

# Merkel Cell Carcinoma

*(Staging for Merkel Cell of the eyelid [C44.1] is not included in this chapter – see Chap. 48, “Carcinoma of the Eyelid”)*

## At-A-Glance

### SUMMARY OF CHANGES

- This is the first staging chapter specific for Merkel cell carcinoma. Merkel cell carcinoma was previously included in the “Carcinoma of the Skin” chapter

### ANATOMIC STAGE/PROGNOSTIC GROUPS

Patients with primary Merkel cell carcinoma with no evidence of regional or distant metastases (either clinically or pathologically) are divided into two stages: Stage I for primary tumors ≤2 cm in size and Stage II for primary tumors >2 cm in size. Stages I and II are further divided into A and B substages based on method of nodal evaluation. Patients who have pathologically proven node negative disease (by microscopic evaluation of their draining lymph nodes) have improved survival (substaged as A) compared with those who are only evaluated clinically (substaged as B). Stage II has an additional substage (IIC) for tumors with extracutaneous invasion (T4) and negative node status regardless of whether the negative node status was established microscopically or clinically. Stage III is also divided into A and B categories for patients with microscopically positive and clinically occult nodes (IIIA) and macroscopic nodes (IIIB). There are no subgroups of Stage IV Merkel cell carcinoma.

Stage 0	Tis	N0	M0
Stage IA	T1	pN0	M0
Stage IB	T1	cN0	M0
Stage IIA	T2/T3	pN0	M0
Stage IIB	T2/T3	cN0	M0
Stage IIC	T4	N0	M0
Stage IIIA	Any T	N1a	M0
Stage IIIB	Any T	N1b/N2	M0
Stage IV	Any T	Any N	M1

### ICD-O-3 TOPOGRAPHY CODES

C44.0	Skin of lip, NOS
C44.2	External ear
C44.3	Skin of other and unspecified parts of face
C44.4	Skin of scalp and neck
C44.5	Skin of trunk
C44.6	Skin of upper limb and shoulder
C44.7	Skin of lower limb and hip
C44.8	Overlapping lesion of skin
C44.9	Skin, NOS
C51.0	Labium majus
C51.1	Labium minus
C51.2	Clitoris
C51.8	Overlapping lesion of vulva
C51.9	Vulva, NOS
C60.0	Prepuce
C60.1	Glans penis
C60.2	Body of penis
C60.8	Overlapping lesion of penis
C60.9	Penis, NOS
C63.2	Scrotum, NOS

### ICD-O-3 HISTOLOGY CODE RANGES

8247

## INTRODUCTION

Merkel cell carcinoma (MCC) is a relatively rare, potentially aggressive primary cutaneous neuroendocrine carcinoma, originally described by Tang and Toker in 1972 as trabecular carcinoma.<sup>1</sup> The mortality rate is twice that observed in melanoma (33% vs. 15%). Although the molecular pathogenesis remains largely unknown, ultraviolet radiation and immune suppression are likely significant predisposing factors. The identification of a novel polyomavirus termed *Merkel cell polyomavirus* in the majority of MCC tumors suggests a viral component in many cases.<sup>2</sup> Merkel cell carcinoma occurs most commonly on sun-exposed skin in fair-skinned individuals older than 50 years with a slight male predominance.<sup>3,4</sup> An increased incidence is also observed in patients with HIV infection, leukemias, and organ transplantation.<sup>4-6</sup> Merkel cell carcinoma is increasing in frequency, rising from 0.15 cases per 100,000 in 1986 to 0.44 cases per 100,000 in 2001. Much of this increase in reported frequency is likely due to increased recognition and improved techniques for diagnosis.<sup>7</sup> Currently in the United States, approximately 1,500 cases of MCC are diagnosed annually.<sup>8</sup> As the US population ages and improved transplantation regimens prolong the lives of organ transplant recipients, the incidence of MCC will likely continue to rise.

Merkel cell carcinoma has a nonspecific clinical presentation, though rapid growth of a firm, red to violaceous, nontender papule or nodule is often noted.<sup>4</sup> Diagnosis is made via biopsy, almost invariably with the aid of immunohistochemistry, classically demonstrating a peri-nuclear dot pattern of cytokeratin-20 staining. The majority of patients present with clinically localized disease. However, the disease can rapidly spread to regional and distant sites. The regional draining nodal basin is the most common site for recurrence.<sup>9</sup> The natural history of the disease is variable but heavily dependent on the stage at time of diagnosis.

Five different staging systems for Merkel cell carcinoma have been described in the literature and all are currently in use.<sup>10-14</sup> Depending on the system used, Stage III MCC could represent local, nodal, or metastatic disease. This situation impedes effective patient–physician communication, data comparison, and outcomes analysis. Therefore, development of a standardized, data-driven staging system is important for improving clinical care and research in this disease. Moreover, a separate staging system for MCC is appropriate given its unique behavior compared with other malignancies that will remain in the “Cutaneous Squamous Cell Carcinoma and other Cutaneous Carcinomas” staging chapter (see Chap. 29). This new staging system is based on an analysis of over 4,700 patients using the National Cancer Database as well as extensive review of the literature.

## ANATOMY

**Primary Sites.** Merkel cell carcinoma is postulated to arise from the Merkel cell, a neuroendocrine cell of the skin.<sup>1</sup> MCC

can occur anywhere on the skin but arises most often in sun-exposed areas. It occurs most commonly on the head and neck, followed by the extremities. In 14% of cases, the primary site remains unknown with MCC presentation in nodal or visceral sites.<sup>4</sup>

**Regional Lymph Nodes.** The draining regional lymph nodes are the most common site of metastasis. Regional lymph node metastasis occurs relatively frequently and early, even in the absence of deep local extension or large primary tumor size. Thirty-two percent of clinically negative draining lymph node basins were in fact positive for microscopic metastases as revealed by sentinel or elective lymphadenectomy.<sup>15</sup> Intralymphatic “in transit” regional metastases also occur but are uncommon. For MCC, an in transit metastasis is defined as a tumor distinct from the primary lesion and located either (1) between the primary lesion and the draining regional lymph nodes or (2) distal to the primary lesion. In contrast to melanoma, for MCC there is no separate subclassification of in transit metastases based on distance from the primary (i.e., no *satellite* metastasis classification). By convention, the term “regional nodal metastases” refers to disease confined to one nodal basin or two contiguous nodal basins, as in patients with nodal disease in combinations of femoral/iliac, axillary/supraclavicular, or cervical/supraclavicular metastases or in primary truncal disease with axillary/femoral, bilateral axillary, or bilateral femoral metastases.

**Metastatic Sites.** Merkel cell carcinoma can metastasize to virtually any organ site. Metastases occur most commonly to distant lymph nodes, followed by the liver, lung, bone, and brain.<sup>16</sup>

## RULES FOR CLASSIFICATION

The primary difference between the definitions of clinical and pathologic nodal staging is whether the regional lymph nodes were staged by clinical/radiologic exam only or by pathologic exam (after partial or complete lymphadenectomy).

**Clinical Staging.** Clinical staging is defined as regional lymph nodes that are staged by clinical inspection and palpation of the involved area and the regional lymph nodes and/or by radiologic studies. For cases without documentation of abnormal regional nodes on physical exam, patients should be considered to not have macroscopic nodal disease.

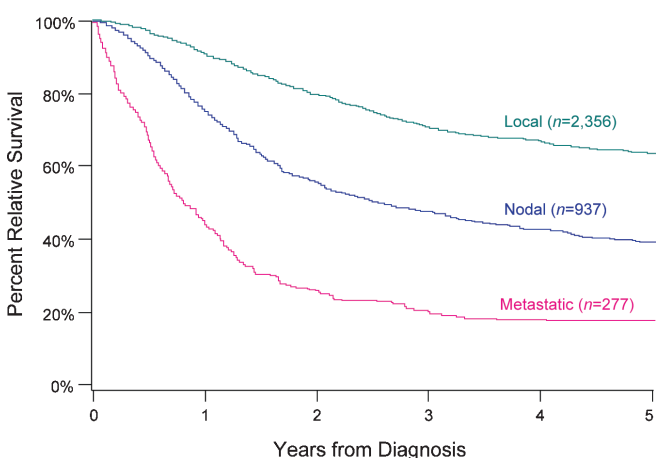
**Pathologic Staging.** Pathologic staging is defined as regional lymph nodes that are staged by focused (sentinel lymph node biopsy), therapeutic, or complete lymphadenectomy. With regard to Merkel cell carcinoma, the distinction between clinical vs. pathologic staging is highly significant. The natural history of MCC is variable and dependent on the pathologic stage at time of presentation.

Sentinel lymph node biopsy should be performed routinely on MCC patients, as approximately 32% of patients without palpable lymph nodes will have positive sentinel lymph node biopsies.<sup>15</sup> Pathologic staging with negative sentinel lymph node biopsy at time of diagnosis is a predictor of improved survival.<sup>12</sup> Despite these issues, approximately two-thirds of MCC patients captured in the National Cancer Database did not have pathologic staging of the regional nodes.

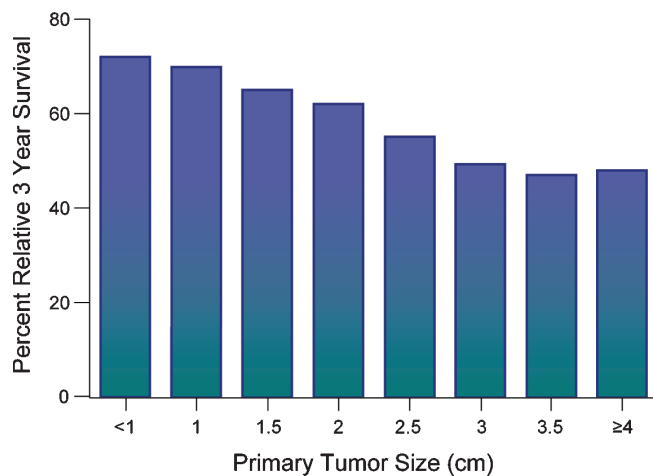
## PROGNOSTIC FEATURES AND SURVIVAL RESULTS

Survival in Merkel cell carcinoma is based on stage at presentation (Figure 30.1). Overall survival relative to an age- and sex-matched population was determined using 4,700 Merkel cell carcinoma patients in the National Cancer Database registry (manuscript in preparation). Tumor size is a continuous variable with increasing tumor size correlating with modestly poorer prognosis (Figure 30.2). True lymph node negativity by pathologic evaluation portends a better prognosis compared with patients whose lymph nodes are only evaluated by clinical or radiographic examination (Figure 30.3). This is in large part likely due to the high rate (33%) of false negative nodal determination by clinical exam alone.<sup>15</sup> Thus, patients should have pathologic evaluation of the draining nodal basin to most accurately predict survival and guide optimal therapy. Percent relative survival based on stage is shown in Figure 30.4.

Profound immune suppression, such as in HIV/AIDS, chronic lymphocytic leukemia, or solid organ transplantation have all been associated with worse survival in MCC.<sup>6,17</sup> Further, immunosuppressed patients frequently present with more advanced disease.<sup>4</sup>



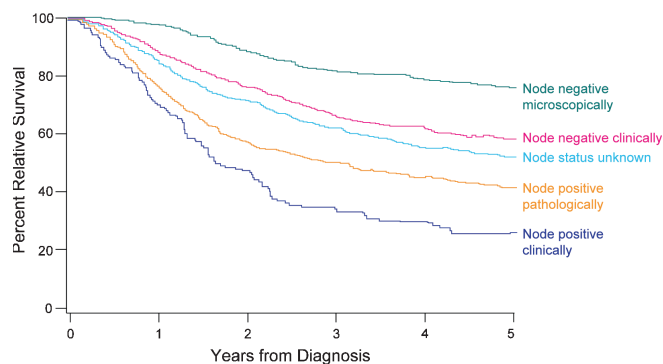
**FIGURE 30.1.** Relative survival for Merkel cell carcinoma by extent of disease at time of diagnosis. Percent relative survival was calculated for cases in the National Cancer Database using age- and sex-matched control data from the Centers for Disease Control and Prevention.



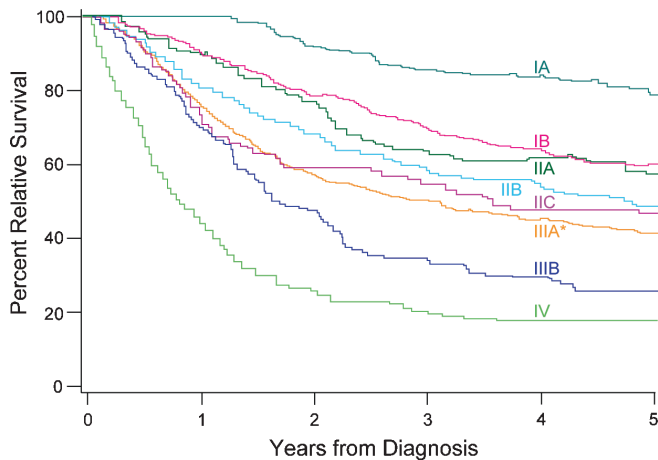
**FIGURE 30.2.** Three-year relative survival for Merkel cell carcinoma based on primary tumor dimension. While increased tumor dimension is associated with worse prognosis, these differences were modest, suggesting that tumor size alone is a poor predictor of survival. Total number of patients was 3,297, and individual groups were as follows: <1 cm = 517, 1 cm = 641, 1.5 cm = 519, 2 cm = 432, 2.5 cm = 288, 3 cm = 291, 3.5 cm = 123, ≥4 cm = 486.

## DEFINITIONS OF TNM

Those patients with MCC presentations where the primary tumor cannot be assessed should be categorized as TX. Patients with Merkel cell carcinoma in situ are categorized as Tis. The T category of MCC is classified primarily by measuring the maximum dimension of the tumor: 2 cm or less (T1), greater than 2 cm but not more than 5 cm (T2), and greater



**FIGURE 30.3.** Relative survival among 4,426 Merkel cell carcinoma patients by node status. Percent relative survival was calculated for cases in the National Cancer Database using age- and sex-matched control data from the Centers for Disease Control and Prevention. Relative survival curves shown are divided into node negative patients (top two lines), nodes status unknown (middle line), and node positive patients (bottom two lines). The curve indicated by “Node positive pathologically” includes pathologic node positive patients with clinical node status negative or unknown. Total number of patients was 4,426, and individual groupings were as follows: node negative microscopically = 630, node negative clinically = 1,726, node status unknown = 1,134, node positive pathologically = 794, node positive clinically = 143.



**FIGURE 30.4.** Relative survival for 2,856 Merkel cell carcinoma patients by stage. Percent relative survival was calculated for cases in the National Cancer Database using age- and sex-matched control data from the Centers for Disease Control and Prevention. Stages are as indicated in the figure except for Stage IIIA which could not be derived using this dataset. The curve marked “IIIA\*” represents pathologically node positive patients, with the clinical node status unknown or negative. It is anticipated that true Stage IIIA patients (clinical node status negative) have better survival than the line marked with “IIIA\*.” Total number of patients was 2,856, and individual substages were as follows: IA = 266, IB = 754, IIA = 124, IIB = 414, IIC = 84, IIIA\* = 794, IIIB = 143, IV = 277.

than 5 cm (T3). Extracutaneous invasion by the primary tumor into bone, muscle, fascia, or cartilage is classified as T4. Inclusion of 2 cm MCC tumors as T1 is consistent with the prior AJCC staging system but differs from other frequently used MCC staging systems<sup>12,14</sup> that categorize 2 cm tumors as T2. The breakdown of T category is conserved from the prior version of AJCC staging for “Carcinoma of the Skin.”

Regional metastases most commonly present in the regional lymph nodes. A second staging definition is related to nodal tumor burden: microscopic vs. macroscopic. Therefore, patients without clinical or radiologic evidence of lymph node metastases but who have pathologically documented nodal metastases are defined by convention as exhibiting “microscopic” or “clinically occult” nodal metastases. In contrast, MCC patients with both clinical evidence of nodal metastases and pathologic examination confirming nodal metastases are defined by convention as having “macroscopic” or “clinically apparent” nodal metastases. Nodes clinically positive by exam and negative by pathology would be classified as pN0. Clinically positive nodes in the draining nodal basin that are assumed to be involved with Merkel cell carcinoma but are without pathologic confirmation (no pathology performed) should be classified as N1b and the pathologic classification would be NX. Then in determining the stage grouping, it would be Stage IIIB defaulting to the higher N category.

Distant metastases are defined as metastases that have spread beyond the draining lymph node basin, including cutaneous, nodal, and visceral sites.

### Primary Tumor (T)

TX	Primary tumor cannot be assessed
T0	No evidence of primary tumor (e.g., nodal/metastatic presentation without associated primary)
Tis	In situ primary tumor
T1	Less than or equal to 2 cm maximum tumor dimension
T2	Greater than 2 cm but not more than 5 cm maximum tumor dimension
T3	Over 5 cm maximum tumor dimension
T4	Primary tumor invades bone, muscle, fascia, or cartilage

### Regional Lymph Nodes (N)

NX	Regional lymph nodes cannot be assessed
N0	No regional lymph node metastasis
cN0	Nodes negative by clinical exam* (no pathologic node exam performed)
pN0	Nodes negative by pathologic exam
N1	Metastasis in regional lymph node(s)
N1a	Micrometastasis**
N1b	Macrometastasis***
N2	In transit metastasis****

\*Clinical detection of nodal disease may be via inspection, palpation, and/or imaging.

\*\*Micrometastases are diagnosed after sentinel or elective lymphadenectomy.

\*\*\*Macrometastases are defined as clinically detectable nodal metastases confirmed by therapeutic lymphadenectomy or needle biopsy.

\*\*\*\*In transit metastasis: a tumor distinct from the primary lesion and located either (1) between the primary lesion and the draining regional lymph nodes or (2) distal to the primary lesion.

### Distant Metastasis (M)

M0	No distant metastasis
M1	Metastasis beyond regional lymph nodes
M1a	Metastasis to skin, subcutaneous tissues or distant lymph nodes
M1b	Metastasis to lung
M1c	Metastasis to all other visceral sites

### ANATOMIC STAGE/PROGNOSTIC GROUPS

Patients with primary Merkel cell carcinoma with no evidence of regional or distant metastases (either clinically or pathologically) are divided into two stages: Stage I for primary tumors  $\leq 2$  cm in size and Stage II for primary tumors  $> 2$  cm in size. Stages I and II are further divided into A and B substages based on method of nodal evaluation. Patients who have pathologically proven node

negative disease (by microscopic evaluation of their draining lymph nodes) have improved survival (substaged as A) compared to those who are only evaluated clinically (substaged as B). Stage II has an additional substage (IIC) for tumors with extracutaneous invasion (T4) and negative node status regardless of whether the negative node status was established microscopically or clinically. Stage III is also divided into A and B categories for patients with microscopically positive and clinically occult nodes (IIIA) and macroscopic nodes (IIIB). There are no subgroups of Stage IV Merkel cell carcinoma.

Stage 0	Tis	N0	M0
Stage IA	T1	pN0	M0
Stage IB	T1	cN0	M0
Stage IIA	T2/T3	pN0	M0
Stage IIB	T2/T3	cN0	M0
Stage IIC	T4	N0	M0
Stage IIIA	Any T	N1a	M0
Stage IIIB	Any T	N1b/N2	M0
Stage IV	Any T	Any N	M1

### PROGNOSTIC FACTORS (SITE-SPECIFIC FACTORS) (Recommended for Collection)

Required for staging	None
Clinically significant	Measured thickness (depth) Tumor base transection status Profound immune suppression Tumor infiltrating lymphocytes in the primary tumor (TIL) Growth pattern of primary tumor Size of tumor nests in regional lymph nodes Clinical status of regional lymph nodes Regional lymph nodes pathological extra-capsular extension Isolated tumor cells in regional lymph node(s)

### HISTOLOGIC GRADE (G)

Histologic grade is not used in the staging of Merkel cell carcinoma.

### HISTOPATHOLOGIC TYPE

While several distinct morphologic patterns have been described for MCC, these have not been reproducibly found

to be of prognostic significance. These histologic subtypes include: intermediate type (most common), small cell type (second most common), and trabecular type (least common but most characteristic pattern of MCC).

### REFERENCES

1. Tang CK, Toker C. Trabecular carcinoma of the skin: an ultrastructural study. *Cancer*. 1978;42(5):2311–21.
2. Feng H, Shuda M, Chang Y, Moore PS. Clonal integration of a polyomavirus in human merkel cell carcinoma. *Science*. 2008;319:1096–100.
3. Agelli M, Clegg LX. Epidemiology of primary Merkel cell carcinoma in the United States. *J Am Acad Dermatol*. 2003;49(5):832–41.
4. Heath ML, Jaimes N, Lemos B, et al. Clinical characteristics of Merkel cell carcinoma at diagnosis in 195 patients: the AEIOU features. *J Am Acad Dermatol*. 2008;58(3):375–81.
5. Kanitakis J, Euvrard S, Chouvet B, Butnaru AC, Claudy A. Merkel cell carcinoma in organ-transplant recipients: report of two cases with unusual histological features and literature review. *J Cutan Pathol*. 2006;33(10):686–94.
6. Penn I, First MR. Merkel's cell carcinoma in organ recipients: report of 41 cases. *Transplantation*. 1999;68(11):1717–21.
7. Hodgson NC. Merkel cell carcinoma: changing incidence trends. *J Surg Oncol*. 2005;89(1):1–4.
8. Lemos B, Nghiem P. Merkel cell carcinoma: more deaths but still no pathway to blame. *J Invest Dermatol*. 2007;127(9):2100–3.
9. Medina-Franco H, Urist MM, Fiveash J, Heslin MJ, Bland KI, Beenken SW. Multimodality treatment of Merkel cell carcinoma: case series and literature review of 1024 cases. *Ann Surg Oncol*. 2001;8(3):204–8.
10. Allen PJ, Zhang ZF, Coit DG. Surgical management of Merkel cell carcinoma. *Ann Surg*. 1999;229(1):97–105.
11. AJCC cancer staging manual. 6th ed. Chicago, IL: Springer; 2002.
12. Allen PJ, Bowne WB, Jaques DP, Brennan MF, Busam K, Coit DG. Merkel cell carcinoma: prognosis and treatment of patients from a single institution. *J Clin Oncol*. 2005;23(10):2300–9.
13. Clark JR, Veness MJ, Gilbert R, O'Brien C J, Gullane PJ. Merkel cell carcinoma of the head and neck: Is adjuvant radiotherapy necessary? *Head Neck*. 2007;29(3):249–57.
14. Yiengpruksawan A, Coit DG, Thaler HT, Urmacher C, Knapper WK. Merkel cell carcinoma. Prognosis and management. *Arch Surg*. 1991;126(12):1514–9.
15. Gupta SG, Wang LC, Penas PF, Gellenthin M, Lee SJ, Nghiem P. Sentinel lymph node biopsy for evaluation and treatment of patients with Merkel cell carcinoma: the Dana-Farber experience and meta-analysis of the literature. *Arch Dermatol*. 2006;142(6):685–90.
16. Voog E, Biron P, Martin JP, Blay JY. Chemotherapy for patients with locally advanced or metastatic Merkel cell carcinoma. *Cancer*. 1999;85(12):2589–95.
17. Buell JE, Trofe J, Hanaway MJ, et al. Immunosuppression and Merkel cell cancer. *Transplant Proc*. 2002;34(5):1780–1.

## MERKEL CELL CARCINOMA STAGING FORM

CLINICAL <i>Extent of disease before any treatment</i>	STAGE CATEGORY DEFINITIONS	PATHOLOGIC <i>Extent of disease through completion of definitive surgery</i>
<input type="checkbox"/> y clinical – staging completed after neoadjuvant therapy but before subsequent surgery	<b>TUMOR SIZE:</b> _____ <b>LATERALITY:</b> <input type="checkbox"/> midline <input type="checkbox"/> left <input type="checkbox"/> right <input type="checkbox"/> bilateral	<input type="checkbox"/> y pathologic – staging completed after neoadjuvant therapy AND subsequent surgery
<input type="checkbox"/> TX <input type="checkbox"/> T0 <input type="checkbox"/> Tis <input type="checkbox"/> T1 <input type="checkbox"/> T2 <input type="checkbox"/> T3 <input type="checkbox"/> T4	<b>PRIMARY TUMOR (T)</b> Primary tumor cannot be assessed No evidence of primary tumor <i>In situ</i> primary tumor Less than or equal to 2 cm maximum tumor dimension Greater than 2 cm but not more than 5 cm maximum tumor dimension Over 5 cm maximum tumor dimension Primary tumor invades bone, muscle, fascia, or cartilage	<input type="checkbox"/> TX <input type="checkbox"/> T0 <input type="checkbox"/> Tis <input type="checkbox"/> T1 <input type="checkbox"/> T2 <input type="checkbox"/> T3 <input type="checkbox"/> T4
<input type="checkbox"/> NX <input type="checkbox"/> N0  <input type="checkbox"/> N1  <input type="checkbox"/> N2	<b>REGIONAL LYMPH NODES (N)</b> Regional lymph nodes cannot be assessed No regional lymph node metastasis Nodes negative by clinical exam* (no pathologic node exam performed) Nodes negative by pathologic exam Metastasis in regional lymph node(s) Micrometastasis** Macrometastasis*** In transit metastasis ****  *Clinical detection of nodal disease may be via inspection, palpation and/or imaging **Micrometastases are diagnosed after sentinel or elective lymphadenectomy ***Macrometastases are defined as clinically detectable nodal metastases confirmed by therapeutic lymphadenectomy or needle biopsy ****In transit metastasis: a tumor distinct from the primary lesion and located either 1) between the primary lesion and the draining regional lymph nodes or 2) distal to the primary lesion	<input type="checkbox"/> NX <input type="checkbox"/> N0 <input type="checkbox"/> cN0 <input type="checkbox"/> pN0 <input type="checkbox"/> N1 <input type="checkbox"/> N1a <input type="checkbox"/> N1b <input type="checkbox"/> N2
<input type="checkbox"/> M0 <input type="checkbox"/> M1 <input type="checkbox"/> M1a <input type="checkbox"/> M1b <input type="checkbox"/> M1c	<b>DISTANT METASTASIS (M)</b> No distant metastasis (no pathologic M0; use clinical M to complete stage group) Metastasis beyond regional lymph nodes Metastasis to skin, subcutaneous tissues or distant lymph nodes Metastasis to lung Metastasis to all other visceral sites	<input type="checkbox"/> M1 <input type="checkbox"/> M1a <input type="checkbox"/> M1b <input type="checkbox"/> M1c

### ANATOMIC STAGE • PROGNOSTIC GROUPS

CLINICAL					PATHOLOGIC				
GROUP	T	N	M		GROUP	T	N	M	
<input type="checkbox"/>	0	Tis	N0	M0	<input type="checkbox"/>	0	Tis	N0	M0
<input type="checkbox"/>	IB	T1	N0	M0	<input type="checkbox"/>	IA	T1	pN0	M0
<input type="checkbox"/>	IIB	T2/T3	N0	M0	<input type="checkbox"/>	IB	T1	cN0	M0
<input type="checkbox"/>	IIC	T4	N0	M0	<input type="checkbox"/>	IIA	T2/T3	pN0	M0
<input type="checkbox"/>	IIIB	Any T	N1b/N2	M0	<input type="checkbox"/>	IIB	T2/T3	cN0	M0
<input type="checkbox"/>	IV	Any T	Any N	M1	<input type="checkbox"/>	IIC	T4	N0	M0
<input type="checkbox"/>	Stage unknown				<input type="checkbox"/>	IIIA	Any T	N1a	M0
					<input type="checkbox"/>	IIIB	Any T	N1b/N2	M0
					<input type="checkbox"/>	IV	Any T	Any N	M1
					<input type="checkbox"/>	Stage unknown			

<b>HOSPITAL NAME/ADDRESS</b>	<b>PATIENT NAME/INFORMATION</b>
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# MERKEL CELL CARCINOMA STAGING FORM

## PROGNOSTIC FACTORS (SITE-SPECIFIC FACTORS)

**REQUIRED FOR STAGING:** None

**CLINICALLY SIGNIFICANT:**

- Measured Thickness (Depth) \_\_\_\_\_
- Tumor Base Transection Status \_\_\_\_\_
- Profound Immune Suppression \_\_\_\_\_
- Tumor Infiltrating Lymphocytes in the Primary Tumor (TIL) \_\_\_\_\_
- Growth Pattern of Primary Tumor \_\_\_\_\_
- Size of tumor nests in regional lymph nodes \_\_\_\_\_
- Clinical Status of Regional Lymph Nodes \_\_\_\_\_
- Regional Lymph Nodes Pathological Extracapsular Extension \_\_\_\_\_
- Isolated Tumor Cells in Regional Lymph Node(s) \_\_\_\_\_

**General Notes:**

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

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

**General Notes (continued):**

**y prefix** indicates those cases in which classification is performed during or following initial multimodality 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.

**r prefix** indicates a recurrent tumor when staged after a disease-free interval, and is identified by the "r" prefix: rTNM.

**a prefix** designates the stage determined at autopsy: aTNM.

**surgical margins** is data field recorded by registrars describing the surgical margins of the resected primary site specimen as determined only by the pathology report.

**neoadjuvant treatment** is radiation therapy or systemic therapy (consisting of chemotherapy, hormone therapy, or immunotherapy) administered prior to a definitive surgical procedure. If the surgical procedure is not performed, the administered therapy no longer meets the definition of neoadjuvant therapy.

**Histologic Grade (G)** (also known as overall grade)

Histologic grade is not used in the staging of Merkel cell carcinoma.

**ADDITIONAL DESCRIPTORS**

**Lymphatic Vessel Invasion (L) and Venous Invasion (V)** have been combined into Lymph-Vascular Invasion (LVI) for collection by cancer registrars. The College of American Pathologists' (CAP) Checklist should be used as the primary source. Other sources may be used in the absence of a Checklist. Priority is given to positive results.

- Lymph-Vascular Invasion Not Present (absent)/Not Identified
- Lymph-Vascular Invasion Present/Identified
- Not Applicable
- Unknown/Indeterminate

**Residual Tumor (R)**

The absence or presence of residual tumor after treatment. In some cases treated with surgery and/or with neoadjuvant therapy there will be residual tumor at the primary site after treatment because of incomplete resection or local and regional disease that extends beyond the limit of ability of resection.

- RX Presence of residual tumor cannot be assessed
- R0 No residual tumor
- R1 Microscopic residual tumor
- R2 Macroscopic residual tumor

Clinical stage was used in treatment planning (describe): \_\_\_\_\_

National guidelines were used in treatment planning  NCCN  Other (describe): \_\_\_\_\_

Physician signature \_\_\_\_\_

Date/Time \_\_\_\_\_

HOSPITAL NAME/ADDRESS

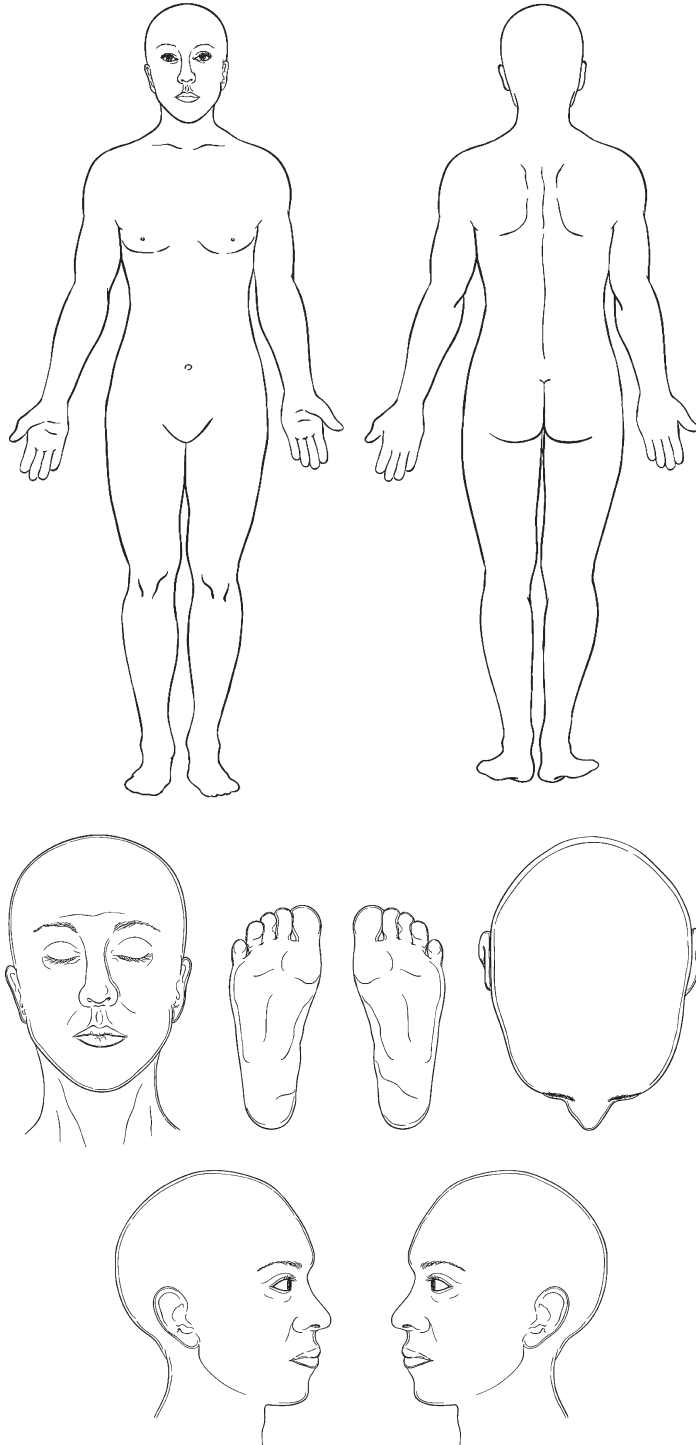
PATIENT NAME/INFORMATION

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# MERKEL CELL CARCINOMA STAGING FORM

## Illustration

Indicate on diagram primary tumor and regional nodes involved.



HOSPITAL NAME/ADDRESS

PATIENT NAME/INFORMATION



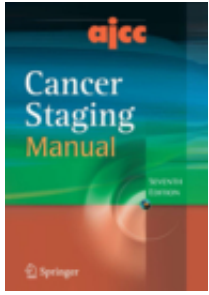
ajcc

# Cancer Staging Manual

SEVENTH  
EDITION



Springer

**AJCC Cancer Staging Manual**

Edge, S.B.; Byrd, D.R.; Compton, C.C.; Fritz, A.G.; Greene, F.L.; Trotti, A. (Eds.)

7th ed., 2010, X, 646 p. 130 illus. With CD-ROM., Softcover

ISBN: 978-0-387-88440-0

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**\$64.95**



## About this book

- The AJCC Cancer Staging Manual (with staging forms included) is the gold standard to help the cancer patient management team determine the correct stage for patients and allowing the most appropriate care plan
- The AJCC Cancer Staging Handbook is the gold standard to help the cancer patient management team determine the correct stage for patients and allowing the most appropriate care plan

The *AJCC Cancer Staging Manual* and *Handbook*, prepared by the American Joint Committee on Cancer, are used by physicians and health care professionals throughout the world to facilitate the uniform description and reporting of neoplastic diseases. Proper classification and staging is essential for the physician to assign proper treatment, evaluate results of management and clinical trials, and to serve as the standard for local, regional and international reporting on cancer incidence and outcome.

The Seventh Edition of the *AJCC Cancer Staging Manual* brings together all the currently available information on staging of cancer at various anatomic sites and incorporates newly acquired knowledge on the etiology and pathology of cancer. As knowledge of cancer biology expands, cancer staging must incorporate these advances. The current revision provides evidence-based staging based upon the established tenets of TNM classification supplemented by selected molecular markers. Relevant markers supported by evidence and of sufficient impact for treatment decisions have been included to define stage, for example Gleason's Score and PSA in prostate cancer.

Organized by disease site into 57 comprehensive chapters, the Seventh Edition features much-anticipated, major revisions to many chapters including breast, colon, prostate, kidney, and others. **There are new primary site chapters** for extrahepatic bile ducts, distal bile duct, cutaneous squamous cell carcinoma, **Merkel cell carcinoma**, and the adrenal gland plus a vastly expanded section on ophthalmologic malignancies.

User-friendly enhancements include:

- **a revised and expanded presentation of the principles and rules of TNM staging**
- a concise summary of changes in the TNM classification and "Staging at a Glance" opening each chapter to provide a snapshot of staging and coding details
- numerous new line drawings illustrating key sites throughout the text
- full color text to highlight elements of TNM, stage groupings and prognostic factors
- a revised user friendly "Staging Form"
- a CD-ROM packaged with each Manual containing printable Staging Forms

The Seventh Editions of the *AJCC Cancer Staging Manual* and *Handbook* remain the essential references for oncologists, pathologists, surgeons, cancer registrars and medical professionals worldwide to ensure that all those taking care of cancer patients are fully versed in the language of cancer staging.

**Written for:**

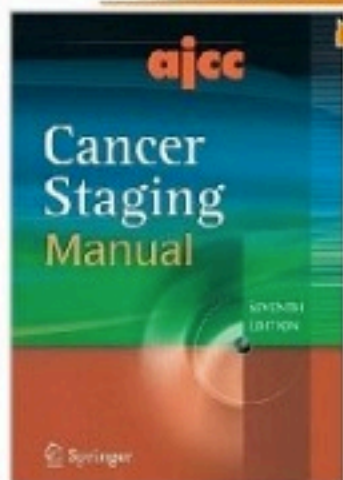
Surgeons, oncologists, pathologists, cancer registrars

**Keywords:**

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