Mediastinal Evaluation in Early Lung Cancer

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'Early' Lung Cancer

| | | No | N1 | N2 | N3 | |
|---|-----|------|------|------|------|--|
| / | T1 | IA | IIB | IIIA | IIIB | |
| | T2a | IB | IIB | IIIA | IIIB | |
| | T2b | IIA | IIB | IIIA | IIIB | |
| | Т3 | IIB | IIIA | IIIB | IIIC | |
| | T4 | IIIA | IIIA | IIIB | IIIC | |
| | M1a | IVA | IVA | IVA | IVA | |
| | M1b | IVA | IVA | IVA | IVA | |
| | M1c | IVB | IVB | IVB | IVB | |

Aim of mediastinal staging:

To detect radiologically occult N2/N3

Lung Cancer Staging

Tissue Diagnosis

MRI Brain

PET CECT





Why stage the mediastinum?

Stage Distribution and 5-year Relative Survival by Stage at Diagnosis for 2001-2007, All Races, Both Sexes

| Stage at Diagnosis | Stage Distribution (%) | 5-year Relative Survival (| |
|--|---------------------------|-------------------------------|--|
| Localized (confined to primary site) | 15 | 5 | |
| Regional (spread to regional lymphnodes) | 22 | 24 | |
| Distant (cancer has metastasized) | 56 | | |
| Unknown (unstaged) | 7 | 1 | |

- Nodal status determines treatment opt
- Significantly affects survival
- 12.5% unforeseen N2 despite staging*

* Bousema et al. Lung Cancer Feb 2020



except:

When can we avoid it?

Invasive mediastinal staging is recommended in ALL patients of NSCLC

- **Peripheral** (outer 1/3 of lung fields)
 - AND
 - T1 (</= 3 cm)
 - AND
- **Node negative** at hilum and mediastinum on PET scan

Why is PET CT not enough?

False Positive

- 18% in our series
- Country endemic for TB, granulomatous disease
- Upstages 1 in 5: may deny curative surgery

- Understages

False Negative

• Usually due to 'shine through' of organs with high uptake

Risk of inadequate treatment



Invasive mediastinal staging

EBUS +/- EUS-B

- Less 'invasive'
- Under conscious sedation
- Recommended initial step
- Rapid on-site examination
- Negligible complications

Mediastinoscopy

- Surgical procedure under GA
- Standard / video (Gold Standard)
- False negative: 2%
- Complications: 6%



Nodal stations addressed

- EBUS: 2,4,7,10,11
- EUS-B: 8,9 (liver, adrenal mets)
- M'scopy: 2,4,7,5
- Blind spots: 5,6



The genesis of EBUS

A prospective controlled trial of endobronchial ultrasound-guided transbronchial needle aspiration compared with mediastinoscopy for mediastinal lymph node staging of lung cancer

Kazuhiro Yasufuku, MD, PbD,^a Andrew Pierre, MD, MSc,^a Gail Darling Thomas Waddell, MD, PhD,^a Michael Johnston, MD,^a Gilda da Cunha William Geddie, MD,^b Scott Boerner, MD,^b Lisa W. Le, MSc,^c and Sha

Mediastinoscopy vs Endosonography for Mediastinal Nodal Staging of Lung Cancer A Randomized Trial

Jouke T. Annema, MD, PhD Jan P. van Meerbeeck, MD, PhD Robert C. Rintoul, FRCP, PhD Christophe Dooms, MD, PhD

Context Mediastinal nodal staging is recommended for patients with resectable nonsmall cell lung cancer (NSCLC). Surgical staging has limitations, which results in the performance of unnecessary thoracotomies. Current guidelines acknowledge minimally invasive endosonography followed by surgical staging (if no nodal metastases are found by endosonography) as an alternative to immediate surgical staging. **ORIGINAL ARTICLE**

Endobronchial Ultrasound versus Mediastinoscopy for Mediastinal Nodal Staging of Non–Small-Cell Lung Cancer

Sang-Won Um, MD, PhD, * Hong Kwan Kim, MD, PhD, † Sin-Ho Jung, PhD, ‡ Joungho Han, MD, PhD, § 1D, * Hye Yun Park, MD, PhD, * Yong Soo Choi, MD, PhD, † Young Mog Shim, MD, PhD, † -Ju Ahn, MD, PhD, || Keunchil Park, MD, PhD, || Yong Chan Ahn, MD, PhD, ¶ ng Choi, MD, PhD, # Kyung Soo Lee, MD, PhD, ** Gee Young Suh, MD, PhD, * yo Chung, MD, PhD, * O Jung Kwon, MD, PhD, * Jhingook Kim, MD, PhD, † and Hojoong Kim, MD, PhD*

All surgeons!

2004



Impressive results!

| | Yasufuku | Won-Um | Mediastinoscopy |
|-------------------|----------|--------|-----------------|
| Sensitivity (%) | 81 | 88 | 81 |
| NPV | 91 | 85 | 78 |
| Accuracy (%) | 93 | 92 | 89 |
| Specificity(100%) | 100 | 100 | 100 |
| PPV | 100 | 100 | 100 |

Are they reproducible?

- Operator dependent, learning curve
- Single centre experiences: False -ve upto 20%!
- High price to pay

- A randomized trial comparing endosonography followed by surgical staging versus surgical mediastinal staging alone in non-small cell lung cancer: The ASTER study.
- K. G. Tournoy, C. A. Dooms, R. C. Rintoul, P. Deleyn, A. G. Nicholson, E. Deschepper...
 - Sensitivity surgical staging alone: 79%
 - Sensitivity EBUS/EUS-B + SS: 94%!
 - Futile thoracotomies: 18% v/s 7%



ESTS 2014 Guideline



EBUS today

- Systematic nodal sampling > selective strategy: SCORE study
- Improving false negative rates: inching closer to M'scopy
- Sensitivity and specificity: 88% and 100%!
- Almost NO complications

Confirmatory M'scopy mandatory?

Why not?

- Limited additional nodal met detection
- Increased morbidity
- Treatment delay and cost
- Micro-met N2: questionable impact on survival

Bousema et al. meta-analysis

- Unforeseen N2 rate: 9.6% v/s 9.9%
- 6% morbidity: RLN palsy, chyle leak
- Unforeseen N2: no impact on survival

MEDIASTrial



Value addition with VAMLA

- Assessing 'resectability' of nodes
- Good clearance for left sided tumours
- Acceptable morbidity: RLN palsy: 2.4%
- The surgeon is enough!
- Can be timed with the resection of primary: if frozen section available



How perfect is perfect?

EBUS/EUS-B + M'scopy:

But...

Sensitivity: 94%!

- We still got it wrong in 9.6%
- 7% still underwent futile thoracotomies
- Cost and time!
- Infrastructure



How feasible is perfect?

Adherence to the mediastinal staging guideline and unforeseen N2 disease in patients with resectable non-small cell lung cancer: Nationwide results from the Dutch Lung Cancer Audit - Surgery

Jelle E. Bousema^a, David J. Heineman^b, Marcel G.W. Dijkgraaf^c, Jouke T. Annema^d, Frank J.C. van den Broek^{a,*}

- EBUS/EUS as initial staging: 43%
- M'scopy as initial staging: 10%
- Confirmatory M'scopy in -ve EBUS: 44%
- Strict invasive mediastinal staging: 19%!

Denmark!

The balancing act



High chance of nodes coming positive

Mediastinoscopy

High chance of nodes coming negative



In conclusion

Nodal status significantly impacts treatment options and prognosis

What you see is not always what you get: invasive staging a must!

• If it's difficult to be perfect, be wise



