

Eppur si muove...

La terapia nel MONDO LINFOMI

Caso clinico 2

Tetiana Skrypets



BARI, 28 GIUGNO 2022

Anamnesi

- Paziente A, 83 anni
- No B-sintomi, solo astenia
- Non fumatore
- Anamnesi patologica remota:
 - Ipertrofia prostatica benigna
 - Ipertensione arteriosa
 - Sofferenza cerebrale ipossica
 - Versamento pericardico organizzato

Diagnosi

- **TAC (16/06/2021)** l/nodi sede sovraclaveare 20x25 cm, versamento pleurico bilaterale, l/nodi in sede ilare dx 22x38 mm, ascellare dx 18x36 mm, l/nodi nelle catene addomino-pelviche 27x57 mm. Milza 15 cm
- **Esami ematici** – HGB 8.06 g/dL
- **Ferro 28 microgr/dL, Ferritina 668 ng/mL, Transferrina 180 mg/dL**
- **Anti-HBsAg** – 68.2 UI/L, **anti HBc (core)** – positive

Biopsia (02/07/2021) – linfonodo inguine destro

DLBCL, non GCB (Hans)

CD20+, BCL6+, BCL2+, IRF4/MUM1 (80%) +, CD30+

CD23+/-, CD5-/CD3- (+ in linfociti T)

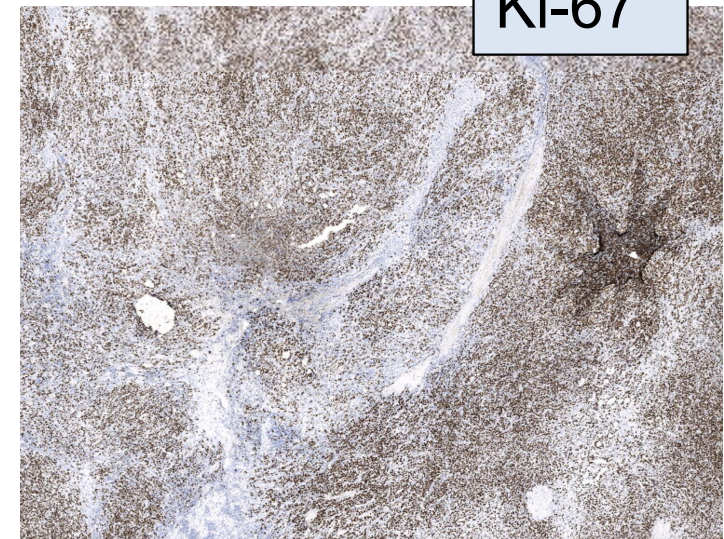
CD10-, EBV/LMP-

C-MYC (30%)

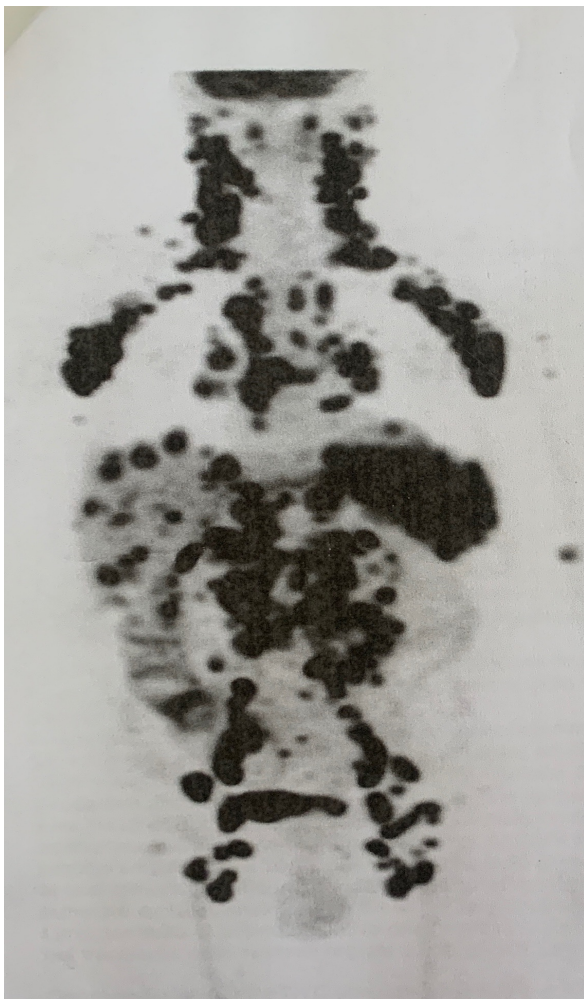
Ki67/Mib1: 85%

EBV (EBER) - negative

*FISH – no riarrangiamento del MYC e BCL2, **ma BCL6 +***



Stadiazione



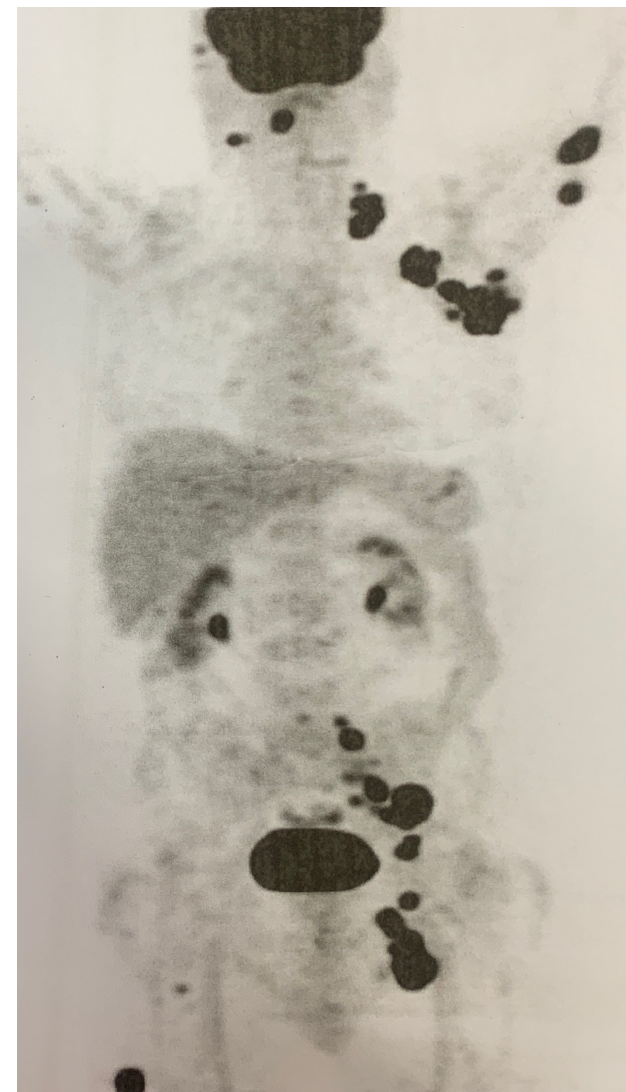
- No infiltrazione osteomidollare
- PET/CT scan (07/2021)
 - Tutte le principali stazioni linfonadali sopra e sottodiaframmatiche (**SUVmax 18.2**)
 - In ambito epatico multiple aree (**SUVmax 18.7**)
 - Milza (**SUVmax 17.7**)

*Linfoma non Hodgkin diffuso a grandi cellule B
(non-GCB), Stadio **IV**,
IPI 4 (età, stadio, ECOG, LDH)*

Terapia – 1 linea

- **6xR-miniCOMP + 2R**
- **PET EOT (07/02/2022)**
 - *laterocervicale dx, sovraclaveare sn, interpettorale ed ascellare sn (SUVmax 34.4)*
 - *Iliaca comune (SUVmax 21.4) ed inguinale omolaterale (SUVmax 26.4)*
 - *muscoli del braccio sinistro (SUVmax 20.7), muscolo gluteo sn (SUVmax 27.5), quadricipite femorale (SUVmax 26.4),*
 - *celiac lymph nodes 13 mm, mesenteric l/n 27x19 mm, right inguinal l/n 31, 38, 27, 23 mm, left inguinal – 13 mm, 17x10 mm with SUVmax 6.2, 13.7, 14.6, 16.3*

DS 5



Terapia – 2 linea

➤ 2xR-Benda dal 28/02/22 al 29/03/22 (Polatuzumab vedotin non disponibile)



National
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NCCN Guidelines Version 4.2022 Diffuse Large B-Cell Lymphoma

SECOND-LINE THERAPY^{d,i,j} (non-candidates for transplant)

Preferred regimens (in alphabetical order)

- GemOx ± rituximab
- Polatuzumab vedotin-piiq ± bendamustine ± rituximab^{k,l}
- Tafasitamab-cxix^m + lenalidomide

Other recommended regimens (in alphabetical order)

- CEOP (cyclophosphamide, etoposide, vincristine, prednisone) ± rituximab
- DA-EPOCH ± rituximab
- GDP ± rituximab or (gemcitabine, dexamethasone, carboplatin) ± rituximab
- Gemcitabine, vinorelbine ± rituximab (category 3)
- Rituximab



GOOD SCIENCE
BETTER MEDICINE
BEST PRACTICE

ESMO > Guidelines > Guidelines by topic > Haematological Malignancies

DIFFUSE LARGE B-CELL LYMPHOMA: ESMO CLINICAL PRACTICE GUIDELINES

Haematological Malignancies

Published in 2015 – Ann Oncol (2015) 26 (suppl 5): v116-v125.

Authors: H. Tilly, M. Gomes da Silva, U. Vitolo, A. Jack, M. Meignan, A. Lopez-Guillermo, J. Walewski, M. André, P. W. Johnson, M. Pfreundschuh, M. Ladetto

PROGRESSIONE

Terapia – 3 linea

➤ 2xPixantrone (1-8 gg, 15 – severa neutropenia)

ARTICLES | VOLUME 13, ISSUE 7, P696-706, JULY 01, 2012

Pixantrone dimaleate versus other chemotherapeutic agents as a single-agent salvage treatment in patients with relapsed or refractory aggressive non-Hodgkin lymphoma: a phase 3, multicentre, open-label, randomised trial

Dr Ruth Pettengell, MD, Prof Bertrand Coiffier, MD, Prof Geetha Narayanan, MD

Fernando Hurtado de Mendoza, MD, Raghunadharao Digumarti, MD, Henry Gomez, MD, et al. Show all authors

> [Acta Haematol.](#) 2021;144(3):259-263. doi: 10.1159/000509923. Epub 2020 Oct 9.

Effectiveness and Safety of Pixantrone for the Treatment of Relapsed or Refractory Diffuse Large B-Cell Lymphoma in Every-Day Clinical Practice: The Italian Cohort of the PIXA Registry

Pier Luigi Zinzani¹, Marco Bregni², Mario Spione³, Manfred Mitterer⁴, Gerardo Musuraca⁵, Annamaria Bugli⁶, Francesco Piazza⁷, Antonello Pinto⁸

- minina cardiotoxicità
- opzioni limitate
- è attualmente approvato in Italia in monoterapia in pazienti adulti con linfomi non Hodgkin B aggressivi recidivati o refrattari ad almeno due linee di terapia

PROGRESSIONE

Terapia – 4 linea...discussione multidisciplinare LymphoTEAM

Paziente anziano

Non eligibile CAR-T

Refrattario



***Bispecifico?
Terapia ad alto
rischio di CRS***

Comorbidità

ARTICLES | VOLUME 21, ISSUE 7, P978-988, JULY 01, 2020

Tafasitamab plus lenalidomide in relapsed or refractory diffuse large B-cell lymphoma (L-MIND): a multicentre, prospective, single-arm, phase 2 study

Prof Gilles Salles, MD   • Johannes Duell, MD  • Eva González Barca, MD • Prof Olivier Tournilhac, MD • Prof Wojciech Jurczak, MD • Prof Anna Marina Liberati, MD • et al. [Show all authors](#) • [Show footnotes](#)

Published: June 05, 2020 • DOI: [https://doi.org/10.1016/S1470-2045\(20\)30225-4](https://doi.org/10.1016/S1470-2045(20)30225-4) •  Check for updates

THE LANCET
Oncology

CLINICAL TRIALS: IMMUNOTHERAPY | NOVEMBER 15 2021

RE-MIND: Comparing Tafasitamab + Lenalidomide (L-MIND) with a Real-world Lenalidomide Monotherapy Cohort in Relapsed or Refractory Diffuse Large B-cell Lymphoma

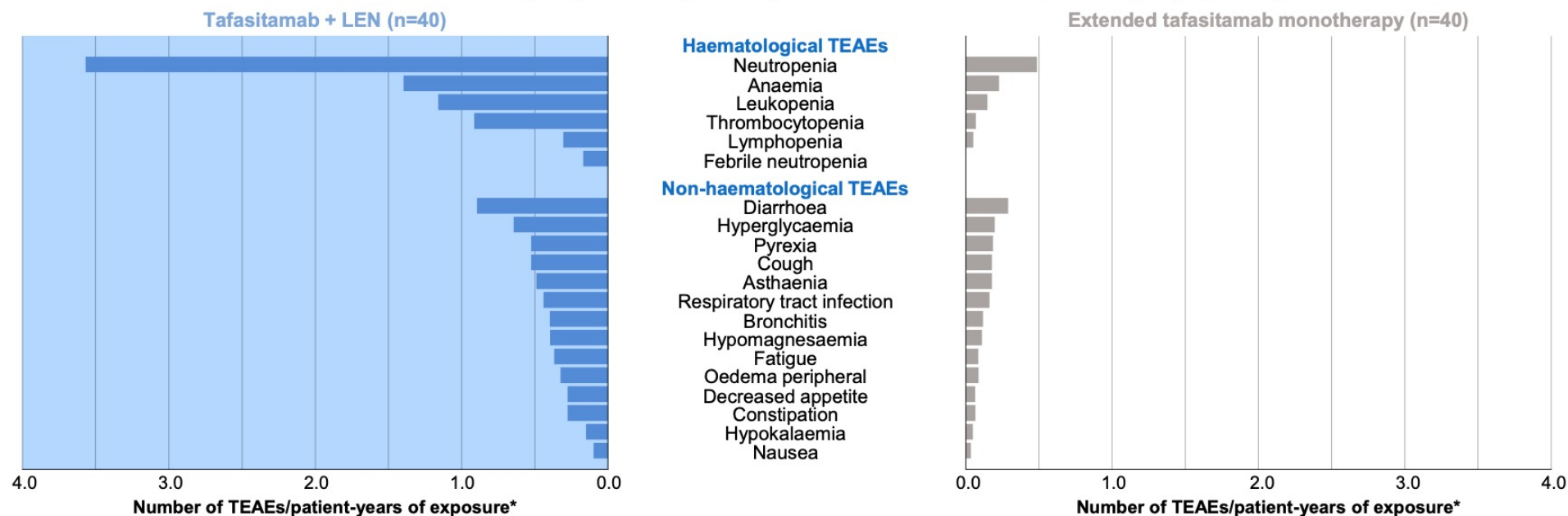
Pier Luigi Zinzani  ; Thomas Rodgers; Dario Marino  ; Maurizio Frezzato; Anna Maria Barbui; Claudia Castellino; Erika Meli; Nathan H. Fowler; Gilles Salles  ; Bruce Feinberg  ; Nuwan C. Kurukulasuriya; Sascha Tillmanns; Stephan Parche; Debarshi Dey; Günter Fingerle-Rowson; Sumeet Ambarkhane; Mark Winderlich; Grzegorz S. Nowakowski 



3-YEAR SAFETY BY L-MIND TREATMENT PHASE

(30 OCT 2020 CUT-OFF)

Numbers of AEs per patient-year by L-MIND treatment phase (any grade)



- Consistent with the 2-year analysis, incidence of TEAEs was lower during the tafasitamab monotherapy phase compared with the combination therapy phase

Conclusioni

- Buon profilo di tossicità dell'associazione tafasitamab+lenalidomide
- Possibilità di richiedere la combinazione per uso nominativo
- Non comorbidità rilevanti
- Dosaggio di lenalidomide sarà modulato in base alla clearance della creatinine
- Terapia di supporto: profilassi della TVP con eparina a basso peso molecolare, profilassi primaria con G-CSF