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An Observational Study of All-cause Mortality among People with HIV Released from an Integrated System of Jails and Prisons, 2007-2014

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Summary

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KBL obtained funding for the project and designed the study, with guidance from FLA and JPM. KBL conducted all data analyses with guidance from and data interpretation by FLA, MMD, MMC, CG, and JPM. KR and DC conducted all SMR data analyses. KBL drafted primary versions of the manuscript, with input from FLA, MMD, MMC, CG, and JPM, KR, and DC.All authors have contributed significantly to the work and reviewed and approved the final version for submission.

Declaration of interests

The authors have no conflicts of interest to disclose.

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Background: People transitioning from prisons or jails have high mortality but data are limited for people living with HIV (PLWH) and no studies have integrated data from both criminal justice and community settings. We aimed to evaluate all-cause mortality among PLWH released from Connecticut's integrated system of prisons and jails.

Methods: We linked pharmacy, custodial, death, case management, and HIV surveillance data (2007-2014) from Connecticut Departments of Corrections and Public Health to create a retrospective cohort. We compared the mortality rate to statewide and national populations, and described and modeled time-to-death from any cause after prison-release using Cox proportional hazards.

Findings: Among 1,350 PLWH, mostly Black or Hispanic men with median age 45 years (IQR 39-50), 184 (184/1350, 13.6%) died during a median 5.2 years (IQR 3.0-6.7) after index release. The crude mortality rate of 2,868/100,000 person-years was 6.97 and 8.47 times higher (by standardized mortality ratio) than that of the general U.S. and statewide populations, respectively. Main reported causes of death were: HIV/AIDS (78/170, 45.9%), drug overdose (26/170, 15.3%), liver-related (17/170,10.0%), cardiovascular disease (16/170, 9.4%), and accidental injury or suicide (13/170, 7.6%). Protective factors for time-to-death were: Black race (adjusted hazard ratio [aHR]=0.52, 95%CI=0.34-0.80), having health insurance (aHR=0.09, 95%CI=0.05-0.17), having

1 long re-incarceration (aHR=0·41, 95%CI=0·22-0·76), and having an increasing proportion of re-incarcerations in which ART was prescribed. Positive predictors of time-to-death were: age 50 years (aHR=3·65, 95%CI=1·21-11·08), lower CD4 count (200-499 cells/ML: aHR=2 μ 54, 95%CI=1·50-4·31; <200 cells/ μ L: aHR=3·44, 95%CI=1·90-6·20), higher medical co-morbidity (aHR=1·86, 95%CI=1·23-2·82), virologic failure before death (aHR=2·76, 95%CI=1·94-3·92), and lacking viral load monitoring (aHR=2·13, 95%CI=1·09-4·18).

Interpretation: To reduce post-release mortality among PLWH, resources are needed to recognize and treat HIV as well as comorbid medical, psychiatric, and substance use disorders, during and following incarceration. Policies that reduce incarceration and support integrated systems of care between prisons and communities can have a significant impact on the survival of PLWH.

Introduction

The United States (US) has the highest incarceration rate worldwide,¹ with at least half of prisoners incarcerated for drug-related offenses, and a disproportionate prevalence of HIV in prisons and jails.² Within-prison HIV-related mortality has declined,³ largely due to expanded antiretroviral therapy (ART) for universal treatment. Nearly all people return to communities following prison-release,⁴ wherein challenges to HIV care continuity and relapse to substance use are well-documented.⁵⁻⁷ Release from prisons is associated with immediate and exceedingly high rates of death,^{8,9} importantly from opioid overdose and liver disease, including Hepatitis C;¹⁰⁻¹² however, data that incorporate both prison and jail release have not been examined.

There are limited data on causes and predictors of post-release mortality among PLWH to inform policies and services. Two small studies from Indonesia and French Guiana found a higher risk of mortality for PLWH returning from prisons compared to the general

population;¹³ most deaths were HIV-related.¹⁴ Causes of death in PLWH can be multifactorial and structural factors, including re-incarceration, may mitigate mortality risk.¹³ Among non-incarcerated PLWH, deaths are more often attributable to non-HIV-related causes, but it is unclear if this pattern holds for incarcerated PLWH.¹⁵

To inform interventions that reduce mortality after release, we previously developed a large cohort study in a resource-rich setting to describe post-release risk of death from all causes among PLWH released from prisons to communities. This study links data from an integrated system that includes both prisons and jails with community-based data to comprehensively assess the complex role that individuals' characteristics and incarceration experiences play in HIV outcomes and risk of death.

Methods

Study design and population

This study took place in Connecticut, a state in the Northeastern United States. The study was conducted in the Connecticut Department of Correction (CTDOC), an integrated system that includes prisons and jails, which were previously described.¹⁶ The CTDOC Research Advisory Committee and institutional review boards at Yale University and Connecticut Department of Public Health (CTDPH) approved all procedures. Participant consent was waived because all data was previously collected and de-identified for analysis.

Data sources

CTDOC custody and pharmacy data were combined with the CTDPH enhanced HIV/AIDS Reporting System (eHARS) surveillance database and CAREWare service utilization database for analyses. We previously created this merged database and examined linkage to care following incarceration (yccr.yale.edu).¹⁶ eHARS includes all reported dates of deaths through December 31, 2014, cross-checked against the National Death Index by July 2016 to capture out-of-state deaths. Primary and secondary causes of death were available for deaths occurring in Connecticut. Individuals included in our cohort (Figure 1) were: 1) adults 18 years old with confirmed HIV; 2) included in all administrative databases; 3) incarcerated in Connecticut at least once for >24 hours; and 4) were admitted and released between January 1, 2007 and December 31, 2014. Each individual's first incarceration during this period was their index incarceration. Individuals were followed from index release date until date of death or censoring on December 31, 2014.

We used 2007-2014 contemporaneous death and person-time data from the Centers for Disease Control and Prevention's Wide-ranging Online Data for Epidemiologic Research (CDC WONDER) to calculate standardized mortality ratios (SMRs) and indirect adjusted mortality rates (IARs) compared to the general population.¹⁷

Outcomes and covariates

The primary outcome was time-to-death from any cause. We described all causes of death, categorized by ICD-10 codes (Appendix page 2). Primary causes of death were further categorized as "HIV/AIDS-related"; "Drug overdose"; "Liver disease, failure, and/or viral

hepatitis"; "Cardiovascular disease"; "Accidental injury, suicide, or homicide"; and "Other causes" (Appendix page 5.) Many of the non-HIV/AIDS-related deaths were secondarily attributed to HIV.

The Behavioral Health Model for Vulnerable Populations provides a framework for understanding modifiable risk factors and enabling resources to inform interventions.¹⁸ We used this model to categorize covariates and hypothesize the role of each in post-release mortality:

<u>Predisposing factors</u> included demographic characteristics. CTDPH data defined the primary source of HIV transmission, with injection drug use representing people who inject drugs (PWID), and CD4 count at the time of HIV diagnosis.

Enabling/disabling resource factors from the CAREWare database included health insurance coverage and transitional case management (TCM) use during the index incarceration (dichotomous) and quantity of TCM visits per year. HIV viral load (VL) from the eHARS database served as a proxy for routine HIV care clinic visits (in prison/jail and in the community). Linkage to HIV care required a VL within 14 days after the index release, as we described previously. Using CTDOC data, we calculated duration of incarcerations with <30 days likely representing jail detentions and 365 days involving prison sentences. Conditions of release were categorized as unsupervised, conditional (e.g., parole or transitional housing), or bonded. Incarceration variables were constructed both for the index incarceration and the most recent incarceration prior to death or censoring. Because HIV care during incarceration was expected to influence risk of death, the "Conditions of release from last incarceration" variable included an additional category for individuals who were re-incarcerated for >30 days by the time of death/censoring. Re-incarceration was otherwise defined as spending >24 hours in a CTDOC facility after index release and analyzed as a continuous rate (number of re-incarcerations per year of follow-up), dichotomously (having at least one re-incarceration lasting 365 days), and categorically (percentage of follow-up time spent re-incarcerated: 0-1%, 2-10%, 11-25%, 26-50%, and 51-100%).

<u>Need factors:</u> Pre-release viral suppression (VL<400 copies/μL) was defined within 90 days before release. Pharmacy records included antiretroviral (ART) and medications prescribed to treat other medical, psychiatric, or substance use co-morbidities; each were coded dichotomously and summed to create a categorical co-morbidity variable (other than HIV). As previously described,¹⁶ CTDOC staff assign psychiatric and addiction severity scores on intake (scale 1-5) to determine services needed during and immediately after incarceration. We analyzed addiction severity scores categorically.We combined psychiatric severity scores with prescription data to create a 4-level composite psychiatric severity variable (1: low, untreated; 2: low, treated; 3: high, untreated; 4: high, treated).

For each re-incarceration, we used pharmacy data to determine whether ART was prescribed. We also used the first and last VL measured during incarceration to determine whether individuals with an initially high VL became virally suppressed during that incarceration and whether suppression was achieved within 90 days pre-release. We calculated the percentage of all re-incarcerations that demonstrated these outcomes and

categorized "ART coverage" by quartiles. Finally, using national guidelines,¹⁹ we used the last VL drawn during the last six months of observation to evaluate viral suppression status and frequency of monitoring.

Statistical analysis

We first calculated the crude mortality rate (CMR). We used indirect standardization to calculate indirect adjusted mortality rates (IARs) and standardized mortality ratios (SMRs) to compare mortality in the cohort with that of the US and Connecticut general populations. Both IARs and SMRs were adjusted for sex, race/ethnicity, and age in years. We calculated basic descriptive frequencies of primary and all causes of death, stratified by primary cause (Appendix page 5). Using Kaplan-Meier curves and log-rank tests, we compared time-todeath during the first year after release by primary causes of death among those who died (Figure 2), and assessed predictors of time-to-death in bivariate analyses (Figure 3) and overall for the entire sample (Appendix page 1). We used Cox proportional hazards to model predictors of time-to-death from any cause to globally assess the impact of release on timeto-death. We incorporated data across multiple incarcerations during follow-up to account for whether an individual tended to be re-incarcerated and engaged in care by virtue of their re-incarceration. We explored the model using both continuous and categorical or calculated variables and only included the latter when continuous terms were not normally distributed. Variables with bivariate associations of p < 0.20 were considered for multivariate models and we generated preliminary parsimonious models with all variables with Type III p < 0.10. Rather than rely on automated selection procedures, we explored relationships between variables with suspected correlation, using chi-squared and Mann-Whitney U tests. When variables were collinear, we selected the variable of greater clinical interest based on prior literature and variable distribution, and factored in the strength of the association with the outcome, using thresholds of significance only to understand how these variables impacted survival to finalize the model. We formally tested each variable for adherence to the proportional hazards assumption. Variables that violated the proportional hazards assumption were included as time-varying covariates in the final model. The effect estimates presented in Table 1 correspond to the first day of follow-up ($t_{C}=0$ days); time-dependent effect estimates were also calculated for discrete time points (t_1 =30 days, t_2 =365 days, t_3 =730 days [2 years], and t_3 =1460 days [4 years]) but the effect estimates did not change direction or significance over time. We used cumulative hazard plots and residual plots as supplementary methods to confirm that this strategy resulted in a robust model. We hypothesized and therefore explored interactions between recidivism, TCM, PWID, race/ ethnicity, and gender. Interaction terms were justified because they provided a clearer explanation of variable effects compared to models that did not include interaction terms. Given Connecticut's high recidivism rate (37-64% return within three years),²⁰ we focused on re-incarceration rates of 0, 0.33, 1, 2, and 3 reincarcerations/year when assessing interactions. Using backward selection, the final parsimonious multivariable model was limited to variables with Type III p < 0.5. Using re-incarceration rates as a covariate may have unintentionally introduced bias, in that people who died early after release from prison had less chance to be re-incarcerated. We therefore conducted a sensitivity analysis that excluded the earliest deaths occurring within 21 days following release. All analyses were performed using SAS version 9.4 (SAS Institute Inc., Gary, NC).

Role of the funding source

The National Institutes of Health played no role in data collection, analysis, data interpretation, drafting of or decision to submit the manuscript for publication.

Results

Table 1 describes the 1,350 PLWH eligible for analysis between January 1, 2007 and December 31, 2014, with most being 40 years old (983/1350, 72.8%), male (1043/1350, 77.3%), Black or Hispanic (1052/1350, 78.0%), and PWID (943/1350, 69.9%).

The sample was followed for a median of $5 \cdot 2$ (IQR $3 \cdot 0 \cdot 6 \cdot 7$) years, contributing a total of 6,418 person-years. After index release, 184 (184/1350, 13 \cdot 6%) individuals died, resulting in a CMR of 2,868 per 100,000 person-years; the IAR was 2,993 per 100,000 person-years (95% CI=2,560-3,425) and 2,964 per 100,000 person years (95% CI=2,536-3,392), compared to the general US and Connecticut populations, respectively. Compared to the general US and Connecticut populations, the SMRs were 6.97 (95% CI=5.96-7.97) and 8.47 (95% CI=7.25-9.69), respectively.

Among the 170 deaths where cause was reported, primary causes included HIV/AIDS (78/170, 45.9%); drug overdose (n=26, 15.3%); accidental injury, homicide, or suicide (13/170, 7.6%); liver (17/170, 10.0%); and cardiovascular (16/170, 9.4%) disease (Appendix page 5). During the first year after index (Figure 2A) or most recent release (Figure 2B), among those who died, time-to-death from drug overdose or accidental injury was significantly shorter than time-to-death from HIV/AIDS complications. Survival was longest for individuals who died from liver or cardiovascular disease. When survival probabilities were assessed after 24-36 months after release, there was no statistically significant difference in time-to-death by primary cause.

The final multivariable Cox model (Table 1) included 1,319 individuals with all data available, who cumulatively contributed 6,276 person-years of follow-up time, during which there were 182 deaths.

<u>Predisposing factors</u> that independently and significantly predicted shorter survival time were age 50 years and lower CD4 count at diagnosis (Figure 3A), despite people having been diagnosed with HIV a median 11·4 years prior to index release. Relative to being White, Black race was protective against time-to-death. In bivariate analysis, PWID had significantly shorter survival (Figure 3B). In the multivariable model, PWID significantly interacted with re-incarceration rate, such that, compared to non-PWID, PWID had worse survival when they were never or infrequently re-incarcerated and better survival when frequently re-incarcerated.

<u>Enabling resource factors</u> that independently predicted longer survival time were having at least one long re-incarceration and having health insurance either consistently or intermittently during follow-up. Health insurance's effect varied over time but did not change direction or significance. Accounting for significant interactions between frequency of TCM visits, re-incarceration rate, and PWID, the following patterns were observed:

compared to those who never received TCM services after release, PLWH using 5 TCM visits/year had worse survival when they were never or infrequently reincarcerated and better survival when frequently re-incarcerated. Increasing reincarceration rate, however was highly hazardous, particularly for PWID and those who either had infrequent (0-2 visits/ year) or extremely frequent (5 visits/year) TCM. The highest hazard of death observed for a 1-unit increase in re-incarceration rate was in non-PWID who did not receive any TCM services. The lowest relative hazard associated with increasing re-incarceration rate was in PWID who received 5 TCM visits/year. Only three deaths (3/182, 1.6% of all deaths included in the final model), occurred within 21 days of release when preventable causes of death including overdose are common,²¹ and there was minimal impact on effect magnitude or direction when these three deaths were removed from the model in a sensitivity analysis.

<u>Need factors:</u> Compared to individuals who were never re-incarcerated, having a higher proportion of re-incarcerations in which ART was prescribed was protective. In contrast, PLWH with 2 other medical co-morbidities, documented virologic failure during follow-up, or who never had a VL drawn in the community after release had significantly shorter survival time.

Discussion

By integrating correctional, public health and mortality data for a large number of PLWH over an 8-year observation period where both prison and jail data were available, we were able to examine the complex contributions to death in prisoners with HIV who transition to the community. We found a high CMR after release (2,868/100,000 person-years), which was higher than prior estimates of general prison populations (720-2,054/100,000 person-years).¹¹ When we adjusted for race, ethnicity, sex, and age, we also found a high SMR compared to US (6·97) and Connecticut general populations(8·47). This was greater than the SMR of 3.6 reported in a study of the general prison compared to the non-institutionalized population in Washington.⁸

To our knowledge, this represents the largest cohort of PLWH from an integrated correctional system that incorporates both jail and prison data. Here, less than half of deaths were HIV/AIDS-related; however, like in non-incarcerated PLWH, a substantial proportion were due to drug overdose and accidents, liver, and cardiovascular disease.¹⁵ Consistent with findings from prisoners, including those without HIV, deaths due to drug overdose and accidents occurred sooner than deaths due to HIV/AIDS; liver and cardiovascular disease occurred later.^{8,12,22} Coercive mobility between prisons and communities elevates the risk of death in PLWH for a number of conditions, including HIV, cardiovascular and liver disease, and substance use disorders (SUDs).

Late HIV diagnosis predisposes PLWH to increased morbidity and mortality.²³ Here, PLWH had been diagnosed for a median of 11.4 years, yet low CD4 count at diagnosis still significantly predicted mortality, perhaps related to discontinuous ART resulting in CD4 returning to its nadir. Similarly, advanced HIV disease during follow-up and medical co-morbidities also predicted post-release mortality, indicating an unmet need for medical care. Despite high levels of ART provision within CTDOC facilities, which previously has been

Health insurance, including coverage of ART medication costs, is associated with better healthcare utilization, viral suppression, and lower mortality.^{25,26} Many states like Connecticut have recently expanded Medicaid programs, which include efforts to enroll patients from the CJ system and to suspend (rather than terminate) Medicaid benefits during incarceration. Nevertheless, at-risk, low-income PLWH and those with other comorbidities continue to face barriers to gaining adequate health insurance. In this cohort, PLWH who had health insurance after release had longer survival times. Yet this setting likely represents the most optimistic scenario where health insurance coverage was high. It is therefore likely that, in the absence of healthcare insurance expansion, post-release mortality rates would be markedly higher.

At least 70% of this cohort were PWID, which is associated with risk of death from drug overdose, poor retention in HIV care, and all-cause mortality.⁵ Longer incarceration increases the likelihood of achieving viral suppression,²⁷ and may also provide sufficient time for initiating treatment for opioid use disorder. Despite evidence that pharmacological treatment for opioid use disorder initiated during incarceration and continued post-release reduces substance use, improves HIV treatment outcomes, and reduces mortality after release,^{24,28} implementation of such strategies globally and in the US are uncommon, including in this study setting. Longer incarceration periods, however, allow sufficient time to provide TCM and address post-release barriers to care like re-activating health insurance and linking to community-based resources including addiction treatment programs. Although recidivism is medically and socially destabilizing, people who are frequently reincarcerated, especially PWID, may be prioritized for transitional care coordination. Consequently, while PWID had reduced survival, higher re-incarceration rates mitigated this effect. The nation's volatile opioid epidemic and related policies that concentrate PWID in CJ systems and contribute to their poor health outcomes²⁹ speak to the need to adequately provide opioid agonist treatments (OAT) in both CJ and community-based healthcare systems.³⁰ OAT reduces drug overdose in both community and CJ settings. Yet, where OAT coverage is absent or under-scaled, drug-related death after release will remain unabated, including in PLWH. Other post-release strategies to reduce the negative consequences of drug use include providing naloxone for overdose prevention, sterile injecting equipment, and screening and treatment for psychiatric and SUDs.

Our finding that Black race was protective against time-to-death among PLWH echo findings in sentenced prisoners elsewhere.¹⁰ Potential explanations for this finding include the disparately protective effect of longer incarcerations, though time spent reincarcerated during follow-up did not attenuate the protective effect of Black race. Alternatively, Black individuals during this time period were at reduced risk for opioid-related overdoses, or

more likely, show greater resilience in the post-release period that could provide some protection against mortality. 31

Several enabling resources and need factors contributed to mortality. Most importantly, TCM, which has been associated with improved linkage to and retention in care in released prisoners with HIV,^{16,32} also reduced mortality. While only 25·2% received TCM services during their index incarceration, the interaction with re-incarceration and PWID suggests a channeling of services by targeting the highest risk individuals for TCM. Though not measured here, receipt of TCM, especially for re-incarcerated PWID, may result in higher levels of entry into addiction treatment, including OAT, after release. For PLWH not receiving TCM, frequent re-incarceration was extremely detrimental, yet TCM was protective for those PLWH who were frequently re-incarcerated. Overall, however, PLWH receiving high-frequency TCM services had reduced survival, potentially due to extreme unmet needs in the community for which TCM services were insufficient to reduce mortality risk. Thus, TCM appears to be directed more towards those with highest need (e.g., PWID, recidivists), but may also be a marker of excessive community-based needs that are challenging to overcome (e.g., housing).

Despite the many important findings, we were unable to fully address changes in some variables over time, either because dynamic data were not available (e.g., housing, PWID status, post-release treatment for psychiatric and SUDs) or because an alternative statistical approach was employed. We relied on death certificates for causes of death, which may have contributed to some misclassification. Findings are from an integrated system that includes both prisons and jails in a single state, so may not be generalizable to all systems, though likely represents a best-case scenario within a resource-rich environment. As with any observational study, unmeasured secular trends in substance use, ART, or incarceration policies may have contributed to mortality. Though compassionate release for "terminal illness" was uncommon during this period, some who may have otherwise died in prison did so in the community following release. This limitation, however, may be mitigated by the fact that deaths related to malignancy and organ failure occurred rarely within 6 months post-release, perhaps because people died sooner from other causes or because they received preventative care during incarceration. We included people released at least once, which potentially biases findings towards people with shorter incarceration periods, though we included all people admitted and released to the CTDOC over eight years of observation, and the majority (69%) of deaths were among people with an index incarceration lasting >30days. Strengths of this study, however, are the inclusion of both jail detainees and sentenced prisoners, the large sample of PLWH, the successful linkage of which allowed for robust characterizization of post-release experiences for both recidivists and non-recidivists over eight years of observation.

In resource-rich settings, incarcerated PLWH are extremely vulnerable to death after release. Furthermore, the majority of deaths are due to treatable conditions including HIV/AIDS, SUDs, and liver disease. The CJ system can, however, assist some at-risk PLWH temporarily interface with needed medical services, but longitudinal engagement in community-based care is key to reducing mortality. Beyond reducing incarcerations, PLWH need improved access to within-prison treatment for HIV, SUDs, viral hepatitis, and other co-morbidities,

with services continued without interruption after release. Improved access to health insurance and case management are two crucial resources that can support the provision of these services in the community.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Research in Context

Evidence before this study

We searched PubMed using the following search terms: 1) "prison", "jail", or "***incarcerat*"; 2) "HIV" or "AIDS"; 3) "mortality" or "death," resulting in 87 matches. We reviewed resulting abstracts and full text to apply the following inclusion criteria: original research articles; published between March 1, 2008 and March 1, 2018; evaluated mortality following prison- or jail-release; and either measured HIV-related deaths, accounted for HIV status, or exclusively sampled people living with HIV (PLWH). Eight published studies met inclusion criteria, and all but one cross-sectional survey used retrospective cohort designs.

In a cross-sectional survey of 102 male prisoners with HIV in Indonesia followed for 24 months after release, the crude mortality ratio was 215.7 per 1,000 person-years and HIV was the most common cause of death. Survival time was inversely correlated with incarceration for a drug-related offense, longer incarcerations, and advanced HIV, whereas addiction treatment was associated with longer survival. Similarly, in a retrospective cohort study of 147 prisoners with HIV released to the community in French Guiana (2007-2013), the standardized mortality ratio (SMR) was 14.8. Age and advanced HIV were associated with death in multivariate models, though just 50% of the sample received antiretrovirals (ART) prior to release.

In the United States, one study from New York City, two from North Carolina, and three from Georgia each linked prison or jail data with death records to assess all-cause mortality, regardless of HIV status. Consistently, across study sites, time periods, and assessments, post-release mortality was high and SMRs were elevated compared to the general population. SMRs were higher for White, compared to Black, former prisoners. Primary causes of death were HIV, homicides, accidents, substance use,HIV, and liver-related disease, accounting for up to 62% of the excess mortality following release in one study. Though some found an elevated risk of cancer-related deaths, a retrospective cohort study from Georgia used matched cancer and death registry data to show that there was no elevated risk of cancer mortality compared to the general population, though PLWH had higher cancer incidence mostly from viral-associated non-AIDS defining cancers. Cumulatively, these studies suggest high risk of death following release from prison overall, but there is limited information specific to PLWH to identify modifiable factors that could inform the development of interventions and policy.

Added value of this study

To our knowledge, this is one of the first studies in a resource-rich setting where ART and insurance coverage is high to describe post-release mortality among PLWH. By merging data from custodial, pharmacy, HIV surveillance, case management, and death indices over 8 years of observation for all PLWH incarcerated within an integrated correctional system, we were able to fully appreciate the impact of incarceration (and reincarceration) on mortality.

Implications of all the available evidence

PLWH transitioning from prisons to communities experience excess risk of death from all-causes compared to the general population, with death primarily due to HIV, drug overdose, liver-related or cardiovascular disease, and accidental injury or suicide. To address this critical health disparity, PLWH require access to health insurance, and early identification and treatment of HIV with ART and substance use disorders with medication assisted therapies, during incarceration and following return to communities.

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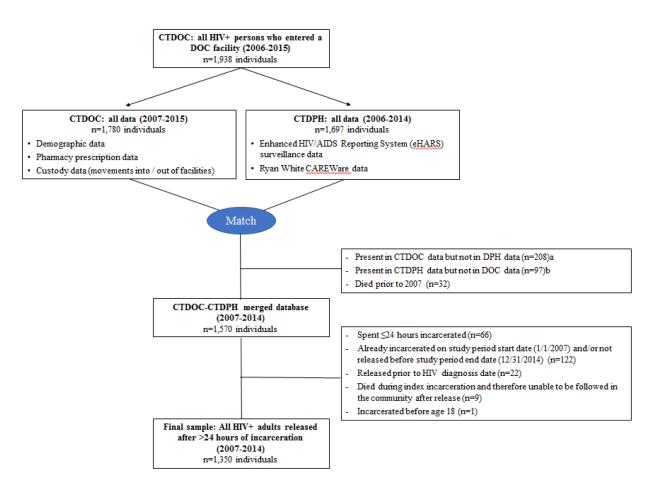
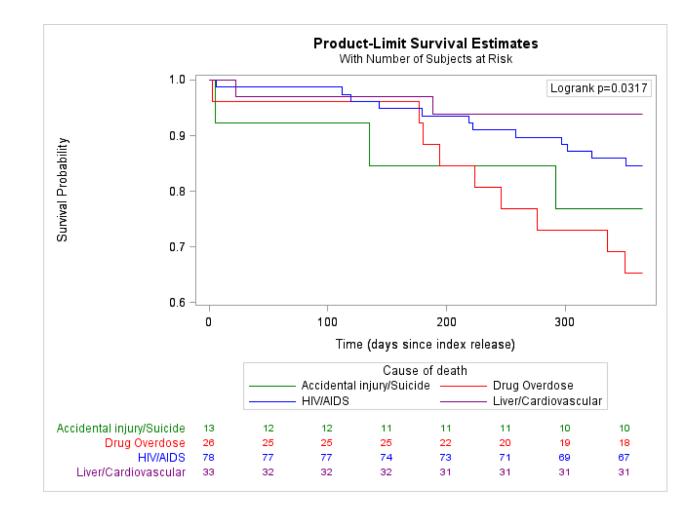


Figure 1. Flow diagram of retrospective cohort

a Resided outside of Connecticut prior to or following incarceration in CTDOC, or first recorded incarceration took place in 2015

b Incarcerated in CTDOC prior to 2007 and was never reincarcerated during 2007-2014



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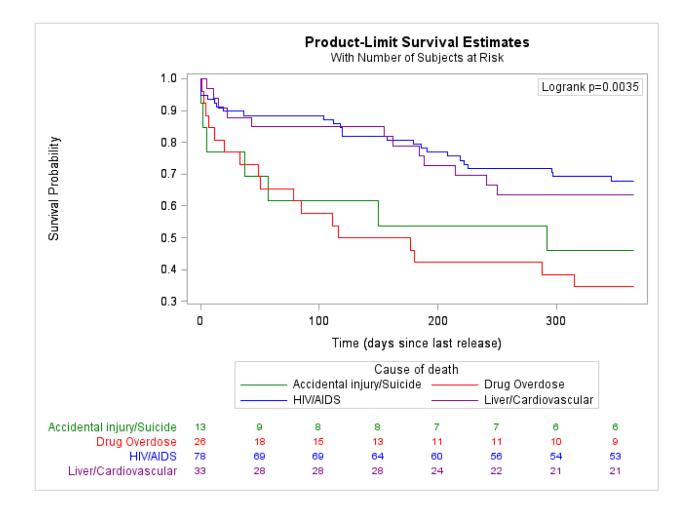
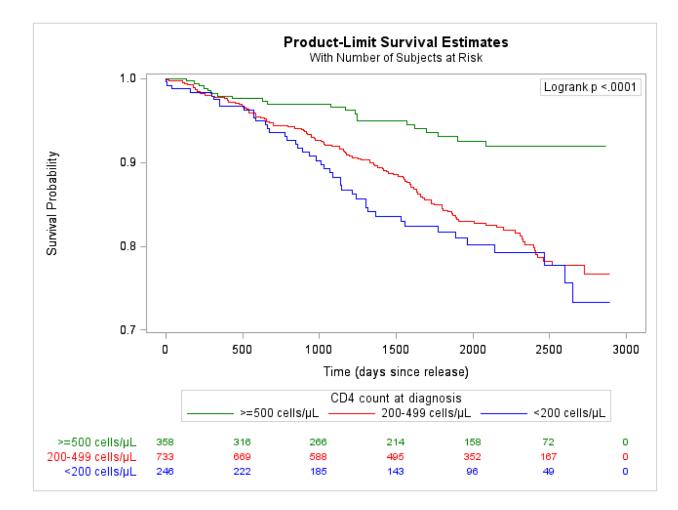


Figure 2.

A. Among those who died, time from index release to death in one year, stratified by major causes of death

B.Among those who died, time from most recent release to death in one year, stratified by major causes of death



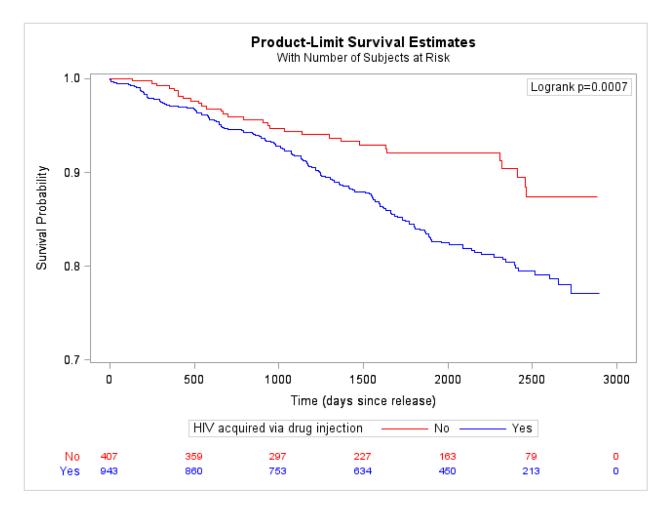


Figure 3. Bivariate association between CD4 count at time of HIV diagnosis and time-to-death from any cause

A.Individuals with missing or unreported CD4 count at time of diagnosis (n=13) were excluded from the bivariate analysis, leaving 1337 individuals for analysis. For 33 individuals missing a CD4 count at time of diagnosis but with CD4 counts available within 3 months of their diagnosis date, their first CD4 counts were used as an approximation of their CD4 counts at time of diagnosis.

B.Bivariate association between prior injection drug use and time-to-death from any cause

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Table 1.

Characteristics of people living with HIV formerly incarcerated in an integrated correctional system^{*} (N=1,350) and predictors of time-to-death from any cause after initial release, 2007-2014

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Predisposing factors Age at time of index release 18-29 18-29 30-39 50 50 617 (50 617 (617 (50 617 (50 617 (50 617 (50 50 617 (50 617 (50 617 (50 617 (617 (50 617 (50 617 (50 618 (619 (610 (50 (610 (610 (610 (610 (610 (610 (610 (610 (610 (610 (610 (610 (610 (114 (8.4%) 253 (18.7%)					
e of index release	↓(8.4%) (18.7%)					
	t (8.4%) (18.7%)					
	(18.7%)	5 (4.4%)	referent		referent	
		25 (9.9%)	1.95 (0.75-5.10)	0.17	1.98 (0.64-6.15)	0.24
	617 (45.7%)	90 (14.6%)	2.96 (1.21-7.29)	0.02	2.83 (0.95-8.40)	0.06
	366 (27.1%)	64 (17.5%)	4.40 (1.77-10.93)	0.001	3.65 (1.21-11.08)	0.02
	307 (22.7%)	31 (10.1%)	referent			
Male 1043	1043 (77.3%)	153 (14.7%)	1.59 (1.08-2.34)	0.02		
Race/Ethnicity						
White 250 (250 (18.5%)	41 (16.4%)	referent		referent	
Black 557 (557 (41.3%)	60(10.8%)	0.62 (0.42-0.92)	0.02	0.52 (0.34-0.80)	0.003
Hispanic 495 (495 (36.7%)	77 (15.6%)	0.89 (0.61-1.30)	0.55	0.96 (0.65-1.43)	0.84
Other 48 (48 (3.6%)	6 (12.5%)	0.68 (0.29-1.59)	0.37	0.69 (0.29-1.68)	0.42
Education level						
< High school 610 (610 (45.2%)	96 (15.7%)	referent			
High school 740 (740 (54.8%)	88 (11.9%)	0.76 (0.57-1.02)	0.06		
Marital status ^d						
Not married 1086	1086 (84.3%)	151 (13.9%)	referent			
Married 203 (203 (15.7%)	32 (15.8%)	1.12 (0.77-1.64)	0.55		
PWID						
407 (407 (30.2%)	31 (7.6%)	referent		referent	
Yes 943 (943 (69.9%)	153 (16.2%)	1.93 (1.31-2.84)	0.001	See below	0.01
PWID × Re-incarceration rate						
PWID \times 0 re-incarcerations/year					2.06 (1.23-3.47)	

	Total n (%) or median (IQR) ^a	n (%) who died ^b	Bivariate HR (95% CI)	<i>p</i> -value	Adjusted HR $(95\% \text{ CI})$ at $t = 0$	<i>p</i> -value
Predisposing factors						
PWID \times 0-33 re-incarcerations/year					1.54 (1.00-2.38)	
PWID \times 1 re-incarceration/year					0.85 (0.48-1.52)	
PWID \times 2 re-incarcerations/year					0.35 (0.11-1.10)	
PWID \times 3 re-incarcerations/year					0.14(0.02 - 0.87)	
CD4 count at time of diagnosis c						
500 cells/mL	358 (26.8%)	21 (5.9%)	referent		referent	
200-499 cells/mL	733 (54.8%)	119 (16.2%)	2.58 (1.62-4.11)	<0.001	2.54 (1.50-4.31)	0.005
< 200 cells/mL	246 (18.4%)	44 (17.9%)	3.08 (1.83-5.18)	<0.0001	3.44 (1.90-6.20)	<0.001
Years since HIV diagnosis at time of index release	11.4	12.9				
Median (interquartile range)	(5.4-17.1)	(7.4-17.9)	1.04 (1.02-1.07)	0.0002		
HIV diagnosed during last incarceration						
No	1282 (95.0%)	183 (14.3%)	referent			
Yes	68 (5.0%)	1 (1.5%)	0.12 (0.02-0.88)	0.04		
Enabling or disabling factors						
Length of index incarceration						
30 days	454 (33.6%)	67 (14.8%)	referent			
31-364 days	697 (51.6%)	102 (14.6%)	0.94 (0.69-1.29)	0.72		
365 days	199 (14.7%)	15 (7.5%)	0.58 (0.33-1.02)	0.06		
Conditions of index release						
Unconditional	783 (58.0%)	116 (14.8%)	referent			
Conditional	351 (26.0%)	40 (11.4%)	0.79 (0.55-1.13)	0.19		
Bonded	216 (16 0%)	28 (13.0%)	0.92 (0.61-1.38)	0.67		
Year of index release						
2007-2008	595 (44.1%)	120 (20.2%)	referent			
2009-2010	388 (28.7%)	47 (12.1%)	0.77 (0.54-1.08)	0.13		
2011-2012	211 (15.6%)	15 (7.1%)	0.79 (0.45-1.38)	0.41		
2013-2014	156 (11.6%)	2 (1.3%)	0.41 (0.10-1.72)	0.22		
Health insurance during follow-up						
Consistently uninsured	365 (27.0%)	102 (27.9%)	referent		referent	

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	Total n (%) or median (IQR) ^a	n (%) who died ^b	Bivariate HR (95% CI)	<i>p</i> -value	Adjusted HR $(95\% \text{ CI})$ at $t = 0$	<i>p</i> -value
Predisposing factors						
Consistently insured	637 (47.2%)	68 (10.7%)	0.28 (0.21-0.38)	<0.0001	0.09 (0.05-0.17)	<0.0001
Intermittently insured	348 (25.8%)	14 (4.0%)	0.10 (0.06-0.17)	<0.0001	001 (0.00-0.04)	<0.0001
Linked to HIV care within 14 days after index release f						
No	1076 (79.7%)	144 (13.4%)	referent			
Yes	274 (20.3%)	40 (14.6%)	1.26 (0.89-1.79)	0.20		
Transitional case management during index release						
No	1010 (74.8%)	138 (13.7%)	referent			
Yes	340 (25.2%)	46 (13.5%)	1.14 (0.82-1.59)	0.45		
Number of case management (TCM) visits per year of follow-up						
0	620 (45.9%)	108 (17.4%)	referent		referent	
>0-2	233 (17.3%)	20 (8.6%)	0.39 (0.24-0.62)	<0.0001	See below	0.38
3-4	146 (10.8%)	14 (9.6%)	0.45 (0.26-0.79)	0.01	See below	0.35
5	351 (26.0%)	42 (12.0%)	0.68 (0.47-0.97)	0.03	See below	<0.0001
Case management (TCM) frequency × Re-incarceration rate						
$>0-2$ TCM visits/year \times 0 re-incarcerations/year					0.77 (0.43-1.37)	
$>0-2$ TCM visits/year $\times 0.33$ re-incarcerations/year					0.64 (0.38-1.07)	
$>0-2$ TCM visits/year $\times 1$ re-incarceration/year					0.43 (0.16-1.16)	
$>0-2$ TCM visits/year $\times 2$ re-incarcerations/year					0.24 (0.03-1.84)	
$>0-2$ TCM visits/year $\times 3$ re-incarcerations/year					0.13 (0.01-3.08)	
3-4 TCM visits/year \times 0 re-incarce rations/year					1.60 (0.59-4.31)	
3-4 TCM visits/year \times 0.33 re-incarcerations/year					1.23 (0.64-2.36)	
$3-4$ TCM visits/year $\times 1$ re-incarceration/year					0.73 (0.26-2.07)	
3-4 TCM visits/year \times 2 re-incarce rations/year					0.33 (0.03-4.35)	
3-4 TCM visits/year \times 3 re-incarcerations/year					0.15 (0.00-10.00)	
5 TCM visits/year \times 0 re-incarcerations/year					2.83 (1.68-4.78)	
5 TCM visits/year \times 0.33 re-incarcerations/year					2.01 (1.28-3.15)	
5 TCM visits/year \times 1 re-incarceration/year					1.00 (0.57-1.75)	
5 TCM visits/year \times 2 re-incarcerations/year					0.35 (0.12-1.01)	
5 TCM visits/year \times 3 re-incarcerations/year					0.12 (0.02-0.63)	

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Total or me (IQ)	Total n (%) or median (IQR) ^a	n (%) who died ^b	Bivariate HR (95% CI)	<i>p</i> -value	Adjusted HR $(95\% \text{ CI})$ at $t = 0$	<i>p</i> -value
Predisposing factors						
Re-incarceration rate						
Median (interquartile range) 0.19 (C	0.19 (0-0.53)	0.19 (0-0.71)	1.60 (1.16-2.20)	0.004	See below	<0.0001
Re-incarce ration rate \times PWID \times Case management (TCM) frequency						
1-unit increase in re-incarceration rate \times non-PWID \times 0 TCM visits/year					21.86 (10.90-43.86)	
1-unit increase in re-incarceration rate \times non-PWID \times >0-2 TCM visits/year	ar				12.13 (3.63-40.52)	
1-unit increase in re-incarceration rate \times non-PWID \times 3-4 TCM visits/year					9.97 (1.81-54.83)	
1-unit increase in re-incarceration rate \times non-PWID x 5 TCM visits/year					7.72 (3.63-16.45)	
1-unit increase in re-incarceration rate \times PWID \times 0 TCM visits/year					8.99 (5.14-15.72)	
1-unit increase in re-incarceration rate \times PWID \times >0-2 TCM visits/year					4.99 (1.62-15.39)	
1-unit increase in re-incarceration rate \times PWID \times 3-4 TCM visits/year					4.10 (0.80-20.89)	
1-unit increase in re-incarceration rate \times PWID \times >5 TCM visits/year					3.17 (1.96-5.14)	
Re-incarcerated for 365 days						
No 1092 (§	1092 (80.9%)	170(15.6%)	referent		referent	
Yes 258 (19.1%)	9.1%)	14(5.4%))	0.26 (0.15-0.45)	<0.0001	0.41 (0.22-0.76)	0.004
Percentage of total follow-up time spent re-incarcerated						
0-1% 605 (4	605 (44.8%)	100(16.5%)	referent			
2-10% 225 (1	225 (16.7%)	21 (9.3%)	0.44 (0.27-0.70)	0.001		
11-25% 196 (1	196 (14.5%)	31 (15.8%)	0.75 (0.50-1.12)	0.16		
26-50% 219 (1	219 (16.2%)	23 (10.5%)	0.48 (0.30-0.75)	0.001		
51-100% 105 ()	105 (7.8%)	9 (8.6%)	0.44 (0.22-0.87)	0-02		
Length of last incarceration						
30 days 347 (2	347 (25.7%)	63 (18.2%)	referent			
31-364 days 769 (5	769 (57.0%)	103 (13.4%)	0.65 (0.48-0.89)	0.01		
> 365 days 234 (17.3%)	7.3%)	18 (7.7%)	0.38 (0.22-0.64)	0.0003		
Conditions of release from last incarceration						
Unconditional 770 (5	770 (57.0%)	122 (15.8%)	referent			
Conditional 291 (2	291 (21.6%)	36 (12.4%)	0.78 (0.54-1.14)	0.20		
Bonded 129 (5	129 (9.6%)	21 (16.3%)	1.21 (0.76-1.92)	0.42		

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	Total n (%) or median (IQR) ^a	n (%) who died ^b	Bivariate HR (95% CI)	<i>p</i> -value	Adjusted HR (95% CI) at $t = 0$	<i>p</i> -value
Predisposing factors						
Currently re-incarcerated	160 (11.9%)	5 (3.1%)	0.19 (0.08-0.46)	0.0002		
Need factors						
Prescribed ART during index incarceration						
No	590 (43.7%)	85 (14.4%)	referent			
Yes	760 (56.3%)	99 (13.0%)	1.15 (0.86-1.53)	0.37		
Virally suppressed within 90 days prior to index release						
No/viral level not reported	588 (43.6%)	89 (15.1%)	referent			
Yes	442 (32.7%)	52 (11.8%)	0.97 (0.69-1.37)	0.85		
Viral load not drawn prior to release	320 (23.7%)	43 (13.4%)	1.01 (0.70-1.45)	0.96		
Number of medical co-morbidities during index incarceration						
0	838 (62.1%)	111 (13.2%)	referent		referent	
Т	294 (21.8%)	37 (12.6%)	1.16 (0.80-1.69)	0.43	1.44 (0.96-2.15)	0.08
2	218 (16.2%)	36 (16.5%)	1.77 (1.21-2.59)	0.003	1.86 (1.23-2.82)	0.003
Psychiatric need during index incarceration						
Low severity score (1-2), untreated	650 (48.2%)	89 (13.7%)	referent			
Low severity score (1-2), treated	58 (4.3%)	5 (2.0%)	0.76 (0.31-1.87)	0.55		
High severity score (3-5), untreated	256 (19.0%)	38 (65.5%)	0.98 (0.67-1.43)	0.90		
High severity score (3-5), treated	386 (28.6%)	52 (13.5%)	1.10 (0.78-1.55)	0.60		
Addiction severity score during index incarceration $^{\mathcal{G}}$						
Low severity/no disorder (score 1-2)	220 (16.7%)	16 (7.3%)	referent			
Moderate severity disorder (score 3)	861 (65.2%)	123 (14.3%)	2.21 (1.31-3.72)	0.003		
Serious disorder (score 4-5)	239 (18.1%)	44 (18.4%)	3.13 (1.77-5.55)	<0.0001		
Treated for an opioid use disorder during index incarceration						
No	1341 (99.3%)	183 (13.6%)	referent			
Yes	9 (0.7%)	1 (11.1%)	2.47 (0.34-17.7)	0.37		
Percentage of all re-incarcerations during which ART was prescribed						
Never re-incarcerated	535 (39.6%)	89 (16.6%)	referent		referent	

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	Total n (%) or median (IQR) ^a	n (%) who died ^b	Bivariate HR (95% CI)	<i>p</i> -value	Adjusted HR $(95\% \text{ CI})$ at $t = 0$	<i>p</i> -value
Predisposing factors						
0-10%	128 (9.5%)	25 (19.7%)	0.87 (0.56-1.36)	0.54	0.34 (0.18-0.65)	0.001
11-50%	97 (7.2%)	15 (15.3%)	0.62 (0.36-1.07)	0.09	0.12 (0.06-0.27)	<0.0001
51-90%	101 (7.5%)	9 (8.7%)	0.33 (0.17-0.66)	0.002	0.04 (0.01-0.11)	<0.0001
91-100%	489 (36.2%)	46 (9.4%)	0.45 (0.31-0.64)	<0.0001	0.08 (0.03-0.21)	<0.0001
Percentage of all re-incarcerations that ended in viral suppression						
Never re-incarcerated	535 (39.6%)	89 (16.6%)	referent			
0-24%	297 (22.0%)	46 (15.5%)	0.71 (0.50-1.01)	0.06		
25-49%	83 (6.2%)	10 (12.0%)	0.47 (0.24-0.90)	0.02		
50-74%	172 (12.7%)	17 (9.9%)	0.40 (0.24-0.67)	0.001		
75-100%	263 (19.5%)	22 (8.4%)	0.40 (0.25-0.64)	0.0001		
Percentage of all re-incarcerations with a documented change from virologic failure to suppression						
Never re-incarcerated	535 (39.6%)	89 (16.6%)	referent			
0-24%	591 (43.8%)	76 (12.9%)	0.60(0.44 - 0.81)	0.001		
25-49%	79 (5.9%)	7 (8.9%)	0.33 (0.15-0.70)	0.004		
50-74%	86 (6.4%)	7 (8.1%)	0.32 (0.15-0.69)	0.004		
75-100%	59 (4.4%)	5 (8.5%)	0.38 (0.16-0.94)	0.04		
Prescribed ART during last incarceration						
No	357 (26.4%)	69 (19.3%)	referent			
Yes	993 (73.6%)	115 (11.6%)	0.62 (0.46-0.84)	0.002		
Virally suppressed within 90 days prior to release from last incarceration						
No/viral level not reported	411 (30.4%)	79 (19.2%)	referent			
Yes	653 (48.4%)	67 (10.3%)	0.56 (0.40-0.77)	0-0004		
Viral load not drawn prior to release	286 (21 .2%)	38 (13.3%)	0.72 (0.49-1.06)	0.10		
Viral suppression status during last incarceration						
Never virally suppressed	398 (29.5%)	77 (19.3%)	referent			
Failure, then suppression	260 (19.3%)	31 (11.9%)	0.60(0.39-0.91)	0.02		
Summession then failure	38 (38%)	6(158%)	0.80 (0.35-1.84)	0.60		

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	Total n (%) or median (IQR) ^d	n (%) who died ^b	Bivariate HR (95% CI)	<i>p</i> -value	Adjusted HR $(95\% \text{ CI})$ at $t = 0$	<i>p</i> -value
Predisposing factors						
Consistent suppression	485 (35.9%)	40 (8.2%)	47 (0.32-0.68)	<0.0001		
Viral load never drawn	169 (12.5%)	30 (17.8%)	1.07 (0.70-1.63)	0.76		
Number of medical comorbidities during last incarceration						
0	653 (48.4%)	92 (14.1%)	referent			
1	340 (25.2%)	39 (11.5%)	0.85 (0.59-1.24)	0.41		
2	357 (26.4%)	53 (14.8%)	1.16 (0.82-1.62)	0.40		
Psychiatric need during last incarceration						
Low severity score (1-2), untreated	589 (43.6%)	81 (13.8%)	referent			
Low severity score (1-2), treated	99 (7.3%)	35 (35.4%)	1.21 (0.81-1.80)	0.35		
High severity score (3-5), untreated	199 (14.7%)	16 (8.0%)	1.14 (0.66-1.94)	0.64		
High severity score (3-5), treated	463 (34.3%)	52 (11.2%)	0.82 (0.58-1.17)	0.28		
Addiction severity score during last incarceration h						
Low severity/no disorder (score 1-2)	122 (9.2%)	19 (15.6%)	referent		referent	
Moderate severity disorder (score 3)	945 (70.9%)	107 (11.3%)	0.70 (0.43-1.13)	0.15	0.65 (0.38-1.12)	0.12
Serious disorder (score 4-5)	265 (19.9%)	56 (21.1%)	1.37 (0.81-2.30)	0.24	1.17 (0.65-2.09)	0.61
Treated for an opioid use disorder during last incarceration						
No	1322 (97.9%)	183 (13.8%)	referent			
Yes	28 (2.1%)	1 (3.6%)	0.28 (0.04-1.97)	0.20		
Viral suppression and timing of last viral load drawn prior to death/censoring						
Viral suppression, within 6 months	840 (62.2%)	79 (9.4%)	referent		referent	
Virologic failure, within 6 months	208 (15.4%)	65 (31.3%)	3.63 (2.62-5.05)	<0.0001	2.76 (1.94-3.92)	<0.0001
Viral suppression, >6 months	160 (11.9%)	17 (10.6%)	1.10 (0.65-1.85)	0.73	0.83 (0.48-1.42)	0.49
Virologic failure, >6 months	87 (6.4%)	11 (12.6%)	1.19 (0.64-2.24)	0.58	0.72 (0.37-1.42)	0.35
Viral load never drawn ⁱ	55 (4.1%)	12 (21.8%)	3.90 (2.13-7.16)	<0.0001	2.13 (1.09-4.18)	0.03

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PWID= people who acquired HIV through injection drug use (people who inject drugs;); non-PWID=people without an injection drug use history

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follow-up. Due to missing data for the CD4 count and addiction severity score variables as specified below, the final parsimonious model includes 1,319 individuals followed for 6,276 person-years, during which there were 182 deaths. Numbers listed are n (%) out of the total number of individuals in the sample (n=1350). Percentages may not sum to 100% due to rounding or missing data as specified below. Bivariate analyses include all 1,350 individuals released during the observation period- During a median follow-up time of 5.2 (IQR 3-0-6.7) years (6,418 person-years), there were 184 deaths during

 b_b Numbers listed are the row n (%) of individuals with the specified characteristic who experienced the outcome of interest (death)

 c_1 fransgender males (n=1) have been included the male category and transgender females (n=3) have been included in the female category

 $d_{\rm Individuals}$ with missing/unreported marital status (n=61) were excluded from the bivariate analysis such that the total n=1289

e^eIndividuals with missing/unreported CD4 count at time of diagnosis (n=13) were excluded from the bivariate analysis such that the total n=1337. For 33 individuals missing a CD4 count at time of diagnosis but with CD4 counts available within 3 months of their diagnosis date, their first CD4 counts were used as an approximation of their CD4 counts at time of diagnosis.

 $f_{\rm T}$ he "No" category includes 25 individuals who were either re-incarcerated or died within 14 days without having a VL drawn in the community.

 g Index incarceration periods where the addiction severity score was never assessed (n=30) were excluded from the bivariate analysis such that the total n=1320.

 $h_{\rm Last}$ incarceration periods where the addiction severity score was never assessed (n=18) were excluded from the bivariate analysis such that the total n=1332.

 \dot{f} wiral load never drawn" category includes 18 individuals who were followed in the community for <6 months prior to death or censoring.