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3	Psychosocial stress and central adiposity: A Brazilian study with users of the
4	public health system
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29 Abstract

Objective: To assess the association between indicators of psychosocial stress
 and central adiposity in adult users of the Unified Health System (SUS) from
 Southeast of Brazil.

33 Methods: This cross-sectional study was conducted with 384 adults (20 to 59 years old) from the city of Alegre, Southeastern Brazil. The simple random 34 sample represented the population using the public health system of the 35 36 municipality. The prevalence of obesity was based on the Body Mass Index, and central adiposity (dependent variable) was measured by waist 37 circumference in centimeters. The independent variables were the following 38 indicators of psychosocial stress: food and nutrition insecurity (yes/no), serum 39 cortisol (µa/dL), symptoms suggestive of depression using the Beck Depression 40 41 Inventory-II \geq 17 (yes/no), and altered blood pressure \geq 130/85 mmHg (yes/no). Univariate linear regression was performed between central adiposity and each 42 stress indicator, and later the models were adjusted for socioeconomic, health, 43 44 and lifestyle variables. All analyses were stratified by rural and urban location. **Results:** The prevalence of weight excess was 68.3%, and 71.5% of individuals 45 presented an increased risk for metabolic complications related to central 46 47 adiposity. Mean waist circumference scores for the rural and urban population were 89.3 ± 12.7 cm and 92.9 ± 14.7 cm, respectively (p = 0.012). Indicators of 48 stress that were associated with central adiposity were: cortisol in the rural 49

50	population and altered blood pressure in the urban population. This occurred
51	both in the raw analysis and in the models adjusted for confounding factors.
52	Conclusion: The associations between stress and adiposity were different
53	between rural (cortisol - inverse association) and urban (altered blood pressure)
54	lifestyles, confirming the influence of local and psychosocial subsistence on the
55	modulation of stress and on how individuals react or restrain stressors. Stress
56	reduction strategies can be useful in public health programs designed to
57	prevent or treat obesity.
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59	Keywords: psychosocial stress, central adiposity, obesity, cortisol.
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62 Introduction

Overweight and obesity are defined as abnormal or excessive 63 accumulation of fat that can be detrimental to health.(1) Weight excess affects 64 all regions of the world and is now appearing as a global epidemic. According to 65 the World Health Organization (WHO), 38.9% of the world population aged 18 66 or more present weight excess and of these, 13.1% are obese.(2) Compared to 67 other WHO regions, the prevalence of weight excess is higher in the Americas 68 (62% for overweight in both sexes, and 26% for obesity in adults over 20 years 69 of age).(3) The prevalence is higher in the United States of America, Mexico, 70 71 and Chile, where weight excess affects between six and seven out of 10 adults.(3, 4) In Brazil, the National Health Survey (NHS) presented a similar 72 scenario, with 56.9% of overweight adults.(5) 73

The term stress has already been mentioned in the literature over the 74 75 years and designates all the non-specific effects of stressors or factors that may act on the body.(6, 7) Stress can be established in the individual in an acute or 76 chronic way, manifested through changes in serum cortisol. Chronic stress can 77 lead to changes in the Hypothalamic-Pituitary-Adrenal (HPA) axis, resulting in 78 altered serum cortisol levels.(8, 9) The frequency of exposure to various 79 stressors can trigger a general adaptation syndrome, composed of three 80 phases: alarm reaction, adaptation phase, and exhaustion phase. (6, 10, 11) 81 The manner in which an individual responds to environmental, economic, social, 82 83 and health adversities is particularly differentiated, and when the possibilities for 84 adaptation are overcome, psychosocial stress becomes a threat to well-85 being.(8, 10, 12)

With cultural changes, globalization, and consequent acceleration of the 86 pace of life, there has been an increase in diseases resulting in psychosocial 87 stress and chronic diseases such as obesity, leading concern to public health 88 authorities.(1, 13-19) The relationship between obesity and psychosocial 89 90 disorders is already well established.(8, 11, 12, 20-22) However, the way 91 different cultures react to the environment and the cause and effect relationship 92 between obesity and stress may be different; these facts along with the issues involved, create a complex concern for authorities. Therefore, this study aimed 93 94 to evaluate the association between indicators of psychosocial stress and central adiposity in adult users of the Unified Health System (SUS) in a city in 95 the Southeast of Brazil. 96

97

98 Materials and Methods

99 Study design, sampling, and data collection

100	This was a cross-sectional study with adults (20 to 59 years, SUS users),
101	which was conducted in Alegre, a city located in the Southeast region of Brazil.
102	It is a municipality with the largest population in the micro-region south of the
103	State of Espirito Santo, Brazil, which has a Human Development Index similar
104	to that of the country (HDI = 0.721 and 0.727, respectively), according to the
105	United Nations Development Program (UNDP).(23)

A population of 10222 individuals, registered in the Primary Care Network of Municipal Health was considered in the study design. For the calculation of the simple random sample, absolute accuracy of 5%, 95% confidence interval, design effect equal to 1 and, in the absence of specific studies in the region, it was estimated that the prevalence of overweight individuals was around 50%. Finally, 10% of losses were added and the sample size calculated was 409 individuals.(24)

The study was approved by the Ethics Committee on Human Research, the Health Sciences Center, Federal University of Espirito Santo, under number 1,574,160 issued on June 03, 2016. The inclusion criteria for participation in the study were the following: not being pregnant, having no cognitive conditions that would interfere with the response to the questionnaires, and declare free consent for participation.

Data collection was performed through an individual interview, with questionnaires that evaluated socioeconomic, health and lifestyle conditions, and food and nutrition insecurity (FNiS), as well as symptoms suggestive of depression. Anthropometric, blood pressure, and blood samples were also collected for cortisol analysis.

The socioeconomic, health, and lifestyle questionnaires were elaborated 124 125 based on the Individual and Domiciliary Registry Files of SUS, Ministry of Health, Brazil. FNiS was evaluated by the application of the Brazilian Food 126 Insecurity Scale (BFIS)(25, 26), a validated instrument comprising 14 questions, 127 128 aimed at families from the same household with and without members under 18 years of age; concerns evaluated were: lack of food at home, and having some 129 130 member of the family spending a whole day without eating in the last three months, among others. The degree of severity of FNiS (mild, moderate, and 131 severe) was grouped in this study. 132

Symptoms suggestive of depression were investigated using the Beck Depression Inventory-II (BDI-II), and total scores were categorized according to the literature: normal, mild, moderate, and severe.(27-29) For the purpose of this study, the following regrouping was used: normal or mild mood disorder (BDI-II <17), and symptoms suggestive of depression (BDI-II \ge 17).

The anthropometric evaluation was performed by qualified professionals 138 in the morning, after participants had fasted for a minimum of eight hours, 139 140 following the technical standards of the Food and Nutritional Surveillance 141 System (SISVAN).(30) Stature was evaluated using Alturexata® stadiometer. with a maximum capacity of 2.10 m and accuracy of 0.5 cm. The weight was 142 measured on a Tanita® bipolar bioimpedance balance, with a BC601® branded 143 144 body fat monitor (with 100 g division and maximum capacity of 150 kg). The Body Mass Index (BMI) was calculated and classified according to WHO 145 146 reference for adults(1), in which low weight individuals present BMI values < 18.5 kg/m², eutrophic range from 18.5 to 24.9 kg/m², overweight from 25.0 to 147 29.9 kg/m², and obese individuals BMI being \geq 30.0 kg/m². Waist circumference 148

(WC) was measured using an inextensible anthropometric tape TBW® (1.5 m 149 150 and 0.1 cm accuracy), with reference to the midpoint between the last rib and 151 the iliac crest. (31, 32) This study considered that increased risk of metabolic complications for men was associated with abdominal obesity with WC values ≥ 152 153 94 cm; and very high risk with WC \geq 102. For women, an increased risk of metabolic complications was considered at/with WC values \geq 80 cm, and very 154 155 high risk, with WC values \geq 88 cm.(1, 32) Blood pressure was measured using an aneroid sphygmomanometer 156 with a G-TECH® Premium model. Both the measurement technique and the 157

blood pressure classification were based on the VI Brazilian Guidelines for
Hypertension from the Brazilian Society of Cardiology (SBC) for individuals 18
years and older(33), considering altered blood pressure (AP) when values are ≥
130/85 mmHg.

Blood collection for cortisol analysis was performed in the morning,
between 7:00 and 9:00 a.m. The samples were collected in a separator gel
tube, kept at room temperature until the clot was retracted, then centrifuged at
2500 rpm for 10 min and refrigerated at 2 ° C to 8 ° C until analysis. The
concentration of cortisol was quantified by the chemiluminescence method and
the reference values for the morning dose were 6.7 to 22.6 µg / dL.(34)

168

169 Data analysis

Data were tabulated and submitted for consistency analysis. Individuals who presented inconsistency of anthropometric data and those who reported the use of corticosteroid medications were excluded from the analysis.

173	For the descriptive analysis, the variables were presented as medians
174	and interquartile ranges, or means and standard deviations (non-parametric or
175	parametric data, respectively, according to Kolmogorov-Smirnov normality test),
176	besides frequencies and proportions. The Mann-Whitney U test or Student's t-
177	test and the chi-square test were used to verify the differences related to the
178	rural and urban locations, considering a level of significance of 5%. In the chi-
179	square test, for the variables that had three or more categories and that
180	presented significant differences, 2 x 2 ratio analysis was performed later, using
181	the Bonferroni correction, which changes the level of significance (p), to avoid
182	type I errors derived from multiple comparisons. After this procedure, the
183	corrected significance level was p<0.016.(35)
184	Univariate linear regression models were used to evaluate the
185	association between stress indicators and central adiposity. All statistical
186	analyses were performed using SPSS $^{ m R}$ software, version 15.0 for Windows
187	(IBM, Munich, Germany).
188	The univariate linear regression analysis consisted of simple models
189	(crude analysis) hierarchically adjusted in three different models: Model 1,
190	analysis adjusted by socioeconomic variables; Model 2, adjusted by
191	socioeconomic and health variables; and Model 3, adjusted by socioeconomic,
192	health, and lifestyle variables.
193	
194	Outcome variable: Central adiposity
195	In this study, the WC in centimeters was chosen as an outcome variable
196	to predict central adiposity from the explanatory variables: it was included in the

to predict central adiposity from the explanatory variables; it was included in themodels as a continuous variable.

198

199 Independent variables

200	The independent variables selected for the univariate analyses were:
201	FNiS (yes/no), serum cortisol (μ g/dL), symptoms suggestive of depression with
202	BDI-II \ge 17 (yes/no), and AP \ge 130/85 mmHg (yes/no), which are directly or
203	indirectly related to psychosocial stress. All these indicators were measured by
204	methods and/or validated instruments with good reproducibility in previous
205	studies.

206

207 Potential confounding variables

Socioeconomic variables (age, sex, schooling, and income classification), health (stress report, one or more comorbidities reported, and self-rated health), and lifestyle (current smoking, current alcohol consumption, and weekly physical or leisure activity) were chosen to adjust the univariate linear regression models based on the knowledge collected from the preexisting literature regarding the relation between psychosocial stress and overweight.(7, 8, 12, 14, 36)

215 Among the socioeconomic variables, it is highlighted that schooling was evaluated in years of study and income was classified according to the Center 216 for Social Policies of the Getúlio Vargas Foundation, which considers low-217 income individuals with a per capita income of less than US\$ 5 per day.(37) In 218 the case of the variable comorbidities, the response options were diabetes, 219 220 hypertension, cardiac palpitation, cardiovascular disease, and metabolic syndrome, conditions known to be associated with obesity and physical 221 stress.(8) The practice of physical activity was added to leisure, and, as a 222

- reference, it was considered that once a week was the minimum frequency to
- "relief/escape" from the stressful routine, but not to be seen as the ideal.
- 225

226 Modeling

- Beyond the WC (outcome variable), the variables age and serum cortisol
- 228 entered the models as continuous variables, and all others as categorical and
- dichotomic, as shown in Table 1.

230

Table 1. Treatment of variables for modeling

riables	Dichotomization
ependent variable	
WC (cm)	continuous variable
dependent variables / Stress indicators	
FNiS	yes* / no
Serum cortisol (µg/dL)	continuous variable
BDI-II ≥ 17	yes* / no
AP ≥ 130/85 mmHg	yes* / no
djustment variables	
Socioeconomic conditions	
Age	continuous variable
Sex	female* / male
Sex Years of study	female* / male < 9* / ≥ 9
Years of study	< 9* / ≥ 9
Years of study Income classification (< \$5.00/day)	< 9* / ≥ 9
Years of study Income classification (< \$5.00/day) Health conditions	< 9* / ≥ 9 low income* / non-low income
Years of study Income classification (< \$5.00/day) Health conditions Stress report	< 9* / ≥ 9 low income* / non-low income yes* / no

Current smoking	yes* / no
Current alcohol consumption	yes* / no
Physical or leisure activities (≤ 1 time/week)	yes* / no

* indicates the reference category. WC: waist circumference; FNiS: Food and Nutritional Insecurity; BDI-II: Beck

233 Depression Inventory-II; AP: Amended Pressure.

234

235 The univariate analyses for the rural and urban populations were performed separately, to evaluate the differences in the behavior of the 236 predictor variables regarding the stratification of the models by location. All 237 238 variables used in the models met the assumption of collinearity. The assumptions of normality, linearity, homoscedasticity, and independence of 239 residues were met in all models that had p <0.05 in the F test. Results are 240 presented as non-standard regression coefficients (β), and their respective 95% 241 242 confidence intervals (CI) and p-values related to the explanatory variable. All 243 analyses were performed using the complete sample, and the significance level was 5% (p < 0.05). 244

245

246 **Results**

A total of 384 individuals participated in the study, 75 men and 309 women, corresponding to 19.5% and 80.5%, respectively. Among the participants, 133 were rural residents, and 251 lived in urban areas. The median age was 42.5 years (Interquartile Ranges - IR=18.0), and there was no significant difference between the urban and rural populations (p=0.82) or between men and women (p=0.23). The prevalence of overweight was 33.1%, and that of obesity was 35.2%.

254 Regarding the risk of metabolic complications related to central adiposity (WC,

in cm), 51.6% of the sample presented increased risk and the risk of 19.9%
substantially increased. The median BMI and WC mean scores were higher in
the urban than in the rural populations. The other characteristics related to
health, socioeconomic conditions, and lifestyle of the population are presented
in Table 2.

260

Table 2. Characteristics of the population according to rural and urban locations

Characteristics	All	Rural	Urban	р*
Health profile				
Nutritional Status - BMI				
Low weight	1.6 (6)	1.5 (2)	1.6 (4)	0.116*
Eutrophy	30.2 (116)	33.1 (44)	28.7 (72)	
Overweight	33.1 (127)	36.8 (49)	31.1 (78)	
Obesity	35.2 (135)	28.6 (38)	38.6 (97)	
BMI (kg/m²) – median (IR)	27.5 (7.8)	26.8 (7.3)	28.4 (8.1)	0.027
Risk metabolic complications - WC				
Low	28.5 (109)	31.8 (42)	26.8 (67)	0.084*
Increased	19.9 (76)	24.2 (32)	17.6 (44)	
Substantially increased	51.6 (197)	43.9 (58)	55.6 (139)	
WC (cm) - mean (±SD)	91.7 (±14.1)	89.3 (±12.7)	92.9 (±14.7)	0.012
Classification of cortisol - % (n)				
Low	10,0 (37)	15,0 (19)	7,4 (18)	0,042*
Normal	87,8 (324)	81,9 (104)	90,9 (220)	
High	2,2 (8)	3,1 (4)	1,7 (4)	
Serum cortisol (µg/dL) - median (IR)	12.1 (6.1)	10.8 (5.5)	12.9 (6.2)	0.006
Depression - % (n)				
BDI-II < 17	76.4 (272)	77.5 (93)	75.8 (179)	0.728
BDI-II ≥ 17	23.6 (84)	22.5 (27)	24.2 (57)	
AP ≥ 130/85 mmHg - % (n)				
No	79.8 (296)	87.5 (112)	75.7 (184)	0.007

Yes	20.2 (75)	12.5 (16)	24.3 (59)	
Stress report - % (n)				
No	23.2 (89)	16.7 (22)	26.6 (67)	0.029
Yes	76.8 (295)	83.3 (110)	73.4 (185)	
Report of comorbidities - % (n)				
No report	65.7 (251)	70.2 (92)	63.3 (159)	0.179
One or more comorbidity	34.3 (131)	29.8 (39)	36.7 (92)	
Use of continuous medication - % (n)				
No	45.1 (173)	46.6 (62)	44.2 (111)	0.630
Yes	54.9 (211)	53.4 (71)	55.8 (140)	
Self-rated health - % (n)				
Good or very good	46.9 (180)	42.1 (56)	49.4 (124)	0.173
Regular or poor	53.1 (204)	57.9 (77)	50.6 (127)	
Socioeconomic profile				
BFIS Classification - % (n)				
FNS	58.2 (223)	60.6 (80)	57.0 (143)	0.493
FNiS	41.8 (160)	39.4 (52)	43.0 (108)	
Age (years) - median (IR)	42,5 (18,0)	42,0 (18,0)	43,0 (18,0)	0,820
Skin color - % (n)				
White	54.7 (205)	57.3 (75)	53.3 (130)	0.461
Non-white	45.3 (170)	42.7 (56)	46.7 (114)	
Marital status - % (n)				
Single	24.7 (95)	25.6 (34)	24.3 (61)	0.819
Not single	75.3 (289)	74.4 (99)	75.7 (190)	
Years of study - % (n)				
≥ 9 years	54.0 (204)	40.9 (54)	61.0 (150)	<0.001
< 9 years	46.0 (174)	59.1 (78)	39.0 (96)	
Income classification - % (n)				
Non-low income (≥ \$5.00/ day)	58.5 (224)	45.1 (60)	65.6 (164)	<0.001
Low income (< \$5.00/ day)	41.5 (159)	54.9 (73)	34.4 (86)	
Lifestyle	. ,	. ,	. ,	
Physical or leisure activities - % (n)				
≥ 1 time/ week	86.1 (322)	84.9 (107)	86.7 (215)	0.640
		- ()	()	•

Non or ≤ 1 time/ week	13.9 (52)	15.1 (19)	13.3 (33)	
Current smoking - % (n)				
No	91.9 (352)	94.7 (125)	90.4 (227)	0.146
Yes	8.1 (31)	5.3 (7)	9.6 (24)	
Current alcohol consumption - % (n)				
No	68.9 (264)	78.8 (104)	63.7 (160)	0.003
Yes	31.1 (119)	21.2 (28)	36.3 (91)	
BMI: Body Mass Index; WC: Waist Circumference; A	AP: Amended Pressu	ure; BFIS: Brazilia	n Food Insecurity	Scale; FN
Food and Nutrition Security; FNiS: Food and Nutrition	on Insecurity; BDI-II:	Beck Depression	Inventory-II. Quan	titative
variables presented in medians and interquartile ran	ges (IR), or means a	and standard devia	ations (±SD) accor	ding to
normality (Kolmogorov-Smirnov test); categorical va	riables presented in	relative (%) and a	bsolute (n) freque	ncies. * p
value for the tests: Mann-Whitney U or Student t and	d chi-square, at 5% s	significance and, v	when three or more	9
categories, corrected by Bonferroni. ** 5% significar	ice, adjusted by Bon	ferroni (p <0.016).		
Tables 3 and 4 present the	results of the	association	between ind	icator
of psychosocial stress and central	adiposity in th	e rural and	urban popul	ations
respectively. In the rural population	n, serum cortis	sol was inve	rsely associa	ated
with central adiposity in all models.	AP was asso	ciated with	adiposity on	ly in tl
crude model, losing significance in	the adjusted	models (Tab	ole 3). The	
evaluation done in the urban popul	ation showed	that only Al	⊃ was assoc	iated
with central adiposity and this occu	irred in all mo	dels (Table	4). The othe	r stres
indicators evaluated – FNiS and Bl	DI-II> 17 – ha	d no associa	ation with ce	ntral
adiposity in the study population (T	ables 3 and 4	!).		
	, , ,	s and centra	al adiposity i	
Table 3. Association between indic	ators of stres		a auposity ii	n the
Table 3. Association between indicrural population	ators of stres		a adiposity ii	n the
	Current smoking - % (n) No Yes Current alcohol consumption - % (n) No Yes BMI: Body Mass Index; WC: Waist Circumference; A Food and Nutrition Security; FNiS: Food and Nutritic variables presented in medians and interquartile ran normality (Kolmogorov-Smirnov test); categorical va value for the tests: Mann-Whitney U or Student t and categories, corrected by Bonferroni. ** 5% significant Tables 3 and 4 present the no of psychosocial stress and central respectively. In the rural population with central adiposity in all models. crude model, losing significance in evaluation done in the urban popul with central adiposity and this occu indicators evaluated – FNiS and BI	Current smoking - % (n) No 91.9 (352) Yes 8.1 (31) Current alcohol consumption - % (n) No 68.9 (264) Yes 31.1 (119) BMI: Body Mass Index; WC: Waist Circumference; AP: Amended Press Food and Nutrition Security; FNIS: Food and Nutrition Insecurity; BDI-II: variables presented in medians and interquartile ranges (IR), or means a normality (Kolmogorov-Smirnov test); categorical variables presented in value for the tests: Mann-Whitney U or Student t and chi-square, at 5% scategories, corrected by Bonferroni. ** 5% significance, adjusted by Bon Tables 3 and 4 present the results of the scategories, corrected by Bonferroni. ** 5% significance, adjusted by Bon Tables 3 and 4 present the results of the scategories, corrected by Bonferroni. ** 5% significance, adjusted by Bon Tables 3 and 4 present the results of the scategories, corrected by Bonferroni. ** 5% significance, adjusted by Bon Current adiposity in all models. AP was associated the central adiposity in the respectively. In the rural population, serum cortists with central adiposity in all models. AP was associated evaluation done in the urban population showed with central adiposity and this occurred in all modiation showed with central adiposity and this occurred in all modiation showed with central adiposity and this occurred in all modiation showed with central adiposity and this occurred in all modiation showed with central adiposity and this occurred in all modiation showed with central adiposity and this occurred in all modiation showed with central adiposity and	Current smoking - % (n) No 91.9 (352) 94.7 (125) Yes 8.1 (31) 5.3 (7) Current alcohol consumption - % (n)	Current smoking - % (n) No 91.9 (352) 94.7 (125) 90.4 (227) Yes 8.1 (31) 5.3 (7) 9.6 (24) Current alcohol consumption - % (n) No 68.9 (264) 78.8 (104) 63.7 (160) Yes 31.1 (119) 21.2 (28) 36.3 (91) BMI: Body Mass Index; WC: Waist Circumference; AP: Amended Pressure; BFIS: Brazilian Food Insecurity : Food and Nutrition Security; BDI-II: Beck Depression Inventory-II. Quan variables presented in medians and interquartile ranges (IR), or means and standard deviations (±SD) accord normality (Kolmogorov-Smirnov test); categorical variables presented in relative (%) and absolute (n) freque value for the tests: Mann-Whitney U or Student t and chi-square, at 5% significance and, when three or more categories, corrected by Bonferroni. ** 5% significance, adjusted by Bonferroni (p <0.016).

	Univariate linear	Univariate linear regression models adjusted						
Stress	regression	for confounding variables						
indicators	Gross analysis	Model 1	Model 2	Model 3				
	β 95%Сl р	β 95%Cl p	β 95%СІ р	β 95%СΙ р				

FNiS	-0.13	-4.62;	0.956	-0.42	-5.21;	0.864	-1.28	-6.21;	0.608	-1.11	-6.34;	0.675
		4.37			4.38			3.65			4.12	
Serum	-0.61	-1.06;	0.010	-0.51	-0.98;	0.033	-0.58	-1.05;	0.015	-0.60	-1.09;	0.018
cortisol		-0.15			-0.04			-0.11			-0.11	
BDI-II ≥ 17	4.62	-0.71;	0.089	5.16	-0.06;	0.053	4.31	-1.45;	0.141	4.95	-1.57;	0.135
		9.96			10.38			10.06			11.47	
AP	6.77	0.22;	0.043	2.68	-4.35;	0.452	-1.72	-9.80;	0.674	-4.71	-13.64;	0.298
		13.33			9.71			6.36			4.22	

281 Complex sample. Univariate linear regression. Significance level of 5% (p <0.05). Dependent variable: Waist 282 circumference (WC) in cm; Stress indicators (independent variables): Serum cortisol (in µg / dL); Food and Nutrition 283 Insecurity – FNiS (yes); Beck Depression Inventory-II – BDI-II ≥ 17 (yes); Amended Pressure – AP ≥ 130/85 mmHg 284 (yes). Gross analysis and univariate linear regression models hierarchically adjusted for confounding factors: Model 1 -285 gross analysis adjusted for socioeconomic variables (sex, age in years, schooling, and income status); Model 2 - gross 286 analysis, adjusted for socioeconomic variables (sex, age in years, schooling, and income status), and health variables 287 (stress report, reporting one or more comorbidities, and self-rated health); Model 3 - gross analysis, adjusted for 288 socioeconomic variables (sex, age, schooling, and income status), health variables (stress report, reporting of one or 289 more comorbidities, and self-rated health), and lifestyle variables (current smoker, current alcohol consumption, and 290 practice of physical activity and/or weekly leisure). β: beta coefficient; 95% CI: 95% confidence interval; p: p-value. The 291 variables used met the assumption of collinearity. The assumptions of normality, linearity, homoscedasticity, and 292 independence of residues were met in all models that had p <0.05 by F test.

293

Table 4. Association between indicators of stress and central adiposity in the

295 urban population

				Univariate linear regression models adjusted for confounding variables								
Stress												
indicators				Model 1			Model 2			Model 3		
	β	95%CI	р	β	95%CI	р	β	95%CI	р	β	95%CI	р
FNiS	-0.66	-4.37;	0.725	-0.62	-4.59;	0.758	-1.22	-5.31;	0.558	-0.65	-4.81;	0.759
		3.04			3.35			2.87			3.51	
Serum	-0.21	-0.62;	0.294	-0.17	-0.59;	0.409	-0.19	-0.61;	0.359	-0.27	-0.69;	0.211
cortisol		0.19			0.24			0.22			0.15	
BDI-II ≥ 17	1.80	-2.66;	0.427	1.59	-3.08;	0.503	0.82	-4.02;	0.740	1.64	-3.27;	0.512
		6.25			6.26			5.65			6.55	
AP	9.66	5.52;	<0.001	8.41	-4.05;	<0.001	7.07	2.59;	0.002	6.66	2.14;	0.004
		13.80			12.76			11.55			11.18	

296 Complex sample. Univariate linear regression. Significance level of 5% (p < 0.05). Dependent variable: Waist 297 circumference (WC) in cm; Stress indicators (independent variables): Serum cortisol (in µg / dL); Food and Nutrition 298 Insecurity – FNiS (yes); Beck Depression Inventory-II – BDI-II ≥ 17 (yes); Amended Pressure – AP ≥ 130/85 mmHg 299 (yes). Gross analysis and univariate linear regression models hierarchically adjusted for confounding factors: Model 1 -300 gross analysis adjusted for socioeconomic variables (sex, age in years, schooling, and income status); Model 2 - gross 301 analysis, adjusted for socioeconomic variables (sex, age in years, schooling, and income status), and health variables 302 (stress report, reporting one or more comorbidities, and self-rated health); Model 3 - gross analysis, adjusted for 303 socioeconomic variables (sex, age, schooling, and income status), health variables (stress report, reporting one or more 304 comorbidities, and self-rated health), and lifestyle variables (current smoker, current alcohol consumption, and practice 305 of physical activity and/or weekly leisure). β: beta coefficient; 95% CI: 95% confidence interval; p: p-value. The variables 306 used met the assumption of collinearity. The assumptions of normality, linearity, homoscedasticity, and independence of 307 residues were met in all models that had p <0.05 by F test.

308

309 **Discussion**

310 This study evaluated the association between indicators of psychosocial stress and central adiposity in adults of a city in Southeast Brazil, and users of 311 312 the Unified Health System (SUS). The health conditions of the Brazilian population, as well as other nations, transcend the concept of absence of 313 diseases resulting from innumerable social, economic, environmental, and 314 315 cultural factors. The main health problems of people living in poverty include 316 increased exposure to environmental risk factors, diseases (especially non-317 communicable diseases), poor nutritional status, difficult access to health 318 services and medicines, as well as other economic, psychosocial, cultural, or health factors, which act as stressing agents, affecting well-being and social 319 320 interaction.(38)

This study was performed on a population that frequently uses the public health system, which presented socioeconomic (mainly in the rural area) and health fragility, including self-perception reports (53.1% reported poor or regular health), which may represent an impairment of the individual and collective

state of well-being, expressing as continuous stress. This, in turn, tends to feed
the same cycle of psychosocial instability and lack of health and well-being in a
chronic way.

The prevalence of weight excess (BMI $\ge 25.0 \text{ kg} / \text{m2}$) found in the present study (68.3%) was equivalent to the current US data and surpassed the weight excess data of the world population (52.0%)(2), of the Brazilian adult population (56.9%)(5), the population of the state of Espirito Santo (52.4%)(39), and of its capital Vitoria (49.7%).(40)

Data from the Family Expense Research - POF 2008-2009(39) showed a lower prevalence of overweight and obesity in rural Brazil (38.8% and 8.8%, respectively), whereas in the present study the prevalence of overweight and obesity were high and statistically similar, both in rural and urban areas (Table 2).

Added to this context, the results were worse when the WC assessment 338 was expanded, with a prevalence of 71.5% at risk of metabolic complications 339 associated with central adiposity, with 51.6% of the individuals at high risk. This 340 341 proportion exceeded the national prevalence stated in the National Health 342 Survey – NHS(5), which was 37.0%, using the same cut points.(1, 32) 343 Based on the literature, the main socioeconomic, health, and lifestyle indicators related to excess weight were considered in the present study (Table 344 345 2)(5, 9, 14, 26, 39, 41), which have also been referred to in studies as psychosocial stressors.(7-9, 11, 42) 346

Socioeconomic variables such as schooling and income are indicators of poverty and, together with the FNiS indicator, may represent a direct relation of hunger, and scarcity, with deficiency and malnutrition states.(25, 43, 44) The

socioeconomic profile traced in this study pointed to a high prevalence of FNiS 350 351 (41.8%) in the general population, characterized by low schooling and income, especially in the rural population (Table 2). The nutritional assessment pointed to 352 30.2% having eutrophy, and only 1.6% having low weight, distancing the 353 354 population from the limit of 5% that would characterize the presence of current 355 malnutrition.(5) On the other hand, it is known that poverty indicators are also 356 related to excess caloric intake(45) and, consequently, to adiposity. Under these conditions, adiposity may even mask situations of nutritional deficiencies, which 357 is often the case in less developed regions with populations of low schooling and 358 359 income.(13, 14, 45)

FNiS was one of the variables chosen to evaluate the association between psychosocial stress and adiposity, since this indicator represents scarcity and is related to fear of hunger, insecurity, fragility, and individual and familial instability, which could trigger the activation of the HPA axis, with chronic stress setting. However, FNiS had no association with central adiposity (Tables 3 and 4).

366 In the context of analysis of health variables, 15% of the individuals living 367 in rural areas had low levels of serum cortisol. Although the prevalence of 368 hypocortisolemia was statistically similar between rural and urban areas, the median serum cortisol was significantly lower in the rural population than in the 369 370 urban population (p = 0.006), according to Table 2. In addition, in evaluating the association of stress indicators with central adiposity, serum cortisol was 371 372 inversely associated with central adiposity in the rural population in all univariate analyses (gross analysis and confounding-adjusted models, as in Table 3). In 373 374 the urban population, serum cortisol was not associated with adiposity (Table

4). The association found in the rural population was contrary to what has
been observed in some studies on the relationship between stress and obesity,
which have justified the weight gain, due to the increase in intake of palatable
and high caloric foods, as a compensatory mechanism resulting in
hypercortisolemia of chronic HPA axis activation(7, 8, 20, 46); or weight gain
due to the deregulation of satiety mechanisms, through alterations in the leptin
system, and also resulting from high levels of cortisol.(7, 12)

On the other hand, the notion that chronic stress acts simply by the direct 382 effect of elevated serum cortisol is becoming less likely relevant in describing 383 384 how the target tissues respond to cortisol, rather than the levels of the hormone 385 itself.(9) Stressors trigger an HPA axis response by regulating corticotrophinreleasing factor and cortisol feedback and, over time, chronic overexposure to 386 387 stressors may result in a blunted stress response with decreased levels of 388 cortisol and other associated neurobiological dysfunctions. (47-49) In the absence of supportive care, stressors experienced during life, especially in early 389 life and developmentally sensitive periods, such as pregnancy, childhood, and 390 391 adolescence, may leave impressions on the neural substrate of emotional and 392 cognitive processes, which may result in a blunted response to cortisol and consequent resilience of the individual to psychosocial stressors.(49) Regarding 393 obesity, hyperactivation of the HPA axis has been observed in obese 394 395 individuals, but also in individuals with paradoxically normal or low plasma cortisol levels. In a study by research colleagues(50), obesity was associated 396 397 with relative insensitivity to glucocorticoid feedback. The authors suggested that this condition is characterized by a decreased response of the mineralocorticoid 398 399 receptor to circulating cortisol.

However, the associations between cortisol and obesity appear to be 400 401 more complex than anticipated(51). Although it promotes changes in the 402 compensatory mechanism of food intake(7, 8, 20, 46), in the leptin system and satiety control mechanism(7, 8), cortisol is clearly not the only peripheral trigger 403 404 of adverse effects, which may explain the controversies in this area. It seems likely that the pattern in which cortisol is secreted in front of the stressors is as 405 406 important as total cortisol secretion. Thus, there is relevance in exploring the social, psychological conditions, health, and lifestyle of a population to identify 407 the main psychosocial stressors and how individuals react and resist to stress. 408 409 Another indicator of psychosocial stress used in the present study was 410 depression using BDI-II scores \geq 17. It is known that early life stress is a risk factor for psychopathology in adulthood, to the extent that the HPA axis was 411 412 described as deregulated in individuals who experienced early psychosocial stress as well as in those with a wide range of psychiatric disorders.(52) In 413 studies done by colleagues(53) related to chronic stress, depression, and 414 415 location of urban neighborhoods, the daily stress of living in neighborhoods 416 where residential mobility and material deprivation prevailed was associated 417 with depression. Another study showing the association between urban versus 418 rural environments and depression in the Scottish population found a higher prevalence of psychotropic medication prescription for anxiety, depression, and 419 420 psychosis in the urban population.(54)

In the present study, a prevalence of depression was found in 23.6% of the population assisted by SUS, with a statistical similarity between rural and urban areas (Table 2). This indicator was not associated with adiposity in any of the models, according to Tables 3 and 4.

In the urban population, the evaluation of health variables pointed to 425 426 higher levels within the following indicators: median BMI and cortisol, mean WC, and prevalence of high blood pressure (24.3% of individuals with AP), 427 corroborating data from the literature related to hypertension and obesity in 428 429 urban areas. (55, 56) Lifestyle assessment also showed higher alcohol consumption in the urban population (Table 2). Finally, AP was the only 430 431 indicator of stress associated with adiposity in the urban population. It is known that among the main risk factors for elevated blood pressure 432 in adults are obesity, high sodium intake, smoking, alcohol consumption, 433 434 psychological factors, and certain personality traits such as stress and 435 anxiety.(57-59) Most studies report obesity as a predictor of arterial hypertension(58-63); however, in this study, it was hypothesized that AP, as a 436 437 psychosocial stress indicator, could predict the increase of WC, which occurred in all models tested (Table 4). It is recalled here that, in the rural population 438 (Table 3), AP was also associated with central adiposity, but this association 439 occurred only in the gross analysis since the adjusted models for confounding 440 441 variables lost their significance.

442 Finally, as part of the initial hypothesis, significant differences were 443 observed between home locations. Socioeconomic conditions were challenging in the rural area regarding income and schooling, despite the statistical 444 445 similarity in the prevalence of FNiS between rural and urban areas. In addition, the report of stress as a personality trait was higher in the population living in 446 447 the rural region which, on the other hand, presented the lowest median for serum cortisol. As mentioned before, the urban population exceeded the rural 448 population regarding the prevalence of AP. Both the literature review data(53, 449

450 54, 64, 65) and the findings of this study reinforce the influence of

451 socioeconomic and livelihood conditions on the different mechanisms of

adaptation of the organism to stressors, and on the different forms of coping

453 and developing resilience.

454

455 **Conclusions**

The stress indicators that were associated with central adiposity were serum cortisol (with an inverse association) in the rural environment and altered blood pressure in the urban area. The study considered several variables known to be associated with stress and weight gain and the models presented robustness in the explanation of the results found.

The observed differences in adiposity prediction regarding housing location reinforce the influence of the local and psychosocial environments on the modulation of stress and on how individuals react to or restrain stressors. Stress reduction strategies can be useful in public health programs designed to prevent or treat obesity.

The study suggests the need for a better evaluation of the use of cortisol 466 as a marker, since both high and low cortisol may explain variations in health 467 468 status. Both levels relate to different forms of coping or resilience to psychosocial stress. Additionally, owing to the possible bidirectionality of the 469 association between stress and adiposity, new paths should be drawn in the 470 471 context of a thorough investigation of the mechanisms that explain psychosocial 472 stress, with hypocortisolemia as a biomarker and adiposity as a consequence of current lifestyle. 473

474

475 Acknowledgments

- 476 We would like to thank the volunteers of this study, the Community
- 477 Health Agents, and the entire Primary Care team of the Municipality of Alegre,
- 478 ES, Brazil.
- 479

480 **Disclosure of potential conflicts of interest**

- 481 No potential conflicts of interest were disclosed. The funders had no role
- in the study design, data collection, analysis, decision to publish, or preparation
- 483 of the manuscript.
- 484

485 **Author Contributions**

- 486 Conceived and designed the experiments: AMAS FVF. Performed the
- 487 experiments: FVF WMB LAAS MJOG JAP ARB CLC JKA HP MMO ABA EASF.
- 488 Analyzed the data: FVF AMAS WMB EBB IDL DRO. Contributed
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- 490 HP ABA DRO EBB IDL AMAS. Wrote the paper: FVF AMAS. Research Project
- 491 Coordinator: AMAS.
- 492

493 **References**

- World Health Organization WHO. Obesity: Preventing and managing the global
 epidemic: Report of a WHO Consultation on obesity. Geneva: World Health Organization
 Technical Report Series, 2000.894 p.
- 497 2. World Health Organization WHO. Global Health Observatory data repositor.498 Overweight and obesity. Available from:
- 499 <u>http://www.who.int/gho/ncd/risk_factors/overweight_obesity/obesity_adults/en/</u>.
- World Health Organization WHO. Global status report on noncommunicable diseases
 2014. Geneva: World Health Organization Technical Report Series, 2014.
- Pan American Health Organization PAHO. Plan of Action for the Prevention of Obesity
 in Children and Adolescents. Washington, D.C.: PAHO, 2015.

504 5. Instituto Brasileiro de Geografia e Estatística - IBGE, Coordenação de Trabalho e 505 Rendimento. Pesquisa nacional de saúde 2013: ciclos de vida – Brasil e grandes regiões. Rio de 506 Janeiro: IBGE, 2015; 92p. 507 Selve H. Stress and the general adaptation syndrome. Br Med J. 1950;1:1384-92. 6. 508 Sinha R, Jastreboff AM. Stress as a common risk factor for obesity and addiction. Biol 7. 509 Psychiatry. 2013;73(9): 827-35. 510 Gundersen C, Mahatmya D, Garasky S, Lohman B. Linking psychosocial stressors and 8. 511 childhood obesity. Obes Rev. 2011;12(5): e54-63. 512 Cohen S, Janicki-Deverts D, Doyle WJ, Miller GE, Frank E, Rabin BS, et al. Chronic stress, 9. 513 glucocorticoid receptor resistance, inflammation, and disease risk. Proceedings of the National 514 Academy of Sciences of the United States of America. 2012;109(16): 5995-99. 515 Fonseca FCA, Coelho RZ, Nicolato R, Malloy-Diniz LF, Silva-Filho HC. The influence of 10. 516 emotional factors on the arterial hypertension. J Bras Psiguiatr. 2009;58(2): 128-34. 517 Isasi CR, Parrinello CM, Jung MM, Carnethon MR, Birnbaum-Weitzman O, Espinoza RA, 11. 518 et al. Psychosocial stress is associated with obesity and diet quality in Hispanic/Latino adults. 519 Ann Epidemiol. 2015;25(2):84-9. 520 Torres SJ, Nowson CA. Relationship between stress, eating behavior, and obesity. 12. 521 Nutrition. 2007;23(11-12): 887-94. 522 Chopra M, Galbraith S, Darnton-Hill I. A global response to a global problem: the 13. epidemic of overnutrition. Geneva: Bulletin of the World Health Organization, 2002.80(12): 523 524 952-58. 525 14. Rtveladze K, Marsh T, Webber L, Kilpi F, Levy D, Conde W, et al. Health and Economic 526 Burden of Obesity in Brazil. PLoS ONE. 2013;8(7): e68785. 527 https://doi.org/10.1371/journal.pone.0068785. 528 Monteiro CA, Mondini L, de Souza AL, Popkin BM. The nutrition transition in Brazil. Eur 15. 529 J Clin Nutr. 1995;49: 105-13. 530 16. Monteiro CA, D'A Benicio MH, Conde WL, Popkin BM. Shifting obesity trends in Brazil. 531 Eur J Clin Nutr. 2000;54:342-46. 532 17. Brasil. Ministério da Saúde. Perspectivas e desafios no cuidado as pessoas com 533 obesidade no SUS: resultados do laboratório de inovação no manejo da obesidade nas Redes 534 de Atenção à Saúde/Ministério da Saúde; Organização Pan-Americana da Saúde. Brasília: 535 Ministério da Saúde. 2014;116 p. 536 Carvalho MC, Martins AA. A obesidade como objeto complexo: uma abordagem 18. 537 filosofico-conceitual Ciência & Saúde Coletiva. 2004;9(4): 1003-12. 538 Chan M. Opening address at the 8th Global Conference on Health Promotion Helsinki, 19. 539 Finland 10 June 2013. Available from : 540 <www.who.int/dg/speeches/2013/health promotion 20130610/en/index.html>. 541 Epel ES, McEwen B, Seeman, T. Stress and body shape: stress-induced cortisol 20. 542 secretion is consistently greater among women with central fat. Psychosom Med. 2000;62: 543 623-32. 544 Ehlert U. Enduring psychobiological effects of childhood adversity. 21. 545 Psychoneuroendocrinology. 2013;38: 1850-57. 546 Associação Brasileira para o Estudo da Obesidade e da Síndrome Metabólica - ABESO. 22. 547 Diretrizes brasileiras de obesidade 2016. 4.ed. São Paulo, SP: ABESO. 2016; 188 p. 548 23. Instituto Brasileiro de Geografia e Estatística - IBGE. Panorama IBGE Cidades. Available 549 from: https://cidades.ibge.gov.br/brasil/es/alegre/panorama. 550 Dean AG, Sullivan KM, Soe, MM. OpenEpi: Open Source Epidemiologic Statistics for 24. 551 Public Health, Versão. Available from: www.OpenEpi.com, updated: April 6, 2013. 552 Segall-Corrêa AM, Marin-Leon L. A Segurança Alimentar no Brasil: Proposição e Usos 25. 553 da Escala Brasileira de Medida da Insegurança Alimentar (EBIA) de 2003 a 2009. Segurança 554 Alimentar e Nutricional, Campinas. 2009;16(2): 1-19.

555 26. Instituto Brasileiro de Geografia e Estatística - IBGE. Pesquisa Nacional por Amostra de 556 Domicílios PNAD. Segurança Alimentar 2013. Rio de Janeiro: IBGE. 2014; 134 p. 557 27. Braver MBJ, Green J, Rawson R. Childhood abuse and current psychological functioning 558 in a university counseling center population. J Couns Psychol. 1992;39: 2-6. 559 28. Burns DD. Feeling good: The new mood therapy. New York: New American Libray, 560 1981. 561 29. Ibrahim MBA. Prevalence of anxiety and depression among medical and 562 pharmaceutical students in Alexandria University. Journal of Medicine. 2014;51:7. 563 Brasil. Ministério da Saúde. Secretaria de Atenção à Saúde. Departamento de Atenção 30. 564 Básica. Orientações básicas para coleta, o processamento e análise de dados e a informação 565 em serviço de saúde. Norma Técnica do Sistema de Vigilância Alimentar e Nutricional – 566 SISVAN. Brasília, DF. 2011;76 p. 567 Lohman T, Roche A, Martorell R. Anthropometric standardization reference manual. 31. 568 Champagne, Illinois, Human Kinetic Books; 1988. 569 32. World Health Organization - WHO. Waist circumference and waist-hip ratio: report of 570 a WHO expert consultation. Geneva, World Health Organization. 2011;47 p. 571 Sociedade Brasileira de Cardiologia - SBC. VI Diretrizes Brasileiras de Hipertensão. Arq 33. 572 Bras Cardiol. 2010;95(1 supl.1): 1-51. 573 34. Instituto de Patologia Clínica Hermes Pardini. Manual de Exames. Hermes Pardini: 574 2015/2016. 573 p. Available from: 575 https://www3.hermespardini.com.br/repositorio/media/site/profissionais da saude/manual 576 exames.pdf. 577 Andy Field. Descobrindo a estatística usando o SPSS. 2.ed. Porto Alegre: Artmed. 2009; 35. 578 684 p. ISBN: 978-85-363-2018-2. 579 Monteiro CA, Conde WL. A Tendência Secular da Obesidade Segundo Estratos Sociais: 36. 580 Nordeste e Sudeste do Brasil 1975-1989-1997. Arg Bras Endocrinol Metab. 1999;43(3): 186-94. 581 37. Neri MC. Fundação Getúlio Vargas - FGV. Centro de Políticas Sociais - CPS. A nova 582 classe média: desigualdade, renda, miséria, classe média, mobilidade trabalhista I. Rio de 583 Janeiro: FGV/IBRE, CPS, 2008; 70 p. 584 Instituto de Pesquisa Econômica Aplicada - IPEA. Brasil em desenvolvimento 2013: 38. 585 estado, planejamento e políticas públicas. Editores: Rogério Boueri, Marco Aurélio Costa. 586 Brasília: Ipea, v. 3 (Brasil: o Estado de uma Nação), 2013 587 39. Instituto Brasileiro de Geografia e Estatística - IBGE. Ministério do Planejamento, 588 Orçamento e Gestão. Pesquisa de Orçamentos Familiares – POF 2008/2009. Antropometria e 589 Estado Nutricional de Crianças, Adolescentes e Adultos no Brasil. Rio de Janeiro: IBGE, 2010; 590 130 p. 591 40. Brasil. Ministério da Saúde. Secretaria de Vigilância em Saúde. Departamento de 592 Vigilância de Doenças e Agravos não Transmissíveis e Promoção da Saúde. Vigitel Brasil 2016: 593 estimativas sobre frequência e distribuição sociodemográfica de fatores de risco e proteção 594 para doenças crônicas nas capitais dos 26 estados brasileiros e no Distrito Federal em 2016. 595 Brasília: Ministério da Saúde, 2017; 160p. 596 41. Rundle A, Richards C, Bader MD, Schwartz-Soicher O, Lee KK, Quinn J, et al. Individual-597 and school-level sociodemographic predictors of obesity among New York City public school 598 children. Am J Epidemiol. 2012;176(11): 986-94. 599 42. Razzoli M, Bartolomucci A. The dichotomous effect of chronic stress on obesity. TEM. 600 2016;27(7): 504-15. 601 43. Graziano da Silva J, Del Grossi ME, Franca CG. O Programa Fome Zero: a experiência 602 brasileira. Brasília: MDA. (NEAD Série Especial, 13), 2010. 603 VALENTE FLS. O direito humano à alimentação: desafios e conquistas. Do combate à 44. 604 fome à insegurança alimentar e nutricional: o direito à alimentação adequada. São Paulo:

605 Cortez Editora, 2002; p. 37- 70.

606 45. Moubarac JC, Batal M, Louzada ML, Martinez Steele E, Monteiro CA. Consumption of 607 ultra-processed foods predicts diet quality in Canada. Appetite. 2017;108: 512-20. 608 46. Björntorp P, Rosmond R. Obesity and Cortisol. Nutrition, 2000; 16(10). 609 47. Gunnar M, Quevedo K. The neurobiology of stress and development. Annu Rev 610 Psychol. 2007;58: 145-73. 611 48. Turner HA, Turner RJ. Understanding variations in exposure to social stress. Health. 612 2005;9(2): 209-40. 613 49. Harkness KL, Stewart JG, Wynne-Edwards KE. Cortisol reactivity to social stress in 614 adolescents: role of depression severity and child maltreatment. Psychoneuroendocrinology. 615 2011;36(2): 173-81. 616 50. Jessop DS, Dallman MF, Fleming D, Lightman SL. Resistance to glucocorticoid feedback 617 in obesity. J Clin Endocrinol Metab. 2001;86(9): 4109-14. 618 John K, Marino JS, Sanchez ER, Hinds TD, Jr. The glucocorticoid receptor: cause of or 51. 619 cure for obesity? Am J Physiol Endocrinol Metab. 2016;310(4): E249-57. 620 52. Palma-Gudiel H, Cordova-Palomera A, Leza JC, Fananas L. Glucocorticoid receptor gene 621 (NR3C1) methylation processes as mediators of early adversity in stress-related disorders 622 causality: A critical review. Neurosci Biobehav Rev. 2015;55: 520-35. 623 Matheson FI, Moineddin R, Dunn JR, Creatore MI, Gozdyra P, Glazier RH. Urban 53. 624 neighborhoods, chronic stress, gender and depression. Soc Sci Med. 2006;63(10): 2604-16. 625 54. McKenzie K, Murray A, Booth T. Do urban environments increase the risk of anxiety, 626 depression and psychosis? An epidemiological study. J Affect Disord. 2013;150(3): 1019-24. 627 55. Sobngwi E, Mbanya JC, Unwin NC, Kengne AP, Fezeu L, Minkoulou EM, et al. Physical 628 activity and its relationship with obesity, hypertension and diabetes in urban and rural 629 Cameroon. International journal of obesity and related metabolic disorders. Journal of the International Association for the Study of Obesity. 2002;26(7): 1009-16. 630 631 Vidya BR, Swetha S, Neerajaa J, Varsha MJ, Janani DM, Rekha SN, et al. An 56. 632 epidemiological survey: Effect of predisposing factors for PCOS in Indian urban and rural 633 population. Middle East Fertil Soc J. 2017;22(4): 313-16. 634 57. Fonseca FCA, Coelho RZ, Nicolato R, Mally-Diniz LF, Silva Filho HC. A influência de 635 fatores emocionais sobre a hipertensão arterial. J Bras Psiguiatr. 2009; 58(2): 128-34. 636 Bezerra VM, Andrade ACS, Medeiros DS, Caiaffa WT. [Arterial prehypertension in slave-58. 637 descendant communities in southeast Bahia State, Brazil]. Cadernos de Saude Publica. 638 2017;33(10): e00139516. 639 59. Bezerra VM, Andrade ACdS, César CC, Caiaffa WT. Comunidades quilombolas de Vitória 640 da Conquista, Bahia, Brasil: hipertensão arterial e fatores associados. Cadernos de Saude 641 Publica. 2013;29(9): 1889-902. 642 60. Schmidt MI, Duncan BB, e Silva GA, Menezes AM, Monteiro CA, Barreto SM, et al. 643 Chronic non-communicable diseases in Brazil: burden and current challenges. The Lancet. 644 2011;377(9781): 1949-61. 645 Davy KP, Hall JE. Obesity and hypertension: two epidemics or one? Am J Physiol Regul 61. 646 Integr Comp Physiol. 2004;286: R803-R813. 647 Hall JE, do Carmo JM, da Silva AA, Wang Z, Hall ME. Obesity-induced hypertension: 62. 648 interaction of neurohumoral and renal mechanisms. Circulation Research. 2015;116(6): 991-649 1006. 63. 650 Jiang SZ, Lu W, Zong XF, Ruan HY, Liu Y. Obesity and hypertension. Exp Ther Med. 651 2016;12(4): 2395-99. Elgar FJ, Arlett C, Groves R. Stress, coping, and behavioural problems among rural and 652 64. 653 urban adolescents. J Adolesc. 2003;26(5): 574-85. 654 Sameem S, Sylwester K. The business cycle and mortality: urban versus rural counties. 65.

655 Soc Sci Med. 2017;175: 28-35.