Title Page

1

- 2 Tuberculosis treatment outcomes after transfer or release from incarceration: A
- 3 retrospective cohort study from Brazil
- 4 Authors and Institutional Addresses
- 5 Yasmine Mabene¹, José Bampi², Everton Ferreira Lemos^{3,4}, Roberto de Oliveira^{3,5},
- 6 Crhistinne Gonçalves^{6,7}, Maria de Lourdes Delgado Alves⁸, Maridiane Coutinho
- 7 Echevarria⁸, Julio Croda^{4,9,10}, Jason R Andrews^{1*}, Yiran E Liu^{1,11,12*}
- 8 *these authors contributed equally
- 9 ¹ Division of Infectious Diseases and Geographic Medicine, Department of Medicine,
- 10 Stanford University, Stanford, California, USA.
- 11 ² Federal University of Mato Grosso do Sul, Infectious and Parasitic Diseases Program,
- 12 Campo Grande, Brazil.
- 13 ³ State University of Mato Grosso do Sul, Dourados, Mato Grosso do Sul, Brazil
- 14 ⁴Fundação Oswaldo Cruz, Science, Technology, Production, and Innovation in Public
- 15 Health, Campo Grande, Brazil.
- ⁵ Post Graduate Program in Health Sciences, Federal University of Grande Dourados,
- 17 Dourados, Dourados, Brazil.
- 18 ⁶ School of Medicine, Federal University of Mato Grosso do Sul, Campo Grande, Mato
- 19 Grosso do Sul, Brazil

21

22

23

24

25

26

27

28

29

30

31

32

33

34

35

36

37

38

39

⁷ Mato Grosso do Sul State Secretariat of Health ⁸ Division of Prison Health Assistance, Agência Estadual de Administração do Sistema Penitenciário, Campo Grande, Mato Grosso do Sul, Brazil ⁹Federal University of Mato Grosso do Sul, Clinical Medicine, Campo Grande, Brazil. ¹⁰ Yale University, Epidemiology of Microbial Diseases, New Haven, United States of America. ¹¹ Department of Epidemiology and Population Health, Stanford University, Stanford, California, USA. ¹² SEICHE Center for Health and Justice, Department of Internal Medicine, Yale School of Medicine, New Haven, USA **Corresponding Author** Yiran Liu, yiran.e.liu@yale.edu **Abstract Background** Tuberculosis (TB) disproportionately affects people deprived of liberty. Prior studies have shown higher TB treatment completion rates among incarcerated individuals compared to the general population. However, little is known about how incarcerationrelated movements such as transfers between facilities or releases to the community affect TB treatment outcomes. Methods

We linked person-level incarceration data with TB notifications data from the Notifiable Disease Information System for the Brazilian state of Mato Grosso do Sul between January 2006 and December 2018. We constructed a cohort of individuals newly diagnosed with drug-susceptible TB who initiated treatment while incarcerated. We compared treatment outcomes between individuals who remained in the same carceral facility and those who were transferred to other facilities or released from incarceration during treatment. We computed the covariate-adjusted relative risk of unfavorable treatment outcomes for individuals transferred or released during treatment.

Results

We identified 1,274 individuals who initiated TB treatment while incarcerated. Of these individuals, 849 (66.6%) remained in the same carceral facility, 259 (20.3%) were transferred to other facilities, and 166 (13.0%) were released to the community during treatment. Among those who remained in the same carceral facility, 72.3% (614/849) were successfully treated within eight months following treatment initiation. In contrast, only 61.0% (158/259) of those who were transferred and 49.4% (82/166) of those who were released achieved TB treatment success within eight months. After adjusting for covariates, the risk of unfavorable treatment outcomes was 1.4 (95% CI: 1.2 to 1.7) times as high for individuals transferred to other facilities and 1.6 (95% CI: 1.3 to 2.0) times as high for individuals released from incarceration, compared to those who remained incarcerated in the same facility during treatment. For individuals released less than two months into treatment, the risk of unfavorable treatment outcomes was twice as high (adjusted relative risk [aRR]:2.1, 95% CI: 1.6 – 2.6).

Conclusions

Transfers between facilities and releases from incarceration are common and may pose barriers to TB treatment completion. Strategies for ensuring continuity of care across carceral facilities and between prison and community health systems are urgently needed to improve TB outcomes for individuals impacted by incarceration.

Keywords

Incarceration, Prisons, Tuberculosis, Treatment, Cascade of care, Linkage to care, Continuity of care, Persons deprived of liberty, Brazil

Background

70

71

72

73

74

75

76

77

78

79

80

81

82

83

84

85

86

87

88

89

90

91

92

In 2023, 10.8 million people fell ill with tuberculosis (TB) worldwide, and 1.25 million people died from TB [1]. Persons deprived of liberty (PDL) face increased risk of TB compared to the general population due to factors such as prison overcrowding. poor ventilation, undernutrition, and limited healthcare access [2-3]. High rates of TB in prisons can also spill over into communities through prison staff, visitors, and individuals released from incarceration [4-5]. Targeted efforts to address TB in prisons are needed to reduce TB morbidity and mortality for PDL and to progress toward population-wide TB elimination. In 2023, the United Nations High-Level Meeting established global targets aimed at ensuring a 90% treatment completion rate for all people with TB [6]. Achieving these targets is especially crucial for vulnerable populations, including PDL. Although previous studies have reported higher TB treatment completion rates for PDL compared to the general population, these rates still fall short of the UN's 90% target [7-12]. More research is needed to identify barriers and facilitators to treatment completion among PDL to improve TB outcomes in this population. Individuals who begin TB treatment while incarcerated may face interruptions to care following transfers between carceral facilities or release from incarceration. Research has shown this for incarcerated people living with HIV, finding that releases from jails and prisons may disrupt the HIV continuum of care [13-15]. Similar challenges have been observed among people undergoing treatment for substance use disorders, including low rates of treatment retention post-release [16-17]. Studies in the United

States on TB preventive therapy for people with latent tuberculosis infection (LTBI) have

found lower completion rates for individuals released during treatment, with some studies reporting rates as low as 20% [18-20]. Far less is known about how carceral movements impact treatment continuity and outcomes for individuals with active TB. Studies in Uganda and Zambia found treatment disruptions and increased rates of loss to follow up for incarcerated individuals who were transferred or released during treatment, as well as for those residing in prisons with shorter incarceration sentences [21-23]. However, there remains a significant knowledge gap regarding TB treatment outcomes and linkage to care among individuals experiencing carceral movements in other regions and settings.

In Brazil, incarceration is the leading driver of the TB epidemic, accounting for approximately 37% of new cases [24]. Brazil, which has the third largest prison population in the world, has the highest number of TB cases among incarcerated individuals [25-26]. Despite this, carceral movements for individuals diagnosed with TB in Brazil are not well documented, and the impact of carceral movements on TB treatment in Brazil remains unknown. PDL in Brazil may reside in closed prisons, semi-open prisons (where individuals leave prison during the day for work and return at night), or in police detention. Varying frequencies of movements and healthcare resources across carceral facilities may affect treatment outcomes and continuity of care. As both TB incidence and incarceration rates rise in Brazil, it is essential to investigate these potential effects to inform strategies for improving TB treatment outcomes for incarcerated individuals.

In this study, we link individual-level incarceration and TB notifications data for the Brazilian state of Mato Grosso do Sul to assess the impact of carceral movements on TB treatment outcomes. We examine the proportion of individuals experiencing transfers or releases during TB treatment and determine the relative risk of unfavorable treatment outcomes compared to individuals who remain in the same carceral facility during treatment.

Methods

Study design

We conducted a retrospective cohort study of individuals who were newly diagnosed with drug-sensitive TB (pulmonary and extra-pulmonary) and who initiated treatment while incarcerated in Mato Grosso do Sul state, Brazil, between January 1st, 2006 and June 30th, 2018. We examined TB treatment outcomes among individuals who were transferred to other carceral facilities and/or released from incarceration during the sixmonth treatment period, compared to those remaining in the same carceral facility. We included individuals who were incarcerated in closed prisons, semi-open prisons, or police detention at the time of treatment initiation. We excluded individuals with prior TB diagnoses, those with drug resistance at the time of diagnosis, and those under the age of 18. The study was approved by the institutional review boards (IRBs) of Federal University of Mato Grosso do Sul (UFMS) (#7.060.054) and Stanford University (#50466).

Data sources

We accessed statewide individual-level data on TB notifications and treatment outcomes between January 1st, 2006 and June 30th, 2020 from *Sistema de Informação de Agravos de Notificação* (SINAN). To determine incarceration status and movements

139

140

141

142

143

144

145

146

147

148

149

150

151

152

153

154

155

156

157

158

159

160

during and after TB treatment initiation, we linked SINAN with Sistema Integrado de Gestão Operacional (SIGO), a database containing movements within the carceral system (i.e., entries, transfers, and releases) in Mato Grosso do Sul from January 1st, 2006 to June 30th, 2018. To account for incomplete data on deaths during the follow-up period, we performed additional linkage with the mortality database, Sistema de Informações Sobre Mortalidade (SIM), accessed between January 1st, 2006 and June 30th, 2020. Data processing and linkage For the incarceration (SIGO) and TB (SINAN) databases, where an individual could have multiple distinct entries (corresponding to multiple carceral movements, TB diagnoses, etc.), we identified records belonging to unique individuals using approximate string matching over name and mother's name, with additional comparisons of identification numbers and dates of birth when available (Text S1, Additional File 1). Approximate string matching was performed using stringdist in R version 4.4.1 [27-28]. We linked unique individuals across the incarceration and TB databases using approximate string matching on shared identifiers (name and mother's name), resulting in a sensitivity of 90% and specificity of 95% (Text S1: SIGO and SINAN database linkage approach, Additional File 1). We performed subsequent filtering steps to exclude false positive matches (Text S1: SINAN and SIGO cohort filtering, Additional File 1). Among matched individuals, we then searched for deaths within the mortality (SIM) database using name, mother's name, and date of birth. Although the TB (SINAN) database includes information on deaths, we additionally matched individuals to the

mortality database (SIM) to account for unreported deaths (Text S1: SIGO, SINAN, and SIM database linkage approach, Additional File 1). Exact criteria for database linkage can be found in (Figure S1, Additional File 1).

Group assignment and outcome determination

161

162

163

164

165

166

167

168

169

170

171

172

173

174

175

176

177

178

179

180

181

182

183

We categorized individuals by whether they experienced carceral movements (transfers to other facilities or release from incarceration) within six months following treatment initiation. Each individual was assigned to one of three groups: 1) individuals who remained in the same carceral facility throughout the treatment period (the "stationary" group); 2) individuals who were transferred to another carceral facility at least once but were not released (the "transferred" group); and 3) individuals who were released from incarceration (the "released" group; this group may also include individuals who escaped incarceration). To avoid exposure misclassification, only movements that occurred prior to ascertainment of treatment outcomes were considered in group assignment (Text S1: Group and Outcome Ascertainment, Additional File 1). In an additional analysis, we stratified the released group into two subgroups based on the timing of release (released within or after two months following treatment initiation). In the primary analysis, we evaluated TB treatment outcomes at eight months post treatment initiation. While treatment completion takes six months, we used the eightmonth point to account for delays in reporting treatment outcomes. We additionally assessed treatment outcomes at two years post treatment initiation. We considered a treatment outcome to be positive if the individual achieved treatment success; all other outcomes were considered unfavorable treatment outcomes. We categorized treatment outcomes in our study using the SINAN guidelines as follows [29]:

185

186

187

188

189

190

191

192

193

194

195

196

197

198

199

200

201

202

- 1. **Treatment success:** Patients who obtained two negative sputum smears (one at any follow-up month and one at the end of treatment) or those who completed treatment without evidence of treatment failure and were discharged based on clinical and radiological criteria, in accordance with Brazilian guidelines [29].
- 2. **No case status update:** Patients whose case had no updates in the TB (SINAN) database following treatment initiation.
- 3. No case status update following treatment discontinuation/transfer and resumption of care: Patients who discontinued treatment or transferred health facilities and were later linked to care (resumed treatment) without further case updates in the TB (SINAN) database.
- Treatment discontinuation with no follow-up record: Patients who interrupted treatment for 30 or more consecutive days and had no subsequent updates in the TB (SINAN) database.
- Transferred health facility with no follow-up record: Patients who were transferred to another health facility and had no subsequent updates in the TB (SINAN) database.
- 6. **Drug-resistant TB (DR-TB):** Patients with confirmed resistance to any antituberculosis drug after treatment initiation.
- 7. **Death (TB):** Patients with tuberculosis as the cause of death.
- 203 8. **Death (non-TB):** Patients with cause of death other than tuberculosis.
- Change of diagnosis/regimen: Patients whose initial TB diagnosis was
 incorrect or patients who changed treatment regimen due to drug intolerance or
 toxicity

Statistical Analysis

207

208

209

210

211

212

213

214

215

216

217

218

219

220

221

222

223

224

225

226

227

228

We used the chi-square test to compare characteristics across groups and to assess differences in the proportion of treatment success across carceral movement groups. To determine the effect of carceral movements on treatment outcomes, we performed multivariable Poisson regression, adjusting for race, age, education, alcohol use disorder, mental illness, incarceration history in the past two years (total duration and unique entries into the carceral system), and facility type at treatment initiation (closed prison, semi-open prison, police detention) [30]. We included facility type at treatment initiation as a covariate due to variation across carceral facilities in healthcare, security level, and movements. Covariates with less than 25% missingness were imputed using missForest version 1.5 in R (Table S1, Additional File 1) [31]. For covariates with greater missingness, we retained "unknown/missing" as a level in the regression. For the regression analysis, we restricted our cohort to individuals who initiated TB treatment after January 1st 2007. This enabled adjustment for incarceration history in the past two years. We conducted two sensitivity analyses. First, we excluded from the regression individuals who died from non-TB causes within the first six months following treatment initiation. This exclusion was made to mitigate potential survival-induced nonexchangeability bias, as individuals in the transferred or released group had to survive long enough to experience carceral movements. In a second sensitivity analysis, we adjusted for individual carceral facility in the regression model to account for potential differences in healthcare across facilities.

Results

229

230

231

232

233

234

235

236

237

238

239

240

241

242

243

244

245

246

247

248

249

250

251

Baseline Cohort Characteristics

After database linkage, we obtained a cohort of 1,274 individuals who initiated TB treatment while incarcerated between January 1, 2006 and June 30, 2018 (Figure 1). Our cohort primarily consisted of male individuals under the age of 30 (Table 1). Nearly half of participants were Black or mixed race (N=584, 45.8%), and the majority (N=690, 54.2%) indicated not having completed high school. Additionally, 86 (6.8%) individuals reported having alcohol use disorder, and less than 1% (9, 0.7%) were reported to have mental illness. The majority of individuals (950, 74.6%) resided in closed prisons at the time of treatment initiation. In the two years preceding treatment initiation, 1021 (80.1%) had been incarcerated for more than one year and approximately one in four individuals (333, 26.1%) had experienced multiple incarcerations. The majority of individuals in the cohort (849, 66,6%) remained in the same carceral facility in the six months following TB treatment initiation. Another 259 (20.3%) individuals were transferred to other carceral facilities, and 166 (13.0%) were released to the community within six months of treatment initiation (Table 1). Individuals who initiated treatment in semi-open prisons were more likely to be released to the community during treatment compared to those who initiated treatment in any other facility type (Figure S2, Additional File 1). At the time of treatment initiation, a higher proportion of individuals in the released group had spent less than one of the past two years incarcerated (31.9%), compared to individuals in the stationary (18.4%) and transferred (17.0%) groups (p value <.001) (Table 1). Lastly, the stationary group had a lower proportion of individuals with multiple prior incarcerations in the last two years

(22.5%), compared to the transferred (34.7%) and released (31.3%) groups (p value <.001). The distributions of the year of TB notification and carceral facility at time of treatment initiation can be found in Tables S2-S3, Additional File 1.

Table 1. Demographic and clinical characteristics of individuals grouped by carceral movement during TB treatment.

Characteristics	Categories	Stationar y N=849	Transfer N=259	Release N=166	p-value	
Facility type	Closed prison	659 (77.6%)	205 (78.5%)	86 (51.8%)	<.001	
	Semi-open prison	100 (11.8%)	30 (11.5%)	58 (34.9%)		
	Police detention	90 (10.6%)	24 (9.2%)	22 (13.3%)		
Sex	Male	827 (97.4%)	254 (98.1%)	158 (95.2%)	0.18	
	Female	22 (2.6%)	5 (1.9%)	8 (4.8%)		
Age	18-29	419 (49.3%)	148 (57.1%)	76 (45.8%)	0.14	
	30-44	335 (39.5%)	94 (36.3%)	69 (41.6%)		
	>45	92 (10.8%)	17 (6.7%)	20 (12.0%)		
	Not reported	3 (0.4%)	0 (0.0%)	1 (0.6%)		
Race	White	281 (33.1%)	82 (31.7%)	50 (30.1%)	0.75	
	Mixed Race	331 (39.0%)	93 (35.9%)	73 (44.0%)		
	Black	59 (6.9%)	18 (6.9%)	10 (6.0%)		
	Asian	12 (1.4%)	4 (1.5%)	1 (0.6%)		
	Indigenous	15 (1.8%)	8 (3.1%)	5 (3.0%)		
	Not reported	151 (17.8%)	54 (20.8%)	27 (16.3%)		
Education	Did not complete high school	453 (53.4%)	152 (58.7%)	85 (51.2%)	0.08	
	Completed high school or more	28 (3.3%)	9 (3.5%)	12 (7.2%)		
	Not reported	368 (43.3%)	98 (37.8%)	69 (41.6%)		
Alcohol use disorder	No	591 (69.6%)	181 (69.9%)	110 (66.3%)	0.45	
	Yes	53 (6.2%)	16 (6.2%)	17 (10.2%)		
	Not reported	205 (24.1%)	62 (23.9%)	39 (24.7%)		
Mental illness	No	647 (76.2%)	202 (78.0%)	122 (73.5%)	0.39	
	Yes	5 (0.6%)	1 (0.4%)	3 (1.8%)		

	Not reported	197 (23.2%)	56 (21.6%)	41 (24.6%)	
Total time incarcerated in	<1 year	156 (18.4%)	44 (17.0%)	53 (31.9%)	<.001
past two years	1-2 years	693 (81.6%)	215 (83.0%)	113 (68.1%)	
Number of distinct incarcerations* in past two years	1	658 (77.5%)	169 (65.3%)	114 (68.7%)	<.001
	>1	191 (22.5%)	90 (34.7%)	52 (31.3%)	

Facility type refers to location at time of TB notification. *Distinct incarcerations defined as periods of incarceration following time in the community. P-values are computed using chi-square test of independence with Monte Carlo simulations for categories with frequencies <5.

TB treatment outcomes by carceral movement group

257

258

259

260

261

262

263

264

265

266

267

268

269

270

271

272

273

274

275

Across the entire cohort, 854 (67.0%) of individuals successfully completed treatment within eight months following treatment initiation. Another 179 (14.0%) individuals were lost to follow up, 187 (14.7%) had no case status updates, and 38 (3.0%) died (Table 2). Among those who remained in the same carceral facility in the six months following TB notification, 614 (72.3%) had successful treatment, 84 (9.9%) were lost to follow up, 110 (13.0%) had no case status updates, and 33 (3.9%) died. Compared to those who remained in the same carceral facility, those who experienced movements during treatment had significantly lower treatment success rates: 61.0% in the transferred group and 49.4% in the released group (p-value <.001). At two years following treatment initiation, success rates remained lower for those transferred (64.9%) or released (54.2%) compared to those who remained stationary (73.7%) (p-value <.001) (Table S4, Additional File 1). Additionally, the released group was less likely to have updates to their treatment status reported in SINAN within eight months of notification, with 22.3% missing case status updates compared to only 13.0% in the stationary group (p-value <.01).

The proportion of individuals who were lost to follow up after being referred to a different health facility was over three times as high for individuals who were transferred to other carceral facilities (11.6%) compared to those who remained in the same location (3.1%) (p-value <.001). For individuals who were released, this proportion was more than twice as high (6.6%) (p-value = .04). Among those released from carceral facilities who were referred to another health facility or experienced treatment discontinuation within eight months of notification, 18 (36.7%) were subsequently linked to care, and 10 (20.4%) successfully completed treatment within the state over a two-year period.

Table 2. TB treatment outcomes eight months following treatment initiation

Treatment Outcome	Stationary Group (N=849)	Transferred Group (N=259)	Released Group (N=166)
Treatment success	614 (72.3%)	158 (61.0%)	82 (49.4%)
No case status update	93 (11.0%)	28 (10.8%)	28 (16.9%)
Referred to different health facility with no follow-up record	26 (3.1%)	30 (11.6%)	11 (6.6%)
Treatment discontinuation with no follow-up record	58 (6.8%)	25 (9.6%)	29 (17.5%)
Referred to different health facility/treatment discontinuation and resumed care with no case status update	17 (2.0%)	12 (4.6%)	9 (5.4%)
Death TB	12 (1.4%)	1 (0.4%)	1 (0.6%)
Death non-TB	21 (2.5%)	1 (0.4%)	2 (1.2%)
DR-TB	1 (0.1%)	1 (0.4%)	1 (0.6%)
Change of diagnosis/regimen	7 (0.8%)	3 (1.2%)	3 (1.8%)

Factors associated with unfavorable treatment outcomes

289

290

291

292

293

294

295

296

297

298

299

300

301

302

303

304

305

306

307

308

309

310

After adjusting for potential confounders, transfer between carceral facilities during TB treatment was associated with a 1.4-fold (95% CI: 1.2 to 1.7) increased risk of unfavorable treatment outcomes compared to the stationary group (Figure 2). For individuals released from incarceration during treatment, the covariate-adjusted risk of unfavorable treatment outcomes was 1.6 (95% CI: 1.3 to 2.0) times as high as individuals in the stationary group. Results were similar in a sensitivity analysis excluding individuals who died from causes other than TB within six months following treatment initiation (Table S6, Additional File 1). Beyond carceral movements, residing in a semi-open prison at the time of treatment initiation was associated with increased risk of unfavorable treatment outcomes (aRR: 1.6, 95% CI: 1.3 to 1.9) (Figure 2). However, this association did not remain after adjusting for individual carceral facility units in the regression (Table S6, Additional File 1). Additionally, higher education level was associated with a significantly reduced risk of unfavorable treatment outcomes (aRR: 0.5, 95% CI: 0.3 to 0.9) (Figure S3, Additional File 1). Treatment outcomes based on time of release

Among individuals in the released group, 40.4% were released less than two months after treatment initiation, during the intensive phase of TB treatment (Figure S4, Additional File 1). The treatment success rate within eight months was significantly lower among individuals released less than two months after initiating TB treatment (35.8%) compared to those who were released two or more months into treatment

(58.6%) (p-value <.01) (Figure 3). Additionally, the proportion of individuals lost to follow up was 37.3% among those released less than two months after treatment initiation, compared to 24.2% among those released two months or later.

Release from incarceration within the first two months of TB treatment was associated with more than twice the risk of unfavorable treatment outcomes, compared to those who were stationary (aRR 2.1, 95%CI: 1.6 to 2.6) (Table S7, Additional File 1). For individuals released two months or more following treatment initiation, risk of unfavorable treatment outcomes was 1.3 (95% CI: 1.0 to 1.8) times that of the

Discussion

stationary group.

Timely diagnosis and treatment completion are essential to reduce morbidity and mortality among TB patients and reduce onward transmission, particularly in high-risk environments like prisons. However, in settings with limited prison healthcare resources and barriers to coordination of care between prisons and communities, carceral facility transfers and release from incarceration may disrupt continuity of care. In this study, we linked incarceration and TB databases to determine the effect of carceral movements on TB treatment outcomes in a Brazilian state. Our findings revealed that approximately one in three incarcerated individuals are transferred and/or released from the facility where they initiated treatment during the six-month TB treatment period. Both releases and transfers were associated with increased risk of unfavorable treatment outcomes, after adjusting for covariates. Individuals released less than two months into treatment had particularly low rates of successful treatment completion. These findings

334

335

336

337

338

339

340

341

342

343

344

345

346

347

348

349

350

351

352

353

354

355

underscore the need for strategies to enhance continuity of care for incarcerated individuals with TB who are transferred between carceral facilities or released from incarceration during treatment. Our findings provide novel insights into barriers to treatment completion among PDL in Brazil, identifying carceral movements as a major risk factor for unfavorable treatment outcomes in this population. Previous literature in the field has indicated that PDL have better TB treatment outcomes compared to the general population [7-12]. Although this may be true for a subset of patients who remain in the same facility during treatment, incarceration is dynamic, and facility transfers and releases are common. To date, studies in Brazil have been unable to test the effect of such movements on TB treatment outcomes due to the lack of detailed incarceration data in TB notifications databases. Our database linkage approach addresses these data limitations, enabling ascertainment of carceral movements during TB treatment and assessment of associated treatment outcomes. Here, we find low TB treatment success rates among individuals who are transferred or released from incarceration during treatment, comparable to those of other highly vulnerable groups including people living with HIV, people who are unhoused, and people who use drugs [32-36]. Moreover, the treatment success rate among all PDL in our cohort, regardless of carceral movement group, was lower than in previous studies from Brazil which report treatment success rates exceeding 80% among PDL [7,10]. This discrepancy may stem from differences in the inclusion or classification of missing outcomes: while many prior studies exclude individuals without reported outcomes, we classify the lack of status update in the TB database as an unfavorable outcome.

357

358

359

360

361

362

363

364

365

366

367

368

369

370

371

372

373

374

375

376

377

Addressing incomplete or delayed reporting in administrative databases is essential to improve monitoring and evaluation of TB treatment outcomes. Our study has several limitations. Missingness within variables in the TB database (SINAN) prevented us from including several covariates in our analysis, such as smoking or drug use, which have been found to be associated with TB treatment outcomes [37-39]. Furthermore, there may have been residual confounding by other socioeconomic or incarceration-related covariates that were not available in our data. Our analysis may be subject to survivorship bias, as individuals in the transferred or released groups had to survive long enough to experience carceral movements. If survival is associated with treatment success, this could bias our results towards the null, leading us to underestimate the relative risk of unfavorable treatment outcomes associated with carceral movements. However, the low proportion of deaths in our cohort suggests minimal impact. Moreover, in our sensitivity analysis excluding individuals with non-TB deaths within six months of treatment initiation, the estimates of relative risk associated with carceral movements remained largely unchanged (Table S6, Additional File 1). Another limitation is that individuals categorized as released in our analysis may have previously experienced transfers that contributed to treatment disruptions. We did not differentiate between individuals released directly from incarceration to the community and those who experienced one or more transfers prior to release. We also do not differentiate between individuals released following the end of their sentence and those who escaped from incarceration. Additionally, we were unable to obtain treatment outcome data for individuals who moved outside the state of

Mato Grosso do Sul during the follow-up period. Finally, the results of this study may not

generalize to other states or countries. Our findings highlight the urgent need for comprehensive strategies to improve continuity of care for individuals diagnosed with TB who are transitioning within and out of the carceral system. Ensuring timely transfer of medical records when individuals move to different carceral facilities is critical, especially for individuals transferred or released early during treatment. Referrals to community health care clinics, case management services, and discharge planning may play pivotal roles in facilitating continuity of care post-release [40-41]. In addition to improving system-level coordination, efforts to support individuals' basic needs and social reintegration are essential. Individuals who are released from incarceration may face social instability, marginalization, and stigma, all of which can hinder care-seeking and access [42-45]. In the present study, we did not distinguish between people who were released versus those who escaped from incarceration, but the latter may experience heightened barriers to accessing healthcare. Further work is needed to understand these challenges and develop strategies to ensure all individuals, regardless of legal status or incarceration history, are supported to access healthcare services without fear of stigma or punishment.

Conclusion

378

379

380

381

382

383

384

385

386

387

388

389

390

391

392

393

394

395

396

397

398

399

Our study found that both releases and transfers from carceral facilities were associated with increased risk of unfavorable treatment outcomes. Furthermore, individuals released earlier within their treatment regimen had higher loss to follow up rates. Our

401

402

403

404

405

406

407

408

409

410

411

412

413

414

415

416

417

418

419

420

421

Ethics approval and consent to participate

results highlight the need for strategies to ensure continuity of care for people with TB who are impacted by incarceration. These strategies can have broader implications for continuity of care after incarceration for health conditions beyond TB, a topic that remains poorly understood in Brazil and many other low- and middle-income countries [46-49]. List of abbreviations TB: Tuberculosis PDL: Persons deprived of liberty aRR: Adjusted relative risk SINAN: Sistema de Informação de Agravos de Notificação SIGO: Sistema Integrado de Gestão Operacional SIM: Sistema de Informações Sobre Mortalidade DR-TB: Drug-resistant Tuberculosis **Additional Materials** File name: Additional file 1 File format: additional_file.pdf Description: This file contains additional information on the methodology of our study as well as supplementary figures and tables. **Declarations**

422 The study was approved by the institutional review boards (IRBs) of Federal University 423 of Mato Grosso do Sul (UFMS) (#7.060.054) and Stanford University (#50466). 424 **Consent for publication** 425 Not applicable 426 Availability of data and materials 427 This study utilized datasets provided by state agencies in Mato Grosso do Sul, Brazil, 428 and we are not authorized to share them directly. The incarceration database was 429 provided by Agência Estadual de Administração do Sistema Penitenciário (AGEPEN, 430 https://www.agepen.ms.gov.br/). The tuberculosis and mortality databases were 431 provided by the Secretaria de Estado de Saúde (https://www.saude.ms.gov.br/). 432 433 **Competing interests** 434 JRA reports funding from the U.S. National Institutes of Health (R01 Al130058 and 435 K24Al182647). All other others declare no competing interests. 436 Funding. JRA is supported by grants from the U.S. National Institutes of Health (R01 437 AI130058 and K24AI182647). 438 **Authors' contributions** 439 YM, YEL, JRA, and JC conceived and designed the study. EFL, MLDA, MCE and CG 440 acquired the data. YM conducted the analysis and wrote the initial draft of the 441 manuscript. JRA, YEL, JC, JB, and EFL provided critical input on analyses. YM and YEL 442 contributed to writing subsequent drafts. All authors were involved in interpreting the 443 data and critically revising the manuscript. All authors approved the final manuscript. 444 445 **Acknowledgements**

446 Not applicable 447 448 References 449 1. World Health Organization. Global Tuberculosis Report 2023: Tuberculosis in 450 Prisons. 2023. https://www.who.int/teams/global-tuberculosis-programme/tb-451 reports/global-tuberculosis-report-2023/featured-topics/tb-in-prisons. Accessed 452 29 Oct 2024. 453 2. Fazel S, Baillargeon J. The health of prisoners. The Lancet. 2011 Mar 454 12;377(9769):956-65. 455 3. Dara M, Grzemska M, Kimerling ME, Reyes H, Zagorskiy A. Guidelines for 456 control of tuberculosis in prisons. 2009. 457 https://www.challengetb.org/publications/tools/ua/Guidelines_for_control_of_TB_i 458 n_prisons.pdf. Accessed 29 Oct 2024. 459 4. Sacchi FP, Praça RM, Tatara MB, Simonsen V, Ferrazoli L, Croda MG, et.al. 460 Prisons as reservoir for community transmission of tuberculosis, Brazil. Emerging 461 infectious diseases. 2015 Mar;21(3):452-455. 462 5. Warren JL, Grandjean L, Moore DA, Lithgow A, Coronel J, Sheen P, et.al. 463 Investigating spillover of multidrug-resistant tuberculosis from a prison: a spatial 464 and molecular epidemiological analysis. BMC medicine. 2018 465 Aug;doi:10.1186/s12916-018-1111-x 466 6. The Stop TB Partnership. UNHLM 2023 Country Targets: Achieving the UNHLM 467 treatment targets country by country. 2023.

468 https://www.stoptb.org/sites/default/files/imported/document/country_targets-469 combined.pdf. Accessed 25 Feb 2025. 470 7. O'Marr J.M. Goncalves C. Arakaki-Sanchez D. Pelissari DM. Costa FD. Croda J. 471 et.al. The effect of incarceration on TB treatment outcomes. The International 472 Journal of Tuberculosis and Lung Disease. 2022 Mar 1;26(3):252-258. 473 8. Singano V, Kip E, Ching'ani W, Chiwaula L. Tuberculosis treatment outcomes 474 among prisoners and general population in Zomba, Malawi, BMC Public Health. 475 2020 May 15;doi:10.1186/s12889-020-08841-z 476 9. Lima LV, Pavinati G, Palmieri IG, Vieira JP, Blasque JC, Higarashi IH, et. al. 477 Factors associated with loss to follow-up in tuberculosis treatment in Brazil: a 478 retrospective cohort study. Revista Gaúcha de Enfermagem. 2023 Nov 479 24;44:e20230077. 480 10. Ryuk DK, Pelissari DM, Alves K, Oliveira PB, Castro MC, Cohen T, et. al. 481 Predictors of unsuccessful tuberculosis treatment outcomes in Brazil: an analysis 482 of 259,484 patient records. BMC Infectious Diseases. 2024 May 27; 483 doi:10.1186/s12879-024-09417-7. 484 11. Faust L, Caceres-Cardenas G, Martinez L, Huddart S, Vidal JR, Corilloclla-Torres 485 R, et.al. Tuberculosis case notifications and outcomes in Peruvian prisons prior 486 to and during the COVID-19 pandemic: a national-level interrupted time series 487 analysis. The Lancet Regional Health–Americas. 2024 May 1;33(100723). 488 12. Medina A, Sussman J, Sosa N, Valdez M, Andrews JR, Croda J, et. al. The 489 concentration of tuberculosis within Paraguay's incarcerated and Indigenous

490 populations, 2018–2022. medRxiv. 2024 Jun 1; 491 doi:10.1101/2024.05.31.24308287 492 13. Westergaard RP, Kirk GD, Richesson DR, Galai N, Mehta SH. Incarceration 493 predicts virologic failure for HIV-infected injection drug users receiving 494 antiretroviral therapy. Clinical Infectious Diseases. 2011 Oct;53(7):725-31. 495 Baillargeon J, Giordano TP, Rich JD, Wu ZH, Wells K, Pollock BH, Paar DP. 496 Accessing antiretroviral therapy following release from prison. Jama. 2009 Feb. 497 25;301(8):848-57. 498 15. Loeliger KB, Altice FL, Desai MM, Ciarleglio MM, Gallagher C, Meyer JP. 499 Predictors of linkage to HIV care and viral suppression after release from jails 500 and prisons: a retrospective cohort study. The lancet HIV. 2018 Feb;5(2):e96-501 e106. 502 16. Bachireddy C, Shrestha R, Bromberg DJ, Azbel L, Kurmanalieva A, Wegman M, 503 et. al. Methadone within prison and linkage to and retention in treatment upon 504 community release for people with opioid use disorder in Kyrgyzstan: Evaluation 505 of a national program. International Journal of Drug Policy. 2022 506 Mar;101:103558. 507 17. Dolan KA, Shearer J, White B, Zhou J, Kaldor J, Wodak AD. Four-year follow-up 508 of imprisoned male heroin users and methadone treatment: mortality, 509 re-incarceration and hepatitis C infection. Addiction. 2005 Jun;100(6):820-8. 510 18. Bock NN, Rogers T, Herron GD, Tapia JR, DeVoe B, Geiter LJ. Acceptability of 511 short-course rifampin and pyrazinamide treatment of latent tuberculosis infection 512 among jail inmates. Chest. 2001 Mar;119(3):833-7.

514

515

516

517

518

519

520

521

522

523

524

525

526

527

528

529

530

531

532

533

534

535

19. Lobato MN, Reves RR, Jasmer RM, Grabau JC, Bock NN, Shang N, et. al. Adverse events and treatment completion for latent tuberculosis in jail inmates and homeless persons. Chest. 2005 Apr;127(4):1296-303. 20. Nolan CM, Roll L, Goldberg SV, Elarth AM. Directly observed isoniazid preventive therapy for released jail inmates. American journal of respiratory and critical care medicine. 1997 Feb;155(2):583-6. 21. Schwitters A, Kaggwa M, Omiel P, Nagadya G, Kisa N, Dalal S, Tuberculosis incidence and treatment completion among Ugandan prison inmates. The International Journal of Tuberculosis and Lung Disease. 2014 Jul 1;18(7):781-6. 22. Lukoye D, Kalamya JN, Awor AC, Gustavson G, Kabanda J, Ferroussier-Davis O, et al. Treatment Outcomes for Tuberculosis Infection and Disease Among Persons Deprived of Liberty, Uganda, 2020. Emerging Infectious Diseases. 2024 Jul;30(7):1402-1405. 23. Hatwiinda S, Topp SM, Siyambango M, Harris JB, Maggard KR, Chileshe C, et.al. Poor continuity of care for TB diagnosis and treatment in Zambian Prisons: a situation analysis. Tropical Medicine & International Health. 2018 Feb;23(2):243-250. 24. Liu YE, Mabene Y, Camelo S, Rueda ZV, Pelissari DM, Johansen FDC, et. al. Mass incarceration as a driver of the tuberculosis epidemic in Latin America and projected effects of policy alternatives: a mathematical modelling study. Lancet Public Health. 2024 Nov;9(11):e841-e851. 25. Martinez L, Warren JL, Harries AD, Croda J, Espinal MA, Olarte RA, et.al. Global, regional, and national estimates of tuberculosis incidence and case detection

537

538

539

540

541

542

543

544

545

546

547

548

549

550

551

552

553

554

555

556

557

558

among incarcerated individuals from 2000 to 2019: a systematic analysis. The Lancet Public Health. 2023 Jul;8(7):e511-9. 26. World Prison Brief. Brazil. https://www.prisonstudies.org/country/brazil. Accessed 29 Oct 2024. 27. Van der Loo MP. The stringdist package for approximate string matching. The R Journal. 2014 Jun 1;6(1):111-122. 28. R Core Team. R: A language and environment for statistical computing. R Foundation for Statistical Computing, Vienna, Austria; 2024. Available from: https://www.R-project.org/ 29. MINISTÉRIO DA SAÚDE SECRETARIA DE VIGILÂNCIA EM SAÚDE. Vigilância epidemiológica da tuberculose: Análise de indicadores operacionais e epidemiológicos a partir da base de dados do Sinan versão 5.0. 2019. http://portalsinan.saude.gov.br/images/documentos/Agravos/Tuberculose/Cadern o de Analise 2019.pdf. Accessed 29 Oct 2024. 30. Zou G. A modified poisson regression approach to prospective studies with binary data. American journal of epidemiology. 2004 Apr 1:159(7):702-6. 31. Stekhoven DJ. missForest: Nonparametric Missing Value Imputation using Random Forest. R package version 1.5. 2022. 32. Silva MR, Pereira JC, Costa RR, Dias JA, Guimarães MD, Leite ICG. Drug addiction and alcoholism as predictors for tuberculosis treatment default in Brazil: a prospective cohort study. Epidemiology & Infection. 2017 Dec;145(16):3516-24. 33. Silva Rodrigues OA, Mogaji HO, Alves LC, Flores-Ortiz R, Cremonese C, Nery JS. Factors associated with unsuccessful tuberculosis treatment among

560

561

562

563

564

565

566

567

568

569

570

571

572

573

574

575

576

577

578

579

580

581

homeless persons in Brazil: A retrospective cohort study from 2015 to 2020. PLOS Neglected Tropical Diseases. 2023 Oct 20:17(10):e0011685. 34. Ranzani OT, Carvalho CR, Waldman EA, Rodrigues LC. The impact of being homeless on the unsuccessful outcome of treatment of pulmonary TB in São Paulo State, Brazil. BMC medicine. 2016;doi:10.1186/s12916-016-0584-8. 35. Prado Junior JC, Medronho RD. Spatial analysis of tuberculosis cure in primary care in Rio de Janeiro, Brazil. BMC Public Health. 2021 Dec;doi:10.1186/s12889-021-11834-1. 36. Sanchez M, Bartholomay P, Arakaki-Sanchez D, Enarson D, Bissell K, Barreira D, et. al. Outcomes of TB treatment by HIV status in national recording systems in Brazil, 2003–2008. PLoS One. 2012 Mar 21;7(3):e33129. 37. Siddigi K, Keding A, Marshall AM, Dogar O, Li J, Hugue R, et. al. Effect of guitting smoking on health outcomes during treatment for tuberculosis: secondary analysis of the TB & Tobacco Trial. Thorax. 2022;77(1):74-8. 38. Wang EY, Arrazola RA, Mathema B, Ahluwalia IB, Mase SR. The impact of smoking on tuberculosis treatment outcomes: a meta-analysis. The International Journal of Tuberculosis and Lung Disease. 2020 Feb 1;24(2):170-5. 39. Pelissari DM, Diaz-Quijano FA. Impact of alcohol disorder and the use of illicit drugs on tuberculosis treatment outcomes: a retrospective cohort study. Archives of Public Health. 2018 Jul;76:1-7. 40. Fuller SM, Koester KA, Maiorana A, Steward WT, Broaddus MR, Lass K, et. al. "I don't have to do this all by myself": systems navigation to ensure continuity of HIV care for persons leaving prison. AIDS and Behavior. 2019;23:14-24.

583

584

585

586

587

588

589

590

591

592

593

594

595

596

597

598

599

600

601

602

603

604

41. Avery A. Ciomica R. Gierlach M. Machekano R. Jail-based case management improves retention in HIV care 12 months post release. AIDS and Behavior. 2019 Apr 15;23(4):966-72. 42. Yanes-Lane M, Dussault C, Linthwaite B, Cox J, Klein MB, Sebastiani G, et. al. Using the barriers and facilitators to linkage to HIV care to inform hepatitis C virus (HCV) linkage to care strategies for people released from prison: findings from a systematic review. Journal of Viral Hepatitis. 2020 Feb;27(2):205-20. 43. Murray EJ, Bond VA, Marais BJ, Godfrey-Faussett P, Ayles HM, Beyers N. High levels of vulnerability and anticipated stigma reduce the impetus for tuberculosis diagnosis in Cape Town, South Africa. Health policy and planning. 2013 Jul 1;28(4):410-8. 44. Courtwright A, Turner AN. Tuberculosis and stigmatization: pathways and interventions. Public health reports. 2010 Jul-Aug;125(Suppl 4):34-42. 45. Woznica DM, Ntombela N, Hoffmann CJ, Mabuto T, Kaufman MR, Murray SM, et. al. Intersectional stigma among people transitioning from incarceration to community-based HIV care in Gauteng province, South Africa. AIDS Education and Prevention. 2021 Jun;33(3):202-15. 46. Bonato PD, Ventura CA, Reis RK, Amaral CD, De Smet S, Grossi S, et. al. Health Education for Women Released from Prison in Brazil: Barriers and Possibilities for Intervention. Social Sciences. 2024 Apr 30;13(5):249. 47. Sgarbi RV, Carbone AD, Paião DS, Lemos EF, Simionatto S, Puga MA, et. al. A cross-sectional survey of HIV testing and prevalence in twelve Brazilian correctional facilities. PLoS One. 2015 Oct 14;10(10):e0139487.

606

607

608

609

610

611

612

613

614

615

616

617

618

619

620

621

622

623

624

625

626

48. Cooper JA, Murphy S, Kirk R, O'Reilly D, Donnelly M. Data linkage studies of primary care utilisation after release from prison: a scoping review. BMC Primary Care. 2024 Aug 7:doi:10.1186/s12875-024-02527-w. 49. Akiyama MJ, Kronfli N, Cabezas J, Sheehan Y, Scheibe A, Brahni T, et.al. The role of low-income and middle-income country prisons in eliminating hepatitis C. The Lancet Public Health. 2022 Jul 1;7(7):e578-9. Figure Titles and Legends Figure 1 Title: Database Linkage between SIGO and SINAN Figure 1 Legend: Identification of individuals who were diagnosed with TB and initiated TB during incarceration, and classification by carceral movements during treatment. See Text S1, Additional File 1 for matching and inclusion criteria. SIGO, Sistema Integrado de Gestão Operacional (incarceration database); SINAN, Sistema de Informação de Agravos de Notificação (TB notifications database). Figure 2 Title: Adjusted relative risks associated with unfavorable treatment outcomes within 8 months of notification (carceral variables) Figure 2 Legend: Adjusted relative risks of unfavorable treatment outcomes evaluated eight months after date of notification for incarceration-related variables. Unfavorable outcomes refer to all outcomes other than treatment success. The duration of previous incarceration and number of prior arrests were recorded in the two years preceding the date of notification. Facility type refers to the facility at the time of TB notification. Adjusted relative risks for date of notification can be found in (Table S5, Additional File 1).

Figure 3 Title: Tuberculosis treatment outcomes stratified by time of release

Figure 3 Legend: Bar plots reporting treatment outcomes of individuals released less
than two months or two months or more from the date of notification. No case status
update includes individuals without any updates to their status in the TB database
following treatment initiation, as well as individuals who were transferred to new health
facilities/discontinued treatment, restarted treatment, and had no subsequent updates to
their case status. Other refers to individuals who experienced change of
diagnosis/regimen or were diagnosed with DR-TB.

Mato Grosso do Sul Incarceration Database (SIGO) 2006-2018 N=144,249 Mato Grosso do Sul TB Notification Database (SINAN) 2006-2018 N=12,417

Fuzzy string matching between record pairs.
Filter matches to obtain best match pairs.



Individuals newly diagnosed with TB and treated during incarceration between January 1st 2006-June 30th 2018 N=1,274

Individuals who remained in the same carceral facility within 6 months following notification N=849 Individuals who were transferred from a carceral facility within 6 months following notification N=259 Individuals who were released from a carceral facility within 6 months following notification N=166



