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A GIS-based approach to assess the influence of the urban built environment on cardiac and respiratory outcomes in older adults

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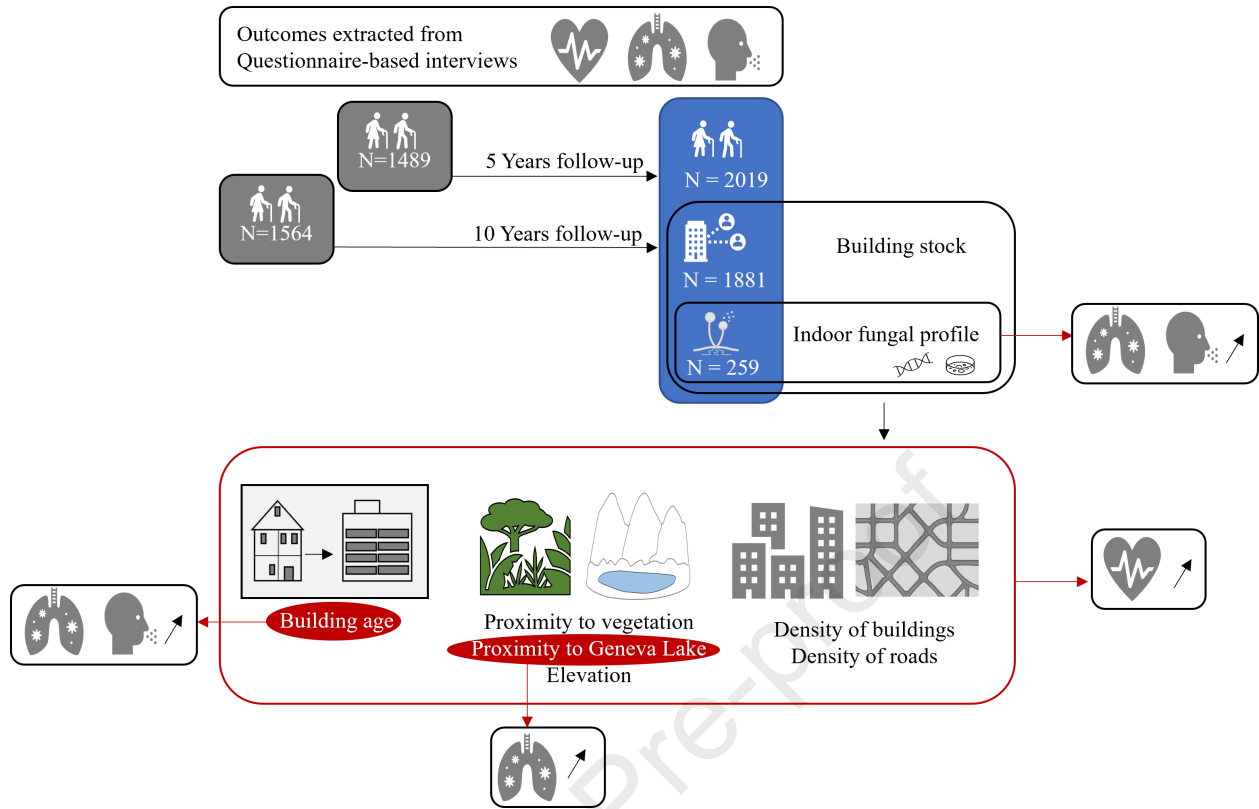
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2 **A GIS-Based Approach to Assess the Influence of the Urban Built**
3 **Environment on Cardiac and Respiratory Outcomes in Older**
4 **Adults**

5

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17 **Keywords:** Building age, Geographic Information System (GIS), Urban landscape, Indoor air
18 quality, Public health, Building associated microbiome, Next Generation Sequencing (NGS)

19 **Abstract**

20 The impact of the urban built environment on cardiovascular and respiratory health has been
21 studied extensively in children and adults. However, limited research exists on this topic in
22 older adults. To fill this gap, we conducted a ten-year retrospective study of a cohort of elderly
23 people living in Lausanne. We extracted cardiac and respiratory health outcomes of people
24 living at the same address. A Geographic Information System (GIS) application was used to
25 join spatial data between participant address and characteristics of the built environment
26 (building age, building density, road density, proximity to vegetation and lake, elevation). To
27 capture the impact of the urban landscape characteristics on the health of older people, the built
28 environment descriptors, were considered alone and aggregated. Principal Component
29 Analysis (PCA) was employed to resolve the multicollinearity among the built environment
30 descriptors, while Agglomerative Hierarchical Clustering was used to reveal underlying
31 patterns and relationships within built environment variables. In addition, to better assess the
32 effectiveness of ventilation in their homes, we characterized indoor air-associated mycobiomes
33 in a representative sample of homes. This involved employing both culture-based methods and
34 metabarcoding of the Internal Transcribed Spacer 1 (ITS1) of the ribosomal DNA. The analysis
35 revealed a significant association between exposure to the built environment as a whole and
36 heart disease in the elderly. A higher prevalence was observed for the cluster of buildings near
37 the lake/highway. The period of building construction showed an association with the
38 prevalence of chronic lung disease and chronic cough, with a higher prevalence observed in
39 houses built between 1975 and 2013. The contribution of indoor air pollution to worsening
40 respiratory health was confirmed by higher *Aspergillus* spore loads and increased relative
41 abundance of *Gloeophyllum* and *Aureobasidium* in indoor air, all of which were associated
42 with respiratory outcomes. This research highlights the crucial influence of the built

43 environment on the health status of the elderly population, with implications for targeted
44 interventions and public health policy development.

45

46 **1 Introduction**

47 The demographic landscape of the Organization for Economic Cooperation and Development
48 (OECD) countries is undergoing profound change, with a significant increase in the proportion
49 of older adults. Recent estimates suggest that by 2060, adults aged 65 and older will represent
50 a staggering 30% of the total population in these countries [1]. This demographic shift has
51 significant implications for public health and healthcare systems. Ensuring a high quality of
52 life for this growing elderly population and containing future healthcare costs have become
53 paramount concerns. Central to these efforts is the recognition that there are substantial benefits
54 to be gained from promoting healthy aging and facilitating the ability of older adults to age in
55 place - remaining at home for as long as possible.

56 Crucially, the built environment, including the design and condition of housing, plays a central
57 role in influencing the health and well-being of older people and has a profound impact on their
58 quality of life. Previous studies have highlighted the complex relationship between the built
59 environment, housing quality and human health. On the one hand, certain environmental
60 factors, such as air pollution and noise pollution steaming from heavy road traffic, have been
61 extensively linked to deterioration in respiratory and cardiovascular health [2-4]. On the other
62 hand, several environmental factors, such as the presence of green spaces near residential areas,
63 have been shown to have a protective effect on mortality, including among the elderly [5]. In
64 addition, the influence of residential density, a factor that promotes walkability and community
65 connectivity, has emerged as a significant contributor to improved health outcomes in older
66 adults [6, 7]. However, the influence of the built environment goes beyond external factors. It

67 is common for older adults to spend a significant portion of their time indoors is commonplace,
68 often due to reduced mobility, weather-related limitations, or environmental barriers for older
69 people in unfamiliar places [8]. This prolonged indoor exposure not only affects their lifestyles,
70 but also raises the specter of increased vulnerability to indoor air pollutants, which have been
71 linked to several health problems, including respiratory disease.

72 Older adults are particularly vulnerable to air pollutants due to normal and pathological aging
73 processes that lead to a decline in respiratory function and immune defenses, making them
74 more susceptible to respiratory infections. In addition to the effects of aging, older adults face
75 a number of chronic diseases, including cardiopulmonary disease, cancer, diabetes, and renal
76 failure. In particular, chronic obstructive pulmonary disease (COPD) and asthma are prevalent
77 in this population, further increasing their vulnerability to the adverse effects of air pollution.
78 When exposed to air pollution, the elderly experience an increased frequency of hospital
79 admissions for asthma and COPD, as well as a higher COPD mortality rate than others [9].

80 Moreover, this vulnerability extends beyond ambient air pollutants to include indoor air
81 pollutants. Previous studies have examined the relationship between indoor air quality and
82 health outcomes in older adults. These studies have demonstrated the increased susceptibility
83 of this population to specific indoor air pollutants, including total volatile organic compounds
84 (TVOC), particulate matter $PM_{2.5}$, and bioaerosols [10, 11]. However, there is limited data on
85 the long-term effects of indoor air pollution on the overall health decline in older adults.

86 Given the multitude of built environments that potentially impact the health of older adults, the
87 challenge is to identify which of these environments promote long-term health and which do
88 not. Deciding on the most effective overall solutions to improve healthy aging becomes a
89 complex task. To effectively address these issues, it is imperative that we shift our focus from
90 short-term assessments to understanding the lasting effects of the built environment on an aging

91 population. What sets our study apart is its innovative approach: we are conducting a
92 retrospective study of individuals living in the same built environments over the course of a
93 decade. This will allow us to assess the long-term impact of the built environment on the
94 collected health outcomes related to the progression of cardiac and respiratory disease in the
95 elderly population. To identify built environments that either significantly worsen the health of
96 older people or, conversely, serve as protective factors, we implemented a dual-method strategy
97 using Principal Component Analysis (PCA) and Agglomerative Hierarchical Clustering. This
98 approach allows us to comprehensively analyze the association of built environment
99 characteristics, join through Geographic Information System (GIS) application with health
100 outcomes in the elderly. PCA helps to address multicollinearity of the environmental variables,
101 while Agglomerative Hierarchical Clustering reveals underlying patterns and relationships
102 within built environment variables. Furthermore, since building age serves as a proxy for
103 indoor exposure, we investigate in a representative number of buildings whether specific
104 construction periods contribute to the development of unhealthy fungi in indoor environments.
105 This involved employing both culture-based methods and metabarcoding of the Internal
106 Transcribed Spacer 1 (ITS1) of the ribosomal DNA.

107

108 **2. Materials and methods**

109 The proposed approach to identify characteristics of the built environment that influence the
110 cardiac and respiratory health of the elderly at a citywide scale includes the following steps: i)
111 extraction of ten years of retrospective data on the cardiac and respiratory health of residents
112 of the Lausanne Cohort 65+ (Lc65+) who have lived in the same location throughout the study
113 period, ii) collection of building age and building environment characteristics based on
114 participants' addresses, iii) investigation of associations between variations in building age and

115 building environment gradient with changes in cardiac or respiratory health, and iv)
116 investigation of whether specific indoor fungal profiles can predict exposure to the built
117 environment and serve as indicators of health deterioration. A visual representation of the
118 methodology is provided in Figure 1.

119 **2.1. Cohort involved**

120 The data used in this study are derived from the Lausanne Cohort 65+ (Lc65+), an ongoing
121 cohort study of 4668 adults aged 65 years and older in Lausanne, Switzerland [12]. The cohort
122 represents a diverse sample of residents of the city of Lausanne in the canton of Vaud, which
123 has a population of approximately 120000. The cohort was established to include individuals
124 born before, during, and after World War II. The study population initially included 1564
125 participants enrolled in 2004, followed by an additional sample of 1489 participants recruited
126 in 2009, and a further 1615 participants recruited in 2014. At the time of enrollment and every
127 three years thereafter, all Lc65+ participants underwent an interview, completed a health
128 questionnaire, and underwent a physical examination. The study was conducted in accordance
129 with the principles of the Declaration of Helsinki and was approved by the Ethics Committee
130 of the Canton of Vaud (initial protocol code 19/04; 2014, amendment approved on June 23,
131 2014). For the purposes of this study, only participants enrolled in 2004 and 2009 who did not
132 move after enrollment were included. Those with incomplete data for the building environment
133 variables were excluded.

134 **2.2. Built environment variables**

135 The city of Lausanne is located in Switzerland. It has a wide range of elevations from 400
136 meters to 700 meters, and a variety of environmental features, including proximity to Lake
137 Geneva (one of the largest lakes in Europe), forests, and highways. The city has a long history

138 of construction dating back to the 1600s. According to the Köppen climate classification
139 system [13], Lausanne's climate is classified as warm-summer ("Dfb"). Due to its proximity to
140 the lake, the average relative humidity in Lausanne varies between 65% and 78% depending
141 on the season. The concentration of various pollutants in Lausanne's ambient air has steadily
142 decreased over the past twenty-five years. For example, the annual average concentration of
143 particulate matter PM₁₀ decreased from 36 $\mu\text{g}\cdot\text{m}^{-3}$ in 1997 to 25 $\mu\text{g}\cdot\text{m}^{-3}$ in 2007, fell below 20
144 $\mu\text{g}\cdot\text{m}^{-3}$ in 2010, and has remained consistently between 16 $\mu\text{g}\cdot\text{m}^{-3}$ and 13 $\mu\text{g}\cdot\text{m}^{-3}$ since 2014.
145 These data are provided by the National Air Pollution Monitoring Network (NABEL) [14].
146 Similarly, the annual average NO₂ concentration was recorded at 60 $\mu\text{g}\cdot\text{m}^{-3}$ in 1991, fell below
147 50 $\mu\text{g}\cdot\text{m}^{-3}$ in 1997, and has remained relatively stable between 40 $\mu\text{g}\cdot\text{m}^{-3}$ and 45 $\mu\text{g}\cdot\text{m}^{-3}$ until
148 2018, according to data from the same network.

149 The characteristics of buildings and the effectiveness of their insulation have changed over the
150 years [15]. Prior to 1944, buildings primarily had cement slab floors with metal joists in
151 unheated areas, wood floors in heated areas, solid walls, and single-pane windows. Between
152 1945 and 1974, concrete slab floors were introduced in both heated and unheated spaces, walls
153 were often uninsulated or partially insulated and windows consisted of two simple panes of
154 glass without insulation. Starting in 1975, insulation improvements were implemented. These
155 included integrated insulation in both heated and unheated floors, mixed insulation in walls,
156 and the use of double-glazed windows. All of the featured buildings had natural ventilation and
157 central heating. Therefore, the construction period was chosen as an indicator of insulation
158 effectiveness and as a proxy for the potential accumulation of indoor pollutants. The
159 participants of our study inhabited a representative sample of dwellings from distinct
160 construction periods, as shown in Table S1.

161 Using the 1881 addresses of the 2019 eligible participants, we collected relevant data from
162 different databases. We used a Geographic Information System (GIS) application (ArcGIS
163 [16]) to comprehensively assess the spatial relationships between built environment features
164 relevant to our investigation (Figure S1) . First, each address was geolocated within the study
165 area to get its geographic coordinates. All GIS analyses were performed on these coordinates.
166 . The building elevation was obtained by extracting the elevation from the swissALTI^{3D} raster
167 dataset [17]. Information on proximity to forests, fields, and parks was obtained from Swiss
168 cadastral data by computing Euclidian distance analyses. Information on proximity to the lake
169 was obtained in the same way from the swissTLM^{3D} database [18]. Building density
170 information was computed in the GIS using the following process: First, the vector data of the
171 swissTLM^{3D} database [18] was converted into raster (gridded) data with a cell size of 2 x 2
172 meters where each cell indicates the presence or absence of a building. Second, a 1500 meter
173 diameter circle was drawn around each address and the number of cells with building presence
174 was counted. The same procedure was applied to street density. In addition, building
175 characteristics such as year of construction, heating system, ventilation system, and any
176 renovation history were obtained from the ‘Registre Cantonal des Bâtiments’ (RCB) database
177 [19], by making a spatial join between the addresses and the RCB’s coordinates, a method
178 commonly used in Geographic Information System (GIS) applications. This method links the
179 spatial information, such as building coordinates, with the relevant building characteristics.

180 To guarantee the privacy and confidentiality of the volunteers, measures were taken to avoid
181 any possibility of identification. Specifically, the explanatory variables characterizing built
182 environment, corresponding to a radial number of pixels on a digital map, were reported in
183 quintiles. These quintiles were determined based on the distribution of variables across all
184 addresses recorded in the Lc65+ database, with each quintile representing a 100 m x 100 m
185 area within 500 m radius.

186 2.3. Quantification of cultivable fungi in settled dust in indoor environments

187 A subsample of 259 dwellings was analyzed to investigate the relationship between building
188 age and the accumulation of indoor pollutants. Indoor fungal load was chosen as an indicator
189 of poor ventilation in buildings because of its proven effectiveness in previous studies [20-22].
190 To capture indoor fungal communities over a representative period of time, we used passive
191 sampling with Electrostatic Dust Collectors (EDCs), a well-established method that has been
192 validated for this purpose [23]. Participant recruitment, sampling protocols, and fungal
193 community characterization have been described in detail elsewhere [24]. Briefly, the study
194 was proposed during follow-up consultations of the Lc65+ cohort, and 287 eligible participants
195 agreed to participate. Each participant was provided with an electrostatic dust collector (EDC)
196 and instructed to place it in their main living room at a height of 1.20 to 1.60m above the floor.
197 Participants were also given a letter with instructions to return the EDC to the laboratory after
198 10 weeks. Upon arrival at the laboratory, dust samples were extracted from the EDC by
199 washing with 0.1% Tween 80 (Merck®, Darmstadt, Germany) in Stomacher™ (AES®,
200 Combourn, France). Cultivable fungal diversity was determined by culturing an aliquot of the
201 harvested liquid on Dichloran Glycerol 18% (DG18) agar medium at 25°C for five days. The
202 overall composition of the fungal communities was characterized using next generation
203 sequencing of internal transcribed spacer 1 (ITS1) with ITS1F and ITS2 universal fungal
204 primers, as previously described [24] (Bioproject: PRJNA1033586, Biosample:
205 SAMN38035622). Cultures revealed that *Penicillium* and *Aspergillus* were the most
206 commonly detected fungal genera. ITS1 metabarcoding analysis revealed the presence of 248
207 different fungal genera indoors [24].

208 2.4. Health outcomes

209 Two primary health outcomes were evaluated: worsening of heart disease or worsening of
210 respiratory disease over a five-year period, with a sensitivity analysis conducted to extend this
211 period to 10 years. These outcomes were determined based on the data collected at enrollment
212 and follow-up interviews. The following measures were considered for each outcome:

- 213 - Self-reported medical diagnoses: Participants were asked if a physician had ever told
214 them about specific chronic respiratory diseases (e.g., asthma, chronic bronchitis, other)
215 and/or heart diseases (e.g., coronary heart disease, other heart disease) within the past
216 12 months. This self-report method allowed for the identification of individuals with
217 existing medical diagnoses.
- 218 - Medication use: Participants were asked about their current medication use for chronic
219 respiratory or cardiac conditions, specifically whether they took medication once a
220 week or more frequently. This information provided insight into the ongoing
221 management and treatment of their respective conditions.
- 222 - Chronic symptoms: Participants were asked to report the presence of chronic symptoms
223 experienced within the past 6 months, including cough and dyspnea. These symptoms
224 served as indicators of respiratory distress and provided additional context regarding
225 participants' respiratory health.

226 **2.5. Covariates**

227 Individual-level covariates assessed at baseline included age, sex, education level, smoking
228 status (categorized as current, former, or never), and household size. These covariates
229 were chosen to capture important demographic and lifestyle factors that could potentially
230 influence the association between the built environment and cardiac and respiratory
231 outcomes in older adults.

232 **2.6. Statistical analysis**

233 Descriptive statistics – frequencies with proportions and means with standard deviations (SD)
234 – were used to describe the sociodemographic and clinical characteristics of respondents at
235 enrollment, 5 years, and 10 years after enrollment. To assess eventual differences in socio-
236 demographic characteristics between participants enrolled in 2004 and 2009 included in the
237 present study, we employed statistical tests tailored to the nature of the variables. Continuous
238 variables, such as age, were compared using a 2-sample t-test. For categorical variables, such
239 as gender or education level, we utilized the chi-squared test.

240 The variables characterizing the built environment around the 2019 addresses of the Lc65+
241 retained participants, as generated by the GIS application, were initially, tested one by one, for
242 their association to health outcomes, assuming a linear effect of the categories, in the absence
243 of a priori hypotheses. However, it is important to note that environmental variables, such as
244 green space and traffic-related variables, may exhibit temporal and/or spatial correlations, and
245 are often correlated with demographic and socioeconomic determinants. When including
246 correlated variables in regression models, it is crucial to consider the potential for erroneous
247 estimation of effect size, and broad confidence intervals, which can lead to inaccurate
248 interpretation. To mitigate the effects of multiple correlated variables, we reduce the
249 dimensionality of the dataset by conducting a principal component analysis (PCA) on the nine
250 built environment characteristics (proximity to forest, field, garden, water body other than lake,
251 the Lake Geneva, road density, building density around the address of interest, elevation, and
252 building construction period), prior to conducting statistical tests for association with health
253 outcomes. PCA is a commonly used method in epidemiological studies [25, 26] that transforms
254 a set of correlated variables into a smaller group of independent variables, called principal
255 components (PCs). These PCs, rather than individual variables, are used in regression analysis
256 to eliminate multicollinearity. However, the main drawback of PCA is its lack of

257 interpretability, as each principal component is a combination of variables, making inference
258 impossible. Furthermore, variables grouped under a single principal component may have
259 opposing effects on health outcomes. To control for such side effects, we also used a method
260 that allowed us to group the addresses based on the similarity of the overall neighborhood
261 descriptors [27, 28], the agglomerative hierarchical clustering (AHC) method. The eight
262 landscape descriptors were used employing the complete linkage method, and measuring
263 cluster distance using the Euclidean distance between the built environment variables
264 generated by the GIS application. This method complements PCA by providing additional
265 insights into the structure of the data and identifying clusters of dwellings with similar built
266 environment characteristics. Considering the number of dwellings with available information
267 on their surrounding characteristics, a maximum of ten clusters were considered in order to
268 maintain sufficient statistical power in each cluster. The dendrogram of cluster analysis is
269 shown in supplementary material Figure S2. The results indicated that five clusters fitted best
270 the data. In the manuscript, we referred to these five distinct neighborhood types (Table S1) as
271 the “built environment score” (BES) to facilitate the presentation of the results. The algorithm
272 was applied to all unique addresses available in the Lc65+ cohort with no missing variables.

273 Multilevel mixed-effects logistic regression analysis was conducted to explore associations
274 between discrete environmental variables and the prevalence of individual health outcomes.
275 For each symptom, selected confounding factors are adjusted using a backward modeling
276 approach. Initially, all factors were included, and subsequently, those found to have no
277 significant influence were iteratively eliminated using a significance level of 5%, considering
278 the large sample size of the population. Additionally, interactions between the remaining
279 factors, gender, and cohort were documented. Separate models were generated for each
280 environment variable considered, including the built environment descriptor, the built
281 environment PCA component, the BES, or the presence/absence of individual fungal species.

282 In our analyses, we systematically quantified the impact on health outcomes using odds ratios
283 (ORs), with adjustments made for age, sex, and smoking status.

284 To investigate the relationship between each symptom and a continuous variable representing
285 the built environment microbiome, encompassing either the overall mycobiome or the relative
286 abundance of individual fungal species, multivariate analysis of variance (MANOVA)
287 followed by multivariate regression was used. For statistical robustness, all fungal
288 concentrations, whether expressed as relative abundances or colony forming units, were log-
289 transformed to approximate a normal distribution. Statistical analyses were performed with
290 STATA version 17.0 (StataCorp LLC, College Station, TX, USA).

291

292 **3. Results**

293 **3.1. Population health status**

294 Among the 2019 participants included in the study (as shown in Table 1), 70% reported good
295 or very good perceived health at enrollment, and this proportion remained relatively constant
296 five years later (70%). However, ten years after enrollment, the proportion reporting good or
297 very good perceived health decreased slightly to 62%. Regarding heart disease, 16% of
298 participants reported having a heart condition (chronic heart disease and/or taking heart
299 medication once a week) at enrollment. This percentage increased to 19% at five years and to
300 28% at ten years. In addition, 13% of these participants reported taking regular medication for
301 their heart condition at enrollment, which increased to 16% at five years and 25% at ten years.
302 The results suggest that the risk of having a heart condition, particularly regular use of heart
303 medication, increases with age (odds ratio [95% confidence interval] = 1.13 [1.09 - 1.18], $p =$
304 0.000; odds ratio [95% confidence interval] = 1.14 [1.09 - 1.18], $p = 0.000$, respectively).

305 Furthermore, the prevalence of heart disease is higher in men than in women, with 26% of men
306 and 17% of women reporting heart disease.

307 In terms of respiratory outcomes, 19% of participants reported having a respiratory condition
308 at enrollment, including chronic lung disease, asthma medication use, and/or chronic cough.
309 This percentage increased slightly to 21% at five years and further to 27% at ten years.
310 Specifically, 7% of participants had chronic lung disease at enrollment, while 4% reported
311 taking asthma medication. After five years, the percentages increased to 9% for chronic lung
312 disease and 5% for asthma medication use. These percentages remained relatively stable ten
313 years after enrollment, with 10% reporting chronic lung disease and 5% taking asthma
314 medication. Those who were active smokers during the entire study period had an increased
315 risk of developing chronic lung disease (odds ratio [95% confidence interval] = 2.80 [1.86 -
316 4.22], $p = 0.000$), regardless of age or sex. Active smokers also had a higher prevalence of
317 chronic cough (odds ratio [95% confidence interval] = 2.02 [1.19 - 3.43], $p = 0.008$). In
318 contrast, the prevalence of asthma medication use was higher in women than in men (odds ratio
319 [95% confidence interval] = 1.89 [1.19 - 3.00], $p = 0.007$).

320 **3.2. Built environment components**

321 Principal component analysis (PCA) of the built environment variables yielded three
322 eigenvalues greater than 1.0, as shown in Table 2. The first principal component (PC1)
323 accounted for 47% of the total variance in the built environment data, while the second and
324 third principal components (PC2 and PC3) explained 12.4% and 11.6% of the variance,
325 respectively. PC1, with an eigenvalue of 4.2, primarily consisted mainly of variables related to
326 proximity to forests and urbanization density (density of buildings and roads in the surrounding
327 area). PC2, with an eigenvalue of 1.1, was mainly influenced by the proximity to the lake. PC3,
328 with an eigenvalue of 1.0, was mainly influenced by the age of the building and the proximity

329 to gardens / to water body other than lake. To effectively integrate the information provided by
330 these three components (urbanization density, proximity to the lake, and age of the building),
331 an agglomerative hierarchical clustering method was applied to the nine built environment
332 variables of the 2019 addresses. This analysis led to the identification of five distinct urban
333 landscape clusters across Lausanne. For the purpose of describing the results, this indicator was
334 named Built Environment Score (BES). Cluster 1 includes dwellings mainly located in rural
335 suburbs surrounded by forests and fields, with low densities of roads and buildings. Cluster 2
336 consists of buildings almost located in urban suburbs with nearby gardens. Cluster 3 groups
337 buildings in the historical town center characterized by high road and building densities, and a
338 lack of vegetation in the surroundings. Cluster 4 includes dwellings located in the urban center
339 with high road and building densities, not far from the lake. Cluster 5 comprises buildings
340 located near the lakeside with low road and building densities but in close proximity to a
341 highway (see Table S2-S9).

342 **3.3. Association between components of the built environment and health outcomes**

343 Regarding heart disease outcomes, none of principal component analysis (PCA) components
344 were significantly associated with an increase in heart disease prevalence (Table 3). Therefore,
345 we researched for an association between the overall characteristics of urban landscape, as
346 described by the five clusters generated by Agglomerative Hierarchical Clustering analysis,
347 and these health outcomes. A worsening of heart disease was positively associated with the
348 built environment score (BES), with individuals living in dwellings within cluster 5 of the
349 urban landscape environment being the most affected (Table 3, Figure S3). The OR was even
350 higher when only weekly medication for heart disease was considered (data not shown).

351 Regarding respiratory outcomes, principal component 2 (proximity to the lake) of the PCA was
352 associated with a higher prevalence of asthma medication use and chronic lung disease (Table

353 2). The principal component 3 (building age, and proximity to the garden /water source other
354 than lake) was negatively associated with the use of asthma medication (Table 3). However,
355 notice that the correlation coefficients between building age / proximity to the garden and PC3
356 were also negatives (Table 2). Therefore, lake proximity and building age were considered as
357 individual variables in the multivariate adjusted model for each respiratory outcome. The
358 analysis confirmed that lake proximity was statistically significantly associated with a higher
359 prevalence of asthma medication use (OR [95% confidence interval] = 1.22 [1.07 - 1.40], $p =$
360 0.004) and chronic lung disease (OR [95% confidence interval] = 1.18 [1.06 - 1.31], $p = 0.001$)
361 (Figure S3). Building age was found to be associated with chronic lung disease (OR [95%
362 confidence interval] = 1.13 [1.04 - 1.23], $p = 0.003$) and chronic cough (OR [95% confidence
363 interval] = 1.12 [1.01 - 1.25], $p = 0.035$), with more recent buildings showing a tendency
364 towards exacerbating respiratory conditions (Figure S3).

365 To further confirm the impact of the built environment on respiratory health, bioaerosols were
366 monitored in a representative sub-sample of 259 homes, as the concentration and composition
367 of bioaerosols are sensitive to the effectiveness of home ventilation. The results showed that
368 individuals exposed to a higher concentration of fungal spores, especially *Aspergillus*, had a
369 higher risk of developing chronic cough (OR [95% confidence interval] = 1.19 [1.04 - 1.36], p
370 = 0.009; OR [95% confidence interval] = 1.25 [1.08 - 1.45], $p = 0.003$, respectively), after
371 adjustment for age, sex, and smoking status. However, no significant associations were
372 observed between fungal community diversity or composition and health outcomes, even after
373 adjusting for proximity to green space (data not shown). Only specific associations between
374 the relative abundance of certain fungal taxa and respiratory outcomes were identified, such as
375 the relative abundance of *Gloeophyllum* and the prevalence of chronic lung disease (OR [95%
376 confidence interval] = 10.38 [1.41 - 76.48], $p = 0.022$), or the relative abundance of
377 *Aureobasidium* and the prevalence of asthma medication use (OR [95% confidence interval] =

378 14.08 [1.32 - 150.64], $p = 0.029$). However, it is important to note that these results are based
379 on a small number of patients with chronic cough (N=7), chronic lung disease (N=8), and
380 asthma medication use (N=2), as the mycobiome was successfully sequenced in the built
381 environment of 136 participants.

382 **4. Discussion**

383 Our study offers several notable contributions to the fields of environmental health, urban
384 planning, and gerontology. First, it allows us to observe trends in the health status of older
385 adults by providing a rare and comprehensive longitudinal dataset spanning a decade. Second,
386 it takes an interdisciplinary approach, integrating data on the built environment, air quality, and
387 health outcomes, to help us better understand the complex interactions among these factors
388 over time. Finally, it exploits the spatial variability of health outcomes within urban areas to
389 identify distinct urban landscape features that are associated with the deterioration of specific
390 health outcomes.

391 The findings of our study on the changing health status of older adults are consistent with
392 previous research showing that perceived health remains relatively stable with age but declines
393 rapidly as death approaches [29]. The observed increase in heart disease prevalence over the
394 ten-year period is consistent with previous research showing an increase in cardiovascular
395 disease with age [30]. The gender disparities found, with higher prevalence in men, are
396 consistent with known gender differences in heart disease incidence and mortality [31]. In
397 addition, our findings of increased use of cardiac medications over time support the notion that
398 older individuals are more likely to require pharmacological interventions to manage cardiac
399 disease [32]. Interestingly, older women in our study had a higher prevalence of asthma
400 medication use than men of similar age, a finding consistent with limited European[33] and
401 extensive Asian research [34]. Several factors may contribute to this discrepancy. Hormonal

402 fluctuations during menopause, common in older women, may influence asthma symptoms by
403 affecting airway inflammation and bronchial hyperresponsiveness [35]. In addition, unique
404 environmental exposures, including indoor air pollutants, allergens, and occupational hazards,
405 may disproportionately affect older women and contribute to the development and exacerbation
406 of asthma. In particular, older women in this age group were more likely to be exposed to
407 indoor air pollutants such as those from gas cooking [36], household use of cleaning products,
408 especially spray cleaners [37], and passive smoking [38], factors that have previously been
409 shown to increase the risk of asthma development and exacerbation. These specific
410 environmental factors may contribute significantly to the prevalence of asthma in older women.

411 One of our key findings highlights the significant impact of the overall built environment,
412 which includes factors beyond proximity to green space or heavy traffic identified in previous
413 studies, on the health of older adults. Specifically, our study finds a significant association
414 between the built environment score (BES) and the prevalence of heart disease. In light of our
415 findings and the existing body of research, it is becoming increasingly clear that a holistic
416 approach is needed when assessing the impact of the built environment on the mental and
417 physical health of older adults. This holistic perspective remains relevant, even as we recognize
418 the importance of identifying specific elements within the built environment that contribute to
419 the creation of healthier living spaces, so that we can take actionable steps to intervene and
420 transform the environment. Previous epidemiological studies have consistently shown robust
421 associations between exposure to air pollutants and the incidence of cardiovascular disease [4,
422 39, 40]. In particular, these studies have highlighted the significant health effects of exposure
423 to PM_{2.5} and NO₂. PM_{2.5} is of particular concern because of its ability to penetrate deeply into
424 the respiratory system and even enter the bloodstream, potentially leading to systemic
425 inflammation and oxidative stress [4, 40, 41]. NO₂, a major component of traffic-related air
426 pollution, has been linked to adverse cardiovascular effects, including increased blood pressure

427 and atherosclerosis [42]. While proximity to busy roads has been shown to increase exposure
428 to noise and traffic-related pollutants such as PM_{2.5} and NO₂ [4], access to green spaces and
429 natural environments has consistently been consistently shown to provide several health
430 benefits, including stress reduction [43], promotion of physical activity [44], improvement of
431 cardiovascular health, and reduction of mortality risk [5, 45, 46]. Similarly, the presence of
432 water bodies in urban environments, such as lakes or rivers, has been associated with positive
433 health outcomes, including improved mental health, and promotion of physical activity [47,
434 48]. In addition, neighborhood walkability, characterized by features such as sidewalks,
435 pedestrian-friendly infrastructure, and proximity to essential services, has been associated with
436 increased levels of physical activity and improved perceptions of physical health [6]. Studies
437 have shown that older adults living in walkable neighborhoods tend to engage in more physical
438 activity, which leads to improved mobility, fitness, and overall perceptions of physical health.

439 Another key finding of this study is the increased prevalence of respiratory conditions among
440 the older adult population over the ten-year period. While it is well-established that aging itself
441 can bring about physiological changes that impact respiratory health, it is essential to consider
442 the potential influence of air pollutants in either exacerbating or mitigating these changes. Our
443 study revealed specific features of the built environment associated with a decline in respiratory
444 health among older adults. Proximity to the lake, which cannot be distinguished from proximity
445 to the highway, was associated with a higher prevalence of asthma medication use, while
446 building age was associated with a higher prevalence of chronic lung disease and chronic
447 cough. These findings suggest that living near a lake/highway or in buildings constructed
448 between 1975 and 2013 may worsen respiratory conditions in older adults.

449 The choice of building materials and the effectiveness of ventilation systems in residential
450 buildings have been shown to have a significant impact on indoor air quality [49, 50]. The age

451 of the building serves as a reliable indicator of the indoor air quality because of the different
452 materials used during each construction period, and the specific emissions they generate. Some
453 materials tend to deteriorate and generate more fine particulate matter (PM_{2.5}) than others [51];
454 also, materials tend to emit higher levels of volatile organic compounds (VOCs) in the first
455 years after the construction than later [50]. It is noteworthy that while the role of PM_{2.5} in
456 asthma is still debated, the effects of VOCs on respiratory health, particularly their association
457 with asthma, have been established in systematic reviews [52]. The accumulation of indoor
458 pollutants in buildings constructed after 1975, compared to those constructed before 1975, was
459 facilitated by improved insulation standards and the lack of implementation of efficient
460 ventilation systems. This oversight often leads to the accumulation of moisture in these
461 buildings, creating favorable conditions for indoor mold growth [22]. Such accumulation is of
462 particular concern for older adults who are already susceptible to respiratory issues due to age-
463 related changes in lung function [53, 54]. Living in damp indoor environments has been
464 recognized as a contributing factor to respiratory health issues [55, 56]. Damp indoor
465 environments can lead to mold growth, which is associated with the release of mycotoxins and
466 allergenic spores. Prolonged exposure to these indoor contaminants can exacerbate respiratory
467 conditions [57, 58]. In contrast, well-designed and maintained ventilation systems, combined
468 with appropriate building materials, have the potential to significantly improve indoor air
469 quality. Adequate ventilation can reduce indoor pollutant concentrations, including particulate
470 matter, VOCs, and allergens, thus promoting a healthier indoor environment [59]. Furthermore,
471 building designs that prioritize the use of low-emitting materials and promote proper air
472 exchange can play a pivotal role in the respiratory health of older populations [60].

473 Our findings suggest that increased insulation in buildings constructed after 1975 may lead to
474 the accumulation of indoor air pollutants in the absence of efficient ventilation systems. To
475 further explore whether the influence of the built environment on respiratory health is

476 associated with inefficient ventilation leading to dampness and mold growth, we characterized
477 mycobiome composition by culture and metabarcoding in a representative subsample of
478 homes. Indeed, mold growth is known to thrive in environments with excess moisture due to
479 inadequate ventilation. Our results showed that individuals exposed to higher concentrations
480 of fungal spores, particularly *Aspergillus*, had an increased risk of developing chronic cough.

481 Numerous studies have shown that indoor air quality, influenced by building age, ventilation
482 systems, and the presence of moisture or dampness, can significantly affect respiratory health.
483 It is worth exploring whether the observed associations between specific fungal taxa and
484 respiratory outcomes are exacerbated or attenuated by characteristics of the built environment
485 such as the presence of green space or the age of the building. The associations between fungal
486 spores load and respiratory outcomes prevalence are consistent with studies highlighting the
487 role of indoor air quality and the mycobiome composition in respiratory health [61, 62].
488 However, the unique associations between specific fungal taxa and respiratory outcomes
489 underscore the complexity of indoor mycobiome interactions with health, and warrant further
490 investigation [63].

491 The strength of our study lies in the use of diverse statistical methods, including hierarchical
492 clustering and principal component analysis (PCA), to explore the relationships between urban
493 landscape characteristics and health outcomes. Hierarchical clustering revealed the importance
494 of considering the built environment as a whole in elucidating its contribution to the long-term
495 development of cardiac outcomes. Conversely, PCA proved to be more effective in identifying
496 the environmental variables that exert an influence on the respiratory health of older adults.
497 This could be explained by the fact that the size of the population followed up in each cluster
498 of urban landscape considered was not large enough to detect a clear association between this
499 built environment score and the worsening of the respiratory health in the elderly. Furthermore,

500 the inclusion of indoor air quality data underscored the importance of collecting such data to
501 refine our understanding of the interplay between indoor and outdoor environmental factors
502 and their impact on respiratory health. This holistic approach highlights the importance of
503 employing various statistical methods to comprehensively capture the multifaceted effects of
504 the built environment on human health. However, while our study provides valuable insights
505 into the association between the built environment and cardiac and respiratory outcomes in
506 older adults, it is important to acknowledge several limitations. First, the study was conducted
507 in a specific geographic location (Lausanne, Switzerland), which may limit the generalizability
508 of the findings to other settings. Future research should aim to replicate these findings in
509 different populations and geographical contexts. Second, some respiratory outcome categories
510 had relatively small sample sizes, resulting in limited statistical power. This limitation may
511 have affected our ability to detect significant associations between the fungal community
512 composition and health outcomes. Future studies with larger sample sizes are warranted to
513 validate these findings and explore the potential role of fungal diversity in respiratory health.
514 Furthermore, although we have discussed the potential mediating role of the built environment,
515 causality cannot be definitively established within the scope of this observational study.
516 Intervention studies would be needed to elucidate causal relationships between specific
517 characteristics of the built environment, air quality, and health outcomes in older adults.
518 Finally, the generalizability of our findings to diverse populations and geographic regions
519 should be approached with caution, as our study focused on a specific urban area. Despite these
520 limitations, our research provides a valuable foundation for future investigation and urban
521 planning initiatives aimed at promoting healthier aging.

522 In conclusion, our study highlights the significant influence of the built environment on cardiac
523 and respiratory health outcomes in older adults. These findings underscore the importance of
524 built environment factors, such as proximity to green space, urbanization density, and building

525 age, in promoting healthy aging. Such findings have profound implications for urban planning
526 and public health interventions aimed at creating age-friendly environments that support the
527 well-being of older populations. Our research offers practical implications for urban planning
528 and policy development, suggesting that designing age-friendly urban environments with better
529 access to green spaces and improved air quality could promote better health and well-being
530 among older populations. Furthermore, the findings have the potential to inform evidence-
531 based policies aimed at improving urban living conditions for older adults, such as initiatives
532 to reduce air pollution, improve green infrastructure, and create age-friendly neighborhoods.
533 By examining health disparities within urban areas, our research also highlights the need for
534 targeted interventions to address the specific health needs of vulnerable populations.
535 Ultimately, this study lays a solid scientific foundation for future investigations into the
536 relationships between the built environment, air quality, and health, and serves as a reference
537 point for designing more focused studies and interventions in this important area of research.

538 **5. Conflict of Interest**

539 The authors declare that the research was conducted in the absence of any commercial or
540 financial relationships that could be construed as a potential conflict of interest.

541 **6. Author Contributions**

542 Conceptualization, H.N-H. and P.W.; methodology, H.N-H. and P.W.; data extraction and GIS
543 analysis, A.H.H.; formal analysis, H.N-H. and P.W; investigation, H.N-H. resources, H.N-H.;
544 writing—original draft preparation, H.N-H. and P.W; writing—review and editing, H.N-H. All
545 authors have read and agreed to the published version of the manuscript.

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552 **9. Supplementary Material**

553 The Supplementary Material for this article can be found online at:

554 **Figure S1.** Geospatial localization of dwellings considered in the study.

555 **Figure S2.** Dendrogram of agglomerative hierarchical cluster analysis

556 **Figure S3.** Marginal predicted mean of health outcomes by built environment descriptor, with
557 95% confidence intervals. **A.** Proportion of heart medication by urban landscape cluster; **B.**
558 Proportion of medication for asthma by quantile of proximity to the lake; **C.** Proportion of
559 chronic lung disease by construction period of the building.

560 **Table S1.** Number of participants residing in buildings from each construction period

561 **Table S2.** Descriptive statistics of road density variable in the five urban landscape clusters.

562 **Table S3.** Descriptive statistics of building density variable in the five urban landscape
563 clusters

564 **Table S4.** Descriptive statistics of elevation variable in the five urban landscape clusters

565 **Table S5.** Descriptive statistics of proximity to Geneva Lake variable in the five urban
566 landscape clusters.

567 **Table S6.** Descriptive statistics of proximity to water other than lake variable in the five
568 urban landscape clusters.

569 **Table S7.** Descriptive statistics of proximity to forest variable in the five urban landscape
570 clusters.

571 **Table S8.** Descriptive statistics of proximity to garden variable in the five urban landscape
572 clusters

573 **Table S9.** Descriptive statistics of proximity to field variable in the five urban landscape
574 clusters.

575 Data Availability Statement

576 The data presented in this study are available on request from the authors. The NGS data are
577 available at NCBI under the Bioproject: PRJNA1033586 and Biosample: SAMN38035622.

578 **10. Declaration of generative AI and AI-assisted technologies in the writing process**

579 During the preparation of this work the authors used ChatGPT-3.5 in order to improve language
580 and readability. After using this tool, the authors reviewed and edited the content as needed
581 and take full responsibility for the content of the publication.

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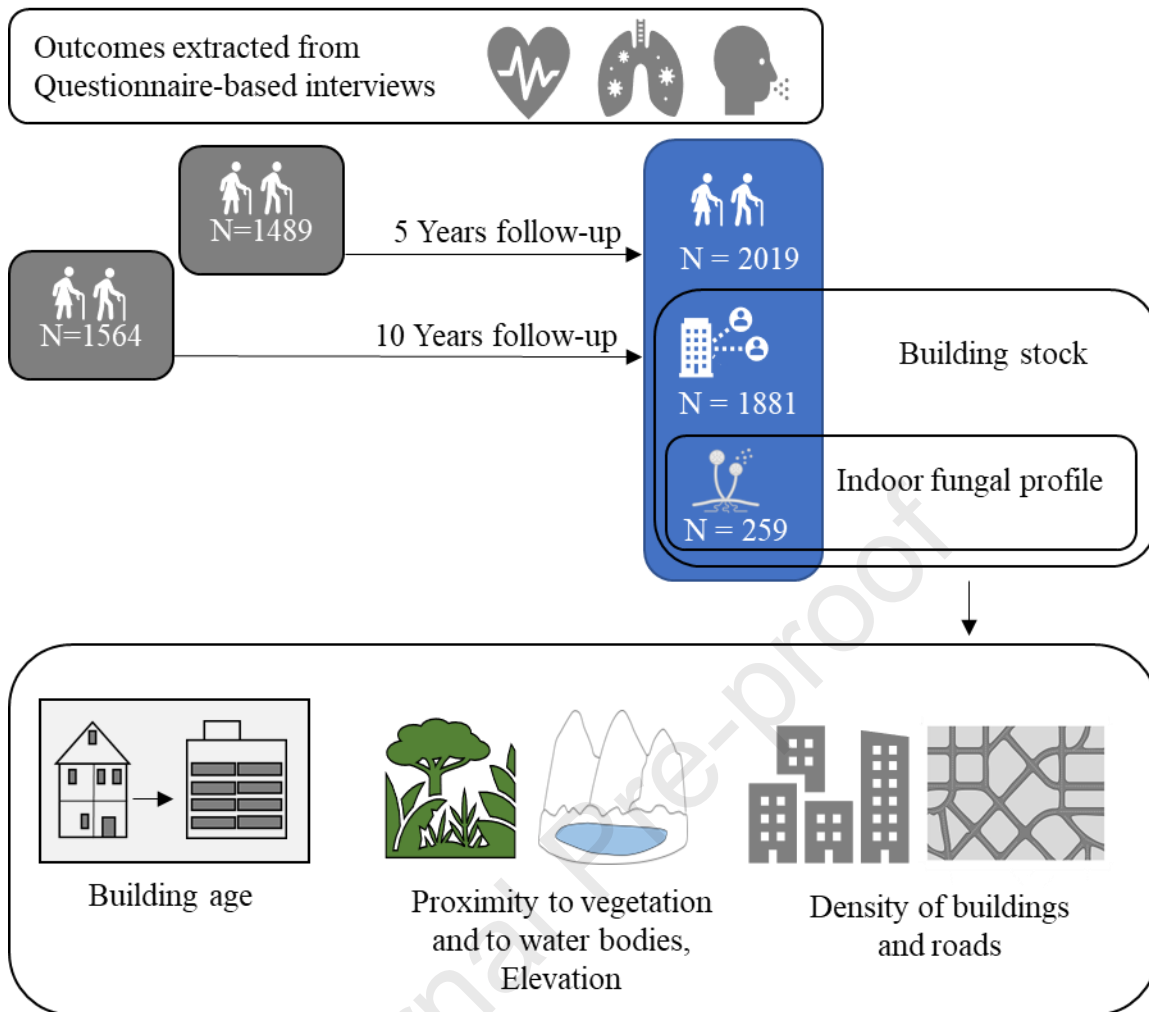
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Figure 1. Study Design: This city-wide study was conducted in Lausanne, a city characterized by significant variations in elevation and environmental characteristics. The Lc65+ cohort, established in Lausanne, is designed to monitor health deterioration in the city's elderly population. We included participants who had not changed their residence since their initial inclusion and extracted cardiac and respiratory outcomes from questionnaire interviews conducted at inclusion, as well as 5 and 10 years later. Participants' addresses were used to retrieve relevant data from various datasets, such as building age, elevation, proximity to green spaces and the lake, and information on building and road density. Indoor fungal profiles were assessed using next-generation sequencing and cultivation in a representative subset of dwellings. Multilevel mixed-effects logistic models were employed to examine the associations between health outcomes and the built environment.

764 **Table 1.** Sociodemographic characteristics of the studied population, N = 2019

Characteristic	Male (N = 784)	Female (N = 1235)
Age, years (Mean \pm SD)	75.2 \pm 2.9	75.2 \pm 2.9
Smoking status		
Non-smokers, N (%)	211 (27%)	622 (51%)
Ex-smokers, N (%)	439 (57%)	438 (36%)
Active smokers, N (%)	124 (16%)	162 (14%)
Education		
Compulsory education	98 (13%)	315 (26%)
Federal Diploma of Vocational ET ¹	339 (43%)	473 (38%)
High school diploma	37 (5%)	135 (11%)
Advanced Federal Diploma of Professional ET ¹	135 (17%)	189 (15%)
University diploma	174 (22%)	120 (10%)
Household size		
single	183 (24%)	711 (59%)
couple	555 (72%)	481 (40%)
more of two	30 (4%)	20 (2%)

¹ ET = Education and Training

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Table 2. Correlation coefficients between the nine built environment variables and the three principal components (PCs) obtained through Principal Component Analyses (PCA) using data extracted from 1881 addresses. Here, a correlation was considered moderate when the correlation coefficient exceeded 0.4 or fell below -0.4, and strong when it exceeded 0.5 or fell below -0.5.

Built environmental variable	PC1	PC2	PC3
Building age	0.25	-0.13	-0.44
Proximity to the forest within a 500 m radius	0.41	-0.07	0.21
Proximity to the field within a 500 m radius	0.34	0.12	0.13
Proximity to the garden within a 500 m radius	0.29	-0.02	-0.54
Proximity to the lake within a 500 m radius	-0.15	0.79	-0.26
Proximity to the water body other than lake within a 500 m radius	0.28	0.38	0.53
Density of roads within a 500 m radius	-0.40	0.05	0.25
Density of buildings within a 500 m radius	-0.41	-0.35	0.08
Elevation	0.37	-0.25	0.21
Eigenvalue	4.24	1.12	1.04
% of variance explained	47.1	12.4	11.6

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774 **Table 3.** Associations between health outcomes and built environmental variables (radius: 500
 775 m) or exposure to moulds

	Cardiac condition	Medication for heart condition	Medication for asthma	Chronic lung disease	Chronic cough
Variable	OR [95% CI]	OR [95% CI]	OR [95% CI]	OR [95% CI]	OR [95% CI]
BES ^{Model1}	1.11 [1.00 – 1.24]*	1.15 [1.03 – 1.28]*	1.01 [0.83 – 1.24]	1.06 [0.91 – 1.22]	0.89 [0.72 1.10]
PC 1 ^{Model2}	0.97 [0.92 – 1.02]	0.95 [0.90 – 1.00]	1.06 [0.96 – 1.16]	1.03 [0.96 – 1.10]	1.06 [0.96 1.12]
PC 2 ^{Model3}	1.05 [0.95 – 1.16]	1.06 [0.96 – 1.17]	1.24 [1.06 – 1.46]**	1.23 [1.08 – 1.39]***	0.94 [0.77 1.15]
PC 3 ^{Model4}	0.96 [0.87 – 1.07]	0.99 [0.89 – 1.11]	0.81 [0.67 – 0.99]*	0.91 [0.79 – 1.05]	0.83 [0.69 1.01]
Building age ^{Model11}	0.97 [0.92 – 1.02]	0.95 [0.90 – 1.01]	1.08 [0.98 – 1.20]	1.13 [1.04 – 1.23]***	1.12 [1.01 – 1.25]*
Proximity to the lake ^{Model5}	1.04 [0.96 – 1.14]	1.05 [0.96 – 1.15]	1.22 [1.07 – 1.40]***	1.18 [1.06 – 1.31]***	0.99 [0.83 1.18]
Total CFU ^{Model6}	0.95 [0.87 – 1.05]	0.94 [0.85 – 1.03]	1.06 [0.88 – 1.28]	0.91 [0.78 1.07]	1.19 [1.04 – 1.36]**
<i>Aspergillus</i> CFU ^{Model7}	0.93 [0.79 – 1.09]	0.94 [0.80 – 1.10]	0.86 [0.49 – 1.52]	0.82 [0.54 1.24]	1.25 [1.08 – 1.45]***
<i>Gloeophyllum</i> abundance ^{Model8}	2.00 [0.54 – 7.44]	2.04 [0.54 – 7.64]	5.80 [0.15 – 217.59]	10.38 [1.41 – 76.48]*	0.53 [0.04 – 7.91]
<i>Aureobasidium</i> abundance ^{Model9}	0.81 [0.38 – 1.75]	0.87 [0.41 – 1.85]	14.08 [1.32 – 150.64]*	1.69 [0.57 4.98]	0.12 [0.00 – 5.55]

OR = odd ratio ; CI = confidence interval ; CFU = colonies forming units

Separate models were constructed to estimate the effect of each environmental variable (BES, PCA component, or built environment variable) on health outcomes. Health effects were systematically quantified using odds ratios (ORs) adjusted for age, sex, and smoking status.

*p < 0.05, **p < 0.01, ***p < 0.005

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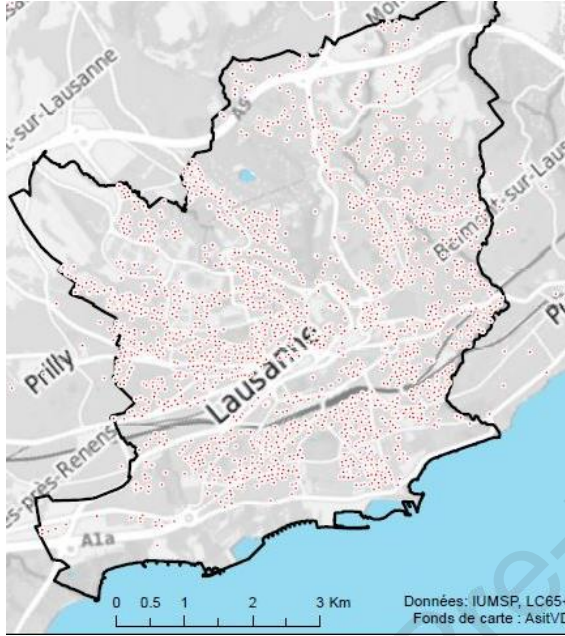
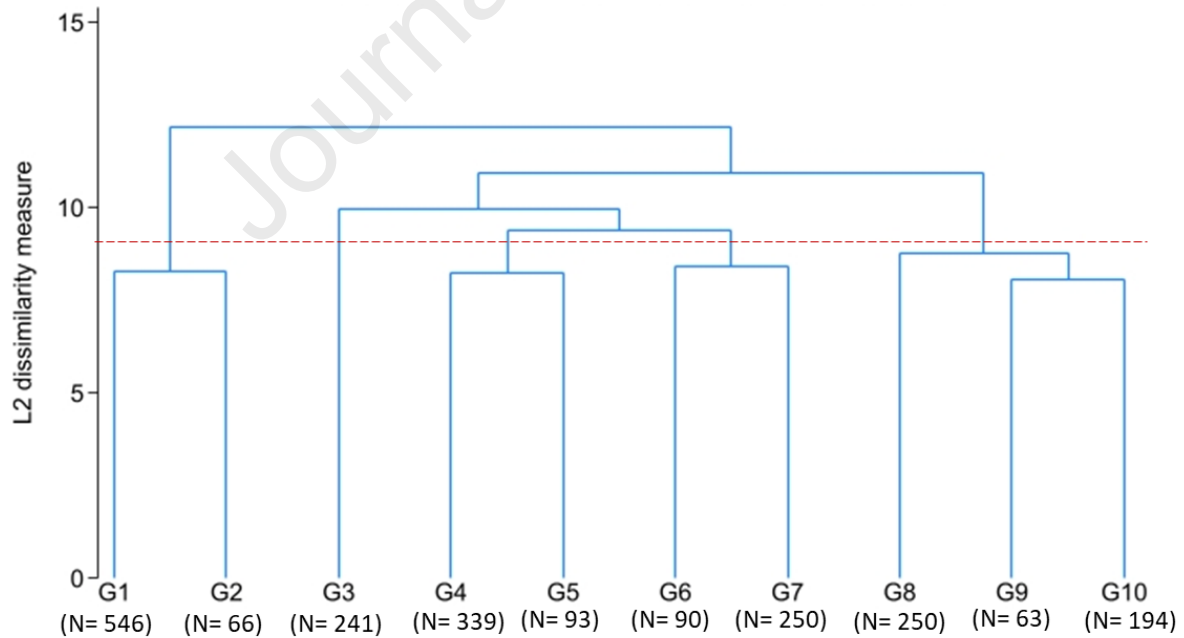


Figure S1. Geospatial localization of dwellings considered in the study.



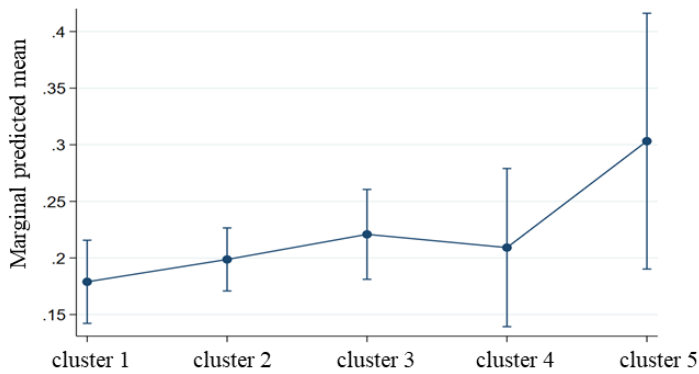
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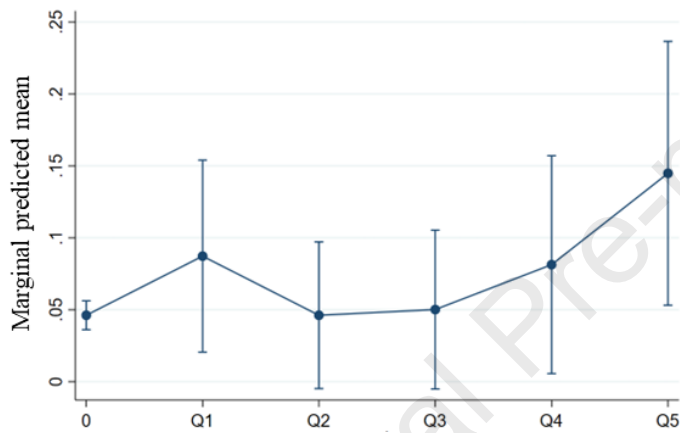
Figure S2. Dendrogram of agglomerative hierarchical cluster analysis

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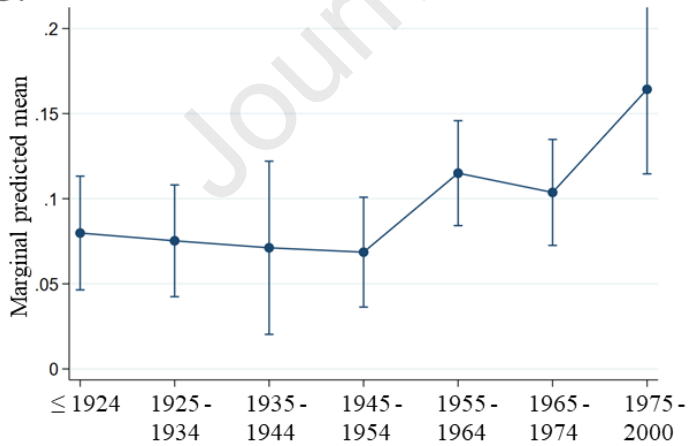
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809 **Figure S3.** Marginal predicted mean of health outcomes by built environment descriptor,
 810 with 95% confidence intervals. **A.** Proportion of heart medication by urban landscape cluster;
 811 **B.** Proportion of medication for asthma by quantile of proximity to the lake; **C.** Proportion of
 812 chronic lung disease by construction period of the building.

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816 **Table S1.** Number of participants residing in buildings from each construction period

Building construction period	Participants
before 1925	258
1925-1934	256
1935-1944	102
1945-1954	257
1955-1964	447
1965-1974	423
1975-2014	258

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Table S2. Descriptive statistics of road density variable in the five urban landscape clusters.

Cluster	Q1	Q2	Q3	Q4	Q5	Total
1 (N)	213	155	74	31	0	473
1 (%)	45.03	32.77	15.64	6.55	0	100
2 (N)	183	219	277	203	34	916
2 (%)	19.98	23.91	30.24	22.16	3.71	100
3 (N)	0	35	61	106	307	509
3 (%)	0	6.88	11.98	20.83	60.31	100
4 (N)	0	0	3	37	98	138
4 (%)	0	0	2.17	26.81	71.01	100
5 (N)	28	11	8	28	6	81
5 (%)	34.57	13.58	9.88	34.57	7.41	100
Total (N)	424	420	423	405	445	2117
Total (%)	20.03	19.84	19.98	19.13	21.02	100

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Table S3. Descriptive statistics of building density variable in the five urban landscape clusters

Cluster	Q1	Q2	Q3	Q4	Q5	Total
1 (N)	295	165	13	0	0	473
1 (%)	62.37	34.88	2.75	0	0	100
2 (N)	97	251	319	211	38	916
2 (%)	10.59	27.4	34.83	23.03	4.15	100
3 (N)	0	0	30	136	343	509
3 (%)	0	0	5.89	26.72	67.39	100
4 (N)	1	5	36	85	11	138
4 (%)	0.72	3.62	26.09	61.59	7.97	100
5 (N)	33	18	29	1	0	81
5 (%)	40.74	22.22	35.8	1.23	0	100
Total (N)	426	439	427	433	392	2117
Total (%)	20.12	20.74	20.17	20.45	18.52	100

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825 **Table S4.** Descriptive statistics of elevation variable in the five urban landscape clusters

Cluster	Q1	Q2	Q3	Q4	Q5	Total
1 (N)	27	55	43	48	300	473
1 (%)	5.71	11.63	9.09	10.15	63.42	100
2 (N)	112	84	224	397	99	916
2 (%)	12.23	9.17	24.45	43.34	10.81	100
3 (N)	130	226	151	2	0	509
3 (%)	25.54	44.4	29.67	0.39	0	100
4 (N)	77	61	0	0	0	138
4 (%)	55.8	44.2	0	0	0	100
5 (N)	81	0	0	0	0	81
5 (%)	100	0	0	0	0	100
Total (N)	427	426	418	447	399	2117
Total (%)	20.17	20.12	19.74	21.11	18.85	100

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827 **Table S5.** Descriptive statistics of proximity to Geneva Lake variable in the five urban
828 landscape clusters.

Cluster	NA	Q1	Q2	Q3	Q4	Q5	Total
1 (N)	440	3	0	30	0	0	473
1 (%)	93.02	0.63	0	6.34	0	0	100
2 (N)	912	2	1	1	0	0	916
2 (%)	99.56	0.22	0.11	0.11	0	0	100
3 (N)	443	63	3	0	0	0	509
3 (%)	87.03	12.38	0.59	0	0	0	100
4 (N)	0	1	60	29	24	24	138
4 (%)	0	0.72	43.48	21.01	17.39	17.39	100
5 (N)	0	0	0	2	38	41	81
5 (%)	0	0	0	2.47	46.91	50.62	100
Total (N)	1,795	69	64	62	62	65	2117
Total (%)	84.79	3.26	3.02	2.93	2.93	3.07	100

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842 **Table S6.** Descriptive statistics of proximity to water other than lake variable in the five
 843 urban landscape clusters.

Cluster	NA	Q1	Q2	Q3	Q4	Q5	Total
1 (N)	0	0	49	82	129	213	473
1 (%)	0	0	10.36	17.34	27.27	45.03	100
2 (N)	531	109	103	101	72	0	916
2 (%)	57.97	11.9	11.24	11.03	7.86	0	100
3 (N)	348	84	40	31	6	0	509
3 (%)	68.37	16.5	7.86	6.09	1.18	0	100
4 (N)	137	1	0	0	0	0	138
4 (%)	99.28	0.72	0	0	0	0	100
5 (N)	27	1	21	10	13	9	81
5 (%)	33.33	1.23	25.93	12.35	16.05	11.11	100
Total (N)	1,043	195	213	224	220	222	2117
Total (%)	49.27	9.21	10.06	10.58	10.39	10.49	100

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845 **Table S7.** Descriptive statistics of proximity to forest variable in the five urban landscape
 846 clusters.

Cluster	NA	Q1	Q2	Q3	Q4	Q5	Total
1 (N)	0	0	0	47	168	258	473
1 (%)	0	0	0	9.94	35.52	54.55	100
2 (N)	14	51	188	274	246	143	916
2 (%)	1.53	5.57	20.52	29.91	26.86	15.61	100
3 (N)	47	239	153	65	5	0	509
3 (%)	9.23	46.95	30.06	12.77	0.98	0	100
4 (N)	30	76	32	0	0	0	138
4 (%)	21.74	55.07	23.19	0	0	0	100
5 (N)	14	19	32	8	8	0	81
5 (%)	17.28	23.46	39.51	9.88	9.88	0	100
Total (N)	105	385	405	394	427	401	2117
Total (%)	4.96	18.19	19.13	18.61	20.17	18.94	100

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848 **Table S8.** Descriptive statistics of proximity to garden variable in the five urban landscape
 849 clusters.

Cluster	NA	Q2	Q3	Q4	Q5	Total
1 (N)	41	106	104	93	129	473
1 (%)	8.67	22.41	21.99	19.66	27.27	100
2 (N)	1	66	261	297	291	916
2 (%)	0.11	7.21	28.49	32.42	31.77	100
3 (N)	319	170	19	1	0	509
3 (%)	62.67	33.4	3.73	0.2	0	100
4 (N)	37	61	25	15	0	138
4 (%)	26.81	44.2	18.12	10.87	0	100
5 (N)	5	18	24	15	19	81

5 (%)	6.17	22.22	29.63	18.52	23.46	100
Total (N)	403	421	433	421	439	2117
Total (%)	19.04	19.89	20.45	19.89	20.74	100

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851 **Table S9.** Descriptive statistics of proximity to field variable in the five urban landscape
852 clusters.

Cluster	NA	Q1	Q2	Q3	Q4	Q5	Total
1 (N)	42	6	65	109	104	147	473
1 (%)	8.88	1.27	13.74	23.04	21.99	31.08	100
2 (N)	540	93	51	49	110	73	916
2 (%)	58.95	10.15	5.57	5.35	12.01	7.97	100
3 (N)	269	72	76	82	10	0	509
3 (%)	52.85	14.15	14.93	16.11	1.96	0	100
4 (N)	99	37	2	0	0	0	138
4 (%)	71.74	26.81	1.45	0	0	0	100
5 (N)	23	35	14	1	5	3	81
5 (%)	28.4	43.21	17.28	1.23	6.17	3.7	100
Total (N)	973	243	208	241	229	223	2117
Total (%)	45.96	11.48	9.83	11.38	10.82	10.53	100

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Highlights:

- Urban landscape impacts heart disease in the elderly;
- Living in homes built between 1975 and 2013 worsens the respiratory health;
- Fungal indicator species link indoor air pollution to respiratory outcomes
- Individuals exposed to a higher concentrations of *Aspergillus* spores had a higher risk of developing chronic cough.

Journal Pre-proof

Declaration of interests

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

The authors declare the following financial interests/personal relationships which may be considered as potential competing interests:

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