## Progression to WHO criteria for antiretroviral therapy in a 7-year cohort of adult HIV-1 seroconverters in Abidjan, Côte d'Ivoire

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**Objective** To estimate the probability of reaching the criteria for starting highly active antiretroviral therapy (HAART) in a prospective cohort of adult HIV-1 seroconverters in Abidjan, Côte d'Ivoire.

**Methods** We recruited participants from HIV-positive donors at the blood bank of Abidjan for whom the delay since the estimated date of seroconversion (midpoint between last negative and first positive HIV-1 test) was  $\leq$  36 months. Participants were offered early trimethoprim-sulfamethoxazole (cotrimoxazole) prophylaxis, twice-yearly measurement of CD4 count and we made standardized records of morbidity. We used the Kaplan–Meier method to estimate the probability of reaching the criteria for starting HAART according to WHO 2006 guidelines.

**Findings** 217 adults (77 women (35%)) were followed up during 668 person-years (PY). The most frequent diseases recorded were mild bacterial diseases (6.0 per 100 PY), malaria (3.6/100 PY), herpes zoster (3.4/100 PY), severe bacterial diseases (3.1/100 PY) and tuberculosis (2.1/100 PY). The probability of reaching the WHO 2006 criteria for HAART initiation was estimated at 0.09, 0.16, 0.24, 0.36 and 0.44 at 1, 2, 3, 4 and 5 years, respectively.

**Conclusion** Our data underline the incidence of the early HIV morbidity in an Ivorian adult population and provide support for HIV testing to be made more readily available and for early follow-up of HIV-infected adults in West Africa.

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Une traduction en français de ce résumé figure à la fin de l'article. Al final del artículo se facilita una traducción al español.

الترجمة العربية لهذه الخلاصة في نهاية النص الكامل لهذه المقالة.

#### Introduction

Experts from the World Health Organization (WHO) and the United Nations Joint Programme on AIDS (UNAIDS) have estimated that at the end of 2005, 65% of the 39 million people with HIV worldwide were living in sub-Saharan Africa.<sup>1</sup>

In Africa, only a small proportion of people who need highly active antiretroviral therapy (HAART) are currently being treated. However, the number who receive HAART has been increasing rapidly since 2004, and will continue to increase within the next few years thanks to national and international funds and initiatives.

In 2003 and 2006, WHO experts defined criteria for initiating HAART in low-resource areas based on clinical criteria and, depending on the availability of laboratory facilities, either CD4 count or total lymphocyte count. Since cohort studies of patients from the time of HIV seroconversion are rare, little is known about the natural history of HIV/AIDS in sub-Saharan Africa and, therefore, about the probability of patients reaching WHO clinical and/or biological HAART starting criteria.

Here, we describe early HIV and non-HIV morbidity, time to first WHOstage II, III or IV clinical events, and time to reach WHO criteria for starting HAART in a cohort of HIV-infected adults recruited shortly after their estimated date of HIV-seroconversion at the national blood bank of Abidjan, Côte d'Ivoire.

#### Methods Patients

Recruitment for the The Primo-CI ANRS 1220 prospective cohort started in 1997 in the HIV care clinic (FonSIDA clinic) of the Centre National de Transfusion Sanguine (CNTS), at the national blood bank of Abidjan.<sup>2</sup> The study protocol was approved by the ethics committees of the national Ivorian program on AIDS and the institutional review board of the French Agency for Research on AIDS (ANRS).

In the Côte d'Ivoire, blood donors are adult volunteers who are not paid. Blood donations are limited to five per year in men and four per year in women. Informed consent to test for HIV, syphilis, hepatitis B and hepatitis C is obtained at each donation. Patients are informed that they can, if they wish, obtain results of these tests at the FonSIDA clinic after a post-test counselling session.

From 9 June 1997 enrolment in the Primo-CI cohort was proposed to each blood donor who: (i) was diagnosed with HIV-1 or HIV-1 and HIV-2 co-infection during a blood donation; (ii) had been HIV-seronegative at the preceding donation; (iii) had returned to the clinic for their HIV test result; and (iv) had a delay of  $\leq 36$  months since the estimated date of seroconversion. We used the midpoint between the last negative and the first positive HIV test to estimate the date of seroconversion.

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After obtaining written informed consent to participate in this study, we asked patients to return to the clinic every six months, or at any time in the case of medical problems. All drugs, investigations, hospital stays and transport to and from the clinic were offered free of charge.

#### Follow-up

We used standardized forms to record baseline and follow up sociodemographic and clinical data. Patients were examined by clinicians at enrolment and every six months. Patients also had free access to services at the study clinic between the six-month visits. Symptomatic patients were managed according to pre-defined standardized algorithms, including laboratory X-ray investigations and standardized first-line treatment regimens for most frequent syndromes.<sup>3</sup> Patients needing parenteral treatment lasting less than 12 hours were managed in the day-care hospital at the study clinic. For life-threatening diseases or for conditions necessitating overnight care, patients were referred to the Treichville University Hospital in Abidjan. All clinical events were reviewed by an event documentation committee. The diagnostic criteria were the same as those used in the Cotrame cohort, a cohort of HIV-infected adults followed up in Abidjan during the same period, under the supervision of the same Franco-Ivoirian team, and for which procedures are described elsewhere.<sup>3,4</sup> Autopsies were not performed.

Tuberculosis prophylaxis was not offered, as it is not recommended by the Côte d'Ivoire program on AIDS and tuberculosis. Trimethoprim-sulfamethoxazole (cotrimoxazole) prophylaxis (800/160 mg daily) was systematically offered to all patients at enrolment. Before February 2001, no antiretroviral treatment was available. Since February 2001, the clinic became one of the centres of the UNAIDS/Côte d'Ivoire initiative for improving access to HIV drugs, and HAART was provided in accordance with international guidelines for resource-limited settings.<sup>5,6</sup>

#### **Biological measurements**

The diagnosis of HIV infection was based on two repeated reactive enzymelinked immunosorbent assays (ELISA) (Murex ICE 1–0-2 (Abbott, North Chicago, IL, USA) and Vironostika HIV Uni-Form II (Organon Teknika BV,

Table 1. Baseline and follow-up characteristics of 217 HIV-1 seroconverters from
the ANRS 1220 cohort study, Abidjan, 1997–2004

Baseline characteristic <sup>a</sup>		
Male sex	140	(65%)
Age in years (median (range))	29	(19 – 53)
Heterosexual transmission	215	(99%)
Estimated time from seroconversion in months (median (IQR))	8.9	(4.9 – 18.0)
Schooling		
Illiterate	13	(6%)
≤ Primary school level	48	(22%)
≥ Secondary school level or above	156	(72%)
Unemployed	93	(43%)
Marital status		
Single	134	(62%)
Free union or married	74	(34%)
Divorced or widowed	9	(4%)
CD4-cell count $\leq$ 350/mm <sup>3</sup>	73	(34%)
HIV-1 RNA level in copies/mL (median (IQR))	4.7	(4.1–5.2)
CD4-cell count in per mm <sup>3</sup> (median (IQR))	461	(333–650)
Haemoglobin in g/L (median (IQR))	12.7	(11.6–14.0)
Hepatitis B surface antigen positive serum	9	(4%)
Body mass index in kg/m <sup>2</sup> (median (IQR))	22.2	(20.5–24.7)
Status at study termination		
HAART non initiated	159	(73%)
Dead	12	(6%)
Lost to follow-up <sup>c</sup>	21	(10%)
Alive	126	(58%)
HAART initiated	58	(27%)
Dead	2	(1%)
Lost to follow-up	0	(0%)
Alive	56	(26%)
Cumulative follow-up (person-years)	668	-
Follow-up in months (median (IQR))	33.5	(18.0, 54.1)

IQR, interquartile range

<sup>a</sup> Data are number (%) unless otherwise stated.

<sup>b</sup> Last contact with study team before 31 December 2004 and no further information on vital status at 31 December 2005.

Boxtel, Netherlands)). Before enrolment, we confirmed HIV infection and differentiated between HIV-1 and HIV-2 using two ELISAs (Murex ICE 1–0-2, and Pepti-Lav 1-2 (Pasteur Diagnostics, Marnes-la-Coquette, France)).

Blood samples were collected at inclusion and every six months thereafter to measure CD4 count (FACScan, Becton Dickinson, Aalst-Erembodegem, Belgium) and plasma HIV-1 RNA level (HIV-1 Monitor, version 1.5; Roche Diagnostics, Indianapolis IN, USA).

Microbiological procedures were the same as those used in the Cotrame cohort.<sup>3,4</sup>

#### **Statistical analysis**

Baseline was the date of inclusion in the cohort. For this analysis, data were censored on 31 December 2004 for patients

who were still living, or the date of death for patients who had died. For patients who were lost to follow-up before 31 December 2004, data were censored at the date of their last contact with the study team. However, we used tracing procedures for up to one year after the date of data censoring. For patients whose last contact with the study team was before 31 December 2004 but who, via the tracing system, were found to be alive or have died during 2005 were considered to be alive on 31 December 2004.

The incidence of a given event was defined as the number of patients with an event per 100 patient-years (PY) of at-risk follow-up. The at-risk period began at enrolment and continued to the date of the first event or to the end of the study period. Patients were censored on HAART initiation, first event or default. Default was the date of the last contact with the cohort team for all patients who were not receiving HAART and whose last contact was before 31 December 2004.

The analyses had three steps. First, we used the Kaplan-Meier method to estimate the probability of survival and event-free survival from enrolment to first WHO stage II, III or IV defining event, and the probability of remaining free of criteria for starting HAART. Criteria for HAART initiation were successively defined in accordance with WHO 2003-2006 recommendations for settings where measurement of CD4 count was possible (WHO stage IV; WHO stage III with CD4 < 350/mm<sup>3</sup>; or CD4 < 200/mm<sup>3</sup>, irrespective of clinical stage);<sup>6,7</sup> and WHO 2003-2006 recommendations for settings where measurement of CD4 count was unavailable (WHO stage III or IV; or WHO stage II and total lymphocyte count < 1200/mm<sup>3</sup>).<sup>6,7</sup>

Second, in order to assess how initiating HAART at earlier stages of HIV disease would affect the probability of starting treatment, we modified WHO criteria and considered the effect of offering HAART to all patients at WHO stage III irrespective of CD4 count, and all patients at stage II with CD4 < 350/mm<sup>3</sup>. Following this option, criteria for starting HAART became: WHO stage III or stage IV; WHO stage II with CD4 < 350/mm<sup>3</sup>; or CD4 < 200/mm<sup>3</sup> irrespective of clinical stage. Throughout the article, we refer to these parameters as "alternative early-initiation criteria".

Third, in a second set of analyses, survival and event-free survival were also estimated from the estimated date of seroconversion (and not from the date of enrolment in the cohort). Incidence rates and survival probabilities were given with 95% confidence interval (95% CI).

#### Results

#### **Patients**

By the end of 2004, 217 seroconverters (216 with HIV-1 and one with HIV-1 and HIV-2 co-infection) were enrolled in the cohort. Median time between the last HIV-negative test and the first HIV-positive test was seven months (interquartile range (IQR) 3–17). Median time from the estimated date of seroconversion to enrolment in the cohort was 9 months (IQR 5–18). Other baseline and follow-up characteristics are

Table 2. Causes of morbidity in the ANRS 1220 cohort study of HIV-1 seroconverters, Abidjan, 1997 – 2004

Classifying diseases	Patients (n)	Events (n)
WHO stage II	38	55
Herpes zoster	22	23
Fungal nail infections	7	8
Angular stomatitis	9	11
Prurigo	12	12
Sinusitis	1	1
WHO stage III	30	44
Oral candidiasis	10	13
Pulmonary tuberculosis	10	10
Severe bacterial diseases <sup>a</sup>	14	18
Vaginal candidiasis	1	3
WHO stage IV	15	20
Severe bacterial diseases <sup>b</sup>	7	8
HIV wasting syndrome	2	2
Extra-pulmonary tuberculosis	4	4
Oesophageal candidiasis	2	2
Cryptococcal meningitis	1	1
HIV encephalopathy	1	1
Kaposi's sarcoma	1	1
Cerebral toxoplasmosis	1	1
Non-classifying diseases	77	316
Malaria	22	25
Non-classifying bacterial diseases <sup>c</sup>	37	59

<sup>a</sup> Including pneumonia with bacteraemia (n=3), pneumococcus meningitis (n=1), bacterial enteritis (n=11), sinusitis with bacteraemia (n=1).

<sup>b</sup> All *Salmonella typhi* bactaeramia.

<sup>c</sup> Including sinusitis (n=4), otitis (n=12), enteritis (n=4), cystitis (n=5), uretritis (n=5), skin abscess (n=18), amygdalitis (n=5), boils (n=6), all with no WHO stage III or IV classifying diseases criteria.

shown in Table 1.

At inclusion, 146 (67%) patients reported symptoms compatible with an acute retroviral infection syndrome in the preceding months, including acute fever (n = 125), rash (n = 17), pharyngitis (n = 31), acute diarrhoea (n = 44) and herpes zoster (n = 5). At baseline examination, 17 patients had polylymphadenopathy. By the end of the study period, 58 patients had started HAART. Of these patients on treatment, none was lost to follow-up and all but one patient (a woman who continued in active follow-up but refused to continue antiretroviral drugs) were still receiving HAART at study termination.

#### Survival

Twelve patients died during follow-up, including eight (67% of deaths) at Centres for Disease Control and Prevention (CDC) clinical stage C. At the time of death, the median last available CD4 count was 94/mm<sup>3</sup> (IQR 19–372) and the median last plasma HIV-1 RNA 5.1 log<sub>10</sub> copies/mm<sup>3</sup> (range: 4.8–6.1).

Causes of death included cerebral toxoplasmosis (n = 1, last CD4 = 9/mm<sup>3</sup>); tuberculosis (n = 4, last CD4 = 4/mm<sup>3</sup>, 27/mm<sup>3</sup>, 22/mm<sup>3</sup> and 374/mm<sup>3</sup>); and pneumococcus meningitis (n = 1, last  $CD4 = 666/mm^3$ ). Cause of death was unknown in the remaining patients  $(n = 6, \text{ last } \text{CD4} = 5/\text{mm}^3, 26/\text{mm}^3,$ 161/mm<sup>3</sup>, 372/mm<sup>3</sup>, 288/mm<sup>3</sup> and 572/mm<sup>3</sup>). Of the seven patients who had a CD4 count less than 200/mm<sup>3</sup> at the time of death, five died before HAART was available in Abidjan and two died after the Côte d'Ivoire HAART programme had begun, but they refused treatment. The probability of survival at 1, 2, 3, 4 and 5 years after enrolment was estimated at 0.99 (95% CI 0.97-1.00), 0.96 (95% CI 0.93-0.99), 0.96 (95% CI 0.93-0.99), 0.93 (95% CI 0.88-0.98) and 0.92 (95% CI 0.86-0.98), respectively. The probability of survival at 1, 2, 3, 4 and 5 years after the estimated date of seroconversion was 1.00 (95% CI 0.98-1.00), 0.99 (95%CI 0.98-1.00),

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0.98 (95% CI 0.95–1.00), 0.96 (95% CI 0.93–1.00) and 0.94 (95% CI 0.89–0.99), respectively.

#### Morbidity

During follow-up, 119 events included in the WHO staging system occurred in 62 patients, including 55 stage II defining events (38 patients), 44 stage III defining events (30 patients), 20 stage IV defining events (15 patients). We recorded 316 events (77 patients) that are not included in the WHO staging system. The incidence of WHO stage II, III and IV defining diseases was 6.1/ 100 PY, 4.7/100 PY and 2.2/100 PY, respectively. Diagnoses are detailed in Table 2. Overall, the most frequent diseases were mild non-classifying bacterial diseases (incidence 6.0/100 PY), malaria (3.6/100 PY), herpes zoster (3.4/ 100 PY), tuberculosis (2.1/100 PY) and severe WHO stage III or IV classifying bacterial diseases (3.1/100 PY). The probabilities of developing malaria, tuberculosis, and/or severe bacterial diseases within the first 5 years of inclusion in the cohort are shown in Fig. 2. Of the 14 episodes of tuberculosis, four led to death and the remaining 10 patients were considered cured according to international criteria. Only one of the 23 of episodes of severe bacterial diseases, and none of the 25 malaria episodes, were considered as a cause of death.

1.0 0.9 0.8-0.7 Probability 0.6-0.5 0.4-0.3 0.2 0.1-0.0 2 3 4 0 Time since enrolment (years)

Fig. 1. Survival in the ANRS 1220 cohort study of HIV-1 seroconverters, Abidjan, 1997–2004

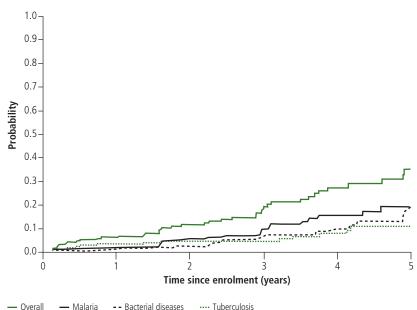
At the onset of the WHO classifying morbidity events, the median last available CD4 count was 319/mm<sup>3</sup> for WHO stage II (IQR 214–472; range 22–994), 243/mm<sup>3</sup> for WHO stage III (IQR 140–527, range 9–908) and 254/mm<sup>3</sup> for WHO stage IV (IQR 146–303, range 9–576). When considering specific diseases, the median last available CD4 count was 388/mm<sup>3</sup> for malaria (IQR 260–517), 303/ mm<sup>3</sup> for severe bacterial diseases (IQR 205–617), and 210/mm3 for tuberculosis (IQR 164–330). The median last available plasma HIV RNA was  $4.9 \log_{10}$ copies/mm<sup>3</sup> for WHO stage II (IQR 4.5-5.3, range 3.6-6.0),  $5.1 \log_{10}$  copies/ mm<sup>3</sup> for WHO stage III (IQR 4.4-5.9, range 2.3-6.6) and  $5.9 \log_{10}$  copies/mm3 for WHO stage IV defining events (IQR 5.0-6.4, range 4.2-6.6).

#### **Criteria for HAART initiation**

The 1-year, 2-year, 3-year, 4-year and 5-year probability of reaching criteria for starting HAART was estimated at 0.09 (95% CI 0.04-0.13), 0.16 (95% CI 0.10-0.22), 0.24 (0.17-0.31), 0.36 (0.28-0.45) and 0.44 (95% CI 0.35-0.55) following WHO 2003-2006 guidelines for settings with available CD4 count, 0.11 (95% CI 0.06-0.16), 0.23 (95% CI 0.16-0.29), 0.36 (95% CI 0.28-0.44), 0.49 (95% CI 0.39-0.58) and 0.56 (95% CI 0.46-0.66) following alternative early-initiation guidelines for settings with available CD4 count, and 0.06 (95% CI 0.03-0.10), 0.13 (95% CI 0.08-0.18), 0.19 (95% CI 0.13-0.26), 0.31 (95% CI 0.22-0.40) and 0.38 (95% CI 0.28-0.49) following WHO 2003-2006 guidelines for settings where CD4 counts were are not available, respectively. Fig. 3 shows the probability of reaching criteria for starting HAART over time from enrolment (overall and by criteria) in settings where CD4 measurement is not available, but total lymphocyte counts are possible.

Table 3 shows the first HAART initiation criterion that was met in our population when applying WHO

Fig. 2. Probability of developing tuberculosis, severe bacterial diseases, and/or malaria in the ANRS 1220 cohort study of HIV-1 seroconverters, Abidjan, 1997–2004



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#### Research Probability of starting HAART in Côte d'Ivoire

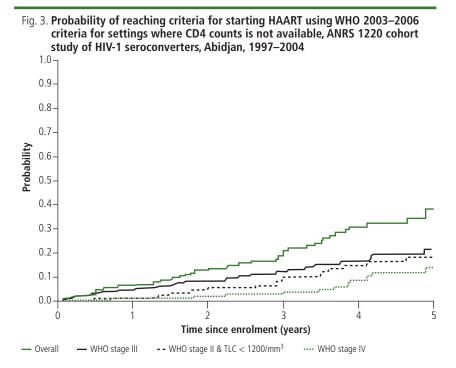
2003–2006 guidelines and alternative early-initiation criteria.

#### Discussion

When the WHO 2003-2006 guidelines for starting HAART in settings with available CD4 count were applied to our cohort of adult HIV-1 seroconverters in Abidjan,<sup>6,7</sup> we noted a probability of reaching criteria for HAART initiation of 9%, 16%, 24%, 36% and 44% at 1, 2, 3, 4 and 5 years, respectively. Unsurprisingly, these probabilities were 19-33% higher when applying alternative early-initiation criteria, which would include all patients to treatment who were at WHO stage III irrespective of CD4 count, and all patients at stage II with CD4 < 350/mm<sup>3</sup>. By contrast, when applying the WHO 2003-2006 guidelines for starting HAART in settings where CD4 counts are not available, the probability of starting HAART was 14-34% lower, which illustrates the difficulty in finding a suitable surrogate for CD4 counts.

Rates of survival, WHO stage IV, and CD4 counts < 200/mm<sup>3</sup> at 5 years' in our Ivorian population did not differ greatly from those reported in industrialized countries.<sup>8,9</sup> Our results do, however, contrast with findings from studies in the Central African Republic, Haiti, Kenya, Thailand, and Uganda that have suggested that median time from seroconversion to first HIV diseases was shorter in these settings than in industrialized countries.<sup>10–14</sup>

In Kenya, estimates of the median duration to CDC stage IV-A and IV-C



disease were 3.5 years and 4.4 years in a cohort of female sex workers.<sup>14</sup> In the Ugandan cohort, only 17% of patients remained symptom-free 6 years after seroconversion.<sup>12</sup> In the Thai cohort, median time to clinical AIDS and to a CD4 cell count < 200/mm<sup>3</sup> was 7.4 years and 6.9 years, respectively.<sup>11</sup> In the study from the Central African Republic, 35% of HIV-1 seroconverters died 3–5 years after seroconversion, and all died at AIDS stage.<sup>13</sup> In Haiti, median time to first HIV symptoms was 3 years and the median survival time to death was 7.4 years.<sup>10</sup> It is well known that there are differences between sub-Saharan Africa and industrialized countries with respect to HIV-morbidity. In African HIV-positive adults, tuberculosis and severe bacterial diseases are the most frequent causes of HIV morbidity.<sup>4,15</sup> They are also early diseases, which can occur in patients with CD4 counts greater than 200 cells/mm<sup>3</sup>. Malaria is also recognized as another HIV-related disease that can occur soon after seroconversion.<sup>16</sup> The early occurrence of both malaria and severe bacterial diseases is one of the

Table 3. Criteria for HAART initiation: first WHO criterion met in ANRS 1220 cohort study of HIV-1 seroconverters, Abidjan, 1997–2004

First criterion	CD4 count measurements <sup>a</sup>		
	Available		Not available
	WHO 2003 & 2006 guidelines <sup>6, 7</sup>	Alternative early- initiation criteria guidelines	WHO 2003 & 2006 guidelines <sup>6, 7</sup>
CD4 < 200/mm <sup>3</sup>	43 (67%) <sup>b</sup>	12 (21%) <sup>c</sup>	-
CD4 < 350/mm <sup>3</sup> & WHO stage III	13 (21%)	_	-
CD4 < 350/mm <sup>3</sup> & WHO stage II	-	21 (37%)	-
WHO stage IV	8 (12%)	6 (11%)	7 (14%)
WHO stage III	-	18 (31%)	25 (49%)
Total lymphocyte count < 1 200/mm <sup>3</sup> & WHO stage II	_	_	19 (37%)

<sup>a</sup> Data are number (%).

<sup>b</sup> Including three patients who were previously at stage III with CD4 count > 350/mm<sup>3</sup>.

<sup>c</sup> including five patients who were previously at stage II with CD4 count > 350/mm<sup>3</sup>.

main reasons that cotrimoxazole prophylaxis is recommended at an earlier stage in Africa than it is in Europe or in the USA.<sup>17</sup>

Since 1994, all HIV-infected adults attending the blood donor centre of Abidjan have been prescribed cotrimoxazole prophylaxis, irrespective of their CD4 count and of their clinical stage. Thus, all patients included in the Primo-CI cohort received cotrimoxazole, including those at WHO stage I and with CD4 > 500/mm<sup>3</sup>, and those included before the 1999 Côte d'Ivoire national consensus on cotrimoxazole. This decision to provide prophylaxis was made by staff at the blood donor medical centre, and although it might have seemed an over-reaction 10 years ago, it has recently been recognized as a valid option.<sup>18</sup> This prophylaxis may have contributed to a decrease in early morbidity in our study group and could explain the difference in morbidity between our cohort and those in other low-resource settings.

#### Limitations

First, for our data to be generalizable, our population should be representative of the overall population of recently HIVinfected adult in Abidjan. Unfortunately, there is no way to compare our selfselected population of urban voluntary blood donors with the overall population of recent seroconverters, as our cohort is the only source of data on recent HIV seroconverters in Abidjan.

Second, most of the data we present here shows probabilities of remaining event-free from the date of inclusion in the cohort, and not from seroconversion. However, our population included recent seroconverters with median time from seroconversion to enrolment estimated at about nine months. Analyses to estimate the probability of remaining event-free from the estimated date of seroconversion systematically gave probabilities that were slightly higher than those calculated using time since enrolment in the cohort. Because the real date of seroconversion was unknown and because the natural history of HIVinfection including the risk of very early mortality and severe morbidity within the first months of seroconversion has never been described in Côte d'Ivoire we chose to use time since inclusion rather than that since the estimated date of seroconversion in our analyses.

Third, we censored follow-up data at HAART initiation, with the potential of informative censoring in survival analyses and morbidity-free probability analyses.<sup>19</sup> On the other hand, this potential bias did not affect estimates of remaining free of criteria for HAART initiation, because all patients who started HAART in the cohort met the HAART starting criteria before their follow-up was censored.

In conclusion, our data provide valuable estimates of the probability of reaching the criteria for HAART initiation by time from first contact in a population of west African recent seroconverters. They support the hypothesis that median time to AIDS and survival in Africa do not differ from those in industrialized countries.20 However, our data also emphasize the significant incidence of the early morbidity in the first years following seroconversion in a west African urban population. This latter finding provides strong support in favour of more readily available HIV testing in this population so that HIV infection may be diagnosed and patients receiving care and follow-up as early as possible within the course of HIV disease.

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Competing interests: none declared.

#### Résumé

# Étude sur 7 ans et sur une cohorte d'adultes dont la séroconversion au VIH-1 est connue de l'évolution vers les critères de l'OMS concernant la mise en route d'un traitement antirétroviral (réalisée à Abidjan en Côte d'Ivoire)

Objectif Estimer la probabilité de remplir les critères de mise en route d'un traitement antirétroviral hautement actif (HAART) dans le cadre d'une étude prospective portant sur une cohorte d'adultes ayant subi une séroconversion au VIH-1, à Abidjan (Côte d'Ivoire). Méthodes Nous avons recruté des sujets parmi les donneurs positifs pour le VIH d'une banque de sang d'Abidjan pour lesquels le délai écoulé depuis la date estimée de la séroconversion (date située à mi-distance entre le dernier test négatif et le premier test positif pour le VIH-1) était  $\leq$  36 mois. Nous avons proposé à ces sujets un traitement prophylactique précoce par le triméthoprimesulfaméthoxazole (cotrimoxazole) et une numération deux fois par an des lymphocytes T CD4. Nous avons procédé à un enregistrement normalisé de la morbidité. Nous avons utilisé la méthode de Kaplan-Meier pour estimer la probabilité de remplir les critères de mise en route d'un traitement de type HAART selon les Directives OMS de 2006.

**Résultats** 217 adultes, dont 77 femmes (35 %), ont été suivis sur 668 personnes-ans. Les maladies les plus fréquemment relevées chez eux étaient des affections bactériennes bénignes (6,0/100 personnes-ans), des paludismes (3,6/100 personnes-ans), des zonas (3,4/100 personnes-ans), des affections bactériennes graves (3,1/100 personnes-ans) et des tuberculoses (2,1/100 personnes-ans). La probabilité de remplir les critères 2006 de l'OMS concernant la mise en route d'un traitement HAART a été estimée à 0,09 ; 0,16 ; 0,24 ; 0,36 et 0,44 respectivement pour les années 1, 2, 3, 4 et 5.

**Conclusion** Nos données font ressortir l'incidence de la morbidité précoce liée au VIH parmi une population adulte ivoirienne et confirment la nécessité de rendre plus accessible le dépistage de ce virus et de pratiquer un suivi précoce des adultes séropositifs en Afrique de l'Ouest.

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## Grado de cumplimiento de los criterios de la OMS para iniciar el tratamiento antirretroviral en adultos infectados de África Occidental: cohorte de siete años de seroconvertidos para el VIH-1

**Objetivo** Estimar la probabilidad de satisfacer los criterios para comenzar la terapia antirretroviral de gran actividad (TARGA) en una cohorte prospectiva de seroconvertidos para el VIH-1 en el banco de sangre de Abidján, África Occidental.

**Métodos** Se invitó a participar en la cohorte a todos los donantes de sangre en los que habían transcurrido  $\leq 36$  meses desde la fecha estimada de seroconversión (punto medio entre la última prueba negativa y la primera positiva para el VIH-1). El seguimiento incluyó la profilaxis temprana con cotrimoxazol, el recuento semestral de CD4 y la documentación normalizada de la morbilidad. Mediante el método de Kaplan–Meier calculamos la probabilidad de cumplir los criterios para iniciar la TARGA de acuerdo con las directrices de la OMS de 2006.

Resultados Se siguió la evolución de 217 adultos (35% de

mujeres) durante 668 años-persona. Las dolencias más frecuentes fueron enfermedades bacterianas leves (incidencia de 6,0 por 100 años-persona [/100 AP]), malaria (3,6/100 AP), herpes zóster (3,4/100 AP), enfermedades bacterianas graves (3,1/100 AP) y tuberculosis (2,1/100 AP). La probabilidad de cumplir los criterios de la OMS de 2006 para comenzar la TARGA se estimó en 0,09, 0,16, 0,24, 0,36 y 0,44 a los uno, dos, tres, cuatro y cinco años, respectivamente.

**Conclusión** Estos datos destacan la incidencia de morbilidad precoz por VIH en una población adulta de África Occidental y respaldan la idea de realizar un cribado temprano de la infección por VIH y un seguimiento temprano de los adultos infectados por el virus en esa parte de África.

#### ملخص

التقدم المحرز تجاه تلبية معايير منظمة الصحة العالمية التي توجب المعالجة مضادات الفيروسات القهقرية في مجموعة من البالغين الذين تحولوا من إيجابية المصل إلى سلبية المصل خلال سبع سنوات، في أبيدجان، كوت ديفوار

**الهدف:** استهدفت هذه الدراسة تقييم احتمالية الوصول إلى المعايير التي الموجودات: تمت متابعة 217 بالغاً (منهم 77 امرأة، أي بنسبة 35%) على تستوجب بدء المعالجة الفعالة بمضادات الفيروسات القهقرية في مجموعة مدى 668 شخص عام. وكانت الأمراض المسجلة الأكثر تكراراً هي أمراض من البالغين الذين تحولوا من إيجابية المصل إلى سلبية المصل خلال سبع جرثومية خفيفة (6 لكل 100 شخص عام)، والملاريا (3.6 لكل 100 شخص سنوات، في أبيدجان، كوت ديفوار.

> **الطريقة:** قمنا بتعيين المشاركين في الدراسة من بين الإيجابيين لفيروس الإيدز، المتبرعين بالدم في بنك الدم في أبيدجان، الذين كانت مدة تأخرهم عن تاريخ التحول من الإيجابية إلى السلبية (النقطة المتوسطة بين آخر اختبار سلبي وأول اختبار إيجابي لفيروس الإيدز من النمط 1) 36 شهراً أو أقل. وأعطي للمشاركين في الدراسة جرعات وقائية من الكوتر يوكسازول، مع تعداد الخلايا اللمفية CD4 مرتين في العام، كما أعددنا سجلات موحدة للمراضة، واتبعنا طريقة كابلان ماير لتقدير احتمالية تلبية المعايير التي توجب المعالجة الفعًالة بمضادات الفيروسات القهقرية وفقاً للدلائل الإرشادية لمنظمة الصحة العالمية لسنة 2006.

الموجودات: تمت متابعة 217 بالغاً (منهم 77 امرأة، أي بنسبة 35%) على مدى 668 شخص عام. وكانت الأمراض المسجلة الأكثر تكراراً هي أمراض جرثومية خفيفة (6 لكل 100 شخص عام)، والملاريا (3.6 لكل 100 شخص عام)، والهربس النطاقي (3.4 لكل 100 شخص عام)، وأمراض جرثومية وخيمة (3.1 لكل 100 شخص عام)، والسل (2.1 لكل 100 شخص عام). وتم تقدير احتمالية تلبية معايير منظمة الصحة العالمية لسنة 2006، التي تستوجب بدء المعالجة الفعّالة بمضادات الفيروسات القهقرية، على النحو التالي: 0.09 في العام الأول، 1.60 في العام الثاني، 204 في العام الثالث، 3.66 في العام الرابع، 0.44 في العام الخامس.

الاستنتاج: تؤكد معطياتنا وقوع إصابات مبكِّرة بفيروس الإيدز بين مجموعة من السكان البالغين في كوت ديفوار، وتدعم هذه البيانات ضرورة القيام باختبار فيروس الإيدز وأهمية المتابعة المبكِّرة للبالغين المصابين بالفيروس في غرب أفريقيا.

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