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ORIGINAL RESEARCH

Racial Disparities in Medication Use During Pregnancy: Results from the NISAMI Cohort

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Purpose: This study aimed to evaluate racial disparities in medication use and associated factors among pregnant women receiving prenatal care at Brazilian Unified Health System primary care health units in the northeast region.

Patients and Methods: A total of 1058 pregnant women in the NISAMI Cohort were interviewed between June 2012 and February 2014. Medicines used during pregnancy were classified according to the Anatomical Therapeutic Chemical (ATC) classification system and ANVISA pregnancy risk categories. Prevalence ratios (crude and adjusted) and 95% confidence intervals (CIs) were estimated using Poisson regression with robust error variance. All analyses were stratified by race (Asian, black, brown/ mixed, Brazilian indigenous, and white).

Results: Approximately 84% of the pregnant women used at least one medication, with a lower proportion among white women. The most reported medications were antianemic preparations (71.08%; 95% CI 68.27–73.72%), analgesics (21.74%; 95% CI 19.36–24.32%), and drugs for functional gastrointestinal disorders (18.81%; 95% CI 16.57–21.28%). Approximately 29% of women took potentially risky medications during pregnancy, with a higher prevalence among Asian and white women. Factors associated with medication use during pregnancy include a greater number of prenatal consultations, higher education levels, health problems, and smoking. In addition, maternal age above 25 years, smoking status, and two or more previous pregnancies were associated with potentially risky medication use during pregnancy.

Conclusion: A high prevalence of medication use during pregnancy was found; however, this prevalence was lower among white women. Nonetheless, black and brown women used antianemic preparations less frequently. This finding suggests that race is a factor of inequity in prenatal care, demanding public policies to mitigate it.

Keywords: drug utilization, racial groups, pregnancy, pharmacoepidemiology, cohort studies

Introduction

Medication use during pregnancy is common in treating acute and chronic conditions through prescription or selfmedication.¹ Among the most commonly used medications by pregnant women described in the literature are medicines prescribed to manage pregnancy and nonpregnancy-related conditions, such as ferrous sulfate, folic acid and vitamins, analgesics, antacids, antibiotics, antiemetics, antihistamines, and nonsteroidal anti-inflammatory agents.^{2–6}

A significant increase in medication consumption by pregnant women has been reported in recent years.^{5,7} The prevalence of at least one medication use during pregnancy reaches proportions greater than 90.0% worldwide.^{8,9} However, medication use might be related to numerous factors, including healthcare access.¹⁰ Furthermore, medicines such as folic acid, insulin, antacids, ondansetron, ranitidine, methyldopa, hydralazine, enoxaparin sodium, nifedipine, and aspirin are more commonly dispensed to pregnant women than to women of childbearing age.¹¹

In addition to the characteristics of healthcare systems, medication use patterns may differ by race because of racism. Racism is a structuring system, historically constituted based on the belief of the inferiority of one race or ethnicity to another. It produces practices, beliefs, behaviors, and prejudices that favor avoidable and unfair inequalities between social groups by obstructing access to goods, resources, services, and opportunities.^{12,13}

Furthermore, institutional racism is a failure by institutions and organizations to provide appropriate and professional services to people because of their race, culture, or ethnicity. Its processes, attitudes, and behaviors cause discrimination due to involuntary prejudice, ignorance, neglect, and racist stereotyping, which disadvantage people of minority groups,¹⁴ resulting in differences in the distribution of services, benefits, and opportunities to different racial groups. Institutional racism, including the effective implementation of public policies and the generation of inequalities and inequities, is established in institutional routines.¹⁵

Due to racism, there are important differences in socioeconomic status, living conditions, education, and healthcare access between black and brown people compared to white people in Brazil. Historically, black and brown Brazilians have a lower socioeconomic status and poorer health outcomes than white Brazilians.^{16,17}

Therefore, racism is a social determinant of health, as it exposes black and brown people to more vulnerable situations of illness and death,^{12,13} which does not differ among pregnant women seeking prenatal care. In Brazil, black and brown people have epidemiological and social vulnerabilities that imply difficulty accessing health services and medications.¹⁸ Similarly, black and brown Brazilian women have difficulties accessing preventive services and are less likely to undergo prenatal care, gynecological consultations, and prenatal care before the fourth month of pregnancy.^{18,19}

The use of medications during pregnancy is a topic of concern because of their potentially harmful effects on the fetus, as many drugs used during pregnancy can cross the placental barrier and cause pharmacological effects.²⁰ However, the use of medicines during pregnancy may be influenced by social determinants of health, such as race. Therefore, this study provides a pertinent discussion in the context of prenatal care and the use of medications during pregnancy stratified by maternal race.

The present study delineates the profile of medication use during pregnancy and associated factors, explaining their differences due to race, which may enhance the literature on the subject, especially because few studies in Brazil have demonstrated inequities in medication use during pregnancy, and there is still a lack of information about race, mainly in regions with well-known disparities in healthcare access.^{4,9}

For example, Costa et al⁴ and Lutz et al⁹ evaluated the factors associated with medication use during pregnancy in two Brazilian regions and reported a lower use of medicines among black women than among white women (PR 0.96, 95% CI 0.90– 1.03; and PR 0.97, 95% CI 0.89–1.06, respectively), although the results were not statistically significant; however, the authors did not stratify their analysis by race group to evaluate potential differences between races and their associated factors; they only used race as an adjustment variable. At the same time, previous studies have explored the use of medicines stratified by the race of pregnant women; however, they limited their analysis to specific drug classes, such as opioids, or low-risk pregnant women.^{21,22}

Moreover, the findings of drug utilization studies can contribute to public health policy decisions and resource allocation²³ which, in this case, could focus on diminishing potential racial disparities in the use of medicines during pregnancy.

Thus, this study aimed to evaluate racial disparities in the prevalence of medication use during pregnancy in women who underwent prenatal care at the Brazilian Unified Health System (BUHS) primary care health units in the northeast region, as well as the factors associated with medication use and potentially risky medication use during pregnancy.

From this perspective, the present study aimed to answer the following questions: "Are there differences in medication use during pregnancy according to maternal race?", and "Considering the maternal race, what factors are associated with medication use during pregnancy by race?".

Materials and Methods

Study Design and Data Source

This was a cross-sectional study nested within the prospective NISAMI Cohort (Núcleo de Investigação em Saúde Materno Infantil - NISAMI). The NISAMI Cohort resulted from the research project entitled "Maternal Risk Factors for

Low Birth Weight, Prematurity and Intrauterine Growth Retardation, in Recôncavo Baiano region", which was performed by the Center for Research in Maternal and Child Health of the Health Sciences Center, Federal University of Recôncavo da Bahia. The research was conducted in all primary care health units in the urban area of Santo Antônio de Jesus city. Pregnant women who sought prenatal care at the Family Health Unit (FHU) through the Brazilian Unified Health System (BUHS) were enrolled from June 2012 to February 2014. These women were followed prospectively throughout pregnancy until delivery.

Santo Antônio de Jesus is a city located in the state of Bahia, Brazil, and had 90,985 inhabitants in 2010, with 79,299 living in the urban area.²⁴ Public health services are provided in 21 FHUs, one basic health unit, and two hospitals.

Study Population

The study population comprised all pregnant women of any gestational age living in urban areas who attended the FHU for prenatal care. Pregnant women attending rural primary care health units were not eligible because of distance and difficulty accessing them. In addition, pregnant women who had not reported their race were excluded from this study. Thus, all pregnant women residing in urban areas, at any gestational age, and who underwent at least one prenatal consultation were included in this study.

Data Collection

Trained interviewers collected data longitudinally during prenatal consultations. A Fieldwork Guidance Manual was prepared to describe in detail all activities related to data collection. Daily visits to prenatal services were carried out from June 2012 to February 2014, where pregnant women were interviewed using a standardized questionnaire with 116 questions grouped into seven axes: sociodemographic characteristics, nutritional information, gynecological-obstetric information, oral health information, blood and biochemical tests, medication use information (before and during pregnancy), and anthropometry. In addition to the self-reported interviews, information related to the obstetric history was obtained from the prenatal services' clinical records.²⁵

The supervisors' team reviewed all the completed questionnaires. Data validation was performed by comparing the information obtained from the questionnaires with that recorded on the prenatal cards. Whenever necessary, the interviewers returned to the patients' homes to confirm the data from the interviews. Quality control of the interviews was performed to identify any inaccuracies, systematic errors, or frauds. Approximately 20% of the pregnant women were revisited by field supervisors who reapplied selected parts of the questionnaires. The data were compared with the original interviews. Questionnaires with inaccuracies that could not be corrected were excluded.²⁶

Outcomes

The Outcomes of interest were medication use and potentially risky medication use through prescription and/or over-thecounter at any time during pregnancy. During the interview, the women were asked,

Have you used any medication during this pregnancy for high blood pressure, diabetes (high blood sugar), pain or colic, nausea or vomiting, cough, vaginal discharge, infection, respiratory problems, and/or other reasons?

If pregnant women answered yes, they provided all their medication names and dosages.

Medications were coded up to the third level (pharmacological subgroup) according to the Anatomical Therapeutic Chemical Classification System (ATC) of the World Health Organization.²⁷ Additionally, medications were classified according to the Food and Drug Administration (FDA) risk classification system adopted by the Agência Nacional de Vigilância Sanitária (ANVISA) to identify potential risks to the fetus due to the use of medications during pregnancy.²⁸ Although the FDA has recommended discontinuation of this classification since 2015, it is still used by ANVISA.

According to this classification, medications were divided into five risk categories: category A, drugs without risk to the fetus; category B, drugs for which animal studies have not demonstrated a fetal risk, but there are no human studies; category C, drugs for which animal studies have shown adverse effects on the fetus, but there are no adequate human studies; category D, drugs for which the experience of use during pregnancy showed an association with malformations, whose risk-benefit should be evaluated; and category X, drugs associated with fetal abnormalities in animal and human

studies and/or whose risk-benefit contraindicates their use in pregnancy.²⁸ Potentially risky medications during pregnancy were those classified as C, D, or X due to adverse effects on the fetus.

Exposure

Information regarding exposure was self-reported maternal race, recorded during the interview, and categorized into five groups: Asian, black, Brazilian indigenous, brown, and white.

Covariates

Covariates were selected after a literature review and included socioeconomic and demographic, maternal, and health service utilization variables: maternal age (≤ 24 ; 25 to 29; 30 to 35; ≥ 36 years); education (≤ 8 ; 9 to 11; ≥ 11 years); marital status (with a partner; without a partner); employment status (with employment; without employment); family income – according to the 2013 minimum wage (one minimum wage in 2013: R\$ 678.00/US\$ 301.33) (MW) (≤ 1 MW; > 1 MW); smoking during pregnancy (yes; no); number of previous pregnancies (< 2; ≥ 2); onset of prenatal care (during the 1st trimester; after the 1st trimester); number of prenatal consultations (≤ 3 ; > 3); history of miscarriage (yes; no); and health problems (yes; no) – anemia, asthma, tuberculosis, pneumonia, diabetes, hypertension, kidney disease, urinary tract infection, and/or bleeding (Table S1).

Statistical Analysis

Descriptive analyses were performed using central tendency and dispersion measures for quantitative variables and frequency distributions for qualitative variables. Percentages and 95% confidence intervals (95% CI) were used for figures. The normality of the data was verified using the Shapiro–Wilk test. Pearson's chi-square and Fisher's exact tests were used to examine the differences between strata. The analyses were stratified by race.

Prevalence ratios (PRs) and their 95% CIs were estimated using Poisson regression with robust error variance using each maternal race as an independent variable and medication use and potentially risky medication use as the dependent variables, categorized as binary variables (yes; no). Multivariate Poisson regression analysis included all variables associated with outcomes at a level lower than 20% in the bivariate analysis and those considered of interest according to the scientific literature. For all tests and the permanence of the variable in the final model, a significance level of 5% was assumed. The final model selection was carried out using backward elimination, consecutively eliminating those variables with lower statistical significance until only those with a significance level of 5% remained.

Analyses were conducted using R software, version 4.1.2²⁹ and the sandwich³⁰ and Imtest³¹ packages.

Ethics Statement

The NISAMI cohort study was conducted by the Federal University of Recôncavo da Bahia under the supervision of Professor Djanilson Barbosa dos Santos. At the time of conducting the research, there was no ethics review board established at this institution; therefore, the research had to be assessed regarding its ethics commitment at the Ethics and Research Committee of the Adventist Physiotherapy Faculty of Bahia (FAFIS), which is close to the Federal University of Recôncavo da Bahia. The research was approved under protocol #4369.0.000.070–10, and this study was conducted in accordance with the Declaration of Helsinki.

Pregnant women were informed about the research purpose, methodology, and confidentiality of the data through an informed consent form that contained explicit information about the nature and objectives of the study. For those aged < 18 years, their parents or guardians provided written informed consent for inclusion in the study. The participants were interviewed only after signing the form.

Results

Population Characteristics

The NISAMI Cohort study enrolled 1099 pregnant women between June 2012 and February 2014. Among these, 1091 had data regarding medication use; 1058 (97.0%) who reported their race during the interviews were included in the

study (Figure 1). Most pregnant women were brown (47.26%), followed by black (37.43%), white (9.64%), Asian (3.69%), and Brazilian indigenous (1.98%). Maternal age ranged from 13 to 45 years (Table S2). The median age of pregnant women of each race was 21 years (IQR = 8) among Brazilian indigenous individuals, 23 years (IQR = 9.5) among Asians, 25 years (IQR = 9) among Black individuals, and 26 years among Brown (IQR = 9) and White (IQR = 10) individuals.

Although not statistically significant, Asian women were more likely to have 24 years or less (51.28%) and less than eight years of education (61.54%) than women of other races, while black (94.44%) and brown (96.60%) women were more likely to smoke during pregnancy. Regarding prenatal assistance and obstetric history, up to three prenatal consultations were more frequent among brown women (63.20%), and a history of miscarriage and health problems were more prevalent among Brazilian indigenous individuals (33.33% and 61.90%, respectively) (Table 1).

Medication Use During Pregnancy

Overall, 84.31% of the women used at least one medication during pregnancy, with a median of 2.0 (IQR = 2) and a maximum of nine medications. However, pregnant white women reported a lower proportion of medication use than other women did (81.37%) (Supplementary Table S3).

Figure 2 shows the proportions of pregnant women who used one, two, or three or more medications at any time during pregnancy. Overall, 32.52% and 32.91% of pregnant women used two or three or more medications, respectively. A small proportion (20.51%) reported using only one medication. The proportion of white pregnant women using three or more medications was lower (32.32%) than that of Asian, black, Brazilian indigenous, and brown pregnant women.

According to the ATC classes, the most frequently reported class was antianemic preparations (B03), including iron preparations, vitamin B12, and folic acid, which were reported by 71.08% (95% CI 68.27–73.72%) of pregnant women.

Analgesics (N02) was the second most frequently reported class (21.74%; 95% CI 19.36–24.32%), followed by drugs for functional gastrointestinal (GI) disorders (A03) (18.81%; 95% CI 16.57–21.28%) and antihistamines for systemic use (R06) (8.60%; 95% CI 7.06–10.54%) (Figure 3, Table 2). Paracetamol was the most reported analgesic, while butylscopolamine, in combination with metamizole, was the main medicine used for functional GI disorders, and dimenhydrinate was used as an antihistamine for systemic use.

The use of antianemic preparations was less prevalent among white (68.63%; 95% CI 59.09–76.82%), black (69.19%; 95% CI 64.48–73.54%), and brown (71.80%; 95% CI 67.70–75.57%) pregnant women than among Asian (76.92%; 95% CI 61.66–87.35%) and Brazilian indigenous (90.48%; 95% CI 71.09–97.35%) pregnant women. Additionally, a lower

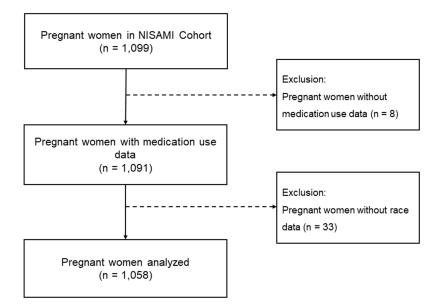


Figure I Study flow diagram.

Table I Characteristics of the Study Population by Race. NISAMI Cohort Study, Brazil, 2012–2014 (N = 1058)

	Asian (n = 39)			ack 396)	Indig	zilian enous = 21)		own 500)		hite : 102)	p value
	n	%	n	%	n	%	n	%	n	%	
Maternal age (years)											0.509 ^a
≤ 24	20	51.28	194	48.99	12	57.14	218	43.60	41	40.20	
25 to 29	9	23.08	97	24.49	5	23.81	141	28.20	32	31.37	
30 to 35	10	25.64	83	20.96	2	9.52	104	20.80	20	19.61	
≥ 36	0	0.00	22	5.56	2	9.52	37	7.40	8	7.84	
Missing	0	0.00	0	0.00	0	0.00	0	0.00	I	0.98	
Education (years)											0.326 ^a
≤ 8	24	61.54	190	47.98	10	47.62	241	48.20	57	55.88	
9 to 11	14	35.90	185	46.72	11	52.38	225	45.00	37	36.27	
>	I	2.56	19	4.80	0	0.00	33	6.60	8	7.84	
Missing	0	0.00	2	0.51	0	0.00	I	0.20	0	0.00	
Marital status											0.421 ^b
Without a partner	9	23.08	59	14.90	4	19.05	93	18.60	15	14.71	
With a partner	30	76.92	337	85.10	17	80.95	406	81.20	87	85.29	
Missing	0	0.00	0	0.00	0	0.00	I	0.20	0	0.00	
Employment status											0.339 ^a
Without employment	19	48.72	190	47.98	10	47.62	271	54.20	56	54.90	
With employment	19	48.72	203	51.26	11	52.38	222	44.40	45	44.12	
Missing	I	2.56	3	0.76	0	0.00	7	1.40	I	0.98	
Family income											0.847 ^b
≤ I MW	9	23.08	90	22.73	4	19.05	114	22.80	18	17.65	
> 1 MW	27	69.23	290	73.23	14	66.67	364	72.80	79	77.45	
Missing	3	7.69	16	4.04	3	14.29	22	4.40	5	4.90	
Smoking during pregnancy											0.096 ^b
No	3	7.69	17	4.29	I	4.76	11	2.20	5	4.90	
Yes	36	92.31	374	94.44	19	90.48	483	96.60	95	93.14	
Missing	0	0.00	5	1.26	I	4.76	6	1.20	2	1.96	
Previous pregnancies											0.630 ^b
< 2	32	82.05	307	77.53	19	90.48	394	78.80	76	74.51	
≥ 2	7	17.95	87	21.97	2	9.52	106	21.20	25	24.51	
Missing	0	0.00	2	0.51	0	0.00	0	0.00	I	0.98	
Onset of prenatal care											0.890 ^b
During the 1st trimester	30	76.92	307	77.53	18	85.71	398	79.60	81	79.41	
After the 1st trimester	8	20.51	67	16.92	3	14.29	77	15.40	15	14.71	
Missing	I	2.56	22	5.56	0	0.00	25	5.00	6	5.88	
Prenatal consultations											0.196 ^a
≤ 3 consultations	19	48.72	238	60.10	10	47.62	316	63.20	64	62.75	
> 3 consultations	20	51.28	147	37.12		52.38	175	35.00	36	35.29	
Missing	0	0.00	11	2.78	0	0.00	9	1.80	2	1.96	
History of miscarriage											0.457 ^b
No	28	71.79	255	64.39	10	47.62	330	66.00	66	64.71	
Yes	8	20.51	89	22.47	7	33.33	105	21.00	17	16.67	
Missing	3	7.69	52	13.13	4	19.05	65	13.00	19	18.63	
Health problems							-				0.080 ^a
No	27	69.23	191	48.23	8	38.10	245	49.00	49	48.04	
Yes	11	28.21	204	51.52	13	61.90	247	49.40	50	49.02	
Missing		2.56	1	0.25	0	0.00	8	1.60	3	2.94	1

 ${\bf Notes:}\ ^{a}p\mbox{-value}$ calculated by chi-square test; $^{b}p\mbox{-value}$ calculated by Fisher's exact test.

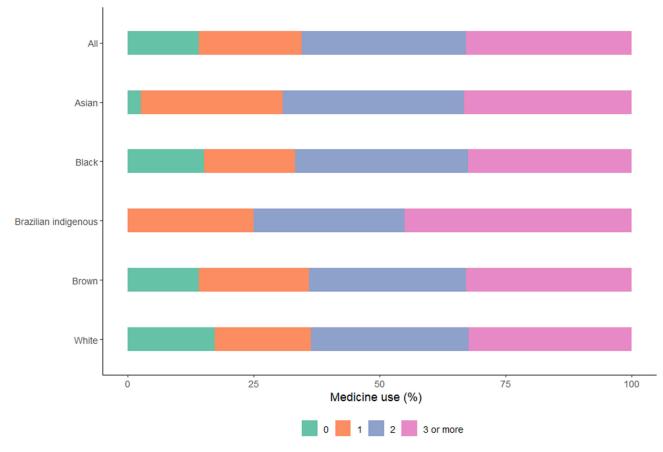


Figure 2 Percentage of patients who used medication during pregnancy by race. NISAMI Cohort Study, Brazil, 2012–2014 (n = 1058).

frequency of analgesic consumption was observed among white women; more specifically, 18.63% (95% CI 12.26–27.27%) of white pregnant women used analgesics during pregnancy versus 21.46% (95% CI 17.70–27.77%) of black pregnant women, 22.00% (95% CI 18.59–25.84%) of brown pregnant women, 25.64% (95% CI 14.57–41.08%) of Asian pregnant women, and 28.57% (95% CI 13.81–49.96%) of Brazilian indigenous women.

Furthermore, white women reported the use of drugs for functional GI disorders more frequently. Remarkably, 22.55% (95% CI 15.52–31.57%) of white women took drugs for functional GI disorders, 20.51% (95% CI 10.78–35.53%) of Asian women, 20.45% (95% CI 16.77–24.70%) of black women, 19.05% (95% CI 7.67–40.00%) of Brazilian indigenous women, and 16.60% (95% CI 13.60–20.11%) of brown women. In contrast, the consumption of antihistamines for systemic use was greater among Brazilian indigenous (9.52%; 95% CI 2.65–28.91%), brown (9.20%; 95% CI 6.97–12.05%), and black (8.59%; 95% CI 6.21–11.76%) pregnant women than among white (7.84%; 95% CI 4.03–14.72%) and Asian (2.56%; 95% CI 0.45–13.18%) pregnant women (Figure 3, Table 2).

The proportion of self-reported medication use decreased during pregnancy from 75.78% in the first trimester to 6.17% in the third trimester (Supplementary Table S4).

Potentially Risky Medication Use During Pregnancy

Among women who used medications during pregnancy, 29.26% (95% CI 26.37–32.33%) used at least one potentially risky medication. The most common potentially risky medications reported were butylscopolamine plus metamizole and isolated metamizole. Asian (31.58%; 95% CI 19.08–47.46%) and white (33.73%; 95% CI 24.48–44.42%) women used potentially risky medications during pregnancy more frequently than black (31.10%; 95% CI 26.33–36.30%), brown (27.19%; 95% CI 23.17–31.62%), and Brazilian indigenous women (20.00%; 95% CI 8.07–41.60%) (Figure 4).



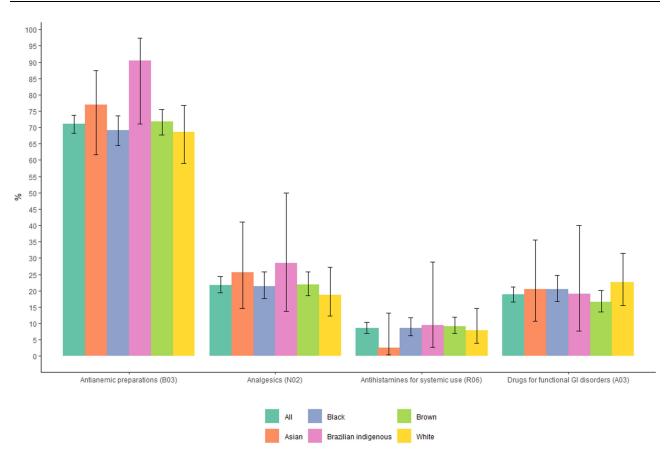


Figure 3 Percentage of the top medication classes taken during pregnancy by race. NISAMI Cohort Study, Brazil, 2012–2014 (n = 892).

Similarly to general medication use, potentially risky medication use during pregnancy decreased from 27.81% in the first trimester to 12.73% in the third trimester (Supplementary Table S5).

Factors Associated with Medication Use During Pregnancy

Table 3 shows the crude prevalence ratios of medication use stratified by race. Thirty-six years or more, access to more than three prenatal consultations, and health problems were associated with medication use by pregnant black women. At least 24 years of age, more than 11 years of education, smoking during pregnancy, onset of prenatal care after the first trimester, more than three prenatal consultations, history of miscarriage, and health problems were associated with medication use by brown women. In contrast, only more than 11 years of education and more than three prenatal consultations were associated with the use of medication by white women.

After the multivariate analysis, access to more than three prenatal consultations (PR 1.27; 95% CI 1.18–1.37) and health problems (PR 1.13; 95% CI 1.04–1.23) remained associated with medication use among black women. For brown women, more than 11 years of education (PR, 1.16; 95% CI 1.05–1.29), smoking during pregnancy (PR, 1.15; 95% CI 1.05–1.26), and more than three prenatal consultations (PR, 1.23; 95% CI 1.15–1.32) were associated with medication use in the multivariate analysis. For White women, the factors associated with medication use during pregnancy were more than 11 years of education (PR 1.18; 95% CI 1.01–1.38) and more than three prenatal consultations (PR 1.36; 95% CI 1.17–1.58) (Table 4).

Factors Associated with Potentially Risky Medication Use During Pregnancy

Variables statistically associated with potentially risky medication use during pregnancy by black women in the bivariate analysis were maternal age between 25 and 29 years and between 30 and 35 years, onset of prenatal care during the first trimester, and more than prenatal consultations. For brown women, the factors associated with the use of these

Table 2 Prevalence of the Use of at Least One Medication During Pregnancy According to Race According to the AnatomicalTherapeutic Chemical Classification System (ATC) Level. NISAMI Cohort Study, Brazil, 2012–2014 (n = 892)

ATC levels	Pregnant Women												
		asian = 39)		lack : 396)	Indi	azilian genous = 21)		own : 500)	White (n = 102)		p value		
	Ν	%	n	%	n	%	n	%	n	%			
Blood and blood forming organs – B	31	79.49	275	69.44	19	90.48	359	71.80	70	68.63	0.190 ^a		
Antianemic preparations – B03	30	76.92	274	69.19	19	90.48	359	71.80	70	68.63	0.231ª		
Iron preparations – B03A	21	53.85	209	52.78	15	71.43	257	51.40	45	44.12	0.207 ^a		
Vitamin B12 and folic acid – B03B	26	66.67	246	62.12	17	80.95	313	62.60	61	59.80	0.451ª		
Alimentary tract and metabolism – A	16	41.03	100	25.25	7	33.33	121	24.20	33	32.35	0.084 ^a		
Drugs for acid related disorders – A02	0	0.00	6	1.52	I.	4.76	12	2.40	2	1.96	0.560 ^b		
Antacids – A02A	0	0.00	4	1.01	1	4.76	10	2.00	Т	0.98	0.397 ^b		
Drugs for peptic ulcer and gastro-esophageal reflux disease (GORD) – A02B	0	0.00	2	0.51	0	0.00	2	0.40	I	0.98	0.642 ^b		
Drugs for functional gastrointestinal disorders – A03	8	20.51	81	20.45	4	19.05	83	16.60	23	22.55	0.466 ^b		
Belladonna and derivatives, plain – A03B	4	10.26	44	11.11	0	0.00	31	6.20	13	12.75	0.022 ^b		
Antispasmodics in combination with analgesics – A03D	3	7.69	28	7.07	3	14.29	31	6.20	8	7.84	0.549 ^b		
Antispasmodics and anticholinergics in combination with other drugs – A03E	I	2.56	19	4.80	0	0.00	20	4.00	I	0.98	0.459 ^b		
Propulsives – A03F	0	0.00	4	1.01	1	4.76	3	0.60	2	1.96	0.205 ^b		
Vitamins – Al I	8	20.51	25	6.31	3	14.29	39	7.80	9	8.82	0.032 ^b		
Multivitamins, combinations – AIIA	7	17.95	23	5.81	2	9.52	31	6.20	8	7.84	0.075 ^b		
Nervous system – N	10	25.64	86	21.72	6	28.57	110	22.00	19	18.63	0.790 ^b		
Analgesics – NO2	10	25.64	85	21.46	6	28.57	110	22.00	19	18.63	0.785 ^b		
Other analgesics and antipyretics – N02B	10	25.64	85	21.46	6	28.57	110	22.00	19	18.63	0.785 ^b		
Respiratory system – R	1	2.56	37	9.34	2	9.52	47	9.40	8	7.84	0.715 ^b		
Antihistamines for systemic use – R06	1	2.56	34	8.59	2	9.52	46	9.20	8	7.84	0.742 ^b		
Antihistamines for systemic use – R06A	1	2.56	34	8.59	2	9.52	46	9.20	8	7.84	0.742 ^b		
Genito urinary system and sex hormones - G	2	5.13	16	4.04	0	0.00	24	4.80	2	1.96	0.699 ^b		
Gynecological antiinfectives and antiseptics – G01	0	0.00	5	1.26	0	0.00	9	1.80	0	0.00	0.737 ^b		
Antiinfectives and antiseptics, excl. combinations with corticosteroids – G01A	0	0.00	5	1.26	0	0.00	9	1.80	0	0.00	0.737 ^b		
Sex hormones and modulators of the genital system – G03	2	5.13	11	2.78	0	0.00	15	3.00	2	1.96	0.823 ^b		
Estrogens – G03D	-	2.56	11	2.78	0	0.00	14	2.80	2	1.96	1.000 ^b		
Antiinfectives for systemic use – J	3	7.69	15	3.79	I I	4.76	12	2.40	4	3.92	0.227 ^b		
Antibacterials for systemic use – J01	3	7.69	14	3.54	1	4.76	12	2.40	3	2.94	0.266 ^b		
Beta-lactam antibacterials, penicillins – J01C	I I	2.56	8	2.02	0	0.00	5	1.00	2	1.96	0.483 ^b		
Other beta-lactam antibacterials – J01D	1	2.56	6	1.52	0	0.00	7	1.40	2	1.96	0.798 ^b		
Other antibacterials – J01X	i	2.56	0	0.00		4.76	3	0.60	0	0.00	0.021 ^b		
Cardiovascular system – C	1	2.56	10	2.53	0	0.00	13	2.60	0	0.00	0.499 ^b		
Antihypertensives - CO2	1	2.56	10	2.53	0	0.00	10	2.00	0	0.00	0.483 ^b		
Antiadrenergic agents, centrally acting – C02A	I	2.56	10	2.53	0	0.00	10	2.00	0	0.00	0.483 ^b		
Agents acting on the renin-angiotensin system – C09	0	0.00	0	0.00	0	0.00	1	0.20	0	0.00	1.000 ^b		
Musculo-skeletal system – M	I	2.56	9	2.27	0	0.00	9	1.80	3	2.94	0.781 ^b		
Anti-inflammatory and antirheumatic products – M01	1	2.56	4	1.01	0	0.00	5	1.00	2	1.96	0.538 ^b		
Anti-inflammatory and antirheumatic products, nonsteroids – M01A	I	2.56	4	1.01	0	0.00	5	1.00	2	1.96	0.538 ^b		
Muscle relaxants – M03	0	0.00	5	1.26	0	0.00	4	0.80	1	0.98	0.850 ^b		
Muscle relaxants, centrally acting agents – M03B	0	0.00	5	1.26	0	0.00	4	0.80		0.98	0.850 ^b		
Other	I	2.56	I	0.25	0	0.00	6	1.20	2	1.96	0.144 ^b		

Notes: ^ap-values were calculated by the chi-square test; ^bp-values were calculated by Fisher's exact test.

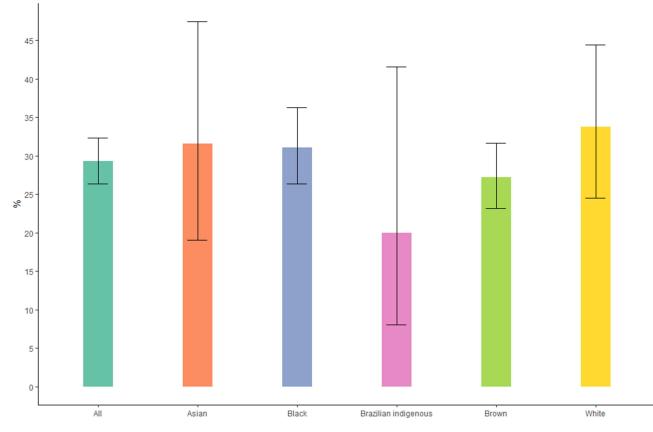


Figure 4 Percentage of potentially risky medication use during pregnancy by race. NISAMI Cohort Study, Brazil, 2012–2014 (n = 261).

medications in the bivariate analysis were maternal age between 25 and 29 years, age between 30 and 35 years, age 36 years or older, employment status, smoking during pregnancy, and more than three prenatal consultations. In addition, two or more previous pregnancies and a history of miscarriage were associated with potentially risky medication use during pregnancy among white women (Table 5).

A greater prevalence of potentially risky medication use during pregnancy among black women aged between 25 and 29 years (PR 1.52; 95% CI 1.01–2.30) and 30 to 35 years (PR 1.74; 95% CI 1.18–2.56) was found in the multivariate analysis. Among brown women, those aged between 25 and 29 years (PR 1.70; 95% CI 1.14–2.53), 30 to 35 years (PR 1.68; 95% CI 1.09–2.58), and 36 years or more (PR 2.37; 95% CI 1.41–3.96) and those who smoked during pregnancy (PR 2.27; 95% CI 1.34–3.84) were identified via multivariate analysis. Moreover, two or more previous pregnancies were associated with potentially risky medication use during pregnancy among white women (PR 2.34; 95% CI 1.34–4.08) (Table 6).

Discussion

The use of medication was highly prevalent in this prospective cohort of pregnant Brazilian women. Approximately 84% of women reported using at least one medication during pregnancy, and approximately one-third reported using at least three medications at any time during pregnancy. Overall, white women reported a lower proportion of medication consumption than Asian, black, Brazilian indigenous, and brown women. The most commonly reported medications were antianemic preparations, analgesics, drugs for functional GI disorders, and antihistamines for systemic use. Additionally, almost 29.3% of women took potentially risky medications during pregnancy, and brown women did. Factors significantly associated with the use of medications during pregnancy included a greater number of prenatal consultations, a higher education level, health problems, and smoking. Additionally, maternal age above 25 years,

Maternal age (years) ≤ 24 1925 to 29930 to 3510 ≥ 36 0Missing0Education (years) \leq ≤ 8 249 to 1113> 111Missing0Marital statusWithout a partner9With a partner29	9 5 9 2 0 2 0 (0 2 4 6 3 3 1 2 0 (0	% 50.00 23.68 26.32 0.00 0.00 63.16 34.21 2.63 0.00	PR 1.00 1.05 1.05 - - 1.00 0.93 1.00	95% CI - 0.95–1.17 0.95–1.17 - - 0.80–1.08	n 162 71 74 21 0	% 49.39 21.65 22.56 6.40 0.00	PR 1.00 0.88 1.07 1.14	95% IC 0.76-1.01 0.97-1.18 1.02-1.28	n 5	% 55.00 25.00	PR	95% IC	n 180	%	PR	95% IC	n	%	PR	95% IC
≤ 24 9 25 to 29 9 30 to 35 0 ≥ 36 0 Missing 0 Education (years) ≤ 8 24 9 to 11 3 > 11 1 Missing 0 Marital status Without a partner 9	9 2 0 2 0 0 0 0 24 6 3 3 1 2 0 0	23.68 26.32 0.00 0.00 63.16 34.21 2.63	1.05 1.05 - - 1.00 0.93	0.95–1.17 0.95–1.17 – –	71 74 21 0	21.65 22.56 6.40	0.88 1.07	0.97-1.18	5			_	180							
25 to 29 9 30 to 35 10 ≥ 36 0 Missing 0 Education (years) 2 ≤ 8 24 9 to 11 13 > 11 1 Missing 0 Marital status 9 Without a partner 9	9 2 0 2 0 0 0 0 24 6 3 3 1 2 0 0	23.68 26.32 0.00 0.00 63.16 34.21 2.63	1.05 1.05 - - 1.00 0.93	0.95–1.17 0.95–1.17 – –	71 74 21 0	21.65 22.56 6.40	0.88 1.07	0.97-1.18	5			_	180							
$30 \text{ to } 35$ 10 ≥ 36 0 Missing 0 Education (years) ≤ 8 24 $9 \text{ to } 11$ 13 > 11 1 Missing 0 Marital status 0 Without a partner 9	0 2 0 (0 (24 6 3 3 1 2 0 (26.32 0.00 0.00 63.16 34.21 2.63	1.05 - - 1.00 0.93	0.95–1.17 – –	74 21 0	22.56 6.40	1.07	0.97-1.18	-	25.00			100	42.55	1.00	-	33	39.76	1.00	-
≥ 36 0 Missing 0 Education (years) 2 ≤ 8 24 9 to 11 13 > 11 1 Missing 0 Marital status 9 Without a partner 9	0 (0 0 (0 24 6 3 3 1 2 0 (0	0.00 0.00 63.16 34.21 2.63	- - 1.00 0.93	-	21 0	6.40			~	20.00	1.09	0.90-1.32	127	30.02	1.09	1.01-1.18	28	33.73	1.09	0.89-1.33
Missing 0 Education (years) 24 ≤ 8 24 9 to 11 13 > 11 1 Missing 0 Marital status Without a partner	0 (24 6 3 3 1 2 0 (0.00 63.16 34.21 2.63	- 1.00 0.93	_	0		1.14	1 02 1 20	2	10.00	1.09	0.90-1.32	86	20.33	1.00	0.90-1.11	15	18.07	0.93	0.69-1.26
Education (years) 24 ≤ 8 24 9 to 13 > Missing 0 Marital status Without a partner	24 6 3 3 1 2 0 (63.16 34.21 2.63	1.00 0.93	_		0.00		1.02-1.20	2	10.00	1.09	0.90-1.32	30	7.09	0.98	0.83-1.16	6	7.23	0.93	0.60-1.44
≤ 8 24 9 to 11 13 > 11 1 Missing 0 Marital status Without a partner 9	3 3 I 2 0 (34.21 2.63	0.93		152		-	-	0	0.00	-	-	0	0.00	-	_	Т	1.20	-	-
9 to II 13 > II I Missing 0 Marital status Vithout a partner	3 3 I 2 0 (34.21 2.63	0.93		152															
> II I I Missing 0 Marital status Without a partner 9	0 0	2.63		0.80-1.08	1,92	46.34	1.00	-	9	45.00	1.00	-	197	46.57	1.00	_	46	55.42	1.00	-
Missing 0 Marital status Without a partner 9	0 0		1.00	0.00-1.00	159	48.48	1.07	0.98-1.18	П	55.00	1.11	0.89-1.38	194	45.86	1.05	0.97-1.14	29	34.94	0.97	0.78-1.20
Marital status Without a partner 9		0.00	1.00	1.00-1.00	15	4.57	0.99	0.77-1.26	0	0.00	-	_	32	7.57	1.18	1.09-1.29	8	9.64	1.24	1.09-1.41
Without a partner 9		0.00	-	_	2	0.61	_	_	0	0.00	-	_	0	0.00	-	_	0	0.00	-	_
With a partner 29	9 2	23.68	1.00	-	49	14.94	1.00	-	3	15.00	1.00	-	75	17.73	1.00	_	8	9.64	1.00	-
	9 7	76.32	0.97	0.90-1.03	279	85.06	1.00	0.88-1.13	17	85.00	1.33	0.73-2.42	347	82.03	1.06	0.95-1.18	75	90.36	1.62	0.99–2.63
Missing 0	0 0	0.00	-	_	0	0.00	_	_	0	0.00	-	_	Т	0.24	-	_	0	0.00	-	_
Employment status																				
Without employment 18	8 4	47.37	1.00	_	153	46.65	1.00	_	10	50.00	1.00	_	231	54.61	1.00	_	46	55.42	1.00	_
With employment 19	9 5	50.00	1.05	0.95-1.18	173	52.74	1.06	0.97-1.16	10	50.00	0.91	0.75-1.11	188	44.44	0.99	0.92-1.07	36	43.37	0.97	0.80-1.18
Missing I	1 2	2.63	_	_	2	0.61	_	_	0	0.00	_	_	4	0.95	_	_	I	1.20	_	_
Family income																				
≤ I MW 8	8 2	21.05	1.00	_	72	21.95	1.00	_	3	15.00	1.00	_	97	22.93	1.00	_	14	16.87	1.00	_
> I MW 27	7 7	71.05	1.12	0.89-1.43	244	74.39	1.05	0.94-1.18	14	70.00	1.33	0.73-2.43	306	72.34	0.99	0.90-1.08	67	80.72	1.09	0.83-1.42
Missing 3	3 7	7.89	_	_	12	3.66	_	_	3	15.00	_	_	20	4.73	_	_	2	2.41	_	_
Smoking during pregnancy																				
No 35	5 9	92.11	1.00	_	309	94.21	1.00	_	18	90.00	1.00	_	407	96.22	1.00	_	78	93.98	1.00	_
Yes 3	3 7	7.89	1.03	0.97-1.09	14	4.27	1.00	0.80-1.25	I	5.00	1.06	0.94-1.18	П	2.60	1.19	1.14-1.23	4	4.82	0.97	0.62-1.53
Missing 0	0 0	0.00	_	_	5	1.52	_	_	I	5.00	_	_	5	1.18	_	_	I	1.20	_	_
Previous pregnancies																				
< 2 31	8 1	81.58	1.00	_	255	77.74	1.00	_	18	90.00	1.00	_	336	79.43	1.00	_	65	78.31	1.00	_
≥ 2 7		18.42	1.03	0.97-1.10	71	21.65	0.98	0.88-1.10	2	10.00	1.06	0.94-1.18	87	20.57	0.96	0.87-1.06	18	21.69	0.84	0.65-1.10
Missing 0		0.00	_	_	2	0.61	_	_	0	0.00	_	_	0	0.00	_	_	0	0.00	_	_
Onset of prenatal care																				
During the 1st trimester 29	9 7	76.32	1.00	_	261	79.57	1.00	_	17	85.00	1.00	_	349	82.51	1.00	_	71	85.54	1.00	_
After the 1st trimester 8		21.05	1.03	0.97-1.11	54	16.46	0.95	0.83-1.08	3	15.00	1.06	0.94-1.19	58	13.71	0.86	0.75–0.98	10	12.05	0.76	0.52-1.10
Missing		2.63	_	_	13	3.96	_	_	0	0.00	_	_	16	3.78	_	_	2	2.41	_	_
Prenatal consultations					_				-				-							
\leq 3 consultations 18	8 4	47.37	1.00	_	180	54.88	1.00	_	9	45.00	1.00	_	248	58.63	1.00	_	47	56.63	1.00	_
> 3 consultations 20		52.63	1.05	0.95-1.18	144	43.90	1.30	1.20-1.40	, LI	55.00	1.11	0.89-1.38	169	39.95	1.23	1.15-1.31	36	43.37	1.36	1.17-1.58
Missing 0		0.00	_	-	4	1.22	-	_	0	0.00	_	_	6	1.42	-	_	0	0.00	-	_

Table 3 Crude Prevalence Ratio of Medication Use According to Race by Study Population Characteristics. NISAMI Cohort Study, Brazil, 2012–2014 (n = 892)

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(Continued)

Table 3 (Continued).

		Asian (n = 38)				Black	c (n = 32	28)	Bra	zilian In	ligenou	is (n = 20)	Brown (n = 423)					White (n = 83)				
	n	%	PR	95% CI	n	%	PR	95% IC	n	%	PR	95% IC	n	%	PR	95% IC	n	%	PR	95% IC		
History of miscarriage																						
No	27	71.05	1.00	-	217	66.16	1.00	-	10	50.00	1.00	-	271	64.07	1.00	-	54	65.06	1.00	-		
Yes	8	21.05	1.04	0.96-1.12	71	21.65	0.94	0.83-1.05	6	30.00	0.86	0.62-1.18	94	22.22	1.09	1.01-1.18	12	14.46	0.86	0.62-1.20		
Missing	3	7.89	-	-	40	12.20	-	-	4	20.00	-	-	58	13.71	-	-	17	20.48	-	-		
Health problems																						
No	26	68.42	1.00	-	147	44.82	1.00	-	7	35.00	1.00	-	199	47.04	1.00	-	37	44.58	1.00	-		
Yes	П	28.95	1.04	0.96-1.12	181	55.18	1.15	1.05-1.26	13	65.00	1.14	0.87-1.50	217	51.30	1.08	1.01-1.17	44	53.01	1.17	0.96-1.41		
Missing	Ι	2.63	-	-	0	0.00	-	-	0	0.00	-	_	7	1.65	-	-	2	2.41	-	-		

Note: Values in bold indicate statistical significance.

		Black	E	Brown	,	White		
	PR	95% CI	PR	95% IC	PR	95% IC		
Medication use								
Education (years)			1.00	-	1.00	-		
≤ 8			1.03	0.94-1.12	0.93	0.76-1.13		
9 to II			1.16	1.05-1.29	1.18	1.01-1.38		
>								
Smoking during pregnancy								
No			1.00	-				
Yes			1.15	1.05-1.26				
Prenatal consultations								
≤ 3 consultations	1.00	-	1.00	-	1.00	-		
> 3 consultations	1.27	1.18-1.37	1.23	1.15-1.32	1.36	1.17-1.58		
History of miscarriage								
No			1.00	-				
Yes			1.09	1.01-1.19				
Health problems								
No	1.00	-						
Yes	1.13	1.04-1.23						
Residual deviance		103.81		116.15		28.65		
Degrees-of-freedom		381		415	96			
Þ		0.999		0.999	0.999			

Table 4 Factors Associated with Medication Use During Pregnancy According to Race inthe Study Population. NISAMI Cohort Study, Brazil, 2012–2014

Notes: Values in bold indicate statistical significance.

smoking status, and two or more previous pregnancies were associated with potentially risky medication use during pregnancy.

The overall prevalence of medication use reported by pregnant women in this study was lower than that previously reported in another Brazilian study. Lutz et al⁹ observed that 92.5% of pregnant women used at least one medication in the Pelotas Birth Cohort Study in southern Brazil. The variation in prevalence between studies may be related to differences in study designs and ways to collect and analyze data. Additionally, region-specific healthcare, socio-economic, and cultural characteristics may be important when comparing different Brazilian regions. Worldwide, the variations in medication use by pregnant women may be even greater, with the prevalence of at least one medicine used reaching 97% in the United States.⁸ In Europe, a Norwegian population-based cohort revealed that 60% of pregnant women used prescription medications between 2005 and 2015.⁵ In contrast, a Chinese population-based cohort revealed that only 11.1% of women used at least one medicine during pregnancy.³²

The lower proportion of women who consumed medication during pregnancy among white women found in our study is different from the findings of several previous studies.^{8,9,22} Studies have previously demonstrated that limited health literacy and reduced access to healthcare lead to increased self-medication,^{33,34} which is commonly practiced by pregnant women due to better accessibility, affordability, and availability of medicines.³⁵ In this study, medication use included both prescribed drugs and self-medication. Consequently, the higher medication use rates among black and brown women may be related to greater self-medication due to limited access to healthcare and health literacy, which are typically lower among black and brown individuals than among white individuals.^{36–38}

Nonetheless, racial differences in medication use prevalence with lower medication use in pregnancy among black women than among women of other races were observed in Brazil and the United States.^{8,9,22} Racism leads to socioeconomic disadvantages that extrapolate and extend to health inequities, making access to health services more

Table 5 Crude Prevalence Ratio of Potentially Risky Medication Use During Pregnancy According to Race According to Study Population Characteristics. NISAMI Cohort Study, Brazil,
2012–2014 (n = 261)

		Asia	n (n =	12)		Blac	k (n = I	02)	В	razilian l	ndigeno	ous (n = 4)		Brow	n (n = I	15)		Wh	ite (n =	28)
	n	%	PR	95% CI	n	%	PR	95% IC	n	%	PR	95% IC	n	%	PR	95% IC	n	%	PR	95% IC
Maternal age (years)																				
≤ 24	7	58.33	1.00	-	39	38.24	1.00	-	1	25.00	1.00	-	34	29.57	1.00	-	9	32.14	1.00	-
25 to 29	2	16.67	0.60	0.15-2.48	26	25.49	1.52	1.01-2.30	1	25.00	2.20	0.12-38.60	40	34.78	1.67	1.12-2.48	12	42.86	1.57	0.76-3.2
30 to 35	3	25.00	0.81	0.25-2.60	31	30.39	1.74	1.18-2.56	1	25.00	5.50	0.41-74.14	28	24.35	1.72	1.12-2.65	3	10.71	0.73	0.22-2.4
≥ 36	0	0.00	-	_	6	5.88	1.19	0.57-2.47	Т	25.00	5.50	0.41-74.14	13	11.30	2.29	1.38-3.83	3	10.71	1.83	0.67-4.9
Missing	0	0.00	-	_	0	0.00	-	-	0	0.00	-	-	0	0.00	-	_	Т	3.57	-	_
Education (years)																				
≤ 8	3	25.00	1.00	_	47	46.08	1.00	-	2	50.00	1.00	-	50	43.48	1.00	_	17	60.71	1.00	-
9 to 11	9	75.00	5.54	0.80-1.08	50	49.02	1.02	0.73-1.42	2	50.00	0.82	0.13-5.18	56	48.70	1.14	0.82-1.58	10	35.71	0.93	0.49-1.7
>	0	0.00	_	_	5	4.90	1.08	0.50-2.30	0	0.00	_	_	9	7.83	1.11	0.60-2.03	T	3.57	0.34	0.05-2.2
Missing	0	0.00	_	_	0	0.00	_	-	0	0.00	_	_	0	0.00	_	_	0	0.00	_	_
Marital status																				
Without a partner	5	41.67	1.00	_	15	14.71	1.00	-	1	25.00	1.00	_	16	13.91	1.00	_	2	7.14	1.00	-
With a partner	7	58.33	0.43	0.18-1.06	87	85.29	1.02	0.64-1.61	3	75.00	0.53	0.07-3.93	99	86.09	1.34	0.84-2.13	26	92.86	1.39	0.39-4.8
Missing	0	0.00	_	_	0	0.00	_	_	0	0.00	_	_	0	0.00	_	_	0	0.00	_	_
Employment status													-				-			
Without employment	6	50.00	1.00	_	46	45.10	1.00	_		25.00	1.00	_	54	46.96	1.00	_	14	50.00	1.00	_
With employment	6	50.00	0.95	0.36-2.46	56	54.90	1.08	0.78-1.49	3	75.00	3.00	0.33-27.06	60	52.17	1.37	1.00-1.87	13	46.43	1.19	0.64-2.2
Missing	0	0.00	_	_	0	0.00	_	_	0	0.00	_	_	1	0.87	_	_	1	3.57	_	_
Family income	-				-				-				-							
≤ I MW	3	25.00	1.00	_	19	18.63	1.00	_	1	25.00	1.00	_	23	20.00	1.00	_	4	14.29	1.00	_
> I MW	8	66.67	0.79	0.26–2.37	79	77.45	1.23	0.80-1.88	3	75.00	0.64	0.09-4.80	89	77.39	1.23	0.82-1.83	24	85.71	1.25	0.51-3.0
Missing	I	8.33	-	_	4	3.92	_	_	0	0.00	_	_	3	2.61	_	_	0	0.00	_	_
Smoking during pregnancy		0.00				0.72				0.00			Ĵ	2.01			Ĩ	0.00		
No	10	83.33	1.00	_	95	93.14	1.00	_	4	100.00	1.00	_	108	93.91	1.00	_	27	96.43	1.00	_
Yes	2	16.67	2.33	0.87–6.23	5	4.90	1.16	0.56-2.40	0	0.00	_	_	6	5.22	2.06	1.17-3.61	0	0.00	_	_
Missing	0	0.00	_	_	2	1.96	-	-	0	0.00	_	_	Ĩ	0.87		_	ĭ	3.57	_	_
Previous pregnancies	· ·	0.00			-					0.00				0.07				0.07		
< 2	П	91.67	1.00	_	80	78.43	1.00	_	3	75.00	1.00	_	86	74.78	1.00	_	17	60.71	1.00	_
≥ 2		8.33	0.40	0.06–2.77	22	21.57	0.99	0.67-1.46	,	25.00	3.00	0.48-18.55	29	25.22	1.30	0.92-1.85		39.29	2.34	1.34-4.0
Missing	0	0.00	-	_	0	0.00	_	_	0	0.00	-	_	0	0.00	-	_	0	0.00	-	
Onset of prenatal care	Ŭ	0.00			Ŭ	0.00			Ů	0.00			Ů	0.00			Ŭ	0.00		
During the 1st trimester	П	91.67	1.00	_	85	83.33	1.00	_	4	100.00	1.00	_	97	84.35	1.00	_	24	85.71	1.00	_
After the 1st trimester		8.33	0.33	0.05–2.30	11	10.78	0.62	0.36-1.09	0	0.00	-	_	12	10.43	0.74	_ 0.44_1.27	24	7.14	0.59	0.16-2.1
Missing	0	0.00	0.33	0.03-2.30	6	5.88	0.62	0.36-1.07	0	0.00	_	_	6	5.22	0.74	0.77-1.27	2	7.14	0.57	0.10-2.1
Prenatal consultations	0	0.00	_	_		5.00	-	_		0.00		_	0	5.22	_	_	_	7.17	-	_
\leq 3 consultations	7	58.33	1.00		62	60.78	1.00			25.00	1.00		74	64.35	1.00	_	14	50.00	1.00	
	5	58.33 41.67	0.64	- 0.24-1.71	62 38	37.25	0.77	- 0.54–1.08	2	75.00	2.45	- 0.27–22.10	74 39	33.91	0.77	- 0.55–1.08	14	50.00	1.00	- 0.71-2.4
> 3 consultations		41.67 0.00				37.25			3	0.00			39 2							
Missing	0	0.00	-	-	2	1.96	-	-	0	0.00	-	-	2	1.74	-	-	0	0.00	-	-

Dovepress

Castro et al

History of miscarriage																				
No	9	75.00	1.00	-	66	64.71	1.00	-	2	50.00	1.00	-	73	63.48	1.00	-	16	57.14	1.00	-
Yes	2	16.67	0.75	0.19-2.90	25	24.51	1.16	0.79–1.68	Т	25.00	0.83	0.08-8.54	28	24.35	1.11	0.77-1.60	6	21.43	1.69	0.83–2.43
Missing	Т	8.33	-	-	П	10.78	-	-	Т	25.00	-	-	14	12.17	-	-	6	21.43	-	-
Health problems [†]																				
No	6	50.00	1.00	-	41	40.20	1.00	-	Т	25.00	1.00	-	50	43.48	1.00	-	13	46.43	1.00	-
Yes	5	41.67	1.97	0.74–5.26	61	59.80	1.21	0.87–1.68	3	75.00	1.61	0.18-14.29	63	54.78	1.15	0.84–1.59	14	50.00	0.91	0.49-1.69
Missing	Т	8.33	-	-	0	0.00	-	-	0	0.00	-	-	2	1.74	-	-	Т	3.57	-	-

Note: Values in bold indicate statistical significance.

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		Black	I	Brown	White			
	PR	95% CI	PR	95% IC	PR	95% IC		
Potentially risky medication use in pregnancy								
Maternal age (years)								
≤ 24	1.00	-	1.00	-				
25 to 29	1.52	1.01-2.30	1.70	1.14-2.53				
30 to 35	1.74	1.18-2.56	1.68	1.09-2.58				
≥ 36	1.19	0.57-2.47	2.37	1.41-3.96				
Smoking during pregnancy								
No			1.00	-				
Yes			2.27	1.34-3.84				
Previous pregnancies								
< 2					1.00	-		
≥ 2					2.34	I.34–4.08		
Residual deviance		232.29		284.40		56.43		
Degrees-of-freedom		324		413		81		
Þ		0.999		0.999 0		0.983		

Table 6 Factors Associated with Potentially Risky Medication Use During Pregnancy According to Race in the Study Population. NISAMI Cohort Study, Brazil, 2012–2014

Notes: Values in bold indicate statistical significance.

difficult and increasing the mortality of minority groups.³⁹ Moreover, institutional racism in health services predominantly affects black people¹⁴ and may result in lower use of medications by black women during pregnancy.

Furthermore, a greater prevalence of use of three or more medications among Brazilian indigenous women was observed in this study. This is possibly related to the increasing biomedical hegemony and overvaluation of the use of medications in Brazilian indigenous health care,⁴⁰ contributing to this finding.

Antianemic preparations were the most commonly used medications for pregnant women, reflecting the adoption of international and national prenatal care recommendations. The supplementation of folic acid, iron, and vitamins during pregnancy, in addition to the adjustment of diet pattern and quality, is important for fulfilling increased nutritional demands during pregnancy, preventing adverse pregnancy outcomes, such as maternal anemia and low birth weight, and promoting fetal development.^{41,42}

The present study revealed a lower prevalence of antianemic preparations among pregnant white and black women. As long as white women have a lower risk of developing anemia during pregnancy,⁴³ this may have impacted the reduced prescription of these supplements to those women. In addition, these results may suggest lower access to health services and inadequate prenatal care among Black women. These findings are concerning, especially for pregnant black women, who are more likely to have anemia than women of other races⁴⁴ due to genetic or unfavorable socioeconomic conditions.³⁹ Previous studies have reported differences in iron preparations, folic acid, and vitamin use during pregnancy according to race, where black and Hispanic women present lower rates of intake of these supplements.^{22,45} Iron supplements, folic acid, and vitamins may improve pregnancy outcomes and minimize the illness burden and cost.⁴⁶ Therefore, further research must be conducted to assess the outcomes related to the lower prevalence of use of these medicines by pregnant women of different races.

Analgesics were used by 21.74% of pregnant women, and paracetamol was the most commonly reported medicine in this group, either alone or in combination. Pregnant women widely use analgesic medications, particularly paracetamol, to treat fever and pain. Despite controversies in the reporting literature about the long-term effects of paracetamol,⁴⁷ it remains a safe analgesic and antipyretic recommended during pregnancy and breastfeeding.⁴⁸

Additionally, a greater frequency of analgesic consumption was observed among Brazilian indigenous women. As discussed before, this may be associated with the medicalization of Brazilian indigenous healthcare in the last few years.

Furthermore, since these medications are available by prescription and over the counter, this finding may be related to self-medication, which may pose risks to the mother and the fetus.

Almost 19% of the women in this cohort used drugs for functional GI disorders, mainly butylscopolamine and metamizole. Butylscopolamine is used to relieve colic, abdominal, and pelvic pain, which is common during pregnancy. Similar to the findings of this research, a high prevalence of butylscopolamine use by pregnant women was previously described in Brazilian studies.^{4,49}

An interesting finding in our study was the relatively low prevalence of antihistamines of systemic use. These medications, predominantly dimenhydrinate, were used by 8.6% of pregnant women. Antihistamines are used to treat allergies, relieve indigestion, and prevent nausea and vomiting during pregnancy.⁵⁰ Dimenhydrinate is commonly used to manage nausea and vomiting during pregnancy and is commonly used as an over-the-counter medication. It acts by blocking H1 receptors in the gastrointestinal tract and decreases stimulation of the vomiting reflex by acting indirectly on the body's vestibular center.⁵¹ A significant variation in the prevalence of antihistamine use during pregnancy has been reported, with antihistamine consumption during pregnancy of 4.5% in Denmark,⁵² 13.7% in Norway,⁵ 17.0% in the United States,²² and 23.3% in Brazil.⁹

Approximately 29% of the women analyzed used at least one potentially risky medication during pregnancy, mainly butylscopolamine combined with metamizole and metamizole alone. Metamizole is an analgesic that is extensively consumed in Brazil but is not available in some European countries or North America because of its association with serious adverse events.²⁶ Additionally, a similar prevalence of the use of potentially risky medications during pregnancy has been reported. For example, web-based research conducted in South America, Europe, North America, and Australia reported that 28% of women used potentially risky medication during pregnancy.⁵³ Likewise, an Ethiopian crosssectional study using clinical charts reported a prevalence of potentially risky medication dispensing of 20%.⁵⁴ Nonetheless, interpretation of the prevalence of consumption of risky medications during pregnancy must be made in light of how the information on medications was obtained (through questionnaires and clinical charts), cultural differences, and the adopted risk classification systems. In addition, surveillance is needed considering the lack of studies on the safety, rationality, and risk-benefit associated with its use by pregnant women.

With regard to medication consumption throughout pregnancy, a decrease in all the medications analyzed was observed. Nonetheless, the scientific literature regarding this finding remains controversial. Similar to our results, a Brazilian prospective cohort revealed a reduction in medication use during pregnancy, from 54.2% to 53.2% and 47.4% during the first, second, and third trimesters, respectively.⁹ In contrast, Trønnes et al⁵³ reported a prevalence of medication consumption of 5.4% in the first trimester, 4.5% in the second trimester, and 6.0% in the third trimester among pregnant women in Europe. Moreover, an Italian cross-sectional study reported that 23.8% of women used some medication in the first trimester, 40.3% in the second trimester, and 14.2% in the third trimester.⁵⁵

In this study, medication use was more prevalent among Black, Brown, and White women who had access to more than three prenatal consultations. This is an expected finding, as more consultations may lead to more access to prescriptions and, consequently, facilitate access to medications. However, a careful interpretation of the rationality of medication use is needed. For example, butylscopolamine combined with metamizole is classified as a potentially risky medication, and despite being labeled as a prescription medication in Brazil, it is usually acquired without a prescription. Further studies need to be conducted to understand whether these medications are prescribed by a health professional or acquired for self-medication.

Furthermore, more than 11 years of education increased the prevalence of medication use during pregnancy by 16% (95% CI 1.05–1.29) among brown women and 18% (95% CI 1.01–1.38) among white women in this cohort. Previous studies have also shown higher rates of medication use during pregnancy among women with higher education.^{8,56,57} In addition, an increase in education can increase socioeconomic status, allowing women to increase their medication consumption.⁵⁶ However, the remaining question is related to whether those who need access to essential medication during pregnancy have access to them.

Additionally, higher rates of medication use during pregnancy have been associated with health problems,³ mainly due to the increase in women with chronic diseases.⁵⁸ The present study revealed that a higher prevalence of medication

use was observed in black women with health problems (PR 1.13; 95% CI 1.04–1.23). In addition, we observed a high prevalence of pain management in this population, which is usually due to acute conditions.

Among brown women, smoking during pregnancy increased the prevalence of medication use by 15% (95% CI 1.05– 1.26). A population-based cohort from Canada between 2002 and 2011 with 225,973 women reported comparable findings. The authors observed that smoking during pregnancy increased the odds of prescription medication use during pregnancy (OR 1.17; 95% CI 1.13–1.21).³

Regarding potentially risky medication during pregnancy, it was observed that a greater prevalence of the use of these medications increases with maternal age among black and brown women. A previous study elucidated the influence of age on the use of any potentially inappropriate medication during pregnancy (category D and X medications only), primarily among women aged at least 35 years.⁵⁹ This is probably related to the increased prevalence of preexisting chronic diseases, such as diabetes, hypertension, asthma, and depression, and increased risk of obstetric complications, such as nausea and vomiting, gestational diabetes, and preeclampsia, that require pharmacotherapy and may lead to the use of potentially risky medications during pregnancy.⁶⁰

Moreover, among white women, two or more previous pregnancies were associated with potentially risky medication use during pregnancy. This finding may suggest more experience with common pregnancy symptoms and less concern about risks associated with medication use during pregnancy.⁶¹

An unexpected finding was the apparent lack of association between the use of potentially risky medications during pregnancy and health problems (comorbidities). This finding differs from those of previous studies, in which pregnant women with health problems, especially chronic diseases, were more likely to use potentially risky medications during pregnancy than those without these problems.^{53,62} Therefore, further examination must be conducted to identify the specific needs and medicines prescribed for women with comorbidities.

The strengths of our study are the control of the quality of the questionnaires, standardization of data collection, and data validation. Another difference of this study was its prospective design, in which interviews were conducted during the prenatal period rather than only at delivery.

Furthermore, another strength of this study is the adoption of Poisson regression with robust variance, which provides correct estimates and is a better alternative for the analysis of cross-sectional data with binary outcomes than logistic regression since the prevalence ratio is more interpretable and easier to communicate than the odds ratio.⁶³

Nevertheless, these limitations cannot be ruled out. Considering that medications may be consumed by prescription and/or self-medication, medication use during pregnancy may be underestimated owing to recall bias and underreporting. In addition, women may not correctly report self-medication and may try to answer what is socially acceptable.

Furthermore, information on the medications' clinical indications was not collected; therefore, there is a possibility of indication bias in the study. In addition, medication use during pregnancy in the present study may not indicate better healthcare because data on pregnant women's previous health status were unavailable.

Additionally, this study used the FDA Pregnancy Risk Categories to categorize potentially risky medications during pregnancy. This classification is still used by ANVISA despite the recommendation to be discontinued in 2015. This categorization may be inappropriate for our population and may not reflect the risks to which pregnant women are exposed.

In addition, because this study was conducted exclusively in the urban area of the city due to the challenges encountered in collecting data from rural areas and only included pregnant women who visited basic health units, it may not be representative of all pregnant women in the city.

To promote health equity in prenatal care, it is crucial to address systemic racism and socioeconomic disparities, which are significant barriers to accessing health care services. Strategies are needed to address unconscious biases among health care workers, improve health literacy, and increase access to prenatal care. It is also important to enhance monitoring and education on drug safety, provide in-depth medication counseling, and promote counseling with healthcare professionals before initiating or discontinuing medication during pregnancy. Public health campaigns and educational programs should target pregnant women to educate them on decision-making regarding the use of medicines and promote safe medication practices.

Conclusion

This study revealed a high prevalence of medication use during pregnancy, supporting the existing scientific literature suggesting a pattern of medicalization among pregnant women. However, this prevalence was lower among white women, which may be related to increased self-medication due to limited health literacy and reduced access to healthcare among black and brown women. Nonetheless, black and brown women used medications indicated during pregnancy as antianemic preparations less frequently, indicating lower access to health services and adequate prenatal care. Therefore, these findings suggest that race is a factor of inequity in prenatal care, demanding public policies to mitigate inequalities.

Ethical Approval

Maternal Risk Factors for Low Birth Weight, Prematurity, and Intrauterine Growth Retardation in Recôncavo da Bahia research was approved by the Adventist Physiotherapy Faculty of Ethics and Research Committee of the Adventist Physiotherapy Faculty Bahia (FAFIS) (Protocol No. 4369.0.000.070-10).

Consent to Participate

Informed consent was obtained from all the participants.

Pregnant women were informed about the confidentiality of the data and were informed about the purpose of the research and its methodology. The interviews were performed only with pregnant women who agreed to participate in the study and signed the informed consent form, which contained explicit information about the study objectives and nature.

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Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

Disclosure

The authors declare no conflicts of interest in this work.

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