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1 2	Disability, Social Functioning and School Inclusion Among Older Children and Adolescents Living with HIV In Zimbabwe
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21	Short heading: Disability and social functioning in HIV-infected children
22	Key words: disability, adolescents, children, Africa, HIV, social functioning
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#### 24 Abstract

25 Objective Increasing numbers of children with HIV are surviving to adolescence and encountering multiple clinical and social consequences of longstanding HIV infection. We 26 aimed to investigate the association between HIV and disability, social functioning and school 27 28 inclusion among 6 to 16-year olds in Zimbabwe. 29 Methods HIV-infected children receiving antiretroviral therapy from a public-sector HIV 30 clinic, and HIV-uninfected children attending primary care clinics in the same catchment area 31 were recruited. Standardised questionnaires were used to collect sociodemographic, social 32 functioning and disability data. Multivariable logistic regression was used to assess the relationship between HIV status and disability and social functioning. 33 34 Results We recruited 202 HIV-infected and 285 HIV-uninfected children. There was no 35 difference in age and gender between the two groups, but a higher proportion of HIV-infected 36 children were orphaned. The prevalence of any disability was higher in HIV-infected than 37 uninfected children (37.6% vs. 18.5%, p<0.001). HIV-infected children were more likely to 38 report anxiety (adjusted odds ratio (aOR) 4.4; 95% CI 2.4, 8.1), low mood (aOR 4.2; 2.1, 8.4) 39 and difficulty forming friendships (aOR 14.8; 1.9, 116.6) than uninfected children. Children 40 with HIV also reported more missed school days, repeating a school year and social exclusion 41 in class. These associations remained apparent when comparing children with HIV and 42 disability to those with HIV but no disabilities. 43 Conclusions Children with HIV commonly experience disabilities, and these are associated with social and educational exclusion. Rehabilitation and support services are needed to 44 45 facilitate educational attainment and social participation in this population.

### Introduction

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47 In 2016, worldwide approximately 160,000 children were newly infected with HIV (1). Of the 48 estimated 2.1 million children aged under 15 years living with HIV, nearly 90% live in Sub-49 Saharan Africa (SSA) (1). The global scale-up of antiretroviral therapy (ART) programmes 50 has meant that increasing numbers of children with HIV who would previously have died in 51 infancy without treatment are now surviving to older childhood and adolescence. However, 52 there is increasing evidence that childhood HIV infection is associated with chronic multi-53 system complications, resulting in hearing, cognitive, mobility and visual impairments (2, 3). 54 HIV may lead to impairments through a variety of mechanisms. For example, HIV-mediated immunosuppression may lead to opportunistic infections such as CMV that can cause visual 55 56 impairment (4). The risk of impairments is increased if initiation of ART is delayed, as is 57 common in many resource-limited settings (5). ART itself may also contribute to impairment; 58 for instance, nucleoside analogue reverse-transcriptase inhibitors (NRTI) commonly used at 59 the time of ART roll out for children in SSA (e.g. stavudine and lamuvidine) is linked to hearing 60 loss (6, 7). Zidovudine has been independently linked to myopathy (8), which may lead to 61 physical impairments. Once established, impairments may not be completely reversed by ART 62 (9) and negatively impact on social functioning and schooling (3, 10). In other words, HIV or 63 its treatment may lead to disability, which is defined as the restriction of participation in society 64 of an individual due to an underlying impairment in combination with attitudinal and 65 environmental and other barriers (11). Socio-economic deprivation, often associated with HIV 66 infection (12), potentially exacerbates disability by further restricting participation in society. 67 To optimise the quality of life and long-term care amongst those living with HIV and their 68 families, HIV programmes need to broaden their focus and address longer-term consequences 69 of HIV infection, including the impact on schooling and social inclusion. Even in the absence 70 of HIV, education and schooling are a major global concern for children and adolescents with disabilities, who are substantially less likely to be enrolled in school and, even when enrolled, lag behind their peers in educational attainment (13). HIV is likely to magnify these issues among children due to poverty resulting from parental ill health, food insecurity and unemployment (14).

We therefore conducted a cross-sectional study to investigate the association between HIV and

disability, social functioning and school inclusion among HIV-infected children compared to uninfected peers in Zimbabwe.

### Methods

79 Study setting and participants

HIV-infected children aged 6 to 16 years and receiving either first or second line ART for at least six months were consecutively recruited from Harare Central Hospital (HCH); this is the largest public-sector hospital in Harare, providing HIV care for more than 3,000 children. This age range was selected because it represents children of school going age. Recruitment was restricted to the first five eligible participants a day for logistical ease. Exclusion criteria were being acutely ill i.e. having a respiratory tract or other acute infection or tuberculosis, not residing in Harare and no guardian consent and/or participant assent.

A comparison group of HIV-uninfected children aged 6-16 years was recruited from primary health care clinics (PHC) in seven high-density communities from the same catchment area served by the clinic from which the HIV-infected participants were enrolled. Provider initiated HIV testing and counselling was offered by the PHCs to all children attending for acute care regardless of the reason for presentation, and those who tested HIV-negative were invited to participate and attend pre-booked appointments for assessments. The same exclusion criteria were applied to HIV-uninfected children.

#### Data collection

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Socio-demographic data including age, sex and orphan status were recorded. Trained research nurses administered standardised questionnaires to collect data on disability, education and social functioning. The Washington Group/UNICEF Child Functioning and Disability 21 Question Set was administered jointly to all children and caregivers by a research nurse to assess disability (15). This question set is validated for children aged 2-17 years. Self-reported functional difficulties were defined as binary variables in the following domains: vision, hearing, walking, speech, learning, memory, self-care, anxiety, low mood, difficulty controlling behaviour, dealing with change, forming friendships and concentration. Disability was defined as reported difficulties in any of the functional domains. Additional information on school and social functioning was collected, including the following: school enrolment, school attendance, repeated school year, problems getting help from teachers and friends, interaction with other children (leadership, play, bullying) and inclusion in lessons and school activities. Caregivers of HIV-infected children were asked additional questions relating to HIV diagnosis, testing, ART history, and children's awareness of diagnosis. At the time of enrolment, CD4 count was determined using an Alere PIMA CD4<sup>+</sup> (Waltham, Massachusetts, USA) and HIV viral load was measured using COBAS Ampliprep/Taqman 48 Version 2.0 (Roche, Rotkreuz, Switzerland).

## 113 Ethics

Ethical approval was obtained from the Medical Research Council of Zimbabwe (MRCZ/A/1856), the Biomedical Research and Training Institute (AP125) Institutional Review Board, Harare Hospital Ethics Committee and the London School of Hygiene and Tropical Medicine (LSHTM) Ethics Committee (8263). All guardians gave written consent, and participants gave assent to participate in the study.

Data management and analysis

Data were collected using paper forms and entered into a Microsoft Access database using optical mark recognition software (Cardiff TELEFORM Intelligent Character, Version 10.7), which has inbuilt quality control checks. Paper forms were manually checked for missing data and inconsistencies before being captured. Further internal and external consistency checks were carried out using database queries.

Data completeness was assessed by summary and descriptive statistics. There was a low proportion of missing data (<6%) in HIV-infected and uninfected children for demographic, clinical, disability and school functioning and social inclusion data. The prevalence of functional difficulties and disability was summarised as frequencies and percentages for each variable by HIV status. Continuous variables were summarised as mean and standard deviation (SD) when normally distributed and median and interquartile range (IQR) when not. Univariable logistic regression analysis was used to compare functional, school and social outcomes between HIV-infected and uninfected children. Multivariable logistic regression was used to adjust each functional outcome of interest for a priori defined variables of age and sex. Orphan status and previous infection/co-morbidity did not significantly affect the fit of the model (p<0.05) on likelihood ratio testing and therefore were excluded. Hence, the final model was adjusted for age and sex alone. All statistical analyses were carried out using Stata v13.0 (College Station, Texas: StataCorp LP).

# **Results**

Baseline characteristics of participants

We recruited 202 HIV-infected children (median age 11 years [IQR 8-13]; 48.0% female) and 285 uninfected children (median age 10 years [IQR 8-13]; 48.8% female). There were no significant differences in age or sex between the two groups, but HIV-infected children were

more likely to be orphaned (p<0.001) (Table 1). Among HIV-infected children, the median age at HIV diagnosis was 5 years [IQR 3-7] and the median CD4 count was 726 cells/  $\mu$ l [IQR 476-941]. The median duration of ART was 2 years [IQR 1-5] and the median age of ART initiation was 8 years [IQR 5-10].

Functioning and disability

The prevalence of any self-reported difficulties in functioning (*i.e.* disability) was higher in HIV-infected children compared to uninfected children (37.6% compared to 18.8% p<0.001) (Table 2). Amongst those with HIV, the most common types of disability were learning (reported by 23.2%) and memory difficulties (reported by 17.8%). Difficulties with seeing (7.7%), hearing (4.8%) and walking (2.5%) were also reported more commonly amongst HIV infected children.

After adjustment for age and sex, the odds of any disability were 2.8 times higher in HIV-infected than HIV-uninfected children (95% CI 1.8, 4.2 p<0.001). HIV-infected children were significantly more likely to report visual (aOR 3.0; 1.3, 6.9), hearing (aOR 3.4; 1.0, 10.5), speech (aOR 3.8; 1.1, 13.9), learning (aOR 3.9; 1.4, 3.4) and memory problems (aOR 3.5; 2.0, 6.6) (Table 2). In addition, HIV-infected children were more likely to report anxiety (aOR 4.4; 2.4, 8.1), low mood (aOR 4.2; 2.1, 8.4) and difficulty forming friendships (aOR 14.8; 1.9, 116.6) compared to their uninfected peers. There was no significant association between age at HIV diagnosis, age of ART initiation, CD4 count, viral load, ART duration or previous comorbidity and disability among HIV-infected children (Table 3).

Schooling and social inclusion

School enrolment rates were high among all children (96.0% in both HIV-infected and uninfected groups). However, children living with HIV were more likely to have repeated a school year (aOR 3.2; 1.6, 3.8) and on average, missed more days of school in the preceding month (mean 0.9 days (range 0-15 days) vs. 0.3 days (range 0-7 days). HIV-infected children more frequently reported not receiving help from teachers (aOR 2.1; 1.2, 3.8) or friends (aOR 3.0; 2.0, 4.5) at school. They were more likely to feel excluded in lessons and activities (aOR 4.7; 2.7, 8.3) and more likely to be physically and verbally bullied by other children (aOR 3.7; 2.2, 6.0). Among children with HIV, those with disabilities were less likely to be enrolled in the same school grade as their age peers (aOR 3.3; 1.7, 6.1) and more likely to repeat a school year (aOR 1.9; 1.0, 3.6) compared to HIV-infected peers without disability. They were also more likely to report that their peers did not look up to them as leaders (aOR 2.1; 1.4, 3.4) and that they experienced violence from their peers (aOR 2.5; 1.3, 4.8) (Table 4). Amongst children with disability, those with HIV were less likely to be enrolled in school, more likely to have needed to repeat a school year and much more likely to have been physically or verbally bullied than disabled children without HIV (Supplementary Table 5).

# **Discussion**

This study demonstrates a high prevalence of physical and cognitive functional difficulties among HIV-infected children compared to their uninfected peers. Children with HIV were more likely to report low mood, anxiety, difficulty forming friendships, repeating a school year and to experience poor social support at school, particularly when HIV and disability coexisted.

Other studies have reported increased physical, sensory and cognitive difficulties in HIV-infected children compared to those uninfected (3,16-22). Developmental delay is strongly

associated with HIV in SSA (2), affecting up to 78% of children (22). Fortunately, in the post-ART era, severe forms of cognitive impairment in children appear to be decreasing; however, the prevalence of mild impairment remains largely unchanged and may even be increasing (23). A number of studies have assessed the prevalence of cognitive (18, 22, 24, 26-29) and motor (18, 19, 21, 23-29) impairments among HIV-infected and uninfected children; however, to date these have largely focused on infants and younger children before school age. Our study highlights both the increased prevalence of learning difficulties among HIV-infected schoolage children, but also shows that learning difficulties are common in uninfected children in Zimbabwe.

This study further demonstrates the additional burden of low mood and anxiety amongst HIV-infected children. There is evidence of a strong bidirectional association between mental health and educational attainment with mood and anxiety disorders having a direct effect on early school leaving, substance misuse and disruptive behavioural disorders (30). Mental health issues impact negatively on treatment compliance and retention in social care and school through the fear of disclosing HIV status and social ostracism (31). Socialising and making friends at school are key protective factors for psychosocial wellbeing in children with HIV, whereas negative peer interactions such as lack of friends, bullying and being beaten by friends have been identified as risks (32). Therefore, school peer support interventions should be adopted as they have been shown to reduce psychological distress, depression, anxiety and anger in children with HIV (33, 34).

Similar to our findings, a recent Malawian cross-sectional study found that a high proportion of HIV-infected school children had hearing impairment identified by extensive audiological testing (10). These children were less likely to attend school and had poorer emotional and school functioning than HIV-infected children without hearing loss. Furthermore, only 40% of

214 caregivers accurately perceived their child's hearing loss, and few had sought treatment, 215 implying that routine screening may be necessary as disability may be underreported (10). Our study found no significant association between HIV disease severity or treatment factors 216 217 and disability. However, previous studies have shown a relationship between CD4 count, viral 218 load at enrolment, ART duration and disability (2,10). The Malawian study mentioned above 219 (10) found hearing loss to be significantly associated with HIV WHO Stage 3 or 4 disease, but 220 not duration of ART or CD4 count. A recent systematic review of disability and HIV in SSA 221 found a significant dose-response relationship between indicators of disease progression (CD4 222 or WHO stage) and disability in 48% of studies (2). The evidence suggests that earlier ART 223 initiation in children may reduce the risk of impairments and consequent disability, but once 224 established, ART alone may not be sufficient to enable children with HIV to lead healthy lives 225 (2).226 Given the high prevalence of physical and sensory impairments amongst children living with 227 HIV, our study underlines the need for increased availability of rehabilitation services to 228 support school age children and adolescents with HIV. Currently, the few existing services are 229 mainly located in urban areas or private health facilities which limits access for many (35). 230 Greater support for children with learning difficulties is required in schools to facilitate social inclusion and educational attainment (36) as learning, remembering, and concentration appear 231 232 to be common in HIV-infected and uninfected children. 233 Although incorporating disability inclusive approaches into HIV treatment and care is likely to 234 increase the social participation and school functioning of children with HIV (37), so far only 235 5 of 18 countries (27%) in Eastern and Southern Africa have recognised the need for specific 236 support services and interventions for people with disabilities in their national strategic 237 responses to HIV and AIDS (38). Although Zimbabwe is one of these countries, the findings

of this study suggest that further work is required to extend services to support school age children with HIV.

To our knowledge, this is the first study to estimate the prevalence of disability and its association with school and social functioning in HIV-infected and uninfected older children in a Sub-Saharan African population. Study limitations include the potential selection bias from non-probability based sampling: selecting the first five children attending the HIV clinic may have led to under-reported disability if children with physical or behavioral disabilities were more likely to attend at clinic later. Alternatively, children with disabilities may have been less likely to go to school and thus be the first to attend. Furthermore, misclassification and/or recall bias from the use of self-reported functional difficulties and disability without contemporaneous clinical measures of the impairments or their cause, coupled with the fact that carers may not accurately perceive their children's functional difficulties, may have also led to under-reported disability.

Unfortunately, socioeconomic data such as household income and size, asset ownership, caregiver education and food security were not available which meant that analyses could not be adjusted for socio-economic status. This is important as poverty and disability are likely to reinforce each other, leading to vulnerability and exclusion. Children who are poor are more likely to become disabled through poor healthcare, malnutrition, or dangerous living conditions. Once disabled, they are more likely to be denied basic resources that would mitigate deepening poverty (39). There is evidence that poverty is a major contributor to poor treatment adherence among in HIV-infected children. (40). Furthermore, evidence from a large cross-sectional study of South African adolescents from deprived urban areas showed that orphanhood by AIDS was significantly related to childhood depression, peer problems, post-traumatic stress and behavioural problems; however, adjusting for poverty indictors in this

study attenuated the association between AIDS-orphanhood and these psychological problems (41).

Although it is evident that disability is common in HIV-infected children and has a major impact on their lives, further research to understand the aetiology of different impairments is needed to inform the design of effective interventions and appropriate rehabilitation services. Examples of the type of interventions for HIV-infected children that could be introduced include: 1) routine screening for impairments 2) linking HIV care to rehabilitation and additional clinical services (*e.g.* ENT in the case of hearing impairment) 3) interventions to promote school inclusion and social acceptance among children with HIV (e.g. through training of parents, teachers and peers).

In conclusion, this study suggests physical and cognitive functional difficulties are common among children with HIV. These difficulties are associated with school exclusion, including impaired educational progress, difficulty forming friendships and reduced ability to participate in lessons and activities. Further work is required to develop tools to better detect and understand the need for rehabilitation and support services within paediatric HIV programmes.

### **Competing interests**

The authors have no competing interests to declare.

### **Authors' contributions**

- RAF and HK designed the study. RR performed the statistical analysis and drafted the report.
- All authors provided feedback on the draft manuscript and approved the final manuscript.

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#### References

- UNICEF 2017. UNICEF Data: Monitoring the Situation of Children and Women; Global and regional trends, July 2017 Current Status and Progress. Available online from https://data.unicef.org/topic/hivaids/global-regional-trends/#
- 2. Banks LM, Zuurmond M, Ferrand R, Kuper H. The relationship between HIV and prevalence of disabilities in sub-Saharan Africa: systematic review (FA). Trop Med Int Health. 2015;20(4):411-29.
- 3. Devendra A, Makawa A, Kazembe PN, Calles NR, Kuper H. HIV and childhood disability: a case-controlled study at a paediatric antiretroviral therapy centre in Lilongwe, Malawi. PLoS One. 2013;8(12):e84024.
- 4. Yust 2004. Retinal and extraocular cytomegalovirus end-organ disease in HIV-infected patients in Europe: a EuroSIDA study, 1994 –2001
- McHugh G, Rylance J, Mujuru H, et al. Chronic Morbidity Among Older Children and Adolescents at Diagnosis of HIV Infection. Journal of Acquired Immune Deficiency Syndromes. 2016;73(3):275-281.
- McNaghten AD, Wan PC, Dworkin MS, Group AASoHDP. Prevalence of hearing loss in a cohort of HIVinfected patients. Arch Otolaryngol Head Neck Surg. 2001;127(12):1516-8.
- 7. Schouten JT, Lockhart DW, Rees TS, Collier AC, Marra CM. A prospective study of hearing changes after beginning zidovudine or didanosine in HIV-1 treatment-naïve people. BMC Infect Dis. 2006;6:28.
- 8. Pasnoor, Mamatha, Richard J. Barohn, and Mazen M. Dimachkie. Toxic Myopathies. Neurologic clinics 32.3 (2014): 647–viii. PMC. Web. 15 May 2017.
- 9. Lowenthal ED, Bakeera-Kitaka S, Marukutira T, Chapman J, Goldrath K, Ferrand RA. Perinatally acquired HIV infection in adolescents from sub-Saharan Africa: a review of emerging challenges. Lancet Infect Dis. 2014;14(7):627-39.
- 10. Hrapcak S, Kuper H, Bartlett P, Devendra A, Makawa A, Kim M, et al. Hearing Loss in HIV-Infected Children in Lilongwe, Malawi. PLoS One. 2016;11(8):e0161421.
- 11. United Nations (UN). Un United Nations Convention on the Rights of Persons with Disabilities <a href="http://www.unorg/disabilities/documents/convention/convention-accessible-pdfpdf">http://www.unorg/disabilities/documents/convention/convention-accessible-pdfpdf</a>. 2006.
- 12. Wabiri N, Taffa N. Socio-economic inequality and HIV in South Africa. BMC Public Health. 2013;13(1):1037.
- 13. Kuper H, Monteath-van Dok A, Wing K, Danquah L, Evans J, Zuurmond M, et al. The impact of disability on the lives of children; cross-sectional data including 8,900 children with disabilities and 898,834 children without disabilities across 30 countries. PLoS One. 2014;9(9):e107300.
- 14. Booysen, F. 2002. Financial responses of households in the free state province to HIV/AIDS-related morbidity and mortality. South African Journal of Economics, 70(7): 1193–1215.
- Washington Group on Disability Statistics, UNICEF. Module on Child Functioning and Disability Available online from <a href="http://www.washingtongroup-disabilitycom/wp-content/uploads/2016/02/wg-unicef-child-disability-background-documentpdf">http://www.washingtongroup-disabilitycom/wp-content/uploads/2016/02/wg-unicef-child-disability-background-documentpdf</a>. 2014.
- 16. Padhani DH, Manji KP, Mtanda AT. Ocular manifestations in children with HIV infection in Dar es Salaam, Tanzania. J Trop Pediatr. 2000;46(3):145-8.
- 17. Taipale A, Pelkonen T, Taipale M, Roine I, Bernardino L, Peltola H, et al. Otorhinolaryngological findings and hearing in HIV-positive and HIV-negative children in a developing country. Eur Arch Otorhinolaryngol. 2011;268(10):1527-32.
- 18. Drotar D, Olness K, Wiznitzer M, Guay L, Marum L, Svilar G, et al. Neurodevelopmental outcomes of Ugandan infants with human immunodeficiency virus type 1 infection. Pediatrics. 1997;100(1):E5.
- 19. Ferguson G, Jelsma J. The prevalence of motor delay among HIV infected children living in Cape Town, South Africa. Int J Rehabil Res. 2009;32(2):108-14.

- Kandawasvika GQ, Ogundipe E, Gumbo FZ, Kurewa EN, Mapingure MP, Stray-Pedersen B. Neurodevelopmental impairment among infants born to mothers infected with human immunodeficiency virus and uninfected mothers from three peri-urban primary care clinics in Harare, Zimbabwe. Dev Med Child Neurol. 2011;53(11):1046-52.
- 21. Msellati P, Lepage P, Hitimana DG, Van Goethem C, Van de Perre P, Dabis F. Neurodevelopmental testing of children born to human immunodeficiency virus type 1 seropositive and seronegative mothers: a prospective cohort study in Kigali, Rwanda. Pediatrics. 1993;92(6):843-8.
- 22. Baillieu N, Potterton J. The extent of delay of language, motor, and cognitive development in HIV-positive infants. J Neurol Phys Ther. 2008;32(3):118-21.
- 23. Heaton RK, Clifford Db Fau Franklin DR, Jr., Franklin Dr Jr Fau Woods SP, Woods Sp Fau Ake C, Ake C Fau Vaida F, Vaida F Fau Ellis RJ, et al. HIV-associated neurocognitive disorders persist in the era of potent antiretroviral therapy: CHARTER Study. Neurology. 2010, 75(23).
- 24. Abubakar A, Holding P, Newton CR, van Baar A, van de Vijver FJ. The role of weight for age and disease stage in poor psychomotor outcome of HIV-infected children in Kilifi, Kenya. Dev Med Child Neurol. 2009;51(12):968-73.
- 25. Boivin MJ, Green SD, Davies AG, Giordani B, Mokili JK, Cutting WA. A preliminary evaluation of the cognitive and motor effects of pediatric HIV infection in Zairian children. Health Psychol. 1995;14(1):13-21.
- 26. Jelsma J, Davids N, Ferguson G. The motor development of orphaned children with and without HIV: Pilot exploration of foster care and residential placement. BMC Pediatr. 2011;11:11.
- 27. McDonald CM, Manji KP, Kupka R, Bellinger DC, Spiegelman D, Kisenge R, et al. Stunting and wasting are associated with poorer psychomotor and mental development in HIV-exposed Tanzanian infants. J Nutr. 2013;143(2):204-14.
- 28. Ruel TD, Boivin MJ, Boal HE, Bangirana P, Charlebois E, Havlir DV, et al. Neurocognitive and motor deficits in HIV-infected Ugandan children with high CD4 cell counts. Clin Infect Dis. 2012;54(7):1001-9.
- 29. Shead GM, Potterton J, Stewart A. Neurodevelopment and growth of institutionalized children with vertically transmitted human immunodeficiency virus. Vulnerable Children and Youth Studies. 2010;5(1):33-43.
- Esch P, Bocqet V, Pull C, Couffignal S, Lehnert T, Graas M, Fond-Harmant, Ansseau M. The downward spiral of mental disorders and educational attainment: a systematic review on early school leaving. BMC Psychiatry201414:237 https://doi.org/10.1186/s12888-014-0237-4
- 31. Rao D, Kekwaletswe TC, Hosek S, Martinez J, Rodriguez F. Stigma and social barriers to medication adherence with urban youth living with HIV. AIDS Care. 2007 Jan;19(1):28-33.
- 32. L. Cluver & F. Gardner. Risk and protective factors for psychological well-being of children orphaned by AIDS in Cape Town: a qualitative study of children and caregivers' perspectives. AIDS Care Vol. 19, Iss. 3,2007a.
- 33. Kumakech E, Cantor-Graae E, Maling S, Bajunirwe F.Peer-group support intervention improves the psychosocial well-being of AIDS orphans: cluster randomized trial. Soc Sci Med. 2009 Mar; 68(6):1038-43.
- 34. Betancourt TS, Meyers-Ohki SE, Charrow A, Hansen N. Mental Health and Resilience in HIV/AIDS-Affected Children: A Review of the Literature and Recommendations for Future Research. *Journal of child psychology and psychiatry, and allied disciplines*. 2013;54(4):423-444. doi:10.1111/j.1469-7610.2012.02613.
- 35. World Health Organization (2010a) Community Based Rehabilitation (CBR) Guidelines. Geneva: WHO. http://www.who.int/disabilities/cbr/guidelines/en/index.html Werner D. (2009) Disabled Village Children. Berkeley, CA.: The Hesperian Foundation. <a href="http://hesperian.org/books-and-resources/">http://hesperian.org/books-and-resources/</a>
- 36. UNESCO, 'Inclusive education: the way of the future', paper prepared for the International Conference on Education, Geneva, 2008.

- 411 37. Nixon SA, Hanass-Hancock J, Whiteside A, Barnett T. The increasing chronicity of HIV in sub-Saharan Africa: Re-thinking "HIV as a long-wave event" in the era of widespread access to ART. *Globalization and Health*. 2011;7:41. doi:10.1186/1744-8603-7-41.
  - 38. Hanass-Hancock J, Strode A, Grant C. Inclusion of disability within national strategic responses to HIV and AIDS in Eastern and Southern Africa. Disability and Rehabilitation. 2011;33(22-23):2389-96.
  - 39. UNICEF Children and Young People with Disabilities Fact Sheet May 2013

- 40. Bermudez LG, Jennings L, Ssewamala FM, Nabunya P, Mellins C, McKay M. Equity in adherence to antiretroviral therapy among economically vulnerable adolescents living with HIV in Uganda. AIDS Care. 2016;28(sup2):83-91. doi:10.1080/09540121.2016.1176681.
- 41. Lucie Cluver, Frances Gardner & Don Operario. Poverty and psychological health among AIDS-orphaned children in Cape Town, South Africa. AIDS Care Vol. 21, Iss. 6,2009.

# **Tables and Figures**

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Table 1. Baseline Characteristics of HIV-infected and HIV-uninfected Children in Zimbabwe

Characteristic	HIV+ n=202 n (%) <sup>a</sup>	HIV- n=285 n (%)	p value
Age	. ,		
6 -11 years	132 (65.4)	165 (57.9)	$0.06^{b}$
12 -16 years	70 (34.6)	32 (42.1)	
Median (IQR) years	11 (8, 13)	10 (8, 13)	0.61 <sup>c</sup>
Sex			
Female	97 (48.0)	139 (48.8)	$0.11^{b}$
Orphan status			
Single orphan	69 (34.2)	25 (8.8)	<0.001 <sup>b</sup>
Double orphan	28 (13.9)	7 (2.5)	
Not orphaned	98 (48.5)	245 (85.9)	
Age at HIV diagnosis			
Median (IQR) years	5 (3, 7)		
Age at ART initiation			
Median (IQR) years	8 (5, 10)		
ART duration			
<1 years	75 (37.1)		
1-5 years	97 (48.0)		
>5 years	30 (14.9)		
CD4			
<200 cells/μl	9 (4.5)		
$200\text{-}500 \text{ cells/}\mu\text{l}$	47 (23.2)		
>500 cells/µl	144 (71.3)		
Median (IQR) cells/μl	726 (476, 941)		
Viral load			
<400 copies/ml	152 (75.2)		
400-5000 copies/ml	14 (7.0)		
>5000 copies/ml	32 (15.8)		
Median (IQR) copies/ml	19 (19, 250)		

Abbreviations: HIV+ HIV-infected, HIV- HIV-uninfected, SD standard deviation, IQR inter quartile range

a) n (%) shown, except for median and IQR shown in italics b) p value from  $\chi^2$  test c) p value from Mann-Whitney U test

Table 2. Domains of Disability and Functioning in HIV-infected and HIV-uninfected Children in Zimbabwe

Outcome	HIV+	HIV-	Crude OR	p value <sup>a</sup>	aOR (95% CI)	p value <sup>a</sup>
Outcome	n=202	n=285	(95% CI)			
	n (%)	n (%)				
Any disability	76 (37.6)	53 (18.8)	2.3 (1.6, 5.3)	< 0.001	2.8 (1.8, 4.2)	< 0.001
Seeing	16 (7.7)	9 (3.1)	2.7 (1.2, 6.0)	0.009	3.0 (1.3, 6.9)	0.009
Hearing	10 (4.8)	4 (1.4)	3.4 (1.1, 10.6)	0.031	3.4 (1.0, 10.5)	0.036
Walking	5 (2.5)	1 (0.4)	7.4 (0.9, 63.5)	0.065	7.4 (0.9, 63.5)	0.055
Speaking	9 (4.3)	3 (1.1)	4.0 (1.1, 14.5)	0.042	3.8 (1.1, 13.9)	0.042
Learning	48 (23.2)	33 (11.6)	2.1 (1.3, 3.2)	0.002	3.9 (1.4, 3.4)	0.001
Memory	37 (17.8)	16 (5.6)	3.6 (2.0, 6.6)	< 0.001	3.5 (2.0, 6.6)	< 0.001
Self-caring	3 (1.5)	1 (0.4)	1.7 (0.4, 8.0)	0.072	1.6 (0.4, 7.8)	0.524
Anxiety	42 (20.3)	14 (5.6)	4.6 (2.4, 8.2)	0.000	4.4 (2.4, 8.1)	< 0.001
Depression	32 (15.5)	12 (4.2)	4.2 (2.1, 8.5)	0.010	4.2 (2.1, 8.4)	0.010
Controlling behaviour	3 (1.5)	1 (0.4)	4.0 (0.4, 39.4)	< 0.001	4.0 (0.4, 39.3)	0.003
Concentration	2 (1.0)	6 (2.1)	0.4 (0.1, 2.2)	0.478	0.4 (0.1, 2.2)	0.311
Accepting change	39 (10.9)	36 (12.6)	1.6 (0.9, 2.6)	0.085	1.5 (1.0, 2.5)	0.075
Making friends	10 (4.8)	1 (0.4)	14.6 (1.9, 115.2)	0.001	14.8 (1.9, 116.6)	0.011

**Abbreviations: HIV+** HIV-infected, **HIV-** HIV-uninfected, **OR** odds ratio, **aOR** age, sex adjusted odds ratio. a) p value from  $\chi^2$  test

Table 3. Difference in HIV Characteristics Amongst HIV-infected Children With and Without Disability in Zimbabwe

437	Disability in Zimbabwe					
Characteristic	HIV+ with disability: n <sup>a</sup> =76	HIV+ without disability: n=126	p value			
Age Median (IQR) years	10.9 (2.6)	10.3 (2.6)				
6-9 years	24 (31.6)	48 (38.1)	0.77	-		
10-12 years	31 (40.8)	50 (39.7)				
13-14 years	15 (19.7)	20 (15.9)				
15-16 years	6 (7.9)	8 (6.4)				
Age at diagnosis						
Median (IQR) years	5.0 (3.0)	5.1 (2.9)		-		
Age of ART initiation						
Median (IQR) years	8 (6, 11)	7 (5, 10)	0.78			
Sex						
Female	35 (46.0)	62 (49.2)	0.66			
				aOR (95% CI)		
CD4 count						
Median (IQR ) cells/uL	736 (513, 914)	720 (459, 910)				
<200 cells/uL	3 (4.0)	6 (4.7)	0.78	1.0		
200-500 cells/uL	15 (19.7)	32 (25.4)		1.4 (0.8, 2.5)		
>500 cells/uL	57 (75.0)	87 (69.1)				
Viral load						
Median (IQR) copies/ml	19 (19, 190)	19 (19, 343)				
<400 copies/ml	57 (75.0)	95 (75.4)	0.16	1.0		
400-5000 copies/ml	2 (2.6)	12 (9.5)		1.1 (0.7, 1.6)		
>5000 copies/ml	14 (18.4)	18 (14.3)				
ART duration						
Median (IQR) years	2 (1,5)	1 (0, 4)				
<1 years	24 (31.6)	51 (40.5)	0.21	1.0		
1-5 years	39 (51.3)	58 (46.0)		1.2 (0.8, 1.9)		
>5 years	13 (17.1)	17 (13.5)		•		
No of hospital admissions in 12 months	` '	. ,				
>1	5 (6.6)	5 (4.0)		1.9 (0.6, 6.1)		
Past history of TB	29 (38.2)	50 (39.7)	0.94	0.9 (0.5, 1.6)		

**Abbreviations HIV**+ HIV-infected, **HIV**- HIV Uninfected, **aOR** odds ratio adjusted for age and sex, **ART** antiretroviral therapy, **TB** tuberculosis, **IQR** inter quartile range.

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a) n shown, except for median and IQR shown in italics

Table 4. School and Social Inclusion at School in HIV-infected and HIV-uninfected Children and in HIV-infected Children with and Without Disability

Characteristic	HIV+ n=202 n (%)	HIV- n=285 n (%)	aOR (95% CI)	HIV+ with disability n=76 n (%)	HIV+ without disability n=126 n (%)	aOR (95% CI)
School inclusion as reported by children and their	carers					
Currently enrolled in school	194 (96.0)	273 (96.0)	0.98 (0.4, 2.5)	71 (93.4)	123 (97.6)	0.3 (0.1, 1.5)
Enrolled in the same grade as peers	102 (50.5)	197 (69.1)	2.4 (1.6, 3.6)	24 (31.6)	78 (61.9)	3.3 (1.7, 6.1)
Ever repeated a year at school	68 (33.7)	53 (18.6)	2.5 (1.6, 3.8)	32 (42.1)	36 (28.6)	1.9 (1.0, 3.6)
Social inclusion at school as reported by children a	nd their care	rs		<u> </u>		
No help from teachers, if problem at school	4 (2.0)	2 (0.7)	2.1 (1.2, 3.8)	2 (2.6)	2 (1.6)	1.7 (0.9, 3.2)
No help from friends, if problem at school	15 (7.4)	3 (1.1)	3.0 (2.0, 4.5)	11(14.5)	4 (3.2)	1.5 (0.9, 2.4)
Child has no friends to play with	2 (1.0)	1 (0.4)	1.8 (0.7, 5.0)	2 (2.6)	1 (0.8)	1.7 (0.8, 5.7)
Friends look up to child as a leader	108 (53.5)	147 (51.6)	1.1 (0.8, 1.6)	44 (57.9)	41 (32.5)	2.1 (1.4, 3.4)
Other children hit, hurt /say nasty things to child	58 (28.7)	28 (9.8)	3.7 (2.2, 6.0)	30 (39.5)	28 (22.2)	2.5 (1.3, 4.8)
Child does not feel included in lessons and activities	6 (3.0)	2 (0.7)	4.7 (2.7, 8.3)	3 (4.0)	3 (2.4)	0.6 (0.1, 3.0)

Abbreviations HIV+ HIV-infected, HIV- HIV Uninfected, aOR odds ratio adjusted for age and sex.

# Supplementary Table 5. School and Social Inclusion at School in Disabled Children With and Without HIV-infection

Characteristic	HIV+ with disability n=76 n (%)	HIV- with disability n=53 n (%)	aOR (95% CI)
School inclusion as reported by children and their carers			
Currently enrolled in school	71 (93.4)	52 (98.1)	1.1 (0.3, 4.5)
Enrolled in the same grade as peers	24 (31.6)	32 (57.1)	0.3 (0.1, 0.6)
Ever repeated a year at school	32 (42.1)	12 (21.4)	3.3 (1.4, 8.0)
Social inclusion as reported by children and their carers			
No help from teachers, if problem at school	2 (2.6)	1 (1.8)	0.1 (0.1, 8.1)
No help from friends, if problem at school	11 (14.5)	1 (1.8)	0.5 (0.0, 0.9)
Child has no friends to play with	2 (2.6)	0 (0.0)	-
Friends look up to child as a leader	26 (34.2)	27 (48.2)	0.5 (0.3, 1.1)
Other children physically or verbally bully	30 (39.5)	3 (5.4)	11.3 (3.9, 39.8)
Child excluded in lessons and activities	3 (4.0)	1 (1.8)	0.4 (0.0, 4.4)