

Hepatitis B infection in People living with HIV/AIDS: A retrospective study of the Efia Nkwanta Regional Hospital

Abstract

Introduction: Comorbidities among people living with HIV/AIDS (PLWHA) increases with disease severity. This prevalence may be attributed to highly active antiretroviral therapy (HAART) toxicity and HIV/AIDS-related infections.

Aim: This study investigated the prevalence of comorbidities among PLWHA and their clinical and biochemical characteristics.

Methods: This study conducted at the Effia-Nkwanta Regional Hospital (ENRH) in the South-Western part of Ghana. A retrospective data of 500 participants (134 males and 366 females) were collected from HIV/AIDS patients on HAART (January 2012 to January 2016).

Sociodemographic characteristics and laboratory data of patients were retrieved from the laboratory database while clinical information was also retrieved from the patients' clinic files.

Data were analyzed with SPSS for both descriptive and inferential analysis.

Results: A total of 96 (19.2%) comorbidities were recorded (N=500). The most prevalent comorbidity was hepatitis B virus infection 33 (34.4%). Among the 96 HIV/AIDS patients who had comorbidities, 27 (28.1%) were males and 69 (71.9%) were females. The systolic blood pressure (SBP) of the HIV/AIDS patients with comorbidities was similar to that of those without comorbidities (113.84 ± 16.73 vs 115.32 ± 15.68). Majority of the participants with comorbidities 59 (61.5%) and those without comorbidities 227 (56.2%) were found to be on the same therapy combination (TDF+3TC+EFV). The decreased CD4 cell count, estimated glomerular filtration rate (eGFR), serum potassium and creatinine were similar in the participants (those with comorbidities and those without comorbidities). None of the demographics, clinical and biochemical parameters were associated with the presence of comorbidities.

Conclusion: The total prevalence of commodities was 19.2% and the prevalent commodity was HBV 33 (34.4%). The comorbidities were common among the females as well as married and old people living with HIV/AIDS. None of the comorbidity associated factors were found to be significant among HIV/AIDS patients on HAART with comorbidities.

Keywords: Comorbidities, HIV/AIDS, Demographic, Clinical, Biochemical.

343 **Introduction**

344 The Human Immunodeficiency Virus (HIV) has been of immense concern over the years and it
345 is reported to be associated with several communicable diseases and non-communicable
346 diseases (Aantjes, 2015). Comorbidities can be defined as the existence of additional distinct
347 disease entities during the clinical course of a patient who has the index disease under study
348 (Cahill &Valadéz, 2013). The index disease in this instance is HIV/AIDS and those infected
349 have been shown to develop comorbidities such as cardiovascular, renal, pulmonary, hepatic and
350 mental diseases as well as non-AIDS defining malignancies at an earlier age than the uninfected
351 (Justice et al., 2010). The HIV/AIDS infection itself greatly compromises immunity and pre-
352 existing chronic medical conditions could also be exacerbated contributing to the comorbidities
353 (Cahill &Valadéz, 2013). Furthermore, the toxicity of the antiretroviral drugs and the
354 interaction between the drugs for the management of the comorbidities and HAART contributes
355 to comorbidities in HIV/AIDS patients. Therefore, the presence of organ damage in patients
356 receiving antiretroviral treatment is not only the expression of treatment toxicity, but also a
357 complex interaction between individual risk factors, HIV/AIDS correlated effects, and
358 antiretroviral drug toxicity. As people living with HIV/AIDS (PLWHA) grow older, they also
359 become more susceptible to developing physical and mental diseases (Olisah, 2011). Individuals
360 with HIV/AIDS have higher prevalence of multimorbidity (Olisah, 2011) including
361 cardiovascular complications such as coronary artery disease, hypertension,
362 hypercholesterolemia, and diabetes (Rodriguez-Penney *et al.*, 2013) as well as cancer and
363 diseases of the liver, kidney, bone (e.g., osteopenia), and nervous system (Deeks& Phillips,
364 2009).

365 Opportunistic infections are claimed to be common among PLWHA who are highly susceptible
366 to various comorbidities in both developed and developing countries (Nduet *al.*, 2011).
367 However, the characteristics of PLWHA on HAART with comorbidity are not well described in
368 Ghana. This study sought to investigate the presence of comorbidities among PLWHA and
369 report their clinical and biochemical characteristics.

370

371 **Materials and method**

372 This was a retrospective hospital-based study conducted from January 2012 to December 2016
373 among HIV/AIDS patients who visited the HIV/AIDS clinic at the Effia-Nkwanta Regional
374 Hospital (ENRH) in the South-Western part of Ghana.

375 **Setting**

376 The hospital offers both general and specialist care services in internal medicine, general
377 surgery, paediatrics, obstetrics and gynaecology, dental and eye care and serves as the main
378 referral facility for the western parts of the country. The hospital admits over 7500 –10,000
379 patients annually.

380 **Study population**

381 The study retrospectively sampled the records of a total number of 500 HIV/AIDS patients
382 receiving HAART in ENRH. Due to the completeness of data within the study period, 134
383 males and 366 females with HIV/AIDS at the HIV/AIDS clinic of the Effia Nkwanta Teaching
384 Hospital were recruited.

385 **Ethical considerations**

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387 The study was approved by the Institutional Review Board of the University of Cape-Coast
388 (IRB-UCC) and the authorities of Effia-Nkwanta Regional Hospital for approval. Besides, all
389 data were anonymized before analyzed.

390 **Inclusion and exclusion criteria**

391 The study included HIV/AIDS patients on HAART and excluded HAART naïve HIV/AIDS
392 patients. Also, HIV/AIDS patients whose folders did not contain adequate information within
393 the stipulated period for the study were excluded.

394 **Collection of data**

395 Retrospective data of 500 HIV/AIDS patients on HAART (134 males and 366 females) were
396 retrieved from the laboratory database and hospital folders. Data of the participants from
397 January
398 2012 to December 2016 were included in this study. Demographic and laboratory data
399 (biochemical and serological findings) of patients were retrieved from the laboratory database.
400 Also, past medical history, family history, social class and clinical examination information
401 were retrieved from the patients' clinic files.

402 **Statistical Analysis**

403 Data were analyzed with SPSS version 16 (SPSS Inc. Chicago). Descriptive statistics were
404 computed with standard methods and were presented as mean and standard deviations (SD).
405 Chi-square test was used to compare the association between categorical variables and
406 independent t-test was used to compare the mean value of some laboratory parameters and
407 socio-demographics. One-way ANOVA was also employed to compare the mean scores of more
408 than two groups and $P < 0.05$ was interpreted as statistically significant.

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410 **Results**

411
412 The data shows that males were older than females ($P = 0.004$) (Table 1). Majority of the
413 participants were married 257 (51.1%) and had been with the condition [400 (80.0%)] and on
414 medication [403 (80.6%) for less than 5 years. SBP ($P = 0.358$) and DBP ($P = 0.882$) were
415 similar among the participants. Majority of the HIV/AIDS patients 292 (58.4%) had normal
416 BMI, 133 (26.6%) were underweight, 56 (11.2%) were overweight and 19 (3.8%) were obese.

417 A total of 96 (19.2%) comorbidities were recorded among the participants. The most prevalent
418 comorbidities were hepatitis B virus infection 33 (34.4%), arthralgia 7 (7.3%), sickle cell
419 disease (SCD) 6 (6.3%), diabetes 6 (6.3%), jaundice 6 (6.3%), chronic diarrhea 5 (5.2%) and
420 visual changes 5 (5.2%) while tuberculosis (TB), insomnia, Kaposi sarcoma, pneumonia, skin
421 rash, slow mentation, anaemia, amnesia and paresthesia were the lowest comorbidities (Table 2).

422 Among the 96 HIV/AIDS patients who had comorbidities, 27 (28.1%) were males and 69
423 (71.9%) were females. A higher proportion of the participants with comorbidities were found
424 within the age group 30 - 39 32 (33.3%) and the majority of them were also married 48 (50.0%).

425 An equal number of HIV/AIDS patients with comorbidities 79 (82.3%) have had HIV/AIDS and
426 had also been on medications for less than 5 years. The SBP of the HIV/AIDS patients with
427 comorbidities was similar to that of those without comorbidities (113.84 ± 16.73 vs $115.32 \pm$
428 15.68). Most of the HIV/AIDS patients with comorbidities had normal BMI (53.1%) and a
429 smaller number of them were obese 5 (5.2%). Also, the majority of the participants with
430 comorbidities 59 (61.5%) and those without comorbidities 227 (56.2%) were found to be on the
431 same therapy combination (TDF+3TC+EFV) (Table 3).

432 None of the urinalysis parameters was found to be significant among HIV/AIDS patients on
433 HAART with and without comorbidities (Table 4). The decreased CD4 cell count, eGFR,
434 potassium and creatinine in patients with comorbidities than those without comorbidities were
435 not significant (Table 5). None of the demographics, clinical and biochemical parameters was
436 associated with the presence of comorbidities (Table 6).

437 **Discussion**

438 HIV/AIDS greatly compromises immunity to make the body susceptible to other diseases. Pre-
439 existing chronic medical conditions are also exacerbated by HIV/AIDS infection. This study
440 sought to investigate the presence of comorbidities among PLWHA and report their clinical and
441 biochemical characteristics. The commonest comorbidity was hepatitis B virus infection 33
442 (34.4%) and the comorbidities had female dominance 69 (71.9%) over the male population 27
443 (28.1%). The modal age range among the HIV/AIDS patients was 30-39 years (35.8%) which is
444 similar to a cross-sectional descriptive study conducted by Nduet *al.*, (2011) in Nigeria, who
445 reported a modal age range of 31- 40 years (38.7%). Again, in this study, 400 (80%) HIV/AIDS
446 infected person was found to have been infected for a duration more than 5 years as at the time
447 of the study. However, an Institution based cross-sectional study by Tesfawet *al.*, (2016) in
448 Ethiopia showed that 390 out of 417 (93%) of the participants had acquired the virus over 2
449 years as at the time of their study.

450 The present study recorded the highest commodities among the HIV/AIDS married group while
451 that of Nduet *al.*, (2011) recorded the highest comorbidities among the divorced group.

452 However, their study was done among all HIV/AIDS patients, irrespective of their employment
453 status while the former study was done among HIV/AIDS workers only attending infectious
454 diseases clinic. Ideally, marriage provides economic and social stability necessary for good

455 health Ross *et al.*, (1990). Therefore, the high prevalence of comorbidities among married
456 population in this study could have resulted from the inclusion of the non- working for married
457 HIV/AIDS population. On the contrary, a divorce which is common among HIV/AIDS
458 discordant couples provides a fertile ground for the development of medical comorbidities as
459 seen in present study (Porter *et al.*, 2004).

460 The prevalence of HIV/AIDS patients on HAART with comorbidities in this study was 19.2%.
461 Majority of the HIV/AIDS patients on HAART had HBV infection 33 (34.4%) infection, whilst
462 anaemia 1 (1%) and amnesia 1 (1%) were the less common comorbidities. These findings are at
463 variance with the previous studies conducted by Nduet *et al.*, (2011), Schouten *et al.*, (2014) and
464 Hareguet *et al.*, (2012). A cross-sectional descriptive study conducted by Nduet *et al.*, (2011), among
465 489 HIV/AIDS positive workers attending HIV clinics in Enugu (Nigeria) revealed that, 53
466 (44.5%), 44 (37%), 9 (7.9%), 5% had hypertension, arthritis, diabetes mellitus and HBV
467 infection as comorbidities respectively. A systematic review of 37 studies by Hareguet
468 *et al.*,(2012) reported on the magnitude and determinants of non-communicable diseases in 30,000
469 PLWHA. They reported the highest prevalence comorbidity in HIV/AIDS patients to be
470 cardiovascular diseases. The difference between the findings in this study and the review study
471 by Hareguet *et al.*, (2012) could be attributed to small sample size (n=500 in this study) against
472 30,000 in the previous study. Also, a cross-sectional study by Schouten *et al.*, (2014) reported a
473 lower prevalence of HBV (3.5%) comorbidity in HIV/AIDS patients in the Netherlands.

474 Again, a prospective Swiss cohort study conducted by Greudet *et al.*, (2000) among 3111 HIV-
475 infected patients reported a higher prevalence of hepatitis C virus infection 1157 (37.2%).
476 Conversely, our study showed a lower prevalence of HCV (2.1%) among the HIV/AIDS
477 patients.

478 Moreover, the prevalence of Kaposi's sarcoma (2.1%) in our study was lower than that of
479 Beralet *et al.*, (1990) in the United States America. The latter study revealed that among persons,
480 with HIV/AIDS, the prevalence of Kaposi's sarcoma was 15% (13 616). On the other hand, the
481 prevalence of tuberculosis recorded in this study (5.7%) was again lower than the previous
482 findings by Tesfawet *et al.*, (2016), (12%) in Ethiopia. The lower prevalence of HCV, Kaposi
483 sarcoma and tuberculosis observed in our study could be associated with retrospective nature of
484 the study, the sample size and the study setting.

485 Our investigation in this study showed co-existence of diabetes and HIV infection. The
486 prevalence of diabetes among the HIV/AIDS patients on HAART was 6 (6.3%). This finding is
487 not consistent with a large prospective cohort study conducted by De Wit *et al.*, (2008) among
488 33,389 HIV positive patients followed at 212 clinics in Europe, the U.S., Argentina, and
489 Australia. In their study, the prevalence of diabetes among HIV/AIDS patients was 952(2.85%).
490 The difference in the prevalence between these two studies could be due the sample size as well
491 as the geographical locations. Prevalence of hepatitis B in people living with HIV/AIDS in
492 Latin America and the Caribbean was also reported by Tengan *et al.* (2017). HBV and HCV Co-
493 infection among HIV/AIDS Patients in the National Hospital of Tropical Diseases, Vietnam was
494 endorsed by Huy *et al.* (2014). A greater proportion of 39 (40.6%) of the HIV/AIDS patients on
495 HAART with comorbidities in this study had a CD4 count level 200 – 499 cell/mm³ which is in
496 contrast with the findings from a cross-sectional study by Schouten *et al.*, (2014) among 540
497 HIV/AIDS patients in Netherlands. This study recorded only 1 (1%) of proteinuria among the
498 HIV/AIDS patients on HAART with comorbidities. This contrasts with previous findings by
499 Dondoet *et al.*, (2013) in Zimbabwe, Galgallo, (2006) in Kenya, Ekuluet *et al.*, (2012) in Congo and

500 Esezoboret *al.*, (2010) in Nigeria who recorded the prevalence of proteinuria to be 16.4%, 30%,
501 23.8% and 20.5% respectively.

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503 The high prevalence of proteinuria could be due to the recruitment of participants with advanced
504 HIV/AIDS as shown by the CD4 count.

505 Our study also showed that the majority of the HIV/AIDS patients 59 (61.5%) with
506 comorbidities used the drug type TDF+3TC+EFV. Markers for renal impairment such as serum
507 urea, creatinine, potassium and sodium showed no significant association between HIV/AIDS
508 patients on HAART with and without comorbidities. None of the HIV/AIDS patients had renal
509 dysfunction in this present study. These findings are contrary to a cohort study conducted by
510 Crum-Cianflone *et al.*, (2010) in California who reported a prevalence of 22 (3%) of renal
511 dysfunction among 717 HIV/AIDS patients on HAART. According to the latter study, the
512 occurrence of the renal dysfunction was associated with duration of tenofovir use.

513 Our study, however, has two major limitations: first, it could not formally tell if the commodities
514 were acquired either before or after the acquisition of the index disease. Finally, the use of a
515 single centre, a retrospective design, limited descriptive information of participants, bias and
516 confounding in our findings will also limit the scope of the outcomes.

517 **Conclusion**

518 In conclusion, the prevalent comorbidity was hepatitis B virus infection. The comorbidities were
519 more common among the females than the males as well as married and old people living with
520 HIV/AIDS. Early and regular screening remains to be the key prevention and control strategy
521 for the HIV/AIDS-associated commodities. The findings warrant coordination of HIV/AIDS and

522 its related commodities in Ghana. A prospective cohort study should consider the extensive
523 evaluation of personal lifestyle factors that contribute to the development of comorbidities in
524 PLWHA

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605 **Table 1: Demographic and clinical characteristics of HIV/AIDS patients on HAART**

Characteristics	Male (n = 134)	Female (n = 366)	Total (n = 500)	P-value
Age (years)	39.81 ± 10.97	36.52 ± 11.31	37.40 ± 11.30	0.004
Age group n (%)				0.001
< 20	1 (12.5)	7 (87.5)	8 (1.6)	
20-29	28 (22.2)	98 (77.8)	126 (25.2)	
30-39	35 (19.6)	144 (80.4)	179 (35.8)	
40-49	43 (39.8)	65 (60.2)	108 (21.6)	
50-59	22 (39.3)	34 (60.7)	56 (11.2)	
≥ 60	5 (21.7)	18 (78.3)	23(4.6)	
Marital status				0.860
Single	39 (26.2)	110 (73.8)	149 (29.8)	
Cohabiting	3 (27.3)	8 (72.7)	11 (2.2)	
Married	70 (27.2)	187 (72.8)	257 (51.1)	
Separated	1 (50.0)	1 (50.0)	2 (0.4)	
Divorced	13 (22.4)	45 (77.6)	58 (11.6)	
Widowed	8 (34.8)	15 (65.2)	23 (4.6)	
Duration of condition				0.013
< 5 years	117 (29.2)	283 (70.8)	400 (80.0)	
≥ 5years	17 (17.0)	83 (83.0)	100 (20.0)	
Duration on medication				0.022
< 5 years	117 (29.2)	286 (71.0)	403(80.6)	
≥ 5years	17 (17.5)	80 (82.5)	97 (19.4)	
Blood pressure (mmHg)				
SBP	113.84 ± 16.73	115.33 ± 15.68	114.93 ± 15.96	0.358
DBP	73.59 ± 10.99	73.78 ± 12.57	73.73 ± 12.16	0.882
BMI (Kg/m²)	19.75 ± 2.40	20.90 ± 4.07	20.60 ± 3.73	0.002
BMI n (%)				0.001
Underweight	36 (27.1)	97 (72.9)	133 (26.6)	
Normal	92 (31.5)	200 (68.5)	292 (58.4)	
Overweight	5 (8.9)	50 (89.3)	56 (11.2)	
Obese	1(5.2)	19 (100)	19 (3.8)	

606 **SBP=Systolic Blood Pressure, DBP=Diastolic Blood Pressure, BMI=Body Mass Index**

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609 **Table 2: Prevalence of comorbidities among HIV/AIDS patients on HAART**

Comorbidities	Frequency	Percentage (%)
Presence of Comorbidities		
Yes	96	19.2
No	404	80.8
Comorbidities		
Hepatitis B virus (HBV)	33	34.4
Hepatitis C virus (HCV)	2	2.1
Herpes zoster	2	2.1
Chronic diarrhea	5	5.2
Diabetes	6	6.3
Sickle Cell Disease (SCD)	6	6.3
STI	3	3.1
Syphilis	3	3.1
TB	4	4.2
Insomnia	3	3.1
Jaundice	6	6.3
Kaposi sarcoma	2	2.1
Pneumonia	2	2.1
Skin rash	3	3.1
Slow mentation	2	2.1
Anaemia	1	1.0
Amnesia	1	1.0
Arthralgia	7	7.3
Paresthesia	3	3.1
Visual Changes	5	5.2

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623 **Table 3: Demographic association with Comorbidities among HIV/AIDS patients on**
 624 **HAART**

Characteristics	Presence of comorbidities		P-value
	Yes (n = 96)	(n = 404)	
Gender			0.744
Male	27 (28.1)	107 (26.5)	
Female	69 (71.9)	297 (73.5)	
Age group n (%)			0.902
< 20	1 (1.0)	7 (1.7)	
20-29	22 (22.9)	104 (25.7)	
30-39	32 (33.3)	147 (36.4)	
40-49	24 (25.0)	84 (20.8)	
50-59	12 (12.5)	44 (10.9)	
≥ 60	5 (5.2)	18 (4.5)	
Marital status			0.002
Single	27 (28.1)	122 (30.2)	
Cohabiting	1 (1.0)	10 (2.5)	
Married	48 (50.0)	209 (51.7)	
Separated	2 (2.1)	0 (0.0)	
Divorced	8 (8.3)	50 (12.4)	
Widowed	10 (10.4)	13 (3.2)	
Duration of condition			0.532
< 5 years	79 (82.3)	321 (79.5)	
≥ 5years	17 (17.7)	83 (20.5)	
Duration on medication			0.641
< 5 years	79 (82.3)	324 (80.2)	
≥ 5years	17 (17.7)	80 (19.8)	
Blood pressure (mmHg)			
SBP	113.84 ± 16.73	115.32 ± 15.68	0.358
DBP	73.59 ± 10.99	73.78 ± 12.57	0.882
BMI n (%)			0.351
Underweight	25 (26.0)	108 (26.7)	
Normal	51 (53.1)	241 (59.7)	
Overweight	15 (15.6)	41 (10.1)	
Obese	5 (5.2)	14 (3.5)	
Type of drug			0.875
AZT+3TC+EFV	27 (28.1)	129 (31.9)	
AZT+3TC+NVP	8 (8.3)	32 (7.9)	
d4T+3TC+EFV	0 (0.0)	3 (0.7)	
TDF+3TC+EFV	59 (61.5)	227 (56.2)	
TDF+3TC+NVP	1 (1.0)	6 (1.5)	
SEPTRIN, VITAFOL	1 (1.0)	7 (1.7)	

626 **Table 4: Urinalysis of HIV/AIDS patients on HAART with comorbidities**

Parameter	Presence of comorbidities	
	Yes (n = 96)	(n = 404)
Protein		
Positive	1 (1.0)	3 (0.7)
Negative	95 (99.0)	401 (99.3)
Glucose		
Positive	0 (0.0)	0 (0.0)
Negative	96 (100)	404 (100)
Presence of Pus cells		
Yes	3 (3.1)	6 (1.5)
No	93 (96.9)	398 (98.5)
Presence of RBCs		
Yes	0 (0.0)	3 (0.7)
No	96 (100)	401 (99.3)
Presence of EC		
Yes	3 (3.1)	6 (1.5)
No	93 (96.9)	398 (98.5)
Presence of cast		
Yes	0 (0.0)	1 (0.2)
No	96 (100)	403 (99.8)
Presence of crystals		
Yes	0 (0.0)	1 (0.2)
No	96 (100)	403 (99.8)

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637 **Table 5: CD4 count and renal function among HIV/AIDS patients on HAART with**
 638 **comorbidities**

Parameter	Presence of comorbidities		P-value
	Yes (n = 96)	No (n = 404)	
CD4 Count cell/mm³	350.64 ± 253.58	382.40 ± 281.07	0.311
CD4 n (%)			0.795
< 200	32 (33.3)	126 (31.2)	
200-499	39 (40.6)	159 (39.4)	
≥ 500	25 (26.0)	119 (29.5)	
Sodium (mmol/L)	139.04 ± 2.12	137.97 ± 15.51	0.723
Potassium (mmol/L)	3.97 ± 0.52	5.29 ± 1.11	0.510
Urea (mmol/L)	7.08 ± 1.98	6.00 ± 0.54	0.459
Creatinine (µmol/L)	78.55 ± 23.58	117.06 ± 10.02	0.061
eGFR mL/min/1.73 m²	93.45 ± 40.29	94.72 ± 37.83	0.856
eGFR n (%)			0.99
≥ 60	31 (86.1)	154 (86.0)	
< 60	5 (13.9)	25 (14.0)	

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UNDER PEER REVIEW

652 **Table 6: Comorbidity associated factors among patients with HIV/AIDS on HAART**

Variables	OR (95%CI)	P-value
Gender		
Male*	1	
Female	0.92 (0.56-1.51)	0.744
Age group n (%)		
< 20*		
20-29	1.48 (0.17-12.65)	0.720
30-39	1.52 (0.18-12.82)	0.698
40-49	2.00 (0.23-17.06)	0.526
50-59	1.91 (0.21-17.06)	0.563
≥ 60	1.94 (0.19-19.74)	0.574
Duration of condition		
< 5 years	1.20 (0.68-2.14)	0.533
≥ 5years*	1	
Duration on medication		
< 5 years	1.15 (0.64-2.05)	0.641
≥ 5years*	1	
BMI n (%)		
Underweight	1.09 (0.64-1.86)	0.740
Normal*	1	
Overweight	1.73 (0.89-3.34)	0.106
Obese	1.69 (0.58-4.90)	0.335
CD4 n (%)		
< 200	1.21 (0.68-2.16)	0.522
200-499	1.17 (0.67-2.04)	0.585
≥ 500*	1	
eGFR		
≥ 60*	1	
< 60	0.42 (0.12-1.44)	0.167

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