Protocol for the Examination of Specimens From Patients With Carcinoma of the Prostate Gland

Protocol applies to invasive carcinomas of the prostate gland.

Based on AJCC/UICC TNM, 7th edition
Protocol web posting date: June 2012

Procedures
• Needle Biopsy
• Transurethral Prostatic Resection
• Suprapubic or Retropubic Enucleation (Subtotal Prostatectomy)
• Radical Prostatectomy

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CAP Prostate Protocol Revision History

Version Code
The definition of the version code can be found at www.cap.org/cancerprotocols.

Version: Prostate 3.2.0.0

Summary of Changes
The following changes have been made since the February 2011 release.

Transurethral Prostatic Resection (TUR), Enucleation Specimen

Tumor Quantitation: TUR Specimens
Deleted the following data elements:
___ Tumor incidental histologic finding in no more than 5% of tissue resected with Gleason score 2 to 6 (cT1a)
___ Tumor incidental histologic finding in more than 5% of tissue resected or Gleason score 7 to 10 (cT1b)

Radical Prostatectomy

Seminal Vesicle Invasion
Optional elements “Right,” “Left,” and “Bilateral” were added, as follows:

Seminal Vesicle Invasion (invasion of muscular wall required) (select all that apply)
___ Not identified
___ Present
       + ___ Right
       + ___ Left
       + ___ Bilateral
___ No seminal vesicle present

Explanatory Notes

B. Gleason Score
The phrase “and radiation therapy” was added to the first sentence.

C. Quantitation of Tumor
The fifth sentence was changed, beginning with “The designation of the proportion (percentage)...”

K. TNM and Stage Groupings
Regional and Distant Lymph Nodes
This section was added.
Surgical Pathology Cancer Case Summary

Protocol web posting date: June 2012

PROSTATE GLAND: Needle Biopsy

Select a single response unless otherwise indicated.

The Gleason grade and score and tumor extent measures should be documented for each positive specimen (container). The essential information in each specimen could be conveyed with a simple diagnostic line such as, “Adenocarcinoma, Gleason grade 3 + 4 = score of 7, in 1 of 2 cores, involving 20% of needle core tissue, and measuring 4 mm in length.” (See “Explanatory Notes.”)

Histologic Type (Note A)
___ Adenocarcinoma (acinar, not otherwise specified)
___ Other (specify): __________________________

Histologic Grade (Note B)
Gleason Pattern
(If 3 patterns present, use most predominant pattern and worst pattern of remaining 2)

___ Not applicable
___ Cannot be determined

Primary (Predominant) Pattern
___ Grade 1
___ Grade 2
___ Grade 3
___ Grade 4
___ Grade 5

Secondary (Worst Remaining) Pattern
___ Grade 1
___ Grade 2
___ Grade 3
___ Grade 4
___ Grade 5

Total Gleason Score: ___

Tumor Quantitation (Note C)
Number cores positive: ___
Total number of cores: ___
and
Proportion (percent) of prostatic tissue involved by tumor: ___%

or

+ Data elements preceded by this symbol are not required. However, these elements may be clinically important but are not yet validated or regularly used in patient management.
Number cores positive: ____
Total number of cores: ____
and
Total linear millimeters of carcinoma: ___ mm
Total linear millimeters of needle core tissue: ___ mm

or

Number cores positive: ____
Total number of cores: ____
and
Proportion (percent) of prostatic tissue involved by tumor: ____% 
and
Total linear millimeters of carcinoma: ___ mm
Total linear millimeters of needle core tissue: ___ mm

+ Proportion (percentage) of prostatic tissue involved by tumor for core with the greatest amount of tumor: ____% 

Periprostatic Fat Invasion (document if identified) (Note D)
+ ___ Not identified
___ Present

Seminal Vesicle Invasion (document if identified) (Note D)
+ ___ Not identified
___ Present

+ Lymph-Vascular Invasion
+ ___ Not identified
+ ___ Present
+ ___ Indeterminate

+ Perineural Invasion (Note E)
+ ___ Not identified
+ ___ Present

+ Additional Pathologic Findings (select all that apply)
+ ___ None identified
+ ___ High-grade prostatic intraepithelial neoplasia (PIN) (Note F)
+ ___ Atypical adenomatous hyperplasia (adenosis)
+ ___ Inflammation (specify type): ___________________________
+ ___ Other (specify): ___________________________

+ Comment(s)

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+ Data elements preceded by this symbol are not required. However, these elements may be clinically important but are not yet validated or regularly used in patient management.
Surgical Pathology Cancer Case Summary

Protocol web posting date: June 2012

PROSTATE GLAND: Transurethral Prostatic Resection (TUR), Enucleation Specimen (Subtotal Prostatectomy)

Select a single response unless otherwise indicated.

Procedure
___ Transurethral prostatic resection (Note G)
___ Enucleation
___ Other (specify): _____________________________
___ Not specified

Specimen Size
Weight: ___ g
Size (enucleation specimens only): ___ x ___ x ___ cm

Histologic Type (Note A)
___ Adenocarcinoma (acinar, not otherwise specified)
___ Other (specify): _____________________________

Histologic Grade (Note B)
Gleason Pattern
(If 3 patterns present, use most predominant pattern and worst pattern of remaining 2)
___ Not applicable
___ Cannot be determined

Primary (Predominant) Pattern
___ Grade 1
___ Grade 2
___ Grade 3
___ Grade 4
___ Grade 5

Secondary (Worst Remaining) Pattern
___ Grade 1
___ Grade 2
___ Grade 3
___ Grade 4
___ Grade 5

Total Gleason Score: __

Tumor Quantitation: TUR Specimens (Note C)
Proportion (percentage) of prostatic tissue involved by tumor: ____%
+ Number of positive chips: ____
+ Total number of chips: ____

+ Data elements preceded by this symbol are not required. However, these elements may be clinically important but are not yet validated or regularly used in patient management.
Tumor Quantitation: Enucleation Specimens (Note C)
Proportion (percent) of prostatic tissue involved by tumor: ____%
+ Tumor size (dominant nodule, if present):
  + Greatest dimension: ___ cm
  + Additional dimensions: ___ x ___ cm

Periprostatic Fat Invasion (document if identified) (Note D)
+ ___ Not identified
  ___ Present

Seminal Vesicle Invasion (document if identified) (Note D)
+ ___ Not identified
  ___ Present

+ Lymph-Vascular Invasion
  + ___ Not identified
  + ___ Present
  + ___ Indeterminate

+ Perineural Invasion (Note E)
  + ___ Not identified
  + ___ Present

+ Additional Pathologic Findings (select all that apply)
  + ___ None identified
  + ___ High-grade prostatic intraepithelial neoplasia (PIN) (Note F)
  + ___ Atypical adenomatous hyperplasia (adenosis)
  + ___ Nodular prostatic hyperplasia
  + ___ Inflammation (specify type): ___________________________
  + ___ Other (specify): ________________________

+ Comment(s)
Surgical Pathology Cancer Case Summary

Protocol web posting date: June 2012

PROSTATE GLAND: Radical Prostatectomy

Select a single response unless otherwise indicated.

Procedure (Note G)
___ Radical prostatectomy
___ Other (specify): __________________________
___ Not specified

Prostate Size (Note G)
Weight: ___ g
Size: ___ x ___ x ___ cm

Lymph Node Sampling (Note G)
___ No lymph nodes present
___ Pelvic lymph node dissection

Histologic Type (Note A)
___ Adenocarcinoma (acinar, not otherwise specified)
___ Prostatic duct adenocarcinoma
___ Mucinous (colloid) adenocarcinoma
___ Signet-ring cell carcinoma
___ Adenosquamous carcinoma
___ Small cell carcinoma
___ Sarcomatoid carcinoma
___ Undifferentiated carcinoma, not otherwise specified
___ Other (specify): __________________________

Histologic Grade (Note B)
Gleason Pattern
If 3 patterns are present, record the most predominant and second most common patterns; the tertiary pattern should be recorded if higher than the primary and secondary patterns but it is not incorporated into the Gleason score.

___ Not applicable
___ Cannot be determined

Primary Pattern
___ Grade 1
___ Grade 2
___ Grade 3
___ Grade 4
___ Grade 5

+ Data elements preceded by this symbol are not required. However, these elements may be clinically important but are not yet validated or regularly used in patient management.
Secondary Pattern
___ Grade 1
___ Grade 2
___ Grade 3
___ Grade 4
___ Grade 5

Tertiary Pattern
___ Grade 3
___ Grade 4
___ Grade 5
___ Not applicable

Total Gleason Score: ___

**Tumor Quantitation (Note C)**
Proportion (percentage) of prostate involved by tumor: ___% and/or
Tumor size (dominant nodule, if present):
  Greatest dimension: ___ mm
  + Additional dimensions: ___ x ___ mm

**Extraprostatic Extension (select all that apply) (Note H)**
___ Not identified
___ Present
    ___ Focal
      + Specify site(s): _______________________
    ___ Nonfocal (established, extensive)
      + Specify site(s): _______________________
    ___ Indeterminate

**Seminal Vesicle Invasion (invasion of muscular wall required) (select all that apply) (Note D)**
___ Not identified
___ Present
    + ___ Right
    + ___ Left
    + ___ Bilateral
___ No seminal vesicle present
Margins (select all that apply) (Note I)
___ Cannot be assessed
+ ___ Benign glands at surgical margin
___ Margins uninvolved by invasive carcinoma
___ Margin(s) involved by invasive carcinoma
   + ___ Unifocal
   + ___ Multifocal
___ Apical
___ Bladder neck
___ Anterior
___ Lateral
___ Postero-lateral (neurovascular bundle)
___ Posterior
___ Other(s) (specify): __________________________

Treatment Effect on Carcinoma (select all that apply)
___ Not identified
___ Radiation therapy effect present
___ Hormonal therapy effect present
___ Other therapy effect(s) present (specify): ________________

Lymph-Vascular Invasion
___ Not identified
___ Present
___ Indeterminate

+ Perineural Invasion (Note E)
+ ___ Not identified
+ ___ Present

Pathologic Staging (pTNM) (Note K)

TNM Descriptors (required only if applicable) (select all that apply)
___ m (multiple)
___ r (recurrent)
___ y (posttreatment)

Primary Tumor (pT)
___ Not identified
___ pT2: Organ confined
   + ___ pT2a: Unilateral, involving one-half of 1 side or less
   + ___ pT2b: Unilateral, involving more than one-half of 1 side but not both sides
   + ___ pT2c: Bilateral disease
pT3: Extraprostatic extension
___ pT3a: Extraprostatic extension or microscopic invasion of bladder neck
___ pT3b: Seminal vesicle invasion
___ pT4: Invasion of rectum, levator muscles and/or pelvic wall (Note J)

Note: There is no pathologic T1 classification. Subdivision of pT2 disease is problematic and has not proven to be of prognostic significance.

+ Data elements preceded by this symbol are not required. However, these elements may be clinically important but are not yet validated or regularly used in patient management.
Regional Lymph Nodes (pN)
___ pNX: Cannot be assessed
___ pN0: No regional lymph node metastasis
___ pN1: Metastasis in regional lymph node or nodes
___ No nodes submitted or found

Number of Lymph Nodes Examined
Specify: ____
___ Number cannot be determined (explain): ______________________

Number of Lymph Nodes Involved
Specify: ____
___ Number cannot be determined (explain): ______________________

Diameter of largest lymph node metastasis: ____ (mm)

Distant Metastasis (pM)
___ Not applicable
___ pM1: Distant metastasis
___ pM1a: Nonregional lymph node(s)
___ pM1b: Bone(s)
___ pM1c: Other site(s) with or without bone disease

Note: When more than 1 site of metastasis is present, the most advanced category is used. pM1c is most advanced.

+ Additional Pathologic Findings (select all that apply)
+ ___ None identified
+ ___ High-grade prostatic intraepithelial neoplasia (PIN) (Note F)
+ ___ Inflammation (specify type): ______________________
+ ___ Atypical adenomatous hyperplasia (adenosis)
+ ___ Nodular prostatic hyperplasia
+ ___ Other (specify): ______________________

+ Ancillary Studies
+ Specify: ______________________
+ ___ Not performed

+ Comment(s)

+ Data elements preceded by this symbol are not required. However, these elements may be clinically important but are not yet validated or regularly used in patient management.
Explanatory Notes

A. Histologic Type
This protocol applies only to carcinomas of the prostate gland. The histologic classification of prostate carcinoma is recommended and shown below.\textsuperscript{1} However, this protocol does not preclude the use of other systems of classification or histologic types. Mixtures of different histologic types should be indicated.

Histologic Classification of Carcinoma of the Prostate

Adenocarcinoma (conventional, acinar)
Special variants of adenocarcinoma and other carcinomas
- Prostatic duct adenocarcinoma
- Mucinous (colloid) adenocarcinoma
- Signet-ring cell carcinoma
- Adenosquamous carcinoma
- Squamous cell carcinoma\textsuperscript{#}
- Basaloid (basal cell) and adenoid cystic carcinoma\textsuperscript{#}
- Urothelial (transitional cell) carcinoma\textsuperscript{#}
- Small cell carcinoma
- Sarcomatoid carcinoma
- Lymphoepithelioma-like carcinoma\textsuperscript{#}
- Undifferentiated carcinoma, not otherwise specified

\textsuperscript{#} This protocol does not apply to these carcinomas.

B. Gleason Score
The Gleason grading system is recommended for use in all prostatic specimens containing adenocarcinoma, with the exception of those showing treatment effects, usually in the setting of androgen withdrawal and radiation therapy.\textsuperscript{2,3} Gleason score is an important parameter used in nomograms, such as the Kattan nomograms,\textsuperscript{4,5} and the Partin tables,\textsuperscript{6} which guide individual treatment decisions. Readers are referred to the recommendations of a recent consensus conference dealing with the contemporary usage of the Gleason system.\textsuperscript{7} The Gleason score is the sum of the primary (most predominant in terms of surface area of involvement) Gleason grade and the secondary (second most predominant) Gleason grade. Where no secondary Gleason grade exists, the primary Gleason grade is doubled to arrive at a Gleason score. The primary and secondary grades should be reported in addition to the Gleason score, that is, Gleason score 7(3+4) or 7(3+4).

In needle biopsy specimens, it is recommended that Gleason scores be assigned for each specimen (container). Alternatively, a Gleason score may be given for each positive intact core in a container.

In needle biopsy specimens where there is a minor secondary component (<5\% of tumor) and where the secondary component is of higher grade, the latter should be reported. For instance, a case showing more than 95\% Gleason 3 and less than 5\% Gleason 4 should be reported as Gleason score 7(3+4). Conversely, if a minor secondary pattern is of lower grade, it need not be reported. For instance, where there is greater than 95\% Gleason score 4 and less than 5\% Gleason 3, the score should be reported as Gleason 8(4+4).

In needle biopsy specimens where more than 2 patterns are present, and the worst grade is neither the predominant nor the secondary grade, the predominant and highest grade should be chosen to arrive at a score (e.g., 75\%, grade 3; 20\%–25\%, grade 4; <5\%, grade 5 is scored as 3+5=8). This approach has been validated in a large clinical series.\textsuperscript{8}
Rules of grading similar to the above apply to transurethral resection and enucleation (simple prostatectomy) specimens.

Tertiary Gleason patterns are common in radical prostatectomy specimens. When Gleason pattern 5 is present as a tertiary pattern, its presence should be recognized in the report. For instance, in a situation where the primary Gleason grade is 3, the secondary is 4 and there is less than 5% Gleason 5, the report should indicate a Gleason score of 7(3+4) with tertiary Gleason pattern 5.

For radical prostatectomy specimens, Gleason score should be assigned to the dominant nodule(s), if present. Where more than one separate tumor is clearly identified, the Gleason scores of individual tumors can be recorded separately, or, at the very least, a Gleason score of the dominant or most significant lesion should be recorded. For instance, if there is a large Gleason score 4(2+2) transition zone tumor and a separate smaller Gleason score 8(4+4) peripheral zone cancer, both scores should be reported, or, at the very least, the latter score should be reported rather than these scores being averaged.

C. Quantitation of Tumor

There are many methods of estimating the amount of tumor in prostatic specimens. For needle core biopsy specimens, it is suggested that the number of positive cores out of the total number of cores always be reported, except in situations where fragmentation precludes accurate counting. The estimated proportion (percent) of prostatic tissue involved by tumor and/or the linear millimeters of the tumor should also be reported. Reporting of the positive core with the greatest percentage of tumor is an option. The designation of the proportion (percentage) of prostatic tissue in transurethral samples is important. When prostate cancer is discovered incidentally (i.e., discovered in specimens submitted for clinically benign disease, usually BPH), the percentage involvement is used to determine the clinical T1 substage, with ≤5% involvement being T1a and >5% being T1b. The Gleason score may also play a factor in the substage. In subtotal and radical prostatectomy specimens, the percentage of tissue involved by tumor can also be “eyeballed” by simple visual inspection. Additionally, in these latter specimens, it may be possible to measure a dominant tumor nodule in at least 2 dimensions and/or to indicate the number of blocks involved by tumor out of the total number of prostatic blocks submitted.

D. Local Invasion in Needle Biopsies

Occasionally in needle biopsies, periprostatic fat is present and involved by tumor. This observation should be noted since it indicates that the tumor is at least pT3a in the TNM system. Furthermore, if seminal vesicle tissue is present (either unintentionally or intentionally, as in a directed biopsy) and involved by tumor, this should be reported since it indicates that the tumor is at least pT3b. Seminal vesicle invasion is defined by involvement of the muscular wall. At times, especially in needle biopsy specimens, it is difficult to distinguish between seminal vesicle and ejaculatory duct tissue. It is important not to overinterpret the ejaculatory duct as seminal vesicle since involvement of the former by tumor does not constitute pT3b disease. If there is doubt as to whether the involved tissue represents the seminal vesicle or the ejaculatory duct, then invasion of the seminal vesicle should not be definitively diagnosed.

E. Perineural Invasion

Perineural invasion in core needle biopsies has been associated with extraprostatic extension in some correlative radical prostatectomy studies, although its exact prognostic significance remains to be determined. Perineural invasion has also been found to be an independent risk factor, in some studies, for predicting an adverse outcome in patients treated with external beam radiation, but not for patients treated with brachytherapy or radical prostatectomy. The value of perineural invasion as an independent prognostic factor has been questioned in a multivariate analysis.
F. Prostatic Intraepithelial Neoplasia
The diagnostic term prostatic intraepithelial neoplasia (PIN), unless qualified, refers to high-grade PIN. Low-grade PIN is not reported. The presence of an isolated PIN (PIN in the absence of carcinoma) should be reported in all biopsy specimens.9 The reporting of PIN in biopsies with carcinoma is considered optional. High-grade PIN in a biopsy without evidence of carcinoma has in the past been a risk factor for the presence of carcinoma on subsequent biopsies, but the magnitude of the risk has diminished, and, in some studies, high-grade PIN was not a risk factor at all, unless multiple cores were positive for PIN.23-26 The reporting of high-grade PIN in prostatectomy specimens is optional.

G. Submission of Tissue for Microscopic Evaluation in Transurethral Resection and Radical Prostatectomy Specimens
Transurethral resection specimens that weigh 12 g or less should be submitted in their entirety, usually in 6 to 8 cassettes.26,27 For specimens that weigh more than 12 g, the initial 12 g are submitted (6 to 8 cassettes), and 1 cassette may be submitted for every additional 5 g may be submitted.

In general, random chips are submitted; however, if some chips are firmer or have a yellow or orange-yellow appearance, they should be submitted preferentially.

If an unsuspected carcinoma is found in tissue submitted, and it involves 5% or less of the tissue examined, the remaining tissue may be submitted for microscopic examination, especially in younger patients.

A radical prostatectomy specimen may be submitted in its entirety or partially sampled in a systematic fashion.28,29 For partial sampling in the setting of a grossly visible tumor, the tumor and associated periprostatic tissue and margins, along with the entire apical and bladder neck margins and the junction of each seminal vesicle with prostate proper, should be submitted. If there is no grossly visible tumor, a number of systematic sampling strategies may be used. One that yields excellent prognostic information involves submitting the posterior aspect of each transverse slice along with a mid anterior block from each side.29 The anterior sampling detects the T1c cases arising in the transition zone and extending anteriorly. The entire apical and bladder neck margins and the junction of each seminal vesicle with the prostate should also be submitted.

H. Extraprostatic Extension
Extraprostatic extension (EPE) is the preferred term for the presence of tumor beyond the confines of the prostate gland.28,30-32 Tumor admixed with fat constitutes extraprostatic extension. Tumor involving loose connective tissue in the plane of fat or beyond, even in the absence of direct contact between the tumor and the adipocytes, indicates EPE. Extraprostatic extension may also be reported when the tumor involves perineural spaces in the neurovascular bundles, even in the absence of periprostatic fat involvement. In certain locations, such as the anterior and apical prostate and bladder neck regions, there is a paucity of fat, and in these locations EPE is determined when the tumor extends beyond the confines of the normal glandular prostate. In the distal apical perpendicular margin section, it is often difficult to identify EPE. Sometimes there is a distinct bulging tumor nodule, which may be associated with a desmoplastic stromal reaction. The specific location(s) and the number of sites (blocks) of EPE are useful to report. Descriptors of EPE (focal versus nonfocal) should be used. Focal EPE equates with only a few neoplastic glands being outside the prostate or a tumor involving less than 1 high-power field in 1 or 2 sections;30 nonfocal (established) EPE is more extensively spread beyond the prostatic edge.

I. Margins
The entire surface of the prostate should be inked to evaluate the surgical margins.28-36 Usually, surgical margins should be designated as “negative” if tumor is not present at the inked margin and as “positive” if tumor cells touch the ink at the margin. When tumor is located very close to an inked surface but is not actually in contact with the ink, the margin is considered negative. Positive surgical
margins should not be interpreted as extraprostatic extension. Intraprostatic margins are positive in the setting of intraprostatic incision (so-called pT2+ disease; Figure 1).\textsuperscript{28} If the surgical margin finding is positive, the pathologist should state that explicitly, although this finding is not relied upon for pathologic staging. The specific locations of the positive margins should be reported, and it should be specified whether EPE or intraprostatic incision is present at each site of margin positivity. There should be some indication of the extent of margin positivity. At the 2009 International Society of Urological Pathology Consensus Conference on Handling and Staging of Radical Prostatectomy Specimens, it was recommended that the extent of a positive margin should be reported as millimeters of involvement.

\textbf{Figure 1.} Surgical incision can create stage pT2+ from either pT2 or pT3 disease.

\textbf{J. Apex and Bladder Neck}

The apex should be carefully examined because it is a common site of margin positivity.\textsuperscript{28-31} At the apex, tumor admixed with skeletal muscle elements does not constitute extraprostatic extension. The apical and bladder neck surgical margins should be submitted entirely, preferably with a perpendicular sectioning technique. Microscopic involvement of bladder neck muscle fibers in radical prostatectomy specimens indicates pT3a disease.\textsuperscript{37}

\textbf{K. TNM and Stage Groupings}

The protocol recommends the use of the TNM Staging System for carcinoma of the prostate of the American Joint Committee on Cancer (AJCC) and the International Union Against Cancer (UICC).\textsuperscript{38}

By AJCC/UICC convention, the designation “T” refers to a primary tumor that has not been previously treated. The symbol “p” refers to the pathologic classification of the TNM, as opposed to the clinical classification, and is based on gross and microscopic examination. pT entails a resection of the primary tumor or biopsy adequate to evaluate the highest pT category, pN entails removal of nodes adequate to validate lymph node metastasis, and pM implies microscopic examination of distant lesions. Clinical classification (cTNM) is usually carried out by the referring physician before treatment during initial evaluation of the patient or when pathologic classification is not possible.

Pathologic staging is usually performed after surgical resection of the primary tumor. Pathologic staging depends on pathologic documentation of the anatomic extent of disease, whether or not the primary tumor has been completely removed. If a biopsied tumor is not resected for any reason (eg, when technically unfeasible), and if the highest T and N categories or the M1 category of the tumor can be
confirmed microscopically, the criteria for pathologic classification and staging have been satisfied without total removal of the primary cancer. pT2, pT3a, and pT3b are illustrated in Figures 2 through 5.\textsuperscript{39}

**Figure 2.** T2a (left) shows tumor involving one-half of one lobe (side) or less whereas T2b (right) shows tumor involving more than one-half of one lobe but not both lobes. Used with permission of the American Joint Committee on Cancer (AJCC), Chicago, Ill. The original source for this material is the *AJCC Cancer Staging Atlas* (2006) edited by Greene et al\textsuperscript{39} and published by Springer Science and Business Media, LLC, www.springerlink.com.

**Figure 3.** T2c tumor involving both lobes (sides). Used with permission of the American Joint Committee on Cancer (AJCC), Chicago, Ill. The original source for this material is the *AJCC Cancer Staging Atlas* (2006) edited by Greene et al\textsuperscript{39} and published by Springer Science and Business Media, LLC, www.springerlink.com.
Figure 4. T3a is defined as a tumor with unilateral extraprostatic extension, as shown in A, or with bilateral extension, as shown in B. Microscopic extension into the bladder neck is also pT3a. Used with permission of the American Joint Committee on Cancer (AJCC), Chicago, Ill. The original source for this material is the AJCC Cancer Staging Atlas (2006) edited by Greene et al39 and published by Springer Science and Business Media, LLC, www.springerlink.com.

Figure 5. T3b tumor invading the seminal vesicle. Used with permission of the American Joint Committee on Cancer (AJCC), Chicago, Ill. The original source for this material is the AJCC Cancer Staging Atlas (2006) edited by Greene et al39 and published by Springer Science and Business Media, LLC, www.springerlink.com.

Regional and Distant Lymph Nodes
Regional Lymph Nodes
The regional lymph nodes are the nodes of the true pelvis, which essentially are the pelvic nodes below the bifurcation of the common iliac arteries. They include the following groups:
- Pelvic, NOS
- Hypogastric
- Obturator
- Iliac (internal, external, or NOS)
- Sacral (lateral, presacral, promontory [Gerota’s], or NOS)

Laterality does not affect the N classification.

Distant Lymph Nodes
Distant lymph nodes lie outside the confines of the true pelvis. They can be imaged using ultrasound, computed tomography, magnetic resonance imaging, or lymphangiography. Although enlarged lymph
nodes can occasionally be visualized on radiographic imaging, fewer patients are initially discovered with clinically evident metastatic disease. In lower risk patients, imaging tests have proven unhelpful. In lieu of imaging, risk tables are many times used to determine individual patient risk of nodal involvement prior to therapy. Involvement of distant lymph nodes is classified as M1a. The distant lymph nodes include the following:

- Aortic (paraaortic lumbar)
- Common iliac
- Inguinal, deep
- Superficial inguinal (femoral)
- Supraclavicular
- Cervical
- Scalene
- Retroperitoneal, NOS

**Primary Tumor (T): Clinical Classification**

<table>
<thead>
<tr>
<th>T-stage</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>T0</td>
<td>No evidence of primary tumor</td>
</tr>
<tr>
<td>T1</td>
<td>Clinically inapparent tumor neither palpable nor visible by imaging</td>
</tr>
<tr>
<td>T1a</td>
<td>Tumor incidental histologic finding in 5% or less of tissue resected</td>
</tr>
<tr>
<td>T1b</td>
<td>Tumor incidental histologic finding in more than 5% of tissue resected</td>
</tr>
<tr>
<td>T1c</td>
<td>Tumor identified by needle biopsy (eg, because of elevated prostate specific antigen [PSA])</td>
</tr>
<tr>
<td>T2</td>
<td>Tumor confined within prostate*</td>
</tr>
<tr>
<td>T2a</td>
<td>Tumor involves one-half of one lobe or less</td>
</tr>
<tr>
<td>T2b</td>
<td>Tumor involves more than one-half of one lobe but not both lobes</td>
</tr>
<tr>
<td>T2c</td>
<td>Tumor involves both lobes</td>
</tr>
<tr>
<td>T3</td>
<td>Tumor extends through the prostate capsule##</td>
</tr>
<tr>
<td>T3a</td>
<td>Extracapsular extension (unilateral or bilateral) including microscopic bladder neck involvement</td>
</tr>
<tr>
<td>T3b</td>
<td>Tumor invades seminal vesicle(s)</td>
</tr>
<tr>
<td>T4</td>
<td>Tumor is fixed or invades adjacent structures other than seminal vesicles: bladder neck, external sphincter, rectum, levator muscles, and/or pelvic wall</td>
</tr>
</tbody>
</table>

* Tumor found in one or both lobes by needle biopsy, but not palpable or reliably visible by imaging, is classified as T1c.

## Invasion into the prostatic apex or into (but not beyond) the prostatic capsule is classified not as T3 but as T2.
The 2009 Anatomic Stage/Prognostic Groups incorporate serum PSA level and Gleason score:

Anatomic Stage / Prognostic Groups

<table>
<thead>
<tr>
<th>Group</th>
<th>T</th>
<th>N</th>
<th>M</th>
<th>PSA</th>
<th>Gleason</th>
</tr>
</thead>
<tbody>
<tr>
<td>I</td>
<td>T1a–c</td>
<td>N0</td>
<td>M0</td>
<td>PSA &lt;10</td>
<td>Gleason ≤6</td>
</tr>
<tr>
<td></td>
<td>T2a</td>
<td>N0</td>
<td>M0</td>
<td>PSA &lt;10</td>
<td>Gleason ≤6</td>
</tr>
<tr>
<td></td>
<td>T1 – 2a</td>
<td>N0</td>
<td>M0</td>
<td>PSA X</td>
<td>Gleason X</td>
</tr>
<tr>
<td>II A</td>
<td>T1 a–c</td>
<td>N0</td>
<td>M0</td>
<td>PSA &lt;20</td>
<td>Gleason 7</td>
</tr>
<tr>
<td></td>
<td>T1 a–c</td>
<td>N0</td>
<td>M0</td>
<td>PSA ≥10 &lt;20</td>
<td>Gleason ≤6</td>
</tr>
<tr>
<td></td>
<td>T2a</td>
<td>N0</td>
<td>M0</td>
<td>PSA ≥10 &lt;20</td>
<td>Gleason ≤6</td>
</tr>
<tr>
<td></td>
<td>T2a</td>
<td>N0</td>
<td>M0</td>
<td>PSA &lt;20</td>
<td>Gleason 7</td>
</tr>
<tr>
<td></td>
<td>T2b</td>
<td>N0</td>
<td>M0</td>
<td>PSA &lt;20</td>
<td>Gleason ≤7</td>
</tr>
<tr>
<td></td>
<td>T2b</td>
<td>N0</td>
<td>M0</td>
<td>PSA X</td>
<td>Gleason X</td>
</tr>
<tr>
<td>II B</td>
<td>T2c</td>
<td>N0</td>
<td>M0</td>
<td>Any PSA</td>
<td>Any Gleason</td>
</tr>
<tr>
<td></td>
<td>T1 – 2</td>
<td>N0</td>
<td>M0</td>
<td>PSA ≥20</td>
<td>Any Gleason</td>
</tr>
<tr>
<td></td>
<td>T1 – 2</td>
<td>N0</td>
<td>M0</td>
<td>Any PSA</td>
<td>Gleason ≥8</td>
</tr>
<tr>
<td>III</td>
<td>T3 a–c</td>
<td>N0</td>
<td>M0</td>
<td>Any PSA</td>
<td>Any Gleason</td>
</tr>
<tr>
<td>IV</td>
<td>T4</td>
<td>N0</td>
<td>M0</td>
<td>Any PSA</td>
<td>Any Gleason</td>
</tr>
<tr>
<td></td>
<td>Any T</td>
<td>N1</td>
<td>M0</td>
<td>Any PSA</td>
<td>Any Gleason</td>
</tr>
<tr>
<td></td>
<td>Any T</td>
<td>Any N</td>
<td>M1</td>
<td>Any PSA</td>
<td>Any Gleason</td>
</tr>
</tbody>
</table>

Note: When either prostate specific antigen (PSA) or Gleason is not available, grouping should be determined by T stage and/or whichever of either the PSA or Gleason is available.

TNM Descriptors
For identification of special cases of TNM or pTNM classifications, the “m” suffix and the “y,” “r,” and “a” prefixes are used. Although they do not affect the stage grouping, they indicate cases needing separate analysis.

The “m” suffix indicates the presence of multiple primary tumors in a single site and is recorded in parentheses: pT(m)NM.

The “y” prefix indicates those cases in which classification is performed during or following initial multimodality therapy (ie, neoadjuvant chemotherapy, radiation therapy, or both chemotherapy and radiation therapy). The cTNM or pTNM category is identified by a “y” prefix. The ycTNM or ypTNM categorizes the extent of tumor actually present at the time of that examination. The “y” categorization is not an estimate of tumor prior to multimodality therapy (ie, before initiation of neoadjuvant therapy).

The “r” prefix indicates a recurrent tumor when staged after a documented disease-free interval, and is identified by the “r” prefix: rTNM.

The “a” prefix designates the stage determined at autopsy: aTNM.
**Additional Descriptors**

**Residual Tumor (R)**
Tumor remaining in a patient after therapy with curative intent (e.g., surgical resection for cure) is categorized by a system known as R classification, shown below.

<table>
<thead>
<tr>
<th>Code</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>RX</td>
<td>Presence of residual tumor cannot be assessed</td>
</tr>
<tr>
<td>R0</td>
<td>No residual tumor</td>
</tr>
<tr>
<td>R1</td>
<td>Microscopic residual tumor</td>
</tr>
<tr>
<td>R2</td>
<td>Macroscopic residual tumor</td>
</tr>
</tbody>
</table>

For the surgeon, the R classification may be useful to indicate the known or assumed status of the completeness of a surgical excision. For the pathologist, the R classification is relevant to the status of the margins of a surgical resection specimen. That is, tumor involving the resection margin on pathologic examination may be assumed to correspond to residual tumor in the patient and may be classified as macroscopic or microscopic according to the findings at the specimen margin(s).

**Lymph-Vascular Invasion**
Lymph-vascular invasion (LVI) indicates whether microscopic lymph-vascular invasion is identified. LVI includes lymphatic invasion, vascular invasion, or lymph-vascular invasion. By AJCC/UICC convention, LVI does not affect the T category indicating local extent of tumor unless specifically included in the definition of a T category.

**References**


27. Sehdev AE, Pan CC, Epstein JI. Comparative analysis of sampling methods for grossing radical prostatectomy specimens performed for nonpalpable (stage T1c) prostatic adenocarcinoma. *Hum Pathol*. 2001;32:494-499.