Prevalence of *Streptococcus* B in pregnant women with preterm birth and its association with adverse outcomes

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Abstract

Objectives: the objective this study was evaluating the prevalence of maternal colonization by Group B beta-hemolytic Streptococcus (GBS) in pregnant women who delivered preterm and its relationship with adverse maternal/perinatal outcomes.

Methods: we carried out a retrospective cohort study with singleton pregnancies with or without a culture for GBS (vaginal-rectum) who delivered spontaneously <37 weeks of gestation.

Results: the study included 190 pregnant women, 53.1% (101/190) did not undergo culture for GBS and 46.8% (89/101) have done. Among the patients who had a culture, 13.5% (12/89) had positive culture for GBS and 86.5% (77/89) had a negative culture. Pregnant women without GBS culture had higher prevalence of preterm birth (74.3 vs. 59.6%, p=0.031) and lower prevalence of antibiotic prophylaxis (27.7 vs. 56.2%, p<0.001) than pregnant women with GBS culture. Higher prevalence of crystalline penicillin G use was observed in pregnant women with positive culture compared to pregnant women with negative culture for GBS (100 vs. 39%, p<0.0001). There was no significant association between pregnant women with or without a culture for GBS or positive and negative GBS cultures and adverse maternal/perinatal outcomes.

Conclusion: No significant association was found between GBS culture or not, GBS positive or negative culture, adequate or inadequate GBS prophylaxis, and the prevalence of adverse maternal/perinatal outcomes.

Key words Group B beta-hemolytic Streptococcus, Screening, Preterm delivery, Antibiotic prophylaxis, Adverse maternal/perinatal outcomes



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Introduction

Group B beta-hemolytic *Streptococcus* (GBS or *Streptococcus agalactie*) is a bacterium that commonly colonizes the lower genital tract of pregnant women and can cause infection and sepsis in neonates and infants <90 days of age.¹ Perinatal infection occurs by vertical transmission when GBS passes from the vagina into the amniotic fluid after the onset of labor or premature rupture of ovular membranes (PROM).² Approximately 50 to 75% of neonates exposed to vaginal GBS become colonized, and 1 to 2% develop early-onset invasive disease.³ Early-onset neonatal sepsis is the most common manifestation and is characterized by infection manifesting by the 6th day of life with symptoms including respiratory distress, pneumonia, and meningitis.⁴

Cultures for screening GBS infection in pregnant women should be obtained by swab of the anal and vaginal areas between 35 and 37 weeks of gestation. The negative predictive value of GBS cultures obtained less than five weeks after delivery is 95 to 98%. Screening and administration of intravenous antibiotic prophylaxis have been associated with a greater than 50% reduction in early-onset neonatal infection. Prophylaxis for GBS with intravenous antibiotics should be given to all pregnant women with a positive vaginal and/or rectal culture for GBS or, if the culture was not obtained in antepartum, with any of the following risk factors: intrapartum fever (>38°C), preterm birth, PROM for more than 18 hours, previous delivery with neonatal GBS infection, or GBS bacteriuria in the current pregnancy.⁵

Intrapartum antibiotic prophylaxis is indicated on admission to the hospital for labor or PROM and is continued every four hours until delivery or negative culture is obtained. The antibiotic of choice is crystalline penicillin at a dose of 5,000.000 IU as a loading dose, followed by 2,500.000 IU maintenance intravenously every four hours until delivery. Patients admitted for preterm birth should have a culture for GBS obtained on admission if this was not done during prenatal care.

Pregnant women who undergo successful inhibition of labor should have prophylaxis discontinued and the culture result checked. If the culture is negative, there is no need to restart prophylaxis if labor occurs within the next five weeks. If the time between the last culture and actual labor is more than five weeks, a new culture should be obtained, and prophylaxis discontinued.⁶

The prevalence of maternal GBS colonization is influenced by several factors, including the gestational

age at which the sample was collected, the culture methods used, race, age (the older the woman, the greater the risk of colonization), parity (the lower the parity, the greater the risk of colonization), and socioeconomic status. These variables explain the variable colonization rate of five to 41% described in the literature.^{7,8}

The objective of this study was to evaluate the prevalence of maternal colonization by GBS in pregnant women who delivered preterm and its association with adverse maternal/perinatal outcomes.

Methods

A retrospective cohort study was carried out at *Mário Palmério Hospital Universitário – Universidade de Uberaba* (UNIUBE), Uberaba, Minas Gerais State, Brazil, by analyzing medical records from March 2016 to March 2023. The inclusion criteria were the following: 1) singleton pregnancies followed at the antenatal clinic admitted to the obstetrics unit, with or without a culture for GBS and whose delivery occurred spontaneously <37 weeks of gestation; 2) absence of chromosomal abnormalities or congenital anomalies diagnosed by prenatal ultrasound or in the postnatal period.

The GBS culture was carried out after vaginal and perianal swabs were routinely taken from all the pregnant women at the antenatal clinic between 35 and 37 weeks of gestation. To collect the swab, pregnant women are asked not to have bathed, douched or used any vaginal topical medication on the day of the test and have not had sexual intercourse in the last 24 hours. To collect the swab, the patients are placed in the lithotomy position using two sterile swabs (Biocon®- Belo Horizonte, Brazil), after consenting. To collect the vaginal sample, the swab is introduced into the introitus, about 2.0 cm, making rotating movements so that it reaches the entire circumference of the vaginal wall. Subsequently, an anal sample was collected by introducing the swab at a distance of 0.5 cm from the anal sphincter, making rotating movements to reach the entire circumference of the region. Immediately after collection, each swab was individually inserted into a tube containing a transport medium called stuart, stored at room temperature until it was sent to the laboratory within a maximum of three days. In the laboratory, the material was inoculated into a specific enrichment medium (Todd Hewitt) that provides essential nutrients for the development of the target microorganism while partially inhibiting other microorganisms in the flora. After 24 hours in this enrichment medium at a temperature of 35 to 37°C, a

manual reading was made to identify and enumerate the growth of GBS.

In the service searched, prophylactic antibiotics are indicated for all pregnant women with a positive culture for GBS who are admitted for induction or management of labor, except those undergoing cesarean section with intact membranes (prophylactic antibiotics are given before skin incision). According to institutional protocol, prophylactic antibiotics are given to all pregnant women with a negative culture ≥five weeks, in labor with PROM for >18 hours. The antibiotic of choice for prophylaxis is crystalline penicillin G at a dose of 5,000.000 IU IV as a starting dose, followed by 2,500.000 IU IV every four hours until delivery. If crystalline penicillin G is unavailable, ampicillin 2.0 grams (IV) is given as a starting dose, followed by 1.0 gram IV every six hours until delivery. In cases of penicillin allergy, clindamycin 900 mg IV is given every eight hours until delivery. In patients undergoing cesarean delivery, cefazolin 2.0 grams IV may also be given as a starting dose, followed by 1.0 gram IV every six hours until delivery. Administration of two doses of any antibiotic within four hours of delivery is considered adequate prophylaxis.9

High-risk pregnancy was considered the following conditions: gestational arterial hypertension, gestational diabetes mellitus, collagen diseases, heart disease, decompensated lung disease, epilepsy, hereditary thrombophilia, antiphospholipid antibody syndrome, chronic kidney disease, thyroid disease, moderate/severe anemia, infection during pregnancy (toxoplasmosis, syphilis, cytomegalovirus, parvovirus, HIV), fetal growth restriction, oligohydramnios, fetal malformations, and chromosomal abnormalities.

To characterize the study population, the following variables were assessed: maternal age, ethnicity, number of pregnancies, number of previous deliveries, weight, height, body mass index (BMI), smoking, pre-existing diseases, gestational age at time of GBS culture, GBS culture result, gestational age at time of hospital admission, preterm birth, PROM, time of PROM, antibiotic prophylaxis for GBS, number of doses of antibiotic prophylaxis for GBS, adequate treatment, type of delivery, gestational age at delivery, birth weight, sex of newborn, APGAR score at the 1st and 5th minute.

The following variables were considered adverse maternal/perinatal outcomes: early-onset neonatal sepsis, maternal admission to the intensive care unit (ICU), chorioamnionitis, admission to the neonatal ICU, APGAR score at the 7th minute <7, early neonatal death (up to 48 hours of life). Early-onset neonatal sepsis was considered to be the presence of laboratory changes such as leukocytes <5,000 or >25,000 cells/mm³ at birth or

>30,000 cells/mm³ at 12-24 hours of life or $\ge 21,000$ cells/mm³ within two days of life, increased immature neutrophils, platelets <150,000/mm³, ratio of immature neutrophils to total neutrophils >0.3, and the presence of signs such as lethargy, irritability, thin pulse, cyanotic extremities, and tachypnea.¹⁰

The GPower 3.1 program was used for sample calculation. To assess the association between a positive GBS culture and the presence of early-onset neonatal sepsis and adverse perinatal outcomes, considering an effect size of 0.2, a test power of 80% and a probability of error α =0.05, a total number of 197 patients will be needed.

The data was transferred to an Excel 2010 spreadsheet (Microsoft Corp., Redmond, WA, USA) and then analyzed using SPSS 20.0 (SPSS Inc., Chicago, IL, USA) and Prisma GraphPad 7.0 (GraphPad Software, San Diego, CA, USA). The quantitative variables were initially subjected to the normality test (D'Agostino Pearson). Variables with a normal distribution were presented as means and standard deviations (SD). Variables with a non-normal distribution were presented as median, minimum and maximum values. Categorical variables were described using absolute and percentage frequencies and represented in tables. The Chi-square test was used to study the difference between categorical variables and their proportions. The Mann-Whitney and Student's t-tests were used to study the effect of the group studied on the continuous variables. Binary logistic regression was used to calculate the odds ratio (OR) for the adverse maternal/perinatal outcomes. The significance level for all tests was α =0.05.

The study was approved by the Research Ethics Committee at UNIUBE (CAAE: 52299421.7.0000.5145).

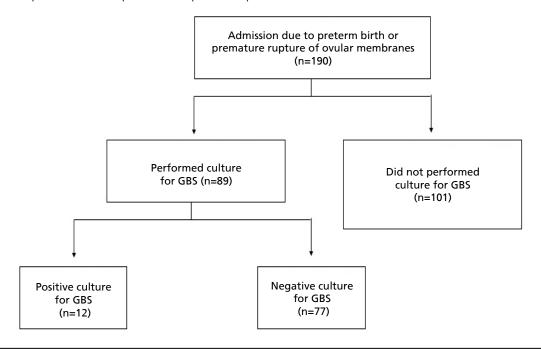
Results

The study included 190 pregnant women admitted to the service due to spontaneous preterm birth and/or PROM who delivered at up to 36 weeks and six days of gestation. Among the patients included, 53.1% (101/190) did not undergo culture for GBS and 46.8% (89/101) have done. Among the patients who had a culture, 13.5% (12/89) had positive culture for GBS and 86.5% (77/89) had negative culture (Figure 1).

Pregnant women who did not have a GBS culture (Group I) had a higher prevalence of preterm birth (74.3 vs. 59.6%, p=0.031) and a lower prevalence of antibiotic prophylaxis (27.7 vs. 56.2%, p<0.001) than pregnant women who had a GBS culture (Group II). There was no significant effect or association between the groups and the other variables analyzed (Table 1).

Figure 1

Flowchart of patients admitted for preterm birth or premature rupture of ovular membranes.



GBS= Group B beta-hemolytic Streptococcus.

Table 1

Clinical characteristics of the population included, who had (Group I) and did not have (Group II) culture for Group B beta-hemolytic *Streptococcus* (GBS). Uberaba-MG, 2016-2023.

Weight (Kg) 76.3 (18.6) 74.1 (8.6) 0.17 0.80 Height (meters) 1.65 (21.0-31.0) 1.70 (22.5-32.3) - 0.17 1.0 38 dy mass index (Kg/m²) 26.0 (7.1) 26.4 (3.6) - 0.07 0.92 Ethnicity 0.56 White 43.6 % (44/101) 38.2% (34/89) - 0.07 0.92 Black 8.9% (9/101) 6.7% (6/89) - 0.06 - 0.07 0.00 - 0.01 0.0% (0/99) - 0.01 0.0% (0/101) 0.0% (0/89) - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.53 - 0.01 0.03 - 0.01 0.03 - 0.01 0.03 - 0.01 0.03 - 0.01 0.03 - 0.01 0.03 - 0.01 0.03 0.03 0.03 0.03 - 0.01 0.03 0.03 0.03 0.03 0.03	Characteristics	Group I (101)	Group II (89)	Cohen's d	p
Height (meters) 1.65 (21.0-31.0) 1.70 (22.5-32.3) - 0.17 1.0 30dy mass index (Kg/m³) 26.0 (7.1) 26.4 (3.6) - 0.07 0.92 Ethnicity 0.56 White 43.6 % (44/101) 38.2% (34/89) Black 8.9% (9/101) 6.7% (6/89) Mixed 47.5% (48/101) 55.1% (49/89) Asian 0.0% (0/101) 0.0% (0/89) Number of pregnancies 2.0 (1.0-7.0) 2.0 (1.1-7.0) - 0.11 0.53 Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4% (56/101) 57.3% (51/89) 5estational age at time of GBS culture (weeks) High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) Preterm birth 74.3% (75/101) 59.6% (53/89) PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) 0.415	Maternal age (years)	24.0 (13-43)	26.0 (15-40)	- 0.07	0.537 †
Body mass index (Kg/m²) 26.0 (7.1) 26.4 (3.6) - 0.07 0.92 Ethnicity 0.56 White 43.6 % (44/101) 38.2% (34/89) Black 8.9% (9/101) 6.7% (6/89) Mixed 47.5% (48/101) 55.1% (49/89) Asian 0.0% (0/101) 0.0% (0/89) Number of pregnancies 2.0 (1.0-7.0) 2.0 (1.1-7.0) - 0.11 0.53 Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) - 0.79 Sestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) 27.7% (28/101) 56.2% (50/89) <0.00	Weight (Kg)	76.3 (18.6)	74.1 (8.6)	0.17	0.806 ∂
Ethnicity 0.56 White 43.6 % (44/101) 38.2% (34/89) Black 8.9% (9/101) 6.7% (6/89) Mixed 47.5% (48/101) 55.1% (49/89) Asian 0.0% (0/101) 0.0% (0/89) Number of pregnancies 2.0 (1.0-7.0) 2.0 (1.1-7.0) -0.11 0.53 Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4 % (56/101) 57.3% (51/89) Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.0415 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415	Height (meters)	1.65 (21.0-31.0)	1.70 (22.5-32.3)	- 0.17	1.0 [†]
White 43.6 % (44/101) 38.2% (34/89) Black 8.9% (9/101) 6.7% (6/89) Mixed 47.5% (48/101) 55.1% (49/89) Asian 0.0% (0/101) 0.0% (0/89) Number of pregnancies 2.0 (1.0-7.0) 2.0 (1.1-7.0) -0.11 0.53 Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4 % (56/101) 57.3% (51/89) Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) 0.415 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415	Body mass index (Kg/m²)	26.0 (7.1)	26.4 (3.6)	- 0.07	0.928 ∂
Black 8.9% (9/101) 6.7% (6/89) Mixed 47.5% (48/101) 55.1% (49/89) Asian 0.0% (0/101) 0.0% (0/89) Number of pregnancies 2.0 (1.0-7.0) 2.0 (1.1-7.0) -0.11 0.53 Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4% (56/101) 57.3% (51/89) Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6% (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Fine of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415	Ethnicity				0.566 §
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Asian 0.0% (0/101) 0.0% (0/89) Number of pregnancies 2.0 (1.0-7.0) 2.0 (1.1-7.0) -0.11 0.53 Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4 % (56/101) 57.3% (51/89) Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Fine of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415	Black	8.9% (9/101)	6.7% (6/89)		
Number of pregnancies 2.0 (1.0-7.0) 2.0 (1.1-7.0) -0.11 0.53 Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4% (56/101) 57.3% (51/89) Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6% (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.0415 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415	Mixed	47.5% (48/101)	55.1% (49/89)		
Number of deliveries 0.79 0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4 % (56/101) 57.3% (51/89) Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00	Asian	0.0% (0/101)	0.0% (0/89)		
0 44.6% (45/101) 42.7% (38/89) ≥ 1 55.4 % (56/101) 57.3% (51/89) Sestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.0415 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415	Number of pregnancies	2.0 (1.0-7.0)	2.0 (1.1-7.0)	- 0.11	0.532 [†]
≥ 1 55.4 % (56/101) 57.3% (51/89) Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Fine of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415	Number of deliveries				0.797 §
Gestational age at time of GBS culture (weeks) 34.7 (21.0-36.9) * High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00	0	44.6% (45/101)	42.7% (38/89)		
High-risk pregnancy 38.6 % (39/101) 41.6% (37/89) 0.67 Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Fime of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415 0.41	≥ 1	55.4 % (56/101)	57.3% (51/89)		
Preterm birth 74.3% (75/101) 59.6% (53/89) 0.03 PROM 73.3% (33/45) 56.8% (25/44) 0.10 Fine of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00	Gestational age at time of GBS culture (weeks)		34.7 (21.0-36.9)		*
PROM 73.3% (33/45) 56.8% (25/44) 0.10 Time of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415 0.41	High-risk pregnancy	38.6 % (39/101)	41.6% (37/89)		0.678 §
Fime of PROM (hours) Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415 0.41	Preterm birth	74.3% (75/101)	59.6% (53/89)		0.031 §
Antibiotic prophylaxis for GBS 27.7% (28/101) 56.2% (50/89) <0.00 Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415 0.41	PROM	73.3% (33/45)	56.8% (25/44)		0.102 §
Gestational age at delivery (weeks) 35.7 (22.4-36.9) 35.9 (25.4-36.9) 0.415 0.41	Time of PROM (hours)				
	Antibiotic prophylaxis for GBS	27.7% (28/101)	56.2% (50/89)		<0.001 §
Type of delivery 0.40	Gestational age at delivery (weeks)	35.7 (22.4-36.9)	35.9 (25.4-36.9)	0.415	0.415 [†]
	Type of delivery				0.409 §

APGAR score at the 5 th min	9.0 (1.0-10.0)	9.0 (1.0-10.0)	0.05	0.838 [†]
APGAR score at the 1st min	8.0 (1.0-9.0)	8.0 (1.0-9.0)	- 0.03	0.703 [†]
Birth weight (grams)	2,378.0 (530.0-3,500.0)	2,520.0 (570.0-3,525.0)	- 0.10	0.438 [†]
Forceps	0.05 (0/45)	2.3% (1/44)		
Cesarean section	51.1% (23/45)	40.9% (18/44)		
Vaginal	48.9% (22/45)	56.8% (25/44)		

PROM= premature rupture of ovular membranes; Student's t-test $^{\circ}$ = mean (standard deviation); Mann Whitney $^{\circ}$ = median (minimum-maximum); Chi-square $^{\circ}$ = percentage (n/N); 'statistical test could not be used due to the absence of at least 3 cases in one of the groups; p<0.05.

There was no significant association between pregnant women with or without a culture for GBS and the presence of early-onset neonatal sepsis (p=0.190), APGAR score at the 1st minute <7 (p=0.940), neonatal ICU admission (p=0.856), neonatal death in the first 48 hours (p=0.924), chorioamnionitis (p=0.184) and maternal ICU admission (p=0.636) (Table 2).

There was no significant association between the presence of positive or negative cultures for GBS and antibiotic prophylaxis (p=0.272). However, a higher prevalence of crystalline penicillin G use was observed in pregnant women with a positive culture for GBS compared to pregnant women with a negative culture ($100 \ vs \ 39\%, p$ <0.0001). There was no significant association or effect between a positive or negative culture for GBS and the other variables (Table 3).

There was no significant association between the presence of positive and negative GBS cultures and the presence of early-onset neonatal sepsis (p=0.732), APGAR score at the 1st minute <7 (p=0.141), neonatal ICU admission (p=0.235), neonatal death in the first 48 hours (p=0.364), chorioamnionitis (p=0.364) and maternal ICU admission (p=0.691) (Table 4).

Considering all the cases included in the study, 33.7% (64/190) underwent adequate prophylaxis for GBS. No significant association was observed between adequate prophylaxis, inadequate prophylaxis and the prevalence of early-onset neonatal sepsis [12.5 (8/64) vs 6.3% (8/126), p=0.171, respectively].

Table 2

Association between pregnant women who did not have (Group I) and have (Group II) culture for Group B beta-hemolytic *Streptococcus* (GBS) and adverse maternal/perinatal outcomes. Uberaba-MG, 2016-2023.

	Group I (101)	Group II (89)	OR (95%CI)	p §
Early-onset neonatal sepsis	5.9% (6/101)	11.2% (10/89)	2.00 (0.69-5.75)	0.190
APGAR score at the 1st min <7	13.9% (14/101)	13.5% (12/89)	0.96 (0.42-2.21)	0.940
Neonatal ICU admission	32.7% (33/101)	31.5% (28/89)	0.94 (0.51-1.74)	0.858
Neonatal death in the first 48 hours	5.9% (6/101)	5.6% (5/89)	0.94 (0.27-3.20)	0.924
Chorioamnionitis	2.0% (2/101)	5.6% (5/89)	2.94 (0.55-15.58)	0.184
Maternal ICU admission	2.0% (2/101)	1.1% (1/89)	0.56 (0.05-6.31)	0.636

CI= confidence interval, ICU= intensive care unit, OR= odds ratio; Chi-Square⁵⁼ percentage (n/N); OR calculated by binary logistic regression.

Table 3

Clinical characteristics of the population who present culture positive and negative for Group B beta-hemolytic *Streptococcus* (GBS). Uberaba-MG, 2016-2023.

Characteristics	Negative culture for GBS (N=77)	Positive culture for GBS (N=12)	Cohen's d	p
Maternal age (years)	27.0 (15-40)	24.0 (16-39)	0.270	0.342 [†]
Ethnicity				0.471 §
White	36.4% (28/77)	50.0% (6/12)		
Black	7.8% (6/77)	0.0% (0/12)		
Mixed	55.8% (43/77)	50.0% (6/12)		
Asian	0.0% (0/77)	0.0% (0/12)		

Number of pregnancies	2.0 (1-7)	2.5 (1-7)	- 0.389	0.386 [†]
Number of deliveries				0.481 §
0	44.2% (34/77)	33.3% (4/12)		
≥ 1	55.8% (43/77)	66.7% (8/12)		
Gestational age at time of GBS culture (weeks)	34.7 (21.0-36.9)	34.6 (30.7-36.6)	- 0.230	0.460 [†]
High-risk pregnancy	41.6% (32/77)	41.7% (5/12)		0.994 §
Antibiotic prophylaxis for GBS	53.2% (41/77)	75.0% (9/12)		0.158 §
Number of doses of antibiotic prophylaxis for GBS	1.0 (0-42)	2.5 (0-24)	0.0009	0.272 [†]
Antibiotic				
Crystalline penicillin G	39.0% (30/77)	100% (12/12)		<0.0001 §
Ampicillin	5.2% (4/77)	0.0% (0/12)		>0.999 [§]
Clindamycin	7.8% (6/77)	0.0% (0/12)		>0.999 §
Ceftriaxone	1.3% (1/77)	0.0% (0/12)		>0.999 §
None	46.7% (36/77)	0.0% (0/12)		0.0013 §
Adequate antibiotic treatment	45.5% (35/77)	75.0% (9/12)		0.057 §
Preterm birth	59.7% (46/77)	58.3% (7/12)		0.926 §
PROM	64.9% (50/77)	66.7% (8/12)		0.907 §
Time of PROM (hours)	7.0 (0.0-2016)	14.0 (0.0-1344)	- 0.137	0.523 [†]
Gestational age at delivery (weeks)	35.9 (25.4-36.9)	35.9 (30.7-36.9)	- 0.124	0.832 [†]
Type of delivery				0.863 §
Vaginal	51.9% (40/77)	58.3% (7/12)		
Cesarean section	46.8% (36/77)	41.7% (5/12)		
Forceps	1.3% (1/77)	0.0% (0/12)		
Birth weight (grams)	2,520 (684-1,500)	2,635 (1,500-3,525)	- 0.298	0.358 [†]
APGAR score at the 1st min	8.0 (1-9)	8.0 (7-9)	- 0.399	0.350 [†]
APGAR score at the 5 th min	9.0 (1-10)	9.0 (8-10)	- 0.241	0.570 [†]

PROM= premature rupture of ovular membranes; Mann Whitney^{†-} median (minimum-maximum); Chi-square⁵⁻ percentage (n/N); p<0.05.

Table 4

Association between positive and negative culture for Group B beta-hemolytic Streptococcus (GBS) and maternal/adverse perinatal outcomes. Uberaba-MG, 2016-2023.

	Negative culture for GBS (N=77)	Positive culture for GBS (N=12)	OR (95%CI)	₽§
Early-onset neonatal sepsis	11.7% (9/77)	8.3% (1/12)	0.68 (0.08-5.96)	0.732
APGAR score at the 1st min <7	15.6% (12/77)	0.0% (0/12)	0.0 (0.0-infinite)	0.141
Neonatal ICU admission	33.8% (26/77)	16.7% (2/12)	0.39 (0.08-1.92)	0.235
Neonatal death in the first 48 hours	6.5% (5/77)	0.0% (0/12)	0.0 (0.0-infinite)	0.364
Chorioamnionitis	6.5% (5/77)	0.0% (0/12)	0.0 (0.0-infinite)	0.364
Maternal ICU admission	1.3% (1/77)	0.0% (0/12)	0.0 (0.0-infinite)	0.691

CI= confidence interval; ICU= intensive care unit; Chi-Square 5 = percentage (n/N); OR calculated by binary logistic regression; p<0.05.

Discussion

In the present study, there was no significant association between pregnant women with or without a culture for GBS or positive and negative GBS cultures and adverse maternal/perinatal outcomes.

In a systematic review and meta-analysis including 45 studies, the estimated risk ratio for preterm birth

with maternal GBS colonization was 1.21 in cohort and cross-sectional studies, and the odds ratio (OR) was 1.85 in case-control studies. In a meta-analysis published in 2022 with 9,778 pregnant Indian women, Ashary *et al.* In women with a positive culture for GBS compared with those without GBS.

In this study, the prevalence of pregnant women who perform culture for GBS between 35 and 37 weeks of gestation was 53.1%. In an Italian study of 2022 that analyzed the regional birth certificate register of the Piedmont Department of Health Policy from 2006 to 2018, following specific recommendations for compliance with international guidelines, the mean proportion of women tested for GBS vaginal-rectal swabs during pregnancy increased from 83.5% in 2006 to 90.7% in 2018, with the largest increase in 2010. 13

The rate of positive culture for GBS (vaginal-rectal) in our population was 13.5% (12/89). The positivity of GBS culture varies among different populations and countries. In a study conducted in two cities in Ghana, Africa, the positive culture for GBS varied from 25.5% (51/200) to 28.0% (56/200). If In a study of 379 Korean pregnant women, the rate of GBS colonization (vaginal-rectal) was 19.8%. If The rate of vaginal colonization by GBS in 200 Pakistani women was 8.5%. If

The prevalence of GBS colonization in 9,778 pregnant women from data of 36 studies (1981 - 2019) was 7.8%.12 A prevalence of positive culture for GBS (vaginal and/or rectum) in different Brazilian regions ranged from 4.2 to 28.4% between 2008 and 2018. ¹⁷ Uberaba is located in the Southwest region of Brazil, has a population of approximately 330,000 inhabitants , and a Municipal Human Development Index (HDI) of 0.772 in 2010, above Brazilian HDI which was 0.699.18 Even though, the prevalence of positive culture for GBS in our population was higher than the prevalence of positive culture of GBS in other less developed countries as Pakistan and lower than higher developed countries as South Korea. Thus, we believe that not only the human development index is related to the prevalence of positive culture for GBS, but other factors should be evaluated to better understand the GBS colonization of the population in each region.

The present study observed a higher prevalence of crystalline penicillin G use in pregnant women with a positive culture for GBS. The antibiotic of choice for prophylaxis in our service is crystalline penicillin G. If crystalline penicillin G is unavailable, ampicillin and clindamycin are the second and third options, respectively. According to the guidelines of the American Academy of Pediatrics and the American College of

Obstetricians and Gynecologists, penicillin, ampicillin, or cefazolin are recommended for prophylaxis of GBS, with clindamycin and vancomycin reserved for cases of significant maternal penicillin allergy. ¹⁹ A European consensus conference in Florence, Italy, attended by 16 experts from different countries representing all major scientific societies, established crystalline penicillin G as the first-line agent for antibiotic prophylaxis of GBS colonization. ²⁰ ACOG Committee Opinion recommends that intravenous penicillin should be the preferred agent for intrapartum prophylaxis, with intravenous ampicillin being an acceptable alternative. First-generation cephalosporins are recommended for women who report a penicillin allergy that indicates a low risk of anaphylaxis or is of uncertain severity. ²¹

In this research, the presence of a positive culture for GBS was not associated with early-onset neonatal sepsis and other maternal/adverse perinatal outcomes. In a Malaysian retrospective cohort study between 2015 and 2020, which included 991 neonates from 44 neonatal intensive care units, the incidence of early-onset neonatal sepsis increased from 0.46 to 0.49/1000 live births during this period. GBS was the most common pathogen (388, 39.2%). Multiple logistic regression analysis showed that Indian ethnicity, chorioamnionitis, gestational age ≥37 weeks, female, spontaneous vaginal delivery, instrumental delivery, and surfactant therapy were significantly associated with increased risk of early-onset neonatal sepsis due to GBS.²² In a Chinese study of 617 pregnant women between 35 and 37 weeks' gestation, the GBS-positive group (560) had a higher incidence of intrauterine infection, postpartum hemorrhage, and fetal distress than the GBS-negative group (57), but similar incidence of PROM, preterm delivery, and meconium-stained amniotic fluid.23 Gad et al.,24 in a retrospective cohort study, evaluated the association between peripartum maternal bacteremia and early neonatal sepsis. Among the 536 maternal blood cultures analyzed, 102 (19.0%) were positive. The most prevalent organisms were GBS (39.2%), followed by Escherichia coli (14.7%) and anaerobes (10.8%). Maternal GBS infection was associates with increased risk of early neonatal sepsis [OR 6.33 (3.02 to 13.25)].²⁴ We believe that the lack of statistical differences between the groups is a consequence of the small sample size in the positive culture for GBS.

In the present study, 33.7% of pregnant women with a positive culture for GBS received adequate prophylaxis, but there were no significant statistical differences between adequate and inadequate prophylaxis and the prevalence of early-onset neonatal sepsis. Zhu et al.²⁵ prospectively evaluated 16,384 pregnant women and 16,634 neonates between May 1, 2019 and April

30, 2020. They observed that adequate intrapartum antibiotic prophylaxis was a protective factor in neonates of pregnant women with GBS colonization. In a retrospective cohort study conducted at 2016, Bianco et al.26 evaluated 902 pregnant women to determine whether complete intrapartum antibiotic prophylaxis for GBS was administered according to the revised Centers for Disease Control and Prevention guidelines. Completely appropriate intrapartum antibiotic prophylaxis for GBS was given to 36.3% of pregnant women, inappropriate prophylaxis was given to 10.4%, and the remaining 45.3% of pregnant women received partially appropriate prophylaxis. Multivariate analysis showed that completely appropriate antibiotic prophylaxis for GBS was significantly more likely in pregnant women with a positive culture at antenatal GBS screening. In our study, we observed a similar rate of appropriate antibiotic prophylaxis for GBS as in the study by Bianco et al.26

Limitations of our study include that it was a retrospective analysis and we were unable to reach the sample size of 197 patients. However, according to post hoc calculation, the power of our study to evaluate the association between a positive GBS culture and the presence of early-onset neonatal sepsis was 78.7%.

In summary, we did not find significant association between GBS culture or not, GBS positive or negative culture, and adequate or inadequate GBS prophylaxis and the prevalence of adverse maternal/perinatal outcomes in pregnant women who delivered preterm.

Authors' contribution

Rezende MC: literature search and medical practices. Rodrigues TP: concept. Freitas APM: data collection or processing. Lopes KS: design. Araujo Júnior E: writing. Peixoto AB: analysis or interpretation. All authors approved the final version of the article and declare no conflicts of interest.

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