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Reproductive factors in relation to breast cancer: A hospital based case control study in Jammu, India

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ABSTRACT

Research Question: To Study the risk factors of Breast cancer , Setting:All the newly registered cases of Breast cancer in one year from Ist May 2004. Study Design:A Hospital based case control study. Methodology:Data was collected by a face to face interview using the prestructured Questionnaire after taking informed consent. Results: 100 cases of Breast Cancer and 100 controls were analyzed. Mean age of breast cancer cases was 50.20 ± 12.49 years. Income ($p > 0.39$), Educational Status ($p > 0.35$) Age at Marriage ($p > 0.36$) Age at First pregnancy ($p > 0.32$) Total No of live Birth ($p > 0.09$) Duration of Breast Feeding ($P > 0.07$) Showed no statistically significant relationship to the risk of Breast Cancer. Dietary History ($p < 0.001$) Smoking History ($p < 0.04$) Physical Activity ($p < 0.001$) Age at Menarche ($p < 0.002$) History of Breast Feeding ($p < 0.04$) History of Abortion ($p < 0.003$) were shown to be Statistically Significant to the risk of Breast Cancer . Conclusion : The risk factor which are often implicated in the risk of Breast Cancer may not hold true in our settings.

Key Words:Breast Cancer, risk factors, cumulative duration of Breast Feeding.

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Introduction

In the year 2000, breast cancer counted for 22% of all new cancers in women making it by far most common cancer in females. In high income countries, in the year 2000 the incidence of breast cancer in women was at least twice that of any other cancer in women, cancer of breast was also commonest tumor among women in low income region, with 470,000 new cases per year¹. Breast cancer causes 376000 deaths in a year worldwide². Out of six population based cancer registries in India, two have reported it as leading cancer site among women and the rest four as the second commonest cancer³. In spite of this, statistical data related to risk factors of Breast cancer is limited and available only from few centers. This study was carried out to identify the risk factors of Breast cancer in this part of country.

Material and Methods

All the newly registered cases of Breast cancer in Government Medical College Jammu during one year period from 1st May 2004 were included in the study. For every case of Breast cancer one control was selected from among women admitted in the hospital for non neoplastic or non hormone related diseases and

matched for age (taking interval of two years) and residence (urban/rural). Data was collected by a face to face interview using a prestructured pretested questionnaire which was modified and adapted from American Cancer Society⁴ after taking informed consent. Some of the elements of the questionnaire were not considered like Ethnicity etc. In place of ethnicity, religion was considered. The study variables included general information, menstrual and reproductive factors, obstetric history, family history and clinical history of breast cancer, diagnostic criteria included microscopic confirmation, evidence obtained at operation or clinical diagnosis. Institutional ethical committee clearance was obtained before starting the study.

The information collected was analyzed using computer software SSPS version 12.0 and Epi – Info version 6.0. Univariate analysis was undertaken to evaluate the relationship of demographic and reproductive factors with breast cancer and results were reported as Crude OR along with 95% confidence interval. Statistical significance of the relationship was ascertained by Chi square test (with Yates correction). Multivariate logistic regression was

employed in a stepwise manner to evaluate the independent and joint effect of the factors found significant on univariate analysis. A p value of <0.5 was considered as statistically significant unless specified otherwise. All p values reported are two tailed.

Results

A total of 103 patients of Breast cancer were registered during one year period. 2 Patients could not be contacted and one male patient was excluded from final results. Final analysis was done on 100 cases and 100 controls. Mean age of breast cancer cases was 50.20 +/- 12.49 years. 77% cases had infiltrating duct carcinoma. Past history of breast lump /pain was present in 22% of cases and 8% of cases gave a family

history of breast / ovarian cancer. The proportion of women in different income categories (p>0.39) and in various educational categories (p>0.35) did not differ significantly. Those who took non vegetarian diet (p<0.001) and had sedentary life style (p<0.001) were at statistically highly significant risk of breast cancer. Those attaining menarche after 13 years of age were at lower risk of breast cancer (p<0.002) than those attaining before 13 years of age (Table 1). Age at marriage did not seem to bear any statistically significant relationship to cancer risk (p>0.36). In both cases and controls one fourth of women (25%) became pregnant for the first time after they were 24 years of age. However the proportion in three age categories did not differ significantly (p>0.32).

Table 1 : Univariate analysis of Reproductive factors in relation to breast cancer

| Reproductive factor | Cases (%) | Controls (%) | OR (95%CI) | P value |
|--|-----------|--------------|-----------------|---------|
| Age at menarche in years | | | | |
| <13 | 25(25) | 11(11) | 1.00(Ref) | <0.002 |
| 13-14 | 45(45) | 36(36) | 0.55(0.22-1.36) | |
| 15 & above | 30(30) | 53(53) | 0.25(0.25-0.62) | |
| Age at marriage in years | | | | |
| <18 | 34(34.69) | 33(33.33) | 1.00(Ref) | >0.36 |
| 18-23 | 48(48.97) | 56(56.56) | 0.83(0.45-1.54) | |
| 24+ | 16(16.32) | 10(10.10) | 1.55(0.62-3.91) | |
| Age at first pregnancy in years* | | | | |
| <18 | 15(15.95) | 9(9.18) | 1.00(Ref) | >0.32 |
| 18-23 | 55(58.51) | 65(66.32) | 0.51(0.21-1.25) | |
| 24 & above | 24(25.53) | 24(24.48) | 1.00(0.31-3.22) | |
| *6 cases & 2 controls did not become pregnant | | | | |
| Total no. of live births | | | | |
| 0 | 8(8) | 2(2) | 1.00(Ref) | >0.09 |
| 1-2 | 28(28) | 24(24) | 0.29(0.06-1.5) | |
| 3+ | 64(64) | 74(74) | 0.22(0.04-1.06) | |
| Cumulative duration of breast feeding (months) | | | | |
| Never | 10(10) | 3(3) | 1.00(Ref) | >0.07 |
| Upto 12 | 1(1) | 1(1) | 0.30(0.01-6.38) | |
| 12-23 | 8(8) | 3(3) | 0.80(0.13-5.09) | |
| >=24 | 81(81) | 93(93) | 0.26(0.07-0.98) | |
| Number of abortions | | | | |
| 0 | 61(61) | 80(80) | 1.00(Ref) | <0.003 |
| 1 | 20(20) | 12(12) | 2.19(0.93-5.19) | |
| 2+ | 19(19) | 8(8) | 3.11(1.19-8.37) | |
| History of menopause | | | | |
| Yes | 58(58) | 57(57) | 1.00(Ref) | 0.8 |
| No | 42(42) | 43(43) | | |

Total number of live births showed no relation to the risk of Breast cancer (p>0.09). Regarding breast feeding, those who had breast fed were at lesser risk of breast cancer (p<0.04). Lactation for more than 24

months was observed to lead to reduced risk of breast cancer. However this relationship failed to achieve statistical significance (p>0.07). 80% of women in controls and 61% of women with breast cancer had no history of abortion. Women, who had undergone

abortion once, were at twice the risk but this difference was not statistically significant. Women with multiple abortions were at higher risk (3 fold) compared to those without abortion history ($p < 0.003$). Almost equal number of women in both the groups had attained menopause. Those attaining menopause at the age 45 - 49 were at slightly higher risk of breast cancer ($p = 0.007$)

Discussion

Majority of the cases (30%) were in the age group of 45 - 54 years with mean age of 50 ± 12.49 . The peak age around 50 years has been reported in some studies^{5, 6}. The 16th Annual report of Hospital cancer registry of Tata Memorial Hospital mentions the lowest age as 19 years and maximum as 86 years with an average age of 48 years which is almost in accordance with our study⁷. No significant association was observed for income and literacy with breast cancer. Few authors have given similar finding⁶ and some have given opposite results^{8, 9, 10}. High fat diet and obesity have been often implicated in the development of Breast cancer. A study conducted by Ingram DM Etal¹¹ has shown that consuming Red meat, savoury meals and starches caused high risk. A statistically significant difference in dietary habits was found among cases and controls in our study. The findings of smoking history and Breast cancer by Fergus J Couchetal¹² are also found in our study. Frisch etal¹³ in their study on college athletes concluded that long term athletic training establishes a life style which somehow lowers the risk of breast cancer and cancer of the reproductive system. Yu-Tang etal¹⁴ have studied the physical activity among 1459 cases and 1556 controls, where they have observed that physical activity has the bearing on cancer breast and the results were significant, which is in accordance with this study.

In this study the comparative data of age of Menarche shows that in term of risk, those attaining after 13 years of age were at lower risk of breast cancer than those attaining before 13 years of age. Our findings are in accordance with their studies which shows that age at menarche was inversely associated with risk of breast cancer^{9, 14, 15, 16}

Age at marriage did not seem to bear any statistically significant relationship to risk of cancer in this study. There are contradictory finding in the literature, where some studies show significant relationship with breast cancer and others are not in agreement^{6, 16, 17, 18}

No significant results were found when analyzed for age at first pregnancy among cases and controls in our study. In a study of 6705 and 5492 controls, La

Vecchia etal¹⁵ concluded that there was no significant interaction between age at menarche, or age at diagnosis, parity and age at first birth. As far as total number of live births in relations to risk of breast cancer is concerned, the findings are similar to those of some authors.^{14, 15}

A study conducted by Ramon JM etal¹⁹ is in conformity with our findings of duration of lactation and breast cancer, however Zheng T etal²⁰ and Becher etal²¹ in their study state that, risk of breast cancer decreases with increased duration of life time lactation experience. In a case control study, Rao DN etal⁶ did not find a significant relationship between abortion and cancer breast. These findings were contradictory to our results.

In the present study those attaining menopause at age 45-49 years were at slightly higher risk of breast cancer. Although it was statistically insignificant. Similar finding have been observed by few m other authors^{14, 16}

Multiple logistic regression was performed and the variables found significant on univariate analysis lost significance. The results could possibly be explained on the basis of lesser power of study owing to small sample size, inadequate measurement of certain variables including age at menarche due to recall bias and deliberate misinformation about the number and type of abortions. All these results point to the need of more rigorous study particularly in the light of contradictory results available from various studies regarding risk factors of breast cancer.²²

Conclusion

The menstrual and reproductive factors associated with breast cancer were not found to be significant in our study. To further evaluate the association and the influence of these factors on the temporal trend of breast cancer incidence, there is a need to conduct a longitudinal population based case control study.

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