Maternal risk factors of molar pregnancy

Original Article

A CASE CONTROL STUDY ON SELECTED MATERNAL RISK FACTORS OF MOLAR PREGNANCY

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ABSTRACT

Background: The incidence of gestational trophoblastic disease varies dramatically in different regions of the world. While there are some geographical and racial variations, perhaps a higher incidence in Africa and Asia. The high incidence of molar pregnancy in some populations have been attributed to nutritional and socioeconomic factors. Some maternal biological factors also associated with molar pregnancy. This study aims to explore the various maternal risk factors associated with molar pregnancy.

Methods: This case control study was carried out among 50 molar and 100 non-molar pregnant mothers admitted at the patient department of obstetrics and gynecology, Rajshahi Medical College Hospital from January to December 2011. The patients of molar pregnancies were selected on the basis of clinically diagnosed confirmed by ultrasonogram and histopathological examination. Controls group were selected as pregnant women of 1st & 2nd trimester. With the ethical approval from the institutional review board (IRB), patients were selected after taking their written consent. A structured questionnaire and a check list were designed with considering all the variables of interest.

Results: Out of 50 cases 48% belongs to the age group below 20 years, 17% belongs to the age group above 34 years. 88% of the control groups are belongs to the age group 20-34 years. Out of 50 cases 4% had history of prior molar pregnancy, none of the control groups had prior history of mole. 46% of molar cases belongs to the blood group A, 82% in case group and 86% of control group were hormonal contraceptive users. Among the socio-demographic risk factors occupation, education, socioeconomic condition, nutritional status of the case and control group were included.

Conclusion: Molar pregnancy largely remains as etiologic dilemma. No pregnancy is immune to develop as hydatidiform mole. As the risk factors for the disease are unclear, an attempt has been made to identify some risk factors of molar pregnancy to compare them with normal pregnancy to find out any modifiable factors.

Keywords: Molar pregnancy, Maternal, Risk factors, Biological, Socio-demographic.

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INTRODUCTION

The incidence of gestational trophoblastic disease (GTD) varies dramatically in different regions of the world. In Europe and North America trophoblastic disease complicates 1 in every 2000-3000 pregnancies. The incidence is as high as 1 in 200-300 live births in South East Asia¹. Molar pregnancy is a pathological condition and has adverse effects on women’s health²,³. In our country, molar pregnancy is not an uncommon event, but is one of the important causes of maternal death. GTD consist of a group of neoplastic disorders arising from placental trophoblastic tissue after normal or abnormal fertilization. The WHO classification of GTD includes hydatidiform mole, invasive mole, choriocarcinoma and placental site trophoblastic tumor. Although maternal age, parity, geographical location, ethnicity, low socioeconomic condition, history of oral contraceptive use, previous history of hydatidiform mole, history of abortion, blood group, radiation and artificial insemination have been suggested as risk factors little is known about its etiology²,³. There are few studies about the relationship between molar pregnancy and husband’s occupation which involves contact with soil and dust⁴. The occurrence of a trophoblastic tumor can be regarded as the result of breakdown in delicate host invader balance. There are also reports of matching leukocyte HLA types between the women and her
partner\(^1\). Women with a history of one hydatidiform mole have 20-fold risk of recurrence\(^6,9\). The risk of persistent trophoblastic disease following hydatidiform mole is about 2-10\%. The risk of recurrence of molar pregnancy is about 1-4\%. There is 1-2\% risk of recurrence of GTD following one molar pregnancy and 25\% following two molar pregnancies\(^6\).

Though exact cause is not known, association of mentioned factors with molar pregnancy is not analyzed in our country. Considering the multidimensional effect of molar pregnancy on women’s health as well as the impending choriocarcinoma, this study was done to find to find out the risk factors of molar pregnancy in the context of our country. So, this study was designed to analyze the risk factors, early detection and prevention of its graves sequlae.

**METHODS**

This case control study was carried out in obstetrics and gynecology department of Rajshahi Medical College Hospital from January to December 2011. Purposive sampling technique was followed in this study. Molar and non-molar pregnant mother of 1\(^{st}\) and 2\(^{nd}\) trimester admitted in the Gynaecology department were the study population. In this study, the number of cases were 50 and the control were 100. The patient of molar pregnancy diagnosed clinically, ultrasonography, HCG reports and by histological examination. Cases of choriocarcinoma excluded. The data were collected in a preformed standard data collection sheet form after taking written informed consent of the patients. Data were analyzed with the help of SPSS software program. Descriptive statistics such as mean, median, and SD were calculated. Chi-square test was done to find out the association of selected maternal risk factors with molar pregnancy. Ethical clearance from the Ethical Review Board of Rajshahi Medical College was taken to carry out the study.

**RESULTS**

For this case control study 50 molar cases and 100 non-molar pregnancies were taken as control group. Risk factors were described by some biological variables and also by some socio-demographical variables.

Table I showed that in case group women with age <20 years and >34 years were in higher and control group women with age 20-34 years were in higher than others. The mean ± SD in case and control were 25.66 ± 8.7 and 24.77 ± 4.04 respectively.

| Age          | Case-50 Frequency (Percent) | Control-100 Frequency (Percent) |
|--------------|-----------------------------|---------------------------------|
|              | N (%)                       | N (%)                           |
| <20 years    | 24(48%)                     | 10(10%)                         |
| 20-34 years  | 09(18%)                     | 88(88%)                         |
| >34 years    | 17(34%)                     | 02(02%)                         |
| Mean±SD      | 25.66 ± 8.7                 | 24.77 ± 4.04                    |

Table II showed in case group there was higher incidence of molar pregnancy in extreme of age, it was 70.6\% in <20 years and 89.5\% in >34 years. In control group 90.7\% were from 20-34 years. The relationship between age and molar pregnancy was statistically significant (p<.001).
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**TABLE-II: Relationship between age and molar pregnancy**

| Age      | Case (Molar) N (%) | Control (Non-molar) N (%) | Total N (%) |
|----------|-------------------|---------------------------|-------------|
| <20 years | 24(70.60%)        | 10(29.40%)                | 34(22.67%)  |
| 20-34 years | 09(9.3%)         | 88(90.70%)                | 97(64.67%)  |
| >34 years | 17(89.50%)        | 02(10.5%)                 | 19(12.66%)  |
| Total N(%) | 50(33.33%)       | 100 (66.7%)               | 150 (100%)  |

Chi-Square value($x^2$) =73.44; df =2 and P=0.000

Table III showed that incidence of molar pregnancy higher in nullipara and para 3 or more. The relationship between parity and molar pregnancy statistically significant (p<.001).

**TABLE-III: Relationship between parity and molar pregnancy**

| Parity of the mother | Case N(%) | Control N(%) | Total N(%) |
|----------------------|-----------|--------------|------------|
| Nullipara            | 24(80%)   | 06(20%)      | 30(20%)    |
| Para(1or2)           | 12(12%)   | 88(88%)      | 100(66.67%)|
| Para 3or more        | 14(70%)   | 06(30%)      | 20(13.33%) |
| Total N(%)           | 50(33%)   | 100(100%)    | 150(100%)  |

$x^2$=61.98; df= 2; P=.000

Table IV showed that 12 molar cases had past history of molar pregnancy. None of the control group had past history of mole (p<.001).

**TABLE-IV: Relationship between past & recent molar pregnancy**

| Past history of mole | Case N(%) | Control N(%) | Total N(%) |
|----------------------|-----------|--------------|------------|
| Yes                  | 12(100%)  | 0(0%)        | 12(8%)     |
| No                   | 38(27.5%) | 100(72.5%)   | 138(92%)   |
| Total N(%)           | 50(33.3%) | 100(66.7%)   | 150(100%)  |

$x^2$=26.08; df= 1; P=.000
Table V showed that women of blood group A and AB had higher association with molar pregnancy 48.9% and 57.6% respectively (p<.001).

**TABLE-V: Relationship between blood group of mother and molar pregnancy**

| Blood Group | Case N(%) | Control N(%) | Total N(%) |
|-------------|-----------|--------------|------------|
| ‘O’         | 2(5%)     | 38(95%)      | 40(26.67%) |
| ‘A’         | 23(48.9%) | 24(51.1%)    | 30(20%)    |
| ‘B’         | 6(20%)    | 24(80%)      | 30(20%)    |
| ‘AB’        | 19(57.6%) | 14(42.4%)    | 33(22%)    |
| Total (%)   | 50(33.3%) | 100(66.7%)   | 150(100%)  |

$\chi^2=30.73; df=3; p=.000$

Table VI revealed association of hormonal contraceptive users with case (molar) and control (non-molar) groups are 32.3% and 39.1% respectively. The relationship between hormonal contraceptive users and molar pregnancy were not significant (p>.001).

**TABLE-VI: Association of molar pregnancy with hormonal contraceptive users**

| Contraceptives | Case N(%) | Control N(%) | Total N(%) |
|----------------|-----------|--------------|------------|
| Users          | 41(32.3%) | 86(67.7%)    | 127(84.67%)|
| Non users      | 09(39.1%) | 14(60.9%)    | 23(15.3%)  |
| Total (%)      | 50(33.3%) | 100(66.7%)   | 150(100%)  |

$\chi^2=0.41; df=1$ and $P=.339$

Table VII showed 72.7% mothers of primary level and 57.9% mothers of illiterate group had relationship with molar pregnancy which were statistically significant (p<.001).

**TABLE-VII: Relationship between education of mothers and molar pregnancy**

| Education          | Case N(%) | Control N(%) | Total N(%) |
|--------------------|-----------|--------------|------------|
| Illiterate         | 11(57.9%) | 8(42.1%)     | 19(12.67%) |
| Primary            | 32(72.7%) | 12(27.3%)    | 44(29.3%)  |
| Secondary 0r above | 7(8%)     | 80(92%)      | 87(58%)    |
| Total (%)          | 50(33.3%) | 100(66.7%)   | 150(100%)  |

$\chi^2=60.92; df=2; P=.000$
Table VIII showed 67.7% mothers of poor social class had a relationship with molar pregnancy which was statistically significant (P<.001).

| Socio-economic condition | Case N (%) | Control N (%) | Total N (%) |
|--------------------------|------------|---------------|-------------|
| Poor                     | 42(67.7%)  | 20(32.3%)     | 62(41.33%)  |
| Middle class             | 7(9.6%)    | 66(90.4%)     | 73(48.67%)  |
| Higher class             | 1(6.7%)    | 14(93.3%)     | 15(10.0%)   |
| Total N(%)               | 50(33.3%)  | 100(66.7%)    | 150(100%)   |

χ²=56.35; df=2 and P=.000

DISCUSSION
The incidence of molar pregnancy is now gradually increasing. The prognosis of the disease varies with specific biological and socio demographic factors as well as treatment regimen and compliance to follow up.

In this study, in case group women with age <20 years and >34 years were significantly higher (P<0.05) compared to control group which correlates with Parazzini F et al11 and the study in Abu Dhabi12.

In this study in case group nullipara or para 2 or more were significantly higher (P<.05) compared to control group which indicates that in these mothers’ 1st pregnancy in nullipara or any pregnancy in para 2 or more ended in GTD which support the statement of a retrospective study of 455 patients of the Royal college Hospital13.

Regarding history of previous molar pregnancy, this study showed that in case group 2.8% mothers had the previous history of mole compared to 0% in controls which supported by Bracken MB, an epidemiologic review14.

In this study 72% cases and 16% controls had the history of abortion. Parazzini F et al reported that the risk factor for molar pregnancy increased with two or more miscarriage15.

Genetic aspects were also considered through the analysis of blood group of cases and controls. In this study, 46% cases had blood group ‘A’ and 38% cases had blood group “AB” which were supported by a study in Italy11.

There was no significant (p>0.05) association of molar pregnancy with hormonal contraceptive users. This study differs with ‘A case control study of Shimuzi T16. Recent use of OCP was found to be unrelated to risk of developing hydatidiform mole14. This study has similarity with the present study.

Illiterate and primary level educated women were significantly (P<0.001) higher in cases compared to control (Table VII). These women knew a little about antenatal care. They were admitted in hospital due to pregnancy complications.

In this study, in case group 86% were poor and in control group 66% were of middle class, mothers of poor socioeconomic condition had a relationship with molar pregnancy (P<0.001). A study in Mexico was done to find out the important etiological factors which showed 22.85% patients of molar pregnancy were of low social class against 8.57% of control group suggesting that low socioeconomic condition was a risk factor17.

CONCLUSION
As the disease are unclear, an attempt had been made to identify some risk factors of molar pregnancy to compare them with normal pregnancy and also to find out any rectification possible or not. It must not be forgotten that the prevention or early diagnosis should be the principal goal. Prevention means the suppression of the pathology leading to the disease (molar pregnancy). In this study age <20 years, nullipara, illiteracy, malnutrition, blood group A or AB, past history of molar pregnancy or abortion were found more common among cases than controls. Most of the factors
are preventable or modifiable. Encouraging later marriage, postponing childbearing up to the age of 20 years will help to reduce the incidence. Nutritional status can be improved by social upgrading and by increasing the level of education. By improving maternal health, we can make a major improvement in the quality of life. Patients with history of spontaneous abortion should be on regular obstetric care which will also help to reduce the occurrence or dreadful complication of molar pregnancy.

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