

# The clinical characteristics and prognosis of ovarian endometrioid carcinoma

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## Abstract

**Objective:** To investigate the clinical characteristics and relative factors of ovarian endometrioid carcinoma.

**Methods:** The clinical data of 36 patients with ovarian endometrioid carcinoma were retrospectively analyzed.

**Results:** Age of the patients with tumor was relatively young (61.3% < 55 years). Major clinical manifestations are pain, bloating, and abnormal vaginal hemorrhage. Ultrasound examination commonly showed cystic-solid lesion. Five-year survival rate of patients in tumor stage I-II was up to 77.9%, stage III to 36.8%, patients undergoing satisfied cytoreductive surgery to 68.5%, patients with ovarian endometrioid carcinoma without endometrial cancer surgery to 87.4%, and the patients with endometrial cancer surgery to 27.4%.

**Conclusion:** Prognosis of ovarian endometrioid carcinoma patients undergoing cytoreductive surgery early is favorable, but prognosis of those patients with endometrial cancer is poor.

## Introduction

With the progress of pathologic diagnosis technology, more and more patients with ovarian endometrioid carcinoma are detected early. For exploring the diagnosis, treatment and prognostic of ovarian endometrioid carcinoma, the clinical data of 36 patients with ovarian endometrioid carcinoma were analyzed retrospectively.

## Material and method

### Subjects

From Jan1, 2007 to Nov 30, 2016, a retrospective analysis was performed on 36 patients with ovarian endometrioid carcinoma. Clinical data were analyzed and follow-up by telephone were made after operation till Mar 31, 2014, 5 patients were lost to follow-up, with a follow-up rate of 86.1%. Thirty-one patients with ovarian endometrioid carcinoma were included in this study.

### Treatment

**Operation:** All the 31 patients with ovarian endometrioid carcinoma received operation. The basic pattern of operation is cytoreductive surgery (including removal of total hysterectomy, bilateral adnexectomy, appendix, omentum majus, pelvic metastases, and pelvic lymphadenectomy). Recurrence within six months is considered unsatisfied cytoreductive surgery.

**Chemotherapy:** 30 cases received additional chemotherapy one or two weeks after operation. Chemotherapy is with TP therapy: carboplatin (CBP) AUC4-5 plus taxol135 mg/m<sup>2</sup>. 1 case gave up for being too old.

### Statistics analysis

Data analysis was performed using the statistical software package SPSS 19.0 (SPSS Inc., Chicago, USA). 5-year survival rate was analyzed

with Kaplan-Meier method and comparison of groups was with Chi-square test or Fisher's test. *P* value < 0.05 was considered significant.

## Results

### Comparison of 5-year survival rate

**Ages:** 5-year survival rate of patients aged less than 55 years was 70.4%, while patients aged 55 years or more was 49.6%. There was no significant difference between them. (*P* = 0.326)

**Tumor stage:** 5-year survival rate of patients in tumor stage I-II from FIGO was 77.9%, patients in tumor stage III was 36.8%, and there was significant difference between them (*P*=0.033). 5-year survival rate of patients in tumor stage I-II was higher than 5-year survival rate of patients in tumor stage III.

**Tumor marker:** 5-year survival rate was 83.3% when preoperative CA125 < 150u/ml; while 5-year survival rate was 54.2% when CA125 ≥ 150u/ml. No statistical differences were found between groups (*P*=0.268).

**Pathologic type:** 5-year survival rate of patients with simple type ovarian endometrioid carcinoma and mixed type ovarian endometrioid carcinoma was 63.3% and 63.6% respectively and there was no significant difference between them (*P*=0.528). 5-year survival rate of patients with endometrial cancer or without was 87.4% and 27.4% respectively, with significant difference (*P* = 0.015).

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**Operation:** 5-year survival rate of patients underwent satisfied cytoreductive surgery and without cytoreductive surgery was 68.5% and 0.0% respectively ( $P < 0.001$ ). The patients were divided into two groups according to chemotherapy course (Group 1: chemotherapy course  $< 6$ , group 2: chemotherapy course  $\geq 6$ ). 5-year survival rate of group 1 and group 2 was 57.8% and 75% respectively. There was no significant difference between them ( $P = 0.581$ ) (Table 1).

## Discussion

### Clinical pathological features

It was reported that the incidence of ovarian endometrioid carcinoma was 3.1% to 24.4% in patients with ovarian cancer [1,2]. The patients with ovarian endometrioid carcinoma accounted for 3.5% (36 cases) of the ovarian malignant tumor at the corresponding period in our study. Ovarian endometrioid carcinoma has two pathologic types: endometrial carcinogenesis and epithelial differentiation of germinal epithelium, first reported by Sampson and Santesson. respectively. Pathologic diagnosis need to detect endometriosis ectopic focus location malignant cells or differentiation histology manifestation of ovary cambium epithelia toward endometrium in our hospital. Among the 36 cases, 4 patients with endometriosis were accompany with ovarian endometrioid adenocarcinoma, up to 12.9%, which was consistent with the study of Sainz de la Cuesta R, ranged from 10% to 28% [2]. Ratio of malignant transformation from endometriosis to cancer was not high. Erzen *et al.* [3] thought that 1/3 of the patients with ovarian endometriosis malignant neoplasms had endometriosis history. This report showed that tissue type of canceration of endometriosis was mainly ovarian endometrioid

carcinoma, accounting for 75% [4], it indicated that endometriosis might undergo probably malignant transformation. Some studies indicated that 40.0% of ovarian endometrioid carcinoma came from of malignant transformation of ovarian endometriosis, and that the incidence of endometriosis (atypical hyperplasia endometriosis) was 37.5%, twice as much as that reported in other reports. Most of them were confirmed after re-slice and review, indicating that diagnosis level of pathologic doctor is of crucial importance. Pathologic doctors need to improve diagnosis level and enhance collaboration with clinicians to make accurate, comprehensive, and careful pathological evaluation. Lee *et al.* [5] argued that combination of routine pathological microscopy and immunohistochemical method can diagnose ovarian cancer malignant transformation from endometriosis or primary ovarian endometrioid carcinoma more accurately, which provided sound identifying method. Our study found that there were 18 cases of ovarian endometrioid carcinoma with endometrial cancer, 1 case with mucinous adenocarcinoma, 7 cases with cystadenocarcinoma and 1 case with clear cell carcinoma, indicating that this cancer was more commonly derived from ovary epithelial cell multi-differentiation. Therefore we concluded that the primary pathological manifestation of this cancer was the cancer cell differentiation of ovary germinal epithelium toward endometrium. According to the diagnosis criterion of double primary carcinoma of uterine corpus and the ovary proposed by Scully and Young [6], our data displayed 14 cases of ovarian endometrioid carcinoma combining with endometrioid carcinoma, accounting for 45.2%, indicating ovarian endometrioid carcinoma was easy intercurrent double carcinoma. The main clinical manifestations of the patients with ovarian endometrioid carcinoma were pain and bloating (80.6%) and vaginal irregular bleeding (54.8%). Tumor size  $> 5$  cm by ultrasound examination made up 78.9%. The bigger tumor usually accompanied with abdominal discomfort, urging the patient to see the doctor. Irregular colporrhagia patients were often required to go to the hospital for check. It made the early stage cancer more easily detected [7]. Consequently, our research shown it was 61.3% in patients with stage I and II. Early detection provided possibility for early treatment. In the late stage, symptoms such as bloating, abdominal pain, and abnormal vaginal bleeding often appeared. The degree of symptoms was associated with the size and location of the tumor, degree of tumor invasion to nearby organs, and the complications [8].

### Treatment

Our treatment was based on traditional operation combined with post operative chemotherapy. It was reported that the occurrence and development of endometrial adenocarcinoma was likely associated with stimulation of estrogen. It was unknown that the development of ovarian endometrioid carcinoma was likely connected with the stimulation of estrin just as endometrial cancer. How to express oestrogen receptor in ovarian endometrioid carcinoma? However, it is rare on this aspect at present.

Progestin can markedly antagonize on the biological effects of estrogen. Experiment *in vitro* also verifies progestin's apparent inhibiting effect on the ovarian cancer cell lines proliferation. Progestin has been reported that it would lead to endometrial carcinoma cell differentiation and change of cancerous endometrium secretion phase, significantly reducing mitosis and inhibiting further dysplasia and malignancy, which indicates that progestin has direct inhibition function on tumor cells [9]. However, these researches are still in the initial stage and have not been put into clinical application. We have not given hormone therapy yet. Since all the patients received radiotherapy at other hospitals, the results of radiotherapy were not evaluated.

**Table 1.** Clinical features of patients

Item	Case (%)	P-value
Age(year)		0.205
≤55	19(61.3)	
>55	12(38.7)	
clinical features		0.426
pain, bloating	25(80.6)	
vaginal bleeding	17(54.8)	
pelvis masses found in physical examination	4(12.9)	
Frequent urination after menopause	1(3.2)	
preoperative CA125 level (U/L)		0.029
≤150	11(35.5)	
>150	20(64.5)	
preoperative ultrasound feature		0.651
cystic lesion	6(19.4)	
solid lesion	8(25.8)	
mixed cystic-solid lesion	17(54.8)	
tumor diameter (cm)		0.473
<5	6(22.1)	
5-10	15(48.9)	
>10	9(29.0)	
FIGO staging		
Stage I	10(32.3)	
Stage II	9(29.0)	
Stage III	12(38.7)	
histological type		0.738
Simple type endometrioid carcinoma	18(58.1)	
Mixed type endometrioid carcinoma	13(41.9)	
concomitant endometrioid carcinoma cases	14(45.2)	
TP chemotherapy course		<0.01
□ < 6	28(90.3)	
≥ 6	3(9.7)	

## Prognosis

Our research found that there was significant difference between the patient with tumor stage I-II and stage III in the 5-year survival rates. Therefore, early diagnosis and early surgical operation is the key to prognosis improvement. Preoperative ultrasonograph showed that the focus of infection was cystic mixed lesion, highly similar in imaging with ovarian endometrial cysts, indicating the lack of characteristic ultrasonographic manifestation. It can only serve as morphologic index of tumor size instead of histological evidence. So far, CA125 is still the preferred marker of ovarian epithelial carcinomas. This research set 150 u/ml as the segmentation point and found that there was no statistical difference between the group of preoperative CA125 < 150 u/ml and group of CA125 ≥ 150 u/ml in prognosis, indicating that preoperative CA125 level could only contribute to diagnosis basis but not as prognosis judgment. Platinum-based combination chemotherapy has become the first-line chemotherapeutic regimen for ovarian epithelial carcinomas. One study [10] suggested that the efficacy of TP therapy was superior to other platinum-based chemotherapy. Yet patients did not receive better prognosis after they experienced 6 or more chemotherapy courses, possibly due to the small sample scale of chemotherapy patients with 6 or more chemotherapy courses thus the difference was not reflected accurately. It need further study in larger scale. It was reported that the ovarian endometrioid carcinoma had a 5-year survival rate of around 40% to 50% and had better prognosis than serouscarcinoma or mucous carcinomas [11]. Among patients in this group, 5-year postoperative survival rate was 54.8%, slightly better than literature. Due to the high pathologic diagnosis requirement, the number of confirmed cases was deficient, avoiding other pathologic type ovarian cancer to be misdiagnosed as endometrioid carcinoma, indicating that strict pathologic diagnosis and distinction of this cancer from ovarian cancer might guarantee a better prognosis for patients. Patients with tumors associated with endometriosis had a higher rate of synchronous endometrial cancer. Cases also demonstrated a lower rate of recurrence and improved 5 year DFS [12]. Women with EEOC and concurrent endometriosis showed distinct characteristics and had longer disease-free survival when compared with those without endometriosis [13]. Ovarian endometrioid carcinoma shows great variability in the onset age. In our research, the median age was 52, and patients less than 55 years old accounted for 61.3%. Our study shown there was no correlation closely between the age of onset and prognosis of ovarian endometrioid carcinoma. The reason might be the small sample size. Hence there was no significant difference. Initial cytoreductive surgery satisfaction degree was closely related to prognosis. Cases of postoperative return of ovarian endometrioid carcinoma within 6 months all died within 5 years. Therefore efforts should be made in perfecting the cytoreductive surgery to improve the prognosis of patients.

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