

Continued late presentation for HIV care across Europe

Amanda Mocroft, Jens D Lundgren and Ole Kirk
for the Late Presentation Working Group of
COHERE

IMIT AMACS ECS-Mothers & ECS-Infants NSHPC-Mothers & NHPS-Infants PISCIS
KOMPNET CASCADE ANRS CO2 SEROCO Frankfurt HIV Cohort Study San Raffaele
ANRS CO1/CO10 EPF UK CHIC Athena ITLR-Mothers & ITLR-Infants Swiss HIV Cohort Study
ICC ANRS CO6 PRIMO Co-RIS MOCHIV-Mothers & MoCHIV-Infants The Italian MASTER Cohort
CHIPS ANRS CO4 French Hospital's Database on HIV HIV-MIP-Mothers & HIV-MIP-Infants
GEMES-Haemo ANRS CO3 AQUITAINE EuroSIDA Madrid Cohort HIV Children VACH
Modena Cohort Study Danish HIV Study ANRS CO8 COPILOTE ICONA St. Pierre
Collaboration of Observational HIV Epidemiological Research Europe
Coordination: Copenhagen HIV Programme (CHIP) & Institut de Santé Publique, d'Epidémiologie et de Développement (ISPED)

Background

- Late Presentation (LP) for HIV defined as CD4 \leq 350/mm³ or AIDS, regardless of CD4 count¹
- Proportion presenting late ~ 50% with considerable variation among different demographic groups²⁻³
- Consequences for the individual and for public health⁴⁻⁵
- Numerous interventions to target HIV testing and increase awareness⁶⁻⁷

¹Antinori et al HIV Med 2011; 12::61-64. ²Mocroft et al, PLoS Med 2013;10(9):E1001510. ³Girardi et al 2004;36:951-959. ⁴Sabin et al AIDS 2004;18:2145-2151. ⁵Quinn et al NEJM 2000;342:921-929. ⁶HIDES Indicator disease testing (<http://hiveurope.eu/Ongoing-Projects/Guidance-HIV-Indicator-Conditions>). ⁷HIV European testing week (<http://hiveurope.eu/Ongoing-Projects/European-HIV-Testing-Week>)

Objectives

- To describe changes in LP in Europe between 2010-2013
- To determine changes over time in new clinical events following HIV+ diagnosis

Study participants

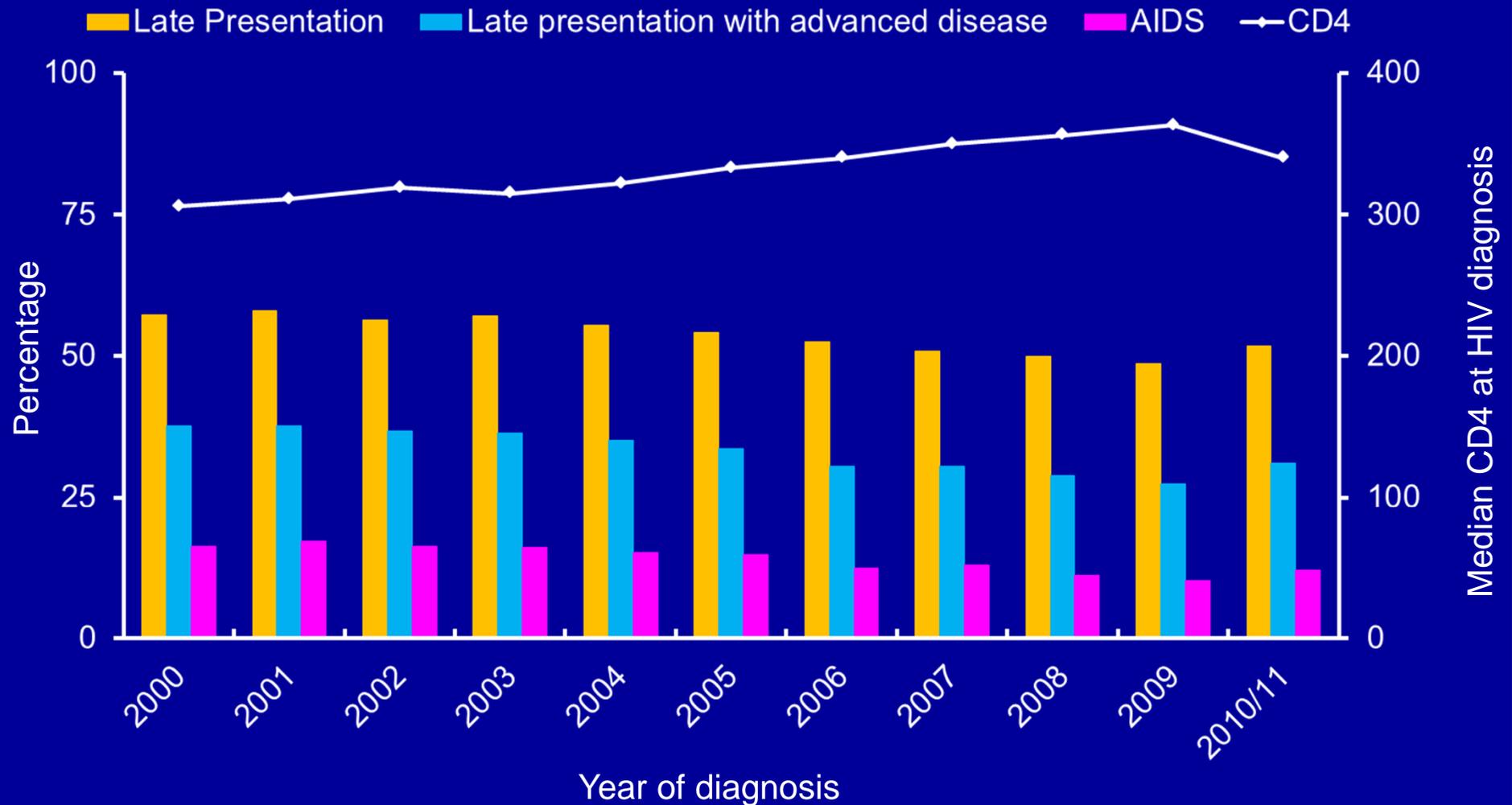
- Participants in COHERE, a collaboration of 33 cohorts from 35 countries testing HIV+ $\geq 1/1/2010$
- CD4 count measured in the 6 months after HIV diagnosis
- Aged > 16 years
- Seroconverter cohorts were excluded

Definitions and methods

- Date HIV+ : date tested HIV+ve or recruitment to cohort/study enrolment where date HIV+ve missing
- LP : $CD4 < 350/mm^3$ in 6 months following HIV+ diagnosis, or AIDS, regardless of CD4 count
- Advanced LP : $CD4 < 200/mm^3$ in 6 months following HIV+ diagnosis, or AIDS, regardless of CD4 count
- Changes over time in proportions of LP assessed using logistic regression
- Kaplan-Meier plots and Cox PH models to examine changes in clinical outcomes (new AIDS/death)

Changes over time in late presentation and CD4 count at HIV-diagnosis : COHERE 2000-2011

Mocroft et al, PLoS Med 2013;10(9):E1001510

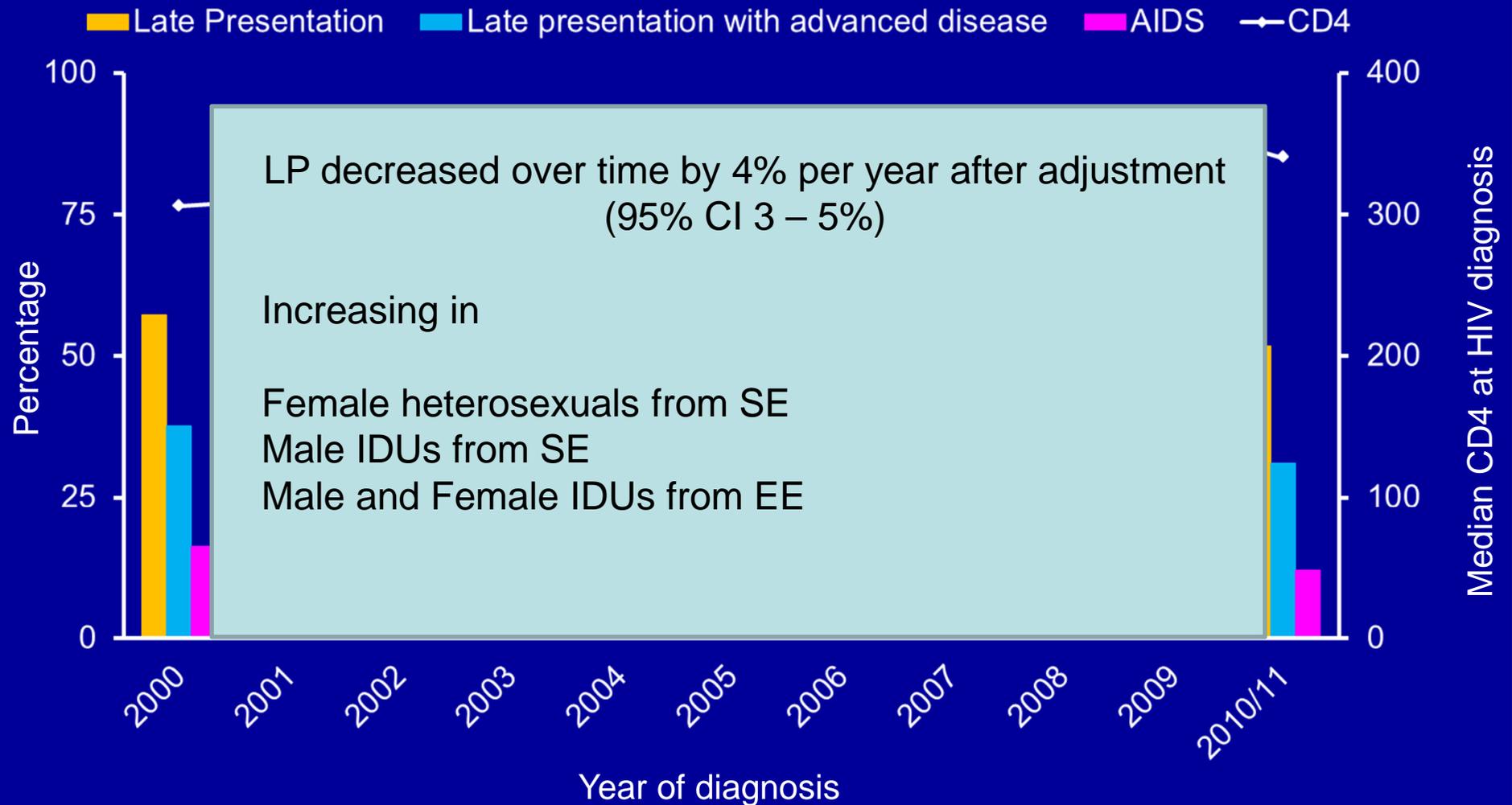


N 7367 7404 8046 7756 8591 8663 8251 8618 9057 7548 3223

Late presentation: diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis. Late presentation with advanced disease: diagnosed with HIV with a CD4 count below 200/mm³ or an AIDS defining event, regardless of CD4 cell count, in the 6 months following HIV diagnosis.

Changes over time in late presentation and CD4 count at HIV-diagnosis : COHERE 2000-2011

Mocroft et al, PLoS Med 2013;10(9):E1001510



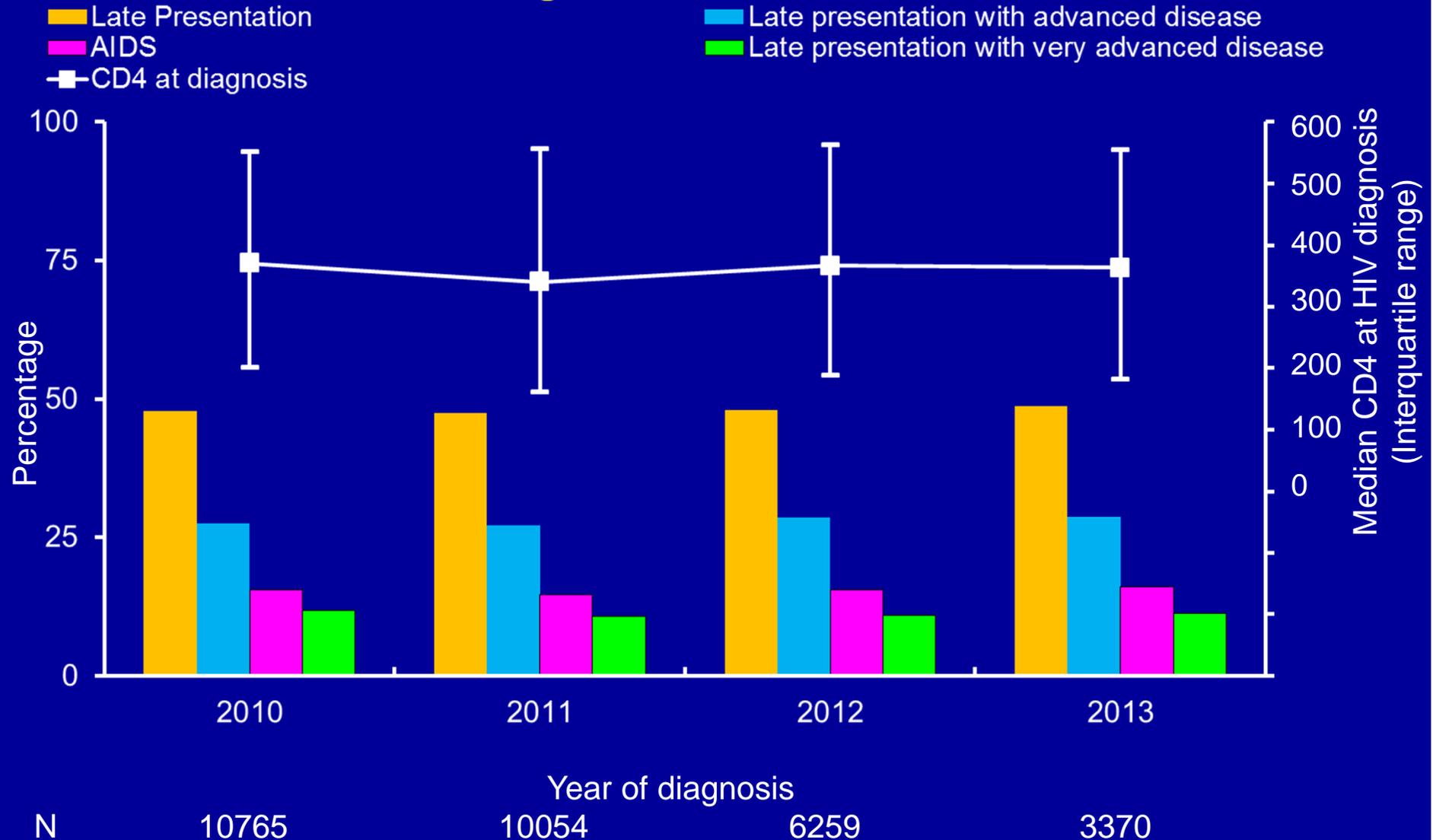
N 7367 7404 8046 7756 8591 8663 8251 8618 9057 7548 3223

Late presentation: diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis. Late presentation with advanced disease: diagnosed with HIV with a CD4 count below 200/mm³ or an AIDS defining event, regardless of CD4 cell count, in the 6 months following HIV diagnosis.

Patient characteristics

		All (N=30448)		LP (N=14582; 47.9%)	
		N	%	N	%
HIV	MSM	15370	50.5	5992	39.0
Exposure	Male heterosexual	4826	15.9	3011	62.4
Group	Female heterosexual	5487	18.0	2864	52.2
	Male injecting drug user	843	2.8	481	57.1
	Female injecting drug user	321	1.1	126	39.3
	Male other	2548	8.4	1493	58.6
	Female other	1053	3.5	615	58.4
European	South	6950	22.8	3133	45.1
Region of	Central	7621	31.6	4789	49.8
Care	North	12570	41.3	6135	48.8
	East	1307	4.3	525	40.2
		Median	IQR	Median	IQR
Age	Years	37	29 -45	39	32 – 48
CD4	/mm ³	368	193 – 555	184	73 – 276

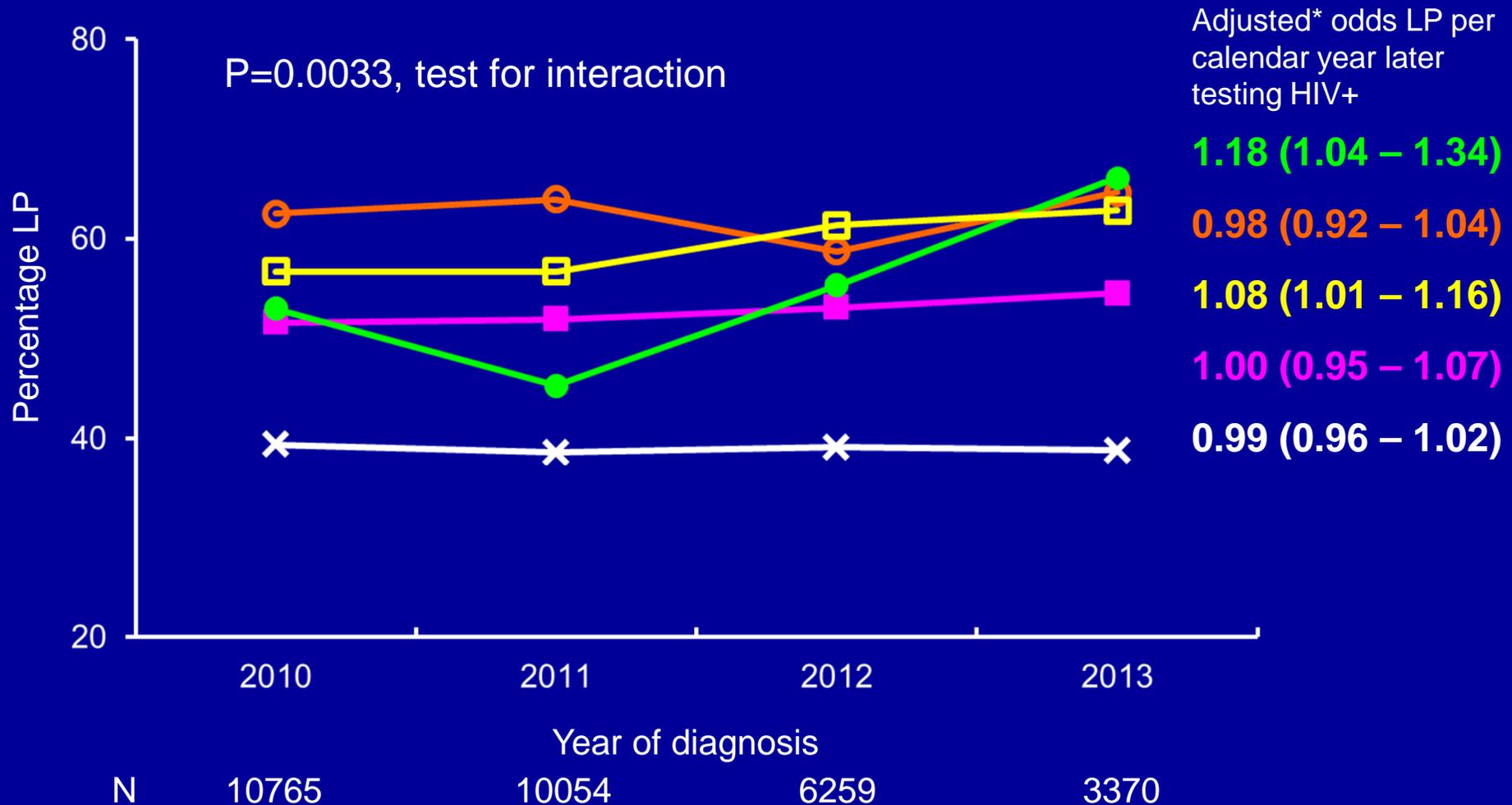
Changes over time in late presentation and CD4 count at HIV-diagnosis : COHERE 2010-2013



Late presentation: diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis. Late presentation with advanced disease: diagnosed with HIV with a CD4 count below 200/mm³ or an AIDS defining event, regardless of CD4 cell count, in the 6 months following HIV diagnosis. Late presentation with very advanced disease: diagnosed with HIV with a CD4 count below 50/mm³ or an AIDS defining event, regardless of CD4 cell count, in the 6 months following HIV diagnosis

Changes over time in late presentation : HIV exposure group

✖ MSM
 ○ M Het
 ■ F Het
 ● IDU
 □ Other



Late presentation: diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis. *Adjusted for age, region of care, region of origin and age

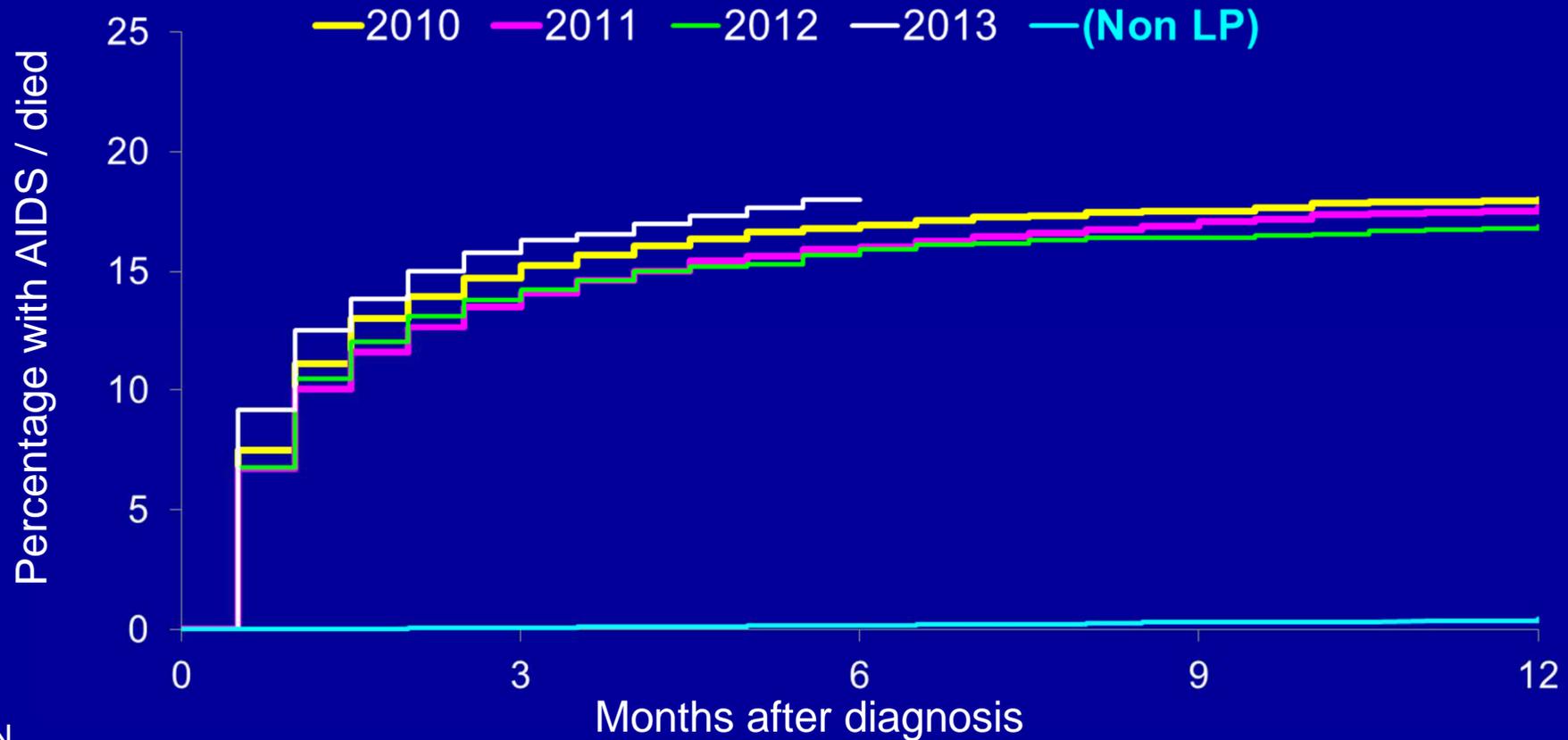
Odds of LP per year later testing HIV+

	Univariate			Multivariate*		
	OR	95% CI	P	OR	95% CI	p
Overall	1.01	0.99 – 1.03	0.40	1.00	0.98 – 1.03	0.84
<u>Region</u>						
South	1.00	0.95 – 1.05	0.95	1.00	0.96 – 1.06	0.87
Central	1.01	0.97 – 1.05	0.64	1.01	0.97 – 1.06	0.61
North	1.01	0.98 – 1.04	0.62	0.98	0.94 – 1.02	0.27
East	1.07	0.91 – 1.25	0.41	1.03	0.88 – 1.21	0.73

P=0.073, test for interaction

Late presentation: diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis. *Adjusted for age, HIV exposure group, region of origin and age

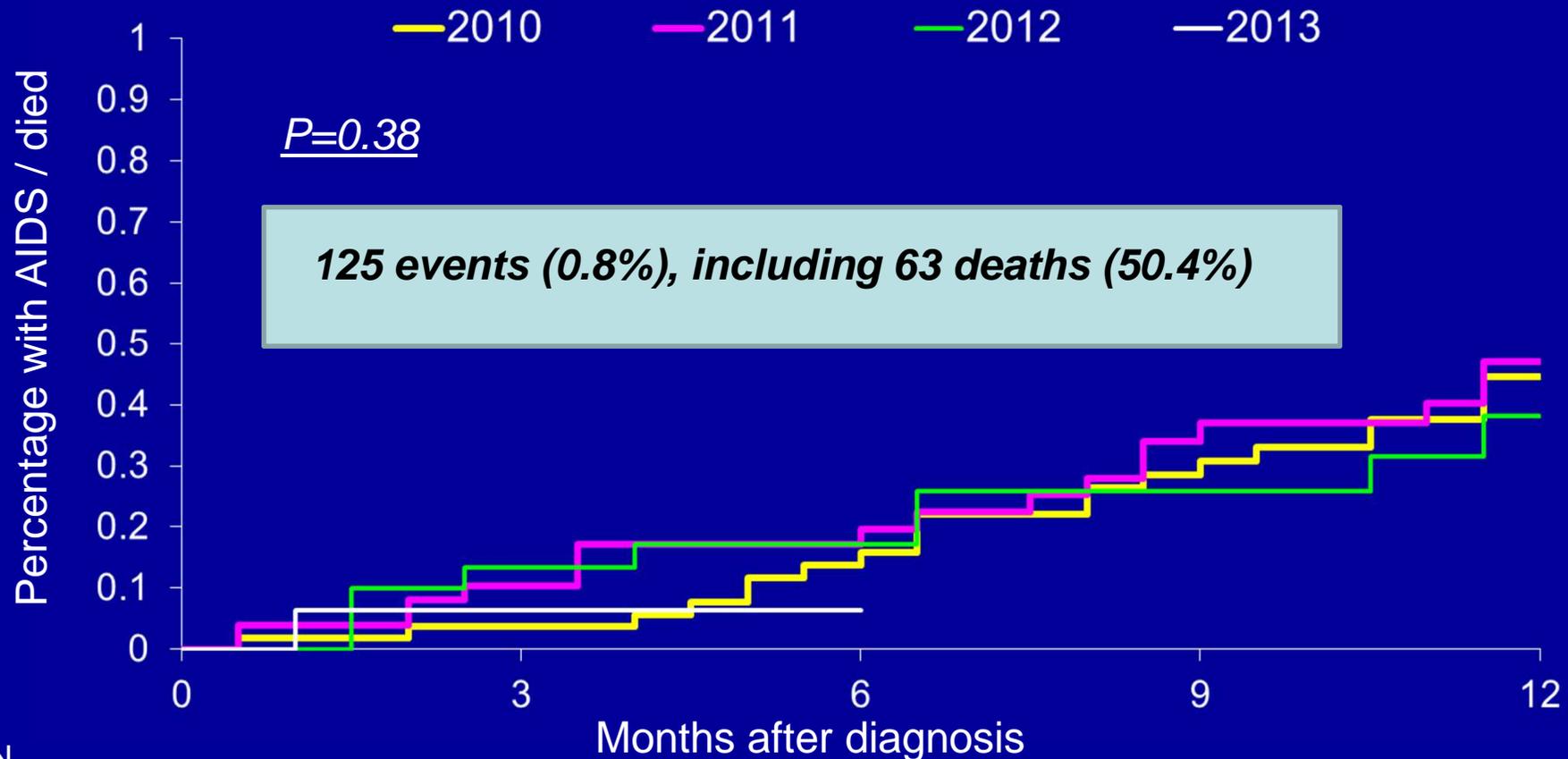
Progression to new AIDS/death up to 1 year after HIV diagnosis in LP and calendar year of testing HIV+



N	2010	2011	2012	2013	(Non LP)
	5130	4736	2988	1613	3796
					3079
					1873
					374
					3326
					2333
					1147
					15

Late presentation (LP): diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis.

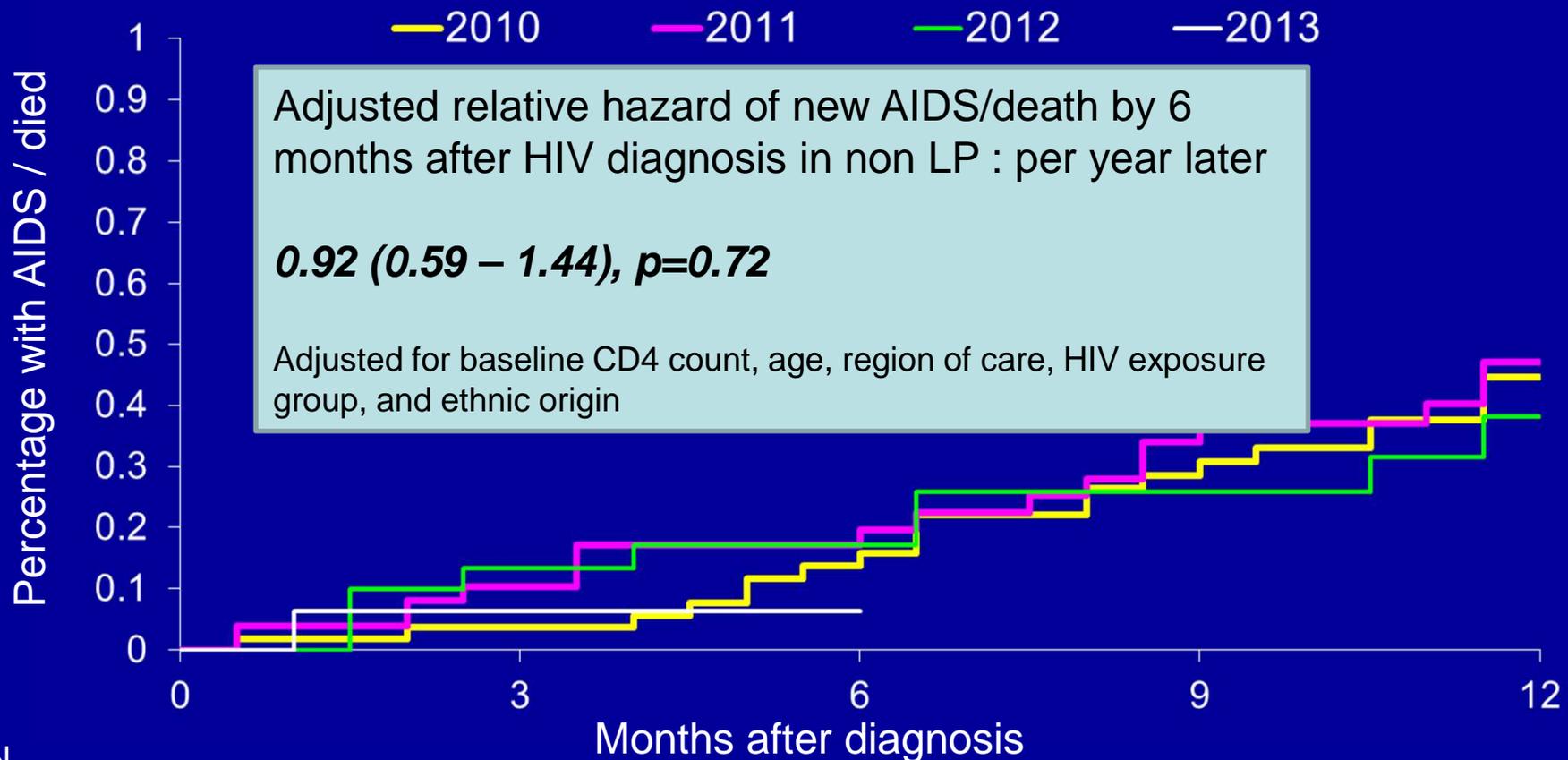
Progression to new AIDS/death up to 1 year after HIV diagnosis in non-LP and calendar year of testing HIV+



N	2010	2011	2012	2013
	5564	5189	3201	1684
				413
				4223
				2878
				1429

Late presentation (LP): diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis.

Progression to new AIDS/death up to 1 year after HIV diagnosis in non-LP and calendar year of testing HIV+



N	2010	2011	2012	2013
	5564	5189	3201	1684
				413
				4223
				2878
				1429

Late presentation (LP): diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis.

Limitations

- Comparatively small numbers from Eastern Europe
- Participants in cohort studies
- Requirement to have CD4 measured
- Date HIV diagnosis unknown for ~10%

Conclusions

- LP remains at 50% and has not decreased since 2010 across Europe, or in any region of Europe, despite the trends seen 2000-2010
- Proportion with advanced LP, very advanced LP, presentation with AIDS alone remained similar over time as did CD4 count at HIV+ diagnosis
- LP remains lowest in MSM (~40%) with some evidence to suggest continued increase over time in IDUs and 'other' group
- Further efforts to reach those not testing required to reduce LP

Acknowledgements



Project leaders and statistical analysis:

Manuel Battegay, Julia Bohlius, Vincent Bouteloup, Heiner Bucher, Alessandro Cozzi-Lepri, François Dabis, Antonella d'Arminio Monforte, Julia del Amo, Maria Dorrucchi, Matthias Egger, Frederik Engsig, Hansjakob Furrer, Ali Judd, Ole Kirk, Olivier Lambotte, Sara Lodi, Sophie Matheron, Laurence Meyer, Jose Miro, Amanda Mocroft, Susana Monge, Fumiyo Nakagawa, Niels Obel, Roger Paredes, Andrew Phillips, Massimo Puoti, Caroline Sabin, Colette Smit, Jonathan Sterne, Rodolphe Thiebaut, Claire Thorne, Carlo Torti, Giota Touloumi, Marc van der Valk, Linda Wittkop, Natasha Wyss

Project Team

Antonella d'Arminio Monforte, Norbert Brockmeyer, Jordi Casabona, Antonella Castagna, Dominique Costagliola, Francois Dabis, Stéphane De Wit, Gerd Fätkenheuer, Hansjakob Furrer Anne Johnson, Ole Kirk, Mario Lazanas, Catherine Leport, Jens Lundgren, Amanda Mocroft, Santiago Moreno, Niels Obel, Frank Post, Peter Reiss, Caroline Sabin, Adriane Skaletz-Rorowski, Ignacio Suarez-Loano, Carlo Torti, Josiane Warszawski, Robert Zangerle

Steering Committee:

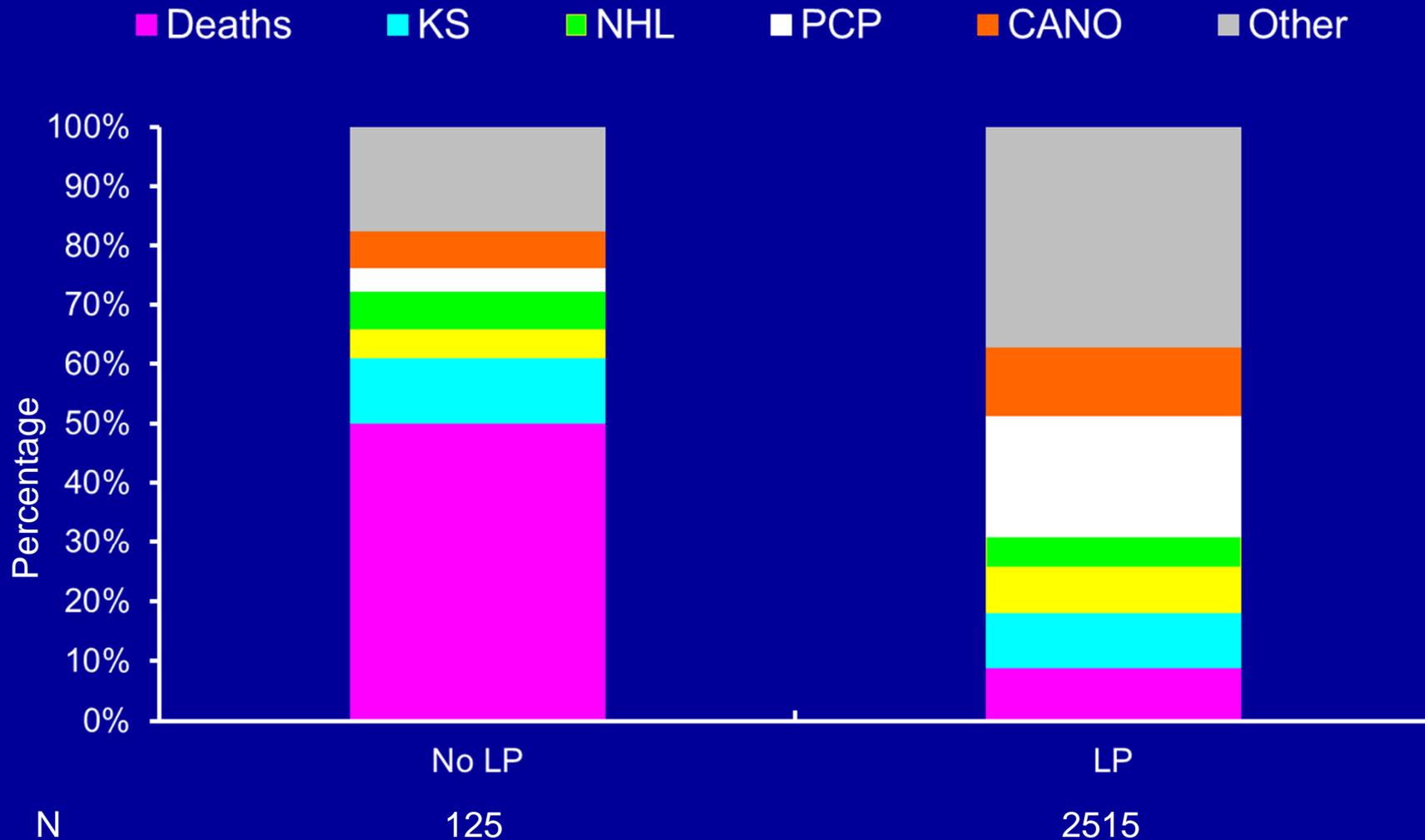
- **Executive committee:** Stéphane De Wit (Chair, St. Pierre Brussels Cohort), Manuel Battegay (SHCS, MoCHIV), Jordi Casabona (PISCIS), Dominique Costagliola (FHDH), Antonella d'Arminio Monforte (ICONA), Julia Del Amo (Co-RIS), Amanda Mocroft (University College London), Dorte Raben (Head, Copenhagen Regional Co-ordinating Center), Geneviève Chêne (Head, Bordeaux Regional Co-ordinating Centre).
- **Contributing cohorts:** Robert Zangerle (AHIVCOS), Giota Touloumi (AMACS), Josiane Warszawski (ANRS CO1 EPF/ANRS CO11 OBSERVATOIRE EPF), Laurence Meyer (ANRS CO2 SEROCO), François Dabis (ANRS CO3 AQUITAINE, and ANRS CO13 HEPAVIH), Murielle Mary Krause (ANRS CO4 FHDH), Jade Ghosn (ANRS CO6 PRIMO), Catherine Leport (ANRS CO8 COPILOTE), Peter Reiss (ATHENA), Ferdinand Wit (ATHENA), Maria Prins (CASCADE), Heiner Bucher (CASCADE), Caroline Sabin (CHIC), Diana Gibb (CHIPS), Gerd Fätkenheuer (Cologne-Bonn), Julia Del Amo (CoRIS), Niels Obel (Danish HIV Cohort), Claire Thorne (ECS), Amanda Mocroft (EuroSIDA), Ole Kirk (EuroSIDA), Christoph Stephan (Frankfurt), Santiago Pérez-Hoyos (GEMES-Haemo), Osamah Hamouda (German ClinSurv), Barbara Bartmeyer (German ClinSurv), Nikoloz Chkhartishvili (Georgian National HIV/AIDS), Antoni Noguera-Julian (CORISPE-cat), Andrea Antinori (ICC), Antonella d'Arminio Monforte (ICONA), Norbert Brockmeyer (KOMPNET), José Ramos (Madrid PMTCT Cohort), Pablo Rojo Conejo (CORISPES-Madrid), Antoni Soriano-Arandes (NENEXP), Manuel Battegay (SHCS), Andri Rauch (SHCS), Cristina Mussini (Modena Cohort), Pat Tookey (NSHPC), Jordi Casabona (PISCIS), Jose M. Miró (PISCIS), Antonella Castagna (San Raffaele), Deborah Konopnick (St. Pierre Cohort), Tessa Goetghebuer (St Pierre Paediatric Cohort), Carlo Torti (Italian Master Cohort), Ramon Teira (VACH), Myriam Garrido (VACH).
- **Paediatric cohort representatives:** Ali Judd, Pablo Rojo Conejo
- **European AIDS Treatment Group:** David Haerry.

Regional Coordinating Centers:

Bordeaux RCC cohorts: Diana Barger, Geneviève Chêne (Head), Christine Schwimmer, Monique Termote
Copenhagen RCC cohorts: Rikke Salbøl Brandt, Maria Campbell, Nina Friis-Møller, Dorte Raben (Head)

Sources of funding: The COHERE study group has received unrestricted funding from: Agence Nationale de Recherches sur le SIDA et les Hépatites Virales (ANRS), France; HIV Monitoring Foundation, the Netherlands; and the Augustinus Foundation, Denmark. COHERE receives funding from the European Union Seventh Framework Programme (FP7/2007-2013) under EuroCoord grant agreement n° 260694. A list of the funders of the participating cohorts can be found on the Regional Coordinating Centre websites at <http://www.cphiv.dk/COHERE/tabid/295/Default.aspx> and <http://etudes.isped.u-bordeaux2.fr/cohere/>.

Clinical events following HIV diagnosis



Late presentation: diagnosed with HIV with a CD4 count below 350/mm³ or an AIDS defining event regardless of the CD4 count, in the 6 months following HIV diagnosis.