Predictive Factors for Pregnancy Hypertension in Primiparous Adolescents: Analysis of Prenatal Care, ABPM and Microalbuminuria

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Objective: To quantify PH prevalence in primiparous adolescents; define predictive factors for the occurrence of PH and its impact on newborns.

Methods: We followed 29 primiparous adolescents from the prenatal period through the 12th week of the puerperium, with a mean of sixteen years of age, served at the Outpatient Facility for Adolescents of Maternidade Escola Assis Chateaubriand (MEAC) of Universidade Federal do Ceará (Fortaleza, Brazil). The pregnant adolescents were divided into two groups, that is, those who remained normotensive (Group I) and those who developed PH (Group II). The variables investigated in the assessment of the value of predictability for the development of PH were anthropometric measures, socioeconomic aspects, smoking habit, inheritance for SAH (father/mother), prenatal tests requested in the first prenatal care visit in addition to microalbuminuria and ambulatory blood pressure monitoring (ABPM) in the 28th week of gestation. The pregnant adolescents were followed up at delivery and late puerperium (12th week after the puerperium). The newborns to the mothers included in our study were assessed at birth according to the Apgar score and the Capurro method, for weight, height and perinatal hypoxia.

Results: The prevalence of PH was 51.7%. Inheritance for SAH presented the highest predictive value for PH with an odds ratio of 10.99. Diastolic arterial pressure equal to or above 70 mmHg at the gestational age of 35 weeks was statistically significant as a predictive value for PH. At ABPM we found a predictive value for PH: diastolic pressure load during alertness, diastolic and systolic pressure load during night sleep, pressure variability and maximum diastolic pressure during sleep. Specifically a maximum diastolic arterial pressure (DAP) at ABPM during the period of night sleep ≥64 mmHg presented an odds ratio of 6 for PH with a sensitivity of 80% and a specificity of 60% for the development of PH.

Conclusion: The research for PH predictive factors in primiparous adolescents showed to be easy to apply and useful to stratify high-risk pregnant women as regards the development of PH.

Key words: Gestational hypertensive, microalbuminuria, ambulatory blood pressure monitoring (ABPM), pregnancy in adolescents.

Gestational hypertensive diseases continue to be the major causes of maternal/fetal mortality in developing countries and account for 60% of direct maternal obstetrics-related deaths.

The specific group of primiparous adolescents is the age group that more strongly correlates with worse maternal/fetal prognosis in large population surveys, and presents high rates of premature delivery, newborns who are small for the gestational age (SGA)2-4 and with a maternal death risk approximately sixty times higher as compared to women in the 20 to 24 age range5-8.

Pregnancy hypertension, defined as the presence of transitory arterial hypertension during pregnancy, with no proteinuria, and with normalized arterial pressure after the 12th week of pregnancy, has been more strongly correlated with recurrence in future pregnancies and with a higher risk for the development of cardiovascular diseases9-10.

In the last years, ambulatory blood pressure monitoring (ABPM) has become a very useful noninvasive method in clinical obstetrics for hypertensive or normotensive pregnant women with a risk factor for developing pregnancy-induced hypertensive disease11.

Microalbuminuria is considered an independent marker of lesions in target organs of hypertensive12 and diabetic13-14 patients. There is no consensus about its predictive value for pregnancy-induced hypertensive disease15-18 in pregnant women, and studies are needed to validate its use in pregnancy with a high risk for the development of pregnancy-induced hypertensive disease.

Methods

Twenty-nine primiparous adolescents were recruited in the outpatient facility of the Service for Adolescents of Maternidade Escola Assis Chateaubriand (MEAC), of Universidade Federal do Ceará (Fortaleza, Brazil) to participate in the study. Participation in the study was on a volunteer basis upon consent by the patients’ legal representative since the patients were all under eighteen years of age (median of sixteen years of age). None of them had diseases and only those who came for at least six visits during the prenatal period remained in
the study. They were separated into two different groups: Group I, those who remained normotensive, and Group II, which comprised those who had a PH diagnosis. All the variables investigated were correlated with the PH diagnosis and assessed as regards their predictive power for PH. The variables investigated as regards PH predictability were: weight (kg), height (m), body mass index (kg/m²), socioeconomic aspects (schooling, marital status, type of dwelling, smoking habit); inheritance for SAH; laboratory tests (hematocrit, hemoglobin, urine analysis, fasting glyceremia) requested in the first prenatal care visit; pressure measurements taken in the first and last prenatal visit; microalbuminuria and ABPM in the 28th week of pregnancy. The adolescents were followed up until delivery (vaginal, cesarean section, forceps) and were later visited at home after the 12nd week of puerperium. The newborns were assessed as regards their condition of birth (perinatal anoxia, APGAR at the first and fifth minutes, Capurro), weight (gr), height (cm) and whether they were AGA (adequate for gestational age), LGA (large for gestational age), SGA (small for gestational age) and the data were correlated with the data of the mothers’ diagnosis of PH. The study was approved by the Research Ethics Committee of the Medical College of the Federal University of Ceará, Brazil (Universidade Federal do Ceará).

In the ABPM, we used DYNAMAP monitors (device for indirect noninvasive automatic mean arterial pressure). Pressure level readings were done using the oscillometric method according to the protocol of the British Hypertension Society19 and to the criteria established by the American Association of Medical Instrumentation20 and by the 3rd ABPM Guidelines of the Brazilian Society of Hypertension21. The ABPM monitor was adjusted to the patient’s waist, on the opposite side of the arm where the blood pressure was measured. The periods of alertness and night sleep of the ABPM were defined and programmed so that the pressure reading could be measured every twenty minutes during alertness and every thirty minutes during night sleep. ABPM was accepted as valid for analysis if a minimum of fourteen measurements had been taken during alertness and seven measurements had been taken during night sleep21. The following ABPM variables were assessed with a predictive value for PH:

- Mean arterial pressure (mmHg) – (MAP);
- Mean SAP (systolic arterial pressure) and mean DAP (diastolic arterial pressure) (mmHg) during alertness and sleep;
- Maximum SAP or DAP value recorded during alertness and sleep (mmHg);
- Systolic and Diastolic pressure variability (standard deviation);
- Maximum systolic and diastolic variation (mmHg) – maximum AP variation as compared with the normal pattern during alertness (SAP of 140 mmHg and DAP of 90 mmHg) and sleep (SAP of 120 mmHg and DAP of 80 mmHg during sleep);
- Systolic pressure load (% of SAP measurements above 140 mmHg) and diastolic (% of DAP measurement above 90 mmHg) during alertness;
- Systolic pressure load (%of SAP measurements above 120 mmHg) and diastolic (% of DAP measurements above 80 mmHg) during night sleep;
- Nocturnal descent – percentage of pressure level reduction during sleep as compared with the alertness period (a fall of pressure levels by or above 10% as compared with pressure levels during alertness).

The data were collected in individual records and stored in an Access 7.0 database. They were analyzed in the Excel and SPSS 7.5 for Windows (Microsoft Corporation) programs. Student’s t test and Mann-Whitney nonparametric tests were used to compare the means of continuous variables between Groups I and II. Categoric variables were analyzed using Fisher’s exact test and the likelihood ratio test, in that a value of p < 0.01 was defined as statistically significant. For the multivariate analysis of risk factors for the development of PH we used logistic regression analysis. Wald’s test was used to assess the significance of the odds ratio (OR) adjusted by logistic regression, with a confidence interval of 95% (CI 95%); the Hopkins scale was used for the OR.

Results

The adolescents’ age ranged from fourteen to eighteen years, and 58.6% were under sixteen. The average gestational age at the first and last prenatal care visits was 13.5 weeks and 35 weeks respectively. Delivery occurred at the mean gestational age of 38 weeks.

Fourteen adolescents (48.3%) remained normotensive (Group I) and fifteen (51.72%) presented PH (Group II) with a mean arterial pressure (AP) of 146.6/97.3 mmHg at delivery (Tab. 1) (Chart 1).

DAP in the last prenatal care visit of patients in Group II presented significant statistical difference as compared with Group I (p < 0.001) (chart 1).

Fourteen patients (48.3%) were underweight in the beginning of pregnancy; nine (31.0%) were within ideal weight limits; and six (20.7%) were overweight (Chart 2) (tab.2). There was no statistically significant difference as regards body weight gain during pregnancy when the two Groups were compared (11.4 ± 3.7 kg in Group I and 12.2 ± 3.5 kg in Group II).

Seventeen adolescents (58.6%) were enrolled in elementary/junior high school; nine were enrolled in high school (31.0%); and three (10.3%) did not attend school. Three patients (10.34%) continued to smoke during pregnancy and two (6.89%) developed PH. Twenty-two adolescents (75.8%) lived with their parents or in-laws. Seven got married as a result of pregnancy and moved with their partners into their own place. We found no correlation between the social variables.
Fourteen adolescents had a family history for SAH and eleven (78.6%) developed PH (p = 0.0092) (Tab. 3). A positive family history for SAH represented a risk (odds ratio) of 10.99 for PH (confidence interval: 1.99-60.57; significance level: 95%).

Laboratory tests requested in the first prenatal visit showed no predictive value for the development of PH (chart 3).

Microalbuminuria with values above normal (> 20 mg/dl) was found in nineteen individuals, of which eleven (57.9%) developed PH (Group II) (Tab. 4).

ABPM measurements were of good technical quality, with the mean of valid measurements being 83.15% for Group I and 85.33% for Group II.

During alertness, there were statistically significant differences between the two Groups as regards diastolic pressure load (p < 0.01) (chart 4). Isolated systolic and diastolic pressure “peaks” in the ABPM period were recorded in the two groups studied, although there was greater predominance of this change in Group II, where 59.0% of the patients presented this alteration on ABPM (Tab. 5). Fisher’s exact test showed no association between this event and PH development.

During night sleep systolic and diastolic pressure loads presented statistically significant differences between the two Groups (p < 0.01) (chart 5). In Group II, 60.9% of the patients presented greater pressure instability represented by the presence of isolated “pressure peaks” (Tab. 6). Fisher’s exact test showed no correlation between the occurrence of hypertensive “peaks” and the development of PH (Tab. 7).

According to the Backward Stepwise model for analysis of multiple linear regression coefficients with a significance level of 45% we can find the predictive value for PH, the mean SAP and DAP during night sleep, maximum DAP during alertness and sleep, DAP pressure variability during sleep and DAP descent during sleep (chart 6). Considering the multivariate analysis with a significance level of 10% among these variables we find the predictability value for PH, diastolic pressure variability and maximum diastolic pressure in the period of night sleep (chart 7).

The odds ratio estimated for DAP variability during sleep was 0.5574 (interval between 0.3145 – 0.9879- confidence interval of 90%) and for maximum DAP during sleep was 1.1347 (CI 1.0215 - 1.2605 with a confidence coefficient
We can assume that the increase in DAP variability during night sleep provides a certain protective effect as to the development of PH, whereas increased maximum DAP during sleep would represent a greater risk factor for the development of PH.

By assessing the maximum DAP during sleep in an isolated fashion we verified that at the cut-off point of DAP ≥ 64 mmHg we observe a dichotomy between the two Groups studied, with a greater association with the development of PH with a sixfold odds ratio (variation of coefficient between 1.172 and 30.72, confidence interval of 95%) (Tab. 8). The sensitivity for this pressure value was 80% and the specificity was 60%, with a positive predictive value (PPV) of 66.7, negative predictive value (NPV) de 75.0, false positive (FP) of 33.33 and false negative (FN) of 25 (chart 8).

Twenty two patients had vaginal delivery, one had forceps delivery and four had cesarean sections due to obstetric reasons. Perinatal anoxia occurred in twelve newborns, two of which belonged to Group II (66.7%), but with no statistically significant difference compared to Group I (statistical test not presented).

Maximum pressure variability during sleep, in turn, did not present any specific cut-off point with a predictive value for PH.
significant differences between the two Groups (Tab. 9).

No statistically significant differences were found between the two Groups studied as regards the other variables analyzed in the newborns (Chart 9) (Tab. 10).

### Discussion

The data found in our study confirm that in the specific group of primiparous adolescents there is a high risk for PH, with an occurrence of 51.7%.

The development of systemic arterial hypertension in women who have developed gestational hypertension is described in many studies in the literature. Chesley & Sibai demonstrated that pregnant women with mean arterial pressure (MAP) in the second quarter of pregnancy equal to or above 90 mmHg presented a higher incidence of development of chronic arterial hypertension evidenced during years of clinical follow up. Therefore, clinical follow-up of these adolescents in the subsequent years will provide information for the early diagnosis of arterial hypertension and for the adoption of nonpharmacological preventive measures for HAS, as well as for the adoption of early appropriate therapy where necessary, thus avoiding co-morbidities associated with the natural progression of arterial hypertension, such as chronic renal failure, left ventricle hypertrophy and heart failure.

According to the 3rd Brazilian Consensus of Arterial Hypertension, for individuals under eighteen and considered normal, pressure values range from SAP < 130 mmHg and DAP < 85 mmHg. Considering that the average height of the adolescents in our study was 161 cm (percentile 50), all of them presented normal mean pressure levels in the first and last prenatal care visits alike.

When the two Groups studied were compared we found a statistically significant difference for DAP measured with the manual sphygmomanometer in Group II. Since absolute pressure levels are considered within normal patterns, they do not have clinical significance in the prenatal period.

**Isolated elevation of systolic or diastolic pressure - Night sleep**

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_Fisher’s exact test (p = 0.054)._  
_**Table 6 - Isolated pressure elevations (“pressure peaks”) during the night sleep period – Groups I and II**_

**Changes of AP in the night sleep period**

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_Fisher’s exact test (p = 0.5977)._  
_**Table 7 - Changes of AP in the night sleep period - Groups I and II**_
There was no correlation between anthropometric variables or the social aspects investigated and PH. Up to the study’s follow-up, pregnancy did not show to have a negative impact on the life of these adolescents. Most of the adolescents in the study, i.e. 89.7% remained in school, and only seven (24.2%) moved in with their partners to their own homes, while most of them, 75.8% continued to live with their parents or with the parents of their partners.

Routine prenatal laboratory tests and microalbuminuria in the 28th gestational week did not show to have any predictive value for PH. Maybe the fact that only one sample for laboratory tests was collected in the beginning of pregnancy contributed to the results found.

ABPM in the 28th gestational week demonstrated to be
the only test that had a predictive value for PH. Brown et al.28 found maximum pressure values of 133/81 mmHg during alertness and 117/68 mmHg during sleep for normotensive pregnant women between the 26th and 30th gestational week.28 The adolescents in our study who developed PH (Group II) presented maximum pressure values of 150.2 ± 11.8/86.5 ± 16.4 mmHg during alertness, and of 118.8 ± 9.7/68.7 ± 11.2 mmHg during the night sleep period, which are therefore above the values deemed normal, specifically in this gestational age. The presence of isolated “pressure peaks” during alertness and night sleep of the pregnant women in Group II evidences that, in fact, there is greater pressure variability and an instability of the circadian rhythm of pressure in those women who come to develop PH.

The diastolic pressure load during alertness in Group II presented statistically significant differences (p < 0.01) with regard to pregnant women in Group I. Diastolic pressure load values during alertness are absolutely normal if we consider the values of the population in general. The population we studied, in turn, is made up of pregnant adolescents in the second half of pregnancy and, because they are in a state of intense systemic vasodilation, we should actually find pressure values which are lower than those of the population in general; therefore, this variable showed to have a predictability value for PH.

Changes in pressure behavior during ABPM during night sleep in pregnant women with PE are very controversial. The presence of diminished nocturnal descent and the inversion of the alertness-sleep pattern, i.e., the fact that pressure levels are higher during sleep than during alertness are among the most significant findings included in the literature29-34. Halligan et al.34 correlate nocturnal descent with the degree of PE severity. In our casuistics, we did not find statistically significant differences as regards nocturnal descent between the two Groups studied, but we observed, through the odds ratio estimated, that the higher the value of maximum DAP during sleep, the higher the risk for PH. Of the pregnant women who presented maximum DAP during sleep ≥ 64 mmHg, 66.7% progressed to PH (odds ratio of 6.0 times for PH). Hermida et al.35 observed that the pregnant women who developed PE or PH had ABPM alterations which became

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Chart 8 - DAP-Maximum during sleep ≥ 64 mmHg – sensitivity, specificity, positive predictive value (PPV), negative predictive value (NPV), false positives (FP), false negatives (FN) and predictability of PH.

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Fisher's exact test (p = 0.2635).

Table 9 - Perinatal anoxia in newborns born to mothers of Groups I and II.
manifest since the first quarter of pregnancy with greater pressure variability and an increase in pressure levels in the last quarter of pregnancy.

As regards the newborns, we did not find any factor that could be correlated with PH, although there were more cases of perinatal anoxia (66.7%) among Group II newborns.

The number of cases studied limits the results found, but may show us that hypertensive gestational diseases, specifically PH, in primiparous adolescents, are prevalent in our population. The investigation of predictive factors for PH may modify the natural progression of hypertensive gestational diseases.

Conclusion

The study to investigate predictive factors for PH in the specific Group of primiparous adolescents showed to be easy to apply and yielded satisfactory results. Heredity of SAH and ABPM in the 28th gestational week were shown to have the highest predictive values for PH. Maximum DAP during sleep and pressure variability during night sleep, as well as the diastolic pressure load during alertness in ABPM had significant predictive values for PH. Maximum DAP during night sleep ≥ 64 mmHg presented a sensitivity of 80%, specificity of 60% with PPV of 66.67 and NPV of 75 for the development of PH.

Potential Conflict of Interest

No potential conflict of interest relevant to this article was reported.

References


