1	Associations of the neighbourhood built and natural environment with cardiometabolic health
2	indicators: A cross-sectional analysis of environmental moderators and behavioural mediators
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# 39 Abstract

40 Background: Most studies examining the effects of neighbourhood urban design on cardiometabolic

41 health focused solely on the built or natural environment. Also, they did not consider the roles of

42 neighbourhood socio-economic status (SES) and ambient air pollution in the observed associations,

and the extent to which these associations were mediated by physical activity and sedentary

- 44 behaviours.
- 45 **Methods:** We used data from the AusDiab3 study (N=4141), a national cohort study of Australian
- 46 adults to address the above-mentioned knowledge gaps. Spatial data were used to compute indices of

47 neighbourhood walkability (population density, intersection density, non-commercial land use mix,

48 commercial land use), natural environment (parkland and blue spaces) and air pollution (annual

49 average concentrations of nitrogen dioxide (NO<sub>2</sub>) and fine particulate matter  $<2.5 \mu m$  in diameter

50 (PM<sub>2.5</sub>)). Census indices were used to define neighbourhood SES. Clinical assessments collected data

on adiposity, blood pressure, blood glucose and blood lipids. Generalised additive mixed models were

52 used to estimate associations.

53 **Results:** Neighbourhood walkability showed indirect beneficial associations with most indicators of

54 cardiometabolic health via resistance training, walking and sitting for different purposes; indirect

b5 detrimental associations with the same indicators via vigorous gardening; and direct detrimental

so associations with blood pressure. The neighbourhood natural environment had beneficial indirect

57 associations with most cardiometabolic health indicators via resistance training and leisure-time

sitting, and beneficial direct associations with adiposity and blood lipids. Neighbourhood SES and air

59 pollution moderated only a few associations of the neighbourhood environment with physical activity,

60 blood lipids and blood pressure.

61 **Conclusions:** Within a low-density and low-pollution context, denser, walkable neighbourhoods with 62 good access to nature may benefit residents' cardiometabolic health by facilitating the adoption of an 63 active lifestyle. Possible disadvantages of living in denser neighbourhoods for older populations are 64 having limited opportunities for gardening, higher levels of noise and less healthy dietary patterns

- 65 associated with eating out.
- 66

*Keywords:* walkability; greenspace; blue space; physical activity; neighbourhood socio-economicstatus; air pollution

69	Funding
70	This work was supported by a program grant ("The environment, active living and cognitive
71	health: building the evidence base") from the Australian Catholic University [grant number
72	ACURF18]. Jonathan E. Shaw is supported by a National Health and Medical Research Council
73	(NHMRC) Investigator Grant [grant number 1173952].
74	The funders had no role in study design, data analysis, interpretation of the results, the
75	decision to publish, or preparation of the manuscript.
76	Ethical statement
77	The AusDiab study was approved by the Alfred Hospital Ethics Committee (ref. no. 39/11)
78	and conducted according to the guidelines of the Declaration of Helsinki. All participants provided
79	written consent prior to partaking in the study.
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## 106 **1. Introduction**

Cardiovascular diseases (CVD) top the list of causes of death in Australia (Global Burden of 107 108 Disease Study, 2019) and globally (Global Burden of Disease Study, 2016). CVD risk can be reduced 109 by tackling related cardiometabolic and behavioural risk factors. The former include obesity, elevated 110 blood pressure, elevated blood glucose and dyslipidaemia (Dahlöf, 2010). An important behavioural 111 risk factor for CVD and the above-mentioned cardiometabolic risk factors is physical inactivity 112 (Balakumar et al., 2016; Cunningham et al., 2020). With 28% of adults being insufficiently physically active (Guthold et al., 2018) and 39% being overweight (Loos & Yeo, 2022) globally, large-scale, 113 long-term sustainable interventions are required to address the high prevalence of CVD and associated 114 115 risk factors.

116 Physical features of residential neighbourhoods, such as residential density and access to amenities and nature, have been identified as important, large-scale modifiable determinants of 117 physical activity and health (Giles-Corti et al., 2022; World Health Organisation, 2009; World Health 118 119 Organization, 2020), especially in ageing populations (World Health Organisation, 2020), who are 120 more susceptible to CVD (World Health Organisation, 2017). Neighbourhood design can impact ambient air pollution (Borrego et al., 2006; Münzel et al., 2018; Wang et al., 2017) and lifestyle 121 behaviours (physical activity and sedentary behaviours), known to affect cardiometabolic risk factors 122 (e.g., obesity and elevated blood glucose) (An et al., 2018; Balakumar et al., 2016; Cunningham et al., 123 2020; Honda et al., 2017; Münzel et al., 2018; Zhang et al., 2019). Specifically, it is well established 124 125 that a physically active lifestyle contributes to better cardiovascular health by exerting beneficial 126 effects on the heart (e.g., lower resting heart rate, improved mitochondrial biogenesis and greater cardiac output), blood vessels (e.g., lower resting blood pressure, vascular resistance and 127 128 atherosclerotic plaque formation), blood (e.g., increased insulin sensitivity and insulin-dependent 129 glucose uptake, better lipid profile) and by reducing systemic inflammation (Nystoriak & Bhatnagar, 130 2018), while exposure to ambient air pollution is deemed to harm cardiometabolic health by increasing oxidative stress and systemic inflammation (Brook, 2008). 131 It is, thus, important to understand how neighbourhood environments are associated with CVD 132

risk factors and related behaviours, and how such risk factors can be reduced through urban and transport planning. In fact, suboptimal urban and transport planning resulting in higher NO<sub>2</sub> concentrations, carbon emissions, loss of green spaces and activity-unfriendly environments have been

blamed for substantially contributing to morbidity and premature mortality (Bird et al., 2018; Mueller
et al., 2021; Nieuwenhuijsen, 2020).

Studies suggest that adult residents of denser areas with better street connectivity and access to
a variety of services and natural features (e.g., greenspace) tend to walk more for utilitarian purposes
(Cerin et al., 2017) and, in general, be more physically active in their leisure time (Georgiou et al.,
2021; Van Cauwenberg et al., 2018). There is also some evidence that, by engaging in more physical
activity as a result of living in walkable, destination-rich environments, older people may reduce time

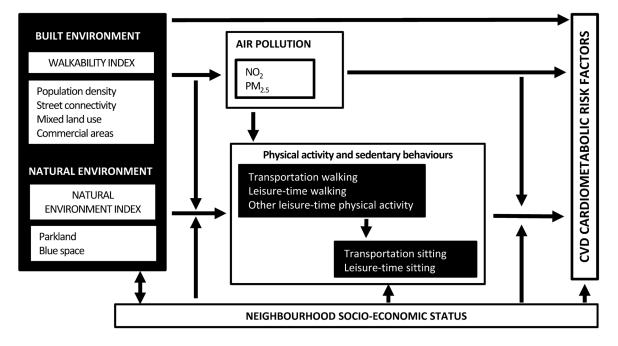
spent sitting (Astell-Burt et al., 2014; Barnett et al., 2015; Cerin et al., 2020; Cerin, Zhang et al.,

- 144 2023). It follows that residents of walkable neighbourhoods with good access to amenities and
- 145 greenspace are likely to have better cardiometabolic health (e.g., lower probability of being
- 146 overweight/obese or having elevated blood glucose) and, hence, lower risk of CVD. In fact, recent
- systematic reviews report beneficial effects of greenspace on CVD mortality (Liu et al., 2022) and
- 148 cardiometabolic risk factors, including elevated blood pressure (Fan et al., 2022), overweight/obesity
- 149 (Rahimi-Ardabili et al., 2021), elevated blood glucose and dyslipidaemia (Dendup et al., 2018;
- 150 Rahimi-Ardabili et al., 2021). Similarly, access to blue space has been reported to be negatively
- associated with abdominal adiposity, blood glucose and dyslipidaemia (Cerin et al., 2022), but, in
- 152 general, findings are mixed (Geneshka et al., 2021). In contrast, the evidence of walkable
- neighbourhoods (typified by higher levels of density, street connectivity and access to services) having
- beneficial effects on overweight/obesity, and elevated blood pressure and glucose has somewhat been
- 155 more consistent (Chandrabose et al., 2019).

156 A major limitation of studies examining built and natural environmental correlates of CVD risk factors, and especially those focusing on neighbourhood walkability, pertains to many of them not 157 158 accounting for ambient air pollution, which is a by-product of urbanisation (Cerin, 2019; James et al., 159 2015). Dense, destination-rich neighbourhoods that promote an active lifestyle are often accompanied by higher volumes of traffic and traffic-related air pollution (Khreis et al., 2023) that contribute to 160 CVD (Huang et al., 2021) and related cardiometabolic (Gaio et al., 2019; Liu et al., 2019; Wang et al., 161 2023) and behavioural risk factors (An et al., 2019). For example, exhaust fumes from vehicles and 162 163 media warnings about poor air quality may deter residents from engaging in outdoor physical activity (An et al., 2019). Conversely, the presence of greenery in a neighbourhood may reduce traffic-related 164 165 air pollution (Hirabayashi & Nowak, 2016). As the neighbourhood natural and built environments 166 impact ambient air pollution, and the latter increases the risk of CVD, to estimate the independent 167 contribution of the neighbourhood built and natural environment on CVD and its risk factors, it is important to consider ambient air pollution. Air pollution may not only directly impact CVD 168 cardiometabolic and behavioural risk factors, it can also determine the strength and direction of 169 170 associations between the neighbourhood environment and risk factors (Howell et al., 2019), as well as 171 associations among risk factors (e.g., physical activity and elevated blood glucose) (D'Oliveira et al.,

- 172 2023; Hou et al., 2021).
- 173 Another important environmental factor that may explain CVD risk factors, as well as modify
- their relationships with the built and natural environment is neighbourhood socio-economic status
- 175 (SES). Neighbourhood SES has been identified as a key determinant of cardiometabolic health
- 176 (Barnett et al., 2022; Carroll et al., 2020; Keita et al., 2014; Mohammed et al., 2019; Tiwari et al.,
- 177 2022; Williams et al., 2012), physical activity (Cerin & Leslie, 2008; Grant et al., 2010; Tiwari et al.,
- 178 2022; Zhu et al., 2021) and sedentary behaviours (Proper et al., 2007), with those living in more
- advantaged neighbourhoods being healthier and more physically active. However, less is known about

- 180 the extent to which neighbourhood SES moderates associations between the neighbourhood
- 181 environment and CVD risk factors, and between CVD behavioural risk factors and cardiometabolic
- 182 risk factors. While, in their recent systematic review, Rigolon and colleagues reported stronger
- 183 beneficial effects of greenspace on cardiometabolic health among residents of more disadvantaged
- areas (Rigolon et al., 2021), the evidence of the moderating role of neighbourhood SES in relation to
- 185 other neighbourhood environmental attributes and CVD behavioural risk factors is mixed and
- inconclusive (Sallis et al., 2009). Understanding the relative importance of environmental and
- 187 behavioural factors that contribute to cardiometabolic health in communities with different levels of
- 188 social disadvantage can inform interventions aimed at reducing health inequalities, one of the key
- 189 goals of the United Nations sustainable development goals agenda (United Nations, 2021).



- Fig. 1. Simplified model of neighbourhood environmental correlates of cardiometabolic risk factors ofcardiovascular disease (CVD).
- 193

To address the above-mentioned knowledge gaps, the aims of this cross-sectional study were three-fold. We examined: (1) associations of the neighbourhood natural and built environment with

196 CVD cardiometabolic risk factors in mid-aged and older Australians, while adjusting for

- 197 neighbourhood SES and ambient air pollution; (2) the mediating roles of domain-specific physical
- 198 activity and sedentary behaviours in these associations; and (3) ambient air pollution and
- 199 neighbourhood SES as moderators of environment-cardiometabolic risk factor associations and related
- 200 physical activity and sedentary behaviour pathways, as depicted in Figure 1. It noteworthy that,
- 201 according to this theoretical model, physical activity is an antecedent of sedentary behaviour because,
- 202 from an evolutionary perspective, the natural tendency of adults (and older adults) is to preserve
- 203 energy (i.e., be inactive) unless they have specific reasons to be active, such as performing activities of

daily living or exercising for health or leisure purposes (Caldwell, 2016; Speakman, 2020). Thus, in
adults and older adults, it makes more sense to assume that physical activity displaces sedentary
behaviour (which, from an evolutionary perspective, is the "default" behaviour) than the opposite.

207

208 **2.** Methods

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# 210 2.1. Study design and participants

211 This study employed data from the third wave of the Australian Diabetes, Obesity and 212 Lifestyle study (AusDiab3) (Dunstan et al., 2002; Tanamas et al., 2013) conducted in 2011-12 and 213 enriched with spatial indicators of neighbourhood SES, walkability, natural environment and ambient 214 air pollution detailed in subsection 2.2.1 (Cerin et al., 2022; Cerin, Barnett et al., 2023). Only data from AusDiab3 (N = 4614) were utilised because spatial data corresponding to earlier AusDiab waves 215 216 were of inadequate quality or unavailable (Cerin, Barnett et al., 2023). The analytical sample (N =217 4141) was restricted to participants who at the time of the assessment were living in urban areas 218 (towns or cities with 10,000 or more inhabitants (ABS, 2017) consisting of 1286 Statistical Areas 1 219 (SA1, the smallest census administrative units in Australia).

Briefly, in 1999-2000, AusDiab recruited and examined adults (25+ years of age) with no 220 physical or intellectual disabilities and who had resided for at least six months in one of 42 randomly-221 222 selected urban areas across Australia. Follow-up assessments were conducted in 2004-05 and 2011-12. Data collection (surveys and cardiometabolic health biomarkers) was done in person at local testing 223 224 sites (Tanamas et al., 2013). AusDiab data collection procedures, and response and attrition rates have been previously reported (Dunstan et al., 2002; Tanamas et al., 2013). The AusDiab study was 225 approved by the Alfred Hospital Ethics Committee (ref. no. 39/11) and conducted according to the 226 227 guidelines of the Declaration of Helsinki. All participants provided written consent prior to partaking 228 in the study.

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## 230 *2.2. Measures*

231 2.2.1. Neighbourhood characteristics (environmental exposures)

A participant's neighbourhood was defined as an area within 1 km from their residential address following the street network, approximating the distance that an able-bodied adult can walk in 10-20 minutes (Adams et al., 2014). This is a frequently used neighbourhood definition (Cerin et al., 2013; Gunn et al., 2017) that, in an Australian context, has yielded stronger associations with walking

- behaviours than definitions of neighbouhood based on shorter (e.g., 0.5 km) or longer (e.g., 1.6 km)
- distances (Gunn et al., 2017). ESRI's ArcGIS v.10.5 software (ESRI, Redlands) was used to generate
- 238 spatial indicators of the neighbourhood environment around participants' residential addresses, the
- spatial distribution of which can be found in the supplementary data (Figure S4). For sensitivity

- analyses purposes, we also computed spatial indicators for 500 m and 1.6 km residential buffers. Nine
- 241 neighbourhood environmental characteristics were computed for each participant's neighbourhood.
- 242 These included four built environment attributes [population density (persons/ha), street intersection
- 243 density (intersections/km<sup>2</sup>), percentage of commercial land use and non-commercial land use mix (an
- entropy score of non-commercial land uses ranging from 0 to 1)], two natural environment attributes
- 245 (percentage of parkland and percentage of blue space), neighbourhood SES (Index of Relative
- 246 Socioeconomic Advantage and Disadvantage, IRSAD) and two ambient air pollution measures
- $[annual average concentrations of nitrogen dioxide (NO_2, unit: ppb) and fine particulate matter < 2.5$
- 248  $\mu$ m in aerodynamic diameter (PM<sub>2.5</sub>; unit:  $\mu$ g/m<sup>3</sup>)]. Concentrations of air pollutants were estimated at
- 249 the participants' residential addresses using satellite-based land-use regression models. The models
- $\label{eq:250} utilised spatial predictors of annual average NO_2 and PM_{2.5} at fixed-site monitors (e.g., roads,$
- 251 industrial emissions), including time-varying information from satellites, to calculate concentrations at
- unmeasured locations (e.g., residential addresses) (Knibbs et al., 2014; Knibbs et al., 2016; Knibbs et al., 2016;
- al., 2018). Cross-validation revealed that the  $NO_2$  model captured 81% of spatial variability in annual
- 254 NO<sub>2</sub> (RMSE: 1.4 ppb), while the PM<sub>2.5</sub> model captured 63% of spatial variability (RMSE:  $1 \mu g/m^3$ ).
- 255 All neighbourhood measures were based on spatial data collected during AusDiab3 assessments
- (2011-12). Detailed descriptions of the environmental exposures and their data sources have been
  published elsewhere (Cerin et al., 2021; Cerin, Barnett et al., 2023) and can be also found in the
  supplementary data.
- 259 For the purpose of this study, we used four composite indices representing the neighbourhood 260 built, natural, socio-economic and air quality environments. In line with the extant literature (Cerin et 261 al., 2016; Frank et al., 2010), we summed the standardised values of population and street intersection 262 densities, percentage of commercial land use and non-commercial land use mix to obtain a walkability 263 index representing the built environment. A natural environment index was computed by summing the 264 standardised values of percentage of parkland and blue space in the neighbourhood. IRSAD is a composite measure of neighbourhood SES tailored to the Australian context and encompasses census-265 derived information on SES indicators such as household income, educational attainment, 266 267 occupational class, housing conditions, and mortgage and rental costs (ABS, 2011). Finally, as estimates of all air pollutants typically included in air quality indices (NSW Department of Planning 268 269 and Environment, n.d.) were not available, an ambient air pollution index was obtained by summing 270 the standardized values of annual average concentrations of NO2 and PM2.5, which, in this study, were
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# 273 2.2.2. Cardiometabolic health indicators (outcomes)

positively correlated (Spearman's  $\rho = 0.27$ ).

The outcomes of this study were a series of cardiometabolic health indicators, including an indicator of adiposity [waist circumference (in cm)], an indicator of hypertension [mean arterial blood pressure (MAP; in mmHg)], an indicator of hyperglycaemia [glycated haemoglobin (HbA1c, in

- 277 mmol/mol)], and three indicators of dyslipidaemia [low-density lipoprotein (LDL) cholesterol
- 278 (mmol/L), high-density lipoprotein (HDL) cholesterol (mmol/L) and triglycerides (mmol/L)]. The
- assessment of cardiometabolic health indicators in the AusDiab study has been previously described in
- detail (Dunstan et al., 2002; Tanamas et al., 2013) and is summarised in the supplementary data.
- 281
- 282 2.2.3. Physical activity and sedentary behaviours (mediators)

283 We used domain-specific measures of physical activity and sedentary behaviours that are deemed to be impacted by different aspects of the neighbourhood environment (Cerin et al., 2017; Van 284 Cauwenberg et al., 2018). These included four measures of physical activity (previous-week 285 286 frequencies of engagement in walking for transport, walking for recreation, vigorous gardening and 287 resistance training) adapted from the Active Australia survey (Australian Institute of Health and Welfare, 2003), and two measures of sedentary behaviour (previous-week average daily hours of 288 289 sitting time for transport and leisure) developed for AusDiab3. The leisure-time sitting measure 290 included time spent on screen-based leisure activities (e.g., TV watching).

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# 292 2.2.4. Socio-demographic and other characteristics (confounders and covariates)

293 Participants self-reported socio-demographic and health-related information, including age, 294 educational attainment (up to secondary; trade or technician certificate; associate diploma or equivalent; bachelor degree or post-graduate diploma), employment status (not working; paid 295 296 employment; volunteer), ethnicity (English vs. non-English background), history of heart problems or stroke (ves; no), annual household income (<AUD 50,000; AUD 50,000 to AUD 99,999; >AUD 297 298 100,000), living arrangements (living with partner and no children; living with partner and children; living alone; other living arrangements), medications (diabetes medication; anti-hypertensive 299 300 medication; blood-lipid lowering medication), sex (female; male) and tobacco smoking status (current 301 smoker; past smoker; never smoked). Residential self-selection was assessed with two variables based 302 on participants' responses to items (on a 5-point scale, ranging from "not at all important" to "very important") gauging the importance of access to recreational facilities and a variety of destinations for 303 304 choosing to live in the current neighbourhood (Cerin et al., 2007; Owen et al., 2007).

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# 306 *2.3. Statistical analyses*

307 Descriptive statistics, including patterns of missing data, were first computed for all variables.
308 Given that a substantial percentage (17%) of participants had missing data on at least one variable and
309 data were not missing completely at random (e.g., missingness was related to participants' age,

- household income, IRSAD and population density) (Cerin et al., 2021; Cerin et al., 2022), 20 imputed
- 311 datasets were created using chained equations. Multiple imputations were performed using the
- package 'mice' (van Buuren & Groothuis-Oudshoorn, 2011) in R version 4.2.0 (R Core Team, 2018).

To estimate potentially-curvilinear associations of environmental exposures with 313 cardiometabolic health indicators (Cerin et al., 2021; Cerin et al., 2022) and examine the mediating 314 roles of physical activity and sedentary behaviours, we employed generalised additive mixed models 315 316 (GAMMs; package 'mgcv' version 1.8.42 in R) (Wood, 2017) with appropriate variance and link 317 functions and random intercepts accounting for spatial clustering at the SA1 level (Wood, 2017). We 318 used GAMMs with Gaussian variance and identity link functions to model waist circumference, MAP 319 and LDL cholesterol. These models produce regression coefficients, b, indicating the difference in the response variable associated with a 1 unit increase in the explanatory variable. Gamma variance and 320 321 logarithmic link functions were used to model glycated haemoglobin, HDL cholesterol, triglycerides, 322 sitting for different purposes and non-zero frequency of physical activity behaviours. The regression 323 coefficients of these models were exponentiated  $(e^b)$  so that they can be interpreted as the proportional increase (if  $e^{b}>1.0$ ) or decrease (if  $e^{b}<1.0$ ) in the response variable associated with a 1 unit increase in 324 the explanatory variable. Lastly, we employed GAMMs with binomial variances and logit link 325 functions to model engagement (yes vs. no) in specific physical activities. The exponentiated values of 326 327 the regression coefficients of these models are odds ratios (OR). Directed acyclic graphs (DAGs), 328 based on previous studies and the authors' expert opinion, guided the selection of a minimal sufficient 329 set of confounders for the regression models (Figure S1 and Table S1). We determined multicollinearity based on the Variance Inflation Factor (VIF) values of the variables included in the 330 331 GAMMs. All VIFs were smaller than 2.44, suggesting no substantial multicollinearity (Sheather, 332 2009).

333 A first set of models examined the independent effects of the neighbourhood built (walkability index), natural (natural environment index), socio-economic (IRSAD) environments and ambient air 334 335 pollution (air quality index) on cardiometabolic health indicators. Within these models, we also tested 336 the moderating effects of medications on the environment-outcome associations and retained 337 interaction terms that were statistically significant at a 0.05 probability level. Here, the word 'effect' is used as a statistical term (i.e., association) that does not provide evidence of causality. Given that air 338 339 pollution may mediate the associations between other environmental attributes and cardiometabolic 340 health (see Figure 1), we also estimated associations unadjusted for the ambient air pollution index as 341 supplementary analyses.

342 A second set of models examined IRSAD and the ambient air pollution index as moderators of 343 the associations between the walkability and natural environment indices and cardiometabolic health indicators by adding two- and three-way interaction terms (and four-way interaction terms in the case 344 of moderation by medication) to the second set of models and retaining the interaction terms that were 345 346 statistically significant at a 0.05 probability level. Statistically significant moderation effects were probed by estimating the associations of the exposures (walkability or natural environment indices) 347 with the cardiometabolic outcomes at different values of the moderators (IRSAD and ambient air 348 349 pollution index).

To examine the roles of physical activity and sedentary behaviours as potential mediators of 350 the above associations and moderating effects, we employed the joint-significant test (MacKinnon & 351 352 Luecken, 2008). We decided to use this test of mediation for multiple reasons. In simulation studies, 353 the joint-significance test displayed the best balance of Type I error and statistical power compared to 354 other tests, such as the product-of-coefficient test derived from structural equation modelling software 355 (MacKinnon et al. 2002). This was an important consideration because the statistical effects of 356 environmental attributes on behaviours and CVD risk factors are typically small and require statistically powerful methods of mediation. Secondly, this study examined multiple mediators. Hence, 357 selecting a test with reasonable Type I error rates was important. Thirdly, because all physical activity 358 359 mediating variables were zero inflated, they required to be modelled as two-part distributions. At 360 present, generalised structural equation models can accommodate zero inflated outcomes but not zero inflated mediators. Counterfactual-based mediation analysis can accommodate single zero inflated 361 362 mediators with normally distributed outcomes but not multiple zero inflated mediators or single zero 363 inflated mediators with outcomes following other distributional assumptions (Jiang et al., 2023).

According to the joint-significance test, mediation of a main effect is statistically confirmed if 364 365 exposure-mediator, exposure-adjusted mediator-outcome and, in the case of serial mediation, 366 mediator-mediator associations are statistically significant. Mediation of a moderation effect 367 (mediated moderation) is confirmed if (1) the moderator of an exposure-outcome association is also a 368 moderator of the exposure-mediator association and the exposure-adjusted mediator-outcome 369 association is statistically significant; or (2) the exposure-mediator association is statistically 370 significant and the moderator of an exposure-outcome association is also a moderator of the exposure-371 adjusted mediator-outcome association (Cerin et al., 2018; Muller et al., 2005). Here, exposure-372 outcome associations mediated by physical activity and/or sedentary behaviours are referred to as 373 indirect statistical effects or associations, while those not mediated by these behaviours represent 374 direct statistical effects or associations. Estimates of direct associations between exposures and 375 outcomes were derived from regression models (separate models for each outcome) adjusted for all 376 mediators (and confounders), which, in this case, were measures of physical activity and sedentary 377 behaviours. Indirect associations were inferred from regression models estimating exposure-mediator 378 and mediator-mediator (here, physical activity-sedentary behaviour) associations (separate models for 379 each mediator), and regression models estimating exposure-adjusted mediator-outcome associations 380 (separate models for each outcome). It is important to note that, although appropriate for the type of data examined in this study, unlike the product-of-coefficient, structural equation models and the 381 382 counterfactual framework (Dzhambov et al., 2020), the joint-significant test does not explicitly 383 quantify the indirect effects of exposures on the outcome, i.e., it does not provide a point estimate and standard error of the effect of an exposure on the outcome via a specific mediator or series of 384 mediators (Cerin, 2010). This is a limitation. However, similarly to the product-of-coefficient test and 385 386 structural equation models (Dzhambov et al., 2020), it can determine parallel and serial mediation

processes and does not require significant total environment-outcome associations to establishmediation (Cerin, 2010).

389

## **390 3. Results**

Table 1 shows the characteristics of the analytic sample. Participants were mainly middle-391 392 aged or older adults (82% aged 50+ years) of English-speaking background. The sample was diverse 393 in household income, educational attainment and neighbourhood environmental characteristics. For example, the ranges of the four environmental indices were -5.1 to 21.1 for the walkability index, -1.1 394 395 to 20.3 for the natural environment index, -5.0 to 7.6 for the air pollution index, and 0 to 10 for 396 IRSAD (distributions reported in Figure S2, supplementary data). The associations between the 397 environmental indices are reported in the supplementary file (Table S2 and Figure S3) and descriptive 398 statistics for environmental attributes based on 500 m and 1.6 km street-network buffers can be found 399 in the supplementary data (Table S6). 400 Only a small percentage of the sample reported taking medications for diabetes (6.3%), while

401 nearly a quarter and a third of the sample were on lipid-lowering and antihypertensive medications, 402 respectively. Walking for recreation was the most and resistance training the least prevalent form of 403 physical activity. On average, participants reported 0.8 h/day of sitting for transport, with 22.8% 404 accumulating  $\geq$ 1 h/day. They also reported an average of 2.6 h/day leisure-time sitting (range: 0-15 405 h/day). The average scores on the residential self-selection scales were around 3.0, indicating that, on 406 average, access to destinations and recreational facilities were "somewhat important" reasons for 407 choosing to live in the current neighbourhood.

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## **409** Table 1. Analytic sample characteristics (N = 4, 141).

Characteristics	Statistics	Characteristics	Statistics
Individual-level socio-demographic	characteristics		
Age, years, $M \pm SD$	$61.1\pm11.4$	Sex, female, %	55.2
Educational attainment, %		Employment status, %	
Up to secondary	32.7	Not employed	30.4
Trade, technician certificate	29.1	Paid employment	52.2
Associate diploma & equiv.	14.5	Volunteering	15.1
Bachelor degree, post-graduate	23.1	Missing data	2.3
diploma			
Missing data	0.6	English-speaking background, %	89.9
Living arrangements, %		Household income, annual, %	
Couple without children	48.2	Up to \$49,999	32.9
Couple with children	26.8	\$50,000 - \$99,999	26.8
Other	22.4	\$100,000 and over	28.9
Missing data	2.4	Does not know or refusal	8.8
-		Missing data	2.7
Residential self-selection – access to destinations, $M \pm SD$	$3.0\pm1.4$	Residential self-selection – recreational facilities, $M \pm SD$	3.1 ± 1.5
Missing data, %	7.8	Missing data, %	7.8

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Population density, persons/ha	$17.4 \pm 10.0$	adius street-network buffers), $M \pm SI$ Street intersection density,	$62.2 \pm 32.2$
		intersections/km <sup>2</sup>	
Percentage of commercial land in	$2.5\pm6.1$	Non-commercial land use mix,	$0.14 \pm 0.11$
residential buffer		entropy score (0 to 1)	
Percentage of parkland in	$11.6 \pm 12.5$	Percentage of blue space	$0.24 \pm 1.9$
residential buffer		(waterbody) in residential buffer	
NO <sub>2</sub> , ppb	$5.5 \pm 2.1$	$PM_{2.5}, \mu g/m^3$	$6.3 \pm 1.7$
NO <sub>2</sub> , $\mu g/m^3$	$10.4\pm4.0$		
Area-level IRSAD, in deciles	$6.4 \pm 2.7$	Walkability index, sum of z- scores	$0.0 \pm 2.5$
Natural environment index, sum	$0.0 \pm 1.4$	Ambient air pollution index,	$0.0\pm1.6$
of z-scores		sum of z-scores	
Physical activity and sedentary beha	viours		
Walking for transport		Walking for recreation	
Times per week, $M \pm SD$	$1.4 \pm 3.5$	Times per week, $M \pm SD$	$2.4 \pm 2.5$
Prevalence, %	29.1	Prevalence, %	61.6
Missing data, %	2.7	Missing data, %	3.0
Vigorous gardening		Resistance training	
Times per week, $M \pm SD$	$0.8 \pm 1.5$	Times per week, $M \pm SD$	$0.9\pm2.3$
Prevalence, %	37.1	Prevalence, %	25.5
Missing data, %	2.6	Missing data, %	2.6
Sitting for transport, h/day, $M \pm SD$	$0.8 \pm 0.8$	Leisure-time sitting, h/day, $M \pm SD$	$2.6 \pm 1.6$
Missing data, %	2.7	Missing data, %	2.8
Cardiometabolic health indicators (c			
Mean arterial blood pressure	$92.0\pm12.3$	Glycated haemoglobin	$39.9\pm6.3$
(MAP), mmHg, $M\pm$ SD		(HbA1C), mmol/mol, $M\pm SD$	
		(HbA1C), %, M±SD	$5.8\pm2.7$
Missing data, %	0.2	Missing data, %	0.5
Waist circumference, cm, $M \pm SD$	$94.6 \pm 14.2$	LDL cholesterol, mmol/L, $M \pm SD$	$3.0 \pm 0.9$
Missing data, %	0.2	Missing data, %	1.4
HDL cholesterol, mmol/L, $M \pm SD$	$1.5 \pm 0.4$	Triglycerides, mmol/L, M±SD	$1.3 \pm 0.9$
Missing data, %	0.3	Missing data, %	0.3
Other health-related variables, %			
Diabetes medication	6.3	Tobacco-smoking status	
Missing data	1.8	Current smoker	7.0
Anti-hypertensive medication	32.0	Previous smoker	35.9
Missing data	1.8	Non-smoker	54.5
Lipid-lowering medication	24.5	Missing data	2.6
Missing data	1.8	Heart problems/stroke history	8.7
		Missing data	1.0

411 Abbreviations: M, mean; SD, standard deviation; IRSAD, Index of Relative Socioeconomic

412 Advantage and Disadvantage; NO<sub>2</sub>, nitrogen dioxide;  $PM_{2.5}$ , particulate matter < 2.5  $\mu$ m; ppb, parts

*3.1. Associations of neighbourhood physical environment attributes with cardiometabolic health* 

*indicators and moderating effects of neighbourhood SES and air pollution* 

<sup>413</sup> per billion; LDL, low-density lipoprotein; HDL, high-density lipoprotein.

- On average, neighbourhood walkability was positively related to MAP (Table 2; Table S5 in 417 the supplementary data for estimates unadjusted for ambient air pollution). However, this association 418 depended on both neighbourhood IRSAD (SES) and ambient air pollution (walkability by IRSAD by 419 air pollution index 3-way interaction: b = 0.045; 95% CI: 0.013, 0.077; p = .007). It was significant in 420 high-SES neighbourhoods, irrespective of air pollution, as well as in low- and average-SES 421 neighbourhoods with low or average ambient air pollution (Table 3). Overall, the associations were 422 423 stronger in low-SES and low-air pollution neighbourhoods (Table 3). Similar moderation effects of 424 neighbourhood IRSAD and air pollution were observed on the associations of walkability with LDL cholesterol (b = 0.003; 95% CI: 0.001, 0.005; p = .012) and triglycerides ( $e^b = 1.002$ ; 95% CI: 1.0004, 425 1.003; p = .014). Positive relationships were found only in low-to-medium SES and low air pollution 426 427 neighbourhoods (Table 3). The association of walkability with HDL cholesterol was also moderated by air pollution ( $e^b = 1.002$ ; 95% CI: 1.0002, 1.004; p = .030) (Table 3). It was positive only at nearly 428 maximum values of the air pollution index ( $e^b = 1.014$ ; 95% CI: 1.001, 1.028; p=.038). The natural 429 430 environment index was negatively related to waist circumference and LDL cholesterol (Table 2; Table S5 for estimates unadjusted for ambient air pollution). Also, it was negatively associated with HDL 431 cholesterol in areas with high air pollution (Table 3) (natural environment index by air pollution index 432 2-way interaction:  $e^b = 0.994$ ; 95% CI: 0.990, 0.998; p = .008). 433
- 434 Neighbourhood IRSAD was negatively related to waist circumference, MAP and triglycerides, and positively related to glycated haemoglobin and HDL cholesterol (Table 2; Table S5 for estimates 435 unadjusted for ambient air pollution). Finally, the air pollution index was negatively associated with 436 437 MAP and HDL cholesterol, while a positive association was observed with glycated haemoglobin (Table 2). Our data did not provide sufficient evidence of moderating effects of medication intake on 438 439 any of the above associations (Table S4, supplementary data). Associations of exposures based on 500 440 m and 1.6 km radii street-network buffers and cardiometabolic health indicators are reported in the 441 supplementary data (Tables S7-S10b). Overall, they were similar to those based on 1 km radius 442 buffers. The only notable differences were 500 m buffers yielding weaker, non-significant moderating 443 effects of ambient air pollution on the associations between the natural environment index and HDL cholesterol (p=.440), and 1.6 km buffers producing significantly stronger negative associations 444 between the natural environment index and MAP (Tables S7 and S9). 445
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452 Table 2. Associations of neighbourhood environment attributes with cardiometabolic health indicators: main effect models (unadjusted for physical activity

453 and sedentary behaviours).

Neighbourhood environment attributes	Waist circumference (cm)	Mean arterial pressure (mmHg)	Glycated haemoglobin (mmol/mol)	HDL cholesterol (mmol/L)	LDL cholesterol (mmol/L)	Triglycerides (mmol/L)
	<i>b</i> (95% CI)	<i>b</i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)	<i>b</i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)
Walkability index	-0.094	0.387	1.001	1.002	0.001	1.005
	(-0.295, 0.107)	(0.174, 0.601)	(0.999, 1.004)	(0.997, 1.007)	(-0.012, 0.014)	(0.996, 1.014)
Natural environment index	-0.476	-0.169	0.999	1.001	-0.022	1.000
	(-0.812, -0.140)	(-0.477, 0.138)	(0.996, 1.002)	(0.994, 1.007)	(-0.041, -0.003)	(0.987, 1.014)
Neighbourhood IRSAD	-0.388	-0.281	1.002	1.009	0.011	0.990
	(-0.581, -0.194)	(-0.463, -0.098)	(1.001, 1.004)	(1.005, 1.013)	(-0.0002, 0.022)	(0.983, 0.997)
Ambient air pollution index	0.335	-0.537	1.004	0.990	0.002	1.005
	(-0.046, 0.717)	(-0.877, -0.197)	(1.000, 1.008)	(0.983, 0.998)	(-0.019, 0.022)	(0.991, 1.018)

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455 Abbreviations: IRSAD, Index of Relative Advantage and Disadvantage; b, unstandardised regression coefficient from model with Gaussian variance and 456 identity link functions;  $e^b$  = exponentiated regression coefficient from models with Gamma variance and logarithmic link functions; CI, confidence intervals. 457 Estimates in bold are significant at a 0.05 two-tailed probability level. All regression coefficients are adjusted for other environmental indices, age, sex,

English-speaking background, educational attainment, household income, living arrangements, work status, neighbourhood self-selection and taking
 medications relevant to a specific outcome (diabetes, hypertension and/or dyslipidaemia).

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Table 3. Associations of neighbourhood physical environment attributes with cardiometabolic health indicators: moderating effects of neighbourhood socio-467

economic status and air quality (unadjusted for physical activity and sedentary behaviours). 468

Moderator	Moderator values	Mean arterial pressure (mmHg)	LDL cholesterol (mmol/L)	Triglycerides (mmol/L)	Moderator	Moderator values	HDL cholesterol (mmol/L)
		<i>b</i> (95% CI)	<i>b</i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)			<i>e<sup>b</sup></i> (95% CI)
Exposure: Walkability index					Exposure: Walkability	, index	
Neighbourhood IRSAD Air pollution index		0.859 (0.473, 1.246)	0.027 (0.002, 0.052)	1.024 (1.007, 1.042)	Air pollution index	M – 1 SD	0.996 (0.990, 1.001)
Neighbourhood IRSAD Air pollution index		0.441 (0.130, 0.753)	0.006 (-0.014, 0.026)	1.007 (0.994, 1.021)	Air pollution index	М	0.999 (0.994, 1.004)
Neighbourhood IRSAD Air pollution index		0.024 (-0.348, 0.395)	-0.016 (-0.040, 0.009)	0.991 (0.974, 1.007)	Air pollution index	M + 1 SD	1.002 (0.997, 1.007)
Neighbourhood IRSAD Air pollution index		0.665 (0.373, 0.957)	0.012 (-0.006, 0.031)	1.014 (1.002, 1.028)	Exposure: Natural en	vironment index	
Neighbourhood IRSAD Air pollution index		0.439 (0.210, 0.669)	0.003 (-0.011, 0.017)	1.006 (0.996, 1.015)	Air pollution index	M – 1 SD	1.004 (0.996, 1.011)
Neighbourhood IRSAD Air pollution index		0.214 (-0.025, 0.453)	-0.007 (-0.022, 0.009)	0.997 (0.987, 1.008)	Air pollution index	М	0.996 (0.988, 1.003)
Neighbourhood IRSAD Air pollution index		0.470 (0.056, 0.884)	-0.002 (-0.029, 0.024)	1.005 (0.987, 1.024)	Air pollution index	M + 1 SD	0.988 (0.976, 0.999)
Neighbourhood IRSAD Air pollution index		0.437 (0.118, 0.756)	-0.001 (-0.020, 0.020)	1.005 (0.991, 1.018)			
Neighbourhood IRSAD Air pollution index		0.404 (0.105, 0.703)	0.002 (-0.017, 0.021)	1.004 (0.991, 1.017)	- CC <sup></sup>		

Abbreviations: IRSAD, Index of Relative Advantage and Disadvantage; b, unstandardised regression coefficient from model with Gaussian variance and 469 identity link functions;  $e^b$  = exponentiated regression coefficient from models with Gamma variance and logarithmic link functions;CI, confidence intervals; 470 M, mean; SD, standard deviation. Estimates in bold are significant at a 0.05 two-tailed probability level. All regression coefficients are adjusted for other 471 environmental indices, age, sex, English-speaking background, educational attainment, household income, living arrangements, work status, neighbourhood 472

self-selection and taking medications relevant to a specific outcome (diabetes, hypertension and/or dyslipidaemia). 473

# 474 3.2. Physical activity and sedentary behaviours as mediators of the associations of neighbourhood 475 physical environment attributes with cardiometabolic health indicators

476 Figures 2-6 show the indirect (behaviour-mediated) and direct (not mediated by physical 477 activity or sedentary behaviours) associations of neighbourhood walkability and natural environment 478 with five cardiometabolic health indicators. Significant indirect associations were found between 479 neighbourhood walkability and all five indicators (Figures 2-6), while direct associations emerged 480 only with respect to MAP (Tables 4 and 5) and triglycerides (Figure 5; Tables 4 and 5). These direct 481 associations were respectively moderated by both IRSAD and the air pollution index, and suggestive 482 of potential detrimental effects. Specifically, positive associations between walkability and MAP were 483 observed in neighbourhoods with above average IRSAD irrespective of the level of air pollution, and 484 in neighbourhoods with below average and average IRSAD that had below average and average levels of air pollution (Table 5). Walkability was also directly positively related to triglycerides, but only in 485 486 neighbourhoods with below average or average IRSAD and low air pollution (Table 5).

487 The indirect associations of the walkability index with waist circumference, glycated 488 haemoglobin and HDL cholesterol were mainly suggestive of potential beneficial effects via higher 489 odds of engagement in walking for transport, walking for recreation and resistance training, and less 490 sitting for transport and leisure-time sitting (Figures 2-4). A similar pattern of indirect associations 491 was also observed for triglycerides, with the exception of sitting for transport, which was unrelated to 492 this particular cardiometabolic health indicator (Figure 5). All indirect associations via engagement in 493 walking for recreation appeared to be channelled through (lower) leisure-time sitting, while this did 494 not hold for the indirect associations via engagement in walking for transport and/or resistance training in the case of waist circumference (Figure 2), HDL cholesterol (Figure 4) and triglycerides (Figure 5). 495 496 The indirect associations via engagement in resistance training were moderated by air pollution, 497 whereby neighbourhood walkability was positively associated with resistance training only at above 498 average values of the air pollution index (Figures 2-5). The only behavioural pathways through which 499 walkability appeared to have detrimental effects on the examined cardiometabolic health indicators 500 were those through frequency of engagement in vigorous gardening (Figures 2-5) and walking for 501 transport (Figure 6) in those engaging in these physical activities. In fact, walkability was predictive of 502 higher frequency of engagement in walking for transport and the latter was positively related to LDL 503 cholesterol (Figure 6). Walkability was also negatively related to frequency of vigorous gardening in 504 those living in areas with below average or average air pollution, while a higher frequency of vigorous gardening was associated with less sitting for leisure (Figures 2-5). 505

The natural environment index was negatively associated with waist circumference directly and via engagement in resistance training and leisure-time sitting (Figure 2; Table 4). Specifically, this environmental index was positively associated with engagement in resistance training and negatively associated with leisure-time sitting, which were, in turn, directly negatively and positively related to waist circumference, respectively. Resistance training was also negatively associated with waist 511 circumference via leisure-time sitting. Similar indirect but not direct associations were found for

- triglycerides (Figure 5; Table 4). Indirect but not direct associations also emerged between the natural
  environment index and glycated haemoglobin (Figure 3; Table 4). However, engagement in resistance
- training was negatively related to this cardiometabolic health indicator only indirectly, via leisure-timesitting (Figure 3).

516 The indirect associations between the natural environment and HDL cholesterol mirrored 517 those found for waist circumference but were, as expected, in the opposite direction because engagement in resistance training was positively, and leisure-time sitting negatively, associated with 518 HDL cholesterol (Figure 4). However, unlike for waist circumference, the natural environment index 519 520 displayed detrimental direct associations with HDL cholesterol, albeit only for those living in 521 neighbourhoods with above average levels of air pollution (Figure 4; Table 5). No significant indirect 522 associations of the natural environment index with MAP or LDL cholesterol were observed (Table 4). A direct negative association was found with LDL cholesterol (Figure 6; Table 4). 523

524 The mediating effects of physical activity and sedentary behaviours in the associations 525 between cardiometabolic health indicators and the neighbourhood environment indices based on the other two street-network buffers (with 500 m and 1.6 km radii) are reported in the supplementary data 526 (Figures S5a-S9b and Tables 11a-11b). In general, the patterns of associations were similar to those 527 observed for 1 km radius buffers, especially in relation to LDL cholesterol (Figures S9a and S9b) and 528 triglycerides (Figure S8a). For the other cardiometabolic health indicators, mediation analyses using 529 530 500 m and 1.6 km radii buffers yielded fewer mediated effects of physical activity and/or sedentary 531 behaviours. Specifically, when using 500 m radius buffers, sitting for transport was no longer a significant mediator of the associations of neighbourhood walkability with waist circumference, 532 533 glycated haemoglobin and HDL cholesterol (Figures S5a, S6a and S7a), and engagement in resistance 534 training was not a mediator of the associations between walkability, the natural environment index and 535 glycated haemoglobin (Figure S6a). When using 1.6 km radius buffers, engagement in walking for 536 recreation was no longer a mediator of the associations between walkability and waist circumference, 537 glycated haemoglobin, HDL cholesterol and triglycerides (Figures S5b, S6b, S7b and S8b). Also, the 538 associations between the natural environment and all these cardiometabolic health indicators were no longer directly mediated by leisure-time sitting. However, additional negative indirect effects of the 539 540 natural environment index through engagement in walking for transport were observed (Figures S5b, 541 S6b, S7b and S8b).

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Table 4. Associations of neighbourhood environment attributes with cardiometabolic health indicators: direct main effects models (adjusted for physical
 activity and sedentary behaviours).

Neighbourhood environment attributes	Waist circumference (cm)	Mean arterial pressure (mmHg)	Glycated haemoglobin (mmol/mol)	HDL cholesterol (mmol/L)	LDL cholesterol (mmol/L)	Triglycerides (mmol/L)
	<i>b</i> (95% CI)	<i>b</i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)	<i>b</i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)
Walkability index	-0.081	0.435	1.002	0.999	0.001	1.007
	(-0.320, 0.159)	(0.203, 0.666)	(0.9996, 1.004)	(0.995, 1.004)	(-0.015, 0.013)	(0.997, 1.017)
Natural environment index	-0.403	-0.204	0.999	0.994	-0.023	1.002
	(-0.749, -0.057)	(-0.518, 0.111)	(0.996, 1.002)	(0.987, 1.002)	(-0.043, -0.002)	(0.989, 1.017)
Neighbourhood	-0.406	-0.307	1.002	1.009	0.007	0.987
IRSAD	(-0.607, -0.206)	(-0.499, -0.115)	(1.000, 1.004)	(1.005, 1.013)	(-0.004, 0.019)	(0.979, 0.995)
Ambient air pollution index	0.370	-0.580	1.004	0.992	0.002	1.008
	(-0.002, 0.742)	(-0.930, -0.231)	(1.001, 1.008)	(0.985, 0.999)	(-0.020, 0.023)	(0.994, 1.022)

545

546 Abbreviations: IRSAD, Index of Relative Advantage and Disadvantage; *b*, unstandardised regression coefficient from model with Gaussian variance and

identity link functions;  $e^b$  = exponentiated regression coefficient from models with Gamma variance and logarithmic link functions; CI, confidence intervals.

548 Estimates in bold are significant at a 0.05 two-tailed probability level. All regression coefficients are adjusted for other environmental indices, age, sex,

549 English-speaking background, educational attainment, household income, living arrangements, employment status, work status, neighbourhood self-selection,

taking medications relevant to a specific outcome (diabetes, hypertension and/or dyslipidaemia) and physical activity and sedentary behaviour

551 Table 5. Associations of neighbourhood physical environment attributes with cardiometabolic health indicators: direct moderating effects of neighbourhood

552 socio-economic status and air quality (adjusted for physical activity and sedentary behaviours).

Moderator	Moderator values	Mean arterial pressure (mmHg)	Triglycerides (mmol/L)	Moderator	Moderator values	HDL cholesterol (mmol/L)
		<i>b</i> (95% CI)	<i>e<sup>b</sup></i> (95% CI)			<i>e<sup>b</sup></i> (95% CI)
Exposure: Walkability index			· · ·	Exposure: Natural environment i	ndex	· · · ·
Neighbourhood IRSAD Air pollution index		0.845 (0.459, 1.231)	1.025 (1.008, 1.042)	Air pollution index	M-1  SD	1.004 (0.996, 1.011)
Neighbourhood IRSAD Air pollution index		0.436 (0.125, 0.747)	1.008 (0.995, 1.022)	Air pollution index	М	0.994 (0.987, 1.001)
Neighbourhood IRSAD Air pollution index		0.027 (-0.343, 0.398)	0.992 (0.976, 1.009)	Air pollution index	M + 1 SD	0.985 (0.973, 0.996)
Neighbourhood IRSAD Air pollution index		0.655 (0.362, 0.947)	1.015 (1.002, 1.028)			
Neighbourhood IRSAD Air pollution index		0.435 (0.204, 0.665)	1.007 (0.997, 1.017)			
Neighbourhood IRSAD Air pollution index		0.215 (-0.025, 0.454)	0.999 (0.989, 1.010)			
Neighbourhood IRSAD Air pollution index		0.464 (0.050, 0.877)	1.004 (0.987, 1.022)			
Neighbourhood IRSAD Air pollution index		0.433 (0.114, 0.752)	1.006 (0.992, 1.019)			
Neighbourhood IRSAD Air pollution index		0.402 (0.103, 0.701)	1.007 (0.994, 1.020)			

Abbreviations: IRSAD, Index of Relative Advantage and Disadvantage; *b*, unstandardised regression coefficient from model with Gaussian variance and

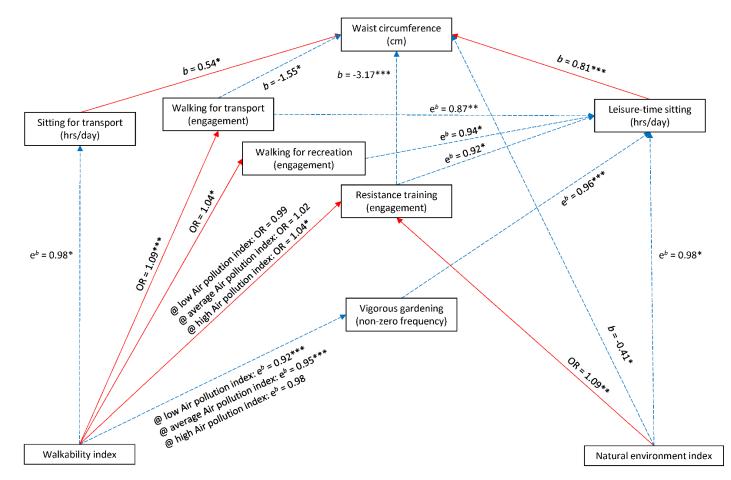
identity link functions;  $e^b$  = exponentiated regression coefficient from models with Gamma variance and logarithmic link functions; CI, confidence intervals;

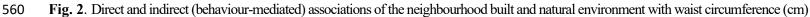
555 M, mean; SD, standard deviation. Estimates in bold are significant at a 0.05 two-tailed probability level. All regression coefficients are adjusted for other

environmental indices, age, sex, English-speaking background, educational attainment, household income, living arrangements, employment status, work

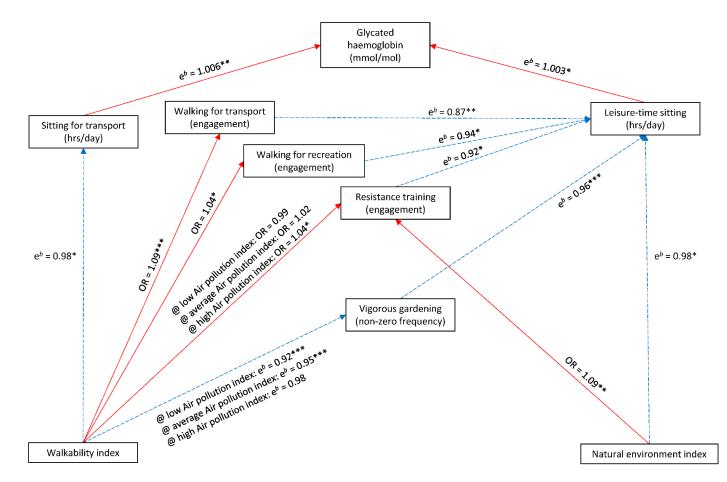
557 status, neighbourhood self-selection, taking medications relevant to a specific outcome (diabetes, hypertension and/or dyslipidaemia) and physical activity and

558 sedentary behaviours.





Arrows linking variables indicate significant associations. Red full arrows denote positive associations, while blue dashed arrows denote negative associations. (a) denotes an association moderated by the Air pollution index and estimates of the association are given at different values of the Air pollution index (low = 1 standard deviation below the mean; average = mean; high = 1 standard deviation above the mean). OR = odds ratio from models with binomial variance and logit link functions (engagement in walking for different purposes and resistance training); *b* = regression coefficient from models with Gaussian variance and identify link functions (waist circumference);  $e^b$  = exponentiated regression coefficient from models with Gauma variance and logarithmic link functions (sitting for different purposes and non-zero frequency of vigorous gardening). \* p<.05; \*\* p<.01; \*\*\*p< .001. Regression coefficients and their 95% confidence intervals are presented in Table 4 and Table S2 (supplementary data). Estimates of path coefficients were obtained using a set of regression models (one for each mediator and cardiometabolic health indicator) rather than simultaneously.



569 Fig. 3. Indirect (behaviour-mediated) associations of the neighbourhood built and natural environment with glycated haemoglobin (mmol/mol)

570 Arrows linking variables indicate significant associations. Red full arrows denote positive associations, while blue dashed arrows denote negative associations. @ denotes an

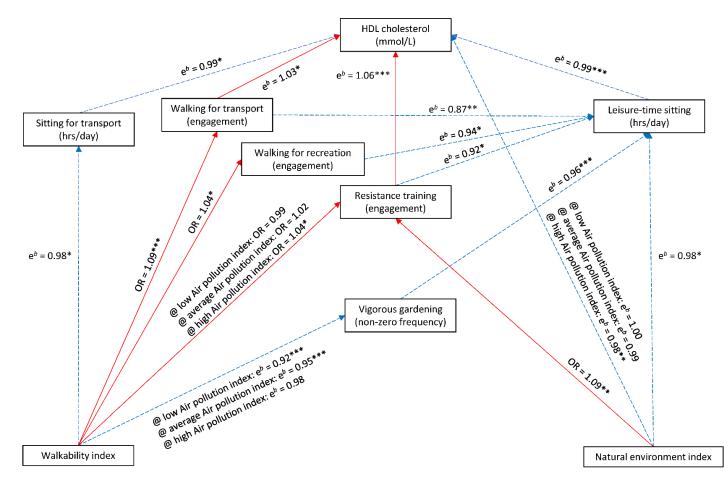
association moderated by the Air pollution index and estimates of the association are given at different values of the Air pollution index (low = 1 standard deviation below the mean;

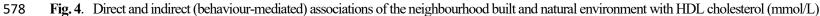
572 average = mean; high = 1 standard deviation above the mean). OR = odds ratio from models with binomial variance and logit link functions (engagement in walking for different 573 purposes and resistance training);  $e^b$  = exponentiated regression coefficient from models with Gamma variance and logarithmic link functions (sitting for different purposes, non-zero

574 frequency of vigorous gardening and glycated haemoglobin). \* p<.05; \*\* p<.01; \*\*\*p<.001. Regression coefficients and their 95% confidence intervals are presented in Table 4 and

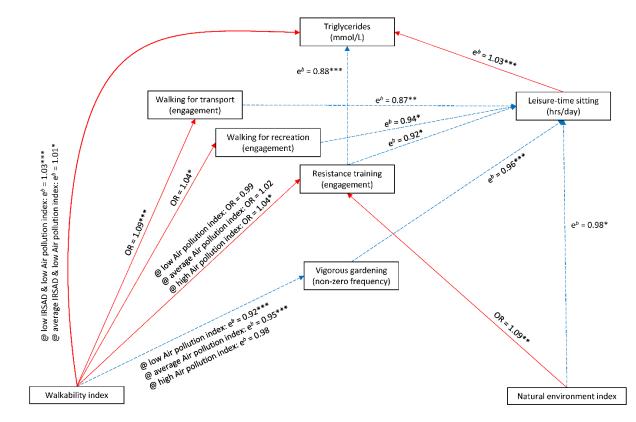
575 Table S2 (supplementary data). Estimates of path coefficients were obtained using a set of regression models (one for each mediator and cardiometabolic health indicator) rather than

576 simultaneously.





- 579 Arrows linking variables indicate significant associations. Red full arrows denote positive associations, while blue dashed arrows denote negative associations. @ denotes an association moderated by the Air pollution index and estimates of the association are given at different values of the Air pollution index (low = 1 standard deviation below the
- association moderated by the Air pollution index and estimates of the association are given at different values of the Air pollution index (low = 1 standard deviation below the mean; average = mean; high = 1 standard deviation above the mean). OR = odds ratio from models with binomial variance and logit link functions (engagement in walking for
- different purposes and resistance training);  $e^b =$  exponentiated regression coefficient from models with Gamma variance and logarithmic link functions (sitting for different
- purposes, non-zero frequency of vigorous gardening and HDL cholesterol). \* p<.05; \*\* p<.01; \*\*\*p<.001. Regression coefficients and their 95% confidence intervals are
- 584 presented in Tables 4, 5 and S2 (supplementary data). Estimates of path coefficients were obtained using a set of regression models (one for each mediator and cardiometabolic
- 585 health indicator) rather than simultaneously.

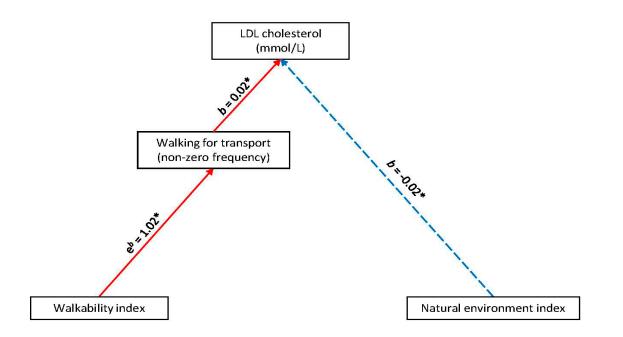


588 Fig. 5. Direct and indirect (behaviour-mediated) associations of the neighbourhood built and natural environment with triglycerides (mmol/L)

589 Arrows linking variables indicate significant associations. Red full arrows denote positive associations, while blue dashed arrows denote negative associations. @ denote

590 associations moderated by the Air pollution index and, where applicable, the Index of Relative Social Advantage and Disadvantage (IRSAD). For associations moderated by Air

- 591 pollution index only, estimates of the association are given at different values of the Air pollution index (low = 1 standard deviation below the mean; average = mean; high = 1 592 standard deviation above the mean). For associations moderated by both IRSAD and Air pollution index, only statistically significant estimates of associations are given for
- specific values of the two moderators (low = 1 standard deviation below the mean; average = mean). OR = odds ratio from models with binomial variance and logit link functions
- (engagement in walking for different purposes and resistance training);  $e^b = exponentiated$  regression coefficient from models with Gamma variance and logarithmic link
- 595 functions (leisure-time sitting, non-zero frequency of vigorous gardening and triglycerides). \* p < .05; \*\* p < .01; \*\*\*p < .001. Regression coefficients and their 95% confidence
- 596 intervals are presented in Tables 4, 5 and S2 (supplementary data). Estimates of path coefficients were obtained using a set of regression models (one for each mediator and cardiometabolic health indicator) rather than simultaneously.



599

- 600 Fig. 6. Direct and indirect (behaviour-mediated) associations of the neighbourhood built and natural environment with LDL cholesterol (mmol/L)
- 601 Arrows linking variables indicate significant associations. Red full arrows denote positive associations, while blue dashed arrows denote negative associations.
- $b = regression \ coefficient \ from \ model \ with \ Gaussian \ variance \ and \ identity \ link \ functions \ (LDL \ cholesterol); e<sup>b</sup> = exponentiated \ regression \ coefficient \ from \ regression \ coefficient \ regression \ coefficient \ from \ regression \ regression \ coefficient \ from \ regression \ coefficient \ from \ regression \ regression$
- model with Gamma variance and logarithmic link functions (non-zero frequency of walking for transport); \* p<.05. Regression coefficients and their 95%
- 604 confidence intervals are presented in Table 4 and Table S2 (supplementary data). Estimates of path coefficients were obtained using a set of regression models (one for
- each mediator and cardiometabolic health indicator) rather than simultaneously.

## 606 **4. Discussion**

607 We examined how neighbourhood built and natural environment characteristics are associated 608 with cardiometabolic risk factors for CVD in Australian mid-aged and older community dwellers, and 609 the extent to which these associations are moderated by ambient air pollution and neighbourhood SES 610 and mediated by physical activity and sedentary behaviours. This study revealed a substantial number 611 of associations of neighbourhood walkability and natural environment with behavioural and 612 cardiometabolic risk factors for CVD that were in the expected direction, but also a few counterintuitive findings especially with respect to neighbourhood walkability. As the associations 613 614 between neighbourhood environment attributes based on residential buffers of different sizes and 615 cardiometabolic health indicators were similar and the mediation effects of physical activity and 616 sedentary behaviours were, as expected (Gunn et al., 2017), stronger when neighbourhood was defined 617 as an area within 1 km from home, our discussion focusses on the findings of exposures based on 1 km 618 radius buffers.

619

# 620 *4.1. Neighbourhood walkability*

621 Higher walkability was related to less sitting for transport which, in turn, showed beneficial associations with several cardiometabolic health indicators, including waist circumference, glycated 622 623 haemoglobin and HDL cholesterol. Higher walkability was also predictive of higher odds of engagement in walking for transport, walking for recreation and resistance training - behaviours that 624 625 were directly and/or indirectly associated, via leisure-time sitting, with better cardiometabolic health 626 (smaller waist circumference, higher HDL cholesterol and lower glycated haemoglobin and triglycerides). More walkable neighbourhoods appear to have a potential beneficial effect on adiposity, 627 628 blood glucose and blood lipids by encouraging an active lifestyle typified by higher levels of transport 629 and leisure-time physical activity and, through these, lower levels of transport and leisure-time sitting. 630 In general, these findings are in line with those of previous studies on environmental correlates of 631 physical activity (Cerin et al., 2017; Van Cauwenberg et al., 2018) and sedentary behaviours (Barnett 632 et al., 2015; Cerin et al., 2020), and those on the effects of these behaviours on cardiometabolic health 633 (Bai et al., 2022; Ballard et al., 2021; Chai et al., 2023; Wood et al., 2022).

A few associations between neighbourhood walkability and risk factors for CVD were counterintuitive, especially with respect to blood pressure (MAP), which was higher in more walkable neighbourhoods. The concept of neighbourhood walkability was coined by urban planners to denote urban spaces with higher levels of density, functional mix and access networks that lend themselves to

- 638 a variety of transport modes and reduce car-dependence (Frank et al., 2010). As expected, and
- evidenced by this study, more walkable neighbourhoods typically encourage active modes of transport
- 640 (e.g., walking for transport) (Cerin et al., 2017) and engagement in leisure-time physical activity (Van
- 641 Cauwenberg et al., 2018), which are beneficial to cardiometabolic health (Bai et al., 2022; Ballard et
- al., 2021; Lee et al., 2021). However, they are also accompanied by higher levels of air pollution

(James et al., 2015) and noise (Salter et al., 2015) that can be detrimental to health (Basner et al., 643 2014; Gaio et al., 2019; Liu et al., 2019; Salter et al., 2015; Wang et al., 2023). Although this study 644 645 partially accounted for ambient air pollution, it did not account for urban noise, and this may explain 646 the positive associations of neighbourhood walkability with MAP observed even after adjustment for 647 antihypertensive medication. In this regard, a large study conducted in Chicago reported that 10-dBA 648 higher residential noise levels corresponded to over 1 mmHg greater systolic and diastolic blood 649 pressure, as well as 20% higher odds of treatment-resistant hypertension (D'Souza et al., 2021). Sleep 650 disruption, oxidative stress and changes in sympathetic tone triggered by affective reactions associated with exposure to noise are thought to be the main mechanisms responsible for these findings (Basner 651 652 & McGuire, 2018; Münzel et al., 2018). Problems arising from urban noise could be mitigated through 653 appropriate urban and traffic planning policies (e.g., pedestrian zones, land use planning) and 654 technological interventions (e.g., installation of double-glazed windows and road resurfacing) (Salter 655 et al., 2015). Clearly, to understand the potential impact of dense, destination-rich neighbourhoods on 656 various cardiometabolic risk factors for CVD, it is important to consider aspects of urban design as 657 well as noise and air pollution exposures.

658 Another seemingly counterintuitive set of findings about the relationship between neighbourhood walkability and risk factors for CVD pertains to its interaction with ambient air 659 pollution. Higher walkability was more strongly positively related to MAP, LDL cholesterol and 660 triglycerides at lower levels of air pollution in more disadvantaged neighbourhoods. It was also more 661 662 strongly negatively related to frequency of gardening at lower levels of air pollution and more strongly 663 positively related to engagement in resistance training at higher levels of air pollution. The annual concentrations of air pollutants in the present study were generally low, i.e., less than half those 664 665 observed in Europe or the U.S (Clifford et al., 2016). Within such context, it is possible that they did 666 not significantly impact on cardiometabolic health (D'Oliveira et al., 2023) but rather acted as proxies 667 for type and quality of neighbourhood destinations that could not be accurately captured by the coarse 668 land use measures employed in the study. For example, we used percentage of commercial land and 669 non-commercial land use mix as measures of access to services incorporated in the neighbourhood 670 walkability index. However, the same percentages of commercial land might have represented popular retail and food outlets encouraging an active lifestyle and attracting visitors from other areas, or 671 672 unfrequently-visited office spaces and warehouses unsupportive of an active lifestyle and of limited 673 interest to residents from other areas. Ambient air pollution may have helped differentiate between the 674 two types of destinations. This would explain a stronger positive association between higher walkability and engagement in resistance training at higher levels of air pollution (Figures 2-5), as 675 676 well as the more beneficial or less detrimental potential effects of walkability on some of the 677 cardiometabolic risk factors for CVD (MAP, LDL cholesterol and HDL cholesterol) at higher levels of 678 air pollution (Table 3), which were attenuated after adjustment for physical activity and sedentary

679 behaviours (potential mediators) (Table 5).

The moderating effects of ambient air pollution on the associations of neighbourhood 680 walkability with MAP, LDL cholesterol and triglycerides depended on neighbourhood SES. They 681 682 were stronger in more disadvantaged neighbourhoods where higher walkability was predictive of 683 worse cardiometabolic health outcomes at lower levels of air pollution. Residents of more 684 disadvantaged neighbourhoods may be more dependent on their local environment as they may not 685 afford visiting destinations outside their neighbourhoods as frequently as their socially advantaged 686 counterparts (Holliday et al., 2017). If, in the context of the present study, air pollution was a proxy for 687 destination relevance and quality uncaptured by the walkability index, we would expect higher neighbourhood walkability to show more beneficial associations at higher levels of air pollution in 688 689 lower SES areas. Having good quality destinations of daily living in the local area, such as a variety of 690 food outlets and essential services, supports an active lifestyle (Cerin et al., 2017; Van Cauwenberg et 691 al., 2018) and a healthier diet (Cooksey-Stowers et al., 2017). Dense, lower SES neighbourhoods with 692 poor-quality destinations of daily living may have access to few low-cost, relatively unhealthy food 693 outlets (e.g., fast food outlets, convenience stores) with a limited choice of fresh produce (Williamson 694 et al., 2017) resulting in unhealthy dietary patterns (Mulrooney & Bell, 2016) and poorer cardiometabolic health (e.g., high LDL cholesterol and triglycerides) (Arnett et al., 2019). These 695 findings suggest that a fine-grained characterisation of the built environment differentiating types of 696 697 destinations that potentially impact cardiometabolic risk factors of CVD through key pathways (e.g., physical activity, diet) is necessary to gain a solid understanding of how neighbourhoods influence 698 CVD. 699

700 We did not find significant moderating effects of neighbourhood SES and air pollution on the 701 associations between behaviours and cardiometabolic risk factors, supporting the notion that being 702 more physically active and less sedentary is beneficial to the cardiometabolic health of mid-aged and 703 older residents living irrespective of area-level advantage and disadvantage. This is understandable 704 given that the effects of an active lifestyle on cardiometabolic health is mainly determined by 705 biological mechanisms (Pinckard et al., 2019) while those of the environment on cardiometabolic 706 health may be also explained by social and behavioural factors (Barnett et al., 2022; Chandrabose et 707 al., 2019; Rigolon et al., 2021). The lack of evidence of the moderating effect of air pollution on 708 behaviour-health associations may be attributed to the generally low annual concentrations of air 709 pollutants found in this study. In fact, a recent systematic review on the impact of air pollution on 710 older adults' health while engaging in physical activity and sedentary behaviours concluded that being 711 physically active in low-pollution environments provide health gains and reduce health risks, while 712 this is less the case if air pollution concentrations are high (D'Oliveira et al., 2023).

A couple of additional findings pertaining to the relationships between neighbourhood
walkability and CVD risk factors require consideration. In areas with low to medium air pollution,
higher walkability was negatively associated with frequency of vigorous gardening which, in turn, was
predictive of less sitting for leisure and, therefore, better cardiometabolic health. In general, dense,

717 walkable areas tend to have fewer and smaller private gardens. Hence, a negative association between higher walkability and gardening was expected. The fact that this effect was not significant in 718 719 neighbourhoods with higher levels of air pollution may be due to residents of low-walkable 720 neighbourhoods avoiding spending time outdoors in their gardens despite having them. This particular 721 finding suggests that air pollution mitigation strategies may be important to ensure older adults' 722 participation in outdoor forms of physical activity such as gardening. In the absence of private 723 gardens, residents of dense, walkable neighbourhoods may benefit from the establishment of urban 724 community gardens, which have been found to confer significant health benefits, including increases 725 in physical activity (Litt et al., 2023).

726 Finally, as expected, higher neighbourhood walkability was positively related to both 727 engagement and frequency of walking for transport. However, while engagement in walking for 728 transport showed beneficial associations with cardiometabolic health, frequency of walking for 729 transport was positively related to LDL cholesterol. Individuals may frequently engage in walking for 730 transport to visit cafés, restaurants or fast-food outlets. In fact, a travel survey conducted in Australia 731 reported that people walked for almost 67% of short trips to bars, pubs, cafés and restaurants (Eady & 732 Burtt, 2019). Eating out more frequently rather than preparing food at home is usually associated with 733 higher energy and fat intake (Lachat et al., 2012) and, consequently, worse cardiometabolic health 734 (Nago et al., 2014). Interventions and policies aimed at making healthy choices more available in 735 food-serving premises may mitigate this problem.

736

## 737 *4.2. Neighbourhood natural environment*

Access to green space is deemed to contribute to better cardiometabolic health by reducing residents' exposure to air pollution and noise, exerting a cooling effect on the environment, facilitating physiological stress recovery, encouraging participation in physical activity and promoting social cohesion (Markevych et al., 2017) which, in turn, can also contribute to stress mitigation (Robinette et al., 2018). Similarly, White and colleagues (2020) have postulated that access to blue spaces may mitigate urban heat, reduce stress, promote positive social relations and encourage physical activity, all of which are beneficial to cardiometabolic health.

In our study, the relationships of risk factors for CVD with the neighbourhood natural
environment were mostly in the expected direction. Higher scores on the natural environment index,
combining information on greenspace (parkland) and blue space, were directly negatively associated
with waist circumference and LDL cholesterol, and, when using 1.6 km radius buffers, also negatively

- associated with MAP. In addition, they were indirectly negatively associated with waist
- circumference, triglycerides and glycated haemoglobin via higher odds of engagement in resistance
- training and less leisure-time sitting, and indirectly positively associated with HDL cholesterol via the
- same behavioural pathways. The fact that many parks (Grigoletto et al., 2021) and beaches (Bliss,
- 2016) have outdoor fitness equipment and many personal trainers run classes in parks may explain the

association between the natural environment and resistance training. Residents spending more time

- outdoors exercising, socialising or relaxing if they have access to nature in the local area (Georgiou et
- al., 2021; Zhang et al., 2021) may be the reason for them engaging in less leisure-time sitting, which,
- in this demographic, is mainly represented by TV viewing (Compernolle et al., 2021). Displacing TV
- viewing with outdoor activities can benefit cardiometabolic health via at least three pathways higher
- rs9 energy expenditure from non-sedentary activities, less snacking and energy-dense food intake
- 760 (Pearson & Biddle, 2011) and higher levels of exposure to UV radiation (Gorman et al., 2017).
- 761 In general, our findings are consistent with those of previous studies reporting beneficial 762 effects of green and blue spaces on cardiometabolic health (Astell-Burt et al., 2020; Dendup et al., 763 2018; Rahimi-Ardabili et al., 2021), physical activity (Astell-Burt et al., 2014; Georgiou et al., 2021; 764 Van Cauwenberg et al., 2018) and sedentary behaviours (Barnett et al., 2015; Cerin et al., 2020). By supporting an active lifestyle, better access to nature in urban areas appears to lead to better 765 766 cardiometabolic health. However, in this study, not all associations between the natural environment 767 index and cardiometabolic health were mediated by physical activity and sedentary behaviours. Direct positive effects were observed on waist circumference, LDL cholesterol and HDL cholesterol. 768 769 Unmeasured leisure-time physical activities (e.g., swimming, surfing, bowling), exposure to UV 770 radiation, which has been associated with reduced risk of obesity and metabolic disease (Gorman et 771 al., 2017), and/or lower stress levels (Catalina-Romero et al., 2013; Sharma et al., 2022; Tomiyama, 772 2019) resulting from spending time in nature (Zhang et al., 2021) may be responsible for these direct effects. 773
- Finally, we observed a moderating effect of air pollution on the association between the natural environment and HDL cholesterol. A negative direct relationship was found only at above average levels of air pollution, suggesting that time spent outdoors in more polluted areas may have an undesirable effect on HDL cholesterol. It is, though, unclear why this effect was observed only in one of the six examined indicators of cardiometabolic health since previous studies have documented worse metabolic health outcomes for those participating in physical activity in more polluted environments (D'Oliveira et al., 2023).
- 781

# 782 *4.3. Strengths, limitations and future studies*

783 This study addressed several important shortcomings of the research on environmental 784 correlates of cardiometabolic risk factors for CVD. It estimated the independent associations of aspects of the neighbourhood built as well as natural environment with six cardiometabolic risk factors 785 786 for CVD while adjusting for neighbourhood SES and ambient air pollution. By doing so, unlike most 787 previous studies focusing on one or two dimensions of the urban environment (e.g., air pollution or built environment), it accounted for four key dimensions - built environment, natural environment, air 788 789 pollution and neighbourhood SES. To better understand how urban design may impact 790 cardiometabolic health across various levels of social and environmental disadvantage, this study also

791 examined the moderating effects of neighbourhood SES and air pollution, and mediating roles of 792 domain-specific physical activity and sedentary behaviours. Such a comprehensive analysis 793 acknowledges the complexities of the real world and the fact that an environmental characteristic may 794 have beneficial as well as detrimental effects on health via different pathways. Methodological 795 strengths include using data from a national study with good geographical coverage and environmental 796 variability, adjustment for neighbourhood self-selection based on self-report measures of reasons for 797 living in a neighbourhood and using directed acyclic graphs to develop analytical models informed by a careful analysis of the relevant literature. 798

799 This study has several limitations. The cross-sectional nature of the data limits our ability to 800 prove causal relationships. The natural environment index did not include information on the quality 801 of parkland areas. The land use variables in the walkability index did not distinguish between 802 destinations that are relevant to daily living from those that are not, making it more difficult to 803 distinguish the beneficial from the detrimental effects of dense, walkable neighbourhoods on 804 cardiometabolic health. We lacked traffic-related noise data and, hence, were unable to distinguish the 805 potential effects of noise on cardiometabolic health from those of co-occurring neighbourhood 806 attributes (e.g., traffic-related air pollution and population density). The participants included in the 807 third wave of AusDiab were healthier than those at baseline. Behaviours were assessed using self-808 reports which are known to have relatively large measurement errors. Information on the usual settings of the physical activity and sedentary behaviours was not available. A substantial proportion of these 809 810 behaviours may have been undertaken outside the neighbourhood. Data on the length of residence in a 811 particular neighbourhood were not available. These methodological issues may have resulted in an underestimation of the associations. 812

813 Future studies would need to determine how aspects of the neighbourhood environment are 814 related to trajectories of cardiometabolic risk factors for CVD across time. While the built and natural 815 environments do not typically change substantially in 5-10 years (the duration of most cohort studies), 816 there is a need for more evidence from longitudinal and quasi-experimental studies that investigate 817 potential effects of the environment on cardiometabolic health and related behaviours. Measures of 818 environmental exposures should encompass all key environmental attributes defining urban environment that may have contrasting direct and behaviour-mediated effects on cardiometabolic 819 820 health. These encompass density, street connectivity, destinations of daily living that impact physical 821 activity and dietary behaviours, green and blue spaces, air pollution, noise and area-level SES. Failure 822 to include all these factors is likely to result in contradictory, counterintuitive or misleading findings. To accurately estimate the effects of the neighbourhood environment on health, it is important to know 823 824 how much time individuals spend outdoors and indoors in their local community. Future studies should collect information on activity spaces using devices (e.g., global positioning system monitors 825 826 or ecological momentary assessment surveys) or map-based interviews (Kestens et al., 2018).

827

## 828 5. Conclusion

Within a relatively low-density and low-pollution context such as that of urban Australia, 829 830 denser, walkable neighbourhoods with good access to nature may benefit residents' cardiometabolic 831 health by facilitating walking for transport and leisure-time physical activity, reducing the need for 832 transport-related sitting (motorised transport), and displacing some indoor leisure-time sitting with 833 more active outdoor pursuits. Possible downsides of living in denser neighbourhoods are having 834 limited opportunities for gardening-related physical activities, exposure to higher levels of noise and air pollution, and exposure to eating-out outlets leading to less healthy dietary patterns with adverse 835 effect on cardiovascular health. Although, within the setting of this study, ambient air pollution 836 837 measures appeared to act as proxies for traffic-attracting destinations of daily living supporting an 838 active lifestyle, our findings suggest that time spent in public open spaces with higher level of air pollution may be associated with less favourable cardiometabolic outcomes (e.g., lower HDL 839 cholesterol) than time spent in less polluted locations. Finally, all the above-mentioned findings were 840 841 generally similar across levels of neighbourhood advantage and disadvantage, although, in a few 842 instances, residents of more disadvantaged neighbourhoods displayed stronger associations indicating that they may be more vulnerable to harmful environmental exposures compared to their more 843 advantaged counterparts. Therefore, socially disadvantaged neighbourhoods should be prioritised in 844 845 environmental public health interventions aimed at enhancing residents' cardiovascular health.

846

# 847 Funding

- This work was supported by a program grant ("The environment, active living and cognitive
  health: building the evidence base") from the Australian Catholic University [grant number
  ACURF18]. Jonathan E. Shaw is supported by a National Health and Medical Research Council
  (NHMRC) Investigator Grant [grant number 1173952].
- 852 The funders had no role in study design, data analysis, interpretation of the results, the853 decision to publish, or preparation of the manuscript.
- 854

## 855 CRediT Authorship Contribution Statement

Ester Cerin: Conceptualization, Data curation, Formal analysis, Funding acquisition, Investigation,
Methodology, Project administration, Software, Supervision, Validation, Visualization, Writing original draft, Writing - review & editing. Yih-Kai Chan: Project administration, Supervision,
Resources, Writing – review & editing. Mark Symmons: Resources, Writing – review & editing.
Maria Soloveva: Resources, Writing – review & editing. Erika Martino: Data curation, Investigation,
Software, Writing – review & editing. Jonathan E. Shaw: Conceptualisation, Data curation, Funding
acquisition, Investigation, Resources, Writing – Review & editing. Luke D. Knibbs: Data curation,

863 Methodology, Resources, Software, Writing – review & editing. Bin Jalaludin: Funding acquisition,

- 864 Writing review & editing. Anthony Barnett: Conceptualisation, Funding acquisition, Validation,
- 865 Writing review & editing.

- 867 Declaration of Competing Interest
  - The authors declare that there is no conflict of interest.
- 868 869

## 870 Acknowledgements

- 871 The AusDiab study, initiated and coordinated by the International Diabetes Institute, and
  872 subsequently coordinated by the Baker Heart and Diabetes Institute, gratefully acknowledges the
  873 support and assistance given by:
- B Atkins, B Balkau, E Barr, A Cameron, S Chadban, M de Courten, D Dunstan, A Kavanagh,
  D Magliano, S Murray, N Owen, K Polkinghorne, T Welborn, P Zimmet and all the study participants.
  Also, for funding or logistical support, we are grateful to: National Health and Medical
  Research Council (NHMRC grants 233200 and 1007544), Australian Government Department of
- c... Itestaten counten (11111100 granto 200200 una 10070 11), rustianun Government Department of
- 878 Health and Ageing, Abbott Australasia Pty Ltd, Alphapharm Pty Ltd, Amgen Australia, AstraZeneca,
- 879 Bristol-Myers Squibb, City Health Centre-Diabetes Service-Canberra, Department of Health and
- 880 Community Services Northern Territory, Department of Health and Human Services Tasmania,
- 881 Department of Health New South Wales, Department of Health Western Australia, Department of
- 882 Health South Australia, Department of Human Services Victoria, Diabetes Australia, Diabetes
- 883 Australia Northern Territory, Eli Lilly Australia, Estate of the Late Edward Wilson,
- 884 GlaxoSmithKline, Jack Brockhoff Foundation, Janssen-Cilag, Kidney Health Australia, Marian & FH
- 885 Flack Trust, Menzies Research Institute, Merck Sharp & Dohme, Novartis Pharmaceuticals, Novo
- 886 Nordisk Pharmaceuticals, Pfizer Pty Ltd, Pratt Foundation, Queensland Health, Roche Diagnostics
- Australia, Royal Prince Alfred Hospital, Sydney, Sanofi Aventis, sanofi-synthelabo, and the Victorian
  Government's OIS Program.
- 889 We are grateful to Mr David H. Lee for contributing to the computation of environmental
- 890 data.
- 891

# 892 Appendix A

- 893 Supplementary data
- 894
- 895 Data availability
- Be Data that support the findings of this study are available on request under a license agreement.
- 897 Written applications can be made to the AusDiab Steering Committee
- 898 (Dianna.Magliano@baker.edu.au).
- 899
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