

La mobilizzazione con G-CSF e plerixafor è preferibile alla mobilizzazione con chemioterapia ai fini di autotrapianto?

....le ragioni del Sì....

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CONVEGNO EDUCAZIONALE GITMO

**HOT QUESTIONS
IN TRASPLANTATION
AND CELLULAR
THERAPIES**

Udine

13-14 novembre 2023

Aula Polifunzionale - Ospedale di Udine

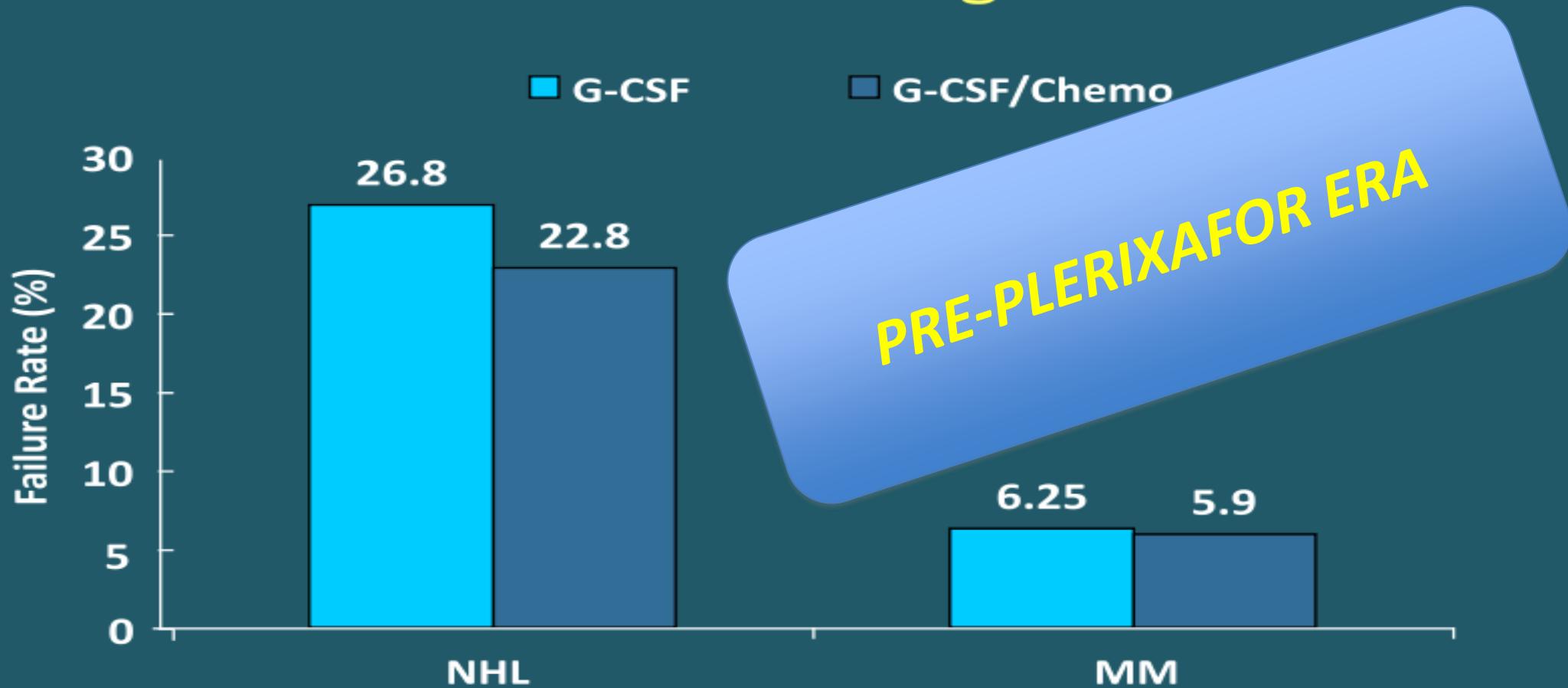
Attilio Olivieri: disclosures

- **Member of Advisory Boards:** Novartis, Jazz Pharma, Incyte
- **Speaker's Bureau:** Therakos: Novartis
- **Ownership of patents in partnership with companies with commercial interests in the health sector: NOTHING TO DECLARE**
- **Equity investments in companies with commercial interests in the health sector: NOTHING TO REPORT**

AGENDA

- *Poor mobilizer: come identificarlo?*
- *Plerixafor, letteratura, real life e la raccolta ideale....*
- *Farmacoeconomia e non solo....*
- *Strategie a confronto nei diversi settings*

Failure Rates of G-CSF ± Chemotherapy Mobilization Regimens



Chemo, chemotherapy; G-CSF, granulocyte colony stimulating factor; MM, multiple myeloma; NHL, non-Hodgkin's lymphoma.

Pusic et al. *Biol Blood Marrow Transplant* 2008;14:1045–1056.

How to identify the “poor mobilizer” ?

- ✓ **parameters proposed to evaluate the extent of mobilization:**
 - ✓ absolute increase (peak) of CD34+ cells in PB
 - ✓ fold increase of CD34+ cells in PB
- ✓ **cumulative aphaeresis yield***
 - ✓ percent candidate patients undergoing ASCT
- ✓ **transplant outcome:** engraftment kinetics
 - ✓ *in a single attempt or with a pre-fixed number of aphaeresis days)



Consensus GITMO su criteri di proven poor mobilizer e predicted poor mobilizer: validazione e refinement su 1317 procedure di mobilizzazione

ORIGINAL ARTICLE

Proposed definition of 'poor mobilizer' in lymphoma and multiple myeloma: an analytic hierarchy process by *ad hoc* working group Gruppo Italiano Trapianto di Midollo Osseo

A Olivieri¹, M Marchetti², R Lemoli³, C Tarella⁴, A Iacone⁵, F Lanza⁶, A Rambaldi⁷ and A Bosi⁸ on behalf of the Italian Group for Stem Cell Transplantation (GITMO)

Table 3 Final definitions of proven and predicted poor mobilizer

A patient with MM or lymphoma candidate to ASCT is a:

Proven poor mobilizer

If he/she received adequate mobilization (G-CSF dose $\geq 10 \mu\text{g}/\text{kg}$ if used alone or $\geq 5 \mu\text{g}/\text{kg}$ after chemo) and he/she shows: peak CD34^+ circulating cell count $< 20/\mu\text{L}$ on days 4-6 after the start of mobilization with G-CSF alone or up to 20 days after chemotherapy and G-CSF
OR in the case of proven poor mobilization, that is: $< 2.0 \times 10^6$ harvested CD34^+ cells per kg (that is, minimum safe dose for each planned ASCT) by ≤ 3 aphereses

Predicted poor mobilizer

If he/she holds at least one major criterion or at least two minor criteria.

Major criteria:

Failed previous mobilization attempt, not otherwise specified.

Previous extensive radiotherapy to marrow bearing tissue.

Full courses of previous therapy, including melphalan, fludarabine or other therapies potentially affecting stem cell mobilization.

Minor criteria:

Advanced phase disease, that is, at least two previous cytotoxic lines

Refractory disease

Extensive BM involvement at mobilization

BM cellularity $< 30\%$ at mobilization

Age > 65 years

Bone Marrow Transplantation
<https://doi.org/10.1038/s41409-017-0051-y>

ARTICLE

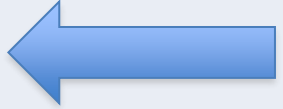
Predicting failure of hematopoietic stem cell mobilization before it starts: the predicted poor mobilizer (pPM) score

Jacopo Olivieri^{1,2} · Immacolata Attolico³ · Roberta Nuccorini³ · Sara Pasquina Pascale³ · Martina Chiarucci¹ · Monica Poiani¹ · Paolo Corradini⁴ · Lucia Farina⁴ · Gianluca Gaidano⁵ · Luca Nassi⁵ · Simona Sica⁶ · Nicola Piccirillo⁶ · Pietro Enrico Pioltelli⁷ · Massimo Martino⁸ · Tiziana Moscato⁸ · Massimo Pini⁹ · Francesco Zallio⁹ · Fabio Ciceri¹⁰ · Sarah Marktel¹⁰ · Andrea Meneghelli¹¹ · Pellegrino Musto¹² · Saveria Capria¹³ · Francesco Merli¹⁴ · Lanza¹⁶ · Giorgina Specchia¹⁷ · Domenico Pastore¹⁷ · Di Nardo¹⁹ · Paolo Perseghin⁷ · Attilio Olivieri¹

End points:

- 1-validate GITMO criteria for pPM;
- 2-improve their predictive ability by building a clinical tool to identify PM **before starting the mobilization attempt!**

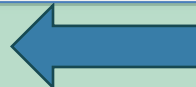
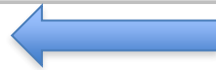
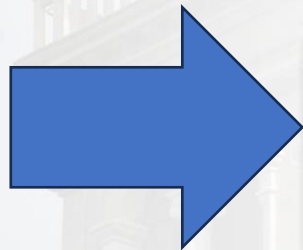
Mobilization outcome in 1318 pts (600 MM; 164HL;554 NHL)

OUTCOME - COLLECTION	
TOTAL HARVEST (CD34 x 10 ⁶ /kg) <2 x 10 ⁶ /kg 2 - 5 x 10 ⁶ /kg >5 x 10 ⁶ /kg	Median 8.9 x 10 ⁶ /kg (range 0 – 63.5) 144 (10.9%) 204 (15.5%) 970 (73.6%)
MOBILIZATION FAILURE	180 pts (13.7%) 
DETERMINANTS OF MOBILIZATION FAILURE	163 / 180 due to LOW CD34 PEAK COUNT (91%) 144 / 180 due to INSUFFICIENT HARVEST (80%) 127 / 180 due to BOTH CRITERIA (71%)
APHERESES CD34 PEAK COUNT	Median 1 aph. (range 1 – 6) Median 85 CD34/mcl (range 0 – 1942)

Olivieri J. et al
BMT 2017

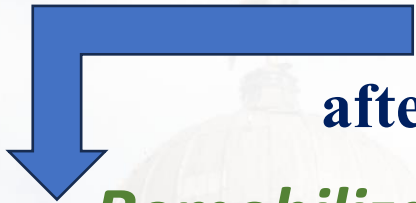
Independent predictive factors for mobilization failure identified by backward variable selection with multiple logistic regression

Predictive factor		β	Odds ratio (95% CI)	Probability (Wald test)
Age class (46-60 years = 1; > 60 years = 2)	BMT 2011	0.3796	1.46 (1.14 - 1.88)	0.003
Diagnosis = NHL		0.5535	1.74 (1.16 - 2.6)	0.007
Disease infiltration \geq 30% at the pre-mobilization BMB	BMT 2011	1.269	3.56 (1.51 - 8.35)	0.004
Number of full chemotherapy courses	BMT 2011	0.5888	1.8 (1.43 - 2.27)	<0.001
At least one previous treatment at risk	BMT 2011	0.7739	2.17 (1.28 - 3.67)	0.004
Pre-mobilization Hb value class (<80 g/l = 1; 80 – 130 g/l = 2)		1.1165	3.05 (1.72 - 5.42)	<0.001
Pre-mobilization WBC < 5 x 10 ⁹ /L		0.7185	2.05 (1.41 - 2.99)	<0.001
Pre mobilization Plt < 170 x 10 ⁹ /L		0.5869	1.8 (1.23 - 2.62)	0.002
Priming with G-CSF alone		2.2513	9.5 (4.75 - 19)	<0.001
Upfront Plerixafor not planned		2.7292	15.32 (5.09 - 46.16)	<0.001
Previous mobilization failure	BMT 2011	1.9059	6.73 (3.67 - 12.34)	<0.001



Plerixafor: phase III 3101 and 3102 studies

MM or NHL patients failing collect. $\geq 0.8 \times 10^6$ CD34+/kg after 2 aphereses
 or $\geq 2 \times 10^6$ CD34+/kg after 4 aphereses ;
 after >7 days of rest considered eligible for randomization



Remobilization rescue protocol

Day	1	2	3	4	5	6	7	8
G-CSF (10 µg/kg)	•	•	•	•	•	•	•	•
Plerixafor (0.24 mg/kg) or Placebo				•	•	•	•	
Apheresis					•	•	•	•

Targets:
 $\geq 2 \times 10^6$ CD34+/kg in ≤ 4 aphereses and $\geq 5 \times 10^6$ CD34+/kg in ≤ 4 aphereses (NHL 3101 study)
 $\geq 6 \times 10^6$ CD34+/kg in ≤ 2 aphereses and $\geq 6 \times 10^6$ CD34+/kg in ≤ 4 aphereses (MM 3102 study),

DiPersio JF, Plerixafor and G-CSF versus placebo and G-CSF to mobilize HSC for AutoHSCT in patients with multiple myeloma. Blood. 2009

DiPersio JF Phase III prospective randomized double-blind placebo-controlled trial of plerixafor plus granulocyte colony-stimulating factor compared with placebo plus G-CSF for mobilization in non-Hodgkin's lymphoma. JCO 2009


Additional plerixafor to granulocyte colony-stimulating factors for haematopoietic stem cell mobilisation for autologous transplantation in people with malignant lymphoma or multiple myeloma (Review)

Hartmann T, Hübel K, Monsef I, Engert A, Skoetz N

- **meta-analysis of AE** did not show a statistically significant difference between the plerixafor and placebo group
- Regarding the primary endpoint of **successful stem cell collection**, significant advantage for those participants randomised to plerixafor
- In **AMD3100-3102 MM**, 95.9% of participants in the plerixafor arm and 88.3% in the placebo arm underwent transplantation.
- In **AMD3100-3101 NHL**, 90% of participants in the plerixafor group but only 55.4% in the control group underwent transplantation.

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Safety and Effectiveness of Plerixafor for Peripheral Blood Stem Cell Mobilization in Autologous Stem Cell Transplantation: Results of a Post-Marketing Surveillance Study

Nobuhiro Tsukada¹ · Momoko Nishikori² · Hiroaki Goto³ · Rie Kanamori⁴ · Satoshi Nishina⁵ · Takashi Seto⁵ · Shinsuke Iida⁶ 

Number of patients with AEs	138 (18.1)
Number of patients with serious AEs	34 (4.5)
Number of patients with ADRs	93 (12.2)
Number of patients with serious ADRs	16 (2.1)

ADRs occurring with a $\geq 1\%$ incidence by PT

System organ class

Blood and lymphatic system disorders	23 (3.0)
Leukocytosis	18 (2.4)
Thrombocytopenia	9 (1.2)
Gastrointestinal disorders	42 (5.5)
Diarrhea	28 (3.7)
Nausea	11 (1.4)
Vomiting	12 (1.6)
Investigations	34 (4.5)
Blood lactate dehydrogenase increased	18 (2.4)
WBC count increased	10 (1.3)
Serum alkaline phosphatase increased	14 (1.8)

Drugs - Real World Outcomes (2022)

	Safety analysis set
Total	764 (100.0)
Age, years, median (min; max)	61.0 (1; 76) ($n = 764$)
Age category, years	
<15	18 (2.4)
15 to <65	522 (68.3)
≥ 65	224 (29.3)
Sex	
Male	417 (54.6)
Female	347 (45.4)
Body weight, kg, mean \pm SD	58.2 \pm 13.5 ($n = 764$)
Diagnosis	
MM	331 (43.3)
ML	360 (47.1)
NHL	344 (45.0)
HL	16 (2.1)
Other ^a	73 (9.6)

The most common ADRs reported in the current study were gastrointestinal disorders (5.5%), laboratory investigations (4.5%), and blood and lymphatic system disorders (3.0%).

Major ADRs included diarrhea ($n = 28$ [3.7%]), leukocytosis ($n = 18$ [2.4%]), blood LDH increased ($n = 18$ [2.4%]), serum ALP increased ($n = 14$ [1.8%]), vomiting ($n = 12$ [1.6%]), and nausea ($n = 11$ [1.4%]).

SCHEDULES FOR PBSC MOBILIZATION

- MOBILIZATION WITH G-CSF ALONE
- G-CSF+CHEMO WITH DISEASE-SPECIFIC SCHEDULES? (E.G. DHAP): Lymphomas
- G-CSF+CHEMO WITH CYTOXAN (2-4 G/M2): MM

HOW AND WHEN ADDING PLERIXAFOR?

- *Upfront....???*
- *On demand or just in time.... (dynamic approach)*
- *Pre-emptive approach (a priori identification of PM....)*
- *Rescue remobilization after failure of 1st attempt*

The «ideal collection»

- Large number of CD34+ (>2 ASCT procedures)...
- ...in one short LK procedure...
- ...withouth need of several days of monitoring...
- ...withouth reaching exagérate Leukocyte count...
- ...*with low PMN contamination...*
- ...*with high immunocompetent cell content...*
- ...*with low/absent tumor cell contamination...*
- ...easy to plan (fixed collection day!)...
- ...no need of toxic mobilizing agents...
- ...no SAE during the mobilization.

**Feasibility
of ASCT**

**Less costs,
better QOL**

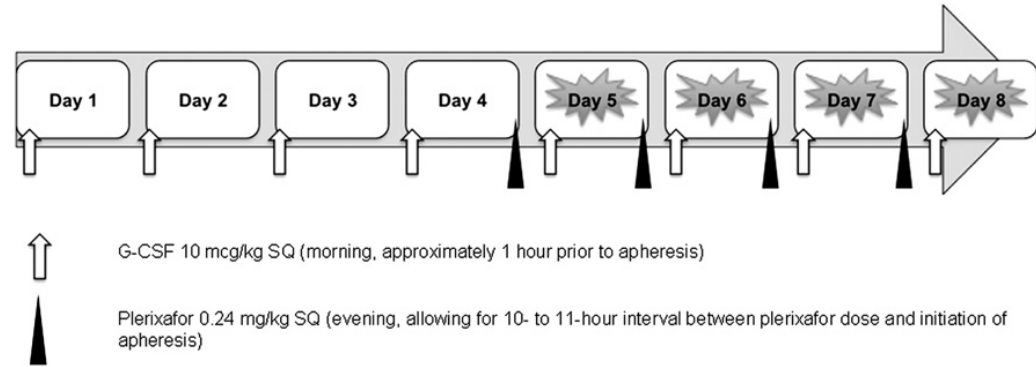
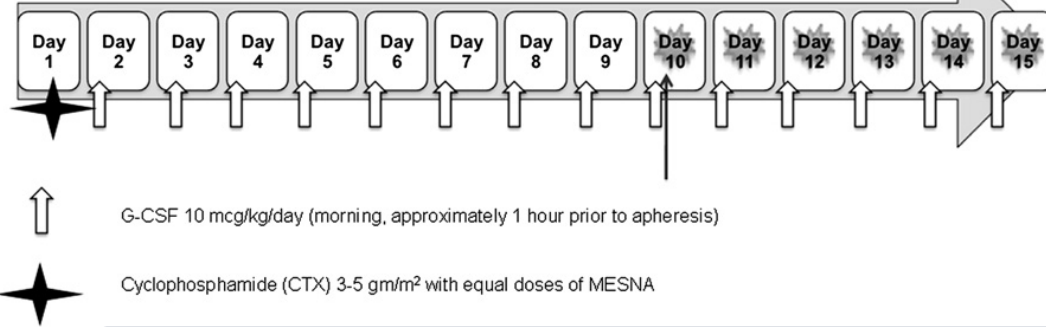
**Minor risk,
Better product**

- **Appropriate
use of
resources**
- **Less
morbidity**

Cost and Clinical Analysis of Autologous Hematopoietic Stem Cell Mobilization with G-CSF and Plerixafor Compared to G-CSF and Cyclophosphamide

Paul Shaughnessy,¹ Miguel Islas-Ohlmayer,¹ Julie Murphy,² Maureen Hougham,¹

G-CSF+plerixafor can mobilize autologous HSC as well as G-CSF+chemotherapy, with similar transplant engraftment outcomes.



Median total cost of mobilization was not different between the plerixafor/G-CSF and control groups (\$14,224 versus \$18,824; P=0.545)

	G-CSF+ Plerixafor	CHEMO+ G-CSF
Median total CD34 ⁺ cells × 10 ⁶ /kg, n (range)	10.7 (3.5-37.9)	11.6 (2.1-69.3)
Number of patients collecting ≥2 × 10 ⁶ CD34 ⁺ cells/kg (%)	33 (100%)	33 (100%)
Number of patients collecting ≥5 × 10 ⁶ CD34 ⁺ cells/kg (%)	31 (94%)	25 (76%)
Number of MM patients collecting ≥ 3 × 10 ⁶ CD 34 ⁺ cells/kg (%)	13/13 (100%)	11/13 (85%)
Number of MM patients collecting ≥6 × 10 ⁶ CD 34 ⁺ cells/kg (%)	20/20 (100%)	18/20 (90%)
Median number of apheresis days (range)	1 (1-4)	1 (1-4)
Number of patients initiating apheresis on scheduled day (%)	33 (100%)	29 (88%)
Number of patients requiring weekend apheresis (%)	0	16 (48%)
Total number of weekend apheresis procedures	0	19

- Kallmeyer et al, BMT, 2011 ; Sinclair et al, J Clin Apher, 2013 : pre-emptive Plerixafor (used in 16% of cases) associated to 91% mobilization success. Incremental cost of 1456 euros/patient.
- Vishnu et al, Transfusion, 2012: estimate of Plerixafor pre-emptive benefit: 19300 \$/patient
- NHS Scotland, Scottish Medicines consortium, 2009: Plerixafor achieves 0.41 QALY at cost of £15,561 compared to G-CSF+CY
- *Shaughnessy et al, Biol Blood Marrow Transplant 2011: similar costs of CTX+G-CSF , compared to Plerixafor*
- Martin et al, J Clin Apher, 2016 Plerixafor upfront in lymphoma patients: significant advantage (-3828 \$;) not confirmed in MM patients (+5245 \$)
- D Laszlo et al Transfus Apher Sci 2020 Mobilization cost components significantly lower for G-CSF+plerixafor vs G-CSF+CTX for hospital stay (3080 euros vs 9653 euros; for mobilizing agent, significant difference with 5470 euros for G-CSF and plerixafor compared with 1140 euros Cy+G-CSF
- Lazzaro et al, 2021, A comparison of chemo-free strategy with G-CSF plus plerixafor on demand versus intermediate-dose cyclophosphamide and G-CSF as PBSC mobilization in newly diagnosed multiple myeloma patients: An Italian explorative cost analysis

**Plerixafor
on demand:
preliminary
experiences
in different
settings
showed
the cost-
effectiveness of
Plerixafor+G-CSF**

Bone Marrow Transplantation (2021) 56:1876–1887

Chemotherapy-based versus chemotherapy-free stem cell mobilization (\pm plerixafor) in multiple myeloma patients: an Italian cost-effectiveness analysis

Carlo Lazzaro¹ · Luca Castagna² · Francesco Lanza³ · Daniele Laszlo⁴ · Giuseppe Milone⁵ · Luca Pierelli⁶ · Riccardo Saccardi⁷

chemotherapy-free mobilization (\pm on-demand plerixafor) is a cost-effective healthcare program for patients with MM who are eligible for autologous SCT in Italy.

Table 3 Base case cost-effectiveness analysis (€2019).

Mobilization schemes	Cost	Effectiveness ^a
CTX 4 g/m ² + G-CSF \pm on-demand PLX	€ 9238.44	0.714
G-CSF \pm on-demand PLX	€ 8039.85	0.766

Parameters	G-CSF (\pm on-demand PLX)	CTX + G-CSF (\pm on-demand PLX)
Demographic and anthropometric parameters		
Age, years, mean (range)	57.62 (51.38, 62.38)	57.62 (51.38, 62.38)
Bodyweight, kg, mean (95% CI)	70.00 (26.34, 134.61) ^a	70.00 (26.34, 134.61) ^a
Height, cm, mean (95% CI)	170.00 (136.68, 203.32) ^a	170.00 (136.68, 203.32) ^a
Mobilization parameters, mean (95% CI)		
Number of vials of on-demand PLX	0.71 (0.00, 3.04) ^a	0.14 (0.00, 1.22) ^a
Days of hospitalization for CTX administration	0	2.00 (0.75, 3.85) ^a
Days of hospitalization for febrile neutropenia	0	0.35 (0.00, 2.04) ^a
Number of RBC transfusions	0	0.24 (0.00, 1.68) ^a
Number of PLT transfusions	0	0.12 (0.00, 1.10) ^a

In this model, patients undergoing chemotherapy-free and chemotherapy-based mobilization received a mean of 0.71 and 0.14 vials of on-demand plerixafor, respectively

Cost analysis of a randomized stem cell mobilization study in multiple myeloma

Ville Varmavuori^{1,2}, Raija Silvennoinen^{3,4}, Pekka Anttila⁴, Marjaana Säily⁵, Marja Sankelo⁶, Mervi Putkonen⁷, Jouni Ahonen⁸, Eija Mahlamäki⁹, Pentti Mäntymaa⁹, Eeva-Riitta Savolainen¹⁰, Kari Remes^{7,11}, Esa Jantunen³

Ann Hematol 2016 Oct;95

MM patients received mobilization with CY+G-CSF (A) or with G-CSF alone (B): 80 MM pts *treated with RVD* for three 21-day cycles and randomized 1:1 (Plerixafor on demand)

The median total costs mobilization were significantly higher in arm A (3855 € vs. 772 €). Cumulative median cost of the mobilization and collection phases was significantly *lower in arm B than in arm A (8524 € vs. 11,622 €, p = 0.012)*.

No significant difference between the arms in the total median costs of ASCT ($n = 59$) (34,997 € in arm A vs. 31,981 € in arm B).



Mobilization with G-CSF alone, a preferable mobilization method for MM patients in terms of mobilization and apheresis costs. In addition, it requires less hospital resource utilization.

A comparison of peripheral blood stem cell collection outcomes for multiple myeloma; mobilization matters in the era of IMiD induction

13 April 2023

eJHaem

Thea Chandler¹ | Christopher Parrish¹ | Marina Karakantza²

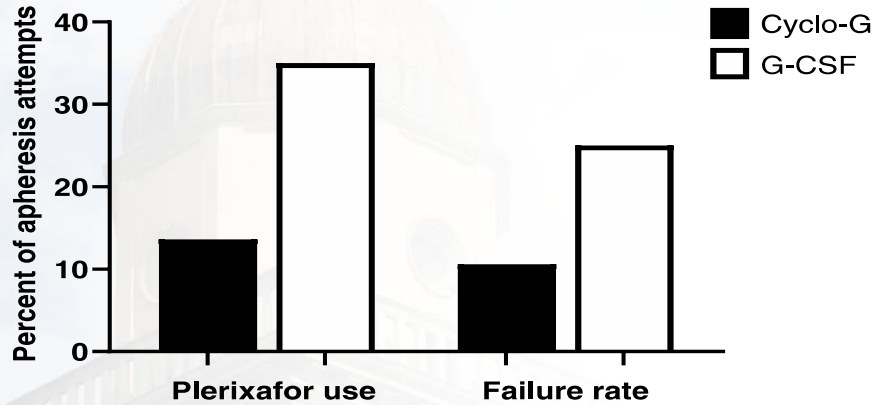
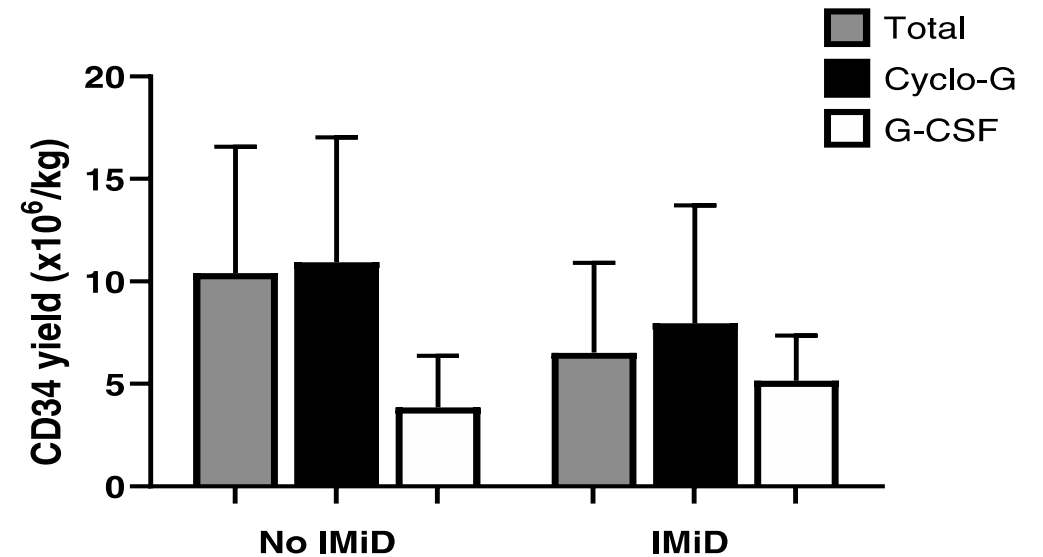


FIGURE 3 Rescue plerixafor use and failure rates by mobilization regimen.

Cyclophosphamide plus G-CSF (cyclo-G) mobilization yielded more CD34⁺ cells (8.94 vs. 4.88 × 10⁶/kg, *p* = < 0.0001) over fewer days (1.6 vs. 2.4 days, *p* = 0.007), and required fewer doses of salvage Plerixafor than G-CSF only (13.6% vs. 35%, *p* = 0.0407). IMiD-containing induction impaired all of these factors. CD34⁺ doses > 8 × 10⁶/kg were more frequent with Cyclo-G (62% vs. 11%, *p* = 0.0001), including for those receiving IMiD 1st line induction (50% vs. 13.3%, *p* = 0.0381). Note that 92.6% of those receiving IMiD-free inductions were mobilized with Cyclo-G.

NHS indicative price of Plerixafor equating to £4882.77 per 20 mg vial (the standard dose for patients up to 84 kg), there are potential

The novel agents used in modern induction regimens (e.g Daratumumab) have been shown to impair yields, increasing the importance of optimizing mobilization regimens in the first instance. Furthermore, as cellular therapies become established in the management of multiple myeloma emerging data highlights the potential benefits of stem cell top up in the management of the haematological toxicities of these therapies. Our findings support re-adoption of Cyclo-G as the gold standard for mobilization to optimize PBSC harvesting and ensure sufficient cells for subsequent ASCTs.



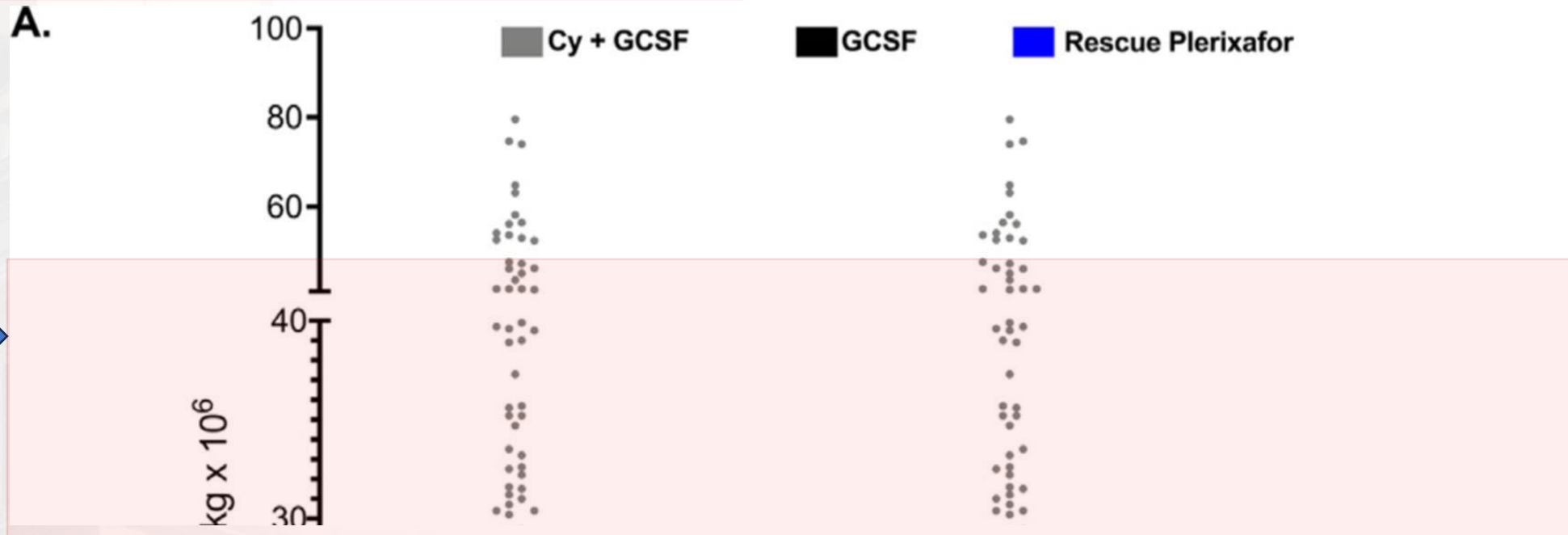
Stem Cell Mobilization in Multiple Myeloma: Comparing Safety and Efficacy of Cyclophosphamide +/- Plerixafor versus Granulocyte Colony-Stimulating Factor +/- Plerixafor in the Lenalidomide Era

Andrew Johnsrud^{1,2}, Abdullah Ladha^{2,3}, Lori Muffly^{1,2}, Parveen Shiraz^{1,2}, Gary Goldstein², Victoria Osgood², Judith A. Shizuru^{1,2}, Laura Johnston^{1,2}, Sally Arai^{1,2}, Wen-Kai Weng^{1,2}, Robert Lowsky^{1,2}, Andrew R. Rezvani^{1,2}, Everett H. Meyer^{1,2}, Matthew J. Frank^{1,2}, Robert S. Nearin^{1,2}, David B. Miklos^{1,2}, Surbhi Sidana^{1,2,*}

Patients stratified into 3 groups:

1. No previous Lena exposure (60);
2. ≤ 6 cycles of Lena (199);
3. ≥ 6 cycles of Lena exposure (138)

Previous lenalidomide exposure for ≤ 6 cycles did not impair cell yield compared with patients without lenalidomide exposure



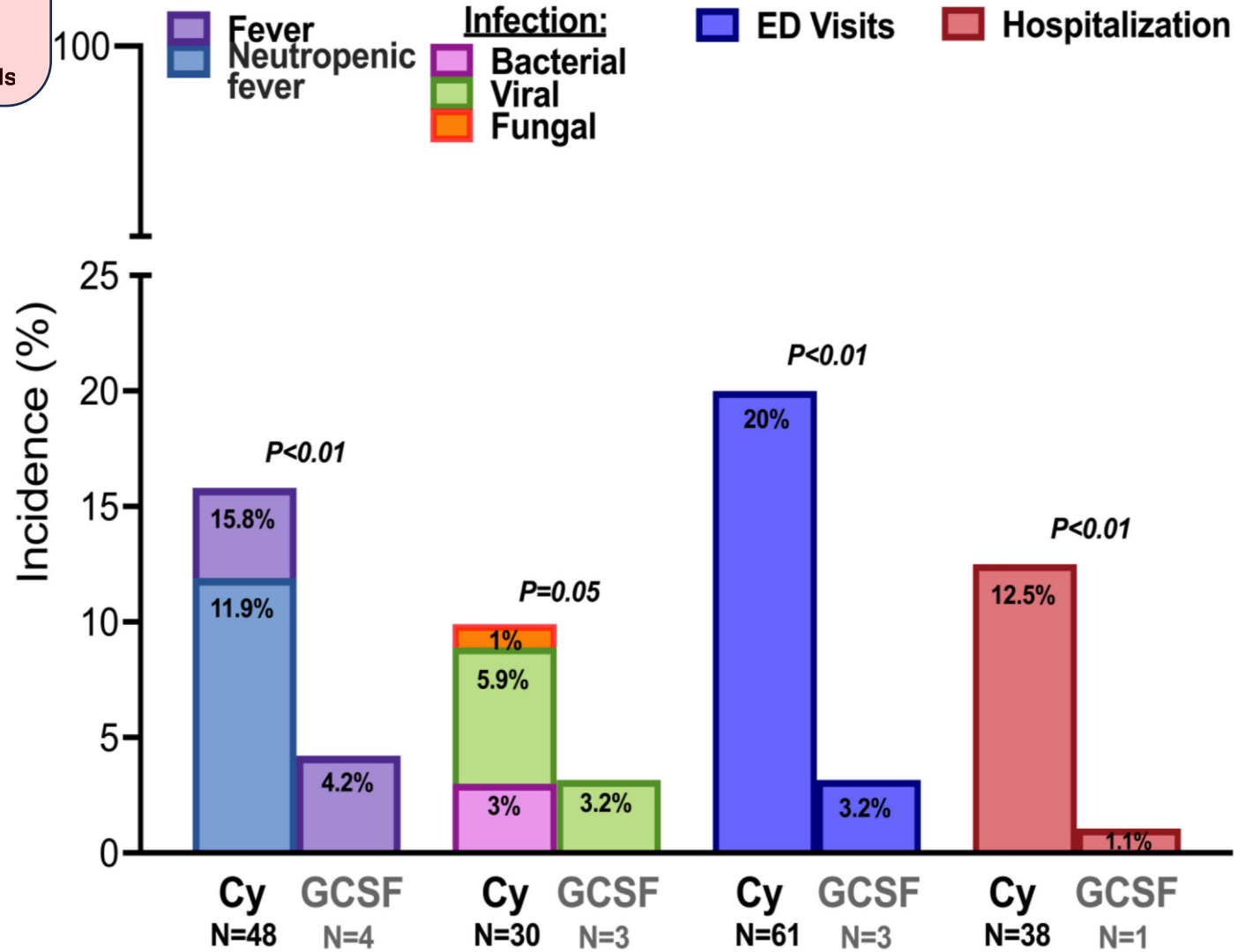
Stem Cell Mobilization in Multiple Myeloma: Comparing Safety and Efficacy of Cyclophosphamide +/- Plerixafor versus Granulocyte Colony-Stimulating Factor +/- Plerixafor in the Lenalidomide Era

Andrew Johnsrud^{1,2}, Abdullah Ladha^{2,3}, Lori Muffly^{1,2}, Parveen Shiraz^{1,2}, Gary Golds

GCSF (+/- P) vs. Cy/GCSF (+/- P)

-20.2% Medicare Reimbursement
P<0.01

-17.4% Institutional Charges
P=0.01



Previous IMiD exposure

- **In patients with lenalidomide exposure;** P+G-CSF and CTX have been shown to be effective approaches.
- Mobilization with G-CSF alone should be avoided in patients with extensive (>4 to 6 cycles) lenalidomide pretreatment.

Strong recommendations in favour of CTX versus P+G-CSF cannot be made owing to a lack of data; controlled prospective trials comparing the 2 strategies are needed.....

Stem cell yield and transplantation in transplant-eligible newly diagnosed multiple myeloma patients receiving daratumumab + bortezomib/thalidomide/dexamethasone in the phase 3 CASSIOPEIA study

Letters to the Editor

Cyrille Hulin et al....HOVON

haematologica | 2021; 106(8)

	D-VTd	VTd
Patients with stem cell mobilization	N=506	N=492
PBSC mobilizing agents, n (%) ^a		
Cyclophosphamide/G-CSF	506 (100)	492 (100)
Plerixafor	110 (21.7)	39 (7.9)
PBSC apheresis performed, n (%)	504 (99.6) ^c	490 (99.6) ^d
Patients with PBSC apheresis performed	N=504	N=490
Number of days of apheresis, mean [SD] (range)	1.9 [0.92] (1–6)	1.4 [0.67] (1–4)
≥5 × 10 ⁶ /kg, n (%)	380 (75.4)	434 (88.6)
Patients who underwent transplantation, n (%)	489 (97.0)	484 (98.8)
Number of CD34 ⁺ stem cells transplanted (10 ⁶ /kg), Mean [SD] (range)	3.6 [1.59]	5.0 [2.80]

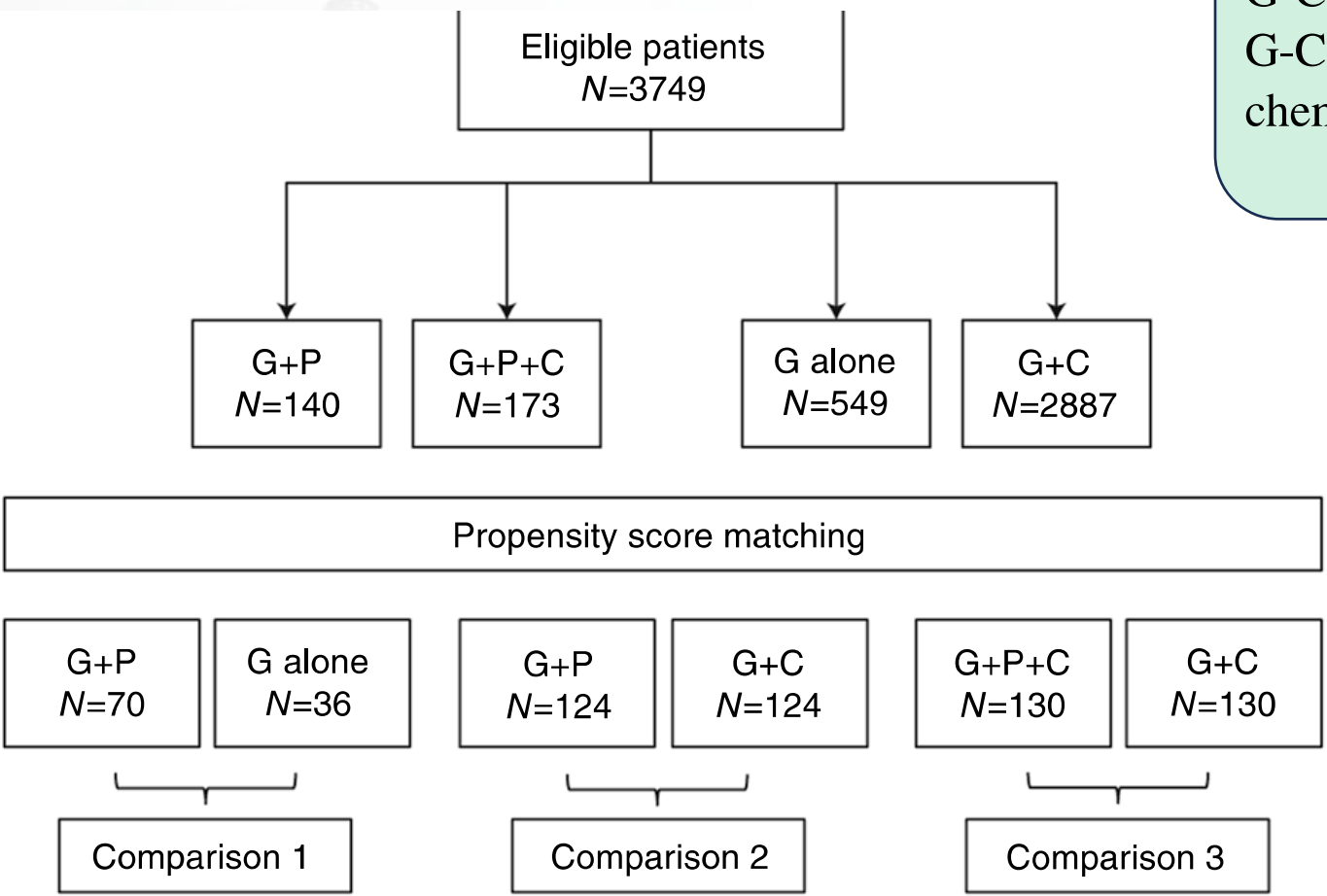


Daratumomab
did not impair
feasibility
and safety of
transplantation
with successful
engraftment,
even though
stem cell yield
was lower

Analysis of data collected in the European Society for Blood and Marrow Transplantation (EBMT) Registry on a cohort of lymphoma patients receiving plerixafor

Anna Sureda¹ · Christian Chabannon² · Tamás Masszi³ · David Pohlreich⁴ · Christof Scheid⁵ ·

G-CSF + plerixafor versus G-CSF alone;
G-CSF + plerixafor versus G-CSF + chemotherapy;
G-CSF + plerixafor + chemotherapy versus G-CSF + chemotherapy.



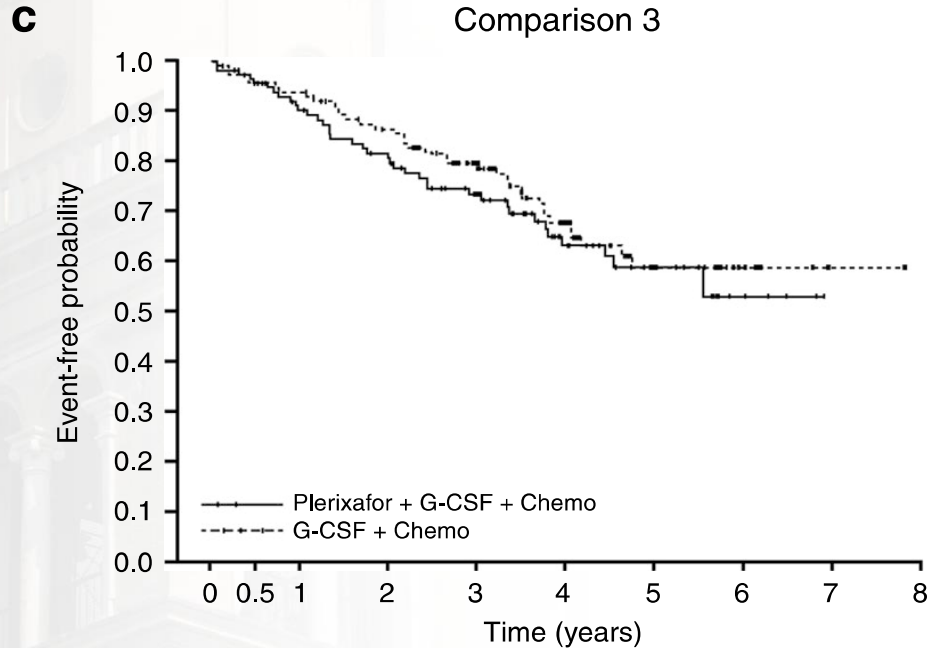
*PFS, OS, and CIR
were numerically
similar
between comparators*

ARTICLE

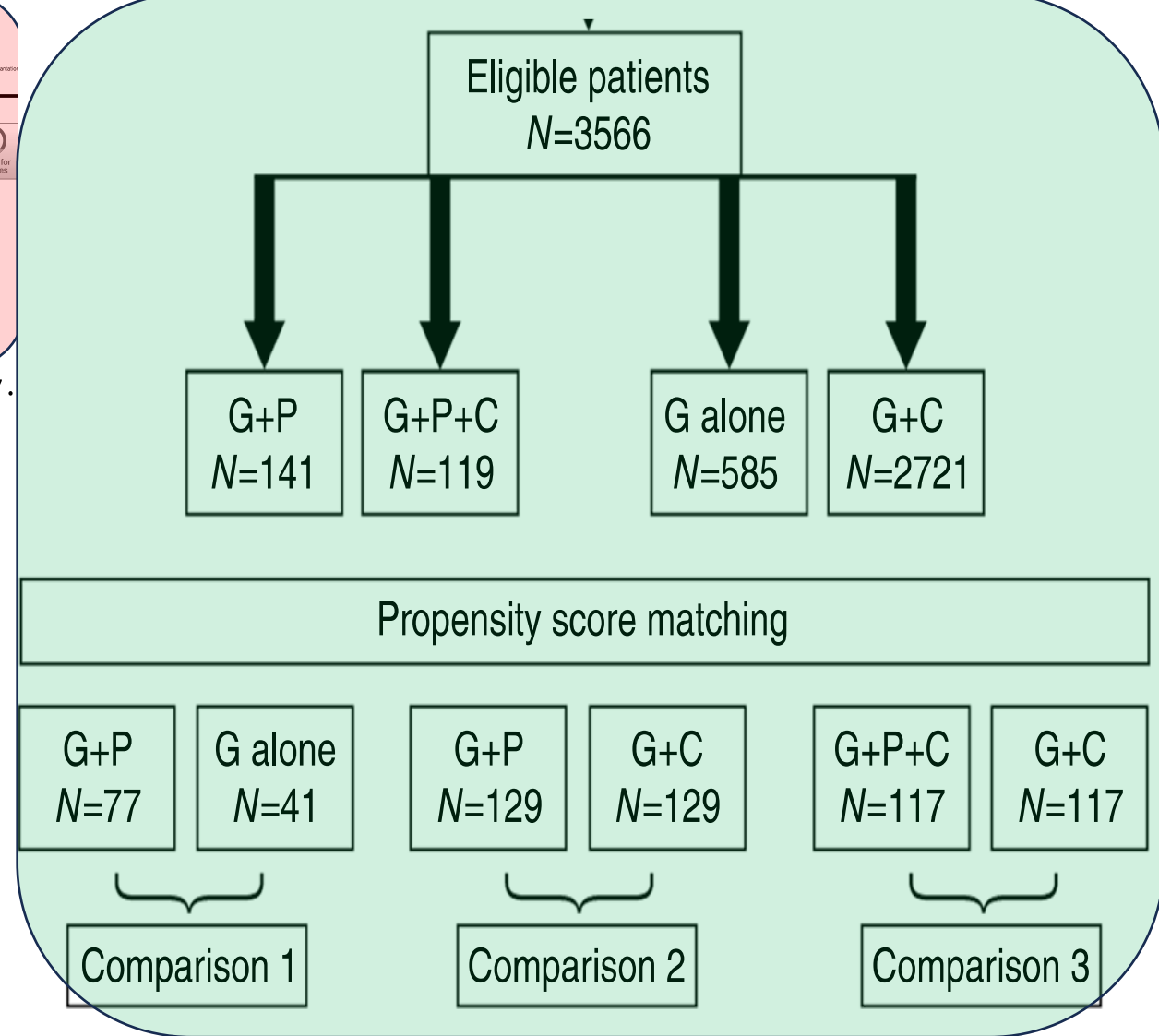


Results from a multicenter, noninterventional registry study for multiple myeloma patients who received stem cell mobilization regimens with and without plerixafor

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	Number at risk	0	0.5	1	2	3	4	5	6	7	8
Plerixafor + G-CSF + Chemo	117	109	97	82	63	36	18	5	0	0	0
G-CSF + Chemo	117	108	106	91	73	45	22	7	1	0	0



Despite propensity scoring, there were more proven poor mobilizers in the plerixafor cohorts, which may have influenced the outcomes

CONCLUSIONS

- Clinical efficacy and safety data of Plerixafor pre-emptive are robust; factors limiting its use are the costs; however the economic analysis must take into account logistical advantages: lower hospitalization rate, fewer complications; better planning and optimization of apheresis sessions.
- Although the general policies regarding ASCT in MM and lymphoma are similar in many centers, there is still great heterogeneity in mobilization strategies....
- The different policies as regard the ASCT and the SC procurement planning not only depend on the different interpretation of some scientific issues regarding ASCT, but also on the availability of economic and logistical resources in different centers.
- G-CSF+Plerixafor «just in time» is recommended over a second mobilization attempt in most currently guidelines!

CHEMO-FREE MOBILIZATION: THE REASONS FORYES!

- **In lymphomas** ASCT indications are declining; CHT+G-CSF mobilization in R/R pts makes hard response evaluation and LK planning: a rescue TX can be carried out without contestual blind mobilization (useless if the patient does not respond completely or in case of disease recurrence);
- G-CSF+Plerixafor **can be easily planned** in selected pts in CR, so avoiding either burdening the logistics either delaying CART. Chemo+G-CSF requires hospitalization and pts should carry out several blood counts to identify the nadir and the rise of WBC/Plt/CD34+.
- **In MM** CTX+G-CSF is certainly effective, improving CD34+ collection in pts heavily exposed to IMiD/Dara; however mobilization is generally performed before Lena (used in maintenance after and not before ASCT);
- CTX+G-CSF can increase the efficacy of mobilization after DARA and could be better in case of tandem ASCT. However, this target is requested less and less frequently.
- CTX+G-CSF does not allow to accurately predict the day of 1st apheresis and it is associated with infections or non-infectious toxicity (so increasing the indirect costs).
- G-CSF+Plerixafor on demand is easy to apply and flexible: different algorithms proposed (from the circulating CD34/WBC ratio to the first LK collection) allow to establish a clear collection calendar.

Thanks....

Simone Angeletti & nurses team

All patients and their families



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