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Background

Prevalence of metabolic syndrome (MetS) in people with HIV (PWH) on ART is higher than in general population; age, BMI, and certain ART regimen have been identified as predictors. Nevertheless, the role of CD4 count at diagnosis has not been investigated yet.

Aim

To estimate the incidence of MetS in PWH enrolled in ICONA cohort who started ART when recently HIV infected (RHI), or when chronically infected with CD4 count above (CHI) or below 200 cells/mm3 (advanced HIV disease, AHI).

Materials & Methods

Study population

PWH enrolled in ICONA who started ART from 2008 and excluded those with MetS or MACE before ART.

Definitions

<u>RHI</u> as reporting acute/primary infection and having started ART within 100 days since diagnosis *and* those with a negative HIV test done within 1 year that have started ART within 6 months since the estimated time of infection. <u>Metabolic syndrome</u> as defined by modified NCEP ATP III criteria or, when

missing waist circumference, we used the following equations:

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for men, waist circumference (cm) = $31.2 + 2.4 \times BMI$ (kg/m2)

for women, waist circumference (cm) = $33.2 + 2.1 \times BMI$ (kg/m2)

Statistical analysis

- Retrospective cohort study
- Incidence rates (IRs) of MetS were calculated as the number of events per 100person-years-of-FU (PYFU) with 95% confidence intervals (95%CIs)
- Kaplan-Meier curves estimated cumulative probabilities of the first incident metabolic syndrome on ART
- Univariable and multivariable Cox proportional hazard models were applied to estimate factors associated with the event, adjusting for age, year of ART start, sex, risk factor for HIV acquisition, ethnicity, HCV and HBV coinfection, ART class of the first line, HIV-RNA and CD8 count at ART initiation.

Results

Population characteristics

- Among 13,034 PWH starting ART after 2008 enrolled in ICONA, 11,137 were included in the analysis after excluding those with a diagnosis of MetS (974, 7.47%) or MACE (63, 0.48%) and those lost-to-follow up.

Incidence and prevalence of MetS

- Overall, 2,058 MetS diagnosis after ART start: prevalence of 18.5% (95%CI 17.8-19.2)
- IR of MetS was 3.96 x 100 PYFU (95%CI: 3.8 -4.1)
- Diagnoses of MetS were more frequent in
- Cumulative probability of MetS was significantly higher in advanced HIV infection than in CHI and RHI (p<.001)

Figure 2. Probability of MetS by KM curve

Table1. Main characteristics of PWH with MetS

	СНІ	AHI	RHI	Total	
	N=7,253	N=3,199	N=685	N=11,137	p-value
	(65.1%)	(28.7%)	(6.2%)	(100.0%)	
Male sex	5,942 (81.9%)	2,493 (77.9%)	649 (94.7%)	9,084 (81.6%)	<0.001
Age	38 [30-46]	43[35-51]	34 [27-43]	39 [31-47]	<0.001
Transmission mode					<0.001
Hetero	2,507 (34.6%)	1,629 (50.9%)	115 (16.8%)	4,251 (38.2%)	
IDU	472 (6.5%)	215 (6.7%)	31 (4.5%)	718 (6.4%)	
MSM	3,805 (52.5%)	1,049 (32.8%)	506 (73.9%)	5,360 (48.1%)	0.004
Ethnicity					<0.001
Asian	82 (1.1%)	60 (1.9%)	7 (1.0%)	149 (1.3%)	
Black	619 (8.5%)	398 (12.4%)	36 (5.3%)	1,053 (9.5%)	
Caucasian	6,087 (83.9%)	2,533 (79.2%)	609 (88.9%)	9,229 (82.9%)	
Hispanic/Latino	401 (5.5%)	183 (5.7%)	27 (3.9%)	611 (5.5%)	
Positive HCV status	687 (10.1%)	291 (9.8%)	43 (6.6%)	1,021 (9.8%)	0.019
					0.007
Positive HBsAg status	325 (5.0%)	170 (5.9%)	18 (2.9%)	513 (5.1%)	
	2015	2016	2017	2015	
Year^	[2012-2018]	[2012-2018]	[2015-2019]	[2012-2018]	< 0.001
Months to ART start	7.8 [2.5-60.9]	1.9 [1.0-4.0]	1.6 [0.7-3.2]	4.0 [1.6-29.9]	<0.001
					< 0.001
AIDS	208 (2.9%)	1,032 (32.3%)	17 (2.5%)	1,257 (11.3%)	
					< 0.001

advanced HIV infection than in RHI and CHI (p <0.001)







MetS associated with advanced HIV infection

- Higher aHR of MetS was found for advanced HIV disease vs CHI at multivariable analysis (Figure 3)
- No difference was observed when comparing calendar period 2008-2015 to 2016-2023 (p-value for interaction= 0.420)

Figure 3. Hazard Ratio (HR) and adjusted Hazarad Ratio (aHR) of MetS



Log HIVRNA >5log ^	1,941 (26.9%)	2,113 (67.3%)	353 (52.3%)	4,407 (40.0%)		CHI	1			1			
CD4 count ^	413 [313-554]	73 [30-136]	484 [347-646]	335 [165-492]	<0.001								
		3,199				AHI	1.67	1.52-1.83	<.001	1.39	1.23-1.57	<.001	
<200	0 (0.0%)	(100.0%)	45 (6.6%)	3,244 (29.1%)									
200-350	2,466 (34.0%)	0 (0.0%)	130 (19.0%)	2,596 (23.3%)		D 1 1 1	0.00		0 4 4 4	4 00	0 77 4 00		
350-500	2,428 (33.5%)	0 (0.0%)	183 (26.7%)	2,611 (23.4%)		KHI	0.83	0.67-1.04	0.111	1.00	0.//-1.29	0.994	
>500	2,359 (32.5%)	0 (0.0%)	327 (47.7%)	2,686 (24.1%)									
					0.001								
Abacavir exposure	1,638 (22.6%)	815 (25.5%)	140 (20.4%)	2,593 (23.3%)		Conclusions							
Same anchor drug													
INSTI	2,538 (35.0%)	1,246 (39%)	308 (45.0%)	4,092 (36.7%)	<0.001	 IR of MetS after ART start was 3.96 x 100 PYFU PWH who start ART with CD4 count < 200 cell/mm3 are at higher risk of developing MetS as 							
NNRTI	1,429 (19.7%)	171 (5.3%)	54 (7.9%)	1,654 (14.9%)	<0.001								
PI	466 (14.6%)	625 (8.6%)	29 (4.2%)	1,120 (10.1%)	<0.001								
Years of follow-up	5.4 [2.4-8.4]	4.8 [1.9-8.2]	4.2 [2.0-6.5]	5.1 [2.2-8.2]	<0.001	compared to CHI and RHI							
^ Demographic and immuno-virological characteristics at ART start					• Risk was inde	 Risk was independent of calendar period, unlikely to be mediated by first-line ART regimen 							

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those starting with CD4 count>200 cells/mm3

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