

9th Edition TNM Staging for Thymic Malignancies

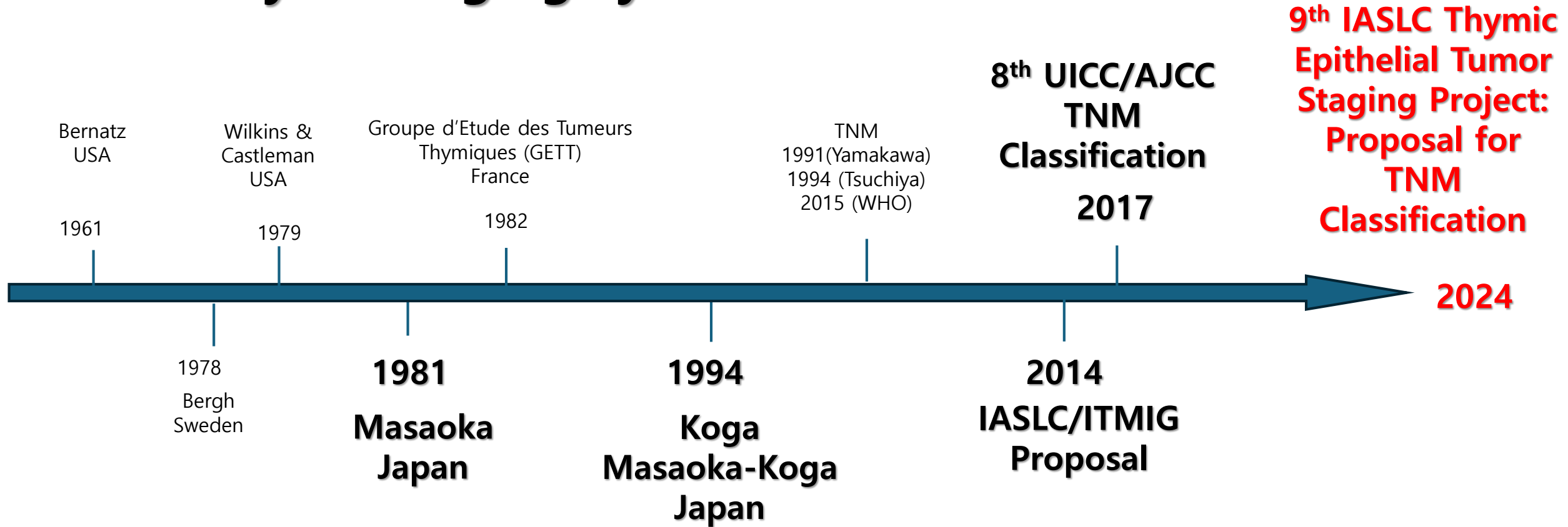
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History of Staging system



The IASLC/ITMIG Thymic Epithelial Tumors Staging Project: Proposal for an Evidence-Based Stage Classification System for the Forthcoming (8th) Edition of the TNM Classification of Malignant Tumors

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Abstract: A universal and consistent stage classification system, which describes the anatomic extent of a cancer, provides a foundation for communication and collaboration. Thymic epithelial malignancies have seen little progress, in part because of the lack of an official system. The International Association for the Study of Lung Cancer and the International Thymic Malignancies Interest Group assembled a large retrospective database, a multispecialty international committee and carried out extensive analysis to develop proposals for the 8th edition of the stage classification manuals. This tumor, node, metastasis (TNM)-based system is applicable to all types of thymic epithelial malignancies. This article summarizes the proposed definitions of the T, N, and M components and describes how these are combined into stage groups. This represents a major step forward for thymic malignancies.

Key Words: Staging, Prognosis, Thymoma, Thymic carcinoma, Stage classification

(*J Thorac Oncol.* 2014;9: S65–S72)

Thymic epithelial malignancies are rare tumors. There have been many obstacles to progress in these diseases. Among these has been the lack of an official, consistent stage classification system put forth by the American Joint Committee on Cancer (AJCC) and the Union for International Cancer Control (UICC)—the bodies responsible for defining stage classification throughout the world. At least 15 different stage classification systems have been proposed and used.¹ These have been largely empirically derived, based on data from small numbers of patients. Perhaps the most widely used have been the Masaoka classification (derived from data on 91 patients),² and the Koga modification of this (based on 76 patients).³ Even among centers using one of these classification systems, often the definitions have been interpreted differently because of vague wording, thus hampering the ability to collaborate effectively.⁴

In 2009, both the nascent International Thymic Malignancies Interest Group (ITMIG) and the International Association for the Study of Lung Cancer (IASLC) recognized the need for a consistent stage classification system for thymic malignancies. These organizations formed a partnership to address this, with ITMIG providing the engagement of the vast majority of clinicians and researchers active in these diseases, and IASLC providing funding for the project and statistical analysis and its expertise in developing proposals for stage classification from its experience in doing this in lung cancer.⁵ A Thymic Domain of the Staging and Prognostic Factors Committee (TD-SPFC) was established collaboratively by IASLC and ITMIG (Appendix 6). IASLC led discussions and received approval from AJCC and UICC to develop proposals for stage classification of thymic malignancies that

8th UICC/AJCC TNM Classification

- In 2009, IASLC and ITMIG collaborated to develop a TNM-based staging system using a large worldwide retrospective database of over 8000 patients.
- Creation of the Thymic Domain of the Staging and Prognostic Factors Committee (TD-SPFC) within the IASLC.
- The proposed TNM classification system for thymic tumors was incorporated into the 8th edition of the TNM classification for thoracic malignancies, approved by UICC and AJCC, becoming effective in 2017 and 2018, respectively.
- Following the release of the eighth TNM edition, the TD-SPFC began working on proposals for the 9th edition, expected in 2024.

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†††See Appendix 1;‡‡‡see Appendices 2, 3, 4;§§§see Appendix 5.

Disclosure: The authors declare no conflict of interest.

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ISSN: 1556-0864/14/0909-0565

Web-based Cross-sectional Survey Questionnaire

| 항목 | 비율(%) |
|--------------------------|------------------|
| 응답 수 | 217 |
| 국가 수 | 37 |
| 대륙 수 | 4 |
| 유용하다고 생각한 비율 | 78 |
| 일상적으로 사용한 비율 | 64 |
| 림프절 지도 인지 비율 | 72 |
| 림프절 지도 사용 비율 | 48 |
| 림프절 지도가 효과적이라고 생각한 비율 | 54 |
| 흉선종 환자의 N1 림프절 절제율 | 50 |
| 흉선 암종 환자의 N1 림프절 절제율 | 66 |
| 흉선종 환자의 N2 림프절 절제율 | 21 |
| 흉선 암종 환자의 N2 림프절 절제율 | 41 |
| 림프절 절제술이 가장 많이 수행된 종양 단계 | T3 (33%) |
| 림프절 절제술이 덜 자주 수행된 종양 단계 | T2 (9%), T1 (8%) |

8th Edition TNM staging system for TET

| TABLE 3. Stage Grouping | | | | |
|-------------------------|-------|-----------|--------------|--------------|
| TABLE 1. T | Stage | T | N | M |
| Category | | | | |
| T1 | I | T1 | N0 | M0 |
| a | II | T2 | N0 | M0 |
| b | IIIa | T3 | N0 | M0 |
| T2 | IIIb | T4 | N0 | M0 |
| T3 | IVa | T any | N1 | M0 |
| | | T any | N0,1 | M1a |
| T4 | IVb | T any | N2 | M0,1a |
| | | T any | N any | M1b |

^aInvolvement of

^bA tumor is cl

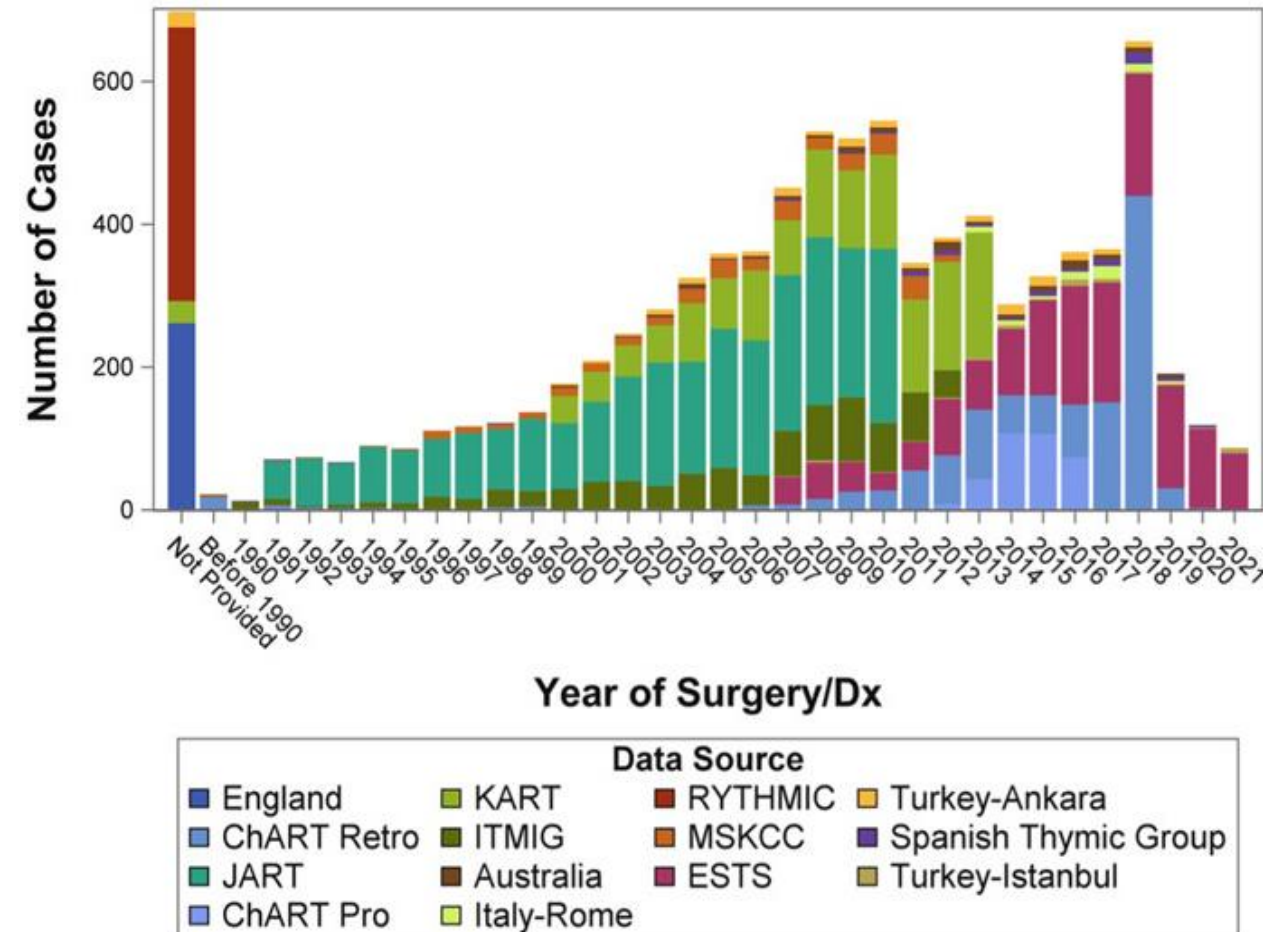
with or without any invasion of structures of lower 1 levels.

^aInvolvement must be pathologically proven in pathologic staging.

IASLC/ITMIC Staging Project – Thymic Domain

| | 8 th Edition (2017) | 9 th Edition (2024) |
|------------------------------|--------------------------------|--------------------------------|
| Periods of Disgnosis | 1990-2012 | 1990-2021 |
| Total atients submitted | 10,808 | 11,347 |
| Regional distribution | | |
| Europe | 2653(33%) | 3113(34%) |
| Asia/Australia | 4043(50%) | 5628(62%) |
| North America | 1383(17%) | 406(4%) |
| South/Central America | 66 | 0 |
| Final enrolled patients | 8145 | 9147 |
| -Thymoma | 7016(86%) | 7662(84%) |
| -Thymic carcinoma | 962(12%) | 1345(15%) |
| -Neuroendocrine thymic tumor | 164(2%) | 140(105%) |
| Treatment modalities | | |
| -Surgery included | 8018 (98%) | 8830(96%) |
| -Non-surgical | 127(2%) | 251(4%) |
| Resection status | | |
| R0 | 6621(81%) | 7647(84%) |
| R+ | 1105(19%) | 1121(16%) |

Data Sources For the 9th TNM staging



Supplemental table 1. Contributing centers.

| Data Source | Total Number of Patients Submitted | Thymoma, Thymic Carcinoma or NETT | Survival Data Available |
|------------------------------|------------------------------------|-----------------------------------|-------------------------|
| Total | 11,347 | 10,567 | 9,147 |
| ChART Prospective database | 625 | 590 | 343 |
| ChART Retrospective database | 1,532 | 1,483 | 1,172 |
| ESTS | 2,305 | 1,739 | 1,411 |
| England | 285 | 283 | 262 |
| Turkey-Istanbul | 77 | 49 | 47 |
| ITMIG | 1,233 | 1,218 | 813 |
| JART | 2,711 | 2,670 | 2,659 |
| KART | 1,363 | 1,360 | 1,357 |
| MSKCC | 322 | 320 | 288 |
| Italy-Rome | 64 | 63 | 63 |
| RYTHMIC | 395 | 385 | 383 |
| Spanish Thymic Group | 124 | 119 | 86 |
| Australia | 114 | 111 | 97 |
| Turkey-Ankara | 197 | 177 | 166 |

NETT: Neuroendocrine Thymic Tumors; ChART: Chinese Alliance for Research in Thymoma; ESTS: European Society of Thoracic Surgeons; ITMIG: International Thymic Malignancies Interest Group; JART: Japanese Association for Research in the Thymus; KART: Korean Association for Research in the Thymus; MSKCC: Memorial Sloan Kettering Cancer Center; Réseau Tumeurs THYMIques et Cancer (RYTHMIC).



**What has changed
in T components?**

T staging of 8th and 9th Edition

TABLE 1. T Descriptors

| Category | Definition (Involvement of) ^{a,b} |
|----------|---|
| T1 | |
| a | Encapsulated or unencapsulated, with or without extension into mediastinal fat |
| b | Extension into mediastinal pleura |
| T2 | Pericardium |
| T3 | Lung, brachiocephalic vein, superior vena cava, chest wall, phrenic nerve, hilar (extrapericardial) pulmonary vessels |
| T4 | Aorta, arch vessels, main pulmonary artery, myocardium, trachea, or esophagus |

^aInvolvement must be pathologically proven in pathologic staging.

^bA tumor is classified according to the highest T level of involvement that is present with or without any invasion of structures of lower T levels.

Table 1. Proposed T Component of Thymic Tumors for the Ninth Edition of the TNM Classification of Malignant Tumors

| T | Description |
|-----|--|
| T1 | Tumor limited to the thymus with or without encapsulation, or directly invades into the mediastinum alone or directly invades the mediastinal pleura but does not involve any other mediastinal structure. |
| T1a | 5 cm or less in its greatest dimension ^a |
| T1b | larger than 5 cm in its greatest dimension ^a |
| T2 | Tumor directly invades the pericardium (either partial or full-thickness), the lung, or the phrenic nerve |
| T3 | Tumor directly invades any of the following: (1) brachiocephalic vein, (2) superior vena cava, (3) chest wall, or (4) extrapericardial pulmonary arteries or veins |
| T4 | Tumor directly invades any of the following: (1) aorta (ascending, arch, or descending); (2) arch vessels; (3) intrapericardial pulmonary artery or veins; (4) myocardium; (5) trachea; or (6) esophagus. |

^aIrrespective of mediastinal pleura invasion. Mediastinal pleura invasion is to be recorded as an “additional histologic descriptor.”

OS by Pathologic T category (proposed ninth TNM) in N0M0R-any

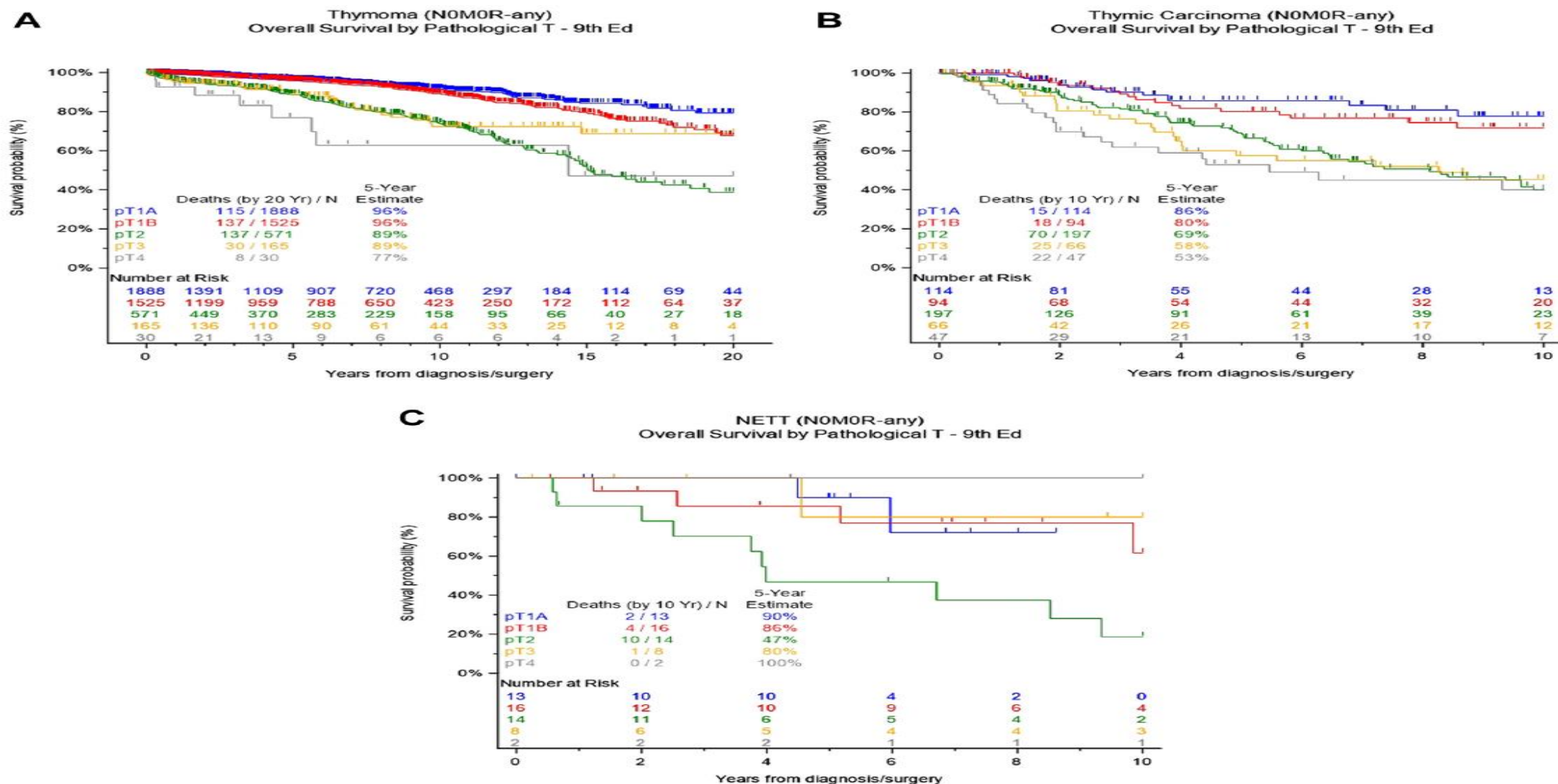


Figure 1. OS by pathologic T category (proposed ninth TNM) in N0M0R-any for (A)Thymoma, (B) Thymic Carcinoma, and (C) NETT. NETT, neuroendocrine thymic tumor; OS, overall survival; R-any, regardless of R status.

FFR by pathologic T category (proposed ninth TNM) in NOMOR0 cases

IASLC Thymic Epithelial Tumor Staging Project for the 9th TNM: T Component

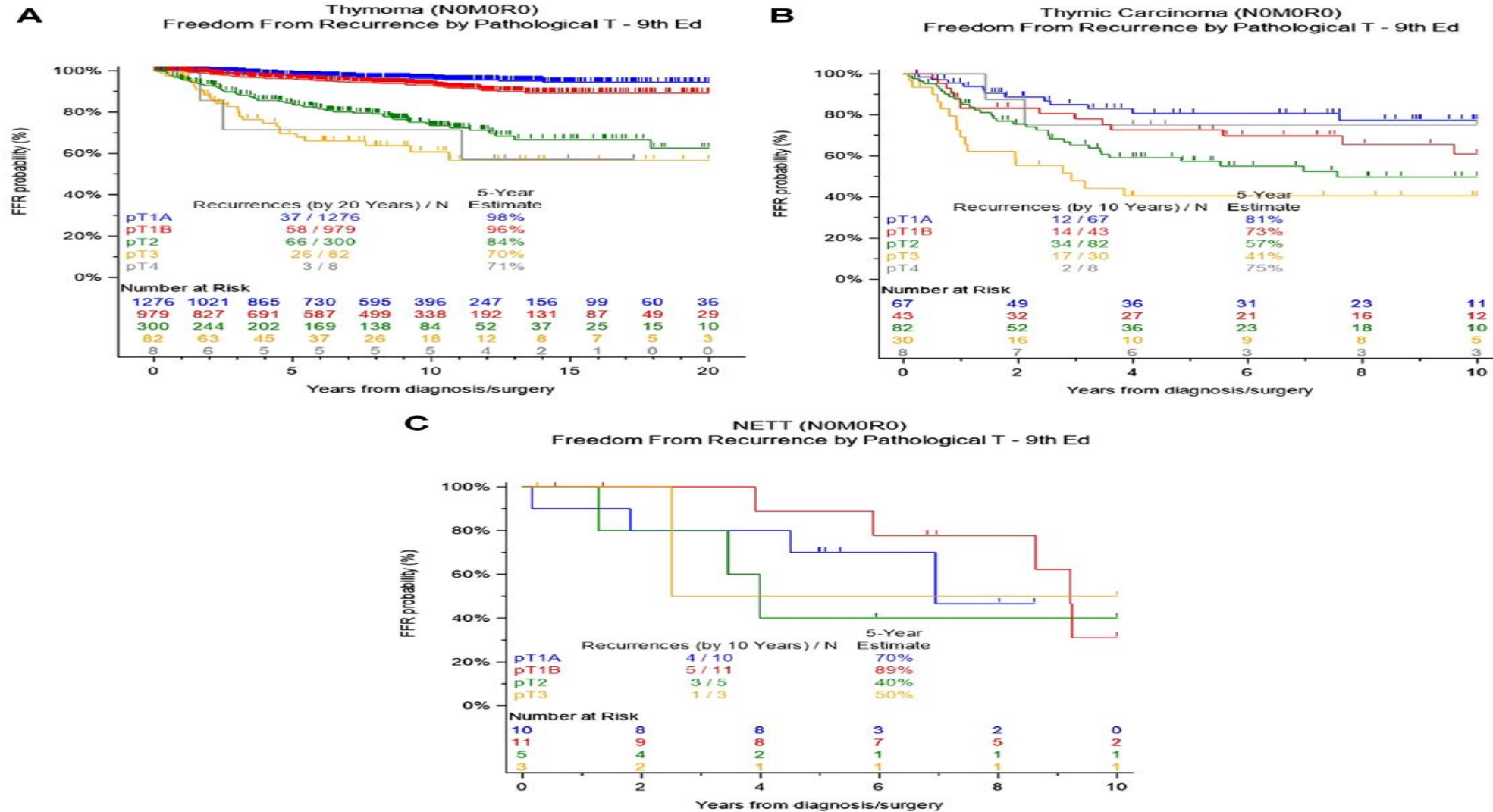


Figure 2. FFR by pathologic T category (proposed ninth TNM) in NOMOR0 cases for (A) Thymoma, (B) Thymic Carcinoma, and (C) NETT. FFR, Freedom-from-recurrence; NETT, neuroendocrine thymic tumor; R0, complete resection.

CIR by pathologic T category (proposed ninth TNM) in N0M0R0 cases

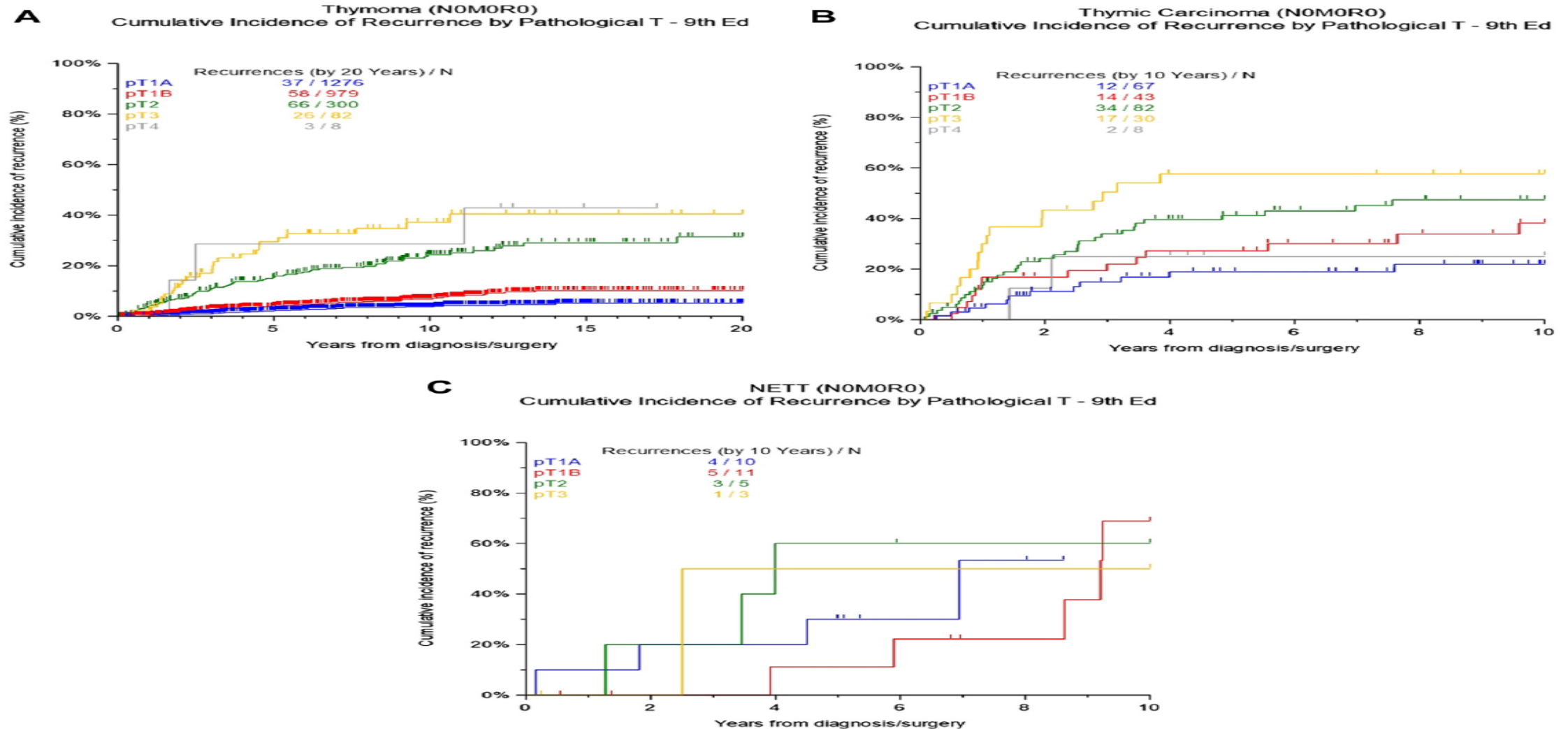
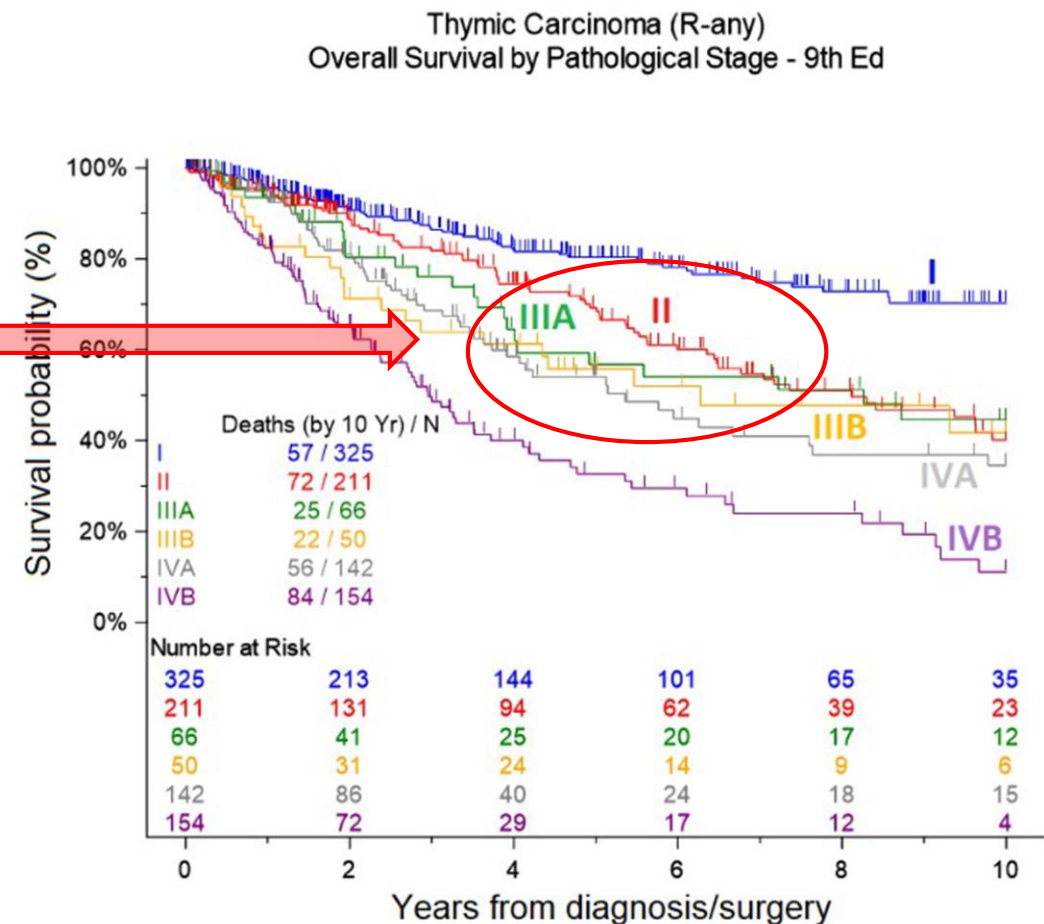
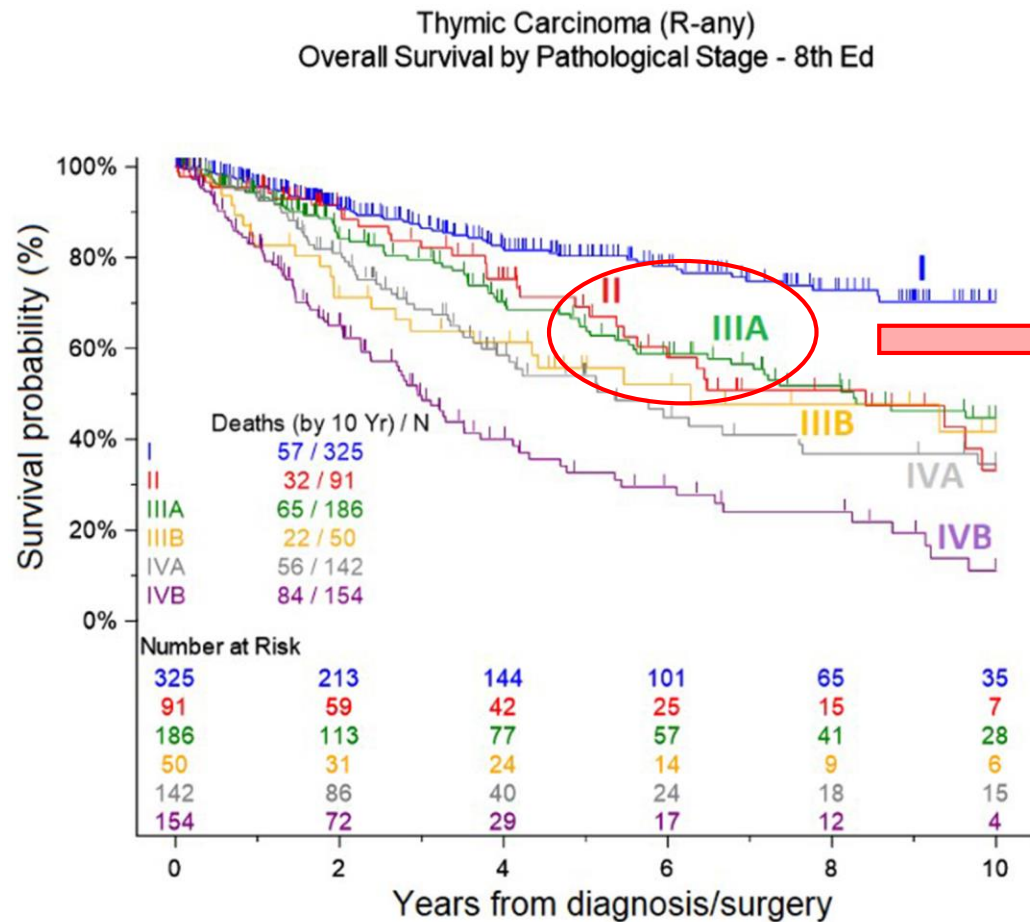


Figure 3. CIR by pathologic T category (proposed ninth TNM) in N0M0R0 cases for (A)Thymoma, (B) Thymic Carcinoma, and (C) NETT. CIR, cumulative incidence of recurrence; NETT, neuroendocrine thymic tumor; R0, complete resection.

Thymic Carcinoma – Overall Survival

Journal of Thoracic Oncology 2023
181655-1671DOI:
(10.1016/j.jtho.2023.09.002)



A 3D rendering of a puzzle with one red piece standing out among many grey pieces. The red piece is in the center, and the text is overlaid on it.

**What has changed in N
and M components?**

N and M Staging of 8th and 9th Edition

| Category | Definition (Involvement of) ^a | N and M Categories | Description |
|---|---|--------------------|---|
| N0 | No nodal involvement | N0 | No nodal involvement |
| N1 | Anterior (perithymic) nodes | N1 | Anterior (perithymic) nodes |
| N2 | Deep intrathoracic or cervical nodes | N2 | Deep intrathoracic or cervical nodes (e.g., paratracheal, subcarinal, aortopulmonary window, hilar, jugular, and supraclavicular nodes) |
| M0 | No metastatic pleural, pericardial, or distant sites | M0 | No metastatic pleural, pericardial, or distant sites |
| M1 | | | |
| a | Separate pleural or pericardial nodule(s) | M1a | Separate pleural or pericardial nodule(s) |
| b | Pulmonary intraparenchymal nodule or distant organ metastasis | M1b | Pulmonary intraparenchymal nodule or distant organ metastasis |
| ^a Involvement must be pathologically proven in pathologic staging. | | | |

No changed

Distribution of clinical and pathologic stage

Supplemental table 2a. Distribution of clinical and pathologic stage. Missing cases included.

| | Thymoma | Thymic Carcinoma | NETT | Overall |
|---------------------------|--------------|------------------|-------------|--------------|
| Clinical Stage | | | | |
| Missing or Not Determined | 5727 (74.7%) | 990 (73.6%) | 114 (81.4%) | 6831 (74.7%) |
| I | 1501 (19.6%) | 162 (12.0%) | 15 (10.7%) | 1678 (18.3%) |
| II | 99 (1.3%) | 20 (1.5%) | 1 (0.7%) | 120 (1.3%) |
| IIIA | 154 (2.0%) | 57 (4.2%) | 6 (4.3%) | 217 (2.4%) |
| IIIB | 31 (0.4%) | 14 (1.0%) | 1 (0.7%) | 46 (0.5%) |
| IVA | 94 (1.2%) | 39 (2.9%) | 2 (1.4%) | 135 (1.5%) |
| IVB | 56 (0.7%) | 63 (4.7%) | 1 (0.7%) | 120 (1.3%) |
| Pathologic Stage | | | | |
| Missing or Not Determined | 2053 (26.8%) | 305 (22.7%) | 51 (36.4%) | 2409 (26.3%) |
| I | 4266 (55.7%) | 325 (24.2%) | 38 (27.1%) | 4629 (50.6%) |
| II | 281 (3.7%) | 91 (6.8%) | 8 (5.7%) | 380 (4.2%) |
| IIIA | 639 (8.3%) | 278 (20.7%) | 18 (12.9%) | 935 (10.2%) |
| IIIB | 34 (0.4%) | 50 (3.7%) | 2 (1.4%) | 86 (0.9%) |
| IVA | 306 (4.0%) | 142 (10.6%) | 16 (11.4%) | 464 (5.1%) |
| IVB | 83 (1.1%) | 154 (11.4%) | 7 (5.0%) | 244 (2.7%) |

For the analysis all cases with valid histologic type and survival data were included.
NETT: Neuroendocrine Thymic Tumors

Supplemental table 2b. Distribution of clinical and pathologic stage. Missing cases excluded

| Summary of data for the stage analysis | | | | |
|--|-------------------|-------------------|----------------|-------------------|
| | Thymoma | Thymic Carcinoma | NETT | Overall |
| Data Available, n/N (%) | | | | |
| Clinical Stage | 1935/7662 (25.3%) | 355/1345 (26.4%) | 26/140 (18.6%) | 2316/9147 (25.3%) |
| Pathological Stage | 5609/7662 (73.2%) | 1040/1345 (77.3%) | 89/140 (63.6%) | 6738/9147 (73.7%) |
| Clinical Stage, n (%) | | | | |
| I | 1501 (77.6%) | 162 (45.6%) | 15 (57.7%) | 1678 (72.5%) |
| II | 99 (5.1%) | 20 (5.6%) | 1 (3.8%) | 120 (5.2%) |
| IIIA | 154 (8%) | 57 (16.1%) | 6 (23.1%) | 217 (9.4%) |
| IIIB | 31 (1.6%) | 14 (3.9%) | 1 (3.8%) | 46 (2%) |
| IVA | 94 (4.9%) | 39 (11%) | 2 (7.7%) | 135 (5.8%) |
| IVB | 56 (2.9%) | 63 (17.7%) | 1 (3.8%) | 120 (5.2%) |
| Pathological Stage, n (%) | | | | |
| I | 4266 (76.1%) | 325 (31.3%) | 38 (42.7%) | 4629 (68.7%) |
| II | 281 (5%) | 91 (8.8%) | 8 (9%) | 380 (5.6%) |
| IIIA | 639 (11.4%) | 278 (26.7%) | 18 (20.2%) | 935 (13.9%) |
| IIIB | 34 (0.6%) | 50 (4.8%) | 2 (2.2%) | 86 (1.3%) |
| IVA | 306 (5.5%) | 142 (13.7%) | 16 (18%) | 464 (6.9%) |
| IVB | 83 (1.5%) | 154 (14.8%) | 7 (7.9%) | 244 (3.6%) |
| n/N=number of cases with data available out of included cases in the 9 th edition database. | | | | |

NETT: Neuroendocrine Thymic Tumors

Bubble depiction of the clinical and pathologic concordance of N

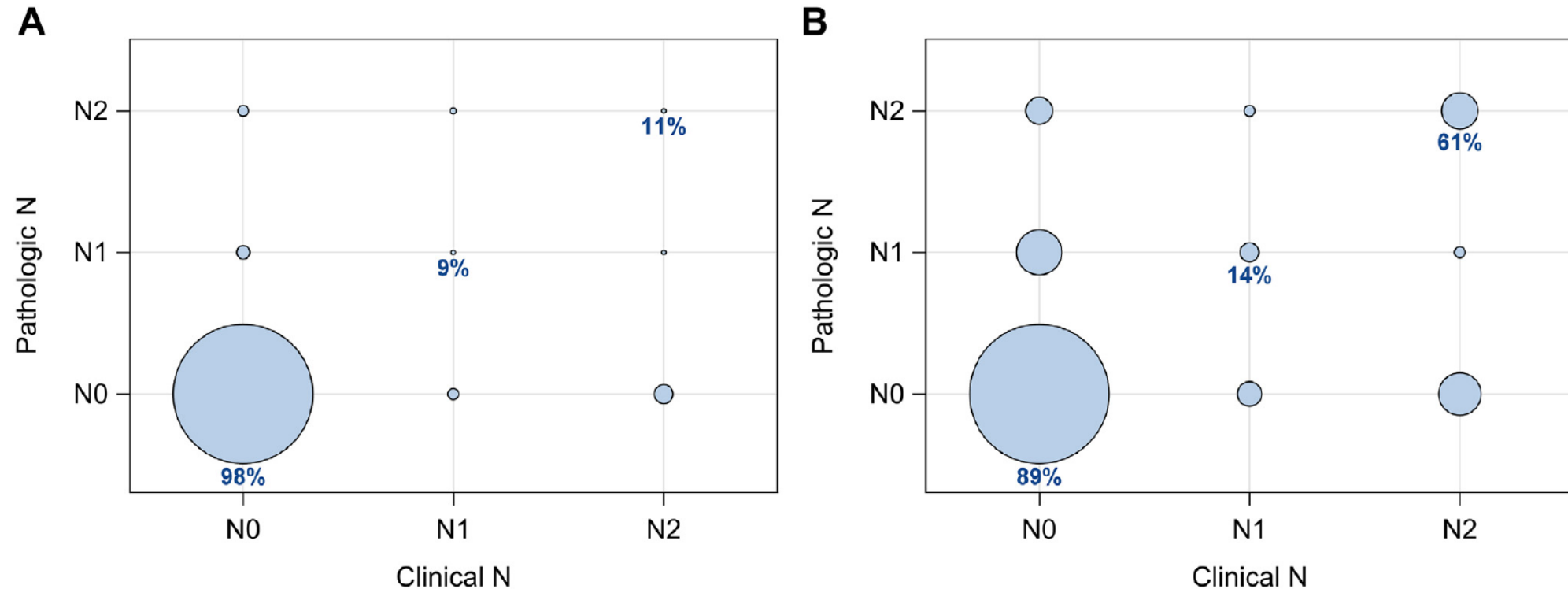


Figure 1. Bubble depiction of the clinical and pathologic concordance of N. The size of the bubble corresponds with the percent of pathologic N patients correctly identified clinically (by imaging). (A) Thymoma. (B) Thymic carcinoma.

Overall survival by clinical N

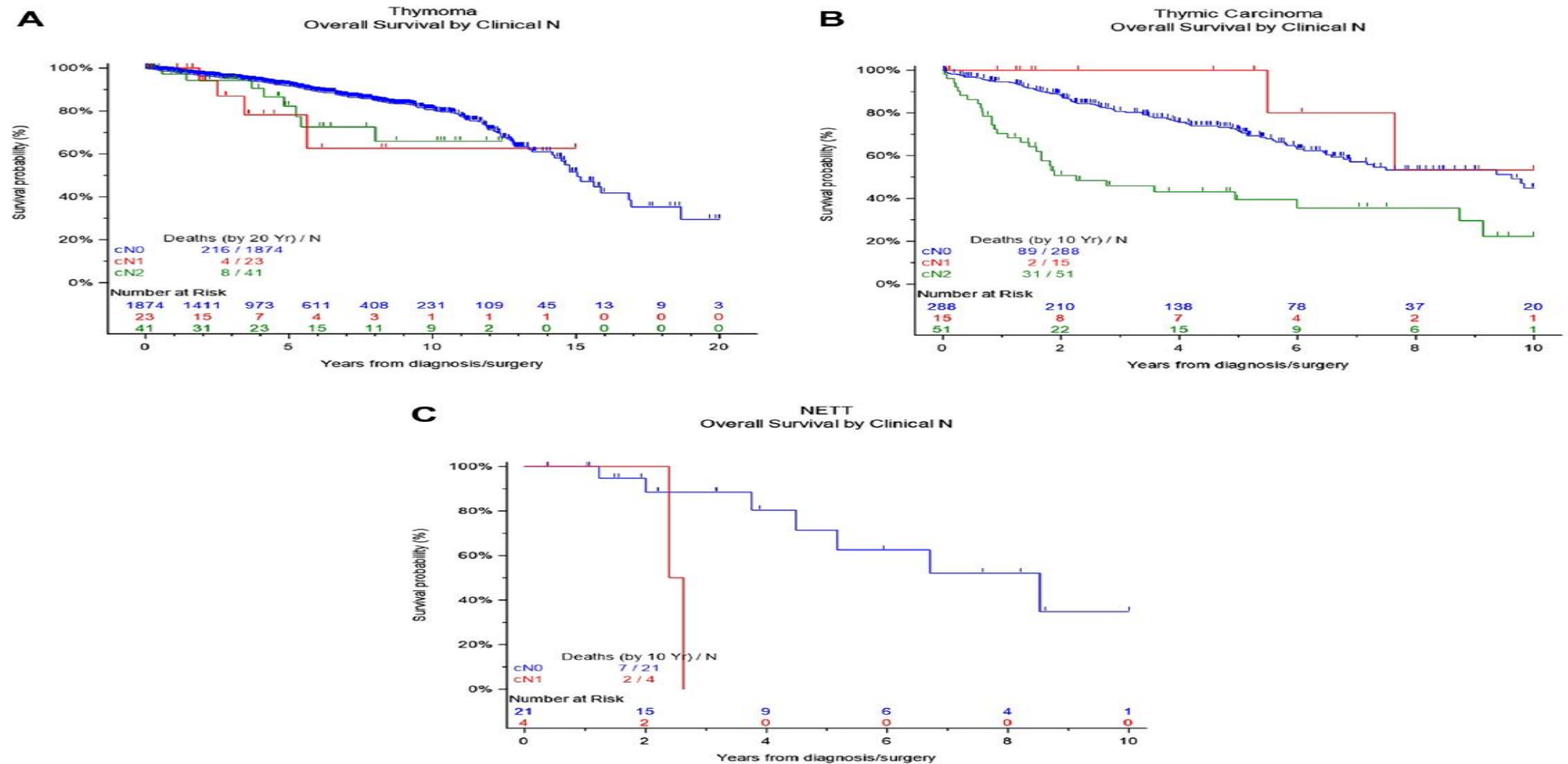


Figure 2. Overall survival by clinical N in (A) thymoma, (B) thymic carcinoma, and (C) NETT. NETT, neuroendocrine thymic tumor.

Overall survival by pathologic N

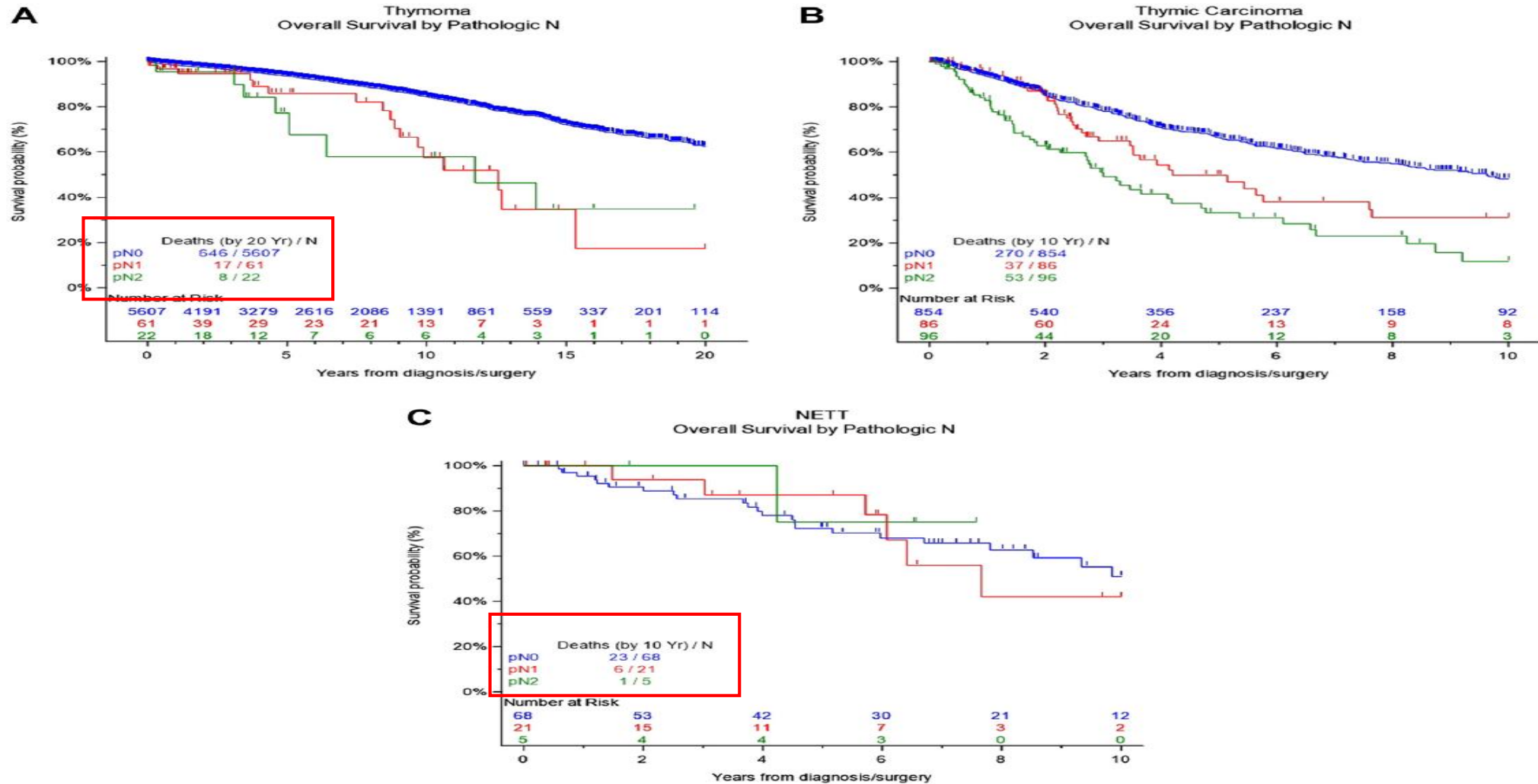


Figure 3. Overall survival by pathologic N in (A) thymoma, (B) thymic carcinoma, and (C) NETT. NETT, neuroendocrine thymic tumor.

Overall survival by clinical M

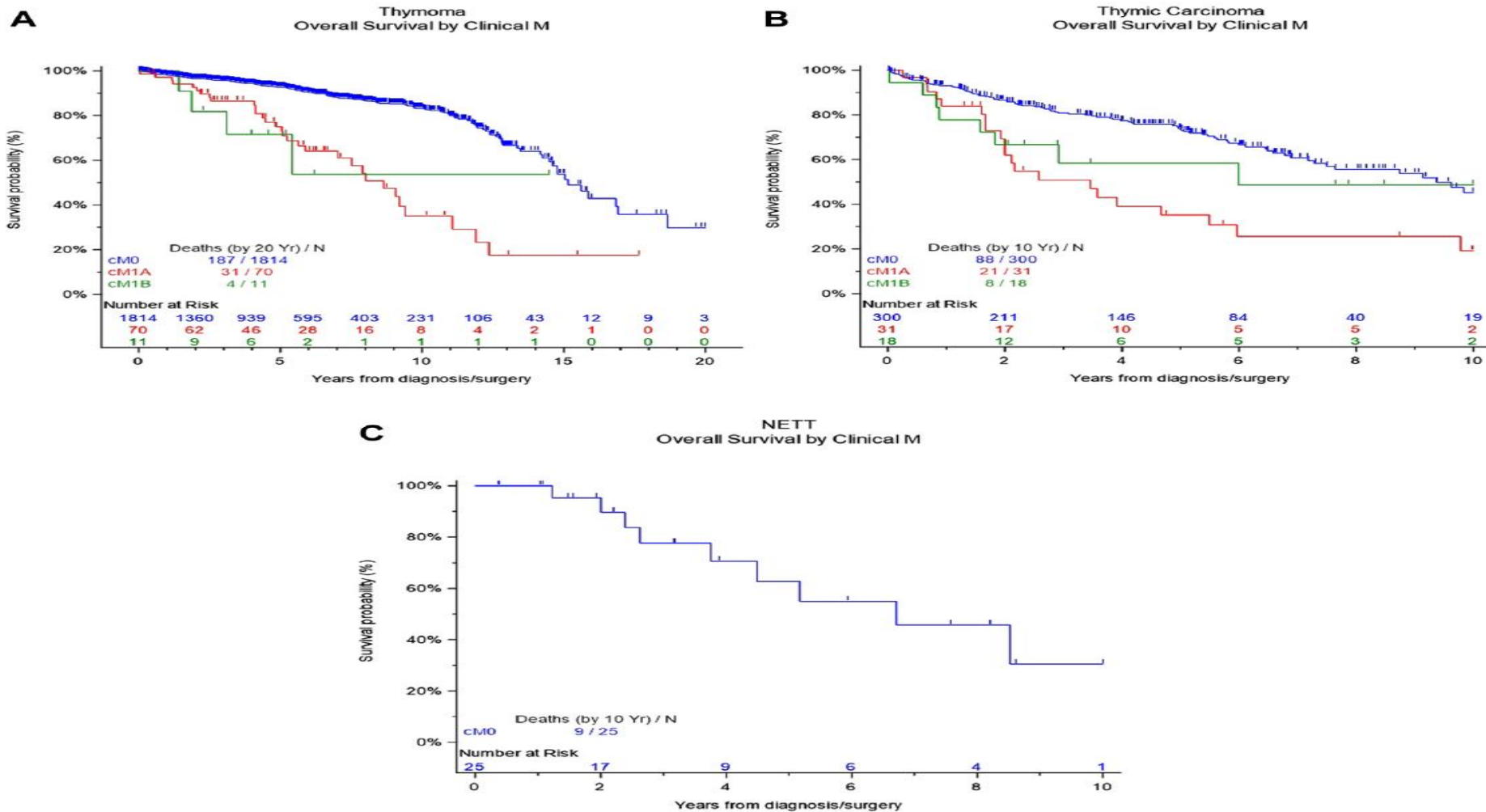


Figure 5. Overall survival by clinical M in (A) thymoma, (B) thymic carcinoma, and (C) NETT. NETT, neuroendocrine thymic tumor.

Overall survival by pathologic M

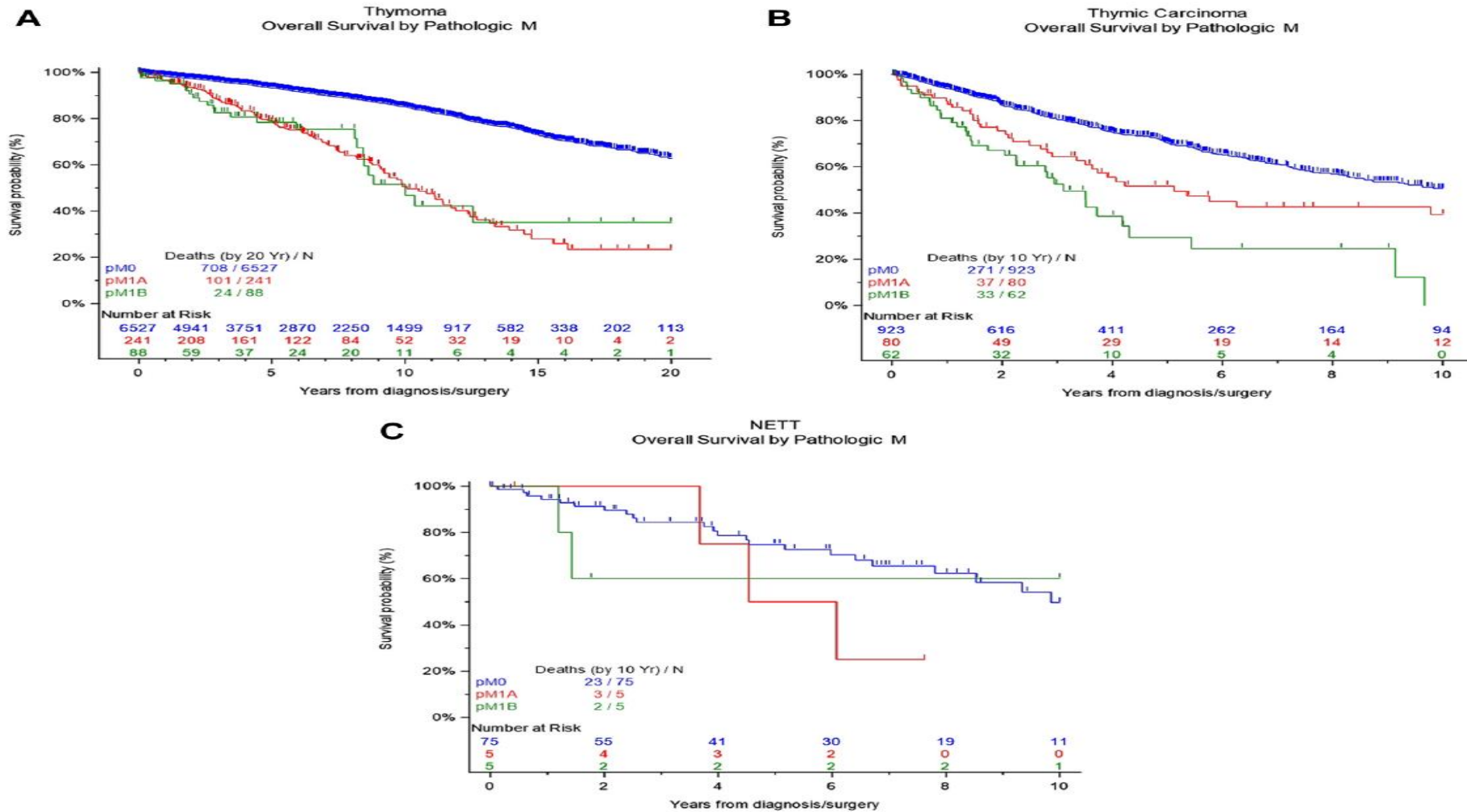


Figure 6. Overall survival by pathologic M in (A) thymoma, (B) thymic carcinoma, and (C) NETT. NETT, neuroendocrine thymic tumor.

Multivaiable Cox Regression Model for OS and FFR (pM staging)

Supplemental Table 4. Results from multivariable Cox regression modeling for Overall survival (OS) and Freedom from recurrence (FFR) by pathological M category, performed separately in thymoma and thymic carcinoma.

| Outcome | Variable in Multivariable Cox Regression | n/N (%) | HR (95% CI) | P-value |
|-------------------------|--|-----------------|--------------------|---------|
| OS in Thymic Carcinoma | pM1a (vs pM0) | 66/693 (10%) | 1.35 (0.91, 2.00) | 0.133 |
| | pM1b (vs pM1a) | 38/693 (5%) | 1.68 (0.99, 2.86) | 0.053 |
| | Age: 65+ (vs <65) | 243/693 (35%) | 1.42 (1.09, 1.84) | 0.009 |
| | Sex: Female (vs Male) | 244/693 (35%) | 0.77 (0.59, 1.01) | 0.058 |
| | Region: Asia/Australia (vs Europe/North America) | 561/693 (81%) | 0.72 (0.53, 0.99) | 0.040 |
| | Performance Status 2+ (vs. 0/1) | 25/693 (4%) | 2.03 (1.08, 3.82) | 0.025 |
| | Surgical resection: R0 (vs no surgery) | 495/693 (71%) | 0.18 (0.11, 0.29) | <0.001 |
| | Surgical resection: R1 (vs no surgery) | 86/693 (12%) | 0.36 (0.22, 0.61) | <0.001 |
| | Surgical resection: R2 (vs no surgery) | 73/693 (11%) | 0.48 (0.29, 0.81) | 0.005 |
| | Surgical resection: RX (vs no surgery) | 4/693 (1%) | 0.52 (0.07, 3.94) | 0.517 |
| OS in Thymoma | pM1a (vs pM0) | 185/5112 (4%) | 3.23 (2.38, 4.55) | <0.001 |
| | pM1b (vs pM1a) | 44/5112 (1%) | 0.39 (0.19, 0.79) | 0.008 |
| | Age: 65+ (vs <65) | 1289/5112 (25%) | 2.71 (2.29, 3.21) | <0.001 |
| | Sex: Female (vs Male) | 2726/5112 (53%) | 0.77 (0.65, 0.90) | 0.001 |
| | Region: Asia/Australia (vs Europe/North America) | 4086/5112 (80%) | 0.43 (0.36, 0.53) | <0.001 |
| | Performance Status 2+ (vs. 0/1) | 193/5112 (4%) | 2.98 (2.18, 4.08) | <0.001 |
| | Surgical resection: R0 (vs no surgery) | 4702/5112 (92%) | 0.35 (0.19, 0.66) | <0.001 |
| | Surgical resection: R1 (vs no surgery) | 250/5112 (5%) | 0.57 (0.30, 1.08) | 0.083 |
| | Surgical resection: R2 (vs no surgery) | 122/5112 (2%) | 1.27 (0.68, 2.35) | 0.456 |
| | Surgical resection: RX (vs no surgery) | 13/5112 (0%) | 2.35 (0.78, 7.06) | 0.118 |
| FFR in Thymic Carcinoma | pM1a (vs pM0) | 13/227 (6%) | 2.63 (1.33, 5.26) | 0.004 |
| | pM1b (vs pM1a) | 3/227 (1%) | 1.11 (0.23, 5.44) | 0.898 |
| | Age: 65+ (vs <65) | 76/227 (33%) | 0.74 (0.46, 1.17) | 0.194 |
| | Sex: Female (vs Male) | 82/227 (36%) | 0.82 (0.51, 1.32) | 0.422 |
| | Region: Asia/Australia (vs Europe/North America) | 190/227 (84%) | 1.57 (0.79, 3.12) | 0.196 |
| | Performance Status 2+ (vs. 0/1) | 5/227 (2%) | 1.47 (0.19, 11.10) | 0.707 |
| FFR in Thymoma | pM1a (vs pM0) | 43/2855 (2%) | 14.3 (9.09, 43.0) | <0.001 |
| | pM1b (vs pM1a) | 9/2855 (0%) | 1.13 (0.46, 2.76) | 0.793 |
| | Age: 65+ (vs <65) | 789/2855 (28%) | 0.82 (0.60, 1.13) | 0.224 |
| | Sex: Female (vs Male) | 1558/2855 (55%) | 0.75 (0.58, 0.98) | 0.036 |
| | Region: Asia/Australia (vs Europe/North America) | 2453/2855 (86%) | 1.04 (0.70, 1.54) | 0.857 |
| | Performance Status 2+ (vs. 0/1) | 74/2855 (3%) | 0.41 (0.10, 1.67) | 0.200 |

HR- Hazard Ratio, 95% CI- 95% Confidence Interval, P-value from Score Chi-Square Test in Cox Regression

Nodal Mapping System

N1 stage – Anterior Region

TABLE 2. Anterior Region (N1) (Anterior Mediastinal and Anterior Cervical Nodes)

| Region Boundaries | Node Groups ^{14, 16} | Node Group Boundaries |
|---|---|---|
| Sup: Hyoid Bone Lat (Neck): Medial Border of Carotid Sheaths Lat (Chest): Mediastinal Pleura | Low Ant Cervical: Pretracheal, Paratracheal, Peri-thyroid, Precricoid/Delphian (AAO-HNS / ASHNS Level 6 / IASLC Level 1) | Sup: inferior border of cricoid Lat: common carotid arteries Inf: superior border of manubrium |
| Ant: Sternum Post (Medially): Great Vessels, Pericardium Post (Laterally): Phrenic Nerve Inf: Xiphoid, diaphragm | Peri-Thymic Prevascular (IASLC Level 3a) Para-aortic, Ascending Aorta, Superior Phrenics (IASLC Level 6) Supradiaphragmatic / Inferior Phrenics / Pericardial (along inferior poles of thymus) | Proximity to thymus Sup: apex of chest Ant: posterior sternum Post: anterior SVC Inf: carina Sup: line tangential to sup border of aortic arch Inf: inf border of aortic arch Sup: inf border of aortic arch Ant: post sternum Post: phrenic nerve (laterally) or pericardium (medially) Inf: diaphragm |

Region and node group boundaries adapted directly from definitions established by AAO-HNS, ASHNS, and IASLC.

AAO-HNS, American Academy of Otolaryngology—Head and Neck Surgery; ASHNS, American Society for Head and Neck Surgery; IASLC, International Association for the Study of Lung Cancer; sup, superior; ant, anterior; inf, inferior; lat, lateral; post, posterior; SVC, superior vena cava.

Table 1. Anterior Region (N1)—Prevascular Mediastinum and Anterior Cervical Lymph Nodes

| Region Boundaries | Node Groups ^a | Node Group Boundaries |
|---|--|--|
| Superior: lower border of cricoid cartilage Lateral (neck): medial border of the carotid sheath/jugular vein Lateral (chest): mediastinal pleura Anterior: sternum Posterior (medially): great vessels, pericardium Posterior (laterally): phrenic nerve Inferior: xiphoid, diaphragm | Low anterior cervical: peritracheal, perithyroid, (AAO-HNS/ASHNS level 6/IASLC level 1) Peri-thymic Prevascular (IASLC level 3a) Para-aortic, ascending aorta, superior phrenic (IASLC level 6) Supradiaphragmatic/inferior phrenic/pericardial (along inferior poles of thymus) | Superior: lower border of the cricoid cartilage Lateral: common carotid arteries Inferior: superior border of the manubrium Proximity to the thymus Superior: apex of chest Anterior: posterior sternum Posterior: anterior SVC Inferior: carina Superior: line tangential to sup border of aortic arch Inferior: inferior border of aortic arch Superior: inferior border of aortic arch Anterior: post sternum Posterior: phrenic nerve (laterally) or pericardium (medially) Inferior: diaphragm |

^aRegion and node group boundaries adapted directly from definitions established by IASLC³¹ and AAO-HNS and ASHNS.³²

AAO-HNS, American Academy of Otolaryngology—Head and Neck Surgery; ASHNS, American Society for Head and Neck Surgery; IASLC, International Association for the Study of Lung Cancer; SVC, superior vena cava.

N2 stage – Deep Region

TABLE 3. Deep Region (N2) (Middle Mediastinal and Deep Cervical Nodes)

| Region Boundaries | Node Groups ^{14, 16} | Node Group Boundaries |
|--|---|--|
| Sup: Level of lower border of cricoid cartilage Anteromedial (Neck): Lateral Border of Sternohyoid, Medial Border of Carotid Sheath Posterolateral (Neck): Anterior Border of Trapezius Ant (Chest): Aortic Arch, Aortopulmonary Window – Ant Border of SVC Post (Chest): Esophagus Lat (Chest): Pulmonary Hila Inf: Diaphragm | Lower Jugular (AAO-HNS / ASHNS Level 4) | Sup: Level of lower border of cricoid cartilage Anteromedial: lat border of sternohyoid Posterolateral: lat border of sternocleidomastoid Inf: clavicle |
| | Supraclavicular/Venous Angle: Confluence of Internal Jugular & Subclavian Vein (AAO-HNS / ASHNS Level 5b) | Sup: Level of lower border of cricoid cartilage Anteromedial: post border of sternocleidomastoid Posterolateral: ant border of trapezius Inf: clavicle |
| | Internal Mammary nodes | Proximity to internal mammary arteries |
| | Upper Paratracheal (IASLC Level 2) | Sup: sup border of manubrium, apices of lungs Inf: intersection of lower border of innominate vein with trachea; sup border of aortic arch |
| | Lower Paratracheal (IASLC Level 4) | Sup: intersection of lower border of innominate vein with trachea; sup border of aortic arch Inf: lower border of azygos vein, sup border of left main pulmonary artery |
| | Subaortic / Aortopulmonary Window (IASLC Level 5) | Sup: inf border of aortic arch Inf: sup border of left main pulmonary artery |
| | Subcarinal (IASLC Level 7) | Sup: carina Inf: upper border of lower lobe bronchus on the left; lower border of the bronchus intermedius on the right |
| | Hilar (IASLC Level 10) | Sup: lower rim of azygos vein on right, upper rim of pulmonary artery on left Inf: interlobar region bilaterally |

Region and node group boundaries adapted directly from definitions established by AAO-HNS, ASHNS, and IASLC.

AAO-HNS, American Academy of Otolaryngology—Head and Neck Surgery; ASHNS, American Society for Head and Neck Surgery; IASLC, International Association for the Study of Lung Cancer; sup, superior; ant, anterior; inf, inferior; lat, lateral; post, posterior; SVC, superior vena cava.

Table 2. Deep Region (N2) (Visceral Mediastinum and Deep Cervical Nodes)

| Region Boundaries | Node Groups ^a | Node Group Boundaries |
|--|---|---|
| Superior: level of lower border of cricoid cartilage Anteromedial (neck): lateral border of sternohyoid, medial border of carotid sheath/jugular vein Posterolateral (neck): anterior border of trapezius Anterior (chest): aortic arch, aortopulmonary window-anterior border of the SVC Posterior (chest): esophagus Lateral (chest): pulmonary hila Inferior: diaphragm | Perijugular (AAO-HNS/ASHNS level 4) | Superior: level of lower border of cricoid cartilage Anteromedial: medial border of the jugular vein and carotid artery Posterolateral: lateral border of sternocleidomastoid Inferior: clavicle Superior: level of lower border of cricoid cartilage Anteromedial: posterior border of sternocleidomastoid Posterolateral: anterior border of trapezius Inferior: clavicle Proximity to internal mammary arteries Superior: superior border of manubrium, apices of lungs |
| | Supraclavicular (AAO-HNS/ASHNS level 5b) | Inferior: intersection of lower border of innominate vein with trachea; superior border of aortic arch Superior: intersection of lower border of innominate vein with trachea; superior border of aortic arch Inferior: lower border of azygos vein, superior border of left main pulmonary artery Superior: inferior border of aortic arch Inferior: superior border of left main pulmonary artery Superior: carina Inferior: upper border of lower lobe bronchus on the left; lower border of bronchus intermedius on the right |
| | Internal mammary arteries Upper paratracheal (IASLC level 2) | |
| | Lower paratracheal (IASLC level 4) | |
| | Subaortic/aortopulmonary window (IASLC level 5) | |
| | Subcarinal (IASLC level 7) | |
| | Hilar (IASLC level 10) | |

^aRegion and node group boundaries adapted directly from definitions established by IASLC³¹ and AAO-HNS and ASHNS.³²

AAO-HNS, American Academy of Otolaryngology—Head and Neck Surgery; ASHNS, American Society for Head and Neck Surgery; IASLC, International Association for the Study of Lung Cancer; SVC, superior vena cava.

Boundaries of the anterior (N1) and deep (N2) lymph node levels (8th)

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Node Map for Thymic Malignancy

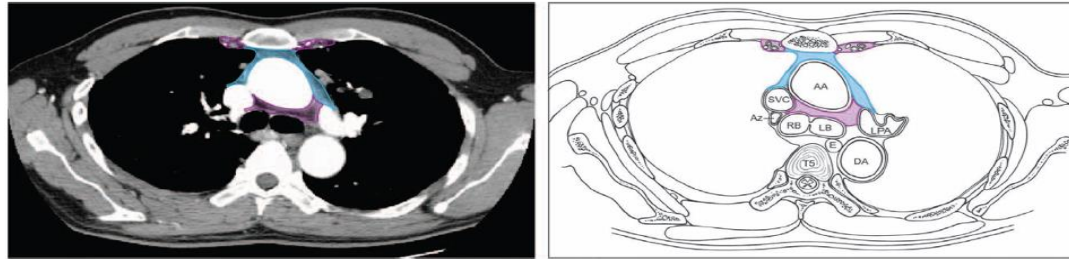


FIGURE 4. Aortopulmonary window level, axial section. Anterior region (blue) and deep region (purple). Note: deep region includes aortopulmonary window nodes. AA, ascending aorta; DA, descending aorta; LPA, left pulmonary artery; SVC, superior vena cava; Az, azygos vein; RB, right main bronchus; LB, left main bronchus.

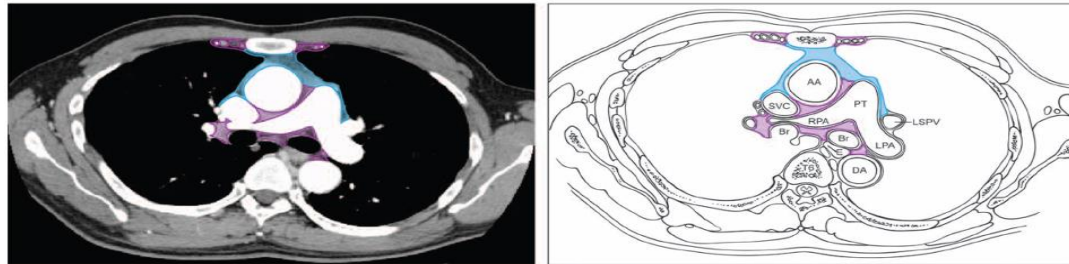


FIGURE 5. Carina level, axial section. Anterior region (blue) and deep region (purple). Note: deep region includes aortopulmonary window nodes. AA, ascending aorta; DA, descending aorta; PT, pulmonary trunk; LPA, left pulmonary artery; RPA, right pulmonary artery; SVC, superior vena cava; LSPV, left superior pulmonary vein; Br, bronchus; E, esophagus.

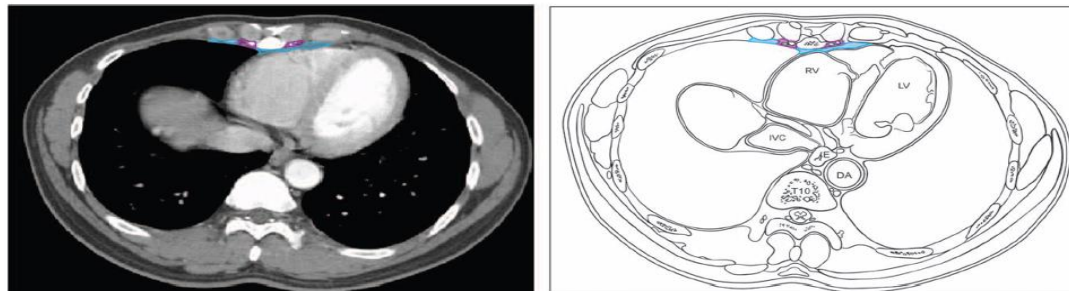


FIGURE 6. Diaphragm level, axial section. Anterior region (blue) and deep region (purple). RV, right ventricle; LV, left ventricle; IVC, inferior vena cava; DA, descending aorta; E, esophagus.

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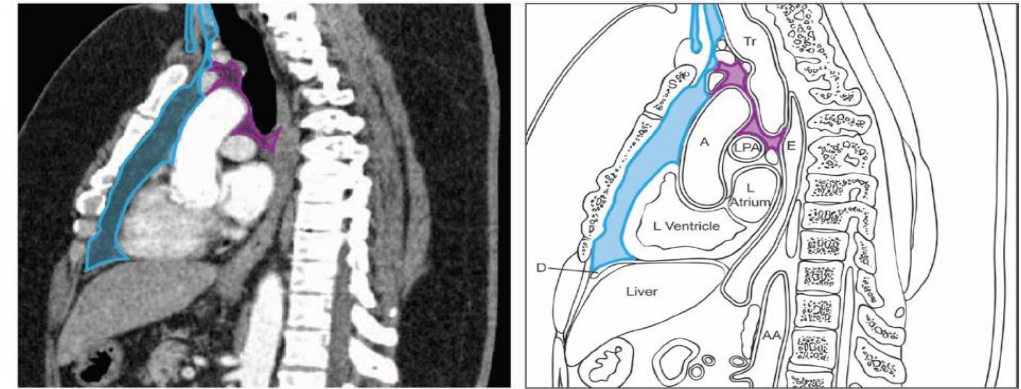


FIGURE 1. Mediastinum, sagittal section. Anterior region (blue) and deep region (purple). Tr, trachea; E, esophagus; LPA, left pulmonary artery; A, aorta; D, diaphragm.

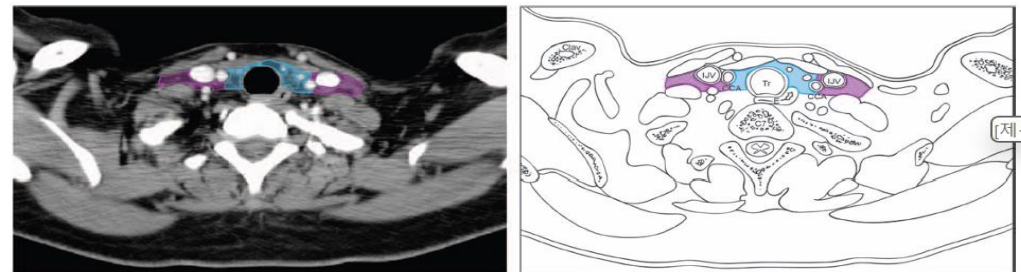


FIGURE 2. Thoracic inlet, axial section. Anterior region (blue) and deep region (purple). CCA, common carotid artery; IJV, internal jugular vein; Tr, trachea; Clav, clavicle; E, esophagus.

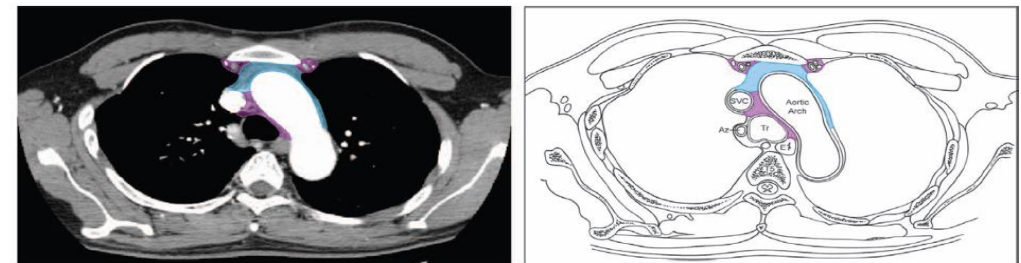


FIGURE 3. Paraaortic level, axial section. Anterior region (blue) and deep region (purple). SVC, superior vena cava; E, esophagus; Tr, trachea.

Boundaries of the anterior (N1) and deep (N2) lymph node levels (9th)

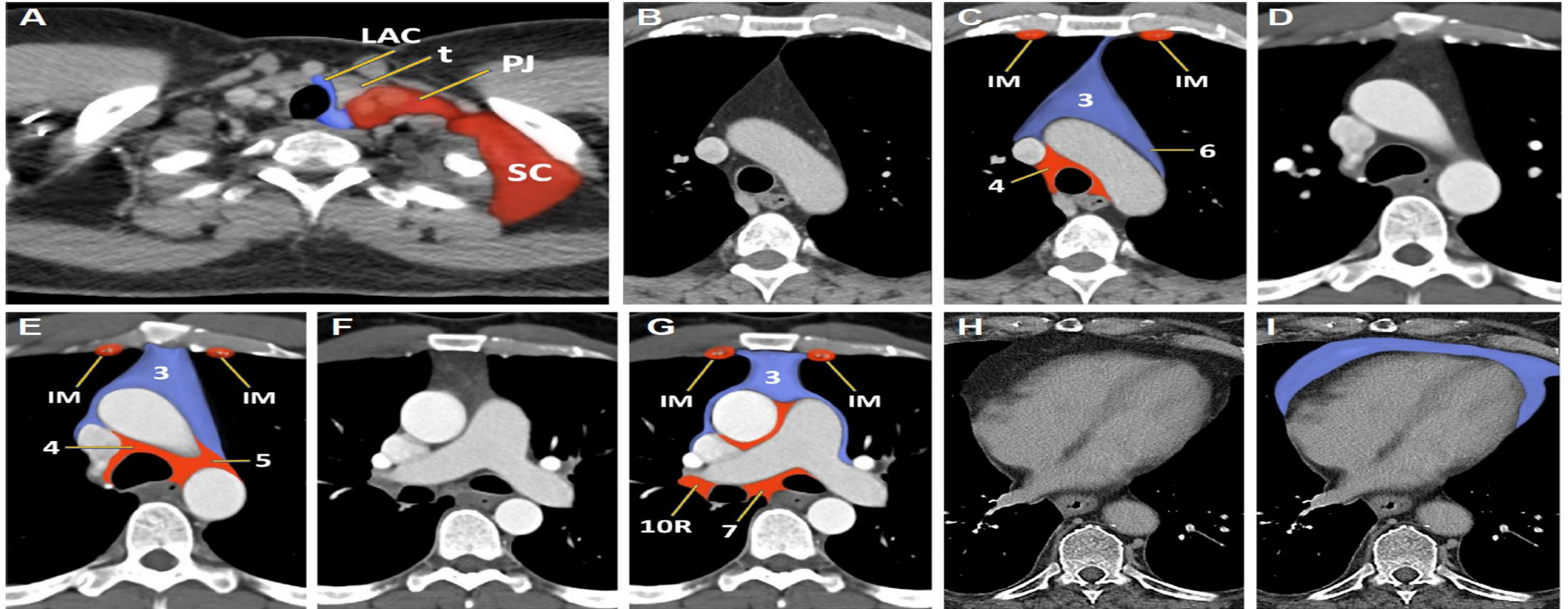


Figure 2. Native and annotated axial computed tomography images revealing the node groups, marking the boundaries of the anterior (N1) and deep (N2) lymph node levels at the levels of the (A) lower neck, (B, C) aortic arch, (D, E) aorto-pulmonary window, (F, G) main pulmonary artery, (H, I) and base of the heart. Boundaries of the anterior (N1) and deep (N2) level are shaded in blue and red, respectively. IASLC, International Association for the Study of Lung Cancer; IM, internal mammary; LAC, low anterior cervical; PJ, perijugular; SC, supraclavicular; t, thyroid gland. Numbers refer to IASLC node map used for lung cancer.³¹ Courtesy of the International Association for the Study of Lung Cancer. Copyright 2023, Aletta Ann Frazier.

Summary of 9th TNM staging system

Table 1. Proposed T Categories for the Ninth Edition of the TNM Stage Classification

| T | Description |
|----|---|
| T1 | A tumor that is limited to the thymus with or without encapsulation or directly invades into the mediastinum alone or directly invades the mediastinal pleura but does not involve any other mediastinal structure T1a: 5 cm or less in its greatest dimension ^a T1b: larger than 5 cm in its greatest dimension ^a (Level 1 structures—thymus, anterior mediastinal fat, mediastinal pleura) |
| T2 | Tumor directly invades the pericardium (either partial or full-thickness), the lung, or the phrenic nerve (Level 2 structures—pericardium, lung, phrenic nerve) |
| T3 | Tumor directly invades any of the following: (1) brachiocephalic vein, (2) superior vena cava, (3) chest wall, or (4) extrapericardial pulmonary arteries or veins (Level 3 structures—brachiocephalic vein, SVC, chest wall, hilar pulmonary vessels) |
| T4 | Tumor directly invades any of the following: (1) aorta (ascending, arch, or descending), (2) arch vessels, (3) intrapericardial pulmonary artery or veins, (4) myocardium, (5) trachea, or (6) esophagus (Level 4 structures—aorta [ascending, arch, or descending], arch vessels, intrapericardial pulmonary artery or veins, myocardium, trachea, esophagus) |

Note: T categories are defined by “levels” of invasion; they reflect the highest degree of invasion regardless of how many other (lower level) structures are invaded.

^aIrrespective of mediastinal pleura invasion. Mediastinal pleura invasion to be recorded as “Additional histologic descriptor.”

SVC, superior vena cava.

Table 1. Proposals for the N and the M Components for the Ninth Edition of the TNM Classification

Descriptors

N0
N1
N2
M0
M1
a
b

- Tumor size** instead of capsular invasion

: T1a ≤ 5cm , T1b >5cm

- Dropped for staging - Encapsulation or not/mediastinal fat or**

pleural invasion (Recorded as an additional histologic descriptor)

- T3 structure (lung, phrenic nerve) -> T2 category**

- N component -> No change**

- M component -> No change**

Note: Involvement must be pathologically proven in pathologic staging.

- Stage Groups -> No change**

Table 3. Proposed Stage Groups of Thymic Tumors for the Ninth Edition of the TNM Classification of Malignant Tumors

| Stage | T | N | M |
|-------|-------|-------|---------|
| I | T1a,b | N0 | M0 |
| II | T2 | N0 | M0 |
| IIIA | T3 | N0 | M0 |
| IVB | T4 | N0 | M0 |
| IVA | T any | N1 | M0 |
| IVB | T any | N0,N1 | M1a |
| IVB | T any | N2 | M0, M1a |
| IVB | T any | N any | M1b |

Note: Any invasion must be histologically confirmed for the pathologic stage.

Limitations and Conclusion

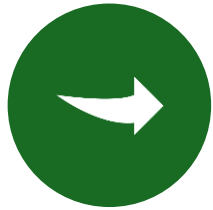
1. Limited Prospective Data
2. Nodal Involvement
3. Imaging and Surgical Correlation
4. Rare Tumor Representation
5. Size and Invasion Clarification

Addressing these limitations requires continued data collection, particularly prospective studies, and further refinement of imaging techniques and surgical practices to enhance the accuracy and applicability of the TNM staging system for thymic epithelial tumors.

9th TNM Staging Proposal for Thymic Epithelial Tumors



1. PRECISE STAGING



2. SUBDIVISION OF
NODAL INVOLVEMENT



3. DATA-DRIVEN
APPROACH



4. GLOBAL
STANDARDIZATION



5. IMPROVED
PROGNOSTIC
ACCURACY



6. PROMOTION OF
MULTIDISCIPLINARY
APPROACH



Pusan National University
Yangsan Hospital



PNUH 양산부산대학교병원
Pusan National University Yangsan Hospital

경청해 주셔서
감사합니다.

Thank you for your
attention