1	Cardiometabolic diseases and associated risk factors in transitional rural
2	communities in tropical coastal Ecuador
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19 Abstract

20 Background

There is a growing epidemic of chronic non-communicable diseases in low and middleincome countries, often attributed to urbanization, although there are limited data from marginalized rural populations. This study aimed to estimate prevalence of cardiometabolic diseases and associated risk factors in transitional rural communities.

25 Methods

A cross-sectional study of Montubio adults aged 18-94 years living in agricultural communities in a tropical coastal region of Ecuador. Data were collected by questionnaires and anthropometry, and fasting blood was analyzed for glucose, glycosylated hemoglobin, insulin, and lipid profiles. Population-weighted prevalences of diabetes, hypertension, and metabolic syndrome were estimated. Associations between potential risk factors and outcomes were estimated using multilevel regression techniques adjusted for age and sex.

32 **Results**

Out of 1,010 adults recruited, 931 were included in the analysis. Weighted prevalences were estimated for diabetes (20.4%, 95% CI 18.3-22.5%), hypertension (35.6%, 95% CI 29.0-42.1%), and metabolic syndrome (54.2%. 95% CI 47.0-61.5%) with higher prevalence observed in women. Hypertension prevalence increased with age while diabetes and metabolic syndrome peaked in the 6th and 7th decades of life, declining thereafter. Adiposity indicators were associated with diabetes, hypertension, and metabolic syndrome.

39 **Conclusion**

We observed an unexpectedly high prevalence of diabetes, hypertension, and metabolic syndrome in these marginalized agricultural communities. Transitional rural communities are increasingly vulnerable to the development of cardiometabolic risk factors and diseases. There is a need for targeted primary health strategies to reduce the burden of premature disability and death in these communities.

45

46 Introduction

Low and middle-income countries (LMICs) are undergoing the epidemiologic transition with a shift in the primary causes of death from those attributable to infectious and childhood diseases to chronic non-communicable diseases (NCDs) in the context of rapidly aging populations [1,2]. Among NCDs, age-adjusted prevalence of cardiovascular diseases peaked over the period 1960 to 1980 in high-income countries (HICs), while an epidemic of these diseases has now emerged in LMICs [3,4], which now account for more than 77% of all deaths attributed to NCDs globally [5].

The large increases in prevalence of NCDs in LMICs have been attributed to economic development and urbanization that have resulted in dramatic changes in where and how people live [6] with consequent changes in living conditions, diet, physical activity, and exposures to psychosocial stressors [7]. The emergence of the NCD epidemic in LMICs is having a major impact on utilization of scarce health care resources and poses a significant hurdle to achieving sustainable development goal targets for poverty reduction and prevention of premature death and disability [5].

Ecuador is an upper middle-income country in Latin America [8] with a diverse multiethnic population of 17 millions [9], life expectancy of 74 years, and a per capita gross domestic product of US\$6,395 in 2022 [10]. NCDs are now the major causes of death in Ecuador: among the five most important causes of death reported in 2019, 4 were NCDs including ischemic heart disease (14.7% of deaths), type 2 diabetes (T2D) (7.1%), cerebrovascular diseases (6.2%), and hypertension (HTN) (4.9%) [11].

66 The NCD epidemic is increasingly present in rural populations [12] through the influence of urbanization that extends even to the most isolated communities [13–15]. While a rural lifestyle is 67 68 presumed to reduce the risk of many NCDs because of potentially protective factors - such as 69 consumption of traditional diets high in complex carbohydrates, increased physical activity, and 70 greater social integration [16] - in many rural areas such protective exposures are being lost. 71 Indigenous and other vulnerable population groups living in transitional settings undergoing 72 environmental degradation, urbanization, and the loss of traditional lifestyles, are at increased risk 73 of cardiometabolic diseases [17].

We carried out a cross-sectional study of adults living in transitional rural communities of Montubios, a marginalized mestizo-derived ethnicity, in an ecologically vulnerable region of coastal Ecuador to estimate prevalence of cardiometabolic NCDs and associated risk factors. Our findings, showing a high prevalence of cardiometabolic diseases, highlight an unmet need for communitybased public health strategies to minimize the growing burden of premature death and morbidity from NCDs among marginalized and indigenous populations living in transitional rural settings.

80 Methods

81 Study design and population

82 We performed a cross-sectional study of adults living in ten adjacent rural agricultural communities 83 in Abdon Calderon parish, Portoviejo district, Manabí province, in tropical coastal Ecuador. The study 84 area (Fig 1) was traditionally inhabited by Montubios, a Spanish-speaking mestizo-derived ethnicity, 85 geographically restricted to the mountainous regions of coastal Ecuador, and who possess distinct 86 socio-cultural characteristics that distinguish them from the dominant mestizo population. 87 Montubios, along with Indigenous and Afro-Ecuadorians, belong to the poorest ethnic groups in 88 Ecuador and live in communities with important unmet health needs [18]. The study communities 89 were located 10 km from a primary health care facility and 30 km from a regional hospital, in a region 90 of Abdon Calderon parish considered to be highly vulnerable to environmental degradation, and 91 hence deprived of access to services and infrastructure investment. Initial contacts were made with 92 the parish council through local political leaders known to the study team and who acted as 93 gatekeepers. Community representatives were involved at all stages of the study including extensive 94 early meetings to inform study design, objectives, and selection of study site; organisation of 95 meetings to explain study objectives and obtain community consent for the study; planning and 96 organization for individual consent and data collection as well as involvement in the data collection 97 process itself; and community meetings to provide individual results and explain study findings. 98 Communities were selected by convenience sampling. Censuses for each community were drawn up 99 by community representatives specifically for this study. All adults aged 18 years and older living in 100 the communities were invited to participate through community assemblies. Exclusions were those

- 101 unwilling to provide informed written consent, pregnant women or those having given birth in the
- 102 previous 3 months. Recruitment took place during scheduled visits to study communities between 6
- 103 and 10 am from 31^{st} July to 12^{th} September 2021.
- 104
- 105 Fig 1. Geographical Context of the Study Area Showing from Left to Right, Locations in Ecuador of
- 106 Manabí Province, Portoviejo District, and Abdon Calderon Parish. The figure was designed using
- 107 shapefiles obtained from the Ecuadorian National Institute of Statistics (INEC),
- 108 (https://www.ecuadorencifras.gob.ec/documentos/web-
- 109 inec/Geografia Estadistica/Micrositio geoportal/index.html#cartograf-histor) and constructed
- 110 using QGIS version 3.14.
- 111

112 Data collection

- 113 Data and clinical samples were collected, and clinical measurements performed as follows using
- 114 standardized protocols: i) questionnaire based on STEPwise approach to NCD risk factor
- surveillance (STEPS) [19] to collect data on symptoms, treatments and diagnoses, and potential risk
- 116 factors; ii) anthropometric measurements including weight (TANITA model BF-60W, Japan), height
- 117 (SECA model 213, Germany), waist and hip circumferences, and bioimpedance (Bodystat model
- 118 1500, UK); iii) blood pressure seated after 15 minutes rest using electronic devices (OMRON model
- 119 7051T, Japan) with daily calibration against a mercury sphygmomanometer and cuffs selected
- 120 according to upper arm circumference. This was repeated after 5 minutes if systolic blood pressure
- 121 (SBP) >130 mmHg and/or diastolic blood pressure (DBP) >85 mmHg; and iv) blood after an

overnight fast to measure glucose, insulin, glycosylated hemoglobin (HbA1c), cholesterol, high-density lipoprotein (HDL), and triglycerides.

124

125 Laboratory analyses

HbA1c was measured with fresh anticoagulated venous blood using a fluorescent immunoassay at a field laboratory (iChroma[™], Boditech Med Incorporated, South Korea). Serum and plasma were separated by centrifugation and stored at -20C for later analysis at a central laboratory in Quito. Biochemical analyses (cholesterol, HDL, and triglycerides) were done using enzymatic colorimetry assays (Human Diagnostics, Germany). Insulin was analyzed using an immunoenzymatic assay (DIAsource ImmunoAssays, Belgium).

132

133 **Definitions**

134 Definitions for outcomes were: i) Type 2 diabetes (T2D) - clinical history of diabetes treatment 135 and/or glycosylated hemoglobin (HbA1c) \geq 6.5% [20]; ii) hypertension (HTN) – clinical history of 136 antihypertensive treatment and/or SBP ≥140 mm Hg or diastolic blood pressure DBP ≥90 mm Hg 137 during 2 separate measurements [21]; and iii) metabolic syndrome (MetS) – 3 or more of the 138 following (using the Harmonized criteria [22]: a) elevated waist circumference of ≥94 cm in men and 139 \geq 88 cm in women [23]; b) triglycerides \geq 150 mg/dL or treatment with triglyceride-lowering drugs; c) 140 reduced high density lipoprotein (HDL) <40 mg/dL in men and <50 mg/dL in women; d) elevated 141 blood pressure \geq 130/85 mmHg or treatment with antihypertensives; and e) fasting glucose \geq 100 142 mg/dL or treatment with glucose-lowering drugs.

143 Definitions for other conditions were: i) Adiposity indicators [24–26]: a) overweight (body 144 mass index \geq 25) and obesity (BMI \geq 30); b) abdominal obesity - waist circumference of \geq 94 cm in 145 men and \geq 88 cm in women [23]; c) elevated weight-to-height (WHtR) - \geq 0.5; d) body fat (free fat 146 mass) was determined using bioimpedance [27] with increased body fat defined as \geq 25% for men 147 and ≥30% for women; e) visceral adiposity index (VAI) was calculated using a model of adipose 148 distribution corrected for triglyceride and HDL levels [23] with high VAI according to age defined as 149 - age <30 – VAI >2.52; age ≥30 & <42 – VAI >2.23; age ≥42 & <52 – VAI >1.92; age ≥52 & <66 – VAI 150 >1.93; age \geq 66 – VAI >2; ii) elevated cholesterol \geq 200 mg/dl; iii) elevated triglycerides \geq 150 mg/dL; 151 iv) low HDL <40 mg/dl in men and <50 mg/dl in women; v) insulin resistance – Homeostatic Model 152 Assessment (HOMA) index >2.5 [28,29]; and vi) and previous medical diagnosis (personal history) of 153 heart attack or stroke. Occupation was categorised into 3 categories: agricultural workers, non-154 agricultural workers, and household chores. Ethnicity was self-identified and categorised into two 155 groups: mestizos and non-mestizos, the latter including Indigenous, Afro-Ecuadorians, Montubios, 156 and White.

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158

159 **Statistical analysis**

160 T2D, HTN, and MetS were analysed as binary outcomes, while blood pressure, glycemia, and 161 HbA1c were analysed as continuous variables. Because of the three-level hierarchical structure of 162 the data (individuals are nested in households and communities), multilevel regression techniques 163 were used to estimate standard errors and P values. We used age and sex data from the census 164 population in district of Portoviejo (S1 Fig) [9] to derive post-stratification weights to inflate

165 misrepresented groups and reduce effect of overrepresented groups in the sample[30]. We derived 166 prevalences (T2D, HTN, and MetS) and means (SBP, DBP, HbA1c and glycemia) stratified by sex, age 167 (up to power of three terms, i.e. non-linear), and their interactions when statistically significant 168 (P<0.05). We estimated age- and sex-adjusted effects of explanatory variables of interest on 169 prevalence of T2D, HTN, and MetS. Sensitivity analyses were performed for missing data but showed 170 consistent findings to those obtained using complete data. Analyses were exploratory and adjusted 171 only for age and sex rather than the construction of multivariable models. All analyses were done 172 using Stata 18 (version 18, Statacorp, College Station, Texas, USA).

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174 Ethics statement

Informed written consent was obtained from each participant and the study protocol was
approved by Bioethics Committee of the UTE University (Comité de Ética de Investigación en Seres
Humanos de la Universidad UTE, CEISH UTE 2019-1121-03) and by the Ecuadorian Ministry of Public
Health (MSP-DIS-2021-0393-O).

179 **Results**

180 Study population and characteristics

181 Of 1,525 adults eligible according to the house-to-house census, 1,010 adults were recruited 182 from 10 communities of which 931 (61.0% of those eligible) provided complete data and were 183 included in the analysis. Total and sex-stratified data on socio-demographic factors, indicators of 184 adiposity, biochemical measurements, and clinical history of NCDs are shown in Table 1. The mean 185 age of the study population was 44.7 years (standard deviation 18.1) and 57.5% were women. Most 186 participants lived with a partner (66.1%); 24.3% were functionally illiterate; 48.3% self-identified as 187 non-mestizo with 41.1% identifying as Montubio, 2.3% as Afro-Ecuadorian, and 0.5% as Indigenous. 188 Most (62.6%) men were agricultural workers while 80.9% of women were housewives; recent 189 alcohol consumption was reported by 62.2% (men 77.6% vs. women 50.8%); current smoking was 190 reported by 7.7% (men 17.2% vs. women 0.8%); men were more likely to do physically demanding 191 work and have contact with agrochemicals. Indicators of adiposity were more frequent in women 192 (for example, presence of abdominal obesity [men 49.5% vs. women 66.4%] and elevated body fat 193 [men 93.1% vs. women 97.9%]). Insulin resistance affected 58.6% of the study population and was 194 more frequent in women. Clinical histories of vascular events were reported by 8.7%. Of those with 195 a clinical history of diabetes or hypertension, only 25.8% had evidence of adequate glucose control 196 (S1 Table) and 34.9% had controlled blood pressure (S2 Table), respectively. A comparison of 197 participation and non-participation using our house-to-house census data showed an increase in 198 participation with age (OR 1.02, 95% CI 1.01-1.03) and among women (OR 5.4, 95% CI 3.7-7.8).

199 Table 1. Baseline frequencies of age and risk factors for chronic non-communicable diseases stratified by sex in 931 adults in a rural parish

200 of Manabi Province, Ecuador.

Variable	Category	Total (n=931)	Women (n=535)	Men (n=396)
		n (%)	n (%)	n (%)
Age distribution and risk factors				
Age group (years)	18-29	229 (24.6)	131 (24.5)	98 (24.8)
	30-39	158 (17.0)	100 (18.7)	58 (14.6)
	40-49	183 (19.7)	107 (20.0)	76 (19.2)
	50-59	149 (16.0)	82 (15.3)	67 (16.9)
	60-69	104 (11.2)	56 (10.5)	48 (12.1)
	≥ 70	108 (11.5)	59 (11.0)	49 (12.4)
Lives with partner	Yes	615 (66.1)	362 (67.7)	253 (63.9)
Functional illiteracy	Yes	226 (24.3)	114 (21.3)	112 (28.3)
Ethnicity	Non-mestizo	450 (48.3)	229 (42.9)	221 (55.8)
	Mestizo	481 (51.7)	306 (57.2)	175 (44.2)
Occupation	Agricultural worker	261 (28.1)	13 (2.4)	248 (62.6)
	Household chores	436 (46.8)	433 (80.9)	3 (0.8)
	Non-agricultural workers	234 (25.1)	89 (16.6)	145 (36.6)
Recent alcohol consumption	Yes	575 (62.2)	270 (50.8)	305 (77.6)
Current smoker	Yes	72 (7.7)	4 (0.8)	68 (17.2)
Physically-demanding work	Yes	319 (34.6)	104 (19.6)	215 (54.8)
Contact with agrochemicals	Yes	253 (27.2)	47 (8.8)	206 (52.0)
Indicators of adiposity and biochemical	risk factors			
Obesity	Not overweight (BMI <25)	322 (34.6)	164 (30.5)	159 (40.2)
	Overweight (BMI 25-29)	367 (39.4)	212 (39.6)	155 (39.1)
	Obese (BMI ≥30)	242 (26.0)	160 (29.9)	82 (20.7)
Abdominal obesity	men ≥94cm; women ≥88cm	551 (59.2)	355 (66.4)	196 (49.5)
Elevated WHtR	≥0.5	791 (85.0)	475 (88.8)	316 (79.8)
Elevated body fat	men (≥25%); women (≥30%)	696 (95.9)	412 (97.9)	284 (93.1)
Elevated VAI	See legend	571 (61.3)	368 (68.8)	203 (51.3)

Elevated cholesterol	≥200 mg/dl	445 (47.8)	270 (50.5)	175(44.2)
Elevated triglycerides	≥150 mg/dL	434 (44.6)	257 (48.0)	177 (44.7)
Low HDL	Men <40 mg/dl; women<50 mg/dl	524 (56.3)	367 (68.6)	157 (39.7)
Insulin resistance	HOMA>2.5	546 (58.6)	372 (69.5)	174 (43.9)
Elevated HbA1c	≥6.5%	210 (22.6)	133 (24.9)	77 (19.4)
Self-report of non-communicable diseases				
Vascular events (stroke or heart attack)	Yes	81 (8.7)	42 (7.9)	39 (9.9)
Diabetes	Yes	133 (14.3)	95 (17.8)	38 (9.7)
Hypertension	Yes	305 (32.9)	200 (37.4)	105 (26.7)
Elevated cholesterol	Yes	364 (39.2)	237 (44.3)	127 (32.3)

Functional illiteracy was defined as 3 or fewer years of formal schooling [29]. Abbreviations: WHtR – waist-to-height ratio; VAI – visceral adiposity index; HDL – high-density lipoprotein. High VAI - age <30 - VAI > 2.52; age $\geq 30 \& <42 - VAI > 2.23$; age $\geq 42 \& <52 - VAI > 1.92$; age $\geq 52 \& <66 - VAI > 1.93$; age $\geq 66 - VAI > 2$. Missing data: alcohol consumption (7); current smoker (1); physical demanding work (8); bioimpedance (increased fat %) (205); history of vascular events (3); history of hypertension (3); history of diabetes (3); and history of elevated cholesterol (3).

207 Population-weighted prevalence of diabetes, hypertension, metabolic

208 syndrome, and vascular events

209 Table 2 shows the crude and population-weighted prevalence of outcomes. Weighted 210 prevalence for T2D, HTN, and MetS were higher among females. Age-prevalence profiles for the 3 211 outcomes by sex are shown in Fig 2 and S1 Table. For T2D, prevalence was greater in women at all 212 ages (Fig 2C) and increased with age reaching a peak of 51% in women and 37% in men at around 75 213 years, after which it declined. Prevalence of HTN increased with age although there was some 214 evidence of an interaction with sex, particularly showing a steep increase in prevalence in women 215 (Fig 2B). For MetS, prevalence increased in both sexes until 57 years in men (70%) and 69 years in 216 women (86%) after which prevalence declined (Fig 2A). Prevalence of MetS was similar in both sexes 217 up to about 40 years, after which prevalence was greater in women. An analysis of 576 participants, 218 for whom blood pressure could be repeated at home, showed a lower average systolic blood 219 pressure in the home setting (average difference, 7.13 mm Hg [95% CI 5.76 - 8.51]) but similar 220 diastolic pressure (average difference 0.17 mm Hg [95% CI -0.82 - 1.16]).

Table 2. Crude and population-weighted prevalence of chronic diseases and means of blood pressure and glucose measures in study

222 population of 931 adults stratified by sex.

Chronic disease	-	Гotal (n=931)		Women (n=535)	Men (n=396)		
	Crude	Estimated weighted	Crude	Estimated weighted	Crude	Estimated weighted	
Binary outcomes	% (n)	% (95% CI)	% (n)	% (95% CI)	% (n)	% (95% CI)	
Type 2 Diabetes	26.3	20.4	29.5	24.5	22.0	15.8	
	(245)	(18.3-22.5)	(158)	(20.9-28.7)	(87)	(12.2-19.4)	
Hypertension	44.7	35.6	46.2	37.7	42.7	33.3	
	(416)	(29.0-42.1)	(247)	(31.0-44.4)	(169)	(26.8-39.7)	
Metabolic syndrome	58.0	54.2	60.9	56.2	54.0	52.1	
	(540)	(47.0-61.5)	(326)	(47.7-64.7)	(214)	(43.3-60.8)	
Vascular events	8.7	4.8	7.9	4.6	9.9	5.0	
	(81)	(2.7-6.9)	(42)	(2.8-6.4)	(39)	(2.2-7.8)	
	Crude	Estimated weighted	Crude	Estimated weighted	Crude	Estimated weighted	
Continuous outcomes	means (n)	means (95% CI)	means (n)	means (95% CI)	means (n)	means (95% CI)	
Systolic blood pressure (mm Hg)	134.0	131.0	132.8	129.2	135.8	132.9	
	(245)	(129.4-132.6)	(158)	(127.1-131.2)	(87)	(131.0-134.9)	
Diastolic blood pressure (mm Hg)	80.8	79.7	80.6	79.5	80.9	80.0	
	(416)	(78.8-80.6)	(247)	(78.3-80.6)	(169)	(78.7-81.2)	
Fasting glucose (mg/dL)	115.0	111.1	118.8	114.6	109.9	107.3	
	(540)	(108.3-113.8)	(326)	(110.3-119.0)	(214)	(104.5-110.1)	
HbA1c (%)	6.4	6.3	6.6	6.4	6.2	6.1	
	(81)	(6.2-6.4)	(42)	(6.3-6.6)	(39)	(6.0-6.2)	

223

Crude sample prevalence and means were derived from clinical history and measurements and weighted estimates (and 95% confidence intervals [CI]) accounted for household and community clustering and were weighted to the census population at district level (Fig S1).

226 Chronic diseases definitions: i) Type 2 diabetes - clinical history of diabetes treatment and/or glycosylated hemoglobin (HbA1c) ≥6.5%; ii)

hypertension – clinical history of antihypertensive treatment and/or SBP \geq 140 mm Hg or DBP \geq 90 mm Hg during 2 separate measurements

[21]; and iii) metabolic syndrome – using the Harmonized criteria [22]; and iv) vascular events- self-reported history of heart attack or

stroke.

230

232 Fig 2. Metabolic syndrome, HTN and diabetes: the age-dependent prevalence stratified by gender. Predictions were from logistic

regression of the outcomes on age, gender and their potential interactions accounting for family cluster and weighted for population

234 characteristics.

Risk exposures associated with diabetes, hypertension, and metabolic

236 syndrome

237 Age-and sex-adjusted associations between potential risk factors and disease outcome are 238 shown in Table 3. All outcomes were strongly positively associated with age. There were age-sex 239 interactions on the risks of HTN (P=0.003) and MetS (P=0.001). Among factors associated with T2D 240 were female sex (male vs. female, adj. OR 0.57, 95% CI 0.41-0.80, P=0.001), non-agricultural work 241 (vs. agricultural, adj. OR 1.74, 95% CI 1.01-2.98, P=0.045), HTN (adj. OR 2.09, 95% CI 1.40-3.11), and 242 abdominal obesity (adj. OR 2.85, 95% CI 1.85-4.39). Among factors associated with HTN were 243 household chores (vs. agriculture, adj. OR 3.67, 95% CI 1.71-7.90) or non-agricultural activities (vs. 244 agriculture, adj. OR 1.92, 95% CI 1.13-3.26); history of vascular events (adj. OR 2.45, 95% CI 1.38-245 4.35); T2D (adj. OR 1.95, 95% CI 1.31-2.90); abdominal obesity (adj. OR 3.12, 95% CI 2.15-4.54); and 246 insulin resistance (adj. OR 3.05, 95% CI 2.10-4.42). MetS was associated with living with a partner 247 (adj. OR 1.55, 95% CI 1.08-2.23), among other factors.

Table 3. Age-and sex adjusted associations between potential risk factors and type-2 Diabetes mellitus, hypertension, and metabolic

syndrome in 931 adults.

Risk factor	Category	Type-2 Diabetes		Hypertension			Metabolic Syndrome			
		n (%)	OR (95% CI)	P value	n (%)	OR (95% CI)	P value	n (%)	OR (95% CI)	P value
Socio-demographic										
Age			1.05 (1.04-1.07)	<0.001		1.10 (1.08-1.11)	<0.001		1.07 (1.06-1.09)	<0.001
Sex	Female	158 (29.5)	1		247 (46.2)	1		326 (60.9)	1	
	Male	87 (22.0)	0.57 (0.41-0.80)	0.001	169 (42.7)	0.80 (0.58-1.12)	0.193	214 (54.0)	0.72 (0.53-0.99)	0.04
Interaction: Age × Sex						0.97 (0.95-0.99)	0.003		0.97 (0.95-0.99)	0.001
Risk exposures										
Lives with partner	No	67 (21.2)	1		124 (39.2)	1		146 (46.2)	1	
	Yes	178 (28.9)	1.46 (1.00-2.13)	0.048	292 (47.5)	1.22 (0.81-1.84)	0.351	394 (64.1)	1.55 (1.08-2.23)	0.017
Functional Illiteracy	No	170 (24.1)	1		262 (37.2)	1		382 (54.2)	1	
	Yes	75 (33.2)	0.72 (0.47-1.09)	0.122	154 (68.1)	1.25 (0.83-1.88)	0.296	158 (69.9)	1.04 (0.67-1.60)	0.878
Ethnicity	Others	120 (26.7)	1		224 (49.8)	1		261 (58.0)	1	
	Mestizo	125 (26.0)	1.04 (0.75-1.45)	0.798	192 (39.9)	0.78 (0.55-1.10)	0.158	279 (58.0)	1.21 (0.90-1.64)	0.213
Occupation	Agricultural workers	55 (21.1)	1		111 (42.5)	1		142 (54.4)	1	
	Household chores	139 (31.9)	1.78 (0.87-3.66)	0.116	227 (52.1)	3.67 (1.71-7.90)	0.001	288 (66.1)	1.54 (0.80-2.95)	0.194
	Non-agricultural workers	51 (21.8)	1.74 (1.01-2.98)	0.045	78 (33.3)	1.92 (1.13-3.26)	0.017	110 (47.0)	1.24 (0.78-1.98)	0.370
Recent alcohol	No	122 (35.0)	1		198 (56.7)	1		234 (67.0)	1	
consumption										
	Yes	120 (20.9)	0.76 (0.52-1.10)	0.146	216 (37.6)	0.80 (0.57-1.12)	0.191	301 (52.3)	0.91 (0.66-1.26)	0.573
Current smoker	No	229 (26.7)	1		388 (45.2)	1		503 (58.6)	1	
	Yes	16 (22.2)	1.08 (0.49-2.38)	0.851	28 (38.9)	0.98 (0.77-1.26)	0.9	37 (51.4)	0.83 (0.44-1.56)	0.561
Physically demanding	No	162 (26.8)	1		269 (44.5)	1		350 (57.9)	1	
work										
	Yes	81 (25.4)	1.13 (0.77-1.67)	0.53	142 (44.5)	1.10 (0.75-1.61)	0.637	185 (58.0)	0.88 (0.61-1.26)	0.476
Contact with	No	185 (27.3)	1		310 (45.7)	1		397 (58.6)	1	
agrochemicals										
	Yes	60 (23.7)	0.93 (0.61-1.42)	0.741	106 (41.9)	0.72 (0.48-1.10)	0.126	143 (56.5)	0.90 (0.60-1.36)	0.631
History of chronic disease	es or risk factors									
Vascular events	No	217 (25.6)	1		360 (42.5)	1		485 (57.3)	1	
	Yes	27 (33.3)	1.22 (0.71-2.08)	0.467	55 (67.9)	2.45 (1.38-4.35)	0.002	54 (66.7)	1.35 (0.74-2.46)	0.330
Hypertension	No	110 (17.7)	1							
	Yes	134 (43.9)	2.08 (1.44- 3.01)	0.001						
Diabetes	No				312 (39.2)	1				
	Yes				103 (77.4)	3.01 (1.79-5.07)	<0.001			
Elevated cholesterol	No	114 (20.2)	1		213 (37.8)	1		284 (50.4)	1	
	Yes	130 (35.7)	1.71 (1.23-2.36)	0.001	203 (55.5)	1.56 (1.08-2.24)	0.018	255 (70.1)	1.62 (1.15-2.27)	0.005

Chronic diseases (histor	y and/or clinical measurements)									
Hypertension	No	79 (15.3)	1							
	Yes	166 (39.9)	2.09 (1.40-3.11)	<0.001						
Diabetes	No				250 (36.4)					
	Yes				166 (67.8)	1.95 (1.31-2.90)	0.001			
Nutritional and laborate	ory measures									
Obesity	Not overweight (BMI<25)	42 (13.0)	1		93(28.9)	1		80 (24.8)	1	
	Overweight (BMI 25-29)	113 (30.8)	2.29 (1.48-3.56)	<0.001	179 (48.8)	1.99 (1.28-3.10)	0.002	248 (67.6)	5.85 (3.86-8.88)	<0.001
	Obese (BMI≥30)	90 (37.2)	3.25 (1.96-5.40)	<0.001	144 (59.5)	4.20 (2.55-6.93)	<0.001	212 (87.6)	27.6 (15.8-48.2)	<0.001
Abdominal obesity	No	44 (11.6)	1		94 (24.7)	1				
	Yes (men≥94 cm; women≥88 cm)	201 (36.5)	2.85 (1.85-4.39)	<0.001	322 (58.4)	3.12 (2.15-4.54)	<0.001			
Elevated cholesterol	No	103 (21.2)	1		174 (35.8)	1		241 (49.6)	1	
	Yes (≥200 mg/dL)	142 (31.9)	1.09 (0.77-1.54)	0.629	252 (54.4)	1.28 (0.89-1.84)	0.175	299 (67.2)	1.37 (0.98-1.92)	0.069
Elevated triglycerides	No	99 (19.9)	1		186 (37.4)	1				
	Yes (≥150 mg/dL)	146 (33.6)	1.66 (1.20-2.32)	0.003	230 (53.0)	1.26 (0.90-1.76)	0.172			
Low HDL	No	93 (22.9)			174 (42.8)					
	Yes (men<40 mg/dL; women<50	152 (29.0)	1.32 (0.93-1.87)	0.126	242 (42.8)	1.40 (0.97-2.02)	0.069			
	mg/dL)									
Insulin resistance	No	46 (11.9)	1		123 (31.9)	1		127 (33.0)	1	
	Yes (HOMA>2.5)	199 (36.4)	3.77 (2.41-5.90)	<0.001	293 (53.7)	3.05 (2.10-4.42)	<0.001	413 (75.6)	8.77 (6.15-12.51)	<0.001

²⁵⁰

251 Odd ratios (ORs) and 95% confidence intervals were estimated using logistic regression and accounting for household and community 252 clustering and weighted for population structure. Age² was strongly associated with all 3 outcomes: type-2 Diabetes (OR 0.9986, 95% CI 253 0.9982-0.9990, P=0.001), hypertension (OR 0.9993, 95% CI 0.9988-0.9999, P=0.028), and metabolic syndrome (OR 0.9992, 95% CI 0.9986-254 0.9997), P<0.001). Associations between outcomes and risk exposures, clinical history, and laboratory measures were adjusted for the 255 sociodemographic variables shown in Table 1 (including age and age² [all outcomes], sex [all outcomes], and the interaction between age 256 and sex [hypertension and the metabolic syndrome]). Functional illiteracy was defined as 3 or fewer years of formal schooling [31]. 257 Outcome definitions: i) Type 2 diabetes - clinical history of diabetes treatment and/or glycosylated hemoglobin (HbA1c) \geq 6.5%; ii) 258 hypertension – clinical history of antihypertensive treatment and/or ≥140 mm Hg or diastolic blood pressure ≥90 mm Hg during 2 separate 259 measurements [21]; and iii) metabolic syndrome – using the Harmonized criteria [22]; and iv) vascular events- self-reported history of 260 heart attack or stroke. Abdominal obesity was measured using waist circumference. Missing data: alcohol consumption (7); current smoker 261 (1); physical demanding work (8); Missing data: alcohol consumption (n=7); current smoker (1); physical demanding work (8); history of: 262 vascular events (3), respiratory disease (3), kidney disease (3), elevated cholesterol (3), diabetes (3) and hypertension (3); and 263 bioimpedance (increased fat %) (205). Abbreviation: BMI – body mass index; HDL – high-density lipoprotein; HOMA - Homeostatic Model 264 Assessment index.

Associations between indicators of adiposity and measurements of systolic and diastolic blood pressure, fasting glucose, and glycosylated hemoglobin

268 Means of systolic blood pressure (SBP) and diastolic blood pressure (DBP), fasting capillary 269 blood glucose and glycosylated hemoglobin (HbA1c) in the study population and stratified by sex are 270 shown in Table 2.

271 There were significant interactions between age and sex for all 4 measurements (P<0.005) 272 (Fig 3 and S3 Table). Predicted values for SBP increased with age being greater in men up to 50 273 years after which values were greater in women (Fig 3A). Predicted DBP increased with age up to 274 50 years and then declined, showing a similar sex interaction as for SBP (Fig 3B). Predicted values 275 of fasting glucose increased with age reaching a plateau after 50 years (Fig 3C): levels were similar 276 in both sexes up to 30 years after which levels diverged becoming greater in women. A similar 277 pattern was seen for HbA1c (Fig 3D). Strong associations were observed between a variety of 278 adiposity indicators and blood pressure levels and glucose measurements (S3 Table). 279 280 Fig 3. SBP, DBP, glycemia and HbA1c predicted levels stratified by gender. Predictions were from

- regression on age, gender and their potential interactions accounting for family cluster and
- 282 weighted for population characteristics.

284 **Discussion**

285 In this cross-sectional study, we estimated prevalence and associated risk factors for 286 cardiometabolic NCDs in adults living in transitional rural communities in tropical Ecuador. These 287 communities, located in an ecologically vulnerable region of coastal Ecuador, were traditionally 288 inhabited by a marginalized population group, the Montubios. Our data showed a high weighted 289 prevalence of T2D, HTN, and MetS affecting 20.4%, 35.6%, and 54.2%, respectively, of inhabitants in 290 these communities. The majority (74.2%) of those with a doctor diagnosis T2D had poorly controlled 291 disease and 14% were unaware of their condition. Similarly, over one-guarter of those with HTN 292 (26.6%) were unaware of their diagnosis before the study. Over 8% of the study population had a 293 history of vascular events (stroke or heart attack). NCDs were more common in females, and largely 294 increased with age, becoming very common at extremes of age (e.g. 70% with HTN by age 80).

295 Weighted prevalence of T2D, HTN and MetS observed here was greater than might be 296 expected from the findings of previous surveys from Ecuador [29,32,33] and elsewhere in Latin 297 America [34-36]. Previous studies have estimated T2D prevalence of 8.7% in the Latin American 298 region [34] and 4.7% in adult Ecuadorians. [37] The most recent national survey from 2018 estimated 299 a T2D prevalence of 7.1%, indicating a likely temporal trend of increasing prevalence among 300 Ecuadorian adults [32]. Other studies have shown that marginalized population groups, particularly 301 indigenous populations, appear to be particularly vulnerable to high rates of T2D: 70% of studies in 302 indigenous populations evaluated in a recent systematic review reported prevalence greater than 303 10%[38]. A recent study of Afro-Ecuadorian adults living in conditions of severe poverty in a rural 304 region of coastal Ecuador, estimated a T2D prevalence of 6.8% [39]. Although differences in 305 estimated prevalence between studies could be explained partly by differences in disease

definitions, mean age of study population, and study design, our estimate of 20.4% wasunexpectedly high.

308 The age-standardized prevalence of HTN declined over the period 1975-2015 in HICs while 309 tending to increase in many LMICs with an overall doubling of numbers of people with HTN globally 310 because of population growth and ageing [6]. Prevalence of HTN in Latin America and the Caribbean 311 declined over the same period [6] although age-standardized prevalence remained high (35.4%) in 312 the region in 2019 [35]. Our estimate of 35.6% HTN prevalence is close to this value and is similar to 313 that reported among Afro-Ecuadorians (36%), another marginalized group living in rural transitional 314 coastal communities [15] among whom HTN was considered the most important cause of death [40]. 315 However, our estimate of HTN prevalence was more than double that reported among adults living 316 in urban and rural settings elsewhere in the country [29,41,42].

317 MetS is known to predict sudden death [43] and increases markedly the risk of cardiovascular 318 diseases [44] - even one to two components of the syndrome have been associated with increased 319 mortality [45,46]. A population-based survey of adults from 7 Latin American cities including Quito 320 estimated prevalence of MetS ranging from 14% in Quito to 27% in Barquisimeto, Venezuela [36]. 321 Within Latin America a high prevalence of MetS has been reported among indigenous populations 322 including 46.7% in the Amazon region of Venezuela [47] and 37% in Brazil, although in Brazil the 323 prevalence in 6 surveys of Indigenous groups varied markedly between 11% in Parana and 66% in 324 Matto Grosso [48]. Our estimate of 54.2% was similar to that obtained (55.7%) in a house-to-house 325 survey of adults living in a (Cholo-)Montubio community in the southern coastal region of Ecuador 326 [49], but greater than that reported (42%) in urban and rural areas in the central Andean region of 327 the country [50] and in a national survey (31.2%) [33]. Differences in prevalence estimates within

Ecuador could represent differences in definitions, sampling methods, as well as differing population
 risks for individual components of the syndrome.

330 Rural communities undergoing rapid changes relating to urbanization processes are of 331 particular relevance for studies examining potential causal links between changes in risk behaviors, 332 development of risk factors, and emergence of cardiometabolic disease. The coastal region of 333 Ecuador, where we did this study, is of particular interest because of unique geographic, cultural, 334 and socio-economic characteristics, likely to affect distributions of risk factors and NCD prevalence. 335 Montubios are recognized in Ecuador as a distinct mestizo-derived ethnicity and represent a 336 historically excluded group living in rural, often low mountainous and isolated regions (<500 m) of 337 coastal Ecuador [51,52]. Montubio communities adhere closely to traditional values, being a 338 patriarchal culture based on tight kinship ties, conservative values, and a close relationship with the 339 land. However, Montubio communities are presently undergoing a rapid transition to a more 340 westernized lifestyle [53], and many communities are found in ecologically vulnerable environments. 341 Traditionally, the inhabitants of these communities lived as agriculturalist (as day-laborers or 342 cultivators of smallholdings) with seasonal work on large coffee plantations. The region consists of 343 mountainous terrain currently undergoing rapid degradation relating to deforestation and soil 344 erosion, resulting in poorly fertile soils for agriculture. There is a high vulnerability to rapid runoff 345 and landslides and the area has been designated as of extreme ecologic risk by the district council 346 and excluded from government and municipal investment in infrastructure and basic services 347 including health care. These factors likely have contributed to a process of acculturation, likely 348 accelerated by occupational shifts from the land to services generally provided outside the 349 communities; the effects of the migration of the young in search of employment either temporarily 350 or permanently; and with the introduction of electricity, the impact of technological innovations

351 such as television and digital media. Such changes have occurred over a period of less than a 352 generation (road access and electricity were introduced 15-20 years previously) and have been 353 accompanied by increased sedentarism and a shift in diet to include increasing quantities of 354 processed foods high in fat, sugar, and salt [29,54] that can be acquired cheaply in local stores.

355 Several studies have shown rural populations in LMICs to have healthier cardiometabolic 356 profiles than urban residents, an effect attributed to greater physical activity more than dietary 357 effects [55–57]. The sedentarization of rural occupations likely has led to changes in cardiometabolic 358 risk factors profiles and disease risk to resemble urban populations. The prevalence of vascular 359 events (8.7% with stroke or heart attack) observed here is somewhat lower than reported from 360 another Montubio population further south in Coastal Ecuador where stroke prevalence increased 361 from 14.1% to 35.2% between 2003 and 2012 [49,58]. Possible explanations are poorer survival rates 362 or the more recent emergence of an unhealthy profile of cardiometabolic risk factors in our 363 population. Environmental degradation may have compounded such effects through more rapid 364 shifts in occupation, diets, and activity levels. Rapid changes in diet and physical activity affecting a 365 population exposed to chronic under-nutrition in childhood likely will saturate metabolic capacity in 366 adulthood [59] and increase vulnerability to the premature development of cardiometabolic risk 367 factors and disease.

Indigenous groups in Latin America appear to be at particular risk of cardiometabolic NCDs particularly diabetes [38] in the context of acculturation, deforestation, and environmental degradation [17]. Other rural marginalized groups might be expected to suffer similar risks although available data are limited [60]. An additional challenge for these populations is inadequate health care access which further contributes to the burden of disability and premature death. Lack of access to health care results in delays in diagnosis and limits treatment access. Even where there is health

374 access, a lack of education and financial resources often limits availability of and adherence to 375 treatment which often needs to be provided regularly for life. This was seen in the present study by 376 the observation that greater than 65% of those with a previous doctor diagnosis of T2D or HTN had 377 poorly controlled glucose or blood pressure, respectively. Lower socioeconomic conditions in LMIC 378 populations have been consistently linked to cardiometabolic NCD prevalence as well as to MetS or 379 its individual components [3,39,61–63]. Further, a rational public health strategy for NCD control 380 and prevention in resource-poor rural populations will require community-based interventions, 381 ideally through primary health care (or community health worker networks) where available and 382 targeted to the specific risk factors and pathologies prevalent in specific populations [64,65].

383

384 Limitations

385 Data collection targeted a marginalized population living in a restricted geographic setting 386 where we believed there to be major unmet health needs. The data can be generalized to similar 387 populations living in coastal Ecuador in conditions of extreme ecologic and social vulnerability. Non-388 participation of healthier members of the communities could have led to overestimation of 389 estimates, likely related to more active, healthier individuals being away for work. Survey non-390 participation was largely explained by absence of working-age adults (20-40 years), particularly men, 391 because of salaried work commitments outside the communities. Under-representation of adults 392 aged 20-40 years that was apparent based on a comparison of age-structure between our house-to-393 house-census and the projected census population for the district, was adjusted for using weighted 394 estimates. The use of questionnaires to collect data on clinical histories and risk factors may be 395 subject to recall or social desirability bias although we used standardized clinical and laboratory396 measurements to measure risk factors and outcomes where possible.

397

398 **Conclusion**

399 We observed a high prevalence of T2D, HTN, and MetS in marginalized Montubio 400 communities in a rural region of coastal Ecuador. Environmental degradation and accelerated 401 urbanization occurring in these transitional communities over a period of approximately 20 years, 402 with increased out-migration, occupational shifts, and changes in diet and physical activity are likely 403 to be important causes of these NCDs. Our data indicate that marginalized rural populations 404 undergoing a rapid transition from traditional to more modern lifestyles in Ecuador and elsewhere 405 in Latin America suffer an unacceptable burden of preventable morbidity. This burden might be 406 reduced by targeted public health strategies including the training and adequate resourcing of 407 community health workers.

408

409 Acknowledgments

The authors thank participant communities and their representatives for their co-operation,
and the support of technicians and health professionals from Universidad Internacional del
Ecuador and the CAMERA study.

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621 Supporting information

622	S1 Fig: Age distribution in the study sample compared to that of the Portoviejo population,
623	stratified by gender. Data were obtained from the 2010 census [9].
624	
625	S1 Table. Clinical history of diabetes and presence of hyperglycemia (Hb1Ac>=6.5%) at time of
626	survey.
627	
628	S2 Table. Clinical history of hypertension and presence of elevated blood pressure (>140/90 mm
629	Hg) at time of survey.
630	
631	S3 Table. Age-and sex adjusted associations between indicators of adiposity and systolic and
632	diastolic blood pressure (mm Hg), glycosylated hemoglobin (HbA1c) (%), and fasting glucose
633	(mg/dL) in 931 adults. Estimates and 95% confidence intervals (CI) were derived from linear
634	regression analyses and accounted for household and community clustering and were weighted for
635	population structure. Associations, i.e. the adjusted mean differences between groups defined by
636	indicators of adiposity were adjusted for the sociodemographic variables shown in the Table
637	(including age and age ² [all outcomes], age ³ [HbA1c and fasting glucose], sex [all outcomes], and
638	the interaction between age and sex [all outcomes]). Abbreviations: BP – blood pressure; WHtR –
639	waist-to-height ratio; VAI – Visceral adiposity index. Definitions for indicators of adiposity:
640	Overweight (BMI≥25); abdominal obesity (waist circumference of ≥94 cm in men and ≥88 cm in
641	women); increased body fat (men (≥25%, women (≥30%); WHtR (≥0.5); elevated VAI (age <30 – VAI
642	>2.52; age ≥30 & <42 – VAI > 2.23; age ≥42 & <52 – VAI >1.92; age ≥52 & <66 – VAI > 1.93; age ≥66
643	– VAI > 2). Missing data: increased fat (205).

S1 Data. Raw data used for analyses.

647 Data availability

648 The data analyzed in this study are provided within the supporting information.





