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Treatment patterns among patients with relapsed/refractory (R/R) diffuse large B-cell lymphoma (DLBCL) who are ineligible for transplantation – a real-world study using French PMSI

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INTRODUCTION

- Diffuse Large B-Cell Lymphoma (DLBCL) is the most frequent non-Hodgkin lymphoma, with more than 5,000 new cases diagnosed each year in France. Rituximab associated immunochemotherapies are the most frequent 1st line treatment (1).
- High-dose chemotherapy associated with autologous stem cell transplantation (ASCT) is the standard of care for 2nd line refractory/relapsed management (R/R-DLBCL). However, due to ASCT eligibility criteria, most R/R patients are ASCT-ineligible, with few therapeutic alternatives (2-3).
- French real-world data on R/R-DLBCL are limited, notably regarding its epidemiology and clinical burden.
 - This study aimed to describe R/R-DLBCL management in ASCT-ineligible patients, notably the number of hospital stays, the length of stay and the inhospital mortality rate.

METHODS

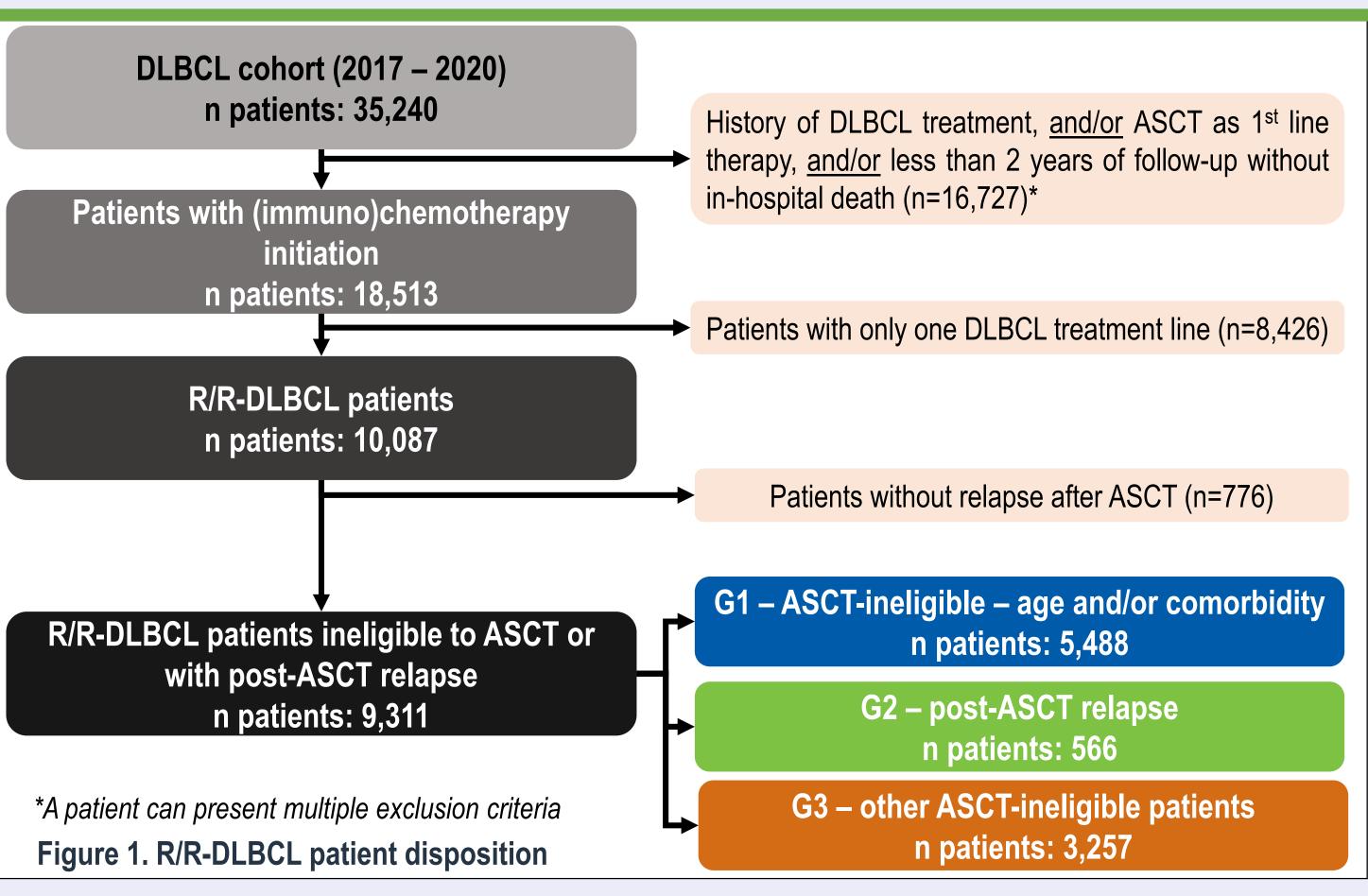
Descriptive, retrospective, longitudinal study using secondary data from French hospital Database (PMSI).

- PMSI database exhaustively includes French hospital-related claims irrespective to healthcare insurance system or hospital settings (public/private). Data from medicine, surgery and obstetric; (MCO); home care (HAD); post-acute care and rehabilitation (SSR) have been used to carry out this study
- DLBCL cohort: patients with ≥1 DLBCL-related stay between 2017 and 2020 (ICD-10 code C83.3). Among them, incident R/R-DLBCL were identified, using validated French national cancer institute (INCa) algorithm and treatment sequences (4).
- Patients with R/R-DLBCL were followed for ≥24 months or until inpatient death.
- Three groups were targeted:
 - (G1) ASCT-ineligible due to age (≥70y) and/or comorbidities,
 - (G2) post-ASCT relapse,
 - (G3) other ASCT-ineligible.

RESULTS

Among the 18,513 newly diagnosed DLBCL patients with sufficient follow-up over 2017-2020, 10,087 (54.5%) relapsed (R/R) before two years

 Most of them (54.4%, n=5,488) were ASCT-ineligible due to old age or comorbidities, while 5.6% had a previous ASCT (Figure 1).



- Recurrence/relapse occurred slightly later among patients from G1 subgroup with a median (Q1–Q3) time to R/R of 6.0 (5.0–14.0) months (Figures 3).
- Length of stays were similar among G1 and G3 subgroups (99.0 (56.0 167.0) and 76.0 (37.0 - 140.0) in median, respectively). Patients from G2 subgroup had slightly more frequent (39.0 (28.0–57.0)) and longer (160.0 (110.0–238.5) days) hospital stays in median, due to ASCT management (Figures 4 and 5).



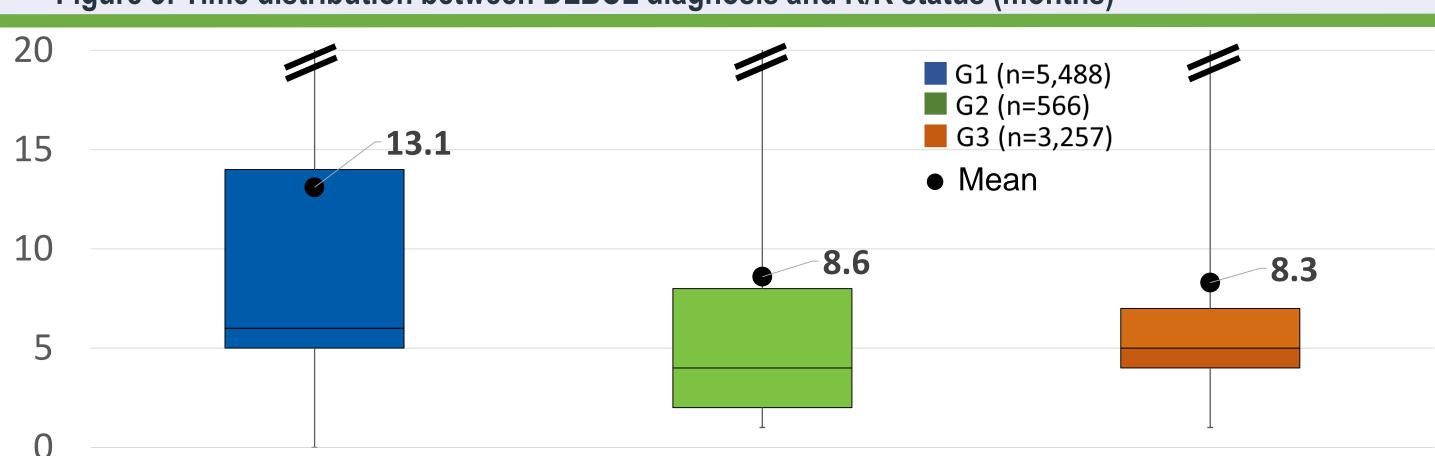


Figure 4: Number of stays, overall stays (dark) and chemotherapy stays (light) over follow-up

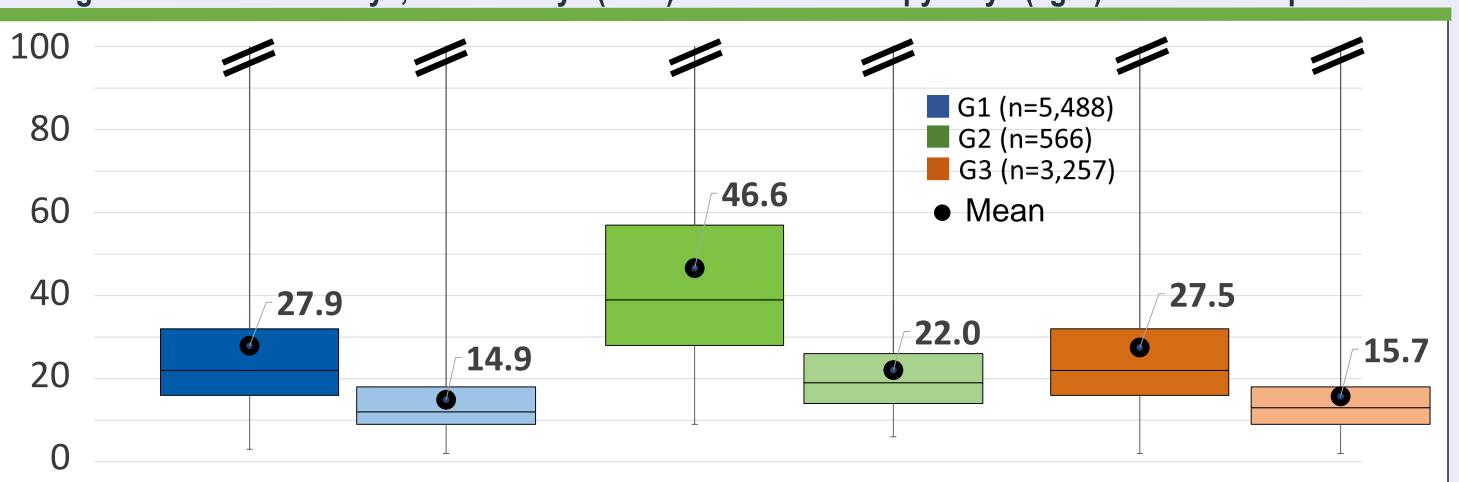
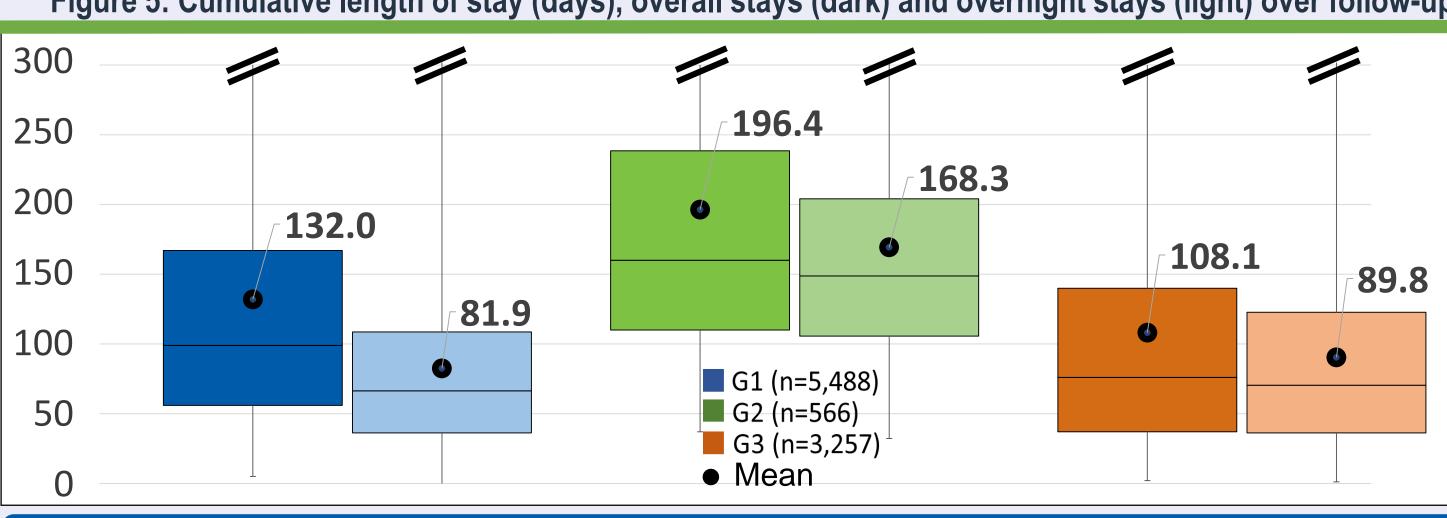


Figure 5: Cumulative length of stay (days), overall stays (dark) and overnight stays (light) over follow-up



Median (Q1–Q3) age at R/R status in G1 was 76.0 (71.0–81.0), higher than other subgroups which had similar median ages around 59 years.

- Sex ratio was around 1.4, with 40% of women (Table 1)
- In G1 subgroup, heart, liver and kidney failures were encountered among around 13% of patients, while being close to 0% in other subgroups (Figure 2 – Table 1).

Figure 2: Age distribution at R/R-DLBCL status (years)

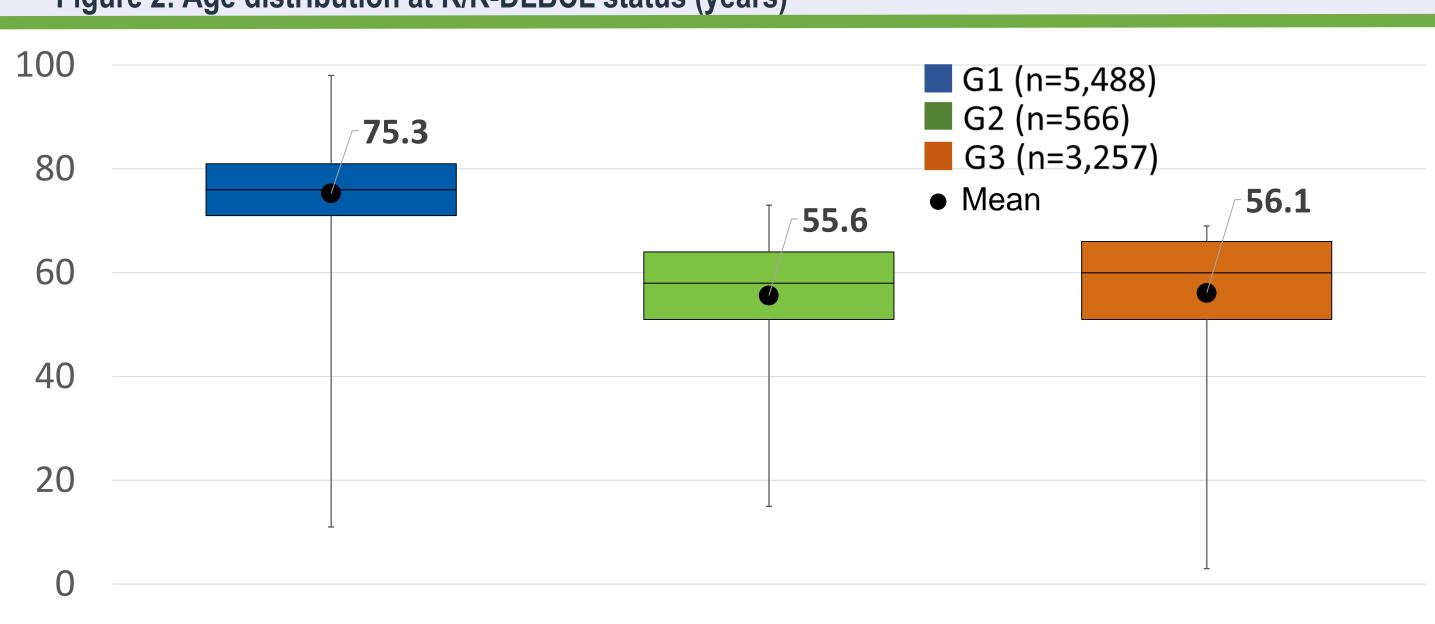


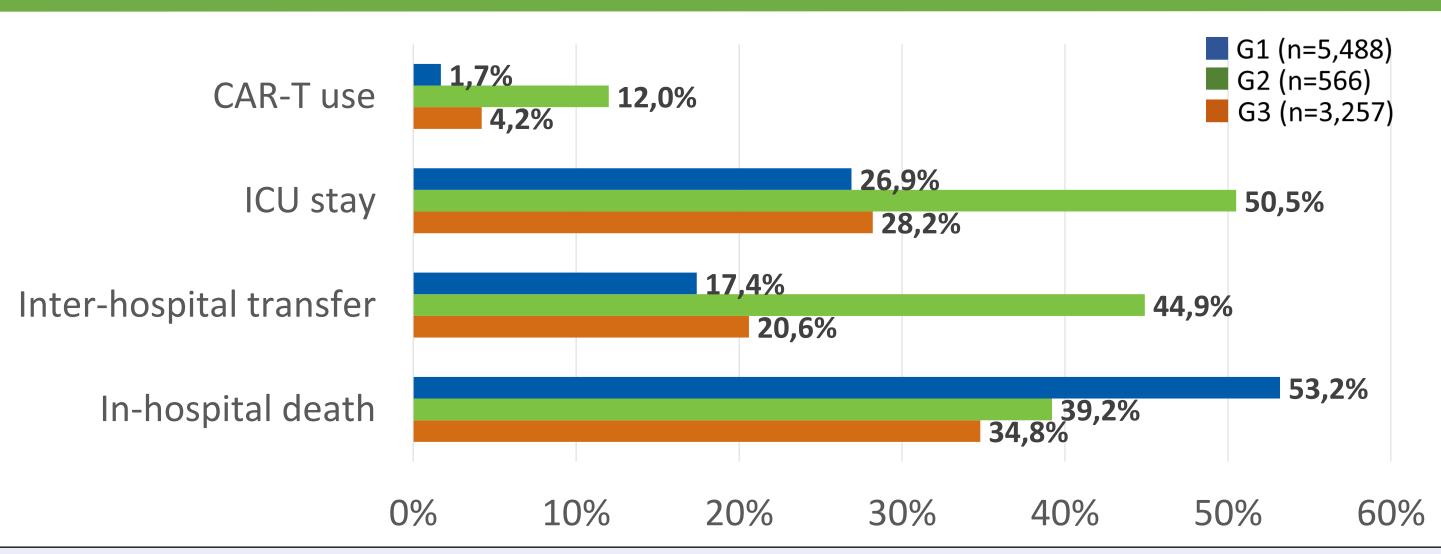
Table 1: Main characteristics of R/R-DLBCL patients

| | G1 (n=5,488) | G2 (n=566) | G3 (n=3,257) |
|---------------------------------|--------------|-------------|--------------|
| Follow-up (months) | 31.0 | 48.0 | 60.0 |
| Median (Q1–Q3) | (17.0–48.0) | (31.0–74.0) | (51.0–66.0) |
| Sex-ratio | 1.3 | 1.5 | 1.5 |
| (% women) | (43.3%) | (39.6%) | (40.5%) |
| Age (years) | 76.0 | 58.0 | 60.0 |
| Median (Q1–Q3) | (71.0–81.0) | (51.0–64.0) | (51.0–66.0) |
| <40 (%) | 1.1% | 7.2% | 12.3% |
| [40-60[(%) | 4.9% | 49.6% | 35.5% |
| ≥60 (%) | 94.0% | 43.2% | 52.2% |
| Medical history of interest (%) | 34.2% | 9.2% | 0.0% |
| Solid transplant (%) | 2.8% | 0.5% | 0.0% |
| Heart failure (%) | 14.9% | 2.3% | 0.0% |
| Hepatic failure (%) | 12.5% | 5.3% | 0.0% |
| Renal failure (%) | 12.7% | 1.8% | 0.0% |

Patients from G2 subgroup seemed more likely to require ICU than patients from other subgroups, due to ASCT itself (Figure 6).

- Similarly, almost 50% of G2 patients had to be transferred to other hospitals, probably for transplantation.
- Patients from G1 subgroup were more likely to die in hospital than other subgroups (Figure 6).

Figure 6: Proportion of patients with ≥1 events of interest over follow-up



Disclosure

Di Blasi: Speaker - Gilead, Novartis, Janssen, Pfizer - Travel and accommodation: Gilead Thieblemont: Speaker - Novartis, BMS/Celgene, Bayer, AbbVie, Gilead Sciences, Roche, Janssen, Kite, Incyte, Amgen, Takeda – Travel and accommodation: Novartis, BMS/Celgene, Bayer, AbbVie, Gilead Sciences, Roche, Janssen, Kite, Amgen, Takeda – Research grant: Janssen, Roche – Consultancy/honoraria: Novartis, BMS/Celgene, Bayer, AbbVie, Gilead Sciences, Roche, Janssen, Kite, Incyte, Amgen, Takeda Haioun: Honoraria - Miltenyi, Roche, Celgene, Incyte, Pfizer, Gilead, Janssen - Travel and accommodation: Roche, Janssen Mayaud: Employee and stock ownership - Incyte Corporation Sackmann Sala: Employee and stock ownership - Incyte Corporation, Diez-Andreu: Employee - stève consultants (commissioned by Incyte Corporation to conduct these analyses), Bugnard: Employee - stève consultants (commissioned by Incyte Corporation to conduct these analyses), Goguillot - Employee - stève consultants (commissioned by Incyte Corporation to conduct these analyses Chillotti: Employee - stève consultants (commissioned by Incyte Corporation to conduct these analyses),), Finzi: Employee and stock ownership - Incyte Corporation, Bénard: Employee and stock ownership - stève consultants (commissioned by Incyte Corporation to conduct these analyses).

CONCLUSION

- This study highlights the extensive hospital management of ASCT-ineligible patients with R/R-DLBCL before 2021.
- R/R-DLBCL patients spend an important time (>100 days) in hospital, notably older and frailer ones, for whom limited therapeutically alternatives exist.
- These hospital patterns reflect the clinical practice before 2021 and could have evolved, notably with the ongoing generalization of CAR-T cell therapy for eligible patients as second line treatment.

Acknowledgement

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