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Validation of the Episodic Disability Framework with Adults Living with HIV

Kelly K. O'Brien^{1,2}; Steven Hanna^{2,3}; Sandra Gardner⁴; Ahmed M. Bayoumi^{5,6,7}; Sergio Rueda⁴; Trevor A. Hart^{8,9}; Curtis Cooper¹⁰; Patricia Solomon²; Sean B. Rourke^{4,5,11}; Aileen M. Davis^{6,12} and the *OHTN Cohort Study Research Team**

¹ Department of Physical Therapy, University of Toronto, Toronto, Ontario, Canada

² School of Rehabilitation Science, McMaster University, Hamilton, Ontario, Canada

³ Department of Clinical Epidemiology and Biostatistics, McMaster University, Hamilton, Ontario, Canada

⁴ The Ontario HIV Treatment Network, Toronto, Ontario, Canada

⁵ Centre for Research on Inner City Health, St. Michael's Hospital, Toronto, Ontario, Canada

⁶ Institute of Health Policy, Management and Evaluation, University of Toronto, Toronto, Ontario, Canada

⁷ Department of Medicine, University of Toronto, Toronto, Ontario, Canada

⁸ Ryerson University, Department of Psychology, Toronto, Ontario, Canada

⁹ Dalla Lana School of Public Health, University of Toronto, Toronto, Ontario, Canada

¹⁰ University of Ottawa, Division of Infectious Diseases, The Ottawa Hospital
Ottawa, Ontario, Canada

¹¹ Department of Psychiatry, University of Toronto, Toronto, Ontario, Canada

¹² University Health Network, Toronto Western Research Institute, Toronto, Ontario, Canada

*The *OHTN Cohort Study Research Team* consists of Dr. Sean B. Rourke (Principal Investigator, University of Toronto and OHTN), Dr. Ann Burchell (Co-Principal Investigator,

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OHTN), Dr. Sandra Gardner (OHTN), Dr. Sergio Rueda (OHTN), Dr. Ahmed Bayoumi and Dr. Kevin Gough, St. Michael’s Hospital; Dr. Jeffrey Cohen, Windsor Regional Hospital; Dr. Curtis Cooper, Ottawa General Hospital; Dr. Don Kilby, University of Ottawa Health Services; Dr. Mona Loutfy and Dr. Fred Crouzat,, Maple Leaf Medical Clinic; Dr. Anita Rachlis and Dr. Nicole Mittmann, Sunnybrook Health Sciences Centre; Dr. Janet Raboud and Dr. Irving Salit, Toronto General Hospital; Dr. Edward Ralph, St. Joseph’s Health Care; Dr. Roger Sandre, Sudbury Regional Hospital; Dr. Marek Smieja, Hamilton Health Sciences, McMaster University Medical Centre; and Dr. Wendy Wobeser, Hotel Dieu Hospital.

Validation of the Episodic Disability Framework with Adults Living with HIV

ABSTRACT

Purpose: To assess the validity of dimensions of disability in the *Episodic Disability Framework*, a conceptual framework derived from the perspective of adults living with HIV.

Methods: We conducted confirmatory factor analyses with 913 adults living with HIV in an observational cohort study called the Ontario HIV Treatment Network Cohort Study (OCS). We tested hypotheses that dimensions of disability in the *Episodic Disability Framework* were represented by a group of measured variables in the observational database.

Results: A model comprised of four latent variables and 43 indicator variables with one cross-loading was superior to models with fewer latent variables and more indicator variables and supported the validity of disability dimensions: physical health symptoms (represented by 21 indicator variables), mental health symptoms (10 variables), difficulties with day-to-day activities (5 variables), and challenges to social inclusion (8 variables). Overall goodness of fit statistics were $\chi^2=2621.50$ ($p<0.001$), Comparative Fit Index=0.912, Tucker Lewis Index=0.907 and Root Mean Square Error of Approximation=0.048. Dimensions of disability correlated with each other ranging from $r=0.44$ (between physical symptoms and challenges to social inclusion) to $r=0.81$ (between physical symptoms and difficulties with day-to-day activities).

Conclusions: This study supports the validity of four disability dimensions in the *Episodic Disability Framework*. This framework provides a new way to conceptualize disability and can lay the foundation for developing a future HIV disability measure for clinical and health services research.

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INTRODUCTION

With improved access to combination antiretroviral therapy, many people living with Human Immunodeficiency Virus (HIV) are living longer [1, 2]. However, individuals can face a variety of health-related consequences of HIV, comorbidities, and the potential side effects of treatment [3-7]. Health challenges experienced by adults living with HIV are complex and can range from symptoms such as fatigue, weakness, pain, and changes in body composition to difficulties participating in society, such as employment [5, 8-10]. Collectively, these health-related challenges may lead to disability.

In an earlier phase of research, we used qualitative methods to develop the *Episodic Disability Framework*, a new conceptual framework of disability derived from the perspective of 38 adults living with HIV [11]. In this *Framework*, disability is multi-dimensional and episodic characterized by periods of wellness and illness. The *Episodic Disability Framework* consists of three main components. The first component includes dimensions of disability: a) symptoms and impairments (e.g. physical and mental health-related challenges), b) difficulties carrying out day-to-day activities (e.g. difficulties with walking, activities of daily living, carrying out household chores), c) challenges to social inclusion (e.g. challenges fulfilling roles as a parent, difficulties engaging in work and school, personal relationships), and d) uncertainty about future health (e.g. worrying about the future and the impact of uncertainty on life decisions) [11]. The second component includes contextual factors: extrinsic factors (social support and stigma) or intrinsic factors (living strategies and personal attributes) that can exacerbate or alleviate dimensions of disability [12]. The third component includes triggers, which are defined as life events that mark major or momentous episodes of disability. Examples include receiving notification of an HIV diagnosis, starting or changing antiretroviral medications, and experiencing a serious illness [11].

The *Episodic Disability Framework* goes beyond existing frameworks such as the *Disablement Process* or *International Classification of Functioning, Disability and Health (ICF)* which were developed prior to the advent of combination antiretroviral therapy or were not developed specifically from the perspectives of adults living with HIV [13-18]. Distinct features of the *Framework* include the description of disability as episodic in nature, the detailed inclusion of contextual factors that influence disability, and the consideration of life events that may initiate a major episode. The *Episodic Disability Framework* may be used by clinicians, adults living with HIV, representatives from community-based service organizations, and researchers to better understand and articulate episodic disability experienced by adults living with HIV. However, the validity of this framework has not been empirically tested.

In this study, we assessed the construct validity of dimensions of disability in the *Episodic Disability Framework*. Specifically, we determined whether dimensions of disability within the *Framework* were represented by a group of measured variables in a large observational cohort of people living with HIV. Validating the *Episodic Disability Framework* will establish a strong conceptual foundation from which to guide the comprehensive assessment of disability, indicate areas to apply rehabilitation interventions that may prevent or mitigate disability, and establish a future measure of disability experienced by adults living with HIV.

METHODS

We evaluated the validity of the *Episodic Disability Framework* using data from the Ontario HIV Treatment Network Cohort Study (OCS) [19, 20]. The OCS is an ongoing longitudinal multi-site clinic-based observational cohort study that has collected clinical and socio-behavioural data from HIV-positive adults who are residents of Ontario and provided informed consent to understand the psychosocial, behavioural and health context of people living with HIV and their

patterns of health services use [19]. Participants complete an interviewer-administered questionnaire at yearly intervals. We focused on participants in three sites in Toronto, Ontario (Toronto General Hospital, St. Michael’s Hospital and Sunnybrook Health Sciences Centre) who completed a 90 to 120 minute questionnaire comprised of detailed health status instruments [21-29] and questions related to employment status, housing, and income. This study was approved by the OCS Governance Committee and by Research Ethics Boards at McMaster University, Hamilton, Ontario, and Ryerson University, Toronto, Ontario, Canada.

OCS Measures

Variables in our measurement model included a combination of summary, subscale and item scores from the OCS questionnaire. The HIV Symptom Index is a 20-item questionnaire that describes the presence and burdensome nature of symptoms experienced by adults with HIV. Each item is scored using a five point ordinal scale ranging from ‘I do not have this symptom’ to ‘I have this symptom and it bothers me a lot’. The HIV Symptom Index demonstrates good construct validity with people living with HIV [26] and developers advise interpreting each symptom separately rather than calculating summary scores. The Center for Epidemiologic Studies for Depression Scale (CES-D) is a 20-item instrument that measures depressive symptomatology [23]. The Scale possesses high internal consistency, adequate test-retest reliability and concurrent and construct validity in the general population [23] and has been used extensively with people living with HIV [30-34]. Item scores can be added to generate a summary score, ranging from 0-60. Higher scores indicate more symptoms and higher frequencies of experiencing depressive symptoms [23]. The Health Utilities Index (HUI) is a generic utility index designed to measure health status and health-related quality of life [22, 35]. Scores range from zero to one with higher scores representing better health. The HUI is

comprised of 41 items, from which domain scores may be calculated for vision, hearing, speech, dexterity, pain, emotion, cognition, mobility, self-care, and ambulation [22, 35]. The SF-36 questionnaire is a generic health-related quality of life instrument [21, 36]. The measurement properties of the SF-36 are well established, demonstrating high internal consistency reliability, content, construct, and predictive validity [24, 25]. Domain scores for the SF-36 include physical function, role limitation physical, bodily pain, vitality, role limitation emotional, mental health, social function, and general health perception that range on a scale from 0-100 with higher scores indicating better quality of life. The Ongoing Problems Stress Index is a component of the National Population Health Survey Stress Questionnaire. The index includes 17 statements that pertain to stressful situations asking whether the statement is true or false. Total scores can range from 0-17 with higher scores representing increased stress [28, 37]. The Medical Outcomes Study Cognitive Functioning Scale includes four items with a six point ordinal scale that measures challenges across reasoning and problem solving, memory, attention, and concentration and thinking [29]. The MOS Cognitive Function Scale scores range from 0-100 with higher scores representing better cognitive function. The scale has been validated for use with people living with HIV [38]. The EQ-5D EuroQoL Questionnaire is a standardized general measure of health related quality of life. The EQ-5D is comprised of five items (mobility, self-care, usual activities, pain and discomfort, and anxiety and depression), each of which is coded on a three point ordinal scale ranging from no problems to severe problems [27, 39]. The EQ-5D demonstrated good construct validity with people living with HIV [40]. Remaining indicator variables for our measurement model were individual items generated specifically for the OCS questionnaire including housing, employment, education, income and

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occupation. This questionnaire was developed and pre-tested with extensive consultation with HIV researchers and community members across Ontario.

Analysis

Our purpose was to assess the construct validity of dimensions of disability in the *Episodic Disability Framework*, a conceptual framework derived from the perspective of adults living with HIV. We conducted a confirmatory factor analysis to assess the construct validity of the dimensions of disability in the *Episodic Disability Framework*. Specifically, we used cross-sectional data to test hypotheses that dimensions of disability (latent variables) in the *Framework* were represented by observed measures in the OCS (indicator variables). We allocated indicator variables to latent variables based on categories that comprised each dimension in the *Episodic Disability Framework* [11, 41]. We further divided symptoms and impairments into physical and mental health symptoms, because while correlated, these variables differ conceptually [42]. Uncertainty was not considered in our analysis because no measures in the OCS captured this dimension. See figure 1 for our *a priori* measurement model.

Insert figure 1 about here

We assessed convergent construct validity by determining the extent to which variables from the OCS represented a hypothesized dimension of disability with factor loadings >0.30. We assessed divergent construct validity by determining the extent to which dimensions of disability were distinct constructs that together comprised the larger construct of disability. We considered correlations between latent variables <0.80 as signifying distinct dimensions of disability [43].

Sample Size Estimation

At least five observations per unknown parameter estimate is required for confirmatory factor analysis [43, 44]. Our primary hypothesized model had 120 unknown parameter estimates (not shown) that included factor loadings and error variances for each of the indicator variables ($n=110$) plus latent variable variances and covariances ($n=10$). Hence, we required at least 600 observations for our analysis.

Confirmatory Factor Analysis

We used Mplus version 6.0 statistical software and the weighted least squares method of estimation, which is appropriate for combinations of continuous and categorical variables and non-normally distributed data [45]. Prior to our analysis, we recoded all 'don't know' or 'refused' item responses to missing values. We assessed the mean, standard deviation, skewness, kurtosis, and frequency of missing values for each variable. All variables were recoded such that higher scores indicated greater disability. Where possible, we included subscale or summary scores to simplify the measurement model. We estimated and reported standardized parameter estimates (or factor loadings) given the variables possessed different units of measurement. Using the weighted least squares method of estimation, missing data were dealt with using pairwise deletion [45].

Confirmatory Factor Models

In our initial model, we included all variables that we hypothesized represented the dimensions of disability in the *Episodic Disability Framework* (figure 1). Confirmatory factor analysis involves a complex estimation of parameters that include factor loadings, variances, covariances and measurement error for indicator and latent variables [43]. Each of these parameters can be freely estimated (derived from the analysis) or fixed (assigned a value by the researcher) and

indicator variables also can be allowed to cross load (represent multiple latent variables) [44]. For our analysis, all error variances were freely estimated. All error covariances were initially fixed to zero and variables were allowed to load on only one latent variable. Latent variables were allowed to correlate freely. We systematically modified the model to reduce complexity and improve fit where possible by removing variables that measured duplicate categories of disability (were highly correlated), lacked variability in scores (e.g. ceiling or floor effect), or when item removal made theoretical and clinical sense. We cross loaded variables, allowing them to load on more than one latent variable, if this made theoretical and clinical sense and the modification indices suggested a large improvement in fit [43].

We used a combination of approaches to evaluate the confirmatory factor analysis (CFA) solution including overall goodness of fit, localized areas of strain (areas of misfit in the solution), and interpretability (magnitude and statistical significance of the parameter estimates) [43]. If the CFA solution demonstrated adequate goodness of fit we considered this as construct validation of the disability dimensions in the *Episodic Disability Framework* [11].

Overall Goodness of Fit: We considered a Root Mean Square Error of Approximation (RMSEA) <0.05 , Comparative Fit Index (CFI) ≥ 0.95 , and Tucker Lewis Index (TLI) ≥ 0.95 to indicate good model fit [46, 47]. We considered the RMSEA as the primary statistic for overall goodness of model fit because it is recommended for confirmatory factor analytical contexts [48]. We reported the chi square statistic (χ^2) but did not consider it a determinant of model fit given its sensitivity to large sample sizes, which can overestimate lack of model fit [43].

Localized Area of Strain: We assessed areas of misfit in the solution by reviewing modification indices (values that indicate how much the chi square statistic would decrease if fixed parameters were allowed to be freely estimated) to identify ways to improve model fit. However, we only

made modifications if they made theoretical and clinical sense [49].

Parameter Estimates: We considered standardized coefficients (i.e., factor loadings) ≥ 0.30 ($p < 0.05$) as ‘representing’ a given dimension of disability [43]. We used chi square difference tests where possible to identify statistically significant improvements in model fit.

We evaluated six CFA solutions. Our primary hypothesized model (Model 1) included a four factor model with all possible OCS variables. Model 2 similarly included all possible variables but had three factors as we collapsed the physical and mental health symptoms into one latent variable. Models 3 and 4 were simplified models with four and three latent variables respectively. We removed variables that lacked variance, were duplicate measures of disability (i.e., measured the same construct), or when their removal made clinical or theoretical sense. Models 5 and 6 included cross loading the SF-36 social function subscale score across three (Model 5) and two (Model 6) dimensions of disability.

RESULTS

We studied 913 adults living with HIV who completed OCS questionnaires between October 2007 and March 2009 in Toronto, Ontario. Participants were primarily male (83%), taking combination antiretroviral therapy (86%), with a median age of 47 years and median length of time since diagnosis of 11.2 years (table 1).

Insert table 1 about here

Indicator Variables

Indicator variables in our original hypothesized model included 37 item, 17 subscale, and three summary scores that spanned continuous ($n=24$) and discrete ordered categorical ($n=33$) levels of measurement (table 2). Skewness and kurtosis values (not shown) indicated non-normality of

the data, supporting our use of the weighted least squares method of estimation.

Insert table 2 about here

Confirmatory Factor Analysis Solutions

Results of all CFA solutions are presented in table 3. Correlation matrices are available on request. Of all the models tested, the four latent variable model with 43 indicator variables and cross loading of the SF-36 social function subscale score onto the physical and mental health symptom and impairment dimensions provided the best fit to the data (Model 6).

Our initial model, Model 1, consisted of four latent variables including physical symptoms and impairments (represented by 25 indicator variables), mental health symptoms and impairments (11 indicator variables), difficulties with day-to-day activities (8 indicator variables), and challenges to social inclusion (13 indicator variables) (table 2). Model 2 consisted of three latent variables represented by 57 indicator variables whereby physical and mental health symptoms and impairments were collapsed into one latent variable. The difference between Models 1 and 2 was significant [chi square difference test (df) = 302.14 (3), $p < 0.001$], indicating that Model 1 with four latent variables possessed better fit than Model 2 (table 3).

Model 3 was a simplified model with 43 indicator variables across four latent variables. Fourteen variables were removed from Model 1 because they were highly skewed, lacked variance, were highly or lowly correlated with other variables in the model, duplicated other measures in the model, or because removing the variable improved model fit as well as made clinical and theoretical sense in accordance with the *Episodic Disability Framework*. We were unable to use a likelihood ratio (i.e., chi-square difference) test to compare Model 3 with Model 1 because the former was not nested within the latter. However, overall goodness of fit statistics for Model 3 improved across all indices including a reduction in the chi square statistic, [χ^2 (df):

3077.75 (854)], increase in CFA (0.889) and TLI (0.883) and reduction in RMSEA (0.053) (table 3). These values still did not achieve our criteria for goodness of fit. High modification indices (>250) suggested the chi square statistic could decrease by more than 250 if the SF-36 social function summary score were free to load on both physical and mental health symptoms and impairments. Chi square difference testing of Model 4 versus Model 3 further confirmed that a four latent variable model better represented the data (table 3).

Model 5 included cross-loading of the SF-36 social function subscale score on physical and mental health symptoms and impairments and challenges to social inclusion. In addition to the modification indices (which indicated the overall chi square statistic would decrease by more than 250 if the SF-36 social function summary score were free to load on both physical and mental health symptoms and impairments), cross loading was considered reasonable based on the phrasing of the subscale items that asked: ‘to what extent has your *physical* and *emotional* problems interfered with your social activities?’ and ‘how much time has your *physical* and *emotional* problems interfered with your social activities?’ We first ran a null model with cross loading the SF-36 social function subscale score across three latent variables (physical symptoms and impairments, mental health symptoms and impairments, and challenges to social inclusion) (Model 5). Results (not shown) showed the SF-36 social function subscale significantly loaded on the symptoms and impairments dimensions only and did not load on challenges to social inclusion. Chi square difference testing that compared the Model 5 (cross loading on three dimensions) to Model 6 (cross loading on the two symptoms and impairments dimensions only) was not significant [χ^2 (df): 0.258 (1); $p=0.612$] indicating that the nested model (Model 6) did not significantly worsen the fit (table 3). Hence, overall goodness of fit in combination with interpretability of the parameter estimates indicated that Model 6 was the best and final model to

represent dimensions of disability in the *Episodic Disability Framework*.

Insert table 3 about here

Final Model - Dimensions of Disability

The final model was comprised of four dimensions of disability and 43 indicator variables (Model 6) (table 4). Overall goodness of fit statistics were $\chi^2 = 2621.50$ ($p < 0.001$); CFI = 0.912, TLI = 0.907, and RMSEA = 0.048, achieving our *a priori* criterion for goodness of fit (RMSEA). All the indicator variables significantly loaded on their hypothesized dimensions (factor loadings ≥ 0.30). To interpret the first factor loading in table 4, one standard deviation increase in physical symptoms and impairments is associated with a 0.78 standard deviation increase in loss of energy. Alternatively, the correlation between loss of energy and physical symptoms and impairments is 0.78.

Insert table 4 about here

The SF-36 social function subscale score cross-loaded on the physical symptoms (0.21) and the mental health symptoms dimensions (0.61). The latent variables (dimensions of disability) were correlated with each other, ranging from $r = 0.44$ (between physical symptoms and impairments and challenges to social inclusion) to $r = 0.81$ (between physical symptoms and impairments and difficulties with day-to-day activities) (figure 2).

Insert figure 2 about here

DISCUSSION

This is the first study to empirically assess the validity of the *Episodic Disability Framework*, a conceptual framework of disability developed from the perspective of adults living with HIV. Results of our confirmatory factor analysis that demonstrated good overall model fit support the validity of the dimensions of disability in the *Episodic Disability Framework* [11]. Although

evidence exists supporting the validation of quality of life frameworks with HIV populations [51, 52], this is the first known validation of an HIV-specific framework of disability.

Our initial model included all possible variables that represented disability in the OCS questionnaire. This initial approach led to an overwhelmingly complex model comprised of duplicate measures of disability. By including summary and subscale scores where possible and removing variables where the process made statistical and clinical sense, we were able to simplify the model. In the final model, dimensions of disability were represented by 35 indicator variables derived from six HIV-specific and generic health status measures as well as eight indicator variables derived from single items (table 4). Existing validated health status measures represented the symptoms and impairments and difficulties with day-to-day activities dimensions, whereas the challenge to social inclusion dimension was represented by individual items in the OCS questionnaire. These observations support results from a content analysis that suggested challenges to social inclusion experienced by adults living with HIV were less well represented in existing health status instruments compared with symptoms and impairments [41].

All indicator variables significantly loaded on their hypothesized dimensions of disability (factor loadings ≥ 0.30). We allowed the SF-36 social function subscale scores to load on both physical symptoms (0.21) and mental health symptoms dimensions (0.61). Despite the factor loading on physical symptoms was <0.30 , we retained this cross loading in the model for two reasons. First, allowing the SF-36 social function summary score to load on both physical and mental health symptoms and impairments improved the model fit by reducing the RMSEA from 0.053 to 0.048 (not shown). Second, we considered this cross loading to be reasonable because the subscale items included phrasing that referred to how physical and emotional problems interfered with social activities.

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Correlations between the latent variables ranged from $r = 0.44$ to $r = 0.81$. High correlations existed between physical symptoms and difficulties with day-to-day activities ($r = 0.81$) and physical and mental health symptoms ($r = 0.79$) suggesting that these latent variables may not be empirically distinct [43]. However, these dimensions of disability were represented by variables that contained items with similar ordered categorical response options such as the HIV Symptom Index and EQ-5D questionnaire, which could explain the high correlations (0.79-0.81) between these dimensions and obscure the discriminant nature between physical and mental health symptoms and impairments and difficulties with day-to-day activities. Hence, we concluded there were four dimensions that comprise the larger construct of disability. The mental health symptoms dimension combined mood (SF-36 mental health subscale score, CES-D) and cognition (MOS cognitive function score) variables. Mood states included items related to depression, and cognition included items related to difficulty focusing attention, problem solving and memory problems. These components of mental health, when measured subjectively, correlate with each other, supporting our decision to collectively consider mental health symptoms as one latent variable in the model [42, 53]. However, treatment strategies to address mental health symptoms may differ depending on whether the impairments are mood or cognition related, suggesting these may be distinct clinical concepts. Future research involving an exploratory factor analysis may help to determine whether mood and cognition should be considered collective or distinct domains (dimensions) in a future measure of disability.

OCS variables possessed different characteristics of measurement which could influence item responses, factor loadings and overall model fit. For example, the timeframe of OCS measures varied, asking about health states within the past four weeks (HIV Symptom Index, SF-36 questionnaire, MOS-HIV Cognitive Function Scale), within the past week (CES-D), or about

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3 today (EQ-5D). Given health states among people living with HIV may fluctuate on a weekly or
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5 even daily basis, responses to item measures may differ depending on the timeframe posed in the
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7 question. High correlations between items may occur because of similar timeframes or response
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9 options rather than their content as it relates to disability. However, we chose to retain these
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11 measures in the model, specifically the EQ-5D subscales in order to fulfill the three indicator
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13 variable minimum required for each latent variable in CFA, which was required for the
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15 difficulties with day-to-day activities dimension [43].
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20 Decisions for using item, subscale, or summary scores in the model could influence
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22 factor loadings and overall indices of model fit. We based our decisions on the construct
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24 measured as it related to the *Episodic Disability Framework* as well as scoring recommendations
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26 established by authors of the original instruments. For example, we used specific items in the
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28 HIV Symptom Index to represent dimensions in the *Framework* because this instrument does not
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30 possess formal summary (or subscale) scores [26]. We used subscale scores of the EQ-5D and
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32 SF-36 questionnaires because they were developed to measure multi-dimensional constructs of
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34 health status or health-related quality of life [21, 22, 27] and contain domains that we
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36 hypothesized represented different dimensions of disability. We used summary scores from the
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38 CES-D, Medical Outcomes Study-HIV Cognitive Functioning Scale and Ongoing Problems
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40 Stress Index because collectively they measured constructs of depressive symptomology [23],
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42 cognition [50] and stress[28] respectively, all of which we hypothesized represented the mental
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44 health symptoms dimension of disability.
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52 When validating the *Episodic Disability Framework*, it is important to consider the
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54 context in which the framework was derived and validated. The participants in the OCS study
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56 possessed similar characteristics to those involved in the qualitative study from which the
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framework was originally developed. The majority of participants in both studies were males, in their 40s, taking combination antiretroviral therapy, who were not currently working. Similarity among the sample participants confirms the types of individuals with whom we validated the dimensions of disability in the *Episodic Disability Framework* and strengthens the validity of this *Framework* for use with adults living with HIV with these characteristics. Generalizability of these findings to the broader HIV population should be carefully considered given the OCS sample may under-represent women (17% in OCS sample versus 22% in Canada) and Aboriginals (2% versus 8%) living with HIV in Canada [54].

Implications for Practice, Research and Policy

The *Episodic Disability Framework* has potential practical implications for clinicians, adults living with HIV, researchers and policy makers. Clinicians can use this *Framework* to frame their assessments of disability to include physical and mental health symptoms impairments as well as consider a patient’s ability to participate in society. Adults living with HIV may use the *Framework* to articulate their health-related challenges to care providers. Researchers may use the *Episodic Disability Framework* to guide the development of a future HIV disability questionnaire. Finally, although we did not empirically assess the episodic nature of disability in this study, policy makers might consider using the *Framework* to recognize the episodic way adults living with HIV might experience disability, which could lead to more flexible income and labor force policies and programs.

Study Limitations

Our study has several limitations. Results are limited to the type and quality of measured variables available and the characteristics of participants who completed the OCS questionnaire. Perhaps because participants had to be well enough to complete a 90 to 120 minute interview,

the spectrum of respondents reporting poor health was limited. Furthermore, all respondents were residents of the Greater Toronto Area, Ontario and the majority was taking antiretroviral medications. Generalization to other individuals and settings remains to be determined. Second, ceiling effects of indicator variables and the absence of uncertainty, a key dimension of disability limited our ability to validate the *Episodic Disability Framework*. Future research should include careful consideration of the distribution of variables prior to their inclusion in the measurement model and include measures of uncertainty to validate this dimension of the *Framework*. Third, a cross-sectional analysis such as this is unable to validate the episodic nature of disability. Longitudinally exploring the episodic nature of disability experienced by adults living with HIV is vitally important, given the importance of time in our *Framework*. Fourth, while our results indicate that dimensions of disability are correlated, the direct relationships between the dimensions are unknown. Our results provide a measurement model which can serve as the foundation for future structural equation model analyses to determine the relationships between the dimensions of disability and whether they differ based on personal or disease characteristics for adults living with HIV. Next, this study specifically focused on validating the first component of the *Episodic Disability Framework*, dimensions of disability. Further empirical testing is needed to validate the contextual factor and trigger components of the *Framework* and determine their relationships with dimensions of disability. Finally, our aim was to validate dimensions of disability within the *Episodic Disability Framework*. We used available data from a rich source of socio-behavioral variables collected in an observational cohort study with adults living with HIV. However, given existing measures in the OCS questionnaire were neither developed, nor validated for the purpose of describing disability, future research will involve the development of a new HIV-specific measure of disability using

content from the newly validated dimensions of disability in the *Framework*.

CONCLUSIONS

Results from this confirmatory factor analysis support the validity of four dimensions of disability experienced by adults living with HIV in the *Episodic Disability Framework*. This is the first known conceptual framework of disability developed from the perspective of adults living with HIV. The *Episodic Disability Framework* can be used to help describe the multi-dimensional nature of disability experienced by adults living with HIV and can lay the foundation for developing a future HIV disability measure for clinical and health services research.

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*The OHTN Cohort Study Research Team consists of Dr. Sean B. Rourke (Principal Investigator, University of Toronto and OHTN), Dr. Ann Burchell (Co-Principal Investigator, OHTN), Dr. Sandra Gardner (OHTN), Dr. Sergio Rueda (OHTN), Dr. Ahmed Bayoumi and Dr. Kevin Gough, St. Michael’s Hospital; Dr. Jeffrey Cohen, Windsor Regional Hospital; Dr. Curtis Cooper, Ottawa General Hospital; Dr. Don Kilby, University of Ottawa Health Services; Dr. Mona Loutfy and Dr. Fred Crouzat,, Maple Leaf Medical Clinic; Dr. Anita Rachlis and Dr. Nicole Mittmann, Sunnybrook Health Sciences Centre; Dr. Janet Raboud and Dr. Irving Salit, Toronto General Hospital; Dr. Edward Ralph, St. Joseph’s Health Care; Dr. Roger Sandre,

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For Peer Review

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Table 1: Characteristics of Participants (n=913)

Characteristic	Number (%)
<i>Gender</i>	
Male	757 (83%)
Female	153 (17%)
Other*	3 (<1%)
Age (median; IQR)	47 years (41-53 years)
Length of Time Since Diagnosis (median; IQR)	11 years (6-17 years)
Taking Antiretrovirals	783 (86%)
<i>Race</i>	
White	588 (64%)
Black / African	158 (17%)
Asian / Latin American / Arab	101 (11%)
Aboriginal	22 (2%)
<i>Employment Status</i>	
Employed (full time or part time)	412 (45%)
On disability leave from work	289 (32%)
Student or retired	106 (12%)
Unemployed	79 (9%)
Volunteering	25 (3%)
<i>Educational Level</i>	
No formal education or less than grade 9	32 (3%)
Partially or fully completed high school education	268 (29%)
Partially or fully completed trade or technical training	31 (3%)
Partially or fully completed college	234 (26%)
Partial or fully completed university	279 (31%)
Post-graduate education	69 (8%)
<i>Gross Personal Income (Canadian dollars)</i>	
Less than \$20,000 per year	386 (43%)
\$20,000 to less than \$40,000 per year	214 (24%)
\$40,000 to less than \$60,000 per year	140 (16%)
\$60,000 to less than \$80,000 per year	70 (8%)
\$80,000 or more per year	85 (9%)

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IQR=interquartile range. Not all characteristics will add to the total n due to missing responses.

*Other=trans, inter-sexed or other.

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Table 2: Description of the OCS Indicator Variables Included in the Initial Hypothesized Measurement Model (Model 1) (n=913)

Dimension of Disability (Latent Variable)	Measures in OCS Questionnaire (Indicator Variables)	Type of Variable (range)	Mean (SD)	% Missing Responses
Physical Symptoms and Impairments (25 OCS variables)	HIV SI – Loss of Energy	Discrete - Ordered Categorical (0-4)	1.78 (1.56)	0.00
	HIV SI – Chills or Sweats	Discrete - Ordered Categorical (0-4)	0.81 (1.31)	0.00
	HIV SI – Feeling Dizzy	Discrete - Ordered Categorical (0-4)	0.82 (1.30)	0.22
	HIV SI - Pain	Discrete - Ordered Categorical (0-4)	1.16 (1.51)	0.00
	HIV SI – Nausea	Discrete - Ordered Categorical (0-4)	0.55 (1.05)	0.11
	HIV SI – Diarrhea	Discrete - Ordered Categorical (0-4)	1.11 (1.47)	0.11
	HIV SI – Difficulty Sleeping	Discrete - Ordered Categorical (0-4)	1.62 (1.64)	0.22
	HIV SI – Skin Problems	Discrete - Ordered Categorical (0-4)	0.89 (1.34)	0.11
	HIV SI – Cough	Discrete - Ordered Categorical (0-4)	0.72 (1.28)	0.11
	HIV SI – Headache	Discrete - Ordered Categorical (0-4)	0.83 (1.32)	0.00
	HIV SI – Loss of Appetite	Discrete - Ordered Categorical (0-4)	0.67 (1.27)	0.22
	HIV SI – Bloating	Discrete - Ordered Categorical (0-4)	1.18 (1.49)	0.11
	HIV SI – Muscle Aches	Discrete - Ordered Categorical (0-4)	1.48 (1.59)	0.11
	HIV SI – Problems with Sex	Discrete - Ordered Categorical (0-4)	1.06 (1.49)	3.50
	HIV SI – Body	Discrete - Ordered Categorical (0-4)	0.99 (1.50)	0.55
	HIV SI – Weight	Discrete - Ordered Categorical (0-4)	0.55 (1.18)	0.11
	HIV SI – Hair Loss	Discrete - Ordered Categorical (0-4)	0.44 (1.09)	0.11
	SF-36 Bodily Pain Subscale Score+	Continuous (0-100)	50.96 (11.62)	0.00
	SF-36 Vitality Subscale Score+	Continuous (0-100)	51.12 (11.94)	0.00
	EQ-5D Pain	Discrete - Ordered Categorical (1-3)	1.57 (0.59)	0.00
	HUI Vision Subscale Score+	Continuous (0-1)	0.01 (0.02)	1.31
	HUI Hearing Subscale Score+	Continuous (0-1)	0.003 (0.02)	1.10
	HUI Speech Subscale Score+	Continuous (0-1)	0.003 (0.02)	0.11
	HUI Dexterity Subscale Score+	Continuous (0-1)	0.003 (0.02)	0.00
	HUI Pain Subscale Score+	Continuous (0-1)	0.10 (0.13)	0.00
Mental Health Symptoms and Impairments (11 OCS variables)	HIV SI – Trouble Remembering	Discrete - Ordered Categorical (0-4)	1.28 (1.49)	0.22
	HIV SI – Felt Sad	Discrete - Ordered Categorical (0-4)	1.68 (1.60)	0.22
	HIV SI – Felt Nervous	Discrete - Ordered Categorical (0-4)	1.38 (1.50)	0.33
	SF-36 Role Emotional Subscale Score+	Continuous (0-100)	54.56 (11.85)	0.00

Difficulties with Day-to-Day Activities (8 OCS variables)	SF-36 Mental Health Subscale Score+	Continuous (0-100)	53.67 (12.34)	0.00
	MOS Cognitive Function Scale Score+	Continuous (0-100)	23.67 (22.96)	0.88
	EQ-5D Anxiety and Depression	Discrete - Ordered Categorical (1-3)	1.56 (0.61)	0.22
	CES-D Total Scale Score	Continuous (0-60)	13.38 (12.12)	3.40
	Ongoing Problems Stress Index Scale Score	Continuous (0-17)	3.45 (2.68)	0.55
	HUI Emotion Subscale Score+	Continuous (0-1)	0.06 (0.11)	1.64
	HUI Cognition Subscale Score+	Continuous (0-1)	0.07 (0.11)	0.00
	SF-36 Physical Function Subscale Score+	Continuous (0-100)	50.38 (9.50)	0.00
	SF-36 Role Physical Function Subscale Score+	Continuous (0-100)	52.38 (10.40)	0.22
	EQ-5D Mobility	Discrete - Ordered Categorical (1-3)	1.23 (0.42)	0.00
Challenges to Social Inclusion (13 OCS variables)	EQ-5D Self Care	Discrete - Ordered Categorical (1-3)	1.05 (0.21)	0.00
	EQ-5D Usual Activities	Discrete - Ordered Categorical (1-3)	1.28 (0.47)	0.00
	HUI Mobility Subscale Score+	Continuous (0-1)	0.01 (0.02)	1.80
	HUI Self Care Subscale Score+	Continuous (0-1)	0.001 (0.02)	0.00
	HUI Ambulation Subscale Score+	Continuous (0-1)	0.003 (0.02)	1.80
	Dwelling	Discrete - Ordered Categorical (1-2)	1.04 (0.18)	0.22
	Belonging in Neighbourhood	Continuous (1-5)	2.13 (1.07)	0.55
	Home Good Location to Live	Continuous (1-5)	1.92 (0.97)	0.33
	Control at Home	Continuous (1-5)	1.96 (0.92)	0.44
	Employment Status	Discrete - Ordered Categorical (0-4)	1.47 (1.52)	0.22
HIV SI = HIV Symptom Index [26]; SF-36=SF-36 questionnaire [21]; MOS=Medical Outcome Study (Cognitive Functioning Scale) [29], EQ-5D=EuroQol (EQ-5D) questionnaire [27], Ongoing Problems Stress Index [28]; CES-D=Center for Epidemiologic Studies Depression Scale [23]; HUI=Health Utilities Index [22]; + indicate reverse scoring of variables so that higher scores indicate greater disability.	Difficulty with Housing Costs+	Discrete - Ordered Categorical (1-4)	1.96 (1.07)	0.66
	Worry about Moving out of Home+	Continuous (1-5)	2.20 (1.18)	0.44
	Educational Level+	Discrete - Ordered Categorical (1-10)	4.88 (2.81)	0.00
	Gross Personal Income+	Discrete - Ordered Categorical (1-10)	7.06 (2.58)	1.97
	Gross Income+	Discrete - Ordered Categorical (1-10)	5.80 (3.08)	5.15
	Sources of Income	Discrete - Ordered Categorical (0-2)	0.52 (0.51)	4.16
	Occupational Category	Discrete - Ordered Categorical (1-7)	4.31 (2.02)	1.53
	SF-36 Social Function Subscale Score+	Continuous (0-100)	55.52 (12.37)	0.22
	HIV SI = HIV Symptom Index [26]; SF-36=SF-36 questionnaire [21]; MOS=Medical Outcome Study (Cognitive Functioning Scale) [29], EQ-5D=EuroQol (EQ-5D) questionnaire [27], Ongoing Problems Stress Index [28]; CES-D=Center for Epidemiologic Studies Depression Scale [23]; HUI=Health Utilities Index [22]; + indicate reverse scoring of variables so that higher scores indicate greater disability.			
	Any variables not identified as subscale or summary scores were individual item scores used in the analysis (n=37).			

Table 3: Comparison of Model Fit

	χ^2 (df)	CFI	TLI	RMSEA	Model comparison	$\Delta\chi^2$ (df)	P-value
Extended Models with all Possible OCS variables (n=57 variables)							
Model 1: Original hypothesized model with all possible OCS variables (n=57) that represent <u>four</u> latent variable dimensions of disability in the Episodic Disability Framework (see Table 1 for indicator variables)	6304.734 (1533)	0.856	0.850	0.058	--	--	--
Model 2: Model with all possible OCS variables (n=57) and <u>three</u> latent variable dimensions of disability collapsing mental health and physical health symptoms into one latent variable.	6990.740 (1536)	0.836	0.829	0.062	M2 vs. M1	302.137 (3)	<0.001*
Simplified Models (n=43 variables)							
Model 3: Simplified model with 43 variables and <u>four</u> latent variable dimensions of disability.	3077.748 (854)	0.889	0.883	0.053	--	--	--
Model 4: Simplified model with 43 variables and <u>three</u> latent variable dimensions of disability.	3762.014 (857)	0.856	0.848	0.061	M4 vs. M3	291.535 (3)	<0.001*
Cross-Loading Social Function Subscale (n=43 variables)							
Model 5: Cross loaded social function subscale of the SF36 onto physical health symptoms and mental health symptoms and challenges to social inclusion.	2626.175 (852)	0.912	0.907	0.048	--	--	--
Model 6: Cross loaded social function subscale of the SF36 onto physical health symptoms and mental health symptoms. **Final Model**	2621.497 (853)	0.912	0.907	0.048	M6 vs. M5	0.258 (1)	0.6118

χ^2 = chi square statistic; df=degrees of freedom; CFI = Comparative Fit Index; TLI= Tucker-Lewis Index; RMSEA = Root Mean Square Error of Approximation;

$\Delta\chi^2$ = chi square difference test statistic; *statistically significant difference in model fit (p<0.05).

Table 4: Final Model (Model 6) – Dimensions of Disability in the Episodic Disability Framework (n=913)

Dimension of Disability (Latent Variables)	Measures in OCS Questionnaire (Indicator Variables)	Standardized Factor Loadings #	95% Confidence Interval
Physical Symptoms and Impairments (n=21 variables)	HIV SI – Loss of Energy	0.78	0.74-0.81
	HIV SI – Chills or Sweats	0.64	0.58-0.69
	HIV SI – Feeling Dizzy	0.68	0.63-0.73
	HIV SI – Pain	0.69	0.64-0.73
	HIV SI – Nausea	0.63	0.57-0.69
	HIV SI – Diarrhea	0.50	0.44-0.56
	HIV SI – Difficulty Sleeping	0.65	0.60-0.70
	HIV SI – Skin Problems	0.51	0.44-0.57
	HIV SI – Cough	0.58	0.52-0.64
	HIV SI – Headache	0.59	0.53-0.65
	HIV SI – Loss of Appetite	0.64	0.58-0.69
	HIV SI – Bloating	0.59	0.54-0.65
	HIV SI – Muscle Aches	0.75	0.71-0.79
	HIV SI – Problems Sex	0.50	0.43-0.56
	HIV SI – Body	0.35	0.28-0.43
	HIV SI – Weight	0.41	0.33-0.48
	HIV SI – Hair Loss	0.47	0.39-0.55
Mental Health Symptoms and Impairments (n=10 variables)	SF-36 Bodily Pain Subscale Score	0.74	0.70-0.78
	SF-36 Vitality Subscale Score	0.82	0.79-0.85
	SF-36 Social Function Subscale Score*	0.21	0.13-0.28
	EQ-5D Pain	0.77	0.73-0.81
	HIV SI – Trouble Remembering	0.67	0.61-0.72
	HIV SI – Felt Sad	0.80	0.76-0.83
	HIV SI – Felt Nervous	0.75	0.71-0.79
	SF-36 Role Emotional Subscale Score	0.78	0.74-0.81
	SF-36 Mental Health Subscale Score	0.82	0.79-0.84
	SF-36 Social Function Subscale Score*	0.61	0.54-0.68

	MOS Cognitive Function Score	0.74	0.71-0.78
	EQ-5D Anxiety and Depression Score	0.79	0.76-0.83
	CES-D Total Score	0.86	0.84-0.89
	Ongoing Problems Stress Index	0.54	0.49-0.60
Difficulties with Day-to-Day Activities (n=5 variables)	SF-36 Physical Function Subscale Score	0.73	0.69-0.76
	SF-36 Role Physical Function Subscale Score	0.86	0.81-0.90
	EQ-5D Mobility	0.76	0.71-0.82
	EQ-5D Self Care	0.61	0.50-0.72
	EQ-5D Usual Activities	0.89	0.85-0.94
Challenges to Social Inclusion (n=8 variables)	Dwelling	0.41	0.21-0.61
	Belonging in Neighbourhood	0.48	0.40-0.55
	Home Good Location to Live	0.50	0.43-0.57
	Control at Home	0.57	0.50-0.64
	Employment Status	0.62	0.54-0.70
	Difficulty with Housing Costs	0.65	0.57-0.73
	Worry about Moving out of Home	0.53	0.45-0.61
	Gross Personal Income	0.59	0.52-0.66

Measurement Model Results

χ^2 = Chi-square (χ^2)=2621.50 (p value<0.0001)

Degrees of freedom (df) =853;

Comparative Fit Index (CFI) =0.912 (ideal is $\geq$ 0.95)

Tucker-Lewis Index (TLI) =0.907 (ideal is $\geq$ 0.95)

Root Mean Square Error of Approximation (RMSEA) =0.048 (good fit indicated by <0.05)

Legend

HIV SI=HIV Symptom Index [26]; SF-36=SF-36 questionnaire [21]; MOS=Medical Outcomes Study (Cognitive Functioning Scale) [31]; Ongoing Problems Stress Index [28]; EQ-5D=EuroQol (EQ-5D) questionnaire [27]; CES-D=Center for Epidemiologic Studies Depression Scale [23];

All standardized parameter estimates were statistically significant ($p < 0.0001$);

*Indicates cross loading of SF-36 social function Subscale score on two latent variables;

Factor loadings ≥ 0.30 are bolded which indicate the variable 'loads' on a given dimension of disability.

Figure 2: Correlations between Dimensions of Disability (Latent Variables) in the Final Confirmatory Factor Analysis Solution (Model 6)