# Residential proximity to transport facilities as urban determinants of individual-level per- and poly-fluoroalkyl substance (PFAS) exposures: Analysis of two longitudinal cohorts in Singapore

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# Summary

**Background** The spatial determinants of human exposures to Perfluoroalkyl Substances (PFAS), a broad class of persistent environmental contaminants affecting pregnancy and early childhood development, amenable to policy are poorly understood because of the diversity of potential sources of exposure. This is especially true for modern, dense urban settings, which contain numerous built environment- and behavioral-related inputs (e.g., road traffic, run-off, wastewater, etc.) but few high-profile sources (e.g., manufacturing, agriculture, military).

**Methods** We quantify exposure based on residential proximity to transport facilities and its parcel size and evaluate the association of exposure with plasma PFAS concentrations analyzed in human blood from two geographically- and demographically- diverse cohorts of pregnant women in urban Singapore (n = 784 aged 19–47 in 2009–2011 and n = 384 aged 18-45 in 2015–2017). To rule out behavioral confounding, we exploit future residential addresses among individuals who moved (negative control exposures). Transport facilities were ground-truthed through automated extraction of Google streetview data.

**Findings** Adjusting for known predictors, within-neighborhood and unobserved spatial heterogeneity, a standard deviation (SD) increase in transport facility exposure (approximately 10,000m<sup>2</sup>) is linked to 0.12, 0.17, 0.11 SD increases in residents' perfluorobutane sulfonic acid (PFBS), perfluorobutanoic acid (PFBA), and perfluorononanoic acid (PFNA) concentrations, respectively in the 2009 cohort. We found similar positive associations in the more recent 2015 cohort.

**Interpretations** Transport facilities are prevalent near residences in urban settings and may be potential sources of PFAS emissions due to their presence in automotive-related lubricants, parts, and materials. Our findings that exposure was robustly related to individual-level concentration over and above behavioral and other factors highlight the importance of monitoring these and other urban sources of exposure.

*Key words and phrases* PFAS, built environment, biomarkers, environmental epidemiology, Singapore, GUSTO, geospatial

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# **1** Introduction

Per- and poly-fluoroalkyl substances (PFAS) include thousands of chemically stable and 2 highly persistent substances used in consumer products and industrial applications,<sup>1</sup> 3 with a long trail of studies suggesting adverse health effects.<sup>2,3</sup> For pregnant mothers, 4 higher PFAS exposure is associated with lower fertility.<sup>4</sup> Prenatal exposure to PFAS (and 5 other persistent organic pollutants) is adversely linked to maternal thyroid functioning.<sup>5</sup> 6 In children, PFAS is linked to airway and respiratory infections,<sup>6,7</sup> emotional-behavioral 7 problems,<sup>8</sup> and cognitive performance.<sup>9</sup> Newborns with in-utero exposure to perfluorooc-8 tanoic acid (PFOA) have lower birth weight.<sup>10-12</sup> 9

Despite the known adverse effects of PFAS in general, <sup>2,3,13</sup> PFAS have essential functions in industrial uses and consumer products, making a blanket ban impractical.<sup>1</sup> Consequently, outside of populations with known occupational exposures, <sup>14,15</sup> PFAS has also been detected in the general population (including mothers and children) across different contexts with less known exposure sources. <sup>4,8,9,16–23</sup>

Many studies on PFAS exposure come from environmental samples taken near wa-15 ter or wastewater, 24-34 and industrial sites. 24-27,29,35 Few studies focus on urban and 16 non-industrial areas, <sup>30</sup> even though significantly higher PFAS has been found in urban 17 areas.<sup>27</sup> Studies with human samples in highly urban contexts are far fewer. Barton et 18 al. (2020) found higher serum PFAS concentrations in adults near affected water dis-19 tricts linked to aqueous film-forming foams from a nearby Air Force base.<sup>36</sup> Across US 20 states, Sun et al. (2018) found higher plasma PFAS in participants living in inland states 21 (as opposed to coastal states).<sup>15</sup> 22

In urban environments, transport facilities (e.g., transit depots and petrol stations) <sup>23</sup> are of potential concern because of the PFAS burden from industrial-related activities <sup>24</sup> and automotive products. Glüge et al. (2020) identify 12 use categories of PFAS in automotive products and applications (car bodies, automotive waxes, windshield wiper fluid, <sup>26</sup> engine and steering systems, engine oil coolers, cylinder head coatings and hoses, electronics, fuel lines, steel hydraulic brake tubes, interior components, and brake pad additives).<sup>1</sup> These use categories are more than any other known industrial branch (such <sup>29</sup> as the chemical or semiconductor industries). <sup>1</sup> A related industrial use is the oil and gas <sup>30</sup> industry, where PFAS are used for oil and gas transportation, oil and gas storage, oil containment, and oil and fuel filtration. <sup>1</sup> These industrial usages are typically used in liquid <sup>32</sup> form. However, there is a growing appreciation for the fact that these PFAS are present <sup>33</sup> in the atmosphere, including gases, aerosols, and particulate matter, <sup>37–39</sup> including in <sup>34</sup> urban areas. <sup>40–47</sup> In particular, PFAS has been detected in air samples near high-profile <sup>35</sup> sites with PFAS contamination. <sup>43,48–51</sup>

This article, therefore, evaluates transport facilities as a potential point source of PFAS 37 exposure in pregnant women drawn from a general population. We use a population-38 and geographically-representative cohort, which measures plasma PFAS concentrations 39 in pregnant women near recruitment and birth (2009–2011). The context is Singapore, 40 a densely built city with fairly concentrated PFAS measures.<sup>20,31,52</sup> Regulatory buffers 41 mean that the known sources of exposure (e.g., heavy industrial zones) locate far from 42 residential areas (Figure A1). On the other hand, transport facilities have high PFAS in-43 dustrial usage but are non-regulated and typically located near residences. Our objective 44 is to systematically investigate the extent to which within-neighborhood spatial proximity 45 to transport facilities based on residence is linked to the PFAS concentrations from our 46 human samples. 47

## 2 Methods

# 2.1 Study population

The study population includes pregnant women from the longitudinal cohort study "Growing Up in Singapore Towards healthy Outcomes" (GUSTO). Families undergoing firsttrimester antenatal dating ultrasound scanning in two hospitals were invited to participate in the study, which aimed to understand pregnancy conditions and early childhood development. In total, 1450 women at 7–11 weeks of pregnancy were recruited into the study between June 2009 and October 2010. The participants are otherwise ethnically diverse and are located across all major areas of Singapore city (Appendix AA).<sup>20</sup> The age

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at delivery of women, near the time of blood collection, is 19–47 years old.<sup>53</sup> We map residences to administrative areas and correlate the recruited participants to the 2010 census count of the adult female population aged 20–49 across neighborhoods (r = .89, <sup>59</sup> p < .001, n = 166 neighborhoods). We also compute correlation at the broader planning area level (r = .92, p < .001, n = 33 planning areas; see Appendix AA), confirming that the cohort is geographically representative.

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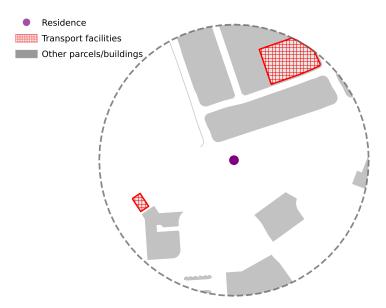
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#### 2.2 PFAS measurements

Part of the inclusion criteria for GUSTO includes the willingness to donate cord blood. 64 Our human measures of PFAS come from the analyses of these cord blood samples that 65 reflect neonatal exposure. In total, GUSTO analyzed PFAS samples for 784 participants. 66 Out of the PFAS measurements, eight had concentrations above limits of detection (LOD) 67 and limits of quantification (LOQ) for at least 95% of participants (Table A1). We subse-68 quently include these eight PFAS measurements in our analyses. These eight PFAS [LOD-69 LOQ] are: perfluorobutane sulfonic acid (PFBS) [0.078–5 ng/ml], perfluorobutanoic acid 70 (PFBA) [0.41–0.5 ng/ml], perfluorononanoic acid (PFNA) [0.016–0.1 ng/ml], perfluorooc-71 tanoic acid (PFOA) [0.009-0.1 ng/ml], perfluorooctane sulfonic acid (PFOS) [0.027-0.1 72 ng/ml], perfluorohexane sulfonic acid (PFHxS) [0.024–0.1 ng/ml], perfluoroundecanoic 73 acid (PFUnDA) [0.011–0.1 ng/ml], and perfluorodecanoic acid (PFDA) [0.01–0.1 ng/ml]. 74 Other PFAS measurements are excluded because of low detection rates. The first two, 75 PFBS and PFBA, are considered shorter-chained 4-carbon PFAS replacements for legacy 76 PFAS, such as the 8-carbon PFOA and PFOA.<sup>2</sup> For the eight included PFAS measures 77 that fall outside the limits of detection or limits of quantification (Table A1), we substitute 78 values based on the corresponding limiting values, as is common.<sup>4,54</sup> 79

# 2.3 Residential exposure to transport facilities

Figure 1 illustrates how we quantify residential exposure to transport facilities for a given residential point. We first create concentric circles around the point location of residences based on a certain radius, such as a 500m radius (approximately 1,750 feet). We



**Figure 1.** Quantifying residential exposure to transport facilities. Residential exposure to transport facilities is computed based on the area of transport facilities parcels intersecting with the concentric circle (500-meter radius) within 500m (approximately 1,640 feet) of the residence. The footprint of the transport facility outside the circle is not counted. Illustration based on a family in the cohort data.

then overlay the concentric circles with the land parcels assigned to transport facilities. <sup>84</sup> The exposure measure is the area of transport facilities within the circle (shaded red in <sup>85</sup> Figure 1). The larger this area is, the higher the measure. We also compute and extend <sup>86</sup> our analyses to alternative radiuses and years for the sensitivity analyses. <sup>87</sup>

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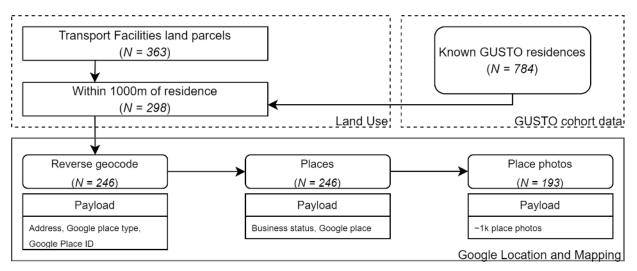
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# 2.4 Auditing land parcels for transport facilities

Figure 2 illustrates how we improve interpretation and construct validity by curating <sup>89</sup> street-level public-contributed images. Using the geo-referenced land parcels, we query <sup>90</sup> the Place API using the Place ID before finally querying the Photos API to obtain the street <sup>91</sup> images of the relevant transport facility parcels. We then manually scan all images. From <sup>92</sup> this approach, most transport facilities appear to be petrol kiosks/gas stations and larger <sup>93</sup> transport depots (e.g., bus depot in Figure 2). Appendix B provides more details. <sup>94</sup>

# 2.5 Maternal baselines

The GUSTO cohort collected demographic characteristics from interviews, which we use <sup>96</sup> to adjust for known predictors of PFAS measurement. These include age at delivery, <sup>36,55,56</sup> <sup>97</sup>



(a) Geocoding land use parcels and retrieving place photos



(b) Woodlands Bus Depot

**Figure 2.** Auditing land parcels for transport facility land use. Panel A shows a schematic representation of geo-referenced parcels used to obtain street-level photos of the location. Panel B shows an example of a large transport depot (with residential buildings in the backdrop).

place of birth/legacy effects, <sup>18,20,21</sup> ethnicity (Chinese, Malay, Indian, or Others/Mixed), <sup>18,20</sup>, <sup>36,55,57</sup> education, <sup>17,20,55</sup> occupation, <sup>14,18,36,58,59</sup> and socioeconomic baselines such as housing <sup>99</sup> type, marital status, maternal monthly income, and household monthly income (see also <sup>100</sup> Appendix AA). <sup>60,61</sup>

# 2.6 Supplementary data

To map participants to areas, we use the 2008 publicly available data on administrative <sup>103</sup> borders (in increasingly fine-grained levels—regions, planning areas, and subzones). We <sup>104</sup> also supplement our analyses using commute time data to construct alternative spatial <sup>105</sup> instruments of exposure (more details on travel time in Appendix D). The analyses with <sup>106</sup> future residences use the 2019 versions of administrative borders and land use (Appendix <sup>107</sup> E). The 2008 version of land use aligns more closely with residences at the time of recruitment, while the 2019 version aligns more closely with future residences (see Section 2.9). <sup>109</sup>

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Our replication study uses a separate younger cohort in Singapore, and these use the 110 2014 versions of administrative borders and land use (more details in Appendix F). 111

#### **Descriptives and summary statistics** 2.7

For basic statistics, we compute the total number, land area, and percentage of land 113 cover across the three plans (2008, 2014, 2019). We map land zones to postal codes 114 to compute the percentage of certain land types near given points of location. To better 115 characterize land use around residential spaces, we also restrict these computations 116 to known public and geocoded buildings (around 10k postal codes and close to 80% of 117 the resident population). Euclidean distance between residences and land parcels is 118 computed both point-to-edge and point-to-centroid. For PFAS measurements, we also 119 compute summary statistics, including quantile categories. Where relevant, we point to 120 visual patterns using spatial plots of residences and transport facilities. 121

#### Statistical analyses 2.8

The estimand of interest is how exposure to transport facilities (Section 2.3), by total sur-123 face area within 500 meters of residence occupied by transport facilities, is linked to PFAS 124 concentration. Our outcome measures are plasma PFAS measurements from cord blood, 125 which passes through to fetuses. We model the relationship between the plasma PFAS 126 concentrations and exposure to transport facilities by fitting a series of multivariable 127 regression models that adjust for variation in PFAS originating from maternal character-128 istics and unobserved and time-invariant spatial heterogeneities. The model we estimate 129 is of the form: 130

$$PFAS_{ic} = \beta(Exposure \text{ to transport facilities})_{ic} + \gamma X_i + \delta_c(administrative \text{ area})_c + \varepsilon_{ic}$$
 (1)

where PFAS measure varies by individual i and the administrative area c. The assumption 131 is that PFAS can be found beyond immediate source sites but has relatively quick decay 132 in exposure past a certain radius.  $^{21,30}$   $X_i$  is the vector of controls for observed maternal 133 characteristics described in Section 2.5 so PFAS can vary by known predictors of PFAS 134

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such as age, ethnicity, education level, and socioeconomic status. Equation (1) adjusts <sup>135</sup> for various levels of administrative areas (details below). <sup>136</sup>

Model 1 is unadjusted (setting  $\gamma = 0$ ,  $\delta_c = 0$ ). Model 2 adjusts for the maternal baselines (setting  $\gamma \neq 0$ ,  $\delta_c = 0$ ). Models 3–5 subsequently adjust for the geographical units (setting  $\gamma \neq 0$ ,  $\delta_c \neq 0$ ) in increasingly fined-grained order: regions (n = 5, see Figure A2), 139 planning areas (n = 32), and subzones (n = 147, hereon *neighborhoods*, see Figure 3). 140 For example, adjusting for the neighborhood unit allows all participants in the same unit 141 to have a constant effect, accounting for differences between neighborhoods, such as 142 drinking water sources or proximity to coasts and water catchments. <sup>15,31,36</sup>

Models 1–4 do not account for neighborhood-level heterogeneities. Model 5 adjusts 144 for neighborhood-level heterogeneities and captures within-neighborhood effects (please 145 see Appendix AA for an example of confounding without this adjustment). Standard 146 errors are clustered at the planning area level to allow for correlation in exposure within 147 towns (especially considering that the planning area delineation heavily informs urban 148 planning).

We focus on PFBS as the unregulated shorter 4-carbon PFAS industrial replacement <sup>150</sup> of the longer-chained legacy PFAS with growing data points raising health concerns (Appendix AB).<sup>2,62–65</sup> Besides having good data coverage (Table A1), PFBS is also known to have high concentrations in the region and can remobilize into the air. From analyses of environmental samples collected from the same city, and likely reflecting industrial shifts to the shorter-chain PFAS, PFBS (followed by PFBA) has the highest concentration, <sup>31</sup> and is one of the most frequently detected PFAS in different matrices with concentrations up to 100 times those reported in other cities.<sup>52,66</sup> A study in the region also reports PFBS as the most concentrated PFSA (perfluorosulfonic acids) in air samples, <sup>49</sup> and air samples from nine Asian cities (albeit excluding Singapore) indicate a relative affinity of PFBS to be aerosolized into fine particulate matter.<sup>52,67</sup> We later extend our evaluations to other PFAS.

#### 2.8.1 Secondary model

A secondary model allows for a stepped effect in exposure to transport facilities. This <sup>163</sup> model is motivated by three pieces of evidence: i) where we see two main types of transport facilities (Section 2.4), gas stations and transport depots, with large differences in <sup>165</sup> footprint; ii) where a standard deviation increase in exposure is larger than the areal footprint of a typical gas station (Table A3), and iii) from detected influential points around a <sup>167</sup> large transport depot (later in Figure 4). Therefore, we use a secondary model as an openended investigation, using indicators for various thresholds of the total area of transport <sup>169</sup> facilities:

$$PFAS_{ic} = \beta^{(t)} \mathbb{1}\{\text{Exposure to transport facilities} > t\}_{ic} + \gamma X_i + \delta_c (\text{administrative area})_c + \varepsilon_{ic},$$

for a range of t values (0 to 20,000m<sup>2</sup>, in steps of 2,000m<sup>2</sup>). Adjustments are otherwise <sup>171</sup> similar to Equation (1).

#### 2.9 Sensitivity analyses

#### 2.9.1 Model specifications

We perform an assortment of sensitivity analyses to ground our non-experimental estimates. First, we include an additional Model 6, which is Model 5 with additional adjustments for income with a smaller sample size (Appendix AB). <sup>60,61</sup> Second, we perform tests to detect influential points using a jackknife approach at the individual level and separately for planning areas and neighborhoods, where participants of an entire area are excluded in each iteration (Appendix AB). Third, in Appendix C, we systematically rule out other geographical variations in transport facilities as simply capturing other land use types (Figure C25). Fourth, we consider and rule out other spatial instruments related to our main spatial instrument (defined in Section 2.3) but should otherwise have no association with PFAS levels (Appendix D).

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(2)

#### 2.9.2 Negative control exposure

As a falsification test, we exploit the trail of participant residences and compute their <sup>186</sup> future exposure to transport facilities using their future addresses (approximately ten <sup>187</sup> years later). The concern is that some unmeasured family-level characteristics related to <sup>188</sup> PFAS exposure also systematically place families near areas with certain spatial proper-<sup>189</sup> ties (e.g., near edges of neighborhoods or near arterial roads) that happen to be where <sup>190</sup> transport facilities are located. In Appendix E, we test if future exposure can predict <sup>191</sup> past plasma concentrations among the subset of participants who have since moved to <sup>192</sup> a different location. <sup>193</sup>

#### 2.9.3 Replication study

Finally, we use a different set of 384 participants (aged 18–45 at recruitment) in the <sup>195</sup> same context from a more recent 2015 Singapore cohort—the S-PRESTO study—as a <sup>196</sup> replication study.<sup>4,68</sup> This cohort is suitable for replication as (i) it also analyzed plasma <sup>197</sup> PFAS measurements from blood samples collected at recruitment and collected basic <sup>198</sup> maternal demographics, and (ii) it allows us to construct the exposure measure the same <sup>199</sup> way using the 2014 transport facilities data. Models are as defined in Equation (1) in <sup>200</sup> Section 2.8 to similarly allow exposure to transport facilities to be associated with PFAS <sup>201</sup> us to examine whether our general findings can be replicated at a different time point. <sup>203</sup> More details in Appendix FB. <sup>204</sup>

# **3** Results

From the PFBS measurements, the median (IQR) is 18.2 (13.28–28.97) ng/ml, and the <sup>206</sup> mean is 24.16 (standard deviation (SD) 15.93) ng/ml (Table A1). These numbers are <sup>207</sup> higher than human samples in other studies, which find a mean of 0.070 ng/ml in US <sup>208</sup> adults, <sup>69</sup> and a median of 0.035 ng/ml in women with GDM in the Shanghai cohort. <sup>54</sup> <sup>209</sup> However, our measurements are consistent with other studies of human and environmen- <sup>210</sup>

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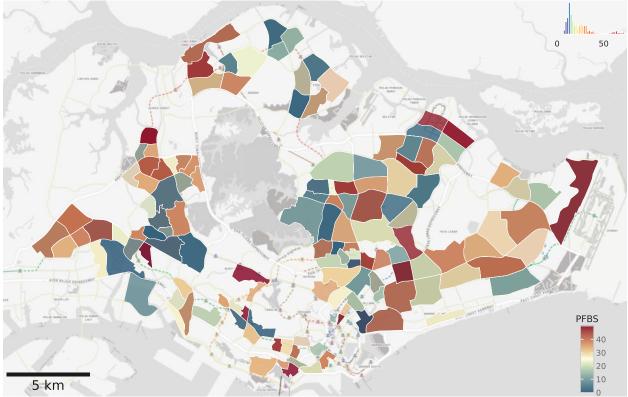


Figure 3. PFBS averaged across neighborhoods (n = 147). Legend indicates underlying distribution.

tal samples in the same region that find fairly concentrated measures of PFAS.<sup>20,31,52,54,67</sup> 211

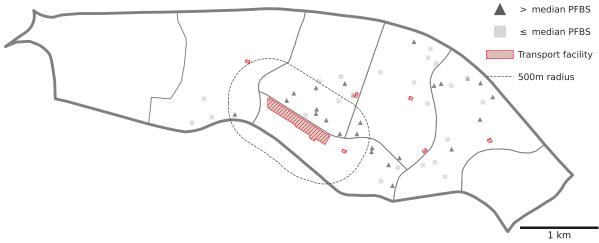
Figure 3 plots the PFBS measurements across the city. One area with high PFBS <sup>212</sup> measurements is the right-most area next to the only commercial airport. Airports are <sup>213</sup> high-profile sites. <sup>13,27,33,36</sup> Another area with high PFBS measurements is in the north- <sup>214</sup> west region, where the residential point from Figure 1 lies. In Appendix AA, Figures A2 <sup>215</sup> to A7 provide more visual aids depicting the spatial distribution of PFAS concentrations <sup>216</sup> around the location of transport facilities. <sup>217</sup>

The example in Figure 1 is a participant with unusually high plasma PFAS concentration, with PFBS near the 95th percentile (between the 75th and 100th percentile for the other seven PFAS measurements). From Figure 1, we observe that this participant lives near two transport facilities as potential points sources of PFAS, one of which is a large bus depot (see also Table A2 and Table B7). While this observation is anecdotal, we next present results that systematically quantify whether living near transport facilities is associated with higher PFAS concentrations.

### 3.1 Main results

We estimate Equation (1) and report the estimates from modeling the plasma PFBS mea-226 surements as dependent on exposure to transport facilities. Models 1–4 estimates, which 227 do not adjust for neighborhood-level heterogeneities, have weak statistical evidence of as-228 sociation, although they are always positive. From Model 5, which additionally adjusts 220 for neighborhood-level heterogeneities, we observe statistical evidence of an association 230 (0.158, SE 0.056, p = .016, Table A3). The estimate from Model 6, which adjusts for in-231 come available for a smaller sample, is comparable (0.149, SE 0.058, p = .016, Table A3). 232 Hence, we observe within-neighborhood associations of PFBS and transport facilities 233 exposure. In Appendix AA, we report statistical evidence of confounding without the ad-234 justments in Model 5 (income and housing types are associated with PFBS, except with 235 the stated adjustments), The estimate 0.153 implies that a  $10,000m^2$  increase in expo-236 sure to transport facilities (approximately 1 SD) is associated with a 1.6 ng/ml increase 237 in PFBS (0.1 SD). We observe similar findings with exposure computed using a 1000m 238 radius (Table A4), but not with the 1500m (Table A5) and 100m radius (Table A6). 239

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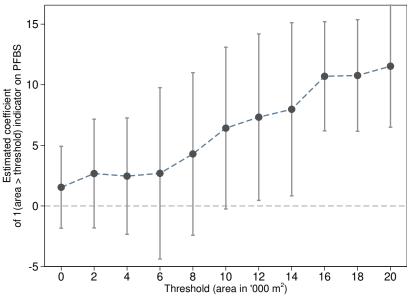


**Figure 4.** Participants around a transit depot in the area identified in the neighborhoods-level influential point analysis (Figure A15). Each triangle and square marker corresponds to a participant living in this area. Lines delineate administrative borders.

The influential point analyses (Appendix A) reveal that a single neighborhood dispro-<sup>240</sup> portionally shrinks the estimate towards zero when excluded (Figure A15). Only one <sup>241</sup> participant changes the estimate by over half the standard error when excluded (Fig-<sup>242</sup>

ure A14). We find no influential planning areas (Figure A15). Figure 4 plots the area <sup>243</sup> corresponding to the influential neighborhoods. A large transport depot with nearby <sup>244</sup> participants having higher detected PFBS. Motivated in part by this finding and that a <sup>245</sup> standard deviation increase in transport facility is larger than the typical land parcels of <sup>246</sup> gas stations (Appendix B), we use the secondary model Equation (2) which flexibly allows <sup>247</sup> for a stepped effect in the exposure to transport facilities. <sup>248</sup>

# 3.2 Thresholds



**Figure 5.** The estimated effect of exposure to an area of transport facilities based on cutoffs from estimating Equation (2). Each point is the point estimate from a model regressing PFBS on an indicator for whether the area of transport facilities within a 500m radius is more than the area indicated on the horizontal axis. For instance, the first point is the estimate of the binary for area of transport facilities > 0 m<sup>2</sup>, the second point for area > 2,000 m<sup>2</sup>, etc. After 20,000 m<sup>2</sup>, fewer than 50 households would be binned into the exposed group. Capped vertical lines are 95% confidence intervals from clustered standard errors. Please see Appendix F for the other PFAS.

Figure 5 reports the estimated coefficients at each equal-sized interval in thresholds <sup>250</sup> for the exposure measure (Section 2.8.1). We observe statistical evidence of association <sup>251</sup> once the threshold reaches 10,000m<sup>2</sup>, which is typically linked to the larger train and bus <sup>252</sup> depots based on our audits (Section 2.4). The estimated coefficients past the 10,000m<sup>2</sup> <sup>253</sup> mark are statistically significant and larger (threshold of 12,000m<sup>2</sup>, 7.326, SE 3.352, p = <sup>254</sup> .037; threshold of 20,000m<sup>2</sup>, 11.530, SE 2.459, p < .001). The estimate at the 20,000m<sup>2</sup> <sup>255</sup> threshold implies a higher PFBS of 11.5 ng/ml (approximately 0.72 of the SD). We observe <sup>256</sup>

### 3.3 Sensitivity analyses

In other sensitivity analyses, we also evaluate other land uses (Appendix C). The spatial <sup>259</sup> distribution of transport facilities does not capture other land use types (Figure C25). <sup>260</sup> To help rule out nuanced spatial confounders and exposure mechanisms, we compute <sup>261</sup> alternative spatial instruments of the exposure defined in Section 2.3 and find no associations between these and PFBS concentrations (Appendix D). We also test and rule out <sup>263</sup> that future exposure using future addresses can predict PFBS concentrations using the <sup>264</sup> subset of GUSTO participants who have relocated since study enrollment (Appendix E). <sup>265</sup>

Ν	< LOD	< LOQ	LOD value	LOQ value		Est. (95% Conf. Int.)	P-val
784	0	0	0.078 ng/mL	5.000 ng/mL		0.117( 0.030 to 0.203)	0.01 <sup>a</sup>
784	0	0	0.016 ng/mL	0.100 ng/mL		0.113( 0.037 to 0.189)	0.005
784	0	0	0.009 ng/mL	0.100 ng/mL		0.063(-0.108 to 0.235)	0.455
783	0	1	0.027 ng/mL	0.100 ng/mL		0.103(-0.027 to 0.234)	0.115
777	1	6	0.024 ng/mL	0.100 ng/mL	-	0.029(-0.021 to 0.080)	0.244
773	11	0	0.410 ng/mL	0.500 ng/mL		0.169( 0.063 to 0.274)	0.003
754	2	28	0.011 ng/mL	0.100 ng/mL	-	0.022(-0.024 to 0.069)	0.334
750	0	34	0.010 ng/mL	0.100 ng/mL		0.072(-0.042 to 0.185)	0.207
	784 784 783 777 773 754	784       0         784       0         784       0         783       0         777       1         773       11         754       2	784       0       0         784       0       0         784       0       0         783       0       1         777       1       6         773       11       0         754       2       28	784       0       0       0.078 ng/mL         784       0       0       0.016 ng/mL         784       0       0       0.009 ng/mL         783       0       1       0.027 ng/mL         777       1       6       0.024 ng/mL         773       11       0       0.410 ng/mL         754       2       28       0.011 ng/mL	784       0       0       0.078 ng/mL       5.000 ng/mL         784       0       0       0.016 ng/mL       0.100 ng/mL         784       0       0       0.009 ng/mL       0.100 ng/mL         783       0       1       0.027 ng/mL       0.100 ng/mL         777       1       6       0.024 ng/mL       0.100 ng/mL         773       11       0       0.410 ng/mL       0.500 ng/mL         754       2       28       0.011 ng/mL       0.100 ng/mL	784       0       0       0.078 ng/mL       5.000 ng/mL         784       0       0       0.016 ng/mL       0.100 ng/mL         784       0       0       0.009 ng/mL       0.100 ng/mL         783       0       1       0.027 ng/mL       0.100 ng/mL         777       1       6       0.024 ng/mL       0.100 ng/mL         773       11       0       0.410 ng/mL       0.500 ng/mL         754       2       28       0.011 ng/mL       0.100 ng/mL	784       0       0       0.078 ng/mL       5.000 ng/mL       0.117(       0.030 to       0.203)         784       0       0       0.016 ng/mL       0.100 ng/mL       0.113(       0.037 to       0.189)         784       0       0       0.009 ng/mL       0.100 ng/mL       0.063(-0.108 to       0.235)         783       0       1       0.027 ng/mL       0.100 ng/mL       0.103(-0.027 to       0.234)         777       1       6       0.024 ng/mL       0.100 ng/mL       0.029(-0.021 to       0.080)         773       11       0       0.410 ng/mL       0.500 ng/mL       0.169(       0.063 to       0.274)         754       2       28       0.011 ng/mL       0.100 ng/mL       0.022(-0.024 to       0.069)

**Figure 6.** Figure reports all estimates from modeling PFAS measurement (from cord blood) as dependent on exposure to transport facilities. Horizontal axis of the plot reports the  $\hat{\beta}$  coefficients from estimating

substance<sup>(scaled)</sup><sub>ic</sub> =  $\beta$ (Exposure to transport facilities)<sup>(scaled)</sup><sub>ic</sub> +  $\gamma X_i + \delta_c$ (neighborhood)<sub>c</sub> +  $\varepsilon_{ic}$ ,

where substance<sup>(scaled)</sup> is the PFAS substance (indicated in the first column). The PFAS substances and the exposure to transport facilities variable are scaled to have a standard deviation of one The specification is otherwise identical to that in Equation (1). Measurements below LOD and LOQ values (indicated in the second and third columns) are first imputed by the LOD/LOQ values (indicated in the fifth and sixth columns) divided by  $\sqrt{2}$ . Gray horizontal lines are the 95% confidence intervals constructed from standard errors clustered at planning areas. Significance levels: <sup>*c*</sup> 0.1 <sup>*b*</sup> 0.05 <sup>*a*</sup> 0.01.

Finally, we extend our analyses to other plasma PFAS measurements that are well-<sup>266</sup> detected. We include the eight PFAS measurements with at least 95% detection rate in <sup>267</sup> our participants (Table A1). Figure 6 reports the estimates. All units are scaled so that <sup>268</sup> Figure 6 reports the standardized regression coefficients. We observe statistical evidence <sup>269</sup> of positive associations in two emerging PFAS: PFBS (0.117, SE = 0.042, p = .010) and <sup>270</sup>

PFBA (0.169, SE = 0.052, p = .003), and the legacy PFNA (0.113, SE = 0.037, p = .005). <sup>271</sup> We also replicated our evaluations using a separate cohort study with separate analytes <sup>272</sup> and detection rates (PFBS is poorly detected and not included). This cohort uses maternal <sup>273</sup> (instead of cord) blood as the participants are not pregnant yet at the time of recruitment. <sup>274</sup> There, we find statistical evidence of positive associations with PFNA, PFOS, and PFHpS <sup>275</sup> (Figure F30, Appendix F). <sup>276</sup>

# **4** Discussion

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Using a sample of pregnant women in Singapore, a densely urban city, this study finds 278 statistical evidence of positive associations between residential proximity to transport 279 facilities and plasma PFAS concentrations. In the eight PFAS measures that were well 280 detected (at least 95% of participants analyzed), we observe statistically significant asso-281 ciations in three PFAS. We find that a one SD increase in exposure to transport facilities 282 (approximately 10,000m<sup>2</sup>) is associated with a 0.12 SD increase in PFBS, a 0.17 SD in-283 crease in PFBA, and a 0.11 SD increase in PFNA. From our systematic evaluation of 284 geo-referenced transport facility images, we find two main types of transport facilities, 285 gas stations and large transport (bus and train) depots, with vast differences in footprint 286 between the two types. An influential point assessment at the neighborhood level reveals 287 that many participants live near a specific transport depot and have above-average PFBS 288 measurements. These results suggest that the larger depots substantially contribute to 289 observed associations, which ties in with the fact that an SD increase in exposure is 290 larger than the footprint of the typical gas station. By modeling a stepped effect, partic-291 ipants with disproportionately large exposures ( $> 10,000m^2$ ) have more than a 0.46 SD 292 increase in PFBS. 293

In line with other findings from the same region, <sup>20,31,52,54,67</sup> we note relatively high <sup>294</sup> concentrations in some PFAS measurements in our sample. Human concentrations of <sup>295</sup> environmental chemicals are generally not well studied in Singapore. However, a few <sup>296</sup> other studies have verified higher PFAS from human samples in the Southeast Asia and <sup>297</sup> East Asia regions, <sup>20,54</sup> as well as high concentrations of PFAS from environmental sam- <sup>298</sup>

ples in the same city, especially PFBS in various matrices (e.g., bulk water, pore water, <sup>299</sup> suspended particles, and benthic sediments).<sup>31,52,67</sup> Specifically, from analyses of urban <sup>300</sup> catchment environmental samples targeting 22 PFAS in Singapore, the most abundant <sup>301</sup> PFAS is PFBS.<sup>31</sup> Another study in Singapore finds a high environmental concentration <sup>302</sup> of PFBS, up to 100 times higher than prior findings.<sup>52</sup> This study exhibits similarly high <sup>303</sup> concentrations, albeit in human samples,<sup>20</sup>, and we add to this by explaining its geo-<sup>304</sup> graphical variation by linking to transport facilities. <sup>305</sup>

Transport facilities are a potential point of PFAS exposure because of the known uses <sup>306</sup> of PFAS products in transport and automotive applications.<sup>1</sup> Some of the uses [functions] <sup>307</sup> of PFAS in automotive-related activities include car body [weather resistance paint, nowax brilliant top coat], engine and steering system [sealants and bearings], and engine <sup>309</sup> oil coolers [heat transfer].<sup>1</sup> Glüge et al. (2020) also identify the oil and gas industry with <sup>310</sup> seven PFAS uses [functions] such as transport [lining of the pipes, oil viscosity reduction], <sup>311</sup> storage [prevention of evaporation loss], and containment [prevent spreading of oil or gas <sup>312</sup> on water].<sup>1</sup> In our setting, residences can be as near as within 18 meters of a transport <sup>313</sup> facility. <sup>314</sup>

Several potential contamination routes exist, such as groundwater contamination <sup>315</sup> by motor vehicle-related fluids, including lubricants, cleaning agents, fire-suppressing <sup>316</sup> chemicals, etc. Additionally, it is increasingly understood that the aerosolization and air- <sup>317</sup> borne dispersal of PFAS are much more substantial than previously appreciated. <sup>40–42,44–47,67</sup><sub>318</sub> A recent extensive field study demonstrated that PFAS can remobilize into the air, am- <sup>319</sup> plifying the risk of atmospheric presence and potential exposure in areas beyond the <sup>320</sup> original point source. <sup>47</sup>

We conduct an array of sensitivity tests to rule out spurious findings and obtain coincided evidence (Section 2.9). Although we cannot definitively state transport facilities as a source of airborne exposure, our sensitivity analyses exhaustively rule out systematic associations with all other land use types (Appendix C), and adjusting for the neighborhood of residence subsume other regional patterns relating to drinking water and proximity to other well-documented sites of contamination (Equation (1)). We also rule out a specific type of behavior regarding residential choice as a confounder with null results when 328 we use future addresses to compute exposure (Appendix E). Finally, we replicated our <sup>329</sup> findings in a separate cohort (Appendix F). <sup>330</sup>

Of the three PFAS with evidence of associations, perfluorobutane sulfonic acid (PFBS) 331 and perfluorobutanoic acid (PFBA) are newer emerging PFAS replacing the legacy PFAS 332 with known adverse health effects such as perfluorooctanoic acid (PFOA) and perfluorooc-333 tanesulfonic acid (PFOS).<sup>2,3,6–8,70–73</sup> Manufacturing companies, for example, no longer 334 manufacture the 8-carbon PFOS by switching to shorter-chained 4-carbon PFBS.<sup>2,62</sup> 335 From environmental samples in the same city, PFBS and PFBA are the two most con-336 centrated short-chain PFAS.<sup>31,52</sup> Using a study on rabbits, Crute, Landon, Garner, et al. <sup>337</sup> (2023) report links between PFBS and adverse maternal outcomes such as renal injury 338 and placenta functioning.<sup>74</sup> However, studies of PFBS and PFBA on human health effects 339 are rare. Three studies using the same sample of children found links between serum 340 PFBS concentration and asthma, immunological markers, and hyperuricemia. <sup>63–65</sup> These <sub>341</sub> emerging PFAS, otherwise, have limited data about health effects, <sup>2,3,13</sup> partly because <sub>342</sub> animal-based studies are limited by the vast differences in toxicokinetic disposition of 343 animals and humans.<sup>3,62,75</sup> 344

The third PFAS with evidence of association, Perfluorononanoic acid (PFNA), has been <sup>345</sup> an understudied legacy PFAS until recent years.<sup>2</sup> Past studies with mice find a link between PFNA and developmental toxicity.<sup>76,77</sup> More recent studies with mothers find an <sup>347</sup> association between PFNA and fertility in mothers, and PFNA and low birth weight, <sup>78–82</sup> <sup>348</sup> suggesting that the understudied PFAS can adversely affect newborns in utero.<sup>2</sup> PFNA <sup>349</sup> has also been reported to contribute to metabolic syndrome in humans.<sup>83</sup>

A primary contribution of the study is how we systematically investigated the nonregulated transport facilities (and other land use types) as potential point sources of PFAS exposure. We link these facilities to plasma PFAS concentrations in a general urban population, a relatively novel approach compared to previous studies. Past studies using human samples typically evaluate known and regulated sources. <sup>15,25,35,36,84</sup> Studies evaluating non-regulated sources using human samples are rare. <sup>42</sup> Moreover, we reduce the likelihood of spurious findings by systematically considering the geographical distribution of other land uses around participants' residences. <sup>351</sup>

Our findings demonstrate that even in a well-planned, densely urban setting, there 359 can be sources of PFAS that are located near dense residences. For a sense of urban den-360 sity in Singapore, there are approximately 3.7 million residents over 126km<sup>2</sup> of purely <sub>361</sub> residential land use parcels in 2010 (approximately 26k residents per 1km<sup>2</sup>). In partic-362 ular, based on the 2010 census, the neighborhood identified as an influential point with 363 the large transport depot (Figure 4) has a population of 30k residents (approximately 364 20k/km<sup>2</sup>). Based on the later 2020 census, density in this neighborhood has increased 365 to 32k/km<sup>2</sup>. On the other hand, land use and transport facilities are static over time. <sup>366</sup> Our computations suggest that total transport facilities changed by less than 0.03% be-367 tween 2008 and 2019 for areas within a 500m radius of known public housing points. <sup>368</sup> This built environmental inertia, contrasted with increasing population density, suggests 369 that the potential risk of PFAS exposure will only intensify with time. 370

A corollary finding that may be of interest is that we observe no associations with the <sup>371</sup> general industrial areas (Appendix C), which are known point sources of PFAS.<sup>24–27,29,35</sup> <sup>372</sup> Given the regulatory buffers, one reason could be that few residents (including cohort <sup>373</sup> study participants) live near those industrial sites. Only one participant resides within <sup>374</sup> 100 meters of an industrial site. In addition, there is a heavy skew in exposure to industrial sites because of their location far away from neighborhoods (Figure A1). Combined <sup>376</sup> with our findings, this urban distribution underscores the need to examine and monitor other potential and under-sampled point sources without such regulatory residential <sup>378</sup> buffers in a highly urban setting. <sup>379</sup>

We note several limitations to our study. First, given our sample and context of analyses, we cannot precisely characterize whether pathways occur indoors at home or outdoors around the neighborhood vicinity. Previous studies have highlighted indoor air pollutants, potentially through heating, ventilation, and air conditioning systems. <sup>37,41,60</sup> We are cognizant that occupation is an important determinant of PFAS exposure, and we adjust for it. However, the cohort occupation classification may not align directly with related occupational hazards. <sup>14,18,36,58,59</sup> Not all PFAS are well detected, and we only report PFAS that were well detected. A related issue is that while PFAS is a large mixture of compounds, we study only a few separately.<sup>72</sup> Future studies should examine PFAS collectively as a mixture in such studies of spatial determinants. Finally, we lack concrete <sup>389</sup> priors about the exposure to transport facilities and thus modeled it linearly with the <sup>390</sup> primary evaluation buffer at 500 meters. The upshot is that we find coincided evidence <sup>391</sup> with different buffers and with different model specifications. <sup>392</sup>

More than half of the world's population lives in urban areas, and the number is rising <sup>393</sup> still. This study addresses a critical research gap in understanding urban determinants of <sup>394</sup> non-classic but ubiquitous contaminants, which may have wide-ranging adverse health <sup>395</sup> effects. We find robust evidence that transport facilities may be an otherwise unrecog-<sup>396</sup> nized source of urban PFAS exposures, a class of chemicals with wide-ranging potential <sup>397</sup> health effects. While Singapore has lower geographical disparities in poverty and minor-<sup>398</sup> ity status, reducing the likelihood of confounding, this also means that our findings may <sup>399</sup> have more stark implications for many other regions with greater segregation and issues <sup>400</sup> of environmental injustice. Since transport facilities are much more prevalent than, e.g., <sup>401</sup> factories or airports in urban areas, the contribution of differential exposure of PFAS and other understudied chemicals to geographic health disparities from these common built <sup>403</sup> environment features warrants far more attention. <sup>404</sup>

#### Data sharing

The GUSTO cohort study was approved by the SingHealth Centralized institutional Review Board (2018/2767/D)7 and by the National Health Care Group Domain Specific Review Board (D/2009/00021). The S-PRESTO cohort study was approved by the SingHealth Centralized institutional Review Board (2019/2143/D). 409

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The data that support the findings in this study are available upon request. Code to reproduce results 410 will be made publicly available online. 411

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Supplementary Materials (Not for Publication) for "Residential proximity to transport facilities as urban determinants of individ level per- and poly-fluoroalkyl substance (PFAS) exposures: Analysis of two longitudinal cohorts in Singapore"					
(Authors here August 2024)	769				
Table of Contents	770				
<b>§A</b> : Supplementary materials for results	771				
<b>§B</b> : Public-contributed street-level pictures of transport facilities					
§C: Other land uses	773				
<u>§D</u> : Other spatial instruments					
SE: Movers and future exposure	775				
§F: Other PFAS	776				
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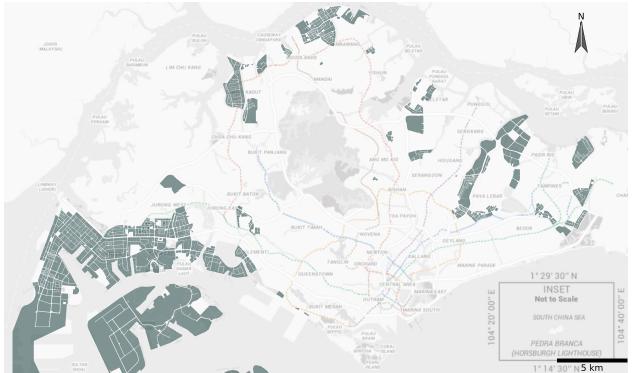
# A. Supplementary materials for results

A.A Descriptive and summary results

This appendix provides supporting tables, figures, and details for descriptive results. All 780 distributions, including spatial distribution of the land parcels of interest, are reported 781 here. 782

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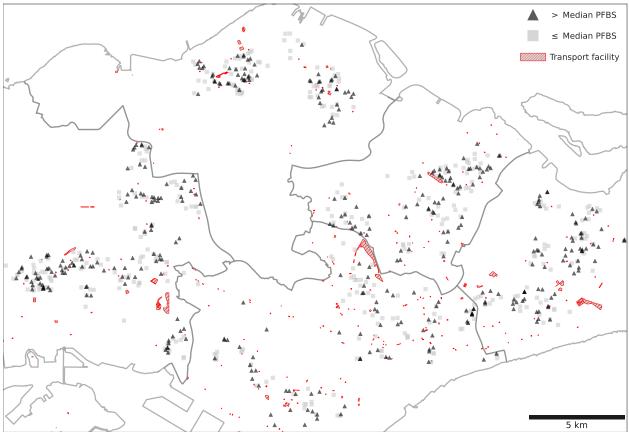
**Figure A1.** General industrial land use parcels (URA Business 2 land use type). From URA Master Plan 2008

Figure A2 plots the land parcels most closely related to heavy industrial sites (Urban Redevelopment Agency "Business 2" land use). Most are scattered on the outskirts of residential areas, in particular the large parcels. A minority of smaller land parcels classified as "Business 2" are near or within neighborhoods (referenced in https: //www.ura.gov.sg/-/media/Corporate/Planning/Master-Plan/MP19writtenstatement.pdf). 787

Table A1 reports all chemicals sent for analysis using the collected cord blood samples, 788 and the number of samples below limits of detection (LOD) or below limits of quantifica-789 tion (LOQ). Figure A2 plots the transport facility land parcels across the island. These 790 are much smaller (in comparison to Figure A1), but tend to be scattered close to resi-791 dential neighborhoods. The lines in the figure delineate the five regions (central, north, 792 north-east, east, and west). Figures A3 to A7 plot the same map by the five regions. Par-793 ticipants with above median plasma PFBS concentrations ( $\sim 18.2$  ng/ml) are shown as 794 black triangles while participants with below median concentrations are in gray squares. 795 Transport facilities are shaded red. The distribution in exposure to transport facilities 796 measure (defined in Section 2.3) is reported in Figure A8 and Figure A9. Table A2 lists 797 the eight types of transport facilities. See Appendix B for how we get more details that 798 most transport facilities are gas stations/petrol kiosks and transit depots. 799

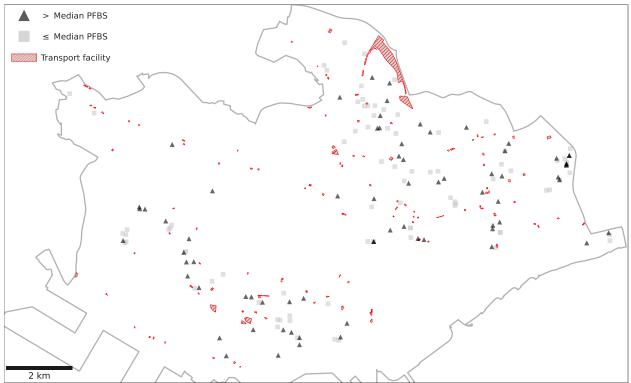
In our other auxiliary descriptive analyses (not documented in this manuscript), we 800 find correlations between the plasma PFBS concentrations and a set of maternal factors. 801 All findings below are correlations of the data without any adjustments, unless reported 802 otherwise, and are subject to data coverage. We find that low-income mothers (less than 803 SGD2000 in monthly income; 67.3%; n = 739) have higher PFBS measurements. Mothers 804 with no university degree (73.1%, n = 772) have higher PFBS measurements. We also find 805 that mothers living in public housing (90.7%; n = 772), which corresponds to more af-806 fordable housing from government housing subsidies, have higher PFBS measurements. 807 We consider public housing as those living in 1–5-room HDB (Housing and Development 808 Board). While the executive condominiums are also under the Housing and Development 809 Board, at least for the first 10 years, we classify these as private (non-public) housing. 810 Living in public housing is associated with approximately 4.01 ng/ml (SE = 1.63; p < .05) 811 higher plasma PFBS concentration. The association between public housing and higher 812 PFBS measurements persists even with the other maternal baselines and attenuates only 813 when we include the neighborhoods areal fixed effects. In this case, the estimated coeffi-814 cient is still positive but no longer statistically significant at conventional levels (adjusted 815  $\hat{\beta}$  = 2.4, SE = 1.67, p > .1). This motivates the importance of including the neighborhood 816 areal fixed effects to adjust for unmeasured spatial heterogeneity. For administrative bor-817 ders and movement between neighborhoods, see Lee, Lim, and Shen 2021,<sup>1</sup> and Lim and 818 Shen 2022.<sup>2</sup> 819

*Correlation of GUSTO participants with base population.* While GUSTO was not designed to be representative by geography, <sup>53</sup> we compute a simple correlation and find that <sup>821</sup>

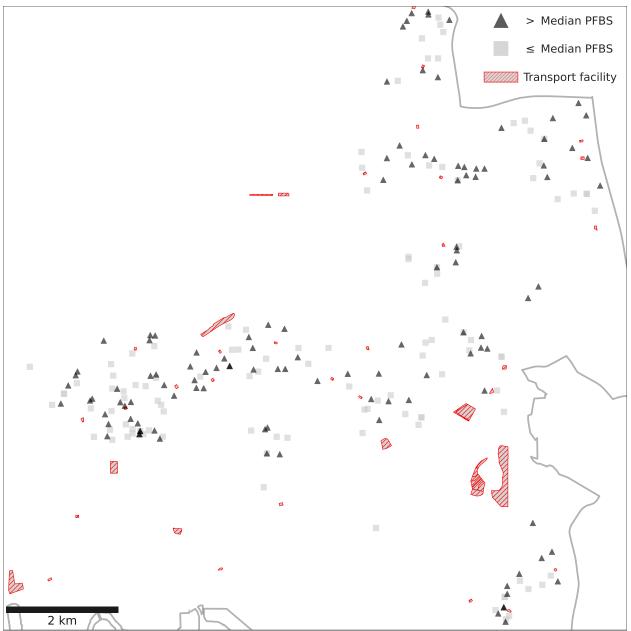


**Figure A2.** Locations of GUSTO residences and transport facilities land use parcels. Transport facilities are shaded in red. Gray lines delineate the five regions (central, west, east, north, north-east). See Figures A3 to A7 for a breakdown of the same figure by the other five regions.

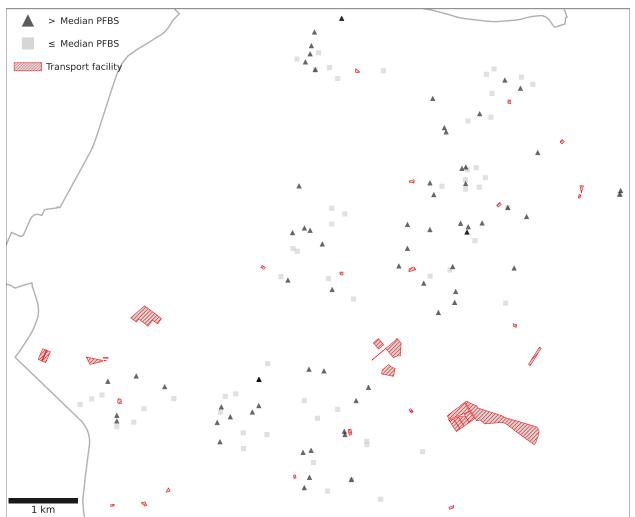
our sample of GUSTO participants vary closely with the base population by region. We 822 start by retrieving the Department of Statistics population census numbers (https://www. 823 singstat.gov.sg/find-data/search-by-theme/population/geographic-distribution) for the 824 2010 census which is available by planning areas and subzones (*neighborhoods* in the 825 main body). The participant age range at delivery is approximately 19–47. We therefore <sup>826</sup> filter the population census figures for females who fall in the six 5-year bins (age 20-827 24, age 25–29, age 30–34, age 35–39, age 40–44, age 45–49). We then ground-truthed 628 the official subzone of each GUSTO participant by first geocoding their location using 829 the postal code before map-matching their point locations to the official 2008 version 830 of the subzone delineation map. This yields a subzone (and planning area) ID tag for 831 each GUSTO participant which we use to match to the subzone tags in the population 832 census figures. Finally, we compute the correlation, by planning areas and separately by 833 subzones, between the number of GUSTO participants and the number of women aged 834 20–49 in the base population (planning area correlation = .895, p-value < .001  $\times$  10<sup>-1</sup>3, 835 planning areas = 33; subzone correlation = .89, p-value <  $.001 \times 10^{-58}$ , subzones = 166; 836 GUSTO participants = 1,488). 837



**Figure A3.** (Central Region) Locations of GUSTO residences and transport facilities land use parcels. Transport facilities are shaded in red. GUSTO residences are in triangles (above median plasma PFBS concentration) and squares (below median plasma PFBS concentrations). See Figure A2 for the map of the whole city. See Figures A4 to A7 for the other four regions.



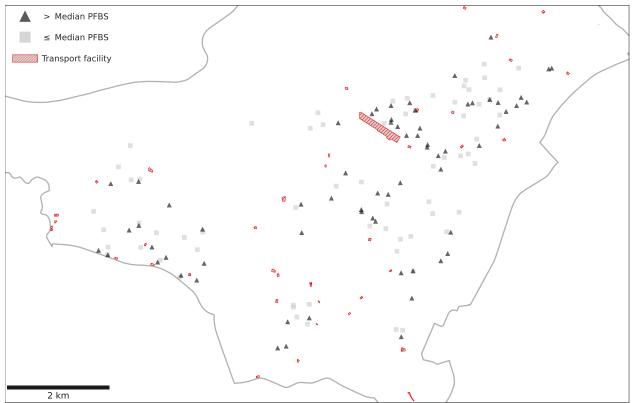
**Figure A4.** (West Region) Locations of GUSTO residences and transport facilities land use parcels. Transport facilities are shaded in red. GUSTO residences are in triangles (above median plasma PFBS concentration) and squares (below median plasma PFBS concentrations). See Figure A2 for all five regions.



**Figure A5.** (East Region) Locations of GUSTO residences and transport facilities land use parcels. Transport facilities are shaded in red. GUSTO residences are in triangles (above median plasma PFBS concentration) and squares (below median plasma PFBS concentrations). See Figure A2 for all five regions.



**Figure A6.** (North Region) Locations of GUSTO residences and transport facilities land use parcels. Transport facilities are shaded in red. GUSTO residences are in triangles (above median plasma PFBS concentration) and squares (below median plasma PFBS concentrations). See Figure A2 for all five regions.

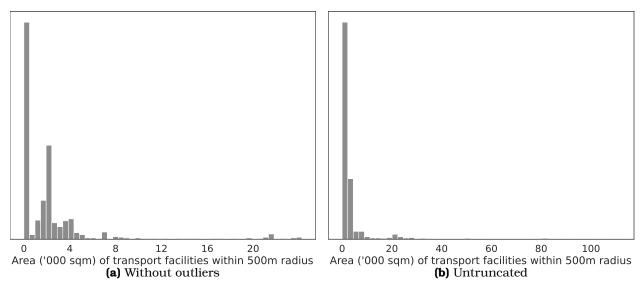


**Figure A7.** (North-East Region) Locations of GUSTO residences and transport facilities land use parcels. Transport facilities are shaded in red. GUSTO residences are in triangles (above median plasma PFBS concentration) and squares (below median plasma PFBS concentrations). See Figure A2 for all five regions.

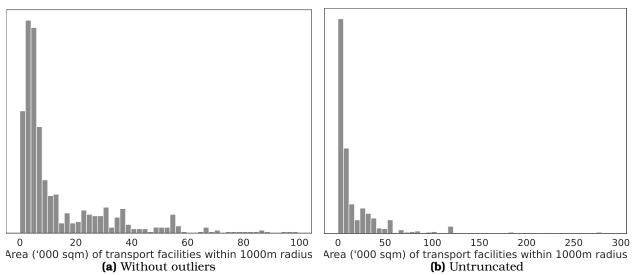
		Not	measure	ed		Summ	ary stati	istics	
	Measured (1)	< LOD (2)	< LOQ (3)	Total (4)	Mean (5)	SD (6)	Q1 (7)	Median (8)	Q3 (9)
PFBS	784	0	0	0	24.162	15.929	13.28	18.23	28.95
MBP	784	0	0	0	2.71	1.61	1.75	2.29	3.122
PFOA	784	0	0	0	1.982	1.544	1.0	1.485	2.415
PFNA	784	0	0	0	0.951	0.528	0.59	0.79	1.212
MEHP	784	0	0	0	10.712	7.614	6.77	9.065	12.24
Oxyben	784	0	0	0	2.269	0.784	1.77	2.11	2.68
PFOS	783	0	1	1	1.326	0.986	0.77	1.11	1.58
PFHxS	777	1	6	7	0.579	0.359	0.33	0.5	0.73
Benzophe	775	9	0	9	57.943	157.356	2.74	6.89	39.045
PFBA	773	11	0	11	2.076	4.34	1.25	1.58	2.06
PFUnDA	754	2	28	30	0.256	0.222	0.14	0.2	0.29
PFDA	750	0	34	34	0.162	0.1	0.11	0.14	0.19
Methylp	663	119	2	121	6.889	7.171	2.39	5.47	9.1
MECPP	655	129	0	129	0.597	0.615	0.29	0.42	0.685
MEP	558	209	17	226	1.93	3.335	0.582	0.9	1.725
PFHpA	555	139	90	229	0.196	0.169	0.11	0.15	0.21
MMP	526	258	0	258	0.911	1.469	0.61	0.75	0.93
MCINP	515	269	0	269	2.972	2.994	1.41	2.15	3.32
Propylp	506	278	0	278	0.366	0.437	0.19	0.25	0.39
BPS	437	326	21	347	0.179	0.22	0.09	0.12	0.19
Ethylp	388	359	37	396	0.26	0.245	0.13	0.2	0.29
PFPeA	207	577	0	577	0.175	0.117	0.12	0.15	0.19
PFDoDA	82	139	563	702	0.109	0.096	0.07	0.09	0.11
BPA	58	726	0	726	10.544	17.001	3.885	5.315	9.608
Butylp	11	756	17	773	0.125	0.084	0.07	0.1	0.13
MCPP	10	661	113	774	0.583	0.411	0.375	0.425	0.488
MBzP	9	773	2	775	1.477	0.756	0.81	1.35	1.85
PFHxA	7	776	1	777	0.933	0.449	0.64	0.85	1.195
MNOP	0	784	0	784	—	—	—	—	

**Table A1.** Completeness of GUSTO EDC measures

Note: Table reports the completeness of PFAS measurements. Measurements in nanograms per millilitre (ng/ML). LOD = limit of detection. LOQ = limit of quantification. Samples flagged as < LOD and < LOQ are imputed with the corresponding LOD/LOQ values divided by  $\sqrt{2}$  in Appendix F.



**Figure A8.** Distribution of area of transport facilities land parcel within 500m radius of residence. Vertical axis is density. Left figure cuts off at approximately 97th percentile. See Figure A9 for the distribution based on a 1000m radius.



**Figure A9.** Distribution of area of transport facilities land parcel within 1000m radius of residence. Vertical axis is density. Left figure cuts off at approximately 97th percentile. See Figure A8 for the distribution based on a 500m radius.

**Table A2.** Eight types of transport facilities

## Facility Type

Car Park Heavy Vehicle Park Trailer Park Bus Depot/Terminal Transport Depot MRT/LRT Marshalling Yard/Depot Driving Circuit/Test Centre Petrol Station/Kiosk

Type of establishment in transport facilities land use parcels. Official source: https://web.archive.org/web/20230207075239/https://www.ura.gov.sg/maps/media/mp/ MPUG\_2014\_for\_map\_legend.htm.

#### **A.B Main results**

838

This appendix supplements figures and tables for the main results where we evaluate the adjusted associations between plasma PFBS concentrations with exposure to transport facilities. Models are as described in Section 2.8 of the main body. 840 841

Table A3 reports the estimates from modeling the plasma PFBS measurements as <sup>842</sup> dependent on exposure to transport facilities. We have GUSTO 784 participants with <sup>843</sup> blood samples sent for PFAS analysis. 13 participants reported postal codes not found <sup>844</sup> from an official source (https://www.onemap.gov.sg/apidocs/apidocs/#search). These 13 <sup>845</sup> participants drop out in Model 1 of Table A3. In Model 2, 29 more participants drop out <sup>846</sup> when we add the maternal baselines. 21 drop out because of missing maternal baselines: <sup>847</sup>

ethnicity(1) <-> age at birth(1) -> highest education(12) <-> housing type(12) -> 848
place of birth(20) -> marital status(21).

The above implies that one participant has missing ethnicity and age at birth data (tally 850 1). 11 more have missing data on education (tally 12). These same 11 participants also 851 have missing housing type data. A further 8 have missing place of birth records (tally 852 20). One more participant also has missing occupation data (tally 21). On top of these 853 21 participants, a further 8 more dropped out as singleton observations.<sup>3</sup> In Model 4, an 854 additional participant drops out as a singleton observation when the planning area fixed 855 effects are included. In Model 5, 21 more participants drop out as singleton observations 856 when the subzone (neighborhood) fixed effects are included. A further 50 participants 857 dropped out in our last model, Model 5, when the income measures with lower coverage 858 were included. 859

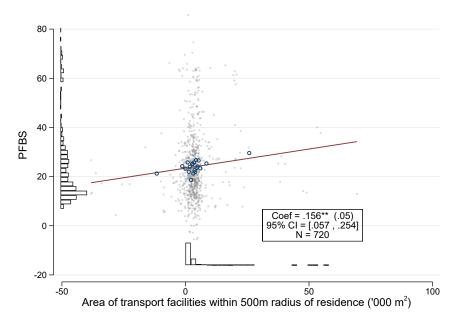
We fail to observe any statistically significant associations in Models 1–4. Model 4, 860 which adjusts for maternal baselines and for the planning areas, is marginally significant. 861 From Model 5, which adjusts for the maternal baselines and for subzone-level effects, we 862 observe a positive and statistically significant association with an estimate of 0.153 (SE 863 0.056, p < 0.05). Hence our modeling captures only within-subzone associations in PFBS 864 and transport facilities exposure but we observe no such associations across regions of 865 the city. See also Lee, Lim, and Shen (2021) and Lim and Shen (2022) for variation in 866 day-to-day movement across neighborhoods.<sup>1,2</sup> Figure A10 plots the estimated fit from 867 Model 5 with the baseline covariates partialed out from both the plasma PFBS concen-868 tration and from the exposure measure. The estimated coefficient of 0.15 implies that a 869  $10,000m^2$  increase in exposure to transport facilities is associated with an approximately 870 1.58 ng/ml increase in PFBS (approximately 0.1 SD). Model 6, which adjusts for maternal 871 and household income, has comparable estimates. Tables A4 to A6 reports the results 872 for the alternative buffer radiuses of 1000m, 1500m, and 100m. Figures A11 to A13 plot 673 the corresponding estimated best-fitted models for Model 5. 874

Figures A14 to A16 report the results from evaluating which points are influential <sup>875</sup> using the change in the estimated coefficients when certain observations are excluded <sup>876</sup> from estimation. As before, we focus on Model 5. Figure A14 reports the results where 877 each point corresponds to the change in estimated coefficient from Model 5 when the 878 observation indexed on the horizontal axis is omitted from the estimation. The rule of 879 thumb we use is to flag runs where the change is larger than  $2/\sqrt{n}$  in absolute terms. More 880 influential points increase than decrease the estimate. One influential point decreases 881 the estimate by more than half the standardized error in the estimate. In Figure A15, 882 we repeat a similar analysis but omit entire groups of individual-level observation by the 883 subzone they are residing in. We detect one influential subzone from this analysis. We 884 visually analyze this subzone in Figure 4. Figure A16 repeats the above analysis for the 885 broader planning areas. We detect no influential planning areas. 886

Figure A17 provides three illustrating examples of participants from our GUSTO sam-887 ple living in the east region. Two participants have exposure to transport facilities, and 888 one does not, as defined using a 500-meter radius using the exposure defined in Sec-889 tion 2.3. The inset in Figure A17 shows the actual gas station identified in the image. 890 The participant with no exposure has a plasma PFBS measurement of 16.93 ng/ml. 891 Based on our estimated (adjusted) coefficient of 0.153 (from Model 5 of Table A3) and the 892 exposure measure of 4.03, together, imply that with all else equal, there would be an esti-893 mated 0.62 ng/ml increase in plasma PFBS. In reality, the plasma PFBS concentrations 894 for the two participants with exposure to transport facilities are 18.42 and 23.05, which 895 is slightly higher (by 5% and by 31%, respectively). 896

Figure A18 provides another illustrating example with a separate pair of participants 897 from the west region. The inset in Figure A18 showing the large transport facility, a bus 898 depot, also shows a heavy industrial area in the backdrop, underscoring the importance 899 of evaluating within-neighborhood variation instead of across-neighborhood variation in 900 our preferred model. The participant in red has exposure to a large transport facility 901 land parcel, while the other participant in gray square has exposure to a smaller trans-902 port facility. This latter participant has a plasma PFBS concentration of 11.27 ng/ml. 903 Based on our estimated (adjusted) coefficient of 0.153 (from Model 5 of Table A3) and the 904 difference in exposure measure of 12.65 (14.653 - 2.003), together, imply that with all 905 else equal, there would be an estimated 1.935 ng/ml increase in plasma PFBS. In reality, 906 the difference in PFBS for these two participants is slightly lower at 1.83. 907

We conduct an array of sensitivity tests. Our analyses are careful about the fact <sup>908</sup> that unmeasured spatial heterogeneities are potentially linked to both unobserved be-<sup>909</sup> havior contributing to PFAS exposure and to the distribution of land use, such as trans-<sup>910</sup> port facilities. We adjust for these spatial properties by modeling participants from the <sup>911</sup> same neighborhoods to be similar and to correct the standard errors to allow for within-<sup>912</sup> neighborhood correlation of land use distribution. We find no associations without ad-<sup>913</sup> justing for neighborhoods (e.g., city-wide and across the five regions). Where possible, we <sup>914</sup> develop alternative exposure measures to rule out the main analyses capturing nuanced <sup>915</sup> unmeasured spatial patterns relating to the geographical location of transport facilities 916 that would be independent of any hypothesized PFAS exposure pathways. Our measure 917 of exposure is justified to the extent that the burden of PFAS directly corresponds to the 918 two-dimensional areal footprint of transport facilities, without accounting for the third 919 dimension of facility height. In Section 2.4 (and Appendix B), we observe that transport <sub>920</sub> facilities tend to have only a single level, and so PFAS burden should correspond most to 921 the land area of transport facilities. We also systematically examine other land use types, 922 in addition to transport facilities, to examine whether we were capturing the spatial cor-923 relation of transport facilities with some other land use types, such as train stations, 924 roads, and general industrial areas (Appendix C). We develop a placebo test using the 925 trail of future residence in the GUSTO cohort and show that exposure using future ad-926 dresses is unable to predict plasma PFAS measurements from blood samples taken years 927 earlier (Appendix E). We also extend our analyses to a separate cohort study with dif-928 ferent plasma PFAS analytes from blood samples collected at a separate time point with 929 broadly similar findings (Appendix F). 930

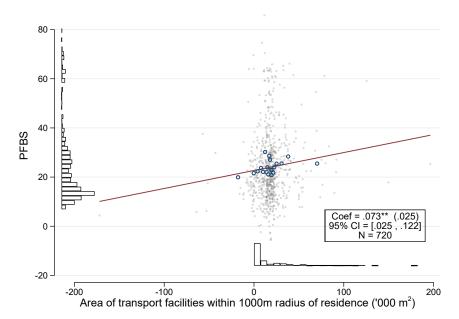


**Figure A10.** PFBS and transport facilities. See also Figure A8 for the distribution of area of transport facilities. Corresponds to column (5) of Table A3. Standard errors clustered at planning areas. Binned data in blue hollow circles. Significance levels: + 0.1 \* 0.05 \*\* 0.01 \*\*\* 0.001.

	Ľ	ependen	t variable	e is PFBS	(ng/mL)	
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 500m buffer	0.052	0.076	0.085	$0.129^{c}$	$0.153^{b}$	$0.149^{b}$
	(0.039)	(0.046)	(0.052)	(0.063)	(0.056)	(0.058)
Constant	$23.935^{a}$	9.980	9.946	14.848	23.386	17.255
	(0.579)	(17.920)	(17.155)	(17.000)	(21.277)	(27.469)
R <sup>2</sup>	0.001	0.056	0.066	0.096	0.193	0.214
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.0	23.7
Std. dev. of X	10.6	10.2	10.2	10.2	10.3	10.6
n(Transport facilities (area) within 500m buffer > 0)	458	439	439	439	426	399
n(Clusters)	_	30	30	29	29	28
Ν	771	742	742	741	720	670

Table A3.	PFBS and	transport	facilities	within	500m	radius	of residence.
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Note: Transport facilities area within 500m buffer is in 1,000 square meters. See Section 2.5 for the maternal and income controls. Figure A10 shows Model 5. Standard errors in (2)–(6) clustered at the planning areas. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

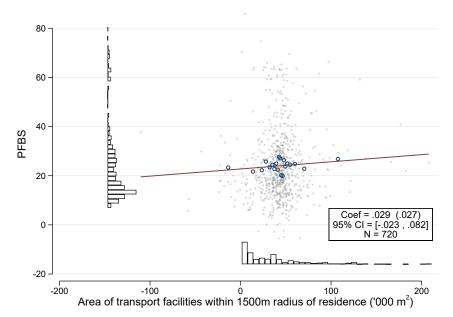


**Figure A11.** PFBS and Transport Facilities area within 1000m radius. Reported estimates are from regressing PFBS on the area (1,000) of transport facilities within 1000m radius of residence, with subzone fixed effects and maternal characteristics partialed out. Model corresponds to column (5) of Table A4. Standard errors clustered at subzones. Raw data in small gray circles. Binned data in blue hollow circles. Fitted line from the estimated model. Marginal histograms indicate the underlying distributions. Significance levels: + 0.1 \* 0.05 \*\* 0.01 \*\*\* 0.001.

	Ι	Dependen	t variable	e is PFBS	(ng/mL)	
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 1000m buffer	0.013	0.019	0.023	$0.046^{b}$	$0.075^{a}$	$0.071^{a}$
	(0.014)	(0.016)	(0.016)	(0.019)	(0.026)	(0.024)
Constant	$23.888^{a}$	10.429	10.372	16.229	25.683	19.761
	(0.624)	(17.754)	(17.061)	(17.107)	(21.334)	(27.395)
$\overline{R^2}$	0.001	0.055	0.065	0.096	0.196	0.216
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.0	23.7
Std. dev. of X	32.1	31.8	31.8	31.8	32.1	32.7
n(Transport facilities (area) within 1000m buffer > 0)	723	694	694	694	678	631
n(Clusters)	_	30	30	29	29	28
Ν	771	742	742	741	720	670

Table A4. PFBS and transport facilities within 1000m radius of residence

Note: Transport facilities area within 1000m buffer is the area (in 1,000 square meters) of the buffer area residence that is allocated to transport facilities land use. Figure A11 shows Model 5. Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

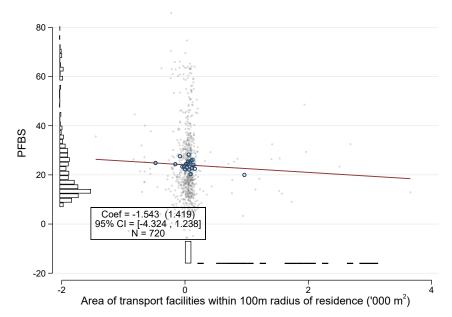


**Figure A12.** PFBS and Transport Facilities area within 1500m radius. Reported estimates are from regressing PFBS on the area (1,000) of transport facilities within 1500m radius of residence, with subzone fixed effects and maternal characteristics partialed out. Model corresponds to column (5) of Table A5. Standard errors clustered at subzones. Raw data in small gray circles. Binned data in blue hollow circles. Fitted line from the estimated model. Marginal histograms indicate the underlying distributions. Significance levels: + 0.1 \* 0.05 \*\* 0.01 \*\*\* 0.001.

	Dependent variable is PFBS (ng/mL)					
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 1500m buffer	0.002	0.003	0.004	0.018	0.033	0.029
	(0.011)	(0.013)	(0.012)	(0.016)	(0.026)	(0.026)
Constant	$24.063^{a}$	10.578	10.420	15.855	24.955	18.952
	(0.762)	(17.422)	(16.705)	(16.921)	(21.523)	(27.304)
$\overline{\mathbf{R}^2}$	0.000	0.054	0.063	0.092	0.190	0.211
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.0	23.7
Std. dev. of X	55.1	54.7	54.7	54.7	55.1	55.1
n(Transport facilities (area) within 1500m buffer > 0)	770	741	741	740	720	670
n(Clusters)	_	30	30	29	29	28
Ν	771	742	742	741	720	670

#### Table A5. PFBS and transport facilities within 1500m radius of residence

Note: Transport facilities area within 1500m buffer is the area (in 1,000 square meters) of the buffer area residence that is allocated to transport facilities land use. Figure A12 shows Model 5. Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

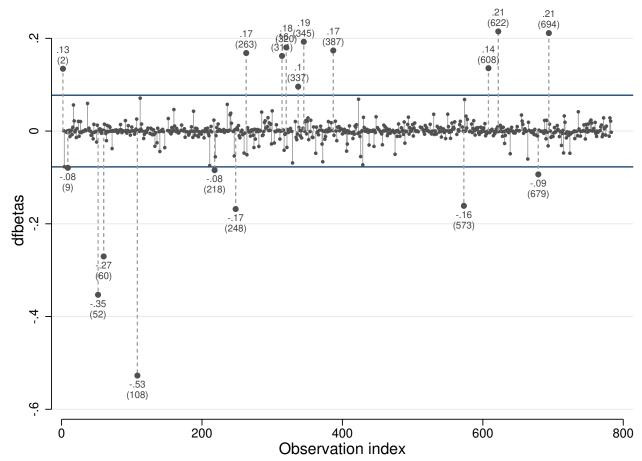


**Figure A13.** PFBS and Transport Facilities area within 100m radius. Reported estimates are from regressing PFBS on the area (1,000) of transport facilities within 1500m radius of residence, with subzone fixed effects and maternal characteristics partialed out. Model corresponds to column (5) of Table A6. Standard errors clustered at subzones. Raw data in small gray circles. Binned data in blue hollow circles. Fitted line from the estimated model. Marginal histograms indicate the underlying distributions. Significance levels: + 0.1 \* 0.05 \*\* 0.01 \*\*\* 0.001.

	Dependent variable is PFBS (ng/mL)					
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 100m buffer	-0.959	-1.766	-1.885	-1.424	-1.569	$-2.899^{c}$
	(1.326)	(1.390)	(1.361)	(1.349)	(1.572)	(1.569)
Constant	$24.186^{a}$	10.252	10.048	15.969	23.216	16.254
	(0.498)	(17.940)	(17.026)	(17.145)	(21.461)	(27.150)
$\overline{R^2}$	0.000	0.055	0.065	0.091	0.189	0.212
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.0	23.7
Std. dev. of X	0.4	0.4	0.4	0.4	0.4	0.4
n(Transport facilities (area) within 100m buffer > 0)	34	34	34	34	33	30
n(Clusters)	_	30	30	29	29	28
Ν	771	742	742	741	720	670

#### Table A6. PFBS and transport facilities within 100m radius of residence

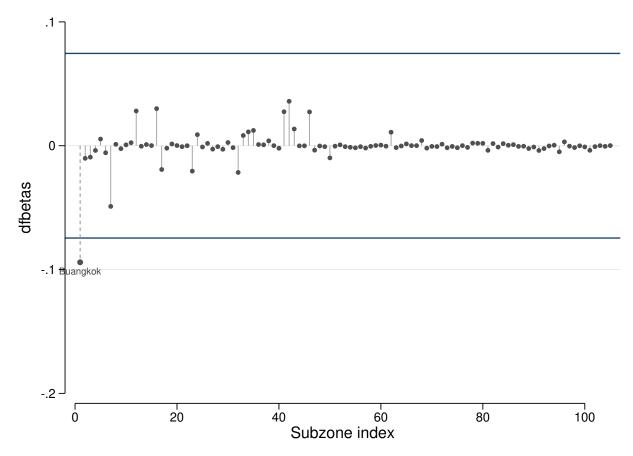
Note: Transport facilities area within 100m buffer is the area (in 1,000 square meters) of the buffer area residence that is allocated to transport facilities land use. Figure A13 shows Model 5. Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.



**Figure A14.** Figure shows the influence of each observation on the estimate of area of transport facilities within 500m radius of residence. Results corresponds to Figure A10 and Table A3. Vertical axis is the dfbeta measure standardized by the standard error of the estimate:

$$dfbetas_{(i)j} = \frac{\hat{\beta}_j - \hat{\beta}_{(i)j}}{SE(\hat{\beta}_j)}.$$
(A1)

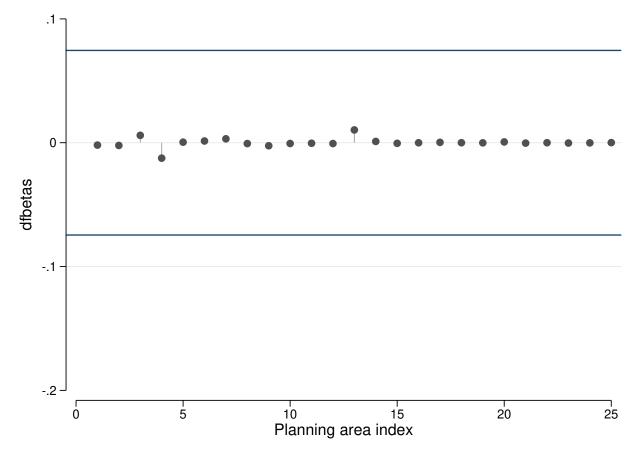
where  $\hat{\beta}_j$  is the estimate from omitting observation *i*. The paired horizontal lines are rule-of-thumb thresholds computed as  $\pm 2/\sqrt{n}$ . Observations falling outside the rule-of-thumb threshold highlighted with the dfbeta and observation index (in parentheses).



**Figure A15.** Figure shows the influence of each subzone on the estimations of model (5) in Table A3. Vertical axis is the dfbetas measure standardized by the standard error of the estimate:

dfbetas<sub>(c)j</sub> = 
$$\frac{\hat{\beta}_j - \hat{\beta}_{(c)j}}{SE(\hat{\beta}_j)}$$
.

where  $\hat{\beta}_c$  is the estimate from omitting all observations from subzone c. The paired horizontal lines are ruleof-thumb thresholds computed as  $\pm^2/\sqrt{n}$ . Subzones falling outside the rule-of-thumb threshold highlighted with the dfbeta and observation index (in parentheses).



**Figure A16.** Figure shows the influence of each planning area on the estimations of model (5) in Table A3. Vertical axis is the dfbetas measure standardized by the standard error of the estimate:

dfbetas<sub>(c)j</sub> = 
$$\frac{\hat{\beta}_j - \hat{\beta}_{(c)j}}{SE(\hat{\beta}_j)}$$
.

where  $\hat{\beta}_c$  is the estimate from omitting all observations from planning area c. The paired horizontal lines are rule-of-thumb thresholds computed as  $\pm^2/\sqrt{n}$ . Planning areas falling outside the rule-of-thumb threshold highlighted with the dfbeta and observation index (in parentheses).



**Figure A17.** Example of three participants with and without exposure. *Image source:* Google Photos.



**Figure A18.** Example of two participants with high and low exposure. *Image source:* https://en.wikipedia.org/wiki/Soon\_Lee\_Bus\_Park#/media/File:Soon\_Lee\_Bus\_Park\_and\_Jurong\_Industrial\_Estate\_aerial\_view.jpg.

#### **B.** Public-contributed street-level pictures of transport facilities

This appendix provides further details about how we attempted to pin down the specific <sup>932</sup> type of transport facility using the publicly available land use data. We know where land <sup>933</sup> parcels for transport facilities are located, but not the specific facility type. <sup>934</sup>

931

To map residences to land use zones of potential hazard in our main analyses, we 935 use the publicly-available vector data on land zones with assigned typology, such as 936 Transport Facility. Within each land use type, there can be further breakdowns of types. 937 For instance, a transport facility can be one of eight subtypes: 1) car park, 2) heavy vehicle 938 park, 3) trailer park, 4) bus depot/terminal, 5) transport depot, 6) MRT/LRT marshalling 939 yard/depot, 7) driving circuit/test centre, or 8) petrol station/kiosk (Table A2). This 940 specificity on the subtypes, however, is not present in the public data. For instance, for 941 a given parcel of land zone assigned for transport facilities, we do not know if this would 942 be a gas station or a transit depot. 943

To address this and to partially scale the solution, we systematically obtain street-944 level public-contributed images geotagged to the location of the transport facilities. There 945 are, in total, 363 land parcels assigned to transport facilities in the 2008 data. We first 946 reverse-geocode all the 298 transport facilities whose centroids are within 1000m of a 947 GUSTO residence, using the centroid coordinate.<sup>1</sup> From this, we obtain the associated 948 formatted address, Google Place ID, and Google Place type. We successfully recovered 949 these for 246 transport facility land parcels. The place types for these 246 are tabulated 950 in Table A2. Each place can have more than one type. The "Establishment" and "Point 951 of Interest" type gets tagged to most (n = 208). Store, food, and convenience stall also 952 get tagged to many transport facility (n  $\geq$  58). This does not rule out our queried land  $_{953}$ parcels as transport facilities. Virtually all gas stations/petrol kiosks in Singapore have 954 convenience shops that also sell food (Figure B19 and Figure B20). Many of the tagged 955 pictures we later see (Figure B20) are also of these known convenience stalls (e.g., 7-956 Eleven, FairPrice Xpress, etc.) commonly found in gas stations, more on the pictures 957 below. Other observations are that 58 are typed as gas stations, 20 as car wash (which 958 can also be found in gas stations, Figure B21), 17 as transit stations, 2 as subway station, 959 and 1 as a bus station. 960

To further gain insights into the transport facility land use, we use the 246 place <sup>961</sup> ID to obtain a list of photo IDs tagged to the place,<sup>2</sup> before finally using the photo IDs <sup>962</sup> to download the photos.<sup>3</sup> We successfully obtained at least one street-level photo for <sup>963</sup> 193 of the transport facility land parcels. In total, we have 1135 photos, and we looked <sup>964</sup>

<sup>&</sup>lt;sup>1</sup>Using the Google Geocoding Service: https://developers.google.com/maps/documentation/javascript/ geocoding.

<sup>&</sup>lt;sup>2</sup>Using the Google Places Service: https://developers.google.com/maps/documentation/places/ web-service/details.

<sup>&</sup>lt;sup>3</sup>Using the Google Photos Service: https://developers.google.com/maps/documentation/places/ web-service/photos.

through all of them. Figure B19 shows an assort of photos related to gas stations while <sup>965</sup> Figure B22 shows an assortment of photos broadly related to transit depots. The Places <sup>966</sup> service also returns the name of the point of interest, if available, but we do not find this <sup>967</sup> to be a particularly clear signal of transport facility type. For instance, one place name we <sup>968</sup> recovered is "LKA Auto Parts Trading Pte Ltd," but upon inspection of the tagged photos, <sup>969</sup> it became apparent that it is a gas station (image D in Figure B19). <sup>970</sup>

Overall, many photos of transport facilities appear to be gas stations/petrol kiosks 971 or transit depots. The assumption here is that the other types of transport facilities (Ta-972 ble A2) are rare and do not have some attribute that makes them less likely to be captured 973 in the places and photos database or less likely to be captured in pictures by the public. 974 Linking the photos to the underlying spatial data, we observe that gas stations occupy 975 relatively small land areas. In Figure B19, the gas station in subfigure (a) has a known 976 land area (and closest centroid-to-point distance to a GUSTO residence point) of 1,190m<sup>2</sup> 977 (414m), the gas station in subfigure (b) is  $800m^2$  (180m), the gas station in subfigure (b) <sub>978</sub> is  $1,862m^2$  (898m), and the gas station in subfigure (d) is  $2,600m^2$  (214m). The land <sub>979</sub> parcels that appear to be transit depots are far larger. The bus depot in subfigure (a) of 980 Figure B22 is 52,608m<sup>2</sup>, and the closest centroid-to-point (not edge-to-point) distance 981 to a GUSTO residence point is 963m. The bus depot in Figure 1 is  $25,981m^2$  with the 982 nearest GUSTO residence within 592m. For reference, one of the larger gas stations we 983 find is 4,055m<sup>2</sup> and is within 976m of a GUSTO residence (Figure B23). All street-level 984 photos (Figures B19 to B23) in this manuscript are public contributed but taken directed 985 via the Google Photos API (unless otherwise stated). 986

	Туре	Count
1	Establishment	208
<b>2</b>	Point Of Interest	208
3	Store	80
4	Food	67
5	Convenience Store	58
6	Gas Station	58
7	Car Wash	20
8	Atm	19
9	Finance	19
10	Transit Station	17
11	Car Repair	14
12	Restaurant	13
13	Grocery Or Supermarket	6
14	Supermarket	6
15	Place Of Worship	4
16	Park	3
17	Meal Delivery	3
18	Parking	3
19	Subway Station	2
20	Cafe	2
21	Bakery	2
22	Clothing Store	1
23	Tourist Attraction	1
24	Street Address	1
25	Health	1
26	General Contractor	1
27	Bus Station	1
28	Car Rental	1
29	Home Goods Store	1
30	Car Dealer	1

Table B7. Types of detected places in transport facilities parcels

Note: Types come from the Google Geocoding API. See Figure 2. For n = 246 Google places. Types are not mutually exclusive, places can have multiple types. Types are not necessarily accurate nor complete.



(a) Cencon Bldg



(b) 76 Yio Chu Kang Rd

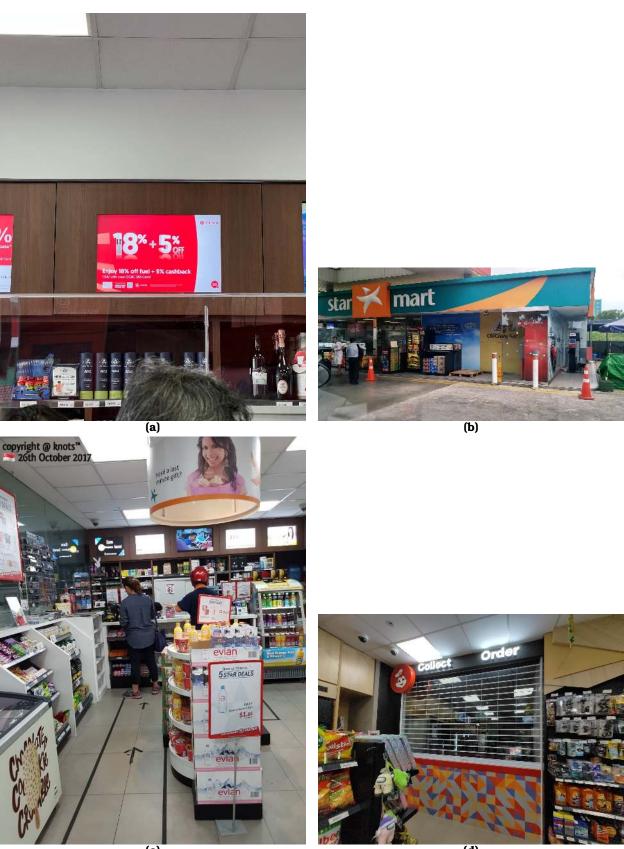


(c) 324 Thomson Rd



(d) 603 Tiong Bahru Rd

**Figure B19.** Petrol Stations. Selected illustrations of petrol stations and kiosks in transport facilities land parcels within 1000m of known (GUSTO) residences. Photos are from the Google Place Photo service which source from photos that are user-submitted and photos from managers of establishments. *Image source:* Google Photos.



(c) Figure B20. Food and convenience shops in gas stations. Image source: Google Photos.

(d)



(a) **Figure B21.** Car wash and car repair services in gas stations. *Image source:* Google Photos.



(a) Bukit Batok Bus Depot



(b) Yio Chu Kang Bus Interchange



(c) Bukit Merah Interchange



(d) (Before) Jurong East Station



(e) St. Michael's Terminal

(f) Bangkit Station

**Figure B22.** Transport depots, terminals, and stations. Selected illustrations of transport depots in transport facilities land parcels within 1000m of known (GUSTO) residences. Photos are from the Google Place Photo service which source from photos that are user-submitted and photos from managers of establishments.

Image source: Google Photos.



**Figure B23.** Larger gas station at 4,055m<sup>2</sup>. *Image source:* Google Photos.

## C. Other land uses

This appendix considers and reports results for all other land use types. The spatial <sup>988</sup> vector data on land parcels and their land uses come from the URA official land use data <sup>989</sup> in Singapore. The master plans are released on a roughly five-year cadence. For the <sup>990</sup> main evaluations, we use the 2008 version, which is closest to when blood samples were <sup>991</sup> collected. In the 2008 version, there are more than 110k delineated land parcels of 32 <sup>992</sup> land types over 776km<sup>2</sup> of land in the 2008 plan. <sup>993</sup>

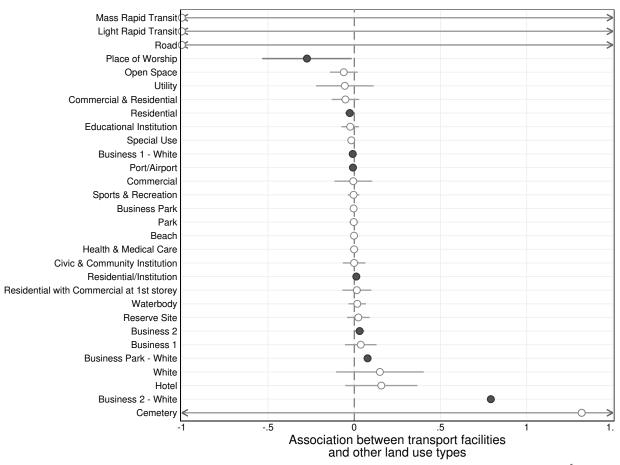
Figure C24 reports the within-subzone correlation of transport facilities with all other 994 land use types. The measures of the land uses are as defined in Section 2.3 of the main 995 article and share the same units. We observe that four land use types—Place of Wor-996 ship, Residential, Business 1 - White, and Port/Airport—are negatively correlated with 997 the transport facility land use. Another four land use types—Residential/Institution, 998 Business 2, Business Park - White, and Business 2 - White—are positively correlated 999 with the transport facility land use. Business 1 and Business 2 broadly refers to light 1000 and heavy industry uses, respectively (see https://web.archive.org/web/20240717071728/ 1001 https://www.ura.gov.sg/-/media/Corporate/Planning/Master-Plan/MP19writtenstatement. 1002 pdf?la=en#page=11.) The estimated coefficients are adjusted for subzone areal effects and 1003 standard errors are clustered at the planning areas. 1004

Figure C25 reports the estimated association between the plasma PFBS concentration <sup>1005</sup> with all the other land uses, in addition to transport facilities. We observe statistically <sup>1006</sup> significant and fairly large estimated coefficients for Business 2 - White and Light Rapid <sup>1007</sup> Transit (LRT). Although it may be interpreted as a signal of potentially hazardous sites, <sup>1008</sup> we also note that in 2008, there were only three Business 2 - White land parcels (14 <sup>1009</sup> today) with a collective land area of 0.09km<sup>2</sup> (0.17km<sup>2</sup> today). For the LRT, in 2008 there <sup>1010</sup> were only 14 parcels (15 today) with a collective land area of 0.01km<sup>2</sup> (0.02km<sup>2</sup> today). <sup>1011</sup>

*General industrial areas.* Factories and industrial plants are the known land parcels <sup>1012</sup> of concern. However, residences near industrial sites in our urban setting are rare. General industrial ("Business 2") areas have a regulatory minimum nuisance buffer of 100m <sup>1014</sup> (approximately 330 feet). In 2008, there were 5,576 land parcels used for heavy industry <sup>1015</sup> with a collective land area of 118km<sup>2</sup> (approximately 15.2% of total land area), but most <sup>1016</sup> are geographically clustered outside residential neighborhoods (Figure A1). In practice, <sup>1017</sup> the industrial areas also usually more than clear the regulatory minimum. The nearest <sup>1018</sup> distance from our participants to a general industrial land use parcel is 45 meters by <sup>1019</sup> edge of the land parcel 70 meters by of land parcel, and only 163 participants have an <sup>1020</sup> industrial land use parcel intersecting within 500m of their residence. For that single <sup>1021</sup> industrial site is small on average—with a land parcel of approximately 2,000m<sup>2</sup>—which <sup>1023</sup> is small relatively to the collectively areas of industrial sites on the outskirts of neigh-<sup>1024</sup> borhoods (see Figure A1). Using distance to centroid, only one residence is within 100 <sup>1025</sup> meters. For distance to edge, 18 residences are within 100 meters.

Transport facilities do not share these regulatory buffers and can locate right next 1027 to residential buildings (Figure A17 and Figures A2 to A7). For distance to centroid, 1028 33 residences are within 100 meters of a transport facility. For distance to edge, 72 1029 residences are within 100 meters of a transport facility. By distance to edge (distance to 1030 centroid), residences in our sample with blood collections live as near as within 18 meters 1031 (40 meters) of a transport facility. There are 363 transport facility land parcels with a 1032 collective land area of 2.3km<sup>2</sup> (approximately 0.3% of total land area). Base on distance 1033 to edge (distance to centroid), the nearest transport facility is within 18 meters (40 meters) 1034 of a residence. While the collective land area is fairly small, these transport facilities are 1035 scattered geographically. For additional context, we use about 120k of known postal 1036 codes and compute the percentage of area attributed transport facilities and industrial 1037 land parcels within a 500m concentric circle as 0.38% (SD 1.1%) and 4.83% (SD 15.79%), 1038 respectively. When we subset postal codes to known public residences, the numbers 1039 change to 0.41% (SD 1.27%) and 1.63% (SD 1.27), suggesting that the industrial areas 1040 tend to be placed further away from residential areas. 1041

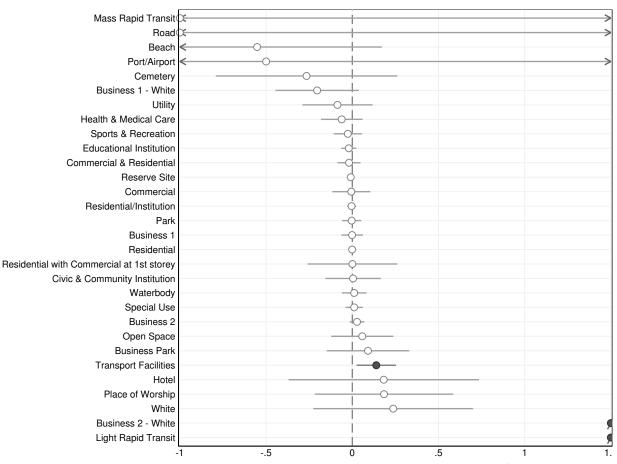
1026



**Figure C24.** Correlation of transport facilities to other land use types. Horizontal axis reports the  $\hat{\beta}$  coefficients from estimating

transport facilities<sub>*ic*</sub> = 
$$\beta$$
(land use)<sup>*j*</sup><sub>*ic*</sub> +  $\delta_c$ (subzone)<sub>*c*</sub> +  $\varepsilon_{ic}$ , (C2)

where (land use)<sup>*j*</sup> is one of the land use types indicated on the vertical axis. All variables are area within 500m radius of residence. Estimates significant at the 5% level have black markers; the remaining estimates have hollow markers. Gray horizontal lines are the 95% confidence intervals constructed from standard errors clustered at planning areas. Arrows indicate truncated confidence intervals.



**Figure C25.** Association between PFBS and land use types. Horizontal axis reports the  $\hat{\beta}$  coefficients from estimating

$$PFBS_{ic} = \beta (land use)_{ic}^{j} + \gamma X_{i} + \delta_{c} (subzone)_{c} + \varepsilon_{ic},$$
(C3)

where  $(\text{land use})^{j}$  is area (units in 1'000m<sup>2</sup>) of the land use type indicated on the vertical axis. Model is equivalent to model (5) of Table A3. Estimates significant at the 5% level have black markers. Gray horizontal lines are the 95% confidence intervals constructed from standard errors clustered at planning areas. Arrows indicate truncated confidence intervals.

#### **D.** Other spatial instruments

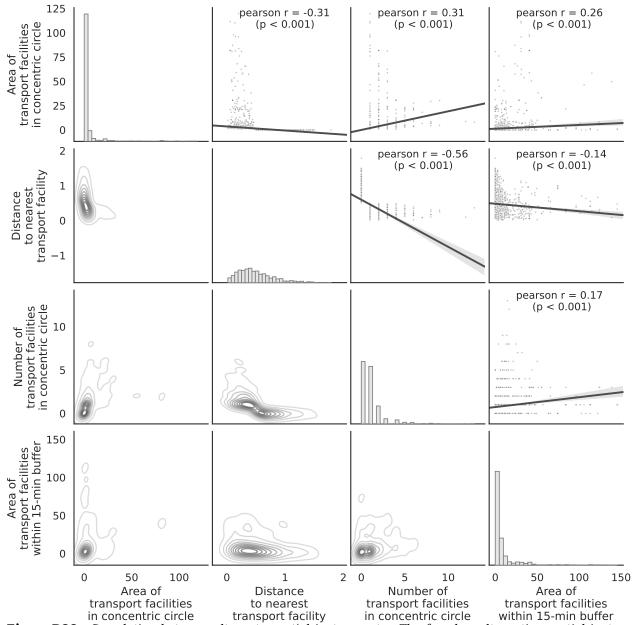
To help examine nuanced spatial patterns in the areal proximity to transport facilities, <sup>1043</sup> and as an auxiliary exploration of the exposure mechanism, this appendix evaluates a <sup>1044</sup> handful of alternative spatial instruments of the exposure measure defined in Section 2.3. <sup>1045</sup>

The four alternative instruments are: i) distance to nearest transport facility by a 1046 point-to-edge distance from the point of residence to the edge of the land parcel, ii) distance to nearest transport facility by a point-to-centroid distance from the point of residence to the geometric centroid of the land parcel, iii) number of transport facilities 1049 within the concentric circle, and iv) accessibility-based area around residence based on 1050 areas within 10–20-minute reach by public transit (including walking). As a caveat, the 1051 public travel time data was collected around 2021. These alternative spatial instruments 1052 that are potentially linked to the main exposure measure defined in Section 2.3 through 1053 spatial patterns relating to where residences and transport facilities according to urban 1054 planning, but potentially have no direct pathway to PFAS exposure. More concretely, if 1055 exposure is also tied to the size of the facility, residences near transport facilities are 1056 likely similar to residences with high exposure as defined in Section 2.3 but otherwise 1057 should not share the same level of exposure. Figure D26 reports the correlation between 1058 the four key alternative instruments (p-values < .001).

Table D8 report the estimates for the different models for the point-to-edge distance <sup>1060</sup> to the nearest transport facility. Models and adjustments are otherwise similar to Equation (1) and Table A3. We observe no association between proximity measured by distance <sup>1062</sup> to nearest transport facility and the plasma PFBS concentration. From Model 5, the estimated coefficient is -0.676 (SE 3.38, p = .843, Table D8), which implies that a standard <sup>1064</sup> deviation decrease in distance to the nearest transport facility (~300m) is related to a <sup>1065</sup> -0.2 decrease in PFBS (SE 0.01). Table D9 reports the estimates for nearest distance to <sup>1066</sup> transport facilities measured as point of residence to the geometric centroid of the land <sup>1067</sup> parcel. We do not observe any statistically significant association here.

Table D10 reports the estimates using the number instead of total area of transport  $_{1069}$  facilities in the concentric circle of 500m radius. The estimated coefficients here are  $_{1070}$  consistently positive but not statistically significant at conventional levels. In Model 5,  $_{1071}$  the estimated coefficient is 0.911 (SE 0.614, p = .149, Table D10), which implies that an  $_{1072}$  additional land parcel for transport facility within the concentric circle is associated with  $_{1073}$  a 0.9 point increase in the plasma PFBS concentration (0.06 of the SD).

Table D12 reports the estimation using a 15-min travel buffer around residences to <sup>1075</sup> compute exposure. The estimated coefficients across the models are negative but not <sup>1076</sup> statistically significant. The estimated association for Model 5 is -0.014 (SE 0.069, p <sup>1077</sup> = .841, Table D12). Using a 10-min buffer reported in Table D11 and a 20-min buffer <sup>1078</sup> reported in Table D13 yield similar null findings.



**Figure D26.** Correlation between alternate spatial instruments. The four key alternative spatial instruments of transport facilities are: (1) area of transport facilities within concentric circle (main instrument in Section 2.3), (2) distance to nearest transport facilities, (3) number of transport facilities in concentric circle, and (4) area of transport facilities within 15-minute of residence. Confidence intervals are bootstrapped (n = 1000). Main diagonal reports the univariate histogram.

	D	ependen	t variable	e is PFBS	(ng/mL)	
	(1)	(2)	(3)	(4)	(5)	(6)
Distance to nearest transport facility	-1.255	-0.284	-0.978	-1.449	-0.676	1.091
	(2.056)	(2.115)	(1.911)	(2.023)	(3.384)	(3.286)
Constant	$24.737^{a}$	11.151	11.619	17.809	24.801	17.482
	(0.945)	(18.944)	(17.973)	(18.122)	(23.320)	(29.096)
$\overline{\mathbb{R}^2}$	0.001	0.054	0.063	0.091	0.188	0.209
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.0	23.7
Std. dev. of X	0.3	0.3	0.3	0.3	0.3	0.3
n(Distance to nearest transport facility > 0)	771	742	742	741	720	670
n(Clusters)	_	30	30	29	29	28
N	771	742	742	741	720	670

## Table D8. PFBS and (point-to-edge) distance to nearest transport facility

Note: The main independent variable—Distance to nearest transport facility—is the nearest distance from the given participant residence to the nearest transport facility land parcel computed as point-to-edge distance (in 1,000 meters). Other than the spatial instrument of exposure to transport facilities, models and adjustments are the same as in Equation (1) and Table A3. Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

	D	ependen	t variable	e is PFBS	(ng/mL)	
	(1)	(2)	(3)	(4)	(5)	(6)
Distance to nearest transport facility	-0.883	0.102	-0.565	-0.810	0.123	1.872
	(2.067)	(2.116)	(1.913)	(1.994)	(3.282)	(3.145)
Constant	$24.590^{a}$	10.845	11.295	17.273	24.012	16.635
	(1.037)	(18.945)	(17.957)	(18.036)	(23.067)	(28.776)
$R^2$	0.000	0.054	0.063	0.090	0.188	0.209
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.0	23.7
Std. dev. of X	0.3	0.3	0.3	0.3	0.3	0.3
n(Distance to nearest transport facility > 0)	771	742	742	741	720	670
n(Clusters)	_	30	30	29	29	28
Ν	771	742	742	741	720	670

## Table D9. PFBS and (point-to-centroid) distance to nearest transport facility

Note: The main independent variable—Distance to nearest transport facility—is the nearest distance from the given participant residence to the nearest transport facility land parcel computed as point-to-centroid distance in meters (in 1,000 meters). Other than the spatial instrument of exposure to transport facilities, models and adjustments are the same as in Equation (1) and Table A3. Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

	D	ependen	t variable	e is PFBS	(ng/mL)	
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities within 500m	0.482	0.538	0.791	$0.879^{c}$	0.911	0.381
	(0.484)	(0.485)	(0.490)	(0.501)	(0.614)	(0.668)
Constant	$23.636^{a}$	10.484	10.127	15.898	24.112	18.389
	(0.803)	(18.292)	(17.457)	(17.614)	(22.027)	(27.758)
$\overline{R^2}$	0.002	0.056	0.067	0.095	0.190	0.209
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.0	23.7
Std. dev. of X	1.4	1.4	1.4	1.4	1.4	1.4
n(Transport facilities within $500m > 0$ )	458	439	439	439	426	399
n(Clusters)	_	30	30	29	29	28
Ν	771	742	742	741	720	670

## Table D10. PFBS and the number of transport facilities

Note: The main independent variable is the number of transport facilities within the 500m concentric circle. Models and adjustments are the otherwise the same as in Equation (1) and Table A3. Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

	D	ependen	t variable	e is PFBS	(ng/mL)	
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 10min	-0.005	-0.005	0.004	0.011	-0.109	-0.099
	(0.177)	(0.158)	(0.145)	(0.149)	(0.113)	(0.103)
Constant	$24.147^{a}$	10.129	9.336	14.461	22.272	16.424
	(0.600)	(18.207)	(17.389)	(17.361)	(21.962)	(26.756)
$R^2$	0.000	0.047	0.056	0.086	0.188	0.212
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	23.9	23.7
Std. dev. of X	4.2	4.2	4.2	4.2	4.0	4.1
n(Transport facilities (area) within 10min > 0)	271	263	263	263	252	230
n(Clusters)	—	28	28	28	28	27
Ν	744	715	715	715	695	646

## **Table D11.** PFBS and transport facilities within 10 min of residence

Note: Transport facilities area is the area (in 1,000 square meters) within 15 minutes of residence that is allocated to transport facilities land use. Maternal characteristics include (i) age at delivery (quadratic), (ii) ethnicity, (iii) education, (iv) occupation, (v) marital status, and (vi) housing type. Income includes mother's income and household income (binned). Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

	D	ependen	t variable	e is PFBS	(ng/mL)	
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 15min	-0.017	-0.005	-0.010	-0.002	-0.014	0.004
	(0.043)	(0.042)	(0.046)	(0.051)	(0.069)	(0.067)
Constant	$24.276^{a}$	10.146	9.255	14.415	22.484	16.453
	(0.727)	(18.424)	(17.561)	(17.491)	(21.991)	(26.754)
$\overline{R^2}$	0.000	0.047	0.056	0.086	0.188	0.212
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	23.9	23.7
Std. dev. of X	16.3	16.3	16.3	16.3	16.2	16.5
n(Transport facilities (area) within 15min > 0)	576	553	553	553	535	496
n(Clusters)	_	28	28	28	28	27
Ν	744	715	715	715	695	646

## Table D12. PFBS and transport facilities within 15 min of residence

Note: Transport facilities area is the area (in 1,000 square meters) within 15 minutes of residence that is allocated to transport facilities land use. Maternal characteristics include (i) age at delivery (quadratic), (ii) ethnicity, (iii) education, (iv) occupation, (v) marital status, and (vi) housing type. Income includes mother's income and household income (binned). Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

	Dependent variable is PFBS (ng/mL)					
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 20min	-0.010	-0.004	-0.012	-0.007	-0.039	-0.028
	(0.017)	(0.018)	(0.021)	(0.021)	(0.036)	(0.034)
Constant	$24.382^{a}$	10.424	9.964	14.543	23.788	18.077
	(0.729)	(18.265)	(17.357)	(17.429)	(21.822)	(26.827)
$\overline{R^2}$	0.001	0.047	0.057	0.086	0.190	0.213
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	23.9	23.7
Std. dev. of X	35.7	36.1	36.1	36.1	35.5	35.9
n(Transport facilities (area) within 20min > 0)	701	672	672	672	653	608
n(Clusters)	_	28	28	28	28	27
Ν	744	715	715	715	695	646

## **Table D13.** PFBS and transport facilities within 20 min of residence

Note: Transport facilities area is the area (in 1,000 square meters) witin 20 minutes of residence that is allocated to transport facilities land use. Maternal characteristics include (i) age at delivery (quadratic), (ii) ethnicity, (iii) education, (iv) occupation, (v) marital status, and (vi) housing type. Income includes mother's income and household income (binned). Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{a}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

#### E. Movers and future exposure

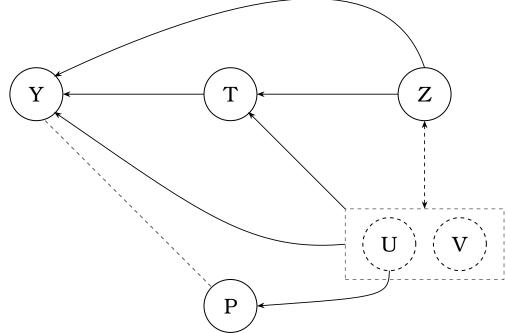
This appendix details how we focus on the subset of known movers in the GUSTO cohort <sup>1081</sup> since the start of the study and assess whether their future exposure is able to predict <sup>1082</sup> their past plasma PFBS concentration. <sup>1083</sup>

The fundamental concern is that the choice of residence is rarely random. Some 1084 unmeasured family characteristics might potentially correlate with lifestyle habits and 1085 non-point PFAS exposures which at the same time determines residence in a way that 1086 systematically leads to residential proximity to transport facilities, such locating near 1087 edges of neighborhoods or arterial roads. To help rule out this channel of confounding, we 1088 exploit the trail of GUSTO residence into the future (around 2022). Figure E27 provides 1089 a graph of this approach where there should be no association between future exposure 1090 and the (past) PFAS measures.<sup>4</sup>

We identify movers based on changes in reported residential addresses. More than <sup>1092</sup> half of the GUSTO participants in our sample (59%) have moved since the start of the <sup>1093</sup> study around 2009. We use the latest known residential address of these "movers" to <sup>1094</sup> compute the exposure to transport facilities, which is as defined in Section 2.3, except <sup>1095</sup> that residence is based on the latest residence and the spatial distribution of transport <sup>1096</sup> facilities is based on the 2019 versions. The plasma PFBS measurement remains the <sup>1097</sup> same version as the one collected during the start of the study around 2009. The regions, <sup>1098</sup> planning areas, and subzones are also based on the 2019 versions by mapping the postal <sup>1099</sup> 1099 codes onto the 2019 vector data.

Table E14 starts by repeating the model estimations defined in Equation (1) and reported in Table A3, but only for participants who moved in the future. We see that the <sup>1102</sup> associations are broadly comparable to the full sample (Table A3). For Model 5, the estimated association is 0.174 (SE 0.073, p = .025, Table E14) which is similar to the <sup>1104</sup> estimated association from the main analyses (0.153, SE 0.056, p = .019, Table A3). The <sup>1105</sup> estimated association from Model 6 is similar (0.174, SE 0.098, p = .090, Table E14). <sup>1106</sup>

Table E15 then reports the estimated associations between future exposure, as a negative control exposure, where we know that future exposure should not be able to predict <sup>1108</sup> pass plasma PFBS concentration through some unmeasured family characteristic(s) determining location choice. We do not see any statistically significant association for any <sup>1110</sup> model. The estimated coefficient under Model 5 is negative but not statistically signifi-1111 cant (-0.057, SE 0.115, p = .623, Table E15). Model 6 has similar null findings (-0.075, <sup>1112</sup> SE = 0.132, p = .576, Table E15).



**Figure E27.** Placebo evaluation via future addresses. Our study is interested in evaluating the relation between plasma PFAS measures (Y) and exposure to transport facilities (T). Our models account for certain (observable) baselines (Z; Section 2.5). Our models omit certain unobserved characteristics (U and V) linked to both Y and T (and potentially Z). Part of this unmeasured baseline may capture families residential choice that might affect T (U). This U should also affect future exposure to transport facilities (P) but otherwise have no relationship to past plasma PFAS concentration (Y). V is an unobserved confounder that is intractable.

	Dep. var. is PFBS					
	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 500m buffer	0.060	$0.102^{c}$	$0.109^{c}$	$0.202^{a}$	$0.174^{b}$	$0.174^{c}$
	(0.053)	(0.058)	(0.061)	(0.049)	(0.073)	(0.098)
Constant	$23.896^{a}$	10.884	9.721	1.972	4.683	12.631
	(0.654)	(24.644)	(24.029)	(21.104)	(25.023)	(32.057)
$R^2$	0.001	0.064	0.071	0.131	0.289	0.316
Maternal baselines		Yes	Yes	Yes	Yes	Yes
Income						Yes
Area fixed effects: Region			Yes			
Area fixed effects: Planning area				Yes		
Area fixed effects: Subzone					Yes	Yes
Mean Dep Var.	24.1	24.1	24.1	24.1	24.1	23.6
Std. dev. of X	9.9	9.9	9.9	9.9	9.9	10.0
n(Transport facilities (area) within 500m buffer > 0)	270	258	258	258	238	219
n(Clusters)	_	30	30	28	27	25
Ν	455	437	437	435	401	367

## Table E14. PFBS and area of transport facilities within 500m radius (Movers only).

Note: Sample includes only known movers in the GUSTO cohort. Table is otherwise identical to Table A3. Dependent variable is PFBS (Perfluorobutanesulfonic acid) of mothers in the GUSTO sample. Transport facilities area within 500m buffer is the area (in 1,000 square meters) of the buffer area residence that is allocated to transport facilities land use. Maternal characteristics include (i) age at delivery (quadratic), (ii) ethnicity, (iii) education, (iv) occupation, (v) marital status, and (vi) housing type. Income includes mother's income and household income (binned). Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{c} 0.1^{b} 0.05^{a} 0.01$ .

**Table E15.** PFBS and area of transport facilities within 500m radius (using future exposure).

	Dep. var. is PFBS					
-	(1)	(2)	(3)	(4)	(5)	(6)
Transport facilities (area) within 500m buffer	-0.069	-0.081	-0.099	-0.107	-0.057	-0.075
	(0.073)	(0.085)	(0.081)	(0.081)	(0.115)	(0.133)
Constant	24.343***	8.837	9.637	13.257	-9.589	-28.506
	(0.814)	(22.177)	(21.944)	(22.725)	(26.343)	(31.363)
$\overline{R^2}$	0.001	0.051	0.054	0.093	0.282	0.336
Maternal characteristics		Yes	Yes	Yes	Yes	Yes
Income						Yes
Region fixed effects			Yes			
Planning area fixed effects				Yes		
Subzone fixed effects					Yes	Yes
Mean of dep. var.	24.2	24.2	24.2	24.1	24.0	23.8
Std. dev. of X	6.5	6.5	6.5	6.5	6.5	6.6
Clusters	_	32	32	26	25	25
Ν	468	449	449	443	395	364

Note: Sample includes only known movers in the GUSTO cohort. The transport facilities area is based on the future addresses of GUSTO participants and not those around time of delivery. Table is otherwise identical to Table A3. See Table E14 for the same table but for movers only using addresses at time of delivery. Dependent variable is PFBS (Perfluorobutanesulfonic acid) of mothers in the GUSTO sample. Transport facilities area within 500m buffer is the area (in 1,000 square meters) of the buffer area residence that is allocated to transport facilities land use. Maternal characteristics include (i) age at delivery (quadratic), (ii) ethnicity, (iii) education, (iv) occupation, (v) marital status, and (vi) housing type. Income includes mother's income and household income (binned). Standard errors in (2)–(6) clustered at subzones. Significance levels:  $^{a}$  0.1  $^{b}$  0.05  $^{a}$  0.01.

#### **F. Other PFAS**

In this appendix, we test and report associations between exposure to transport facilities <sup>1115</sup> and the other well-measured PFAS concentrations. <sup>1116</sup>

#### **F.A GUSTO PFAS measurements**

The different PFAS substances have different measurement ranges (see for example, Figure 6). For ease of reporting, we scale the PFAS measurements by dividing by the standard deviation

substance<sup>(scaled)</sup> = 
$$\frac{\text{substance}}{\sigma_{(\text{substance})}}$$
 (F4)

so that the interpretation of regression coefficients is how much of a standard deviation  $^{1121}$  change in an input is associated with some standard deviation change in the output. For  $^{1122}$  the estimates in Figure 6, where we estimate the association of exposure to transport  $^{1123}$  facilities and the other PFAS substances, we also scale the area of transport facilities  $^{1124}$  within residence for ease of reporting in the same scale. Importantly, we note that this  $^{1125}$  linear scaling of the measurements does not aid causal inference, and separately, does  $^{1126}$  not affect the *t*-statistics or *p*-values.  $^{1127}$ 

Figure F28 reports the correlation between perfluorobutane sulfonic acid (PFBS) and <sup>1128</sup> the other plasma PFAS concentrations. All measurements are scaled as described above, <sup>1129</sup> and the correlation is conditional on area (subzones) of the GUSTO participant. Mea- <sup>1130</sup> surements below the limit of detection (LOD) and the limit of quantification (LOQ) are <sup>1131</sup> first imputed by the LOD/LOQ values divided by  $\sqrt{2}$ . Of the other seven PFAS measure- <sup>1132</sup> ments with detection rates for at least 95% of participants (Table A1), we note a positive <sup>1133</sup> correlation between the plasma PFBS concentration and perfluorohexane sulfonic acid <sup>1134</sup> (PFHxS) perfluoroundecanoic acid (PFUnDA), and perfluorodecanoic acid (PFDA). <sup>1135</sup>

To test the association between the other seven plasma PFAS concentrations and exposure to transport facilities around area, we estimate

substance<sup>(scaled)</sup><sub>ic</sub> = 
$$\beta$$
(Exposure to transport facilities)<sup>(scaled)</sup><sub>ic</sub>

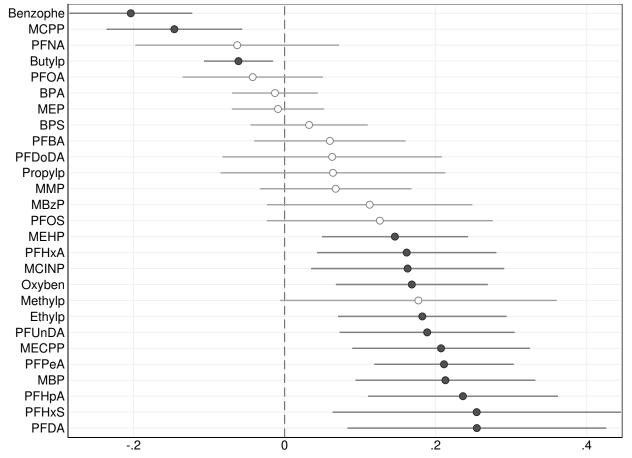
$$+\gamma X_i + \delta_c (\text{subzone})_c + \varepsilon_{ic}, \quad \text{(F5)}$$

where substance<sup>(scaled)</sup> is the PFAS substance (indicated in the first column). The spec-<sup>1136</sup> ification is otherwise identical to that in Equation (1) (to model (5) of Table A3). The <sup>1137</sup> PFAS substances and the exposure to transport facilities variable are scaled to have a <sup>1138</sup> standard deviation of one so that  $\hat{\beta}$  is interpreted as how a standard deviation change in <sup>1139</sup> exposure to transport facilities is associated with a  $\hat{\beta}$ -times standard deviation change in <sup>1140</sup> the substance. Figure 6 reports the results.

Figure F29 reports the results from Equation (2) for all of the eight PFAS measures <sup>1142</sup> with scaled units. We see that higher thresholds are linked to higher associations for <sup>1143</sup>

1114

1117

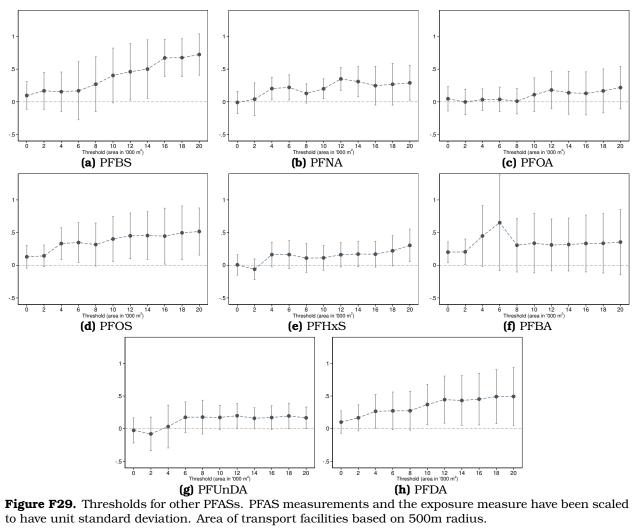


PFBS in Figure 5 (and in Figure F29 with scaled units). We observe similar patterns with <sup>1144</sup> PFNA, PFOS, and PFDA. <sup>1145</sup>

**Figure F28.** Correlation of PFBS (measured from cord blood) with other PFAS substances. Horizontal axis reports the  $\hat{\beta}$  coefficients from estimating

 $substance_{ic}^{(scaled)} = \beta PFBS_{ic}^{(scaled)} + \delta_c (subzone)_c + \varepsilon_{ic},$ 

where substance<sub>*ic*</sub> is one of the PFAS substances indicated on the vertical axis. Estimates significant at the 5% level have black markers; the remaining estimates have hollow markers. Gray horizontal lines are the 95% confidence intervals constructed from standard errors clustered at planning areas. Measurements that are below LOD and LOQ values are first imputed (Table A1) and then scaled to have a standard deviation of one.



#### **F.B S-PRESTO PFAS measurements**

We use participants from S-PRESTO, a separate cohort study in Singapore, as sup- <sup>1147</sup> plementary data for additional cross-sectional analyses. The key relevant detail to the <sup>1148</sup> present study is that S-PRESTO recruited participants between 2015–2017, and there- <sup>1149</sup> fore offers a different time point for evaluation. <sup>1150</sup>

The S-PRESTO (Singapore Preconception Study of Long-Term Maternal and Child Outcomes) recruited N = 1,039 ethnically-diverse (Chinese, Malay, or Indian) participants <sup>1152</sup> aged 18–45 who intended to get pregnant and deliver in Singapore between February <sup>1153</sup> 2015–October 2017. Similar to GUSTO, S-PRESTO is inclusive but there were exclusion <sup>1154</sup> criteria for to-be mothers with existing treatments and conditions (e.g., undergoing fertility treatment or had diabetes) so that the sample included mostly healthy mother-child <sup>1156</sup> pairs.<sup>4,68</sup>

Plasma PFAS concentrations from S-PRESTO comes from a preconception plasma 1158 blood sample collected at enrollment around 2015–2017. S-PRESTO collected only ma- 1159 ternal blood so we lack the same PFAS measurements from cord blood as a reflection of 1160 neonatal exposure. The samples were sent for testing around 2021 for 15 PFAS: perfluo-1161 rohexanesulfonic acid (PFHxS), linear and branched perfluorooctanesulfonic acid (PFOS), 1162 linear perfluorooctanoic acid (PFOA), perfluorononanoic acid (PFNA), perfluoroheptane- 1163 sulfonic acid (PFHpS), perfluorodecanoic acid (PFDA), perfluoroheptanoic acid (PFHpA), 1164 N-methylperfluorooctanesulfonamido acetic acid (NMeFOSAA), 6:2 polyfluoroalkyl phos- 1165 phate ester (6:2 PAP), 6:2 fluorotelomer phosphate diester (6:2 diPAP), perfluorobutane- 1166 sulfonic acid (PFBS), 6:2 fluorotelomer sulfonate (6:2 FTS), N-ethyl perfluorooctanesul- 1167 fonamidoacetic acid (NEtFOSAA), and perfluorodecanesulfonic acid (PFDS), and perflu-1168 orooctanesulfonamide (PFOSA). The measurements data came with below LOD values 1169 replaced to half of the LOD value and this is what we use. Table F16 reports the com- 1170 pleteness of PFAS detection and the corresponding LOD values ranked by completeness. 1171 For this set of analyses, we include the first eight measurements (seven PFAS) with de-1172 tection in at least 34% of the participants. The remaining PFAS are excluded. The seven 1173 included PFAS (LOD) are: PFHxS (0.10 ng/ml), PFOS (0.20 ng/ml), PFOA (0.50 ng/ml), 1174 PFNA (0.50 ng/ml), PFHpS (0.20 ng/ml), PFDA (0.50 ng/ml), and PFHpA (0.20 ng/ml). 1175

As far as possible, we adjust for maternal baselines in the S-PRESTO evaluations so <sup>1176</sup> that they are comparable to our evaluations using the GUSTO sample. For the S-PRESTO <sup>1177</sup> evaluations, we adjusts for age of mother during recruitment (flexibly in quadratics), ed- <sup>1178</sup> ucation, employment, marital status, ethnicity, and household income. We note that <sup>1179</sup> these adjustments do not map exactly to those in the GUSTO sample. Household in- <sup>1180</sup> come, for instance, would have different stratas compared to those asked in the earlier <sup>1181</sup> GUSTO recruitment. All other specifications and adjustments for unmeasured spatial <sup>1182</sup> heterogeneity are otherwise similar as defined in Equation (1). We are unable to focus on <sup>1183</sup> the PFBS (perfluorobutanesulfonic acid) since only one sample had successful detection <sup>1184</sup>

Full analyte name	Shorthand	Measured	< LOD	LOD value
Perfluorohexane-1-sulphonic acid	PFHxS	384	0	0.1 ng/ml
Perfluorooctanesulfonic acid	PFOS-Linear	384	0	0.2 ng/ml
Perfluorooctanesulfonic acid	PFOS-Branched	383	1	0.2 ng/ml
Perfluorooctanoic acid	PFOA-Linear	374	10	0.5 ng/ml
Perfluorononanoic acid	PFNA	233	151	0.5 ng/ml
Perfluoroheptanesulfonic acid	PFHpS	186	198	0.2 ng/ml
Perfluorodecanoic acid	PFDA	159	225	0.5 ng/ml
Perluoroheptanoic acid	PFHpA	130	254	0.2 ng/ml
2-(N-Methylperfluorooctanesulfonamido) acetic acid	NMeFOSAA	6	378	0.1 ng/ml
6:2 Fluorotelomer phosphate monoester	6:2 PAP	5	379	0.1 ng/ml
6:2 Fluorotelomer phosphate diester	6:2 diPAP	2	382	0.1 ng/ml
Perfluorobutanesulfonic acid	PFBS	1	383	0.2 ng/ml
6:2 Fluorotelomer sulfonic acid	6:2 FTS	0	384	0.1 ng/ml
2-(N-Ethylperfluorooctanesulfonamido) acetic acid	NEtFOSAA	0	384	0.2 ng/ml
Perfluorodecanesulfonic acid	PFDS	0	384	0.1 ng/ml
Perfluorooctanesulfonamide	PFOSA	0	384	0.1 ng/ml

Table F16. Completeness of S-PRESTO plasma PFAS measurements

Note: Table reports the completeness of plasma PFAS measurements in the S-PRESTO sample (n = 384). Measurements in nanograms per millilitre (ng/ML). LOD = limit of detection.

above the LOD (Table F16). These estimates are reported in Figure F30.<sup>7</sup> 1185 The exposure to transport facilities measure is as defined in Section 2.3, except that 1186 we map the residence of the S-PRESTO participants to the 2014 land use plans. Likewise, the regions, planning areas, and subzones are also based on the 2014 versions by 1188 mapping the postal codes onto the 2014 vector data. 1189

Full analyte name	N	< LOD				
			LOD value		Est. (95% Conf. Int.)	P-value
Perfluorohexane-1-sulphonic acid	384	0	0.1 ng/ml		0.038(-0.022 to 0.098)	0.207
Perfluorooctanesulfonic acid	384	0	0.2 ng/ml		0.100( 0.025 to 0.176)	0.012 <sup>b</sup>
Perfluorooctanesulfonic acid	383	1	0.2 ng/ml		0.061(-0.040 to 0.161)	0.222
Perfluorooctanoic acid	374	10	0.5 ng/ml		0.028(-0.052 to 0.108)	0.473
Perfluorononanoic acid	233	151	0.5 ng/ml		0.112( 0.011 to 0.213)	0.031 <sup>b</sup>
Perfluoroheptanesulfonic acid	186	198	0.2 ng/ml		0.097( 0.012 to 0.182)	0.027 <sup>b</sup>
Perfluorodecanoic acid	159	225	0.5 ng/ml		0.031(-0.014 to 0.077)	0.171
Perluoroheptanoic acid	130	254	0.2 ng/ml		0.124( 0.042 to 0.205)	0.005 <sup>a</sup>
	Perfluorooctanesulfonic acid Perfluorooctanoic acid Perfluorononanoic acid Perfluoroheptanesulfonic acid Perfluorodecanoic acid	Perfluorooctanesulfonic acid 383 Perfluorooctanoic acid 374 Perfluorononanoic acid 233 Perfluoroheptanesulfonic acid 186 Perfluorodecanoic acid 159	Perfluorooctanesulfonic acid3831Perfluorooctanoic acid37410Perfluorononanoic acid233151Perfluoroheptanesulfonic acid186198Perfluorodecanoic acid159225	Perfluorooctanesulfonic acid38310.2 ng/mlPerfluorooctanoic acid374100.5 ng/mlPerfluorononanoic acid2331510.5 ng/mlPerfluoroheptanesulfonic acid1861980.2 ng/mlPerfluoroheptanoic acid1592250.5 ng/mlPerfluoroheptanoic acid1302540.2 ng/ml	Perfluorooctanesulfonic acid       383       1       0.2 ng/ml         Perfluorooctanoic acid       374       10       0.5 ng/ml         Perfluorononanoic acid       233       151       0.5 ng/ml         Perfluoroheptanesulfonic acid       186       198       0.2 ng/ml         Perfluoroheptanesulfonic acid       159       225       0.5 ng/ml         Perfluoroheptanoic acid       130       254       0.2 ng/ml	Perfluorooctanesulfonic acid       383       1       0.2 ng/ml       0.061(-0.040 to 0.161)         Perfluorooctanoic acid       374       10       0.5 ng/ml       0.028(-0.052 to 0.108)         Perfluorononanoic acid       233       151       0.5 ng/ml       0.112( 0.011 to 0.213)         Perfluoroheptanesulfonic acid       186       198       0.2 ng/ml       0.097( 0.012 to 0.182)         Perfluorodecanoic acid       159       225       0.5 ng/ml       0.031(-0.014 to 0.077)

**Figure F30.** Figure reports estimates for PFAS measurements from the S-PRESTO cohort study. Horizontal axis of the plot reports the  $\hat{\beta}$  coefficients from estimating

 $PFASic^{(\text{scaled})} = \beta(\text{Exposure to transport facilities})ic^{(\text{scaled})} + \gamma X_i + \delta_c(\text{subzone})c + \varepsilon ic,$ 

where  $PFAS_{ic}^{(scaled)}$  is the PFAS substance (indicated in the first column). Transport facilities and regional tags are based on the 2014 versions. All models adjust for mother's age (in quadratics), employment status, education, marital status, ethnicity, and household income collected from the S-PRESTO sample. Gray horizontal lines are the 95% confidence intervals constructed from standard errors clustered at planning areas. Significance levels: <sup>*c*</sup> 0.1 <sup>*b*</sup> 0.05 <sup>*a*</sup> 0.01.

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