

Trends in proportion of older HIV-infected people in care in Latin America and the Caribbean: a growing challenge

Short Paper

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Abstract

We aimed to quantify the proportion of people receiving care for HIV-infection that are 50 years or older (older HIV patients) in Latin America and the Caribbean between 2000 and 2015 and to estimate the contribution to the growth of this population of people enrolled before (<50yo) and after 50 years old (yo) (≥ 50 yo). We used a series of repeated, cross-sectional measurements over time in the Caribbean, Central and South American network (CCASAnet) cohort. We estimated the percentage of patients retained in care each year that were older HIV patients. For every calendar year, we divided patients into two groups: those who enrolled before age 50 and after age 50. We used logistic regression models to estimate the change in the proportion of older HIV patients between 2000 and 2015. The percentage of CCASAnet HIV patients over 50 years had a threefold increase (8% to 24%) between 2000 and 2015. Most of the growth of this population can be explained by the increasing proportion of people that enrolled before 50 years and aged in care. These changes will impact needs of care for people living with HIV, due to multiple comorbidities and high risk of disability associated with aging.

The proportion of people living with HIV (PLWHIV) that are older than 50 years has been growing worldwide and, according to predictive models, up to 50% of PLWHIV in high-income countries will be over 50 years old (yo) in the next few years [1, 2]. Globally, an estimated 110 000 people aged 50 years or over acquire HIV every year [3]. The latter and increases in life-expectancy of PLWHIV contribute to this demographic change [1]. This trend poses important challenges to health systems, such as the provision of care for co-morbidities resulting from natural aging, effects of chronic inflammation, polypharmacy and lack of guidelines for comprehensive care of elderly patients [4].

According to UNAIDS, between 13% and 15% of adults living with HIV in Latin America and the Caribbean are currently aged 50 or more [5]. In fact, this region has experienced one of the fastest and largest growths of PLWHIV older than 50 years [5]. Nonetheless, these figures come from mathematical models with limited empirical validation [6]. Previous studies have found discrepancies in model estimates that were likely due to lack of representativeness among certain subpopulations, such as local region, gender or age groups [6–8]. More information about the growth of this population can contribute to improved precision and accuracy of regional estimates of the aging population living with HIV. Moreover, as older PLWHIV are more likely to have comorbidities than similarly aged people without HIV, [9] accurate estimates of the numbers older than 50 and temporal trends in our region would help to build models of care and costs for this group in the coming years. Also, PLWHIV ≥ 50 yo and aging in care may contribute differently to the burden of non-communicable diseases than those who are diagnosed or infected later in life [9–12]. Therefore, in this study we aimed to quantify the proportion of people receiving care for HIV that are 50 years or older (≥ 50 yo) in a multi-site cohort of clinics in Latin America and the Caribbean between 2000 and 2015 and to estimate the contribution that people enrolled before or after 50 years of age had on the growth of this population over time.

We used data from all adult patients (≥ 18 years), actively receiving care for HIV during each calendar year (2000–2015) in centers affiliated with the Caribbean, Central and South American network for HIV Epidemiology (CCASAnet). CCASAnet comprises a consortium of HIV health care centers from seven countries (Argentina, Brazil, Chile, Haiti, Honduras,

Mexico and Peru) established in 2006 that share anonymised clinical data to study the epidemiology of HIV-infection in our region [10]. For every calendar year between 2000 and 2015, we generated a cohort of adult patients retained in care for that year. We defined 'retained in care' as those patients who had at least one visit after 15 July for that year and assessed whether they were ≥ 50 yo (i.e., older HIV patients) on that date. Among those classified as older HIV patients, we further distinguished between those who were younger than 50 years when HIV care was initiated and reached 50 during his/her care (<50yo at enrolment) and those who initiated care at 50 or older (≥ 50 yo at enrolment). Each year, patients <50yo at enrolment increased either because patients turned 50 that year or because they turned 50 in previous years and survived and were retained in care at least up to that year. Similarly, patients ≥ 50 yo at enrolment may have enrolled in the cohort that year or enrolled after turning 50 in previous years, survived and retained in care up to that year.

Centers from the CCASAnet cohort included in this study have different periods of enrolment of their patients. The periods included by country are as follows: Argentina from January 1996 to January 2015, Brazil from January 1996 to December 2015, Chile from January 1996 to December 2013, Haiti from

November 1999 to November 2012, Honduras from January 1996 to September 2015, Mexico from November 1997 to December 2015 and Peru from February 1996 to August 2015.

We calculated the percentage of older HIV patients relative to the total number of patients retained in care by the calendar year, the distribution of older patients in care who enrolled <50yo at enrolment vs. ≥ 50 yo at enrolment and the percentage of patients entering the group of older HIV each year. We compared socio-demographic characteristics based on age at enrolment using Wilcoxon Rank Sum tests for continuous variables and Chi-square tests for discrete variables. Separate logistic regression models were fit to assess trends in the proportion of older HIV patients over time for each center and overall; calendar year was included in the model using natural splines with three knots. Analyses were performed at INCMNSZ-Mexico using R Statistical Software (www.R-project.org). Analysis scripts are available at biostat.mc.vanderbilt.edu/ArchivedAnalyses.

Among 24 317 adult patients retained in care at any given year during the study period, 5505 were older than 50 years of age. Of these, there were 2789 (51%) <50yo at enrolment who aged in care and 2716(49%) already ≥ 50 yo at enrolment. Demographic and clinical characteristics at enrolment in care for these two groups

Table 1. Comparison of demographic and clinical characteristics at enrolment in CCASAnet centers between patients that were younger than 50 years at enrolment (<50 years at enrolment) and patients enrolled at 50 years or older (≥ 50 year at enrolment) (2000–2015)

Characteristics ^a	<50 years at enrolment (n = 2789)	≥ 50 years at enrolment (n = 2716)	P-value
Patient age at enrolment in care (years)	46 (42–48)	55 (52–59)	NA ^b
Male	1901 (68%)	1797 (66%)	0.121
Naive at enrolment in care	2245 (80%)	2200 (81%)	0.63
CD4 at enrolment in care, (cells/ μ l)	188 (75–338)	196 (85–341)	0.049
CD4 at entrance into the fifties cohort, (cells/ μ l)	442 (290–622)	196 (85–341)	<0.001
AIDS or CD4<200 at enrolment in care, n (%)	1485 (53%)	1407 (52%)	0.297
Probable route of transmission, n (%)			
Heterosexual	1021 (37%)	952 (35%)	
Homosexual	547 (20%)	351 (13%)	
Other	319 (11%)	193 (7%)	
Unknown	902 (32%)	1220 (45%)	
Calendar year of enrolment	2004 (2002–2007)	2008 (2005–2011)	<0.001
ART initiation, n (%)	2736 (98%)	2628 (97%)	0.002
Time from enrolment to ART initiation (months) ^c	0.09 (0–2.76)	0 (0–0.82)	<0.001
Centre ^d , n (%)			<0.001
CMH/FH-Argentina	632 (23%)	578 (21%)	
FC-Brazil	694 (25%)	461 (17%)	
FA-Chile	355 (13%)	226 (8%)	
GHESKIO-Haiti	590 (21%)	882 (32%)	
IHSS/HE-Honduras	124 (4%)	97 (4%)	
INCMNSZ-México	123 (4%)	125 (5%)	
IMTAvH-Perú	271 (10%)	347 (13%)	

^aContinuous variables are reported as medians (interquartile range).

^bNot applicable.

^cPatients who started ART prior to enrolment were assigned a time of 0 months.

^dFH-Argentina: Fundación Huesped, Buenos Aires, Argentina ; FC- Brazil: Instituto Nacional de Infectologia Evandro Chagas-Fiocruz, Rio de Janeiro, Brazil; FA-Chile: Fundación Arriarán, Santiago, Chile; IHSS/HE-Honduras: Instituto Hondureño de Seguridad Social and Hospital Escuela, Tegucigalpa, Honduras; INCMNSZ-Mexico: Instituto Nacional de Ciencias Médicas y Nutrición Salvador Zubirán, Mexico City, Mexico; IMTAvH-Perú: Instituto de Medicina Tropical Alexander von Humboldt, Lima, Peru.

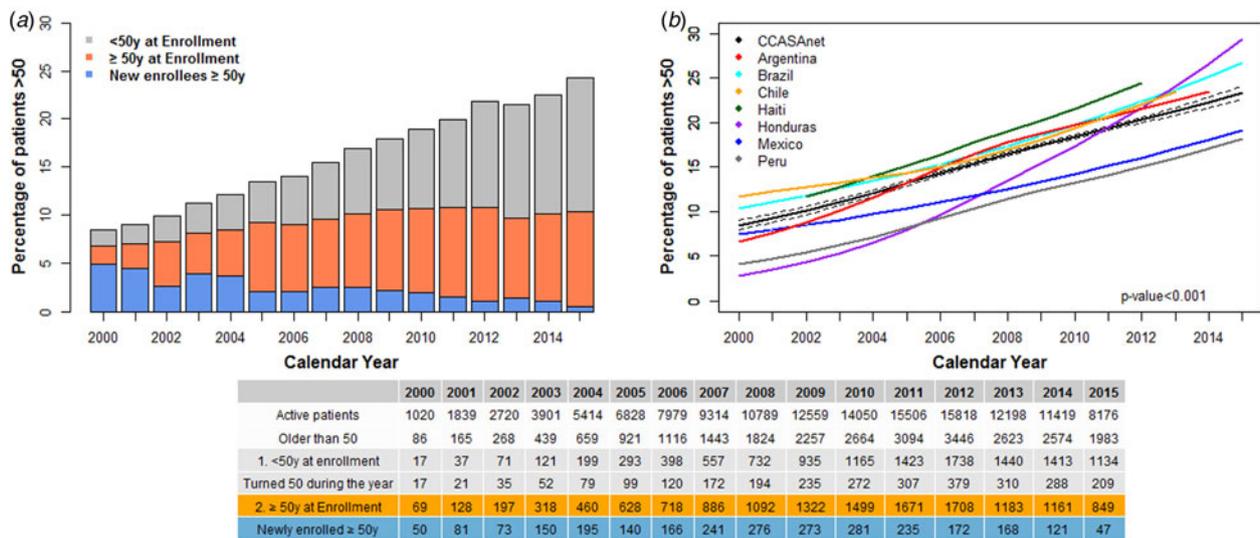


Fig. 1. (a) Percentage of HIV + patients older than 50 years actively receiving care by a group of age and calendar year and (b). The trend of percentage of patients older than 50 among active patients by site and calendar year. (a) Groups of age at enrollment were: (1) <50yo at enrolment and (2) ≥50 at enrolment. In the <50yo at enrolment group we included active patients who enrolled in care with less than 50 years and aged to over 50 while in care (gray). In the ≥50 at enrolment group, we included patients who enrolled in care at 50 years or older (blue + orange). Percentage of new patients enrolled older than 50 at each year are shown by the blue bar. (b) The increase in the percentage of patients older than 50 along time is statistically significant for each site (P -values for a test of tendency are <0.001 for each country and for the whole cohort). The curves are predictions from logistic models adjusted by year and site. The absolute number of all active patients, those who are older than 50 and their classification according to age at enrolment by year, are shown in the table; decreases in active patients from 2013 to 2015 were due to lack of available data from Haiti.

of patients are presented in Table 1. The percentage of male patients, frequency of heterosexual transmission and frequency of AIDS at enrollment were fairly similar between groups. In contrast, people aging in care had a slightly, but statistically lower median CD4 count at enrolment than people who enrolled in care after 50yo.

The percentage of older HIV patients receiving care in CCASAnet increased from 8% in 2000 to 24% in 2015 (Fig. 1a). This upward trend was observed at all CCASAnet centers (Fig. 1b and Table S1). Among all the older HIV patients, the percentage who were <50yo at enrolment substantially increased over time, from 20% ($n = 17/86$) in 2000 to 57% ($n = 1134/1983$) in 2015. Moreover, the percentage of <50yo at enrolment that turned 50yo each calendar year, ranged from 57% ($n = 21/37$) in 2001 to 18% ($n = 209/1134$) in 2015. Additionally, the percentage of ≥50yo at enrolment that was newly enrolled each year also decreased over time, ranging from 72% ($n = 50/69$) in 2000 to 5% ($n = 47/849$) in 2015.

In summary, we observed that by 2015, 24% of patients receiving care for HIV in seven centers in the Caribbean and Latin America were older than 50 years. This figure is a threefold increase in the last 15 years and higher than the UNAIDS mathematical model estimates of 15% [5]. Notably, clinics in Haiti, Honduras and Peru had the fastest upward trend in the relative proportion of people older than 50yo receiving care over time. People turning 50yo each year and the newly enrolled after 50yo, decreased proportionally over time. Thus, in our cohort, most of the growth of this population can be explained by the increasing survival and retention in care of all patients regardless of their age at enrolment, but to a greater extent by those who enrolled younger and aged in care (see Fig. 1a). By the end of the observation period, those enrolled in previous years that survived and were retained in care accounted for the majority of older HIV patients.

While we cannot be certain why our estimates differ from the UNAIDS mathematical model, we posit discrepancies may be due to differences in available data. For example, centers in the CCASAnet cohort typically started provision of care early during the HIV epidemic and were already fully operating during the expansion of universal access to ART in our region between 2002 and 2004, before most patients in the region were started on ART [11]. If this resulted in higher survival rates, our findings, despite coming from a large number of individuals, might not fully represent the whole region. Alternatively, the parameters used by UNAIDS could be affected by the lack of availability and quality of specific surveillance and survey data in each country. In contrast, our data are collected in real settings for a long period of time and include large numbers of patients, which allow us to perform appropriate descriptions of demographic trends among HIV patients in care along time in the region.

Our findings suggest that current models may be underestimating the proportion of older PLWHIV in Latin America and the Caribbean. In any case, with improvements in life expectancy among patients on ART and continued new HIV diagnoses (and new HIV infections) in those aged over 50 years, [13] we can anticipate this trend towards an older HIV population to continue. This demographic shift will impact the needs of care due to the multiple comorbidities and the high risk of disability associated with ageing [14]. The former implies specialised care in tertiary facilities and higher costs. Estimating such costs through modelling studies could help plan an integral approach to the care of older PLWHIV and their future needs.

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Conflicts of interest. Y Caro-Vega, PF Belaunzarán-Zamudio, Jessica L Castilho, B Crabtree-Ramírez, BE Shepherd, F Mejia, MJ Giganti, D Padgett, Beatriz Grinsztejn, Jean W Pape and P Patterson have no conflicts of interest to declare. Catherine C McGowan reports grants from the US NIH. Dr Sierra-Madero reports personal fees and non-financial support from Gilead, non-financial support from MSD, grants from BMS, grants from Pfizer, personal fees from Jansen all outside the submitted work.

References

1. Kirk JB and Goetz MB (2009) Human immunodeficiency virus in an aging population, a complication of success. *Journal of the American Geriatrics Society* 57, 2129–2138.
2. Smit M *et al.* (2015) Future challenges for clinical care of an aging population infected with HIV: a modelling study. *Lancet Infectious Diseases* 15, 810–818, (Erratum in: *Lancet Infectious Diseases* 2015; 15: 998).
3. Joint United Nations Programme on HIV/AIDS (2014) The Gap Report 2014. People aged 50 years and older. [Internet]. Geneva. Available at http://www.unaids.org/sites/default/files/media_asset/12_Peopleaged50year_sandolder.pdf.
4. Martin CP, Fain MJ and Klotz SA (2008) The older HIV-positive adult: a critical review of the medical literature. *The American Journal of Medicine* 121, 1032–1037.
5. Joint United Nations Programme on HIV/AIDS (2013) HIV and aging. A special supplement to the UNAIDS report on the global AIDS epidemic 2013. Available at http://www.unaids.org/sites/default/files/media_asset/20131101_JC2563_hiv-and-aging_en_0.pdf.
6. Silhol R *et al.* (2017) Empirical validation of the UNAIDS spectrum model for subnational HIV estimates: case-study of children and adults in Manicaland, Zimbabwe. *Aids (London, England)* 31(suppl. 1), S41–S50.
7. Michael D *et al.* (2014) Does the spectrum model accurately predict trends in adult mortality? Evaluation of model estimates using empirical data from a rural HIV community cohort study in north-western Tanzania. *Global Health Action* 7(1), 21783. eCollection 2014. Erratum in: *Global Health Action* 2014;7:24079.
8. Eaton JW *et al.* (2015) Assessment of epidemic projections using recent HIV survey data in South Africa: a validation analysis of ten mathematical models of HIV epidemiology in the antiretroviral therapy era. *Lancet Global Health* (10), e598–e608. doi: 10.1016/S2214-109X(15)00080-7.
9. Guaraldi G *et al.* (2015) Aging with HIV vs. HIV seroconversion at older age: a diverse population with distinct comorbidity profiles. *PLoS ONE* 10(4), e0118531. Available from: <https://doi.org/10.1371/journal.pone.0118531>.
10. McGowan CC *et al.* (2007) Cohort profile: Caribbean, Central and South America network for HIV research (CCASAnet) collaboration within the international epidemiologic databases to evaluate AIDS (IeDEA) programme. *International Journal of Epidemiology* 36, 969–976.
11. Belaunzarán-Zamudio PF *et al.* (2015) Monitoring of HIV treatment in seven countries in the WHO region of the Americas. *Bulletin of the World Health Organization* 93, 529–539.
12. Belaunzarán-Zamudio PF *et al.* (2017) Increased age-adjusted mortality and incidence of non-AIDS defining events among people living with HIV enrolled after 50yo and aging in care in Latin America. A CCASAnet cohort study [Abstract O133]. HIV & hepatitis in the Americas 2017. *Journal of the International AIDS Society* 20(suppl. 2), 3.
13. Sabin CA and Reiss P (2017) Epidemiology of aging with HIV: what can we learn from cohorts? *Aids (London, England)* 31(suppl. 2), S121–S128.
14. Ávila-Funes JA *et al.* (2016) Correlates of prevalent disability among HIV-infected elderly patients. *AIDS Research & Human Retroviruses* 32, 155–162.