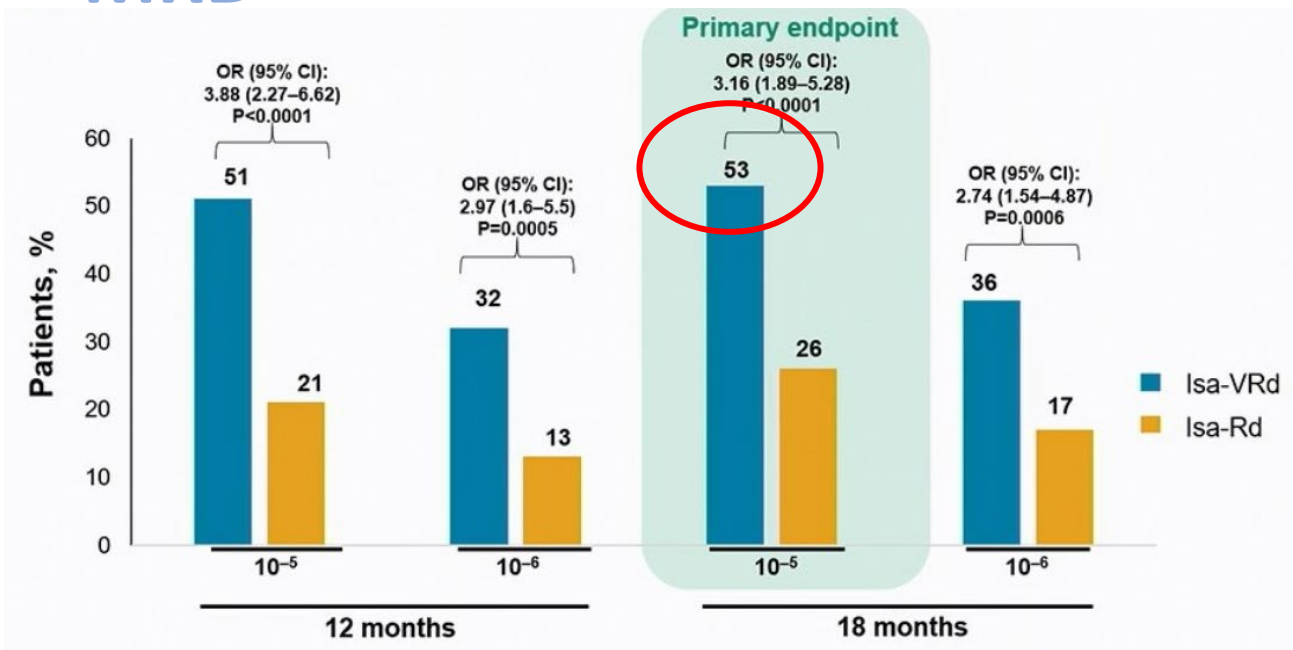
# BENEFIT: IsaVRd vs IsaRd

N=135 each group  
 Median age: 73.2 and 73.6 yo ;  $\geq 75$ : 31 and 36%  
 R ISS 3: 8% each  
 HR : 8 and 10%



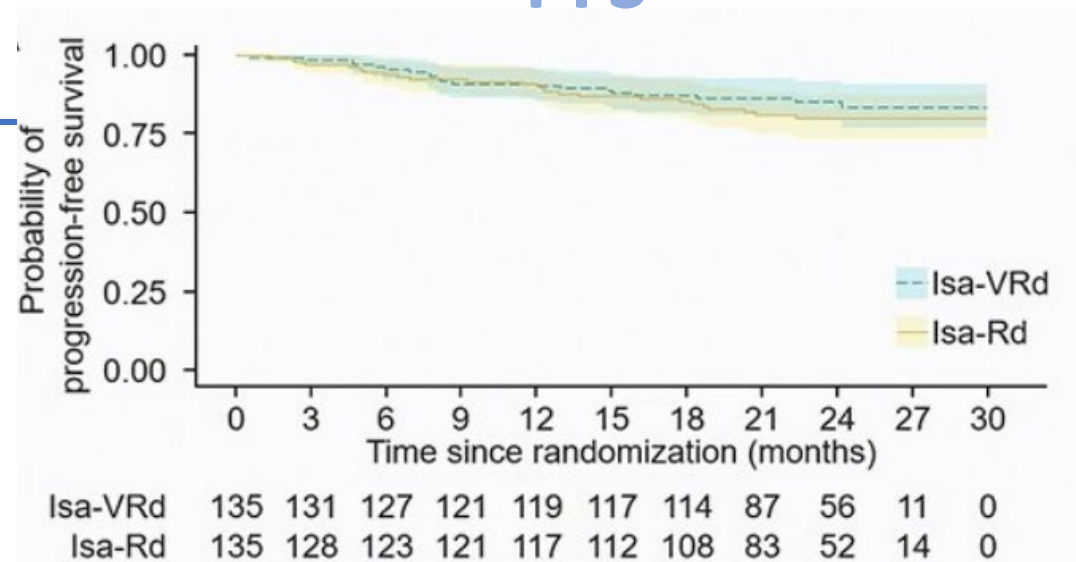
# BENEFIT: IsaVRd vs IsaRd

## MRD



**Estimated 24 months PFS:**  
**85.2% for Isa-VRd**  
**80.0% for Isa-Rd**

## PFS



Event, no. of patients (%)	Isa-VRd (n=135)		Isa-Rd (n=135)	
	Any Grade	≥Grade 2	Any Grade	≥Grade 2
<b>Nonhematologic adverse events (cont'd)</b>				
Eye disorders	20 (15)	10 (7)	19 (14)	12 (8)
SPMs	6 (4)	6 (4)	6 (4)	6 (4)
<b>Infections and infestations</b>				
Infection of other types	61 (45)	48 (36)	48 (36)	35 (28)
Infection of the respiratory system	65 (48)	47 (35)	64 (47)	54 (40)
Covid-19	55 (41)	34 (24)	59 (44)	31 (23)
<b>Nervous system disorders</b>				
Peripheral neuropathy	70 (52)	37 (27) <sup>†</sup>	38 (28)	13 (10) <sup>†</sup>
Other	38 (28)	19 (14)	41 (30)	17 (13)



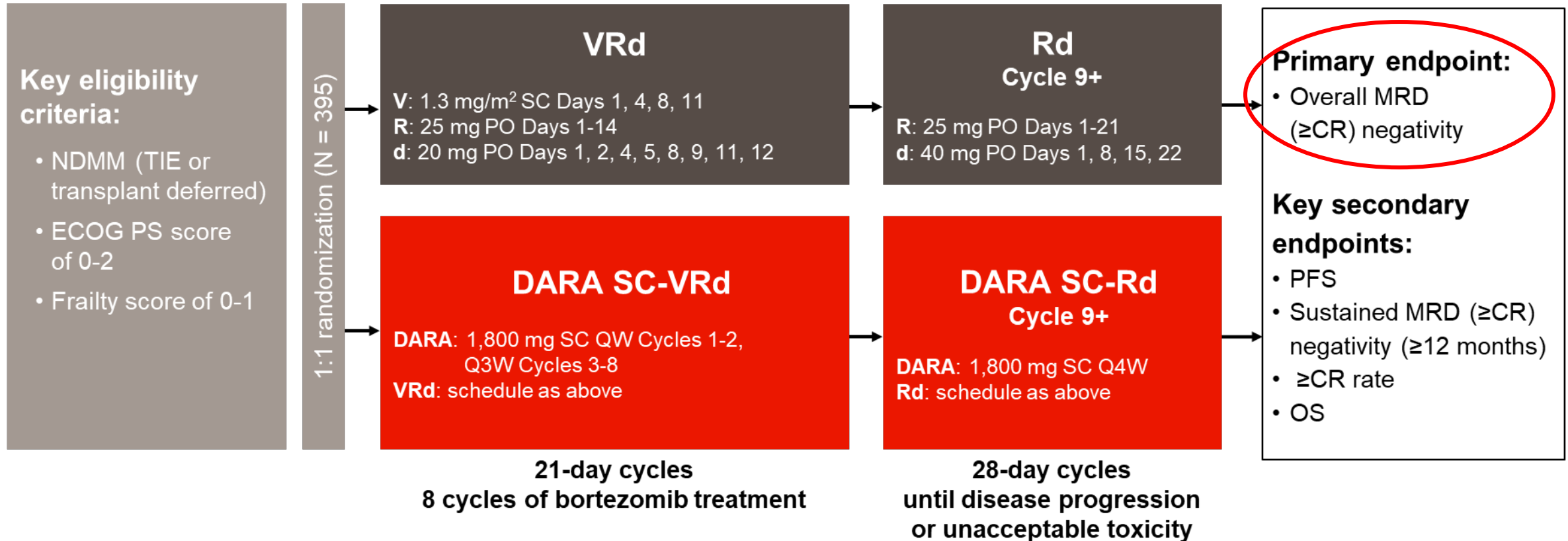
# CEPHEUS: DVRd vs VRd

N=197 et 198

Median age: 70

Transplant ineligible 73% ; transplant deferred 27%

HR 13%



# CEPHEUS: DVRd vs VRd

N=197 et 198

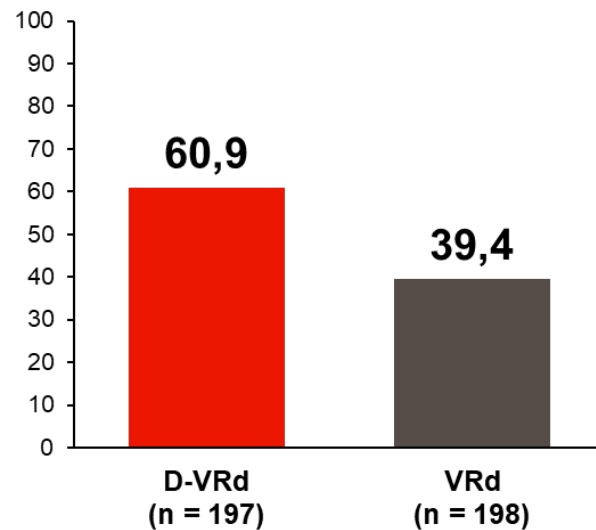
Median age: 70

Transplant ineligible 73% ; transplant deferred 27%

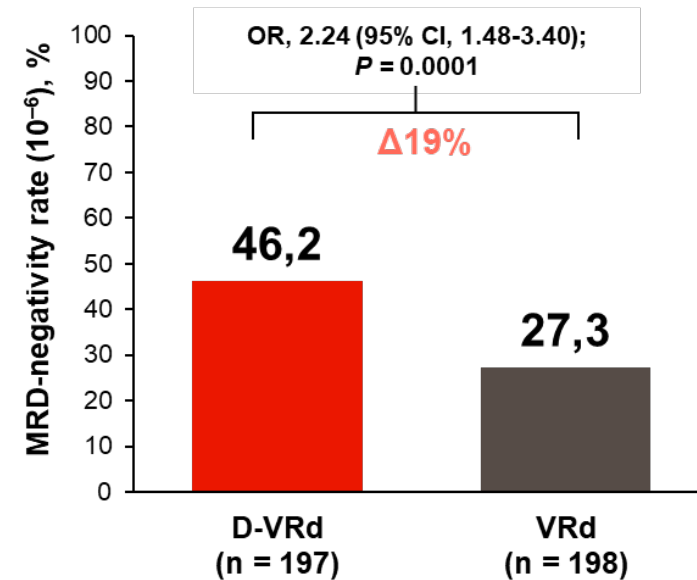
HR 13%

## MRD

**Primary endpoint**  
Overall MRD-negativity rate ( $10^{-5}$ )



Overall MRD-negativity rate ( $10^{-6}$ )



Durée médiane du traitement : **56.3 mois** vs 34.3 mois  
Taux de  $\geq$ CR **81.2%** avec **DVRd** vs 61.6% pour VRd  
Sustained MRD negative  $\geq$ 12 mois = **48.7%** vs 26.3%

# CEPHEUS: DVRd vs VRd

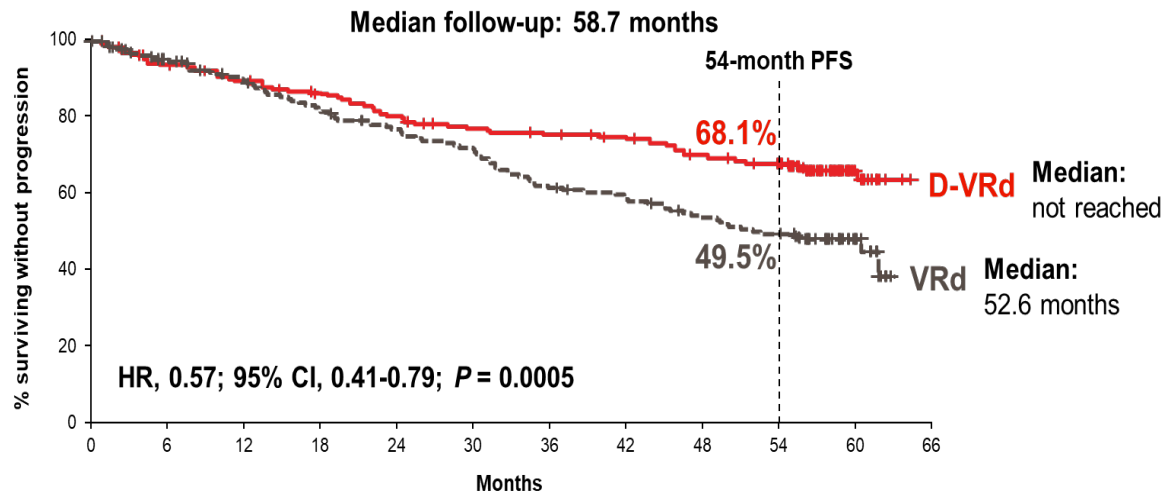
N=197 et 198

Median age: 70

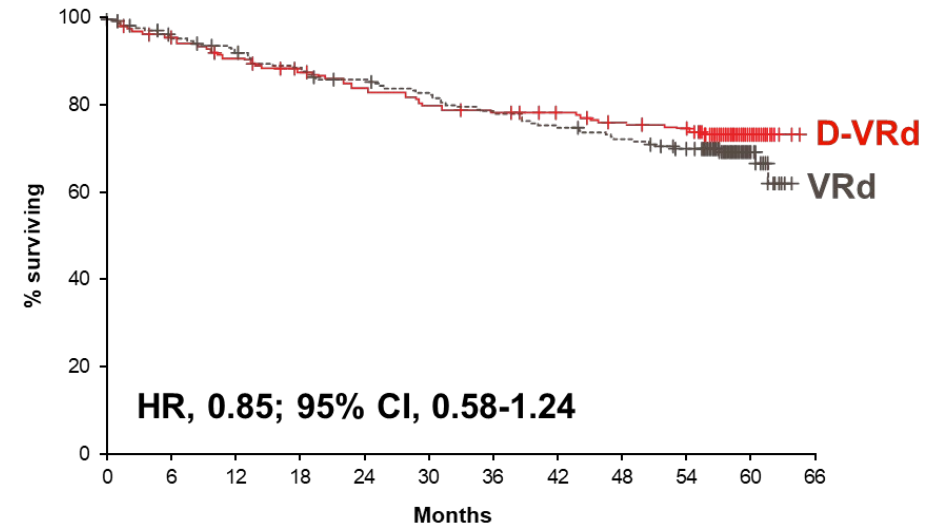
Transplant ineligible 73% ; transplant deferred 27%

HR 13%

## PFS



## OS



## AE

	Any grade	Grade 2	Grade 3 or 4	Any grade	Grade 2	Grade 3 or 4
Peripheral sensory neuropathy	110 (55.8)	60 (30.5)	16 (8.1)	119 (61.0)	70 (35.9)	16 (8.2)

Any grade 3 or 4 TEAE: **92.4** vs **85.6%**  
 TEAE leading to discontinuation: **7.6%** vs **15.9%**



# CC-220-MM-001: Iber + Dara + dex

N=75

Median age: 75 yo ; 59% de  $\geq 75$  yo

HR: 41.3%

R ISS 3 18.7%

## Phase 1: dose escalation

Cohort A IBER
Cohort B IBER + DEX
Cohort E IBER + DARA + DEX
Cohort F IBER + BORT + DEX
Cohort G IBER + CFZ + DEX

## Phase 2: dose expansion<sup>a</sup>

Cohort D IBER <sup>b</sup> + DEX
Cohort I (post BCMA) IBER <sup>b</sup> + DEX
Cohort J1 (NDMM TNE) IBER + BORT + DEX
<b>Cohort K (NDMM TNE) IBER + DARA + DEX</b>

## Key eligibility criteria

- Adults ( $\geq 18$  years of age) with NDMM
- Previously untreated symptomatic MM<sup>a</sup>
- No ASCT planned for initial therapy or ASCT ineligible<sup>b</sup>
- Measurable disease

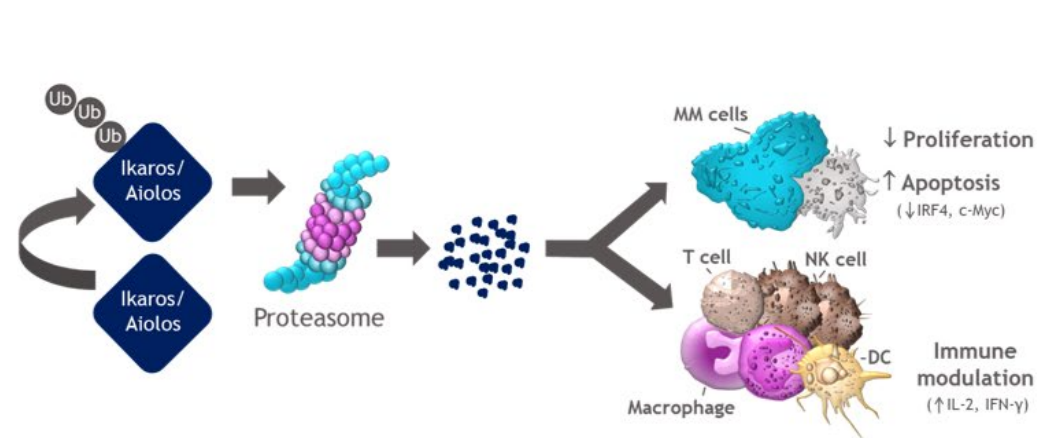
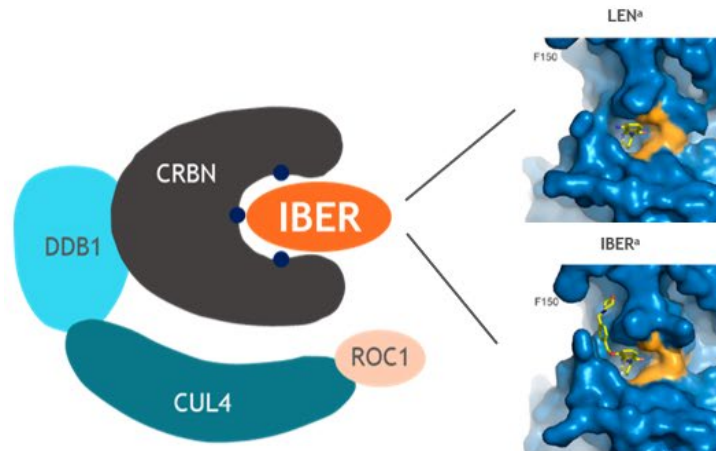
## Treatments<sup>c</sup>

n = 25	IBER <sup>d</sup> (1.0 mg) + DARA <sup>e</sup> + DEX <sup>f</sup> 28-day cycles
n = 25	IBER <sup>d</sup> (1.3 mg) + DARA <sup>e</sup> + DEX <sup>f</sup> 28-day cycles
n = 25	IBER <sup>d</sup> (1.6 mg) + DARA <sup>e</sup> + DEX <sup>f</sup> 28-day cycles

IBER: D1-21 of each cycle  
DARA: 1800 mg SC, QW C1-2, Q2W C3-6, Q4W C  $\geq 7$   
DEX: 40 mg (20 mg if  $> 75$  years of age) QW

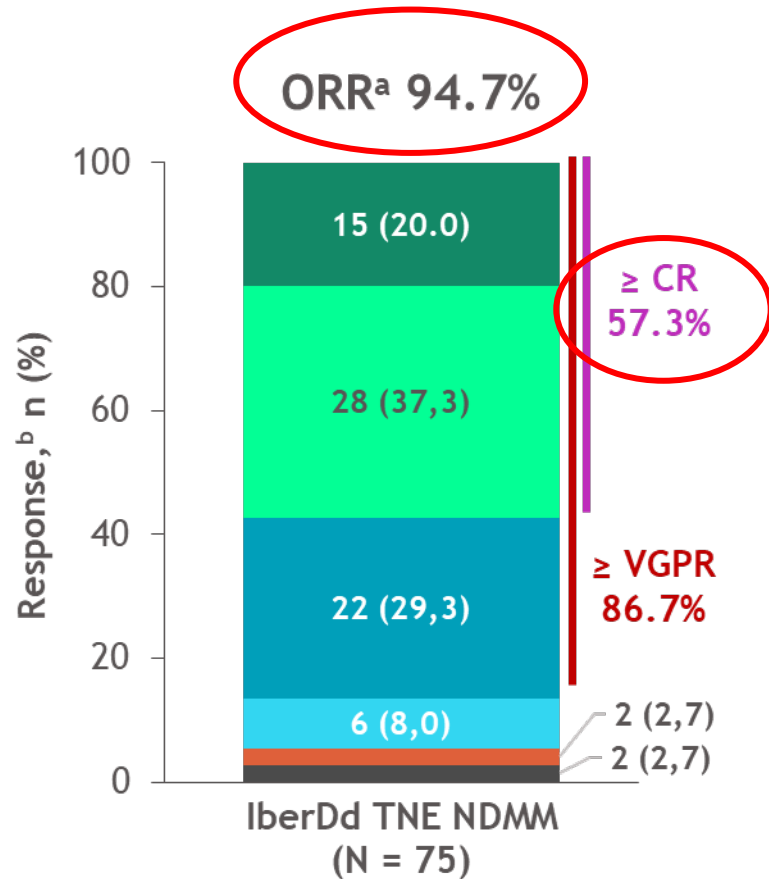
## Objectives

- Evaluation of the preliminary efficacy and safety of IberDd in NDMM-TNE
- PK assessment, MRD evaluation, biomarkers
- MRD evaluation during treatment for patients with  $\geq$  VGPR



# CC-220-MM-001: Iber + Dara + dex

N=75  
 Median age: 75 yo ; 59% de  $\geq 75$  yo  
 HR: 41.3%  
 R ISS 3 18.7%



## MRD

MRD negativity rates<sup>c</sup> evaluated in patients with  $\geq$  VGPR

	Efficacy-evaluable population (N = 73)	Patients with $\geq$ VGPR (N = 65)
MRD negativity rate, <sup>c</sup> n (%)	28 (38.4)	28 (43.1)

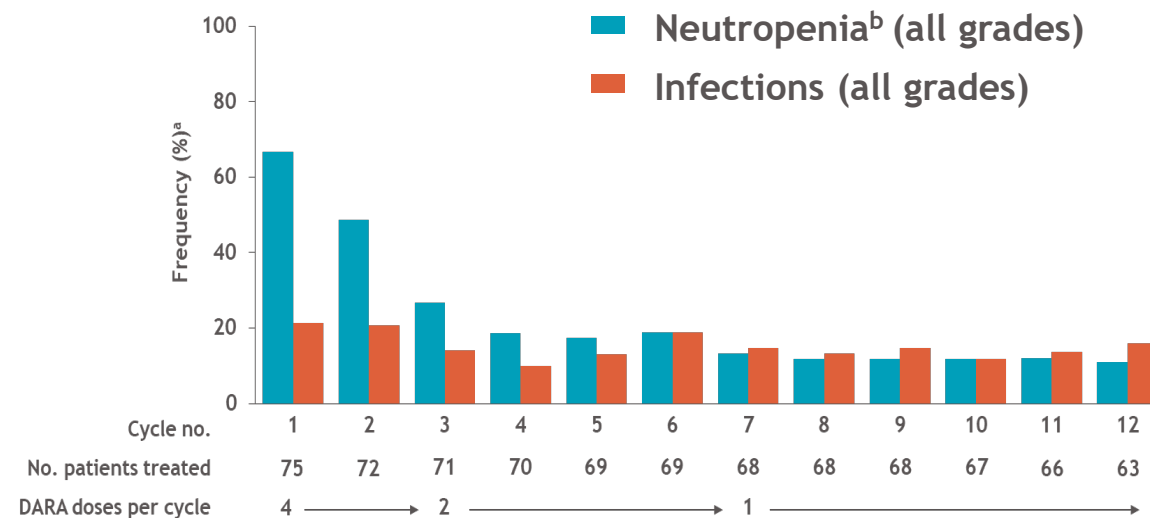
**FUP median: 13.9 mois**  
 Au cutoff, 83% des patients toujours sous traitement  
 durée médiane du traitement = 13.5 mois  
 En médiane, 14 cycles reçus

# CC-220-MM-001: Iber + Dara + dex

N=75  
 Median age: 75 yo ; 59% de  $\geq 75$  yo  
 HR: 41.3%  
 R ISS 3 18.7%

Most common ( $\geq 25\%$ all grade) TEAEs and events of interest, <sup>a</sup> n (%)	IberDd TNE NDMM (N = 75)		
	All grade	Grade 3	Grade 4
Any TEAE	75 (100)	35 (46.7)	36 (48.0)
<b>Hematologic TEAEs</b>			
Neutropenia	60 (80.0)	26 (34.7)	30 (40.0)
Febrile neutropenia	9 (12.0)	9 (12.0)	0
Anemia	25 (33.3)	8 (10.7)	0
Lymphopenia	24 (32.0)	10 (13.3)	5 (6.7)
Leukopenia	17 (22.7)	8 (10.7)	3 (4.0)
Thrombocytopenia	16 (21.3)	2 (2.7)	1 (1.3)
<b>Non-hematologic TEAEs</b>			
Constipation	26 (34.7)	0	0
Rash <sup>b</sup>	24 (32.0)	2 (2.7)	0
Fatigue	20 (26.7)	1 (1.3)	0
Pyrexia	20 (26.7)	2 (2.7)	0
Peripheral edema	20 (26.7)	0	0
Nausea	19 (25.3)	2 (2.7)	0
Diarrhea	17 (22.7)	0	0
<b>Infections</b>			
COVID-19 <sup>c</sup>	24 (32.0)	4 (5.3)	0
Pneumonia <sup>d</sup>	19 (25.3)	17 (22.7)	2 (2.7)

## AE



**Neutropénies plus fréquentes les 2 premiers cycles, puis diminution**