A Study of Histopathological Spectrum of Ovarian Neoplastic and Non-Neoplastic Lesions at Teaching Hospital, Ahmedabad

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ABSTRACT

Background: Ovarian carcinoma is one of the most common gynecologic cancers that ranks third after cervical and uterine cancer. Ovarian lesions are neoplastic and non-neoplastic and many neoplastic lesions are asymptomatic and possess great challenge to the gynecological oncologist.

Aims & Objectives: To analyze frequency, age distribution and histopathological spectrum of ovarian lesions at teaching hospital, Ahmedabad.

Materials & Method: This study was undertaken between period of 1st January 2018 to 29th February 2020 at Department of Pathology, GMERS Medical college, sola, Ahmedabad. H and E stained slides were examined by light microscopy and histopathological type of lesions were classified according to World Health Organization (WHO) classification -2014..

Results: There were total 182 cases of ovarian lesions. 56(30.76%) cases are neoplastic, among them only 6(10.71%) were malignant. Surface epithelial tumor (n=27 cases, 48.21%) were most common neoplastic lesion while sex cord-stromal tumor (n=8 cases, 14.29%) were least common. Most common age group affected for neoplastic lesions was 31-40 years.

Conclusion: We found that ovarian lesions affect wide variation of age starting from 11 years young patient to 65 years old patients. Non neoplastic lesions were almost double in prevalence than neoplastic lesions. Histopathological analysis according to WHO classification reveal that surface epithelial tumors and germ cell tumors were forms the majority of neoplastic lesions.

Keywords: Ovarian Lesion, Surface Epithelial Tumor, Germ Cell Tumor

Introduction

The ovaries are paired intra-pelvic organ of the female reproductive system. The term ‘Ovary’ was introduced in the 17\textsuperscript{th} century. Even Shushruta has described ovarian neoplasms in his book “Shushruta and Ayurveda”. Ovarian neoplastic and non-neoplastic lesions present a great challenge to gynecological oncologist. Certain non-neoplastic lesions of the ovary often form a pelvic mass and potentially mimic an ovarian neoplasm. Their proper diagnosis and classification are therefore important to allow appropriate therapy. Ovarian tumors are one of the major health problems faced by clinicians. Ovarian carcinoma is one of the most common gynecologic cancers that ranks third after cervical and uterine cancer. Recent improvement in patient survival in developed countries is attributed to an earlier stage at diagnosis.

Ovarian tumors are often difficult to detect until they are advanced in stage or size, as symptoms are vague and insidious. Identification of various histologic patterns of ovarian tumors is important for diagnosis as well as prognosis. Benign conditions and some non neoplastic conditions are common during reproductive years while malignant ones tend to occur during perimenopausal and post menopausal age. Incidence of ovarian cancer is higher in postmenopausal, unmarried and low parity women. Pregnancy (especially if before age 25 years) and use of oral contraceptives are associated with a diminished risk. Because of complexity in structure of ovary owing to its unique embryological development, there are various classification schemes for ovarian neoplasms. However, The World Health Organization Histological Classification of ovarian tumors separates ovarian neoplasms according to the most probable tissue of origin: surface epithelial tumors (65%), germ cell tumors (15%), sex cord-stromal tumors (10%), metastatic tumors (5%) and miscellaneous tumors. Surface epithelial neoplasms are further classified according to cell type (serous, mucinous, endometrioid, clear cell, transitional cell and epithelial- stromal) and degree of atypia (benign, borderline or malignant). Screening for ovarian cancers is aided by serum tumor markers like Alpha fetoprotein, Lactate dehydrogenase, Cancer Antigen 125, Placental Alkaline Phosphatase and radiodagnostic techniques like Ultrasonography, Computed Tomography, Positron Emission Tomography scan and Magnetic Resonance Imaging.

More recently, development of techniques like image guided Fine Needle Aspiration Cytology, fluid cytology...
have helped immensely in early diagnosis of ovarian carcinomas.\[^4\] Intraoperative frozen section evaluation is crucial for determining the required extent of surgery, that is, cystectomy for benign tumors, oophorectomy or limited surgical removal for borderline tumors in younger patients to preserve fertility or extensive surgical removal for ovarian carcinomas. Touch imprint smears also aid in differentiating between benign and malignant conditions.

Present study was carried out with an aim to derive distribution, common age presentation and histopathological spectrum of various non-neoplastic and neoplastic lesions of ovary.

**Materials and Methods**
This study was undertaken using existing patient data retrieved from the registers and records of the histopathological laboratory, Department of Pathology, GMERS Medical College and hospital, Sola, Ahmedabad. A total of 182 ovarian specimens were studied for histopathological evaluation from 1\(^{st}\) January 2018 to 29\(^{th}\) February 2020, either as solitary specimens or as part of total abdominal hysterectomy (TAH) specimens.

Relevant clinical information regarding the age of patients, clinical features, radiological findings and provisional diagnosis were obtained. Gross examination of received specimen was done for various parameters like size, external surface, and consistency and cut sections with contents of cyst. The tissues were processed by routine paraffin techniques of tissue processing and sections stained with Haematoxylin and Eosin were taken for microscopic examination. The non-neoplastic and neoplastic lesions from representative sections were studied and classified according to World Health Organization (WHO) classification 2014. Acquired data were tabulated and analyzed.

**Results**
Out of the total 182 ovarian lesions 126(69.23%) were nonneoplastic and 56(30.76%) were neoplastic lesions. 56 neoplastic lesions are further classified in benign (48 cases,85.72%), borderline (02 cases,3.57%) and malignant lesions (06 cases,10.71%). The distribution of the cases amongst these categories is given in Table 1.

Non neoplastic lesions are common in 21-30 years of age group while the most common age group for all ovarian benign neoplastic lesions was found to be 31-40 years followed by 21-30 years, i.e, reproductive age group while Malignant ovarian neoplasm is mainly found in age more than 40 years. (table 2)

Out of total 126 non neoplastic lesions most common lesion was simple serous cyst 49 (38.88%), followed by chocolate cyst 35(27.77%) and Corpus luteal cyst 16(12.69%) Whereas least common was Developmental cyst and nonspecific oophoritis, One case (0.79%) of each reported. 5 cases(3.96%) show normal histology.

Out of total 56 neoplastic lesions, 27 cases (48.21%) were of surface epithelial tumors, 21 cases (37.5%) were of germ cell tumors, 8 cases (14.29%) were of sex cord stromal tumors [chart 1].

Out of 27 cases of surface epithelial tumors, 23(85.18%) were benign, 2(7.41%) cases were borderline and 2(7.41%) were malignant tumors. Serous cystadenoma (9 cases,39.13%) are most common benign neoplasm of surface epithelial tumors. There are 2 cases of borderline surface epithelial tumors, 1 case was of seromucinous type and 1 case was of mucinous type. There are 2 malignant surface epithelial tumors, 1 case was seromucinous type cystadenocarcinoma and 1 case was endometroid carcinoma [figure 2].

We found total 21 germ cell tumors, 20 cases (95.24%) were benign and 1 case (4.76%) was malignant. Out of total 20 benign cases, 19 cases (95%) were Mature cystic teratoma while one was Mature solid teratoma which was associated with gliomatosis peritonei [figure 4]. Malignant case was of a dysgerminoma [figure 1].

Total 8 cases of sex cord stromal tumors are reported. Out of them, 05(62.5%) were benign and 03 (37.5%) were malignant. In benign lesions fibroma comprised of 50% of cases while granulosa cell tumor adult type [Figure 3] comprised of all the malignant cases.

**Discussion**
Ovarian cancer is the second leading cause of mortality among all gynecological cancers.\[^7\] Due to similar clinical presentations, there is confusion in the diagnosis of non-neoplastic and neoplastic lesions of ovary although it is diagnosed as a mass or cystic lesion on ultrasonography and hence removed prophylactically in routine oophorectomies and hysterectomies.\[^5\]

In present study, total 182 ovarian specimens were received in our pathology department, Out of which 126(69.23%) cases were non neoplastic lesion while 56(30.76%) were neoplastic lesion. This result is comparable with the Indian study done by Sawant and Mahajan et al and study in Nepal done by Tadayon M et al in which they got 70% and 61% non neoplastic lesions and 30% and 39% neoplastic lesions respectively.\[^7,15\]
### Table 1: Distribution of ovarian lesions.

| Distribution ovarian lesions | Non-neoplastic lesions | Neoplastic lesions | Total |
|-----------------------------|------------------------|--------------------|-------|
| Number of cases (Percentage %) | 126 (69.23%) | 48 (85.72%) | 02 (3.57%) | 06 (10.71%) | 182 (100%) |
| Total | 126 (69.23%) | 56 (30.76%) |

### Table 2: Age wise distribution of neoplastic lesions.

| Age group | Surface epithelial tumor | Germ cell tumor | Sex cord stromal tumor | Total |
|-----------|--------------------------|-----------------|------------------------|-------|
|           | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant | Benign | Borderline | Malignant |
| 11-20     | 03   | 00     | 00     | 03   | 00     | 01     | 00     | 00     | 01     | 00     | 00     | 00     | 03     |
| 21-30     | 02   | 01     | 00     | 08   | 00     | 00     | 04     | 00     | 00     | 01     | 00     | 00     | 15     |
| 31-40     | 07   | 00     | 01     | 07   | 00     | 00     | 01     | 00     | 00     | 01     | 00     | 00     | 16     |
| 41-50     | 08   | 00     | 01     | 02   | 00     | 00     | 00     | 00     | 00     | 01     | 00     | 00     | 12     |
| 51-60     | 01   | 00     | 00     | 00   | 00     | 00     | 00     | 00     | 00     | 00     | 00     | 00     | 03     |
| >60       | 02   | 01     | 00     | 00   | 00     | 00     | 00     | 00     | 00     | 00     | 00     | 00     | 03     |
| TOTAL     | 23   | 02     | 02     | 20   | 00     | 01     | 05     | 00     | 03     | 56     |

### Table 3: The distribution of various nonneoplastic lesions.

| Non-neoplastic lesions of ovary | Number of cases | Percentage (%) |
|---------------------------------|-----------------|----------------|
| Simple Serous Cyst              | 49              | 38.88%         |
| Endometriosis/Chocolate cyst    | 35              | 27.77%         |
| Corpus luteal cyst              | 16              | 12.69%         |
| Follicular cyst                  | 7               | 5.55%          |
| Congestion                       | 5               | 3.96%          |
| Normal histology                 | 5               | 3.96%          |
| Torsion                          | 3               | 2.38%          |
| Polycystic disease               | 2               | 1.58%          |
| Ectopic gestation                | 2               | 1.58%          |
| Vestigial/Developmental cyst     | 1               | 0.79%          |
| Nonspecific oophoritis           | 1               | 0.79%          |
| TOTAL                            | 126             | 100%           |

### Table 4: Spectrum of Surface epithelial tumors, Germ cell Tumors and Sex cord stromal tumors.

| Type of ovarian tumors | Surface epithelial tumors | Germ cell tumors | Sex cord stromal tumors |
|------------------------|----------------------------|------------------|-------------------------|
| Benign                 | Histopathological type     | Total benign cases (n=5, 85.18%) | Percentage | Histopathological type | Total benign cases (n=20, 95.24%) | Percentage | Histopathological type | Total benign cases (n=5, 62.5%) | Percentage |
| Serous cystadenoma     | 09                         | 39.13%           | 19                      | 95%            | 04                        | 80%         | 48                       | 85.72% |
| Mucinous cystadenoma   | 08                         | 34.78%           | 01                      | 5%             | 01                        | 20%         |
| Serous papillary cystadenoma | 03                       | 13.04%           |                         |                |                           |             |                          |        |

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## Type of ovarian tumors

| Type of Tumor | Surface Epithelial Tumors | Germ Cell Tumors | Sex Cord Stromal Tumors |
|---------------|---------------------------|------------------|------------------------|
| Benign        |                           |                  |                        |
| Histopathological type | Total benign cases (n=23, 85.18%) | Percentage | Total benign cases (n=20, 95.24%) | Percentage | Total benign cases (n=5, 62.5%) | Percentage | Total |
| Benign transitional cell cystic ovarian carcinoma | 01 | 4.34% |                        |                        | 48 (85.72%) |                        |        |      |
| Cystadenofibroma | 01 | 4.34% |                        |                        |                        |                        |        |      |
| Seromucinous cystadenoma | 01 | 4.34% |                        |                        |                        |                        |        |      |
| Borderline    |                           |                  |                        |
| Histopathological type | Total borderline cases (n=2, 7.41%) | Percentage | Total borderline cases (n=0) | Percentage | Total borderline cases (n=0) | Percentage | Total |
| Seromucinous tumor | 01 | 50% |                        |                        |                        |                        |        |      |
| Mucinous tumors | 01 | 50% |                        |                        |                        |                        |        |      |
| Malignant     |                           |                  |                        |
| Histopathological type | Total malignant cases (n=2, 7.41%) | Percentage | Total malignant cases (n=1, 4.76%) | Percentage | Total malignant cases (n=3, 37.5%) | Percentage | Total |
| Seromucinous cystadenocarcinoma | 01 | 50% | Dysgerminoma | 01 | 100% | Granulosa cell tumor of ovary – Adult type | 03 | 100% | 6 (10.71%) |
| Endometroid carcinoma | 01 | 50% |                        |                        |                        |                        |        |      |
| Grand total   |                           |                  |                        | 56 (100%) |

### DISTRIBUTION OF OVARIAN TUMORS

- Surface epithelial tumors: 48.21%
- Germ cell tumor: 37.5%
- Sex cord tumors: 14.29%
Fig. 1: DYSGERMINOMA. A: High power view of dysgerminoma shows Polygonal cells with moderate pleomorphic nuclei. B: Low power view of sheets of polygonal cells and lymphocytic infiltrate.

Fig. 2: ENDOMETRIOID CARCINOMA OF OVARY. A: Scanner view showing presence of endometrial glands in ovarian stroma. B: High power view of endometrial gland.

Fig. 3: GRANULOSA CELL TUMOR – ADULT TYPE A: Gross Picture of Granulosa cell Tumor, B: Microscopic Picture of Granulosa cell tumor shows Coffee bean nuclei and mitotic figures.
Out of total 56 neoplastic conditions, 48 cases (85.71%) were benign, 2 (3.57%) were borderline tumors and 6 (10.71%) were malignant in our study. The various Indian studies done by Gupta et al, Pilli et al, Mankar and Jain et al, kanthikar S.N. et al and one sri lankan study done by M. Thirukumar et al show similar distribution of neoplastic lesions.

Ovarian tumor may occur at any age, including infancy and childhood. Incidence rate, however increases with age, with the greatest number of new cases being diagnosed beyond 4th and 5th decade. In present study, benign tumors were more commonly seen in the age group 20-40 years, while malignant tumors presented commonly after 40 years of age. Similar observations were made in study done by Mondal et al. In our study, the age of the patients are ranging from minimum 11 years to maximum 65 years. The age range varied in different studies. Indian Studies done by Chanu et al, and Danish et al age range is 18-83 years, 5-72 years respectively.

Out of total 126 non neoplastic lesions, Simple serous cyst found most common (38.88%) lesion followed by endometriosis (27.77%) and corpus luteal cyst(12.69%). Sri lankan study done by M. Thirukumar et al shows 60.8% cases of endometriosis, 22.9% cases of corpus luteal cyst and 16.3% follicular cyst. Difference in percentage wise distribution of various non neoplastic lesions may be due to different geographical region where study conducted.

Histopathologically, surface epithelial tumors (48.21%) were the most common category of ovarian tumors encountered, followed by germ cell tumors (37.5%), sex cord stromal tumor (14.29 %). Various Indian studies done by Pilli et al, Kar et al, Kanthikar et al, Nirali et al and Shah et al show similar histopathological distribution of neoplastic ovarian tumors. However, there is difference in percentage of various tumors which depends upon the prevalence of this tumors in different geographic area.

From total 27 surface epithelial tumors, 23(85.18%) were benign tumors, 2(7.41%) were malignant and 2(7.41%)

Fig. 4: MATURE SOLID TERATOMA WITH GLIOMATOSIS PERITONEI. A: Gross picture of mature solid teratoma. B: Low power view of the mature solid teratoma which contains mature adipose tissue and sebaceous glands. C: Gross picture of Peritoneum. D: Low power view shows the implantation of glial tissue in Peritoneum.
were borderline tumors in present study. Most common benign surface epithelial tumor was serous cystadenoma (39.13%) followed by Mucinous cystadenoma (34.78%). These findings are correlated well with study done in Nepal by Pradhan A et al in which they reported benign serous and mucinous cystadenoma 46.15% and 38.4% respectively. Germ cell tumor is the second most common tumor found in our study. Total 21 cases of germ cell tumor were reported out of which 20 (95.24%) were benign and 1 (4.76%) was malignant. Of all 20 benign tumors, 19 (95%) were mature cystic teratoma and one was mature solid teratoma. The mature solid teratoma was associated with gliomatosis peritonei, which is the rare condition to be associated with mature teratoma. Incidence of mature cystic teratoma is comparable to other Indian studies done by Shah et al (83.64%) and Pradhan A et al (92.1%).

Sex cord stromal tumors constitute the 14.28% of total neoplastic lesions. Of total 8 cases, 5 (62.5%) were benign, 3 (37.5%) were malignant cases. All three Malignant tumors were Granulosa cell tumor- Adult type. This data is comparable to the study done by Mankar and Jain et al in which they reported 9.34% cases of sex cord stromal tumors were reported.

Table 5: Comparison of neoplastic tumor type with other studies.

| Neoplastic lesion | Gupta et al.[12] | Pilli et al.[9] | Mankar and Jain et al [19] | Kanthikar S.N. et al.[1] | M. Thirukar et al.[4] | Present study |
|-------------------|-----------------|-----------------|-----------------------------|--------------------------|------------------------|--------------|
| Benign            | 72.9%           | 75.2%           | 63.04%                      | 78.57%                   | 80.1%                  | 85.71%       |
| Borderline        | 4.1%            | 2.8%            | 5.84%                       | 1.42%                    | 3.7%                   | 3.57%        |
| Malignant         | 22.9%           | 21.8%           | 31.12%                      | 20%                      | 16.2%                  | 10.71%       |

Table 6: Comparison of the histopathological spectrum of neoplastic ovarian tumors.

| Study            | Year | Surface epithelial tumors | Germ cell tumors | Sex cord stromal tumors |
|------------------|------|---------------------------|------------------|-------------------------|
| Pilli et al [9]  | 2001 | 71%                       | 21%              | 07%                     |
| Kar et al [10]   | 2005 | 79%                       | 16%              | 1.50%                   |
| Kanthikar et al [9] | 2014 | 67.14%                  | 22.85%           | 5.71%                   |
| Niral et al [11] | 2015 | 73.8%                     | 17.8%            | 6.1%                    |
| Shah et al [10]  | 2016 | 64.98%                    | 25.35%           | 8.29%                   |
| Present study    | 2020 | 48.21%                    | 37.5%            | 14.29%                  |

**Conclusion**

We found that ovarian lesions affect wide variation of age starting from 11 years young patient to 65 years old patients. Non neoplastic lesions were almost double in prevalence than neoplastic lesions and more commonly found in age group of 21 to 30 years. Histopathological analysis according to WHO classification reveal that surface epithelial tumors and germ cell tumors were forms the majority of neoplastic lesions.

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