



ASCO 2018

Highlights

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FOLLICULAR LYMPHOMA

- **PHASE III RELEVANCE study (abstract 7500) – Nathan Fowler**

- R2 (rituximab + lenalidomide) vs R-chemo followed by rituximab maintenance in patients with ND FL
- 1,030 patients included
- Efficacy results were similar (ORR, CRu, DoR)
- No superiority of R2 vs R-chemo in terms of PFS at 120 weeks
- More Grade 3 / 4 neutropenia in the R-chemo arm (50%) compared with R2 (32%)
- Longer follow-up needed for mature PFS and OS data

Summary – Expert's interview

- **Current advances in FL – Sonali Smith**

- Results of the RELEVANCE trial: longer follow-up needed
- New PI3K inhibitor approved in the US (copanlisib)

Expert's interview

- **Polatuzumab vedotin in combination with BR (abstract 7507) – Laurie Sehn**

- Randomized controlled trial against BR – 80 R/R FL patients
- Increase of low grade AEs (peripheral neuropathy) and grade 3-5 AEs (neutropenia, anemia and febrile neutropenia [but no infection])
- No difference in CRR, mPFS and mOS

Expert's interview

- **Phase 1b/2 of Hu5F9-G4 (abstract 7504) – Ranjana Advani**

- Hu5F9-G4: first-in-class anti-CD47 antibody (5F9)
- 7 patients with R/R FL
- MTD not reached
- In combination with rituximab: CRR = 43%, mDoR not reached

Summary

DLBCL

- **TRANSCEND NHL 001 study (abstract 7505) – Jeremy Abramson**

- JCAR017 (lisocabtagene maraleucel)
- 102 patients included with R/R aggressive NHL
- DLBCL cohort: ORR = 80%, CRR = 59%
- 93% patients in remission at 6 months remained in remission at the latest follow-up
- Most important Grade 3 / 4 events = neutropenia and fatigue, CRS: 37%, NE: 23%

Summary

- **Durable responses with axi-cel in R/R DLBCL in the ZUMA-1 trial (abstract 3003) – Frederick Locke**

- 108 patients treated with axi-cel (axicabtagene ciloleucel), follow-up: 15.4 months
- ORR = 82%, CRR = 58%, mDoR = 11.1 months
- Responders at 3 months have 80% chance to maintain their response at 1 year. Some PR converted into CR at 1 year

Summary - Expert's interview

- **Polatuzumab vedotin in combination with BR (abstract 7507) – Laurie Sehn**

- Randomized controlled trial against BR – 80 R/R DLBCL patients
- Increase of low grade AEs (peripheral neuropathy) and grade 3-5 AEs (neutropenia, anemia and febrile neutropenia [but no infection])
- CRR: 40% (vs 15%), mPFS: >6 months (vs 2 months), mOS: >11 months (vs <5 months)

Expert's interview

- **Phase 1b/2 of Hu5F9-G4 (abstract 7504) – Ranjana Advani**

- Hu5F9-G4: first-in-class anti-CD47 antibody (5F9)
- 15 patients with R/R FL
- MTD not reached
- In combination with rituximab: CRR = 33%, mDoR not reached

Summary

- **Final analysis of the AHL2011 LYSA trial (abstract 7503) – Olivier Casasnovas**

- 823 patients included with untreated HL – early PET-driven treatment de-escalation after 2 cycles of BEACOPPesc vs not PET-monitored strategy
 - mFU: 50.4 months – 4-y PFS: 87.1% vs 87.4%; 5-y PFS: 85.7% vs 86.2%; 4-y OS: 97.1% vs 96.9%; 5-y OS: 96.4% vs 95.2%

Summary – Expert’s interview – G. Salles’s interview

- **Novel therapies for R/R HL beyond transplant – Catherine Diefenbach**

- Not all relapsed patients are the same
 - 1st relapse and ASCT = SoC but for patients in CR2
 - BV: 35% CRR in 2nd line salvage prior to SCT, effective in consolidation post AST (mPFS: 42.9% vs 24.1%, but no difference in mOS)
 - BV + B: CRR: 76%, 18-months PFS: 75%
 - Nivolumab: CRR between 9% and 17%
 - BV + ipi: CRR = 48%, 1-y PFS = 58%
 - BV + nivo: CRR = 67%, 6-months PFS = 93%
 - Other new agents tested (mTORi, HDACi)
 - Allo-SCT: 3-y OS = 50% but 20% mortality

Expert’s interview

- **Risk-adapted treatment strategies for advanced-stage HL – Joseph Connors**

- Brentuximab vedotin improves effectiveness of CT (improvement of curability?)
- New agents tested in poorest prognostic patients

Expert’s interview

CLL

- **Ibrutinib and venetoclax in treatment-naïve CLL (abstract 7502) – William Wierda**

- Undetectable MRD in 77% patients after 6 cycles and 79% after 12 cycles
- No new safety signals

Summary – Expert's interview

- **High and durable MRD- with venetoclax and rituximab in R/R CLL (abstract 7508) – Peter Hillmen**

- MRD-: VenR = 84% vs BR = 23%
- 83% of patients maintained MRD-, regardless of risk features

Expert's interview

- **The potential of venetoclax for patients with 17p deletion – Michael Hallek**

- Studied in high-risk patients (17p deleted. P53 mutated or complex karyotype)
- Venetoclax in combination with rituximab (Murano trial) or ibrutinib (Captivate trial) or obinutuzumab induce long-lasting remissions (several years in the Murano trial) and MRD-

Expert's interview

- **Role of chemo-immunotherapy in CLL – Susan O'Brien**

- Relapse setting;
 - no role for chemotherapy
 - Retreatment with FCR or BR
 - Ibrutinib for 17p and 11q deleted patients
 - VenR
- Front-line setting: SoC = FCR
 - Patients with IgVH mutation: FCR
 - Fit unmutated and elderly patients : ibrutinib

Expert's interview

MISCELLANEOUS

FL

- **Retrospective analysis on biopsy-proven FL transformation in the rituximab era – Marek Trněný**
 - 1200 patients with FL, 546 relapses, 370 tFL; risk of transformation is ≈10%
 - Rituximab might reduce the risk of transformation

Expert's interview

- **Definition and management of high-risk FL patients – Stefano Luminari**
 - The most important prognostic factor is the duration of response
 - Early relapses have to be considered as aggressive disease (intensified treatment with ASCT)

Expert's interview

DLBCL

- **Real-time CoO subtype identification by genetic profiling (abstract 7548) – Greg Nowakowski**
 - Feasible on a global scale (mean time: 2.6 days) – ROBUST phase III trial
 - Incidence of ABC (355-38% in the US, 40% in Europe and 60% in Asia)

Expert's interview

- **Tailored therapy for aged patients with aggressive lymphoma – Raoul Cordoba**
 - 3 different prognostic factors identified: age, IPI and co-morbidities to define 3 risk-groups: robust (to be treated as younger patients), vulnerable or frail

Expert's interview

DLBCL, FL

- **Overview of immunotherapy – Carol Jacobson**
 - TRANSCEND study: CRR = 50%
 - Phase I monoclonal antibody anti CD47 + rituximab in FL, DLBCL
 - ORR = 50%, CR in 1/3 of patients. Very good safety profile

Expert's interview

MISCELLANEOUS CONT'D

CTCL

- **Novel therapies for CTCL – Francine Foss**

- BV (ALCANZA trial) efficient even in patients with very low expression of CD30
- Mogamulizumab (anti CCR4), especially efficient in SZ
- Other investigational agents (targeting CD25 or mRNA155)

Expert's interview

WM

- **Acalabrutinib in monotherapy (abstract 7501) – Roger Owen**

- Efficient in treatment-naïve patients: 93% ORR, 24-months duration of response rate = 90%, 90% 24-m PFS, 24-m OS = 92%
- Activity in R/R patients: 93% ORR, 24-months duration of response rate = 84%, 82% 24-m PFS, 24-m OS = 89%
- Most common grade ≥3 AEs: neutropenia, pneumonia and low respiratory tract infection

Summary

CLL, FL and WM

- **Potential for CT-free future – Bruce Cheson**

- FL: R2 regimen; WM: acalabrutinib; CLL: doublets with rituximab idelalisib, ibrutinib or venetoclax

Expert's interview



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