



Gestational weight gain and macrosomia in a cohort of mothers and their children

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Abstract

Objective: To identify co-variables potentially associated to infant macrosomia, including excessive gestational weight gain.

Methods: A cohort was investigated consisting of 230 pairs of mothers and children, residents of the City of Rio de Janeiro. Fetal macrosomia, defined as a birth weight $\geq 4,000$ grams was treated as the dependent variable. Statistical analysis of the relation between macrosomia and co-variables involved stratified analysis and multivariate logistic regression, which estimates odds ratios (OR) and 95% confidence intervals (CI 95%).

Results: The incidence of excessive gestational weight gain was 29.1%, varying from 10% for women under 20 years to 63.6% for women who had given birth to children with macrosomia. Macrosomia incidence was 4.8% for the group as a whole, 10.4% for women with excessive gestational weight gain and 2.5% for women with normal gestational weight gain. Women with excessive gestational weight gain and ≥ 20 years exhibited a chance of developing macrosomia that was 5.42 times greater (CI 95%: 1.11 - 26.34). Within the final multivariate logistic regression model, only excessive gestational weight gain (OR = 5.83, CI 95%: 1.51 - 22.48) remained associated to infant macrosomia.

Conclusions: Considering that excessive gestational weight gain was the only predictor related to macrosomia, it is important that preventive programs that take account of this predictor be implemented, avoiding undesirable fetal outcomes. Future studies should include a wider variety of macrosomia predictors and additional birth outcomes.

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Introduction

A number of different studies, both domestic and international, have been undertaken over recent decades with the objective of investigating the determinants, the magnitude and the evolution over time of low birth weight.¹⁻⁶

In general, these studies have revealed that insufficient gestational weight gain, low maternal age, short maternal stature and alcohol and tobacco consumption, among other factors, are determinants of low birth weight.

There is consensus in the literature that adequate weight gain during gestation is associated with a satisfactory fetal outcome.⁷⁻⁹ In general, it is assumed that normal fetal growth is a positive function of gestational weight gain, modified by pre-gestational nutritional status.¹⁰ During the last decade, the Institute of Medicine - IOM,⁷ with the objective of optimizing birth weight, made recommendations for weight gain rates to be differentiated according to maternal nutritional status, measured by pre-gestational BMI. Thus, greatest gains should occur among women with low pre-gestational BMI (< 19.8 kg/m²), with intermediate ones for women with normal pre-gestational BMI (> 19.8 and ≤ 26.0 kg/m²) and lowest gains among women with pre-gestational BMI defined as overweight (> 26.0 and ≤ 29 kg/m²) and obese (BMI > 29.0 kg/m²).⁷

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Recent data on the magnitude of low birth weight in the Brazilian population revealed that the problem continues to have epidemiological relevance and that the magnitude itself is currently stable.⁴ In contrast, studies of the pattern of distribution and of the determinants of macrosomia remain rare for the Brazilian population.

Recently, certain international studies¹¹⁻¹² and other domestic ones¹³⁻¹⁴ have demonstrated a strict relation between excessive gestational weight gain and macrosomia. The definition of macrosomia is still under debate, but, in general, it has been defined as birth weight values $\geq 4,000$ g, $\geq 4,500$ g or as values for birth weight above the 90th percentile of a reference curve for gestational age and sex.¹⁵ The occurrence of macrosomia has been associated with an increased risk of caesarians, trauma during labor and infant morbidity, in particular when it is associated with gestational diabetes.¹¹ The principal determinants of macrosomia include advanced maternal age, multiparity and pre-gestational obesity, in addition to excessive gestational weight.¹⁶

Against this background, the objective of the present study consists of evaluating the association between excessive gestational weight gain, according to the IOM's recommendations, and macrosomia, based on data obtained from a cohort of mothers and their children, followed-up for 9 months postpartum.

Methods

Four hundred and seventy-nine women were enrolled and followed for 9 months postpartum in a cohort study undertaken in the city of Rio de Janeiro, between May 1999 and April 2001. Observations were made in four waves, at approximately 15 days and at 2, 6 and 9 months. It was at these points that anthropometric data and all co-variables were collected. Women were recruited from three different locations, namely, at the area's central maternity hospital, during the prenatal routine, and during routine BCG immunization; these last two locations are within the Marcolino Candau municipal health center.

The eligibility criteria established for the cohort were: aged 15 to 45 years, first interviewed less than 30 days postpartum, free of chronic diseases, a gestational age of more than 35 weeks at birth, single birth and resident within the study area.

The pattern of losses was evaluated in terms of the distribution of final study completion rates [number of women completing the study/number of women enrolled into the cohort] with respect of a number of important co-variables. Differences in final follow-up rates were evaluated using the chi-square test for proportions. Additional details of cohort recruitment, inclusion of participants, exclusion criteria, anthropometric data collection and the pattern of losses can be obtained from previous publications.¹⁷⁻²¹

Macrossomia (birth weight $\geq 4,000$ grams) was defined as the dependent variable. Birth weight was provided by the mother at the first interview and the value given was checked against the baby's records whenever available. The

consistency of the birth weight variable was tested against the weight values obtained at the first observation, with no inconsistent values being identified; i.e., observed values for birth weight were always lower than those obtained during the first wave of observations.

The following co-variables were included in the analysis: sex of child (male, female), age of mother (< 20 , ≥ 20 years), mother's skin color (white, mulatto, black), marital status (married, with partner, single), education (< 8 , ≥ 8 years' schooling), total family income (< 500 , ≥ 500 Reais), previous caesarian delivery (no, yes), previous abortion (no, yes), parity (1, ≥ 2 children), age of mother at birth of first child (< 17 , ≥ 17 years) and gestational age at birth (< 41 , ≥ 41 weeks). Skin color was based on the interviewer's observations and gestational age calculated based on the date of last menstruation (DLM). This information was obtained during the second wave of observations and has been used systematically to estimate gestational age in the absence of ultrasound.²² Pre-gestational Body Mass Index [weight (kg)/stature (m)²] was obtained during the first wave of observations and was calculated from the pre-gestational weight reported by the mother and stature measured with a Harpenden stadiometer (Harpenden Inc.), using standardized techniques.²³

Gestational weight gain was the study's primary co-variable. The variable was reported by the mother during the third observation wave, at 6 months postpartum and was analyzed according to the weight gain categories recommended by the Institute of Medicine.⁷ The IOM recommend weight gain rates that are differentiated according to pre-gestational nutritional status, measured by body mass index. Gains in weight of more than 18 kg, were considered excessive for women with pre-gestational BMI < 19.8 kg/m², gains over 16kg for women with pre-gestational BMI between 19.8 and 26 kg/m², greater than 11.5 kg, for women with pre-gestational BMI > 26 and ≤ 29 kg/m² and gains ≥ 6 kg were considered excessive for women whose pre-gestational BMI had been > 29 kg/m². Because data for gestational weight gain were only obtained during the third wave of follow-up (6 months postpartum), there were losses and only 230 women could be included in the present analysis.

The statistical analysis initially involved testing the differences between the co-variables described above in proportions of excessive weight gain using the chi-square test. Next, the incidence rates of macrosomia were calculated for the overall group of children, according to the selected co-variables, and stratified by excessive gestational weight gain or not. Relative risk ratios (RR) were then calculated in strata, overall, and adjusted for gestational weight, as were their respective 95% confidence intervals (95% CI). The RR values for the strata were compared using the Mantel & Haenzel test. A selection was made of reference categories for the variables taking the lowest incidence of macrosomia into account. Finally, an unconditional logistic regression analysis was performed, controlling for confounding variables with macrosomia as the dependent variable. Variables were included in the final model if they exhibited statistical significance in the bivariate analysis.

The project was submitted to and approved by the Ethics Committee at the Collective Health Studies Nucleus – NESC (*Núcleo de Estudos de Saúde Coletiva*), at the *Universidade Federal do Rio de Janeiro* (UFRJ) and is in compliance with the ethical principles of non-maleficence, beneficence, justice and autonomy, contained in Resolution 196/96 of the National Health Council (*Conselho Nacional de Saúde*).²⁴ All participants signed a consent form, obtained in a free and spontaneous, once necessary explanations had been made.

All analysis was carried using the computer program SPLUS 2000 (MathSoft). Values for p of less than 0.05 were defined as statistically significant.

Results

Comparative analyses revealed that the incidence of macrosomia and means for birth weight did not differ between the children who participated in the study (n = 230) and those considered as losses or those for whom mothers did not have data for gestational weight gain available (n = 175) (results not shown). Table 1 shows data for the final rate of follow-up according to a number of different important study variables. It can be observed that the pattern of losses is random since for all of the variables tested chi-square for proportions was greater than 0.05.

Of the 230 women studied, 29.1% presented excessive gestational weight gain, 34.4% normal gestational weight gain and 36.5% gestational weight gain below that recommended by the IOM. Table 2 contains data on the frequency of excessive gestational weight gain according to selected variables. The greater incidence rates of excessive gestational weight gain were observed among married women (35.8%), those whose age at first birth was < 17 years (36.6%), with gestational age \geq 41 weeks (38.1%) and with birth weights \geq 4,000 grams (63.6%). No statistically significant differences were observed in the incidence of excessive gestational weight gain between the various variable categories chosen, with the exception of birth weight \geq 4,000 grams.

Table 3 contains data for the incidence rates of macrosomia for the group of children as a whole and stratified by excessive and normal gestational weight gain and by selected variables. The general incidence of macrosomia was 4.8%, with rates of 10.4% for the children of women with excessive gestational weight gain and 2.5% for the children of women with normal gestational weight gain. The greatest macrosomia incidence rates were observed among the children of women with excessive gestational weight gain and ages \geq 20 years (50.0%), with total family incomes \geq 500 Reais (15.2%) and among children born with gestational ages \geq 41 weeks (18.8%).

Table 1 - Distribution of frequency for the selected variables for losses, complete follow-up and final rate of follow-up (Rio de Janeiro, 1991-2001)

| Variables | Initial number of observation | Follow-up losses n | Complete follow-up | Final rate of follow-up % | p |
|--|-------------------------------|--------------------|--------------------|---------------------------|--------|
| Age group (years) | | | | | |
| 18-24 | 196 | 95 | 101 | 51.5 | |
| 25-29 | 114 | 47 | 67 | 58.7 | |
| 30-45 | 95 | 33 | 62 | 65.3 | 0.0753 |
| Marital status | | | | | |
| Single | 74 | 25 | 49 | 66.2 | |
| Partner | 230 | 102 | 128 | 55.6 | |
| Married | 101 | 48 | 53 | 52.5 | 0.1680 |
| Skin color | | | | | |
| White | 154 | 59 | 95 | 61.7 | |
| Mulatto | 179 | 79 | 100 | 55.9 | |
| Black | 72 | 37 | 35 | 48.6 | 0.1711 |
| Child's sex | | | | | |
| Male | 199 | 88 | 111 | 55.8 | |
| Female | 206 | 87 | 119 | 57.8 | 0.6863 |
| Pre-gestational BMI (kg/m ²) | | | | | |
| < 19.8 | 85 | 39 | 46 | 54.1 | |
| 19.8-26.0 | 249 | 103 | 146 | 58.6 | |
| 26.1-29.0 | 47 | 23 | 24 | 51.1 | |
| > 29.0 | 24 | 10 | 14 | 58.3 | 0.7424 |
| Macrossomia | | | | | |
| Yes | 20 | 9 | 11 | 55.0 | |
| No | 385 | 166 | 219 | 56.9 | 0.8683 |

BMI = body mass index.

Table 4 contains figures for relative risk (RR) of macrosomia, stratified, overall and adjusted for gestational weight gain. The chance of having a child with macrosomia was 5.42 (95% CI: 1.11 - 26.34) times greater in women with excessive gestational weight gain and aged 20 years or more. Values for RR did not, however, differ between gestational weight gain strata. Values for RR were not statistically significant for any of the other variables.

Table 2 - Incidence of excessive gestational weight gain according to selected variables (Rio de Janeiro, 1999-2001)

| Variables | n | Gestational weight gain * | | p |
|---|-----|---------------------------|------|--------|
| | | n | % | |
| Child's sex | | | | |
| Male | 111 | 29 | 26.1 | 0.4103 |
| Female | 119 | 38 | 31.9 | |
| Mother's age (years) | | | | |
| < 20 | 20 | 2 | 10.0 | 0.0867 |
| ≥ 20 | 210 | 65 | 31.0 | |
| Skin color | | | | |
| White | 95 | 33 | 34.7 | 0.1867 |
| Mulatto | 100 | 23 | 23.0 | |
| Black | 35 | 11 | 31.4 | |
| Marital status | | | | |
| Married | 53 | 19 | 35.8 | 0.3293 |
| Partner | 128 | 37 | 28.9 | |
| Single | 49 | 11 | 22.4 | |
| Educational level (years) | | | | |
| < 8 | 102 | 30 | 29.4 | 0.9503 |
| ≥ 8 | 128 | 37 | 28.9 | |
| Total family income (reais) | | | | |
| < 500 | 113 | 34 | 30.1 | 0.8656 |
| ≥ 500 | 117 | 33 | 28.2 | |
| Previous caesarian delivery | | | | |
| No | 143 | 27 | 31.0 | 0.7292 |
| Yes | 87 | 40 | 28.0 | |
| Previous abortion | | | | |
| No | 153 | 40 | 26.1 | 0.2107 |
| Yes | 77 | 27 | 35.1 | |
| Parity (number of children) | | | | |
| 1 | 110 | 29 | 26.4 | 0.4599 |
| ≥ 2 | 120 | 38 | 31.7 | |
| Age of mother at birth of first child (years) | | | | |
| < 17 | 41 | 15 | 36.6 | 0.3323 |
| ≥ 17 | 189 | 52 | 27.5 | |
| Gestational age (weeks) | | | | |
| < 41 | 188 | 51 | 27.1 | 0.2200 |
| ≥ 41 | 42 | 16 | 38.1 | |
| Birth weight (g) | | | | |
| < 4,000 | 219 | 60 | 27.4 | 0.0159 |
| ≥ 4,000 | 11 | 7 | 63.6 | |
| Total | 230 | 67 | 29.1 | |

* Gestational weight gain classified according to the Institute of Medicine (IOM, 1990).

Results of the final logistic regression model reveal that excessive gestational weight gain was the only statistically significant predictor, being connected with a 5.83 (95% CI: 1.51 - 22.48) times greater chance of developing macrosomia. Maternal age (RR = 4.38, 95% CI: 0.76 - 25.28) was retained in the final model in order to control residual confounding (Table 5).

Discussion

The results of the present study revealed that women with excessive gestational weight gain presented a 5.83 (95% CI: 1.51 - 22.48) times greater chance of giving birth to a child with macrosomia.

Excessive gestational weight gain has been described as being an important risk factor for a series of unfavorable fetal and maternal outcomes.^{7,8,20,21} In the present study it was observed that almost a third of the mothers studied exhibited excessive gestational weight gain. These values coincide with figures reported by Nucci et al.¹⁴ for a sample of more than 3,000 women studied at six Brazilian state capitals. The incidence rates of excessive gestational weight gain varied from 10% for mothers less than 20 years old to 63.6% for those who had had children with macrosomia.

The incidence of macrosomia in the present study was just 4.8%. This value could be considered low when compared with Native American populations, where figures reach high values from 16 to 31%,²⁵⁻²⁷ or even when compared with the American population in general, which exhibits rates of the order of 10%.²⁸ Comparisons with European countries such as Germany in 1999 (10.1%)²⁹ and Denmark in 1999 (20.0%),³⁰ demonstrate that the values observed here are still low with respect of those populations. In Brazil, studies of the magnitude and even of the determining factors of macrosomia are still rare. The results of one study with 2,275 pairs of mothers and children revealed an incidence of 5.3%,¹³ while Nucci et al.³¹ despite not reporting figures for frequency, observed a 61% greater risk of macrosomia among women with BMI of between 25.0 and 29.9 kg/m.²

Taking the context of nutritional transition into account, it is important to point out that the incidence of macrosomia and its historical tendencies should be considered new and relevant indicators. In this sense it is interesting to observe that the incidence of macrosomia observed in the present study was low. These values place Brazil at an intermediate level in terms of nutritional transition, for which one of the most advanced manifestations is certainly an elevated frequency of macrosomia.²⁹

Among the many factors that determine macrosomia, international studies have highlighted multiparity, pre-gestational overweight or obesity, advanced maternal age, prolonged gestation, excessive gestational weight gain and the occurrence of gestational diabetes as being the most important predictors.^{11,16,29} All of these potential predictors, with the exception of gestational diabetes were investigated in the present study, however, only excessive gestational weight gain was significantly associated with macrosomia. Maternal age presented a significant risk only when

Table 3 - Incidence rates of macrosomia according to gestational weight gain stratified by the variables selected (Rio de Janeiro, 1999-2001)

| Variables | Gestational weight gain * | | | | | |
|---|---------------------------|------|--------|-----|-------|------|
| | Excessive | | Normal | | Total | |
| | n | % | n | % | n | % |
| Child's sex | | | | | | |
| Male | 3 | 10.3 | 4 | 4.9 | 7 | 6.3 |
| Female | 4 | 10.5 | 0 | 0 | 4 | 3.4 |
| Mother's age (years) | | | | | | |
| ≥ 20 | 1 | 50.0 | 1 | 5.6 | 2 | 10.0 |
| < 20 | 6 | 9.2 | 3 | 2.1 | 9 | 4.3 |
| Skin color | | | | | | |
| White | 4 | 12.1 | 2 | 3.2 | 6 | 6.3 |
| Mulatto | 3 | 13.0 | 2 | 2.6 | 5 | 5.0 |
| Black | 0 | 0 | 0 | 0 | 0 | |
| Marital status | | | | | | |
| Married | 2 | 10.5 | 1 | 2.9 | 3 | 5.7 |
| Partner | 5 | 13.5 | 2 | 2.2 | 7 | 5.5 |
| Single | 0 | 0 | 1 | 2.6 | 1 | 2.0 |
| Educational level (years) | | | | | | |
| ≥ 8 | 4 | 10.8 | 3 | 3.3 | 7 | 5.5 |
| < 8 | 3 | 10.0 | 1 | 1.4 | 4 | 3.9 |
| Total family income (reais) | | | | | | |
| ≥ 500 | 5 | 15.2 | 1 | 1.2 | 6 | 5.1 |
| < 500 | 2 | 5.9 | 3 | 3.8 | 5 | 4.4 |
| Previous caesarian delivery | | | | | | |
| Yes | 4 | 14.8 | 2 | 3.3 | 6 | 6.9 |
| No | 3 | 7.5 | 2 | 1.9 | 5 | 3.5 |
| Previous abortion | | | | | | |
| Yes | 3 | 11.1 | 1 | 2.0 | 4 | 5.2 |
| No | 4 | 10.0 | 3 | 2.7 | 7 | 4.6 |
| Parity (number of children) | | | | | | |
| 1 | 3 | 10.3 | 2 | 2.5 | 5 | 4.5 |
| ≥ 2 | 4 | 10.5 | 3 | 2.4 | 6 | 5.0 |
| Age of mother at birth of first child (years) | | | | | | |
| < 17 | 1 | 6.7 | 1 | 3.8 | 2 | 4.9 |
| ≥ 17 | 3 | 11.5 | 6 | 2.2 | 9 | 4.8 |
| Gestational age (weeks) | | | | | | |
| ≥ 41 | 3 | 18.8 | 1 | 3.8 | 4 | 9.5 |
| < 41 | 4 | 7.8 | 3 | 2.2 | 7 | 3.7 |
| Total | 7 | 10.4 | 4 | 2.5 | 11 | 4.8 |

* Gestational weight gain classified according to the Institute of Medicine (IOM, 1990).

unadjusted. For the remaining predictors, no effect was observed from parity or advanced gestational age, although the incidence of macrosomia for women giving birth after ≥ 41 weeks and with excessive gestational weight gain was 18.8%, more than three times greater than the mean global incidence of 4.8%.

Certain limitations of the present study merit discussion. In first place should be considered the significant reduction in observation numbers resulting from follow-up losses, since information on gestational weight gain was only

obtained during the third follow-up wave, at 6 months postpartum. Nevertheless, previous analyses of the same population²⁰ and of others performed for the present study, reveal a pattern of random losses, thus guaranteeing that the 230 women and children analyzed here, are representative of the initial group of 479 pairs of mothers and children. One further limitation is with respect of the fact that data for gestational weight gain were reported by the mothers, which potentially could generate some type of bias. Nevertheless, we believe that the use of this information

Table 4 - Relative risk (RR) of macrosomia, stratified, overall and adjusted for gestational weight gain for the selected variables (Rio de Janeiro, 1999-2001)

| Variables † | Excessive | Gestational weight gain * | | Adjusted |
|---|-------------------|---------------------------|------------------|-------------------|
| | | Normal | Overall | |
| Relative risk (95% CI) | | | | |
| Child's sex | 0.98 (0.24-4.05) | - ‡ | 1.88 (0.57-6.29) | 2.13 (0.62-7.33) |
| Mother's age (years) | 5.42 (1.11-26.34) | 2.69 (0.29-24.47) | 2.33 (0.52-9.71) | 3.64 (0.89-14.85) |
| Educational level (years) | 1.08 (0.26-4.46) | 2.37 (0.25-22.3) | 1.39 (0.42-4.69) | 1.41 (0.43-4.60) |
| Total family income (reais) | 2.58 (0.54-12.36) | 0.31 (0.03-2.95) | 1.16 (0.37-3.79) | 1.19 (0.38-3.73) |
| Previous caesarian delivery | 1.98 (0.48-8.13) | 1.72 (0.25-11.87) | 1.97 (0.61-6.18) | 1.88 (0.60-5.88) |
| Previous abortion | 1.11 (0.27-4.57) | 0.75 (0.08-7.07) | 1.14 (0.34-3.76) | 0.98 (0.30-3.24) |
| Parity (number of children) | 1.02 (0.25-4.20) | 0.99 (0.14-6.85) | 1.10 (0.35-3.50) | 1.01 (0.32-3.16) |
| Age of mother at birth of first child (years) | 1.73 (0.23-13.28) | 0.57 (0.06-5.26) | 0.98 (0.22-4.15) | 1.13 (0.26-4.88) |
| Gestational age (weeks) | 2.39 (0.60-9.58) | 1.76 (0.19-16.2) | 2.56 (0.80-8.48) | 2.18 (0.67-7.06) |

* Gestational weight gain classified according to the Institute of Medicine (IOM, 1990).

† Reference categories: sex (female), mother's age (<20 years), educational level (<8 years), total family income (<500 reais), previous caesarian delivery (no), previous abortion (no), parity (one child), age of mother at birth of first child (<17years), gestational age (<41 weeks).

‡ There were no cases of macrosomia in this sample.

Table 5 - Final logistic regression model for macrosomia (Rio de Janeiro, 1999-2001)

| Variable | OR | 95%CI | p |
|-------------------------------------|------|------------|--------|
| Excessive gestational weight gain * | | | |
| No | 1.00 | | |
| Yes | 5.83 | 1.51-22.48 | 0.0104 |
| Mother's age (years) | | | |
| < 20 | 1.00 | | |
| ≥ 20 | 4.38 | 0.76-25.28 | 0.0982 |

* Gestational weight gain classified according to the Institute of Medicine (IOM, 1990).

does not invalidate the conclusions of the study since the results that involve this variable are highly consistent, in other words, the OR value is high (5.83) in addition to being backed up by a high level of statistical significance (p value = 0.0104). Another justification is based on the fact that previous analyses²⁰ revealed significant differences between the different gestational weight gain categories for the maternal obesity outcome. Furthermore, values that are available for 3082 expectant mothers resident in six Brazilian state capitals, between 1991 and 1995, reveal a mean gestational weight gain of 12.7 kg,¹⁴ which value is very close to the figure observed in the present study (12.9 kg), which points in the opposite direction to the presence of a bias. The low number of macrosomia cases (n = 11) does not go as far as being a limitation, despite potentially

configuring a sample problem. Nevertheless, even with this reduced number of cases, it was possible to demonstrate an effect from excessive gestational weight gain as a factor potentially associated with macrosomia.

With the objective of increasing the consistency and relevance of the results presented here, similar analyses were performed taking extended length at birth as the dependent variable. This was defined as a length for age index as above 2 z-scores. It is interesting to note that, in contrast with what was observed for birth weight, no statistically significant results were observed for extended length at birth, for which reason these results are not presented here.

In synthesis, based on the results of the present investigation, it was observed that excessive gestational weight gain, according to the Institute of Medicine recommendations, is a factor that is potentially associated with macrosomia. While the figures for macrosomia incidence were low, it is fundamental that the magnitude, determinants and historical tendencies of this condition be monitored in a systematic manner, bearing in mind that macrosomia is already a significant public health problem in countries that are at more advanced stages of nutritional transition.

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