

Adrenal Gland

**Protocol applies to adrenal cortical carcinoma.
Pheochromocytoma, neuroblastoma, and other adrenal
medullary tumors of childhood are excluded.**

*Protocol revision date: January 2004
No AJCC/UICC staging system*

Procedures

- **Cytology** (No Accompanying Checklist)
- **Incisional Biopsy**
- **Excisional Biopsy**
- **Adrenalectomy**

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Surgical Pathology Cancer Case Summary (Checklist)

Protocol revision date: January 2004

Applies to invasive carcinomas only

No AJCC/UICC staging system

ADRENAL CORTICAL CARCINOMA: Resection

Patient name:

Surgical pathology number:

Note: Check 1 response unless otherwise indicated.

MACROSCOPIC

Specimen Type

Subtotal adrenalectomy

Total adrenalectomy

Other (specify): _____

Not specified

Laterality

Right

Left

Not specified

Tumor Size

Greatest dimension: ___ cm

*Additional dimensions: ___ x ___ cm

Cannot be determined (fragmented specimen)

Tumor Weight

Specify: ___ g

MICROSCOPIC

Pathologic Staging

Primary Tumor

I: Confined to gland, 5 cm or less

II: Confined to gland, greater than 5 cm

III: Extraglandular extension without other organ involvement

IV: Distant metastasis or extension into other organs

Cannot be determined

- 2 * Data elements **with asterisks** are **not required** for accreditation purposes for the Commission on Cancer. These elements may be clinically important, but are not yet validated or regularly used in patient management. Alternatively, the necessary data may not be available to the pathologist at the time of pathologic assessment of this specimen.

Regional Lymph Nodes

- Cannot be assessed
 - No regional lymph node metastasis
 - Regional lymph node metastasis
- Specify: Number examined: ____
Number involved: ____

Distant Metastasis

- Cannot be assessed
 - Distant metastasis
- *Specify site(s), if known: _____

Margins

- Margins uninvolved by tumor
 - Margin(s) involved by tumor
- Specify site(s) of involvement: _____
- Involvement by tumor cannot be determined

***Venous (Large Vessel) Invasion**

- * Absent
- * Present
- * Indeterminate

***Additional Pathologic Findings (check all that apply)**

- * None identified
- * Tumor necrosis
- * Hyperplasia
- * Adenoma
- * Other (specify): _____

***Comment(s)**

* Data elements **with asterisks** are **not required** for accreditation purposes for the Commission on Cancer. These elements may be clinically important, but are not yet validated or regularly used in patient management. Alternatively, the necessary data may not be available to the pathologist at the time of pathologic assessment of this specimen.

Background Documentation

Protocol revision date: January 2004

I. Cytologic Material

A. Clinical Information

1. Patient identification
 - a. Name
 - b. Identification number
 - c. Age (birth date)
 - d. Sex
2. Responsible physician(s)
3. Date of procedure
4. Other clinical information
 - a. Relevant history (Note **A**)
 - b. Relevant findings (eg, hormonal and imaging studies) (Note **B**)
 - c. Clinical diagnosis
 - d. Procedure (eg, fine-needle aspiration [FNA])
 - e. Anatomic site(s) (eg, right/left adrenal gland, related sites)

B. Macroscopic Examination

1. Specimen
 - a. Unfixed/fixed (specify fixative)
 - b. Number of slides received
 - c. Quantity and appearance of fluid specimen
 - d. Other (eg, cytologic preparation from tissue)
 - e. Results of rapid smear review
2. Material submitted for microscopic evaluation (eg, direct smear, cytocentrifuge preparation, touch or filter preparation, cell block)
3. Special studies (Note **C**)

C. Microscopic Evaluation

1. Adequacy of specimen (if unsatisfactory for evaluation, specify reason)
2. Tumor, if present
 - a. Histologic type, if possible (Note **D**)
 - b. Other descriptive features (eg, nuclear atypia, necrosis)
3. Additional pathologic finding, if present
4. Results/status of special studies (specify) (Note **C**)
5. Comments
 - a. Correlation with intraoperative consultation, as appropriate
 - b. Correlation with other specimens, as appropriate
 - c. Correlation with clinical information, as appropriate

II. Incisional Biopsy (Any Surgical Approach Less Than Complete Adrenal Excision)

A. Clinical Information

1. Patient identification
 - a. Name
 - b. Identification number
 - c. Age (birth date)
 - d. Sex

2. Responsible physician(s)
3. Date of procedure
4. Other clinical information
 - a. Relevant history (Note **A**)
 - b. Relevant findings (eg, hormonal and imaging studies) (Note **B**)
 - c. Clinical diagnosis
 - d. Procedure (eg, fine-needle biopsy, core biopsy, incisional biopsy)
 - e. Operative findings
 - f. Anatomic sites (eg, right/left adrenal gland, related sites)

B. Macroscopic Examination

1. Specimen
 - a. Organ(s)/tissue(s) (specify)
 - b. Unfixed/fixed (specify fixative)
 - c. Number of fragments
 - d. Dimensions
 - e. Weight, if appropriate
 - f. Orientation, if indicated by surgeon
 - g. Descriptive features
 - h. Results of intraoperative consultation
2. Tumor(s), if identified
 - a. Location
 - b. Dimensions
 - c. Descriptive features (eg, hemorrhage/necrosis)
 - d. Relationship to margins, if appropriate
3. Additional pathologic findings, if present (eg, hyperplasia)
4. Tissue submitted for microscopic evaluation
 - a. Tumor
 - b. Margin(s), if appropriate
 - c. Nodules
 - d. Other lesions
 - e. Frozen section tissue fragment(s) (unless saved for special studies)
5. Special studies (specify) (Note **C**)

C. Microscopic Evaluation

1. Tumor, if present
 - a. Histologic type (Note **D**)
 - b. Descriptive features (eg, nuclear atypia, necrosis) (Note **E**)
 - c. Venous/lymphatic vessel invasion
2. Additional pathologic findings, if present (eg, hyperplasia)
3. Results/status of special studies (specify) (Note **C**)
4. Comments
 - a. Correlation with intraoperative consultation, as appropriate
 - b. Correlation with other specimens, as appropriate
 - c. Correlation with clinical information, as appropriate

III. Complete Excision (Including Laparoscopically Removed Glands)**A. Clinical Information**

1. Patient identification
 - a. Name
 - b. Identification number

- c. Age (birth date)
- d. Sex
2. Responsible physician(s)
3. Date of procedure
4. Other clinical information
 - a. Relevant history (Note **A**)
 - b. Relevant findings (eg, hormonal and imaging studies) (Note **B**)
 - c. Clinical diagnosis
 - d. Procedure (eg, laparoscopically removed gland) (Note **F**)
 - e. Operative findings
 - f. Type of specimen (adrenal excision with or without surrounding soft tissues)
 - g. Anatomic site(s) of specimen (eg, right/left adrenal gland, related sites)

B. Macroscopic Examination

1. Specimen
 - a. Organ(s)/tissue(s) included
 - b. Unfixed/fixed (specify fixative)
 - c. Dimensions
 - d. Weight
 - e. Orientation, if indicated by surgeon
 - f. Descriptive features
 - g. Results of intraoperative consultation (Note **C**)
2. Tumor(s)
 - a. Dimensions (3)
 - b. Weight (Note **G**)
 - c. Descriptive features (eg, color, consistency, hemorrhage, necrosis)
 - d. Extent of invasion (Note **H**)
3. Margins, relationship to and distance from tumor, as appropriate
4. Regional lymph nodes, if submitted
 - a. Number
 - b. Location, if designated by surgeon
5. Additional pathologic findings, if present
6. Tissue(s) submitted for microscopic evaluation
 - a. Tumor, adequate sampling of all areas
 - b. Nodules
 - c. Margins of resection
 - d. All lymph nodes
 - e. Other lesions
 - f. Frozen section tissue fragment(s) (unless saved for special studies)
 - g. Other organs/tissues (eg, liver biopsy)
7. Special studies (specify) (Note **C**)

C. Microscopic Evaluation

1. Tumor
 - a. Histologic type (Note **D**)
 - b. Descriptive features (eg, nuclear atypia, mitotic rate, necrosis) (Note **E**)
 - c. Extent of invasion (Note **H**)
 - d. Venous/lymphatic vessel invasion
2. Margins, as appropriate
3. Regional lymph nodes (Note **H**)
 - a. Number (location, if possible)

- b. Number involved by tumor
- 4. Additional pathologic findings, if present (eg, hyperplasia)
- 5. Result/status of special studies (specify) (Note **C**)
- 6. Other organs/tissues
- 7. Comments
 - a. Correlation with intraoperative consultation, as appropriate
 - b. Correlation with other specimens, as appropriate
 - c. Correlation with clinical information, as appropriate

Explanatory Notes

A. Relevant History

Endocrine manifestations, such as hypertension, change in body habitus, feminization, or virilism, are important. Also of import are family history, previous surgery for adrenal tumors (both benign and malignant) or other endocrine organs, other tumors that may metastasize to the adrenal gland, and endocrine or other therapies. Hyperplastic adrenal tissue may re-grow if previously excised incompletely.

B. Endocrine Status

Laboratory findings are important in the evaluation of an adrenal mass that is not obviously a high-grade carcinoma because the absence of evidence of hormonal excess in the presence of an enlarged adrenal gland usually indicates that the tumor is an incidental finding (“incidentaloma”) and not a functioning adenoma.¹

C. Special Procedures

Special procedures may include frozen sections, cytologic imprints, immunohistochemical stains, histochemical stains, electron microscopy, flow cytometry, molecular studies, and cytogenetic studies. If such studies are performed in another laboratory, either extrainstitutional or intrainstitutional, the laboratory should be identified.

D. Histologic Type

The following histologic classification of adrenal tumors has been modified from Page et al.²

Histologic Classification of Adrenal Tumors

Cortical Tumors

- Adenoma
- Carcinoma
- Myelolipoma
- Miscellaneous

Medullary Tumors[#]

- Pheochromocytoma[#]
- Neuroblastoma[#]
- Ganglioneuroblastoma[#]
- Ganglioneuroma[#]

[#] Not covered in protocol.

E. Histologic Grade

Adrenal cortical tumors are not usually graded on histologic grounds. Severe nuclear atypia, high mitotic count, vascular invasion, tumor necrosis, and other microscopic features may, in combination, support a diagnosis of adrenal cortical carcinoma over adenoma and should be recorded, but no precise clustering of histologic features is considered diagnostic of malignancy. However, when several malignant features are present together (eg, highly atypical nuclei, sheet-like growth, necrosis, and many mitoses), the risk of distant metastases is increased.³⁻⁶ In some studies, specific combinations of features, such as mitotic rates of 6 or more per 50 high-power fields (HPF) along with atypical mitosis and venous invasion, have been found to correlate with metastasis or recurrence of adrenal cortical carcinomas.⁴ Other studies have shown that mitotic rates greater than 20 per 50 HPF are associated with decreased survival, suggesting that a high mitotic index may be an important adverse prognostic factor.⁵

Although this protocol does not cover medullary tumors, it should be noted that pheochromocytoma is usually diagnosed preoperatively by pharmacologic means. No pathologic criteria for differentiation of benign from malignant pheochromocytomas have been defined. Metastatic disease is considered the only irrefutable proof of malignancy.

F. Laparoscopic Surgery

An entire adrenal tumor may be removed laparoscopically, but with this technique, the gland may become fragmented. This anatomic information, including maximal diameter of the resected tumor, should be provided by the surgeon.

G. Weight

Accurate weights of adrenal cortical neoplasms are important.⁶ Although tumor mass cannot be used as the sole criterion for malignancy, adrenal cortical neoplasms weighing less than 50 g are almost always benign, whereas the weight of malignant tumors is usually greater than 100 g.

H. Staging

The staging system proposed by MacFarlane⁷ and modified by Sullivan et al⁸ and Henley et al⁹ is most commonly used for adrenal cortical carcinomas. The American Joint Committee on Cancer (AJCC) and the International Union Against Cancer (UICC) have no published TNM staging system for malignancies of the adrenal gland.

Stage	Extent	Size
I	Confined to gland	5 cm or less
II	Confined to gland	Greater than 5 cm
III	Extends out of gland without involving adjacent organs	Any
IV	Distant metastasis or involvement of adjacent organs	Any

*Tumors of the Adrenal Gland and Extra-Adrenal Paraganglia*¹⁰ proposes the following staging system.

Primary Tumor (T)

- T1 Tumor 5 cm or less, no invasion
 T2 Tumor greater than 5 cm, no invasion
 T3 Tumor of any size, locally invasive but not involving adjacent organs
 T4 Tumor of any size with invasion of adjacent organs

Regional Lymph Nodes (N)

- N0 Negative regional nodes
 N1 Positive regional nodes

Distant Metastasis (M)

- M0 No distant metastasis
 M1 Distant metastasis

Stage Definitions

Stage I	T1	N0	M0
Stage II	T2	N0	M0
Stage III	T1,T2	N1	M0
	T3	N0	M0
Stage IV	Any T	Any N	M1
	T3,T4	N1	M0

References

1. Kloos RT, Gross MD, Francis IR, Dorobkin M, Shapiro B. Incidentally discovered adrenal masses. *Endocrine Reviews*. 1995;16:460-484.
2. Page DL, DeLellis RA, Hough AJ Jr. *Tumors of the Adrenal; Atlas of Tumor Pathology*. 2nd Series, Fascicle 23. Washington, DC: Armed Forces Institute of Pathology; 1986.
3. Hough AJ, Hollifield JW, Page DL, Hartmann WH. Prognostic factors in adrenocortical tumors: a mathematical analysis of clinical and morphologic data. *Am J Clin Pathol*. 1979;72:390-399.
4. Weiss LM. Comparative histologic study of 43 metastasizing and non-metastasizing adrenocortical tumors. *Am J Surg Pathol*. 1984;8:163-169.
5. Weiss LM, Medeiros LJ, Vickery AL. Pathologic features of prognostic significance in adrenal cortical carcinoma. *Am J Surg Pathol*. 1989;13:202-206.
6. Medeiros LJ, Weiss LM. New developments in the pathologic diagnosis of adrenal cortical neoplasms: a review. *Am J Clin Pathol*. 1992;97:73-83.
7. MacFarlane DA. Cancer of the adrenal cortex: the natural history, prognosis, and treatment in a study of fifty-five cases. *Ann R Coll Surg Engl*. 1958;23:155-186.
8. Sullivan M, Boileau M, Hodges CV. Adrenal cortical carcinoma. *J Urol*. 1978;120:660-665.
9. Henley DJ, van Heerden JA, Grant CS, Carney JA, Carpenter PC. Adrenal cortical carcinoma: a continuing challenge. *Surgery*. 1983;94:926-931.
10. Lack E. *Tumors of the Adrenal Gland and Extra-Adrenal Paraganglia*. AFIP Fascicle No. 19. Third Series. Washington DC: American Registry of Pathology; 1997.

Bibliography

Dehner LP. Neoplasms of the adrenal cortex: preoccupation bordering on obsession [editorial; comment]. *Am J Clin Pathol.* 1994;101:557-558.

Katz RL, Patel S, Mackay B, Zornoza J. Fine-needle aspiration cytology of the adrenal gland. *Acta Cytol.* 1984;28:269-282.

Lack EE, ed. *Pathology of the Adrenal Glands. Contemporary Issues in Surgical Pathology.* Vol. 14. New York: Churchill Livingstone; 1990.

Lloyd RV. *Endocrine Pathology.* New York: Springer-Verlag; 1990.

Peritoneum

Protocol applies to all primary borderline and malignant epithelial tumors, and malignant mesothelial neoplasms of the peritoneum.

*Protocol revision date: January 2004
No AJCC/UICC staging system*

Procedures

- **Cytology** (No Accompanying Checklist)
- **Biopsy** (No Accompanying Checklist)
- **Resection**

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Surgical Pathology Cancer Case Summary (Checklist)

*Protocol revision date: January 2004
Applies to primary borderline tumors,
carcinomas, and mesotheliomas only
No AJCC/UICC staging system*

PERITONEUM: Resection

Patient Name:

Surgical pathology Number:

Note: Check 1 response unless otherwise indicated.

MACROSCOPIC

Tumor Site(s)

Specify, if known: _____

___ Not specified

Organ(s) Included (if applicable):

Specify: _____

Tumor Size

Greatest dimension: ___ cm

*Additional dimensions: ___ x ___ cm

___ Cannot be determined (see Comment)

MICROSCOPIC

Histologic Type

___ Malignant mesothelioma, epithelial

___ Malignant mesothelioma, sarcomatous (spindle cell)

___ Malignant mesothelioma, biphasic

___ Malignant mesothelioma, other (specify): _____

___ Serous borderline tumor (of low malignant potential)

___ Serous carcinoma

___ Other malignant tumor of Mullerian type (specify):

___ _____
___ Desmoplastic small round cell tumor

___ Other (specify): _____

___ Malignant tumor, type cannot be determined

2

* Data elements **with asterisks** are **not required** for accreditation purposes for the Commission on Cancer. These elements may be clinically important, but are not yet validated or regularly used in patient management. Alternatively, the necessary data may not be available to the pathologist at the time of pathologic assessment of this specimen.

Histologic Grade

- Not applicable (borderline neoplasms and mesotheliomas)
- GX: Cannot be assessed
- G1: Well differentiated
- G2: Moderately differentiated
- G3: Poorly differentiated
- Other (specify): _____

***Venous/Lymphatic (Large/Small Vessel) Invasion (V/L)**

- * Absent
- * Present
- * Indeterminate

***Additional Pathologic Findings (check all that apply)**

- * None
- * Ferruginous bodies
- * Endosalpingiosis
- * Endometriosis
- * Other (specify): _____

***Comment(s)**

* Data elements **with asterisks** are **not required** for accreditation purposes for the Commission on Cancer. These elements may be clinically important, but are not yet validated or regularly used in patient management. Alternatively, the necessary data may not be available to the pathologist at the time of pathologic assessment of this specimen.

Background Documentation

Protocol revision date: January 2004

I. Cytologic Material

A. Clinical Information

1. Patient identification
 - a. Name
 - b. Identification number
 - c. Age (birth date)
 - d. Sex
 - e. Location (eg, ward, clinic, office)
2. Responsible physician(s)
3. Date of procedure
4. Clinical information
 - a. Relevant history
 - (1) present/past occupation
 - (2) asbestos exposure
 - (3) radiation exposure
 - (4) evidence of tumor(s) elsewhere
 - (5) prior tumor(s)
 - (6) prior operation(s)
 - b. Relevant findings (eg, radiologic studies, laboratory data, ascites [duration])
 - c. Clinical diagnosis
 - d. Operative findings (eg, unifocal, multifocal, or diffuse)
 - e. Type(s) or site(s) of specimen(s)
 - (1) ascitic fluid
 - (2) peritoneal washings (specify site[s])
 - (3) brushings (specify site[s])
 - (4) fine-needle aspirate (specify site[s])
 - (5) cytology preparation of tissue (touch preparation) (specify site[s])
 - (6) other

B. Macroscopic Examination

1. Specimen
 - a. Unfixed/fixed (specify fixative)
 - b. Number of slides received, if appropriate
 - c. Quantity and appearance of fluid specimen, if appropriate
 - d. Other (eg, cytologic preparation from tissue)
2. Material submitted for microscopic evaluation (eg, smear, cytocentrifuge, touch or filter preparation, cell block)

C. Microscopic Examination

1. Adequacy of specimen for evaluation (if unsatisfactory or limited, specify reason)
2. Tumor, if present
 - a. Histologic type, if possible (Note **A**)
 - b. Other characteristics (eg, grade, necrosis) (Note **B**)
3. Additional cytologic findings
4. Special studies (specify) (eg, cytochemistry, immunocytochemistry, electron microscopy, asbestos fiber count) (Note **C**)

5. Pathologic stage (Note **D**)
6. Comments
 - a. Correlation with intraoperative consultation, as appropriate
 - b. Correlation with other specimens, as appropriate
 - c. Correlation with clinical information, as appropriate

II. Biopsy

A. Clinical Information

1. Patient identification
 - a. Name
 - b. Identification number
 - c. Age (birth date)
 - d. Sex
 - e. Location (eg, ward, clinic, office)
2. Responsible physician(s)
3. Date of procedure
4. Clinical information
 - a. Relevant history
 - (1) present/past occupation
 - (2) asbestos exposure
 - (3) radiation exposure
 - (4) evidence of tumor(s) elsewhere
 - (5) prior tumor(s)
 - (6) prior operation(s)
 - b. Relevant findings (eg, radiologic studies, laboratory data, ascites [duration])
 - c. Clinical diagnosis
 - d. Procedure
 - e. Operative findings (eg, unifocal, multifocal, or diffuse)
 - f. Anatomic site(s) of specimen(s)

B. Macroscopic Examination

1. Specimen
 - a. Unfixed/fixed (specify fixative)
 - b. Number of pieces
 - c. Size or size range
 - d. Descriptive features
2. Submit entire specimen(s) for microscopic evaluation
3. Results of intraoperative consultation

C. Microscopic Examination

1. Tumor
 - a. Histologic type (Note **A**)
 - b. Histologic grade, if appropriate (Note **B**)
 - c. Other features of possible prognostic or therapeutic significance
2. Additional pathologic findings (eg, endosalpingiosis and relation to tumor, if pertinent)
3. Special studies (specify) (eg, histochemistry, immunohistochemistry, electron microscopy, asbestos fiber count) (Note **C**)
4. Pathologic stage (Note **D**)
5. Comments
 - a. Correlation with intraoperative consultation

- b. Correlation with other specimens, as appropriate
- c. Correlation with clinical information, as appropriate

III. Resection

A. Clinical Information

1. Patient identification
 - a. Name
 - b. Identification number
 - c. Age (birth date)
 - d. Sex
 - e. Location (eg, ward, clinic, office)
2. Responsible physician(s)
3. Date of procedure
4. Clinical information
 - a. Relevant history
 - (1) present/past occupation
 - (2) asbestos exposure
 - (3) radiation exposure
 - (4) evidence of tumor(s) elsewhere
 - (5) prior tumor(s)
 - (6) prior operation(s)
 - b. Relevant findings (eg, radiologic studies, laboratory data, ascites [duration])
 - c. Clinical diagnosis
 - d. Procedure
 - e. Operative findings (eg, unifocal, multifocal, or diffuse)
 - f. Anatomic site(s) of specimen(s)

B. Macroscopic Examination

1. Specimen
 - a. Organs/tissues received (specify)
 - b. Unfixed/fixed (specify fixative)
 - c. Number of pieces
 - d. Dimensions (measure attached tissues individually)
 - e. Orientation of specimen, if indicated by surgeon
 - f. Descriptive features
 - (1) peritoneal tissue(s)
 - i. outer surface (normal, adhesions, roughening, granularity, tumor)
 - ii. sectioned surface(s)
 - iii. size of tumor, if different from size of entire specimen; descriptive features; identification of areas for special study (eg, resection margin[s], if pertinent)
 - iv. other lesions
 - (2) organ(s) removed
 - i. outer surface (normal, adhesions, roughening, granularity, tumor)
 - ii. sectioned surface
 - (a) tumor(s) dimension(s) and distribution on or within organ, descriptive features, identification of areas for special study (eg, resection margin[s], if pertinent)
 - (b) other lesions
2. Tissues submitted for microscopic evaluation

C. Microscopic Examination

1. Tumor
 - a. Histologic type (Note **A**)
 - b. Histologic grade, if appropriate (Note **B**)
 - c. Other features of possible prognostic or therapeutic significance (eg, localized, diffuse)
2. Status of resection margins, if pertinent
3. Additional pathologic findings (eg, endosalpingiosis)
4. Special studies (specify) (eg, histochemistry, immunohistochemistry, electron microscopy, asbestos fiber count)
5. Pathologic stage (Note **D**)
6. Comments
 - a. Correlation with intraoperative consultation, as appropriate
 - b. Correlation with other specimens, as appropriate
 - c. Correlation with clinical information, as appropriate

Explanatory Notes**A. Histologic Type**

This protocol refers only to primary borderline and malignant epithelial tumors of the peritoneum. Secondary tumors, for example, those causing pseudomyxoma peritonei (almost always of appendiceal origin), are not addressed. However, in some cases "peritoneal spread" of a serous borderline tumor may actually reflect a primary peritoneal tumor rather than a metastasis from the ovary.

Classification of Peritoneal Tumors**Benign**

- Adenomatoid tumor
- Benign multicystic mesothelioma (multilocular peritoneal inclusion cyst)
- Mesothelial cyst(s) (unilocular) (free or attached)
- Well-differentiated papillary mesothelioma
- Solitary fibrous tumor (fibrous mesothelioma) (usually benign)

Malignant

- Diffuse malignant mesothelioma
 - Epithelial type
 - Sarcomatous type
 - Biphasic type
 - Rare types[#]
- Serous tumor of borderline malignancy (of low malignant potential)^{1,2 ##}
- Serous carcinoma^{3-5 ###}
- Malignant tumors of other Mullerian types
- Desmoplastic small round cell tumor
- Sarcomas

[#] Rare types include desmoplastic, small cell, lymphohistiocytoid, deciduoid, and undifferentiated types.

^{##} When this tumor involves the extraovarian peritoneum significantly and the ovarian surface minimally or not at all, it is generally considered to be of peritoneal origin.

The Gynecological Oncology Group has adopted the following criteria for the diagnosis of primary peritoneal serous carcinoma:

1. Both ovaries are either normal in size or enlarged by a benign process. In the judgement of the surgeon and the pathologist, the bulk of the tumor is on the peritoneum, and the extent of tumor involvement at 1 or more extraovarian sites is greater than that on the surface of or within either ovary.
2. Microscopic examination of the ovaries reveals: (a) no tumor; (b) tumor confined to the surface epithelium, with no evidence of cortical invasion; (c) tumor involving the ovarian surface and the underlying cortical stroma, but less than 5 x 5 mm in diameter; or (d) tumor less than 5 x 5 mm within the ovarian substance, with or without surface involvement.
3. The histologic and cytologic characteristics of the tumor are predominantly serous and similar or identical to those of ovarian serous papillary carcinoma of any grade.
4. If an oophorectomy has been performed in the past, a confident diagnosis of primary peritoneal serous carcinoma requires 1 of the following: (a) a pathology report to document the absence of carcinoma in the ovarian specimen, with review of all the slides if the oophorectomy has been performed within 5 years of the current procedure; (b) if the oophorectomy has been performed more than 5 years before the current procedure, the pathology report of the specimen should be obtained, and the slides should be reviewed if still available. The peritoneal tumor should be interpreted in light of the ovarian findings.

B. Histologic Grade

There is no established grading system for malignant mesotheliomas. Serous and other Mullerian-type tumors can be graded according to the criteria used for similar tumors in the female genital tract as shown below. (For further detail, see Ovary protocol.)

Grade X	Cannot be assessed
Grade 1	Well differentiated
Grade 2	Moderately differentiated
Grade 3	Poorly differentiated (tumors with minimal differentiation seen in very small foci)

C. Special Studies

Histochemical, immunohistochemical, and electron microscopic studies are helpful to routine microscopic evaluation in the diagnosis of mesothelioma. These tumors are usually mucicarmine and Pas-D negative. They may be positive for Alcian Blue or Colloidal Iron stains. Mesotheliomas usually are positive for different keratins, including cytokeratins 5/6, EMA, thrombomodulin, WT1, and calretinin. They are usually negative for CEA, B72.3, BER-EP4, and CD15 (Leu-M1), although they may be positive for single antibodies. In all these cases, a panel of antibodies is recommended. (For further detail, see Thoracic Mesothelium protocol.)

D. Staging of Peritoneal Tumors

There is no widely accepted staging system for peritoneal tumors, but their extent may have prognostic significance.⁶ Thus, it is important to determine whether a mesothelioma is unifocal, multifocal, or diffuse⁷; and whether there are lymph node or

distant metastases. Peritoneal serous carcinomas are generally staged as though they were stage II to stage IV ovarian cancers. (For further detail, see Ovary protocol.)

References

1. Bell DA, Scully RE. Serous borderline tumors of the peritoneum. *Am J Surg Pathol*. 1990;14:230-239.
2. Biscotti CV, Hart WR. Peritoneal serous micropapillomatosis of low malignant potential (serous borderline tumors of the peritoneum): a clinicopathologic study of 17 cases. *Am J Surg Pathol*. 1992;16:467-475.
3. Gilks CB, Bell DA, Scully RE. Serous psammocarcinoma of the ovary and peritoneum. *Int J Gynecol Pathol*. 1990;9:110-121.
4. Bloss JD, Liao S, Buller RE, et al. Extraovarian peritoneal serous papillary carcinoma: a case-control retrospective comparison to papillary adenocarcinoma of the ovary. *Gynecol Oncol*. 1993;50:347-351.
5. Piver MS, Jishi MF, Tsukuda Y, Nava G. Primary peritoneal carcinoma after prophylactic oophorectomy in women with a family history of ovarian cancer: a report of the Gilda Radner Familial Ovarian Cancer Registry. *Cancer*. 1993;71:2751-2755.
6. Wittenkind CH, Henson DE, Hutter RVP, Sobin LH. *TNM Supplement. A Commentary on Uniform Use*. 2nd ed. New York: Wiley-Liss; 2001.
7. Goldblum J, Hart WR. Localized and diffuse mesotheliomas of the genital tract and peritoneum in women: a clinicopathologic study of nineteen true mesothelial neoplasms, other than adenomatoid tumors, multicystic mesotheliomas, and localized fibrous tumors. *Am J Surg Pathol*. 1995;19:1124-1137.

Bibliography

- Antman KH, Pass HI, Schiff PB. Benign and malignant mesothelioma. In: DeVita VT Jr, Hellman S, Rosenberg SA, eds. *Cancer. Principles and Practice of Oncology*. 5th ed. Philadelphia, Pa: JB Lippincott-Raven; 1997:1853-1878.
- Battifora H, McCaughey WTE. *Tumors of the Serosal Membranes. Atlas of Tumor Pathology*. 3rd series. Fascicle 15. Washington, DC: Armed Forces Institute of Pathology; 1995.
- Scully RE, Young RH, Clement PB. *Tumors of the Ovary, Maldeveloped Gonads, Fallopian Tube, and Broad Ligament. Atlas of Tumor Pathology*. 3rd series. Fascicle 23. Washington, DC: Armed Forces Institute of Pathology; 1997.
- Wick MR. Immunophenotyping of malignant mesothelioma [editorial]. *Am J Surg Pathol*. 1997;21:1395-1398.
- Young RH, Clement PB, McCaughey ET. Solitary fibrous tumors ("fibrous mesotheliomas") of the peritoneum: report of three cases and a review of the literature. *Arch Pathol Lab Med*. 1990;114:493-495.