BMJ Open Opioid substitution therapy as a strategy to reduce deaths in prison: retrospective cohort study

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ABSTRACT

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Objectives: To describe deaths in prison among opioid-dependent people, and examine associations between receipt of opioid substitution therapy (OST) and risk of death in prison.

Design: Retrospective cohort study.

Setting: Adult prisons in New South Wales (NSW), Australia.

Participants: 16 715 opioid-dependent people who were received to prison between 2000 and 2012.

Interventions: Opioid substitution therapy.

Primary outcome measures: Natural and unnatural (suicide, drug-induced, violent and other injury) deaths in prison.

Results: Cohort members were in prison for 30 998 person-years (PY), during which time there were 51 deaths. The all-cause crude mortality rate (CMR) in prison was 1.6/1000 PY (95% CI 1.2 to 2.2/1000 PY), and the unnatural death CMR was 1.1/1000 PY (95% CI 0.8 to 1.6/1000 PY). Compared to time out of OST, the hazard of all-cause death was 74% lower while in OST (adjusted HR (AHR): 0.26; 95% CI 0.13 to 0.50), and the hazard of unnatural death was 87% lower while in OST (AHR: 0.13; 95% CI 0.05 to 0.35). The all-cause and unnatural death CMRs during the first 4 weeks of incarceration were 6.6/1000 PY (95% CI 3.8 to 10.6/1000 PY) and 5.5/1000 PY (95% CI 2.9 to 9.4/1000 PY), respectively. Compared to periods not in OST, the hazard of all-cause death during the first 4 weeks of incarceration was 94% lower while in OST (AHR: 0.06; 95% CI 0.01 to 0.48), and the hazard of unnatural death was 93% lower while in OST (AHR: 0.07; 95% CI 0.01 to 0.53).

Conclusions: Mortality of opioid-dependent prisoners was significantly lower while in receipt of OST.

INTRODUCTION

Deaths in prison are a serious public health issue, raising questions as to the quality of care and supervision provided by correctional authorities. Reflecting such concerns, deaths in prison usually result in extensive postmortem inquiries and may also lead to litigation against correctional authorities and healthcare providers.¹

Strengths and limitations of this study

- This study is based on a large, statewide cohort with mortality outcomes determined via population-based registries.
- Cohort members were not randomly allocated to treatment.
- Data regarding onset of opioid dependence and current opioid dependence were not available.

Unnatural deaths in prison are of particular concern due to their preventable nature. Unnatural deaths include suicides, violent or drug-induced deaths and other injury-related deaths. Studies in developed countries have found that unnatural deaths comprise 48– 59% of all deaths in prison, with suicide being the most common cause of unnatural death.^{2–5} These figures suggest considerable opportunities to reduce unnatural deaths in prison.

Opioid-dependent people commonly experience imprisonment,⁶ and there are several reasons to believe that opioiddependent prisoners may be at particular risk of unnatural death in prison. Drug withdrawal has been implicated as a possible trigger for suicide in the first days of incarceration.⁷ ⁸ Additionally, use of illicitly obtained opioids while in prison⁹ carries with it the risk of overdose. To the best of our knowledge, deaths in prison specifically among opioid-dependent people have not been described previously. Furthermore, no studies have considered whether treatment for opioid dependence during incarceration reduces mortality risk, as it does among opioid-dependent people residing in the community.¹⁰ In this study, we describe deaths in prison in a cohort of opioiddependent people and examine whether receiving opioid substitution therapy (OST) reduces the risk of death in prison.

METHODS Setting

New South Wales (NSW) is the most populous state in Australia and houses around one-third of the country's prisoners, or 9645 people at 31 June 2012.¹¹ Health services in NSW prisons are provided under the Ministry of Health, and there is a well-established prison OST programme that operates as part of the statewide opioid treatment programme.^{12 13} People who enter prison while in OST may continue treatment while incarcerated, and OST can be started during incarceration if clinically indicated.¹² Clinical indications for OST are the same as in community settings.¹³ Most prisoners receiving OST are prescribed methadone due to the longer dosing procedure associated with buprenorphine and concerns about diversion of buprenorphine. In 2012, 16% of NSW prisoners were prescribed OST.^{11 12} Coverage of OST (the proportion of opioid-dependent prisoners receiving treatment) has been estimated at 43%.¹

Data sources and linkage

Data for this retrospective cohort study were extracted from administrative datasets used to record OST programmes, offending and incarceration, and mortality. The Pharmaceutical Drugs of Addiction System (PHDAS) is a database of all methadone and buprenorphine recipients in NSW since 1985. The PHDAS records each patient's full name, date of birth, gender and clinical variables including dates of OST entry and exit. Identifying variables in this dataset are considered to be of high accuracy as proof of identity must be shown to the prescribing doctor before a prescription can be issued. Data used for linkage were for the calendar years 1985–2010.

Offending and incarceration data were sourced from the NSW Bureau of Crime Statistics and Research (BOCSAR) Re-offending Database (ROD). This database includes dates and types of offences in criminal cases heard by local, district and supreme courts of NSW, and dates of receptions to and discharges from the custody of Corrective Services NSW. All available data were provided for linkage; this included offending data for the period 1 December 1993 to 31 December 2011 and incarceration data for the period 1 January 2000 to 31 March 2012.

The National Death Index (NDI) is a database held by the Australian Institute of Health and Welfare (AIHW), containing mortality data collected from all the Registry of Births, Deaths and Marriages in each Australian State and Territory. Causes of death are recorded in the NDI by expert clinical coders on the basis of information contained in death certificates and, where available, coronial files, using the International Statistical Classification of Diseases and Related Health Problems 10th Edition (ICD-10). Dates of death were available for the entire period for which incarceration data were available (1 January 2000 to 31 March 2012); however, for administrative reasons, causes of death were available only for deaths occurring up to 31 December 2010.

Linkage between the OST and offending/incarceration data was performed by BOCSAR staff, and linkage between the OST and mortality data was undertaken by AIHW staff. Linkage was completed using probabilistic linkage software. Variables used for matching purposes included full name, date of birth, gender, and date and state of last known contact where available. These linked datasets were forwarded to the investigators with identifiers removed.

Cohort and data definitions

From the linked datasets, we identified a cohort of opioid-dependent people (people who were recorded in the PHDAS as having received OST at some point during 1986-2010) who had been incarcerated at least once (as recorded in the ROD). In defining receipt of OST, we excluded temporary treatment programmes (usually interstate visitors) and treatment episodes that were part of a buprenorphine clinical trial (during which the individual may have been allocated to placebo). Periods in the custody of Corrective Services NSW were determined using reception and discharge dates in the ROD. We used the first recorded OST episode as a proxy for the onset of opioid dependence and excluded prison episodes that occurred prior to the first OST episode; all subsequent prison episodes for an individual were included. If an individual's first OST episode started while incarcerated, that prison episode was included. Given the chronic and relapsing nature of opioid dependence,¹⁵¹⁶ we assumed that all cohort members remained opioid-dependent from their first episode of OST until death or the end of follow-up.

Deaths in prison were defined as deaths that occurred while the individual was recorded as being in the custody of Corrective Services NSW. This definition does not stipulate that a deceased individual was physically in prison at the time of death. Some decedents may have been receiving care in hospital at the time of death, while still in the custody of Corrective Services NSW. Deaths with known causes (ie, all deaths occurring in the calendar years 2000-2010) were categorised as natural (ie, disease-related) or unnatural deaths. We used the Corrective Services NSW definition of 'unnatural deaths', which includes suicides, drug-induced deaths, violent deaths and deaths due to accidental injuries or injuries in which no intent (ie, suicidal or violent) could be determined.³ ICD-10 codes for unnatural deaths are provided in the online supplementary materials.

Periods of OST while in prison, and treatment status (in OST/out of OST) at the time of death, were determined using treatment entry and exit dates from the PHDAS. New treatment episodes were defined when an individual started treatment 7 or more days after discharge from a previous treatment episode. A change in medicine (methadone to buprenorphine, or vice versa) was considered a continuous episode if there were less than 7 days between ceasing one medicine and startingthe other.

Data analysis

Statistical analyses were undertaken in SAS V.9.3. We used descriptive statistics to examine demographic characteristics and incarceration. Participants with any record of receiving OST in prison were compared with those who did not receive OST in prison using χ^2 tests for categorical variables and the Wilcoxon rank-sum test for continuous variables. We constructed a frequency histogram to illustrate time from entry to prison until death. For dates for which comparable data were available (financial years 2000–2001 to 2009–2010), we compared the total number of unnatural deaths among prisoners in NSW³ with the number of unnatural deaths in the opioid-dependent cohort.

We calculated all-cause and cause-specific crude mortality rates (CMR) with Poisson 95% CI for all time in prison and for the first 4 weeks in prison. Rates were also calculated by treatment status (in/out of OST). Person-years (PY) used in the calculation of CMRs accrued whenever participants were in the custody of Corrective Services NSW. Prison episodes where the participant was received and released on the same day were counted as 1 day. PY ceased to accrue at death on 31 March 2012 for all-cause mortality rates, and on 31 December 2010 for cause-specific mortality rates. CMR ratios with 95% CI were calculated to assess unadjusted risk of death in treatment compared with that out of treatment. The rate ratio (RR) was not calculated if there were zero deaths in either the in-treatment or out-of-treatment condition.

We used Cox regression to examine the adjusted association between OST exposure and two in-prison mortality outcomes: deaths from all causes and unnatural deaths. Two models were developed for each outcome: the first considering mortality risk across all time in prison, and the second examining mortality risk during the first 4 weeks of incarceration. All prison episodes were included, with days in prison counted as discontinuous intervals at risk. Although participants could have multiple observations within the dataset, the internal computations of a Cox model are such that when there is only one event of interest per person (in this case, death), there is no need for adjustments for multiple observations.¹⁷ OST exposure while in prison was coded as a time-dependent variable, with periods out of treatment coded as 0 and periods in treatment coded as 1. Demographic variables included in the models were gender, indigenous (ie, Aboriginal or Torres Strait Islander) status and age at entry to prison; this latter variable increased as necessary with subsequent incarcerations of an individual. We also included variables that described prior experiences of incarceration during our observation periods, namely, the number and duration (days) of prior incarcerations.

These variables increased with each incarceration per person. We also included dichotomous variables that recorded if a participant had been charged with a drug, property or violent offence prior to incarceration. These were included because of possible associations between death in prison (particularly suicide) and prior offending (particularly violent offending).¹⁸ ¹⁹ The offending history variables could change from no to yes with subsequent incarcerations. Observations with missing covariate data were excluded from the analyses. We tested each variable for its bivariate association with mortality and entered all variables into a multivariate model. We tested the proportional hazards assumption for each static predictor variable in the multivariate model by including an interaction between it and log (time) in the model. All variables in both models were found to satisfy the proportional hazards assumption.

RESULTS

The cohort comprised 16 715 opioid-dependent people who had been incarcerated at least once between January 2000 and March 2012 (table 1). The majority (79%; n=13 199) of the cohort was male, and 30% (n=5011) were identified as indigenous. The median age at first entry to prison during our observation period was 30 years (ranging from 16 to 64 years).

Cohort members were received in prison a median of two times (ranging between 1 and 34). Duration of incarceration ranged from 1 day to the entire length of follow-up (12 years and 3 months), with a median duration of 71 days. Most participants (76.9%; n=12 852) received OST at some point while incarcerated. Compared to participants who received OST in prison, participants who did not receive OST in prison were significantly older, less likely to be indigenous and have fewer incarcerations of shorter duration (table 1).

Deaths in prison

Fifty-one cohort members died in prison (table 2). Most were men (90%; n=46) and non-indigenous (73%; n=37). The median age at death was 34 years (ranging between 20 and 54). Only 22% (n=11) of decedents were receiving OST at the time of death. Cause of death was available for 43 deaths in prison. Of these, nearly three-quarters (74%; n=32) were unnatural deaths: 19 suicides, 6 drug-related deaths, 2 violent deaths and 5 deaths due to other injuries. Between the financial years 2000–2001 and 2009–2010, unnatural deaths in this cohort accounted for 35% of all unnatural deaths in NSW prisons³ (see online supplementary materials for further details).

One-third of all deaths in prison (n=17) occurred during the first 4 weeks of incarceration (figure 1). Of these, three-quarters (13/17; 76%) were unnatural deaths: 10 suicides, 2 other injury-related deaths and 1 drug-related death.

 Table 1
 Demographic characteristics of opioid-dependent persons received to prison, New South Wales, Australia, 2000–2012 (n=16 715)

Characteristic	Total (n=16 715)	Ever received OST in prison (n=12 852)	No OST in prison (n=3863)	p Value
 Male, n (%)	13 199 (79)	10 109 (78.7)	3090 (80)	0.07
Age (years) at first observed entry to prison,	30 (16–64)	29 (16–63)	32 (17–64)	<0.0001
median (minimum–maximum)				
Indigenous,* n (%)	5011 (30)	4208 (32.8)	803 (20.8)	<0.0001
Number of prison episodes, median	2 (1–34)	3 (1–34)	1 (1–23)	<.0001
(minimum–maximum)				
Duration (days) of prison episodes, median	71 (1–4473)	77 (1–4473)	46 (1–4473)	<0.0001
(minimum–maximum)				
Any receipt of OST in prison	12 852 (76.9)	-	-	-

*Indigenous status missing for 10 individuals. p Values refer to the comparison between participants who did or did not receive OST in prison (χ^2 test for categorical variables; Wilcoxon rank-sum test for continuous variables).

OST, opioid substitution therapy.

Crude mortality rates

The all-cause CMR in prison was 1.6/1000 PY (95% CI 1.2 to 2.2/1000 PY) (table 3). The all-cause CMR was significantly lower while in OST (CMR: 0.7/1000 PY; 95% CI 0.3 to 1.2/1000 PY), compared to time out of OST (CMR: 2.7/1000 PY; 95% CI 2.0 to 3.7/1000 PY) (RR: 0.24; 95% CI 0.12 to 0.47).

Deaths from natural causes occurred at similar rates whether in (0.3/1000 PY; 95% CI 0.1 to 0.8/1000 PY) or out (0.5/1000 PY; 95% CI 0.2 to 1/1000 PY) of OST (RR: 0.68; 95% CI 0.21 to 2.21). There was, however, a marked effect of OST on rates of unnatural death. The in-treatment CMR for unnatural deaths was 0.3/1000 PY (95% CI 0.1 to 0.8/1000 PY), compared to 2.2/1000 PY (95% CI 0.4 to 3.2/1000 PY) while out of OST (RR: 0.15; 95% CI 0.06 to 0.39). Of the specific unnatural causes of death, suicides were significantly less frequent while in OST (CMR: 0.2/1000 PY; 95% CI 0.04 to 0.6/1000 PY) than while not in OST (CMR: 0.2/1000 PY;

95% CI 0.04 to 0.6/1000 PY) (RR: 0.15; 95% CI 0.04 to 0.52). Of the six observed drug-induced deaths and five other injury deaths, none occurred during OST, but both violent deaths occurred while in OST.

During the first 4 weeks of incarceration, the all-cause CMR was 6.6/1000 PY (95% CI 3.8 to 10.6/1000 PY). Of the 17 cohort members who died during the first 4 weeks of incarceration, only one was receiving OST at the time of death. The in-treatment all-cause CMR in the first 4 weeks of incarceration was 0.8 deaths/1000 PY (95% CI 0.02 to 4.3/1000 PY), significantly lower than the CMR of 12.7/1000 PY (95% CI 7.2 to 20.6/1000 PY) observed among out-of-treatment cohort members (RR: 0.06; 95% CI 0.01 to 0.45). Across all time in prison, the natural mortality rate did not differ by OST status, but the in-treatment CMR for unnatural deaths (0.8/1000 PY; 95% CI 0.02 to 4.4/1000 PY) was significantly lower than the out-of-treatment CMR for unnatural deaths (11/1000 PY; 95% CI 5.7 to 19.2/1000 PY) (RR: 0.07;

Table 2Demographic characteristics and death of opioid-dependent persons who concerns New South Wales, Australia, 2000–2012	lied in prison,
Characteristic	N (%)
Male	46 (90)
Age (years) at death, median	34 (20–54)
(minimum–maximum)	
Indigenous	14 (27)
In OST at time of death	11 (22)
Deaths from natural causes*	11 (26)
Unnatural deaths*	32 (74)
Suicide	19 (44)
Drug-related	6 (14)
Violent	2 (5)
Other injury-related	5 (12)
*Cause of death unknown for eight decedents. based on n=43.	Percentage is

OST, opioid substitution therapy.

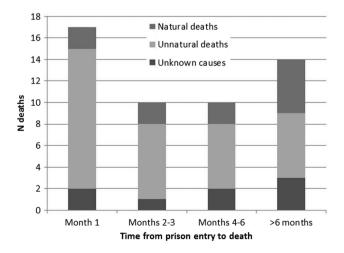


Figure 1 Deaths from natural, unnatural and unknown causes, by time in prison prior to death among opioid-dependent people who died in prison, New South Wales, Australia, 2000–2012 (n=51).

Table 3 All-cause and cause-specific crude mortality rates per 1000 person-years in prison among opioid-dependent people in New South Wales, Australia, 2000–2012 (n=16 715)

	Total ti	me in pr	ison		First	4 weeks	in prison	
Cause of death	ΡΥ	Deaths	Crude mortality rate (95% CI)	Rate ratio (95% CI)	PY	Deaths	Crude mortality rate (95% CI)	Rate ratio (95% CI)
All cause	30 988	51	1.6 (1.2 to 2.2)		2571	17	6.6 (3.8 to 10.6)	
Out of OST	14 548	40	2.7 (2 to 3.7)		1263	16	12.7 (7.2 to 20.6)	
In OST	16 440	11	0.7 (0.3 to 1.2)	0.24 (0.12 to 0.47)	1308	1	0.8 (0.02 to 4.3)	0.06 (0.01 to 0.45)
Unnatural	27 839	32	1.1 (0.8 to 1.6)		2374	13	5.5 (2.9 to 9.4)	
causes*								
Out of OST	12 462	27	2.2 (1.4 to 3.2)		1094	12	11.0 (5.7 to 19.2)	
In OST	15 377	5	0.3 (0.1 to 0.8)	0.15 (0.06 to 0.39)	1279	1	0.8 (0.02 to 4.4)	0.07 (0.01 to 0.55)
Suicide	27 839	19	0.7 (0.4 to 1.1)		2374	10	4.2 (2 to 7.7)	
Out of OST	12 462	16	1.3 (0.7 to 2.1)		1094	9	8.2 (3.8 to 15.6)	
In OST	15 377	3	0.2 (0.04 to 0.6)	0.15 (0.04 to 0.52)	1279	1	0.8 (0.02 to 4.4)	0.10 (0.01 to 0.75)
Drug-induced	27 839	6	0.2 (0.08 to 0.5)		2374	1	0.4 (0.01 to 2.3)	-
Out of OST	12 462	6	0.5 (0.2 to 1)		1094	1	1.0 (0.02 to 5.1)	
In OST	15 377	0	-	-	1279	0	-	
Violence	27 839	2	0.07 (0.01 to 0.3)		2374	0	-	-
Out of OST	12 462	0	-		1094	0	-	
In OST	15 377	2	0.1 (0.02 to 0.5)	-	1279	0	-	
Other injuries	27 839	5	0.2 (0.06 to 0.4)		2374	2	0.8 (0.1 to 3)	-
Out of OST	12 462	5	0.4 (0.1 to 0.9)		1094	2	1.8 (0.2 to 6.6)	
In OST	15 377	0	-	-	1279	0	-	
Natural causes	27 839	11	0.4 (0.2 to 0.7)		2374	2	0.8 (0.1 to 3)	-
Out of OST	12 462	6	0.5 (0.2 to 1.0)		1094	2	1.8 (0.2 to 6.6)	
In OST	15 377	5	0.3 (0.1 to 0.8)	0.68 (0.21 to 2.21)	1279	0	-	

includes suicide, drug-induced, violent and other injury death

OST, opioid substitution treatment; PY, person-years.

95% CI 0.01 to 0.55). The same pattern was observed in relation to suicide deaths (in-OST CMR: 0.8/1000 PY, 95% CI 0.02 to 4.4/1000 PY; out-of-OST CMR: 8.2/1000 PY, 95% CI 3.8 to 15.6/1000 PY; RR: 0.10, 95% CI 0.01 to 0.75).

Adjusted effect of OST on deaths in custody

In the first 4 weeks of a prison episode, the all-cause mortality hazard was 94% lower while in OST, compared to being out of OST (adjusted HR (AHR): 0.06; 95% CI 0.01 to 0.48), regardless of gender, indigenous status, age, incarceration history or offending history (table 4). A similar finding was observed in relation to unnatural deaths; during the first 4 weeks of a prison episode, while in OST, the hazard of unnatural death was 93% lower than while not in OST (AHR 0.07; 95% CI 0.01 to 0.59).

Across all time in prison, the all-cause mortality hazard during periods of OST was reduced by threequarters compared with periods not in OST (AHR: 0.26; 95% CI 0.13 to 0.50), regardless of gender, indigenous status, age, incarceration history or offending history. Across all time in prison, compared with periods not in OST, the hazard of unnatural death was 87% lower while in OST (AHR: 0.13; 95% CI 0.05 to 0.35).

DISCUSSION

We have demonstrated a very strong association between receipt of OST and lowered mortality among opioiddependent prisoners. After adjusting for demographic and criminal history factors, compared to time not in OST, being in OST was associated with a 74% lower hazard of dying in prison. The association between mortality risk and OST was driven by the reduction in unnatural deaths, with periods in OST associated with an 87% decrease in mortality hazard compared to time not in OST. One-third of deaths in prison in this cohort occurred during the first 4 weeks of incarceration; in these initial weeks in prison, compared to time not in OST, being in OST was associated with a 94% lower allcause mortality hazard and a 93% lower hazard of unnatural death. Although the CIs around the HRs were wide, even the uppermost limits of the intervals suggested substantial reductions in mortality while in OST. For example, across all time in prison, there was at least a 65% reduction in unnatural mortality hazard while in OST compared to time not in OST.

The cohort included in this study was drawn from a large, statewide administrative dataset of people receiving treatment for opioid dependence. Although precise data are not available on the representativeness of this cohort, in sentinel surveillance studies of people who inject drugs in NSW (98% of whom have a history of illicit opioid use), almost 60% of participants are currently in OST, and more than 80% have a history of OST,²⁰ suggesting that the cohort is highly representative of the NSW opioid-dependent population. The subset of this population that had been incarcerated was included

	Total time in prison	n			First 4 weeks in prison	ison		
	All-cause deaths		Unnatural deaths		All-cause deaths		Unnatural deaths	
Predictor	HR (95% CI)	Adjusted HR (95% CI)	HR (95% CI)	Adjusted HR (95% CI)	HR (95% CI)	Adjusted HR (95% CI)	HR (95% CI)	Adjusted HR (95% CI)
Receipt of OST*	0.25 (0.13 to 0.48)	0.25 (0.13 to 0.48) 0.26 (0.13 to 0.5)	0.15 (0.06 to 0.39)	0.13 (0.05 to 0.35)	0.15 (0.06 to 0.39) 0.13 (0.05 to 0.35) 0.07 (0.01 to 0.54) 0.06 (0.01 to 0.48) 0.08 (0.01 to 0.59) 0.07 (0.01 to 0.53)	0.06 (0.01 to 0.48)	0.08 (0.01 to 0.59)	0.07 (0.01 to 0.53)
Male	1.21 (0.48 to 3.04)	1.21 (0.48 to 3.04) 0.94 (0.36 to 2.41) 0.75 (0.29 to 1.96) 0.57 (0.22 to 1.54)	0.75 (0.29 to 1.96)	0.57 (0.22 to 1.54)	0.66 (0.22 to 2.04) 0.54 (0.16 to 1.76) 0.48 (0.15 to 1.57) 0.34 (0.1 to 1.18)	0.54 (0.16 to 1.76)	0.48 (0.15 to 1.57)	0.34 (0.1 to 1.18)
Indigenous	0.64 (0.34 to 1.18)	0.64 (0.34 to 1.18) 0.84 (0.44 to 1.59) 0.92 (0.44 to 1.9)		1.02 (0.48 to 2.21)	1.02 (0.48 to 2.21) 0.70 (0.24 to 1.98) 0.71 (0.23 to 2.13) 1.13 (0.37 to 3.46) 0.98 (0.3 to 3.18)	0.71 (0.23 to 2.13)	1.13 (0.37 to 3.46)	0.98 (0.3 to 3.18)
Age at prison reception	1.06 (1.03 to 1.1)	1.06 (1.03 to 1.1) 1.05 (1.01 to 1.09) 1.03 (0.98 to 1.07)	1.03 (0.98 to 1.07)	1.02 (0.97 to 1.07)	1.00 (0.93 to 1.07)	1.00 (0.93 to 1.07) 0.98 (0.91 to 1.06) 0.99 (0.91 to 1.07) 0.99 (0.91 to 1.07)	0.99 (0.91 to 1.07)	0.99 (0.91 to 1.07)
No of prior incarcerations	1.01 (0.91 to 1.12)	1.01 (0.91 to 1.12) 1.09 (0.99 to 1.21) 1.10 (0.97 to 1.24) 1.13 (0.99 to 1.28)	1.10 (0.97 to 1.24)	1.13 (0.99 to 1.28)		0.94 (0.79 to 1.12) 0.96 (0.81 to 1.14) 0.99 (0.78 to 1.24) 1.01 (0.8 to 1.27)	0.99 (0.78 to 1.24)	1.01 (0.8 to 1.27)
Duration of prior	0.98 (0.96 to 1)	0.98 (0.96 to 1)	1.01 (0.98 to 1.04)	1.01 (0.98 to 1.04) 1.01 (0.98 to 1.04)	0.98 (0.95 to 1.01) 0.99 (0.95 to 1.02) 1.02 (0.97 to 1.06) 1.02 (0.97 to 1.07)	0.99 (0.95 to 1.02)	1.02 (0.97 to 1.06)	1.02 (0.97 to 1.07)
incarcerations†								
Any prior drug offences	0.95 (0.53 to 1.7)	0.95 (0.53 to 1.7) 0.97 (0.53 to 1.76) 0.96 (0.46 to 2.01) 0.92 (0.43 to 1.95) 1.12 (0.36 to 3.51) 1.21 (0.38 to 3.82) 0.84 (0.25 to 2.78) 0.91 (0.27 to 3.07)	0.96 (0.46 to 2.01)	0.92 (0.43 to 1.95)	1.12 (0.36 to 3.51)	1.21 (0.38 to 3.82)	0.84 (0.25 to 2.78)	0.91 (0.27 to 3.07)
Any prior property offences	0.47 (0.25 to 0.92)	0.47 (0.25 to 0.92) 0.76 (0.37 to 1.57) 0.71 (0.29 to 1.75) 0.81 (0.31 to 2.14) 0.59 (0.13 to 2.69) 0.70 (0.15 to 3.36) 1.04 (0.13 to 8.39) 1.15 (0.14 to 9.67)	0.71 (0.29 to 1.75)	0.81 (0.31 to 2.14)	0.59 (0.13 to 2.69)	0.70 (0.15 to 3.36)	1.04 (0.13 to 8.39)	1.15 (0.14 to 9.67)
Any prior violent offences	0.53 (0.3 to 0.94)	0.53 (0.3 to 0.94) 0.69 (0.37 to 1.26) 0.65 (0.31 to 1.34) 0.70 (0.32 to 1.5)	0.65 (0.31 to 1.34)	0.70 (0.32 to 1.5)	0.49 (0.18 to 1.33)	0.49 (0.18 to 1.33) 0.49 (0.18 to 1.37) 0.64 (0.2 to 1.98)	0.64 (0.2 to 1.98)	0.56 (0.17 to 1.83)
*Time-dependent variable.								
THR refers to change in hazard associated with 28-day increase in duration of prior incarcerations.	d associated with 28-	day increase in durat	tion of prior incarcera	ations.				
OST, opioid substitution treatment	nent.							

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in this study. As an observational study, cohort members were not randomly allocated to treatment. Although there were some differences between prisoners who did and did not receive OST, these variables did not affect the association between OST and mortality. We did not have clinical indicators of current opioid dependence, and it is possible that some members of our cohort may have ceased all opioid use, thereby reducing their baseline mortality risk. However, given the chronic and relapsing nature of opioid dependence,¹⁵ ¹⁶ this is unlikely to have been the case for any substantial proportion of the cohort. Inclusion of opioid-abstinent individuals would, at any rate, result in conservative mortality estimates. Finally, we used the first episode of OST as a proxy for the onset of opioid dependence. It is possible that some cohort members had previous incarcerations during which they were opioid dependent, but had not vet entered OST.

To the best of our knowledge, this is the first study of its kind. Prior studies have reported on all-cause or cause-specific mortality in prison among total prisoner populations.^{4 21 22} We have extended on these studies by analysing mortality in opioid-dependent prisoners-a group with known mortality risk factors. Compared with opioid-dependent populations at liberty, mortality rates were low in opioid-dependent prisoners.¹⁰ This is most likely a result of limited access to illicit opioids and low exposure to other common causes of death in this population, such as motor vehicle accidents.²³ Highlighting the vulnerability of this population, however, 74% of deaths in prison were due to unnatural causes, compared to 48-59% of deaths in studies of total prisoner populations.^{2–5} The great majority of deaths occurred while prisoners were not receiving OST. Although this protective effect of OST against mortality is known,¹⁰ this is the first time that this association has been observed in a prison setting. Prior studies of OST in correctional settings, including a clinical trial²⁴ and pro-spective cohort studies,^{25 26} have not reported any deaths during OST, but these were not powered to detect differences in mortality rates during periods in and out of treatment. Receipt of OST in prison also reduces illicit opioid use and injecting drug use,^{24 27} and is associated with reduced institutional drug charges.²⁸ Opioid-dependent prisoners who receive OST are more likely than their untreated peers to enrol in OST following release,²⁹ and continuation of OST on release is associated with reduced reincarceration³⁰ and reduced postrelease mortality.³¹

Although we have demonstrated a very strong protective effect of OST against death in prison for opioiddependent prisoners, several questions remain to be answered. This study was undertaken in a jurisdiction with relatively high OST coverage.¹⁴ Our understanding of the relationship between OST and deaths in prison could benefit from further observational or simulation studies that assess the impact of varying levels of OST coverage on deaths. Further, this study was not able to determine why it is that OST should be associated with reduced unnatural deaths. Given that drug withdrawal symptoms have been implicated as a trigger for suicide in the first week of prison,^{7 8} the association may, in part at least, be due to alleviation or prevention of opioid withdrawal. If it is the case that averting withdrawal symptoms is sufficient to prevent some prison suicides, there are important implications for the clinical management of withdrawal symptoms of prisoners dependent on nonopioid drugs. Research examining why there are fewer suicides among opioid-dependent prisoners in OST is needed, as is research examining whether management of withdrawal from non-opioid drugs has a similar impact on suicides in prison.

The implication of these findings for correctional authorities and prison medical providers is clear; ensuring a high coverage of OST in correctional settings will help to minimise unnatural deaths among opioiddependent prisoners. Given the particularly high mortality risk observed during the first 4 weeks of incarceration, it is critical that opioid-dependent prisoners can access OST in a timely manner. Opioid dependence and enrolment in OST immediately prior to entry to prison should be assessed during intake medical examinations. Prisoners enrolled in OST prior to prison entry should be able to continue this treatment without interruption, and other opioid-dependent prisoners should be assessed for OST and offered treatment if clinically indicated. Although there are challenges to the implementation of OST programmes in prisons and other correctional settings, these can be managed through strong executive leadership, ongoing training and education for health and custodial staff, and careful attention to issues of safety and security.³²

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Contributors SL completed the analysis and led the writing and is responsible for the overall content as guarantor. TD provided statistical advice and contributed to the writing. NG, MF, LB, AG and JK contributed to the writing. LD oversaw the larger project for which these data were obtained and contributed to the writing.

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