Histological tumour type (Required)

Reason/Evidentiary Support

All ovarian epithelial malignancies and borderline tumours should be typed according to the WHO classification.¹ There are 5 major subtypes of primary ovarian carcinoma, high-grade serous, clear cell, endometrioid, mucinous and low-grade serous.²⁻⁵ There are also other uncommon minor subtypes, those listed by the WHO including malignant Brenner tumour, seromucinous carcinoma and undifferentiated carcinoma.¹ Carcinosarcoma is a mixed epithelial and mesenchymal malignancy but is included in the category of epithelial malignancies in this dataset since most are of epithelial origin and histogenesis.⁶

Although management of ovarian carcinoma is, at present, largely dependent on tumour stage and grade, accurate typing will almost certainly become more important in the future with the introduction of targeted therapies and specific treatments for different tumour types. This is in part because, although clinically often considered as one disease, there is an increasing realisation that the different morphological subtypes of ovarian carcinoma have a different pathogenesis, are associated with distinct molecular alterations and have a different natural history, response to traditional chemotherapy and prognosis. Tumour typing may also be important in identifying or initiating testing for an underlying genetic predisposition; for example, high-grade serous carcinoma may be associated with underlying BRCA1/2 mutation while endometrioid and clear cell carcinomas can occur in patients with Lynch syndrome. The most common ovarian carcinoma is high-grade serous carcinoma (approximately 70%) followed by clear cell and endometrioid. Mucinous and low-grade serous are less common. Approximately 90% of advanced stage ovarian carcinomas (stage III/IV) are high-grade serous in type. Serous are less common.

Most primary tubal carcinomas are high-grade serous or endometrioid and most primary peritoneal carcinomas are of high-grade serous type. As discussed in the sections on tumour site, it may be difficult to ascertain the origin of a high-grade serous carcinoma since multiple sites are often involved.

Mixed ovarian carcinomas are now considered to be uncommon. The current 2014 WHO classification does not include a category of mixed carcinoma² but the prior classification stated that a diagnosis of mixed carcinoma should only be made if the minor component represents more than 10% of the neoplasm. However, it is recommended that all different morphological subtypes in an ovarian carcinoma are documented, even if they comprise less than 10% of the neoplasm. As stated, mixed carcinomas in the ovary are uncommon, the most prevalent combination being clear cell and endometrioid (both of these tumour types often arise in endometriosis). Most neoplasms which were previously classified as mixed serous and endometrioid and mixed serous and clear cell represent high-grade serous carcinomas with pseudoendometrioid areas and areas of cytoplasmic clearing respectively. In such cases, immunohistochemical markers, especially WT1, may be useful (see **Note 20 IMMUNOHISTOCHEMICAL MARKERS**).

Borderline tumours should also be typed according to WHO criteria. The most common subtypes are serous and mucinous (intestinal type). Seromucinous, endometrioid, clear cell and Brenner subtypes also occur.

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