

15° CONGRESSO NAZIONALE



## Italian Conference on AIDS and Antiviral Research

Presidenza del Congresso:

F. Ceccherini Silberstein, M. Formisano, S. Lo Caputo, A. Saracino

BARI 14-16 GIUGNO 2023



Promosso da  
**SIMIT**  
Società Italiana di Malattie Infettive e Tropicali

From prevention to cure: ready for new challenges

# LOWER AIDS-RELATED HOSPITALIZATIONS IN WOMEN LIVING WITH HIV MULTIDRUG RESISTANCE: RESULTS FROM THE PRESTIGIO REGISTRY

R. Papaioannu Borjesson<sup>1</sup>, L. Galli<sup>2</sup>, R. Lolatto<sup>2</sup>, B Menzaghi<sup>3</sup>, M. Feasi<sup>4</sup>, R. Gulminetti<sup>5</sup>, C. Fornabaio<sup>6</sup>, A.M. Cattelan<sup>7</sup>, S. Bonora<sup>8</sup>, F. Lagi<sup>9</sup>, M. Zazzi<sup>10</sup>, A. Castagna<sup>1,2</sup>, on behalf of PRESTIGIO Study Group

1. San Raffaele Vita-Salute University, Milan, Italy 2. Infectious Diseases Department, San Raffaele Scientific Institute, Milan, Italy 3. Unit of Infectious Diseases, ASST della Valle Olona, Busto Arsizio, Italy 4. Ente Ospedaliero Ospedali Galliera, Genova, Italy 5. Department of Medical Sciences and Infectious Diseases, Fondazione IRCCS Policlinico San Matteo, Pavia, Italy 6. ASST Cremona, Cremona, Italy 7. Azienda Ospedale-Università Padova, Padova, Italy 8. Unit of Infectious Diseases, Department of Medical Sciences, University of Turin, Torino, Italy 9. Infectious and Tropical diseases Unit, Azienda Ospedaliera Universitaria Careggi, Florence, Italy  
10. Department of Medical Biotechnologies, University of Siena, Siena, Italy



# Disclosure Statement

---

Dr. Rebecka Papaioannu Borjesson has no financial relationships with commercial entities to disclose



# Background

---

- Today, people living with HIV (PLWH) who have a complete control of viral replication and a good immunologic profile thanks to antiretroviral therapy (ART) have the same life expectancy as the general population [1].
- PLWH are still at risk of morbidities and hospitalizations [2].
- Non-AIDS-defining events are becoming a much more common cause of hospitalization than AIDS-defining events [3, 4].

1. PA Volberding, SG Deeks, *Antiretroviral therapy and management of HIV infection*, Lancet, 2010

2. R Lazar, L Kersanske, Q Xia, D Daskalakis, SL Braunstein, *Hospitalization Rates Among People With HIV/AIDS in New York City*, 2013, Clin Infect Dis, 2017

3. P Mendes Luz, M Bruyand, S Ribeiro, F Bonnet, RI Moreira, M Hessamfar, D Perreira Campos, C Greib, C Cazanave, VG Veloso, F Dabis, B Grinsztejn, G Chêne; IPEC/FIOCRUZ Cohort and the Aquitaine ANRS CO3 Study Group, *AIDS and non-AIDS severe morbidity associated with hospitalizations among HIV-infected patients in two regions with universal access to care and antiretroviral therapy, France and Brazil, 2000-2008: hospital-based cohort studies*, BMC Infect Dis, 2014

4. SA Berry, JA Fleishman, RD Moore, KA Gebo; HIV Research Network, *Trends in reasons for hospitalization in a multisite United States cohort of persons living with HIV, 2001-2008*, J Acquir Immune Defic Syndr, 2014



# Objectives

---

- To analyze the **incidence of hospitalization in male and female people living with 4-drug class resistant HIV (4DR-PLWH)** enrolled in the PRESTIGIO Registry.
- To analyze the **causes of hospitalization.**



# Material and Methods

---

- **178 4DR-PLWH** from the PRESTIGIO Registry
- **PRESTIGIO Registry:** Italian registry which collects clinical and laboratory data on individuals with documented genotypic resistance to:
  - *nucleoside reverse transcriptase inhibitors (NRTIs)*
  - *non-NRTIs (NNRTIs)*
  - *protease inhibitors (PIs)*
  - *either genotypic resistance to integrase strand transfer inhibitors (INSTIs) or an history of virological failure [defined as two consecutive measurements of viral load (VL)  $\geq 50$  copies/mL] to an INSTI-based regimen.*
- Follow-up: from the date of the first 4DR evidence (baseline) until death/loss-to-follow-up/freezing date (December 31st, 2022).
- Hospitalization: hospital admission for any reason with  $\geq 1$  overnight stay.



# Statistical analysis

---

- Data were reported as median (IQR) or frequency (%).
- Mann-Whitney test used to compare baseline characteristics.
- Poisson regression model was used to model incidence rates (IR) with 95% confidence intervals (CI).



# Baseline characteristics

Characteristic	Overall (n=178)	Males (n=132)	Females (n=46)	P-value
Age, years	48.86 (43.87 - 53.64)	49.8 (45.62 - 54.07)	46.41 (33.18 - 52.5)	0.004
Years since HIV diagnosis	21.17 (16.79 - 24.87)	21.1 (16.43 - 25.43)	21.17 (17.4 - 24.11)	0.735
Years since ART start	17.49 (13.85 - 20.31)	17.65 (14.11 - 20.7)	16.74 (10.83 - 18.96)	0.114
Calendar year of 4DCR evidence	2014 (2011 - 2016)	2014 (2011 - 2016)	2014 (2011 - 2015)	0.626
Total number of primary resistance mutations	15 (12 - 19)	16 (12 - 19)	14 (11 - 18)	0.270
Nadir CD4+, cells/mm <sup>3</sup>	82 (21 - 199)	73 (16 - 189)	96 (42 - 203)	0.317
BL HIV-RNA, log <sub>10</sub> copies/mL	3.45 (2.31 - 4.39)	3.45 (2.25 - 4.37)	3.45 (2.44 - 4.4)	0.936
BL CD4+, cells/mm <sup>3</sup>	395 (200 - 596)	404 (206 - 630)	319 (181 - 540)	0.135
BL CD4+/CD8+ ratio	0.37 (0.2 - 0.61)	0.35 (0.2 - 0.59)	0.41 (0.24 - 0.73)	0.408



# Incidence of hospitalization

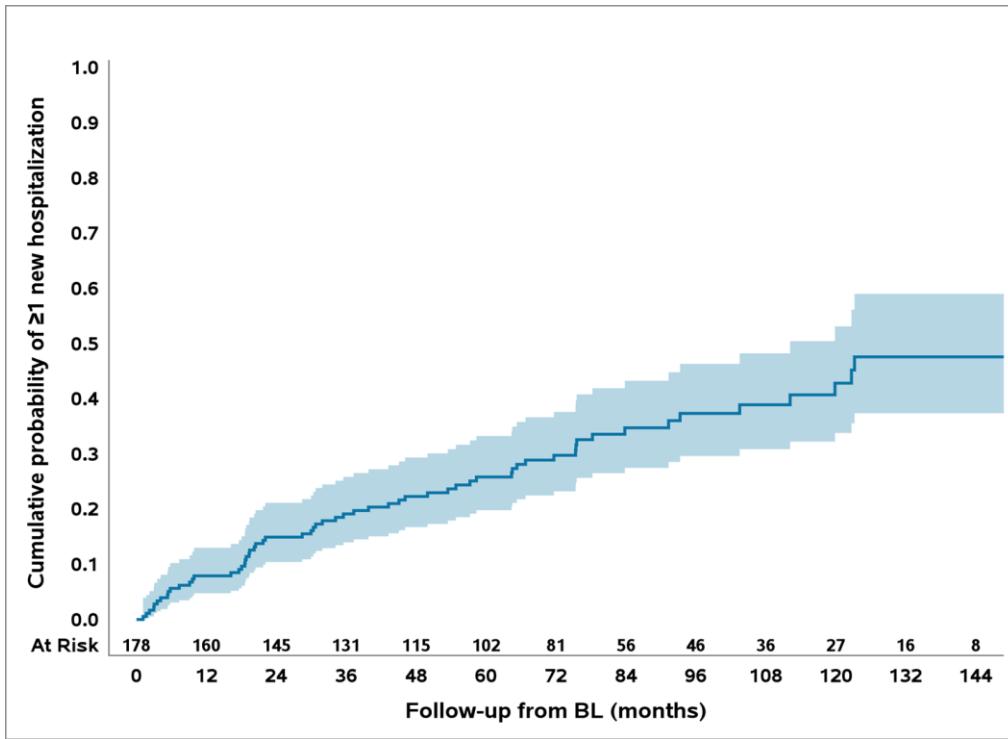
The overall incidence rate of hospitalization was 9.43/100-PYFU (95%CI=7.76-11.10)

MALES (933 PYFU)			FEMALES (361 PYFU)			IRR F/M (95%CI)	P-value (M vs F)
Number of patients	Number of events	Incidence rate (95%CI) per 100-PYFU	Number of patients	Number of events	Incidence rate (95%CI) per 100-PYFU		
43	94	10.08 (8.23-12.34)	17	28	7.75 (5.35-11.23)	0.77 (0.51-1.17)	0.223

The median duration of hospitalization was 8 days (IQR 4-20)



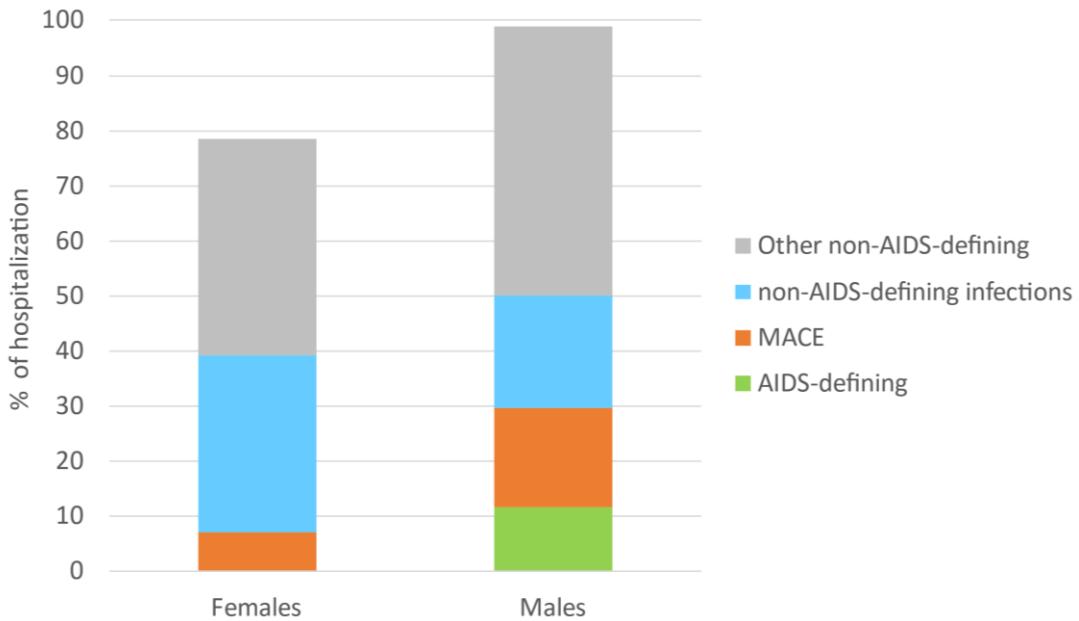
# Cumulative probabilities of new hospitalizations



- 31/178 (17.4%) had 1 hospitalization.
- 13/178 (7.3%) had 2 hospitalizations.
- 10/178 (5.6%) had 3 hospitalizations.
- 3/178 (1.7%) had 4 hospitalizations.
- 3/178 (1.7%) had  $\geq 5$  hospitalizations.
- After 7 years from baseline, 34.7% of people were estimated to have had one or more hospitalizations.



# Most frequent causes of hospitalization



- n=28, 23%
- n=19, 15.6%
- females had no hospitalizations for AIDS-defining events compared to males (0% vs 11.7% (n=11), p=0.049)
- gender differences for non-AIDS-defining events
  - MACEs: 17 (18.1%) in M and 2 (7.1%) in F, p=0.097
  - non-AIDS-defining infections: 19 (20.2%) in M and 9 (32.1%) F, p=0.084



# Immuno-virologic profile at hospitalization

Variables	Males (n=132)	Females (n=46)	p-value
HIVRNA, copies/mL	108 (22-17100)	36 (0.9-541)	0.049
CD4, cells/mm <sup>3</sup>	232 (70-621)	580 (427-729)	0.030
CD4/CD8 ratio	0.35 (0.2 - 0.59)	0.41 (0.24 - 0.73)	0.4080

Immuno-virologic profile at hospitalization was better in females than in males



# Conclusions

---

- Women living with HIV multidrug resistance seem to have a lower incidence of AIDS-defining events hospitalizations
- Women tended to have fewer hospitalizations for MACEs and more hospitalizations for non-AIDS-defining infections than males.





# Acknowledgements

**STEERING COMMITTEE:** Antonella Castagna (Coordinator), Vincenzo Spagnuolo, Laura Galli, Franco Maggiolo, Leonardo Calza, Emanuele Focà, Filippo Lagi, Giovanni Cenderello, Antonio Di Biagio, Giulia Marchetti, Stefano Rusconi, Adriana Cervo, Roberta Gagliardini, Stefano Bonora, Anna Maria Cattelan, Maurizio Zazzi, Maria Mercedes Santoro

**VIROLOGY TEAM AND BIOLOGICAL BANK:** Maurizio Zazzi, Maria Mercedes Santoro, Andrea Galli, Francesco Saladini, Daniele Armenia

**STUDY COORDINATORS:** Elisabetta Carini, Sabrina Bagaglio

**STATISTICAL AND MONITORING TEAM:** Laura Galli, Riccardo Lolatto, Sara Diotallevi

**ENROLLING CENTERS:** *ANCONA*: Marcello Tavio, Alessandra Mataloni Paggi; *BARI*: Annalisa Saracino, Flavia Balena; *BERGAMO*: Franco Maggiolo, Laura Comi, Daniela Valenti, Claudia Suardi; *BOLOGNA*: Leonardo Calza, Malerba Federica; *BRESCIA*: Francesco Castelli, Emanuele Focà, Davide Minisci, Francesca Pennati, Anna Celotti, Francesca Brognoli; *BUSTO ARSIZIO*: Barbara Menzaghi, Maddalena Farinazzo; *CATANIA*: Bruno Cacopardo, Maurizio Celestia, Michele Salvatore Paternò Raddusa, Carmen Giarratana; *CATANZARO*: Carlo Torti, Paolo Fusco, Gabriele Bruno; *CREMONA*: Angelo Pan, Paola Brambilla, Chiara Fornabaio; *FIRENZE*: Alessandro Bartoloni, Filippo Lagi, Susanna Giachè, Francesca Vichi, Francesco Maria Fusco, Alessio Bellucci, Elisa Mirabelli, Paola Corsi, Seble Tekle Kiros, Filippo Ducci; *FOGGIA*: Teresa Santantonio, Sergio Lo Caputo, Sergio Ferrara, Marianna Narducci; *GENOVA*: Emanuele Pontali, Marcello Feasi, Antonio Sarà, Matteo Bassetti, Antonio Di Biagio, Sabrina Bianchi; *MILANO*: Antonella Castagna, Vincenzo Spagnuolo, Elisabetta Carini, Sabrina Bagaglio, Laura Galli, Riccardo Lolatto, Andrea Galli, Rebecka Papaioannu, Tommaso Clemente, Sara Diotallevi, Spinello Antinori, Tiziana Formenti, Andrea Giacomelli, Giulia Marchetti, Lidia Gazzola, Federica De Flaviis, Massimo Puoti, Cristina Moioli, Federico D'Amico; *MODENA*: Cristina Mussini, Adriana Cervo, Enrica Roncaglia, Giulia Nardini, Barbara Beghetto; *NAPOLI*: Elio Manzillo, Amedeo Lanzardo; *PADOVA*: Anna Maria Cattelan, Maria Mazzitelli; *PALERMO*: Antonio Cascio, Marcello Trizzino; *PARMA*: Elisa Fronti, Diletta Laccabue; *PAVIA*: Roberto Gulminetti, Andrea Zuccarini; *PERUGIA*: Daniela Francisci, Elisabetta Schiaroli, Giuseppe De Socio; *REGGIO EMILIA*: Elisa Garlassi, Romina Corsini; *ROMA*: Roberta Gagliardini, Marisa Fusto, Loredana Sarmati, Vincenzo Malagnino, Tiziana Mulas, Simona Di Giambenedetto, Silvia Lamonica; *SANREMO*: Giovanni Cenderello, Rachele Pincino; *SIENA*: Mario Tumbarello, Massimiliano Fabbiani, Francesca Panza, Ilaria Rancan; *TORINO*: Giovanni Di Perri, Stefano Bonora, Micol Ferrara; *VERONA*: Marina Malena, Marta Fiscon

**SUPPORTED BY:** ViiV Healthcare, Gilead Sciences, Theratechnologies, MSD

