

Surgery in Early stage NSCLC

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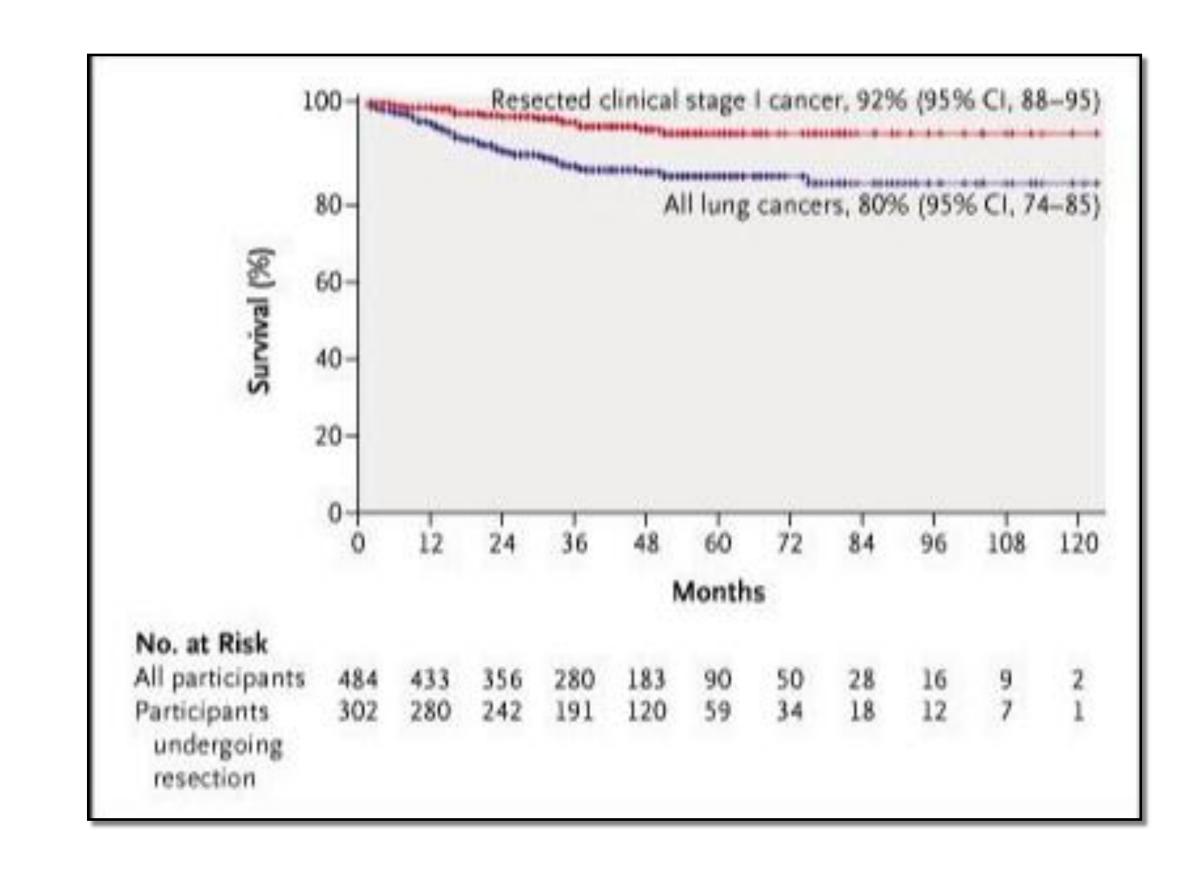
Points for discussion

- Why surgery
- Approach- MIS versus thoracotomy
- Lobectomy versus segmentectomy
- Systemic nodal dissection versus sampling
- Postoperative outcome
- Role of adding systemic therapy to surgery

Why surgery in early stage

- Early stage
 - -stage I and II and select IIIA
 - -T1-3/N0-1

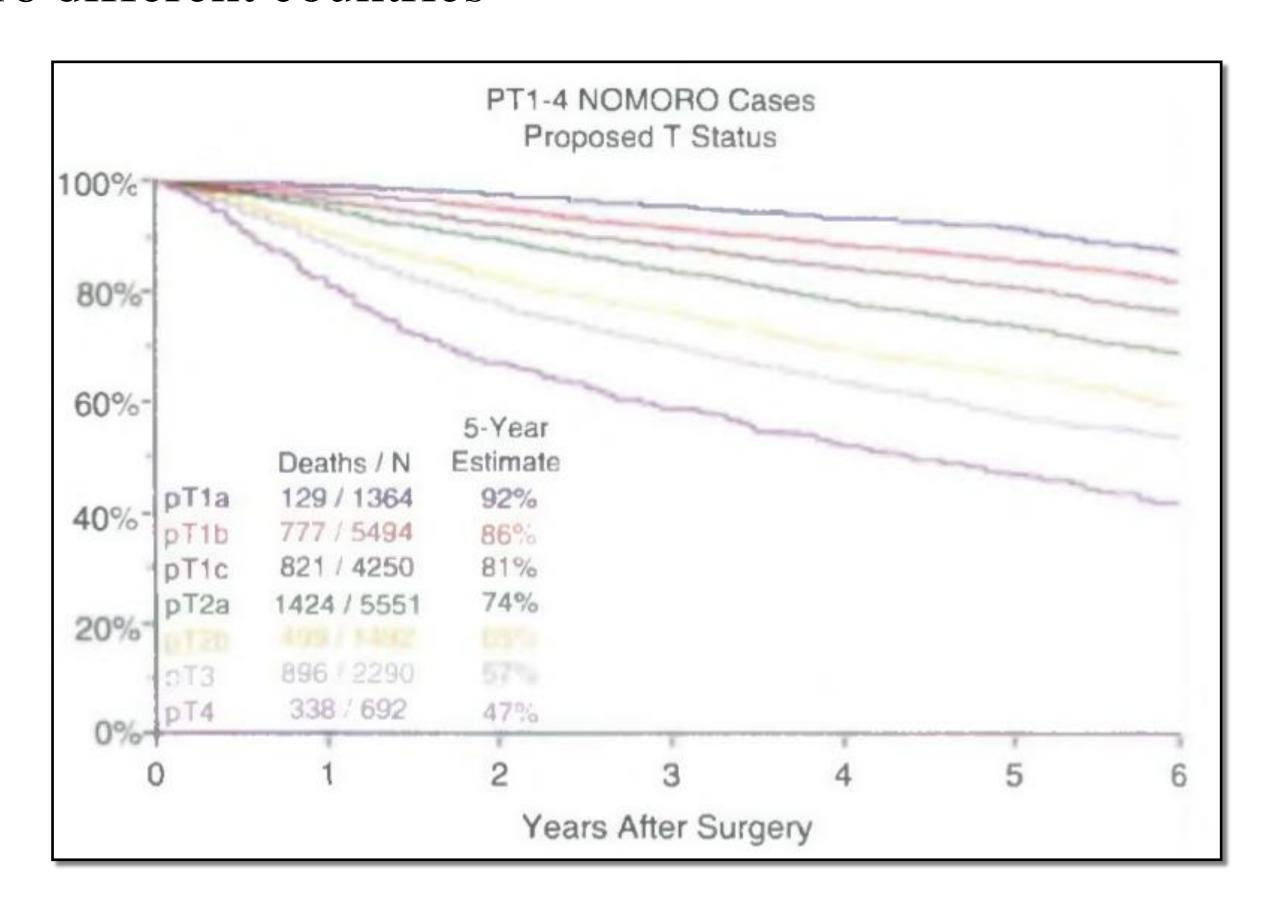
- Outcome of surgery
- I- ELCAP, NEJM, 2006
 - -10 yr OS- 92%



Outcome

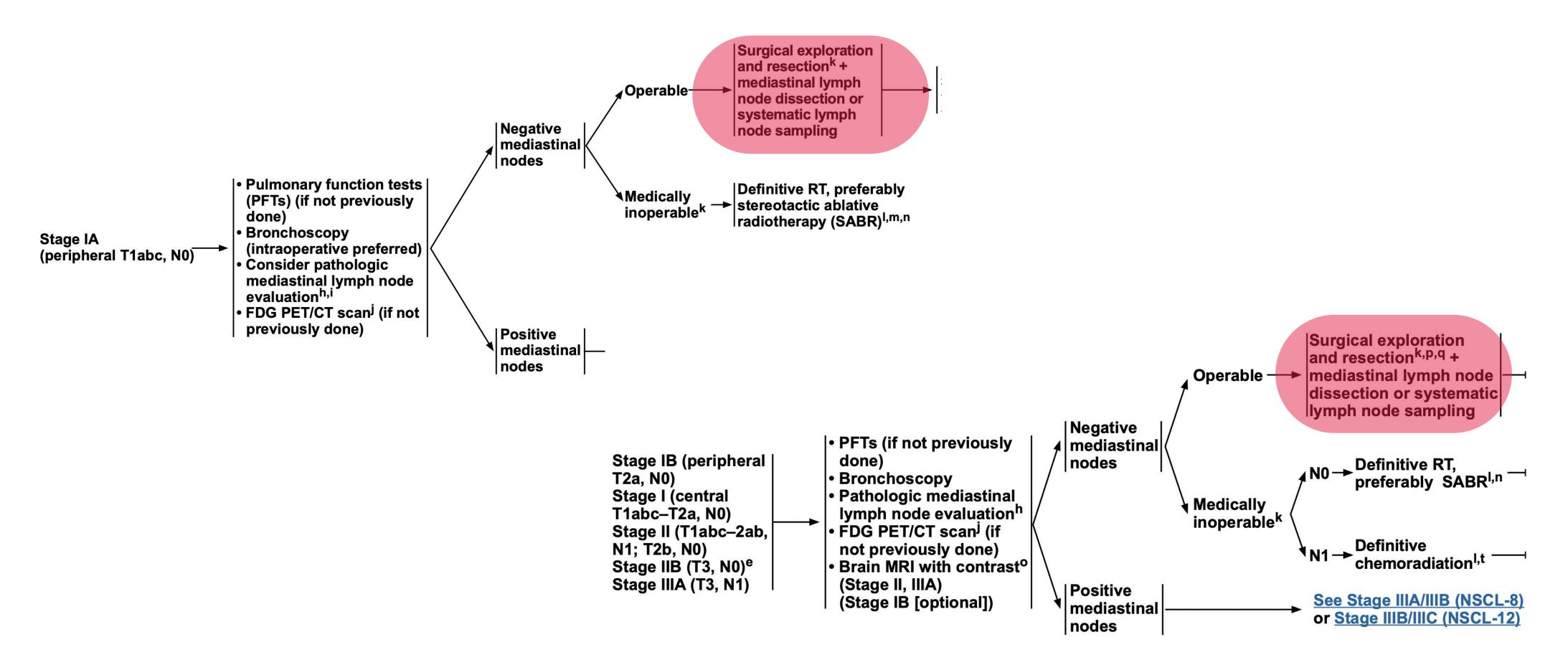
• AJCC 8th edition- 35 databases from 16 different countries

Early stage lung cancer-Surgery has excellent outcome.



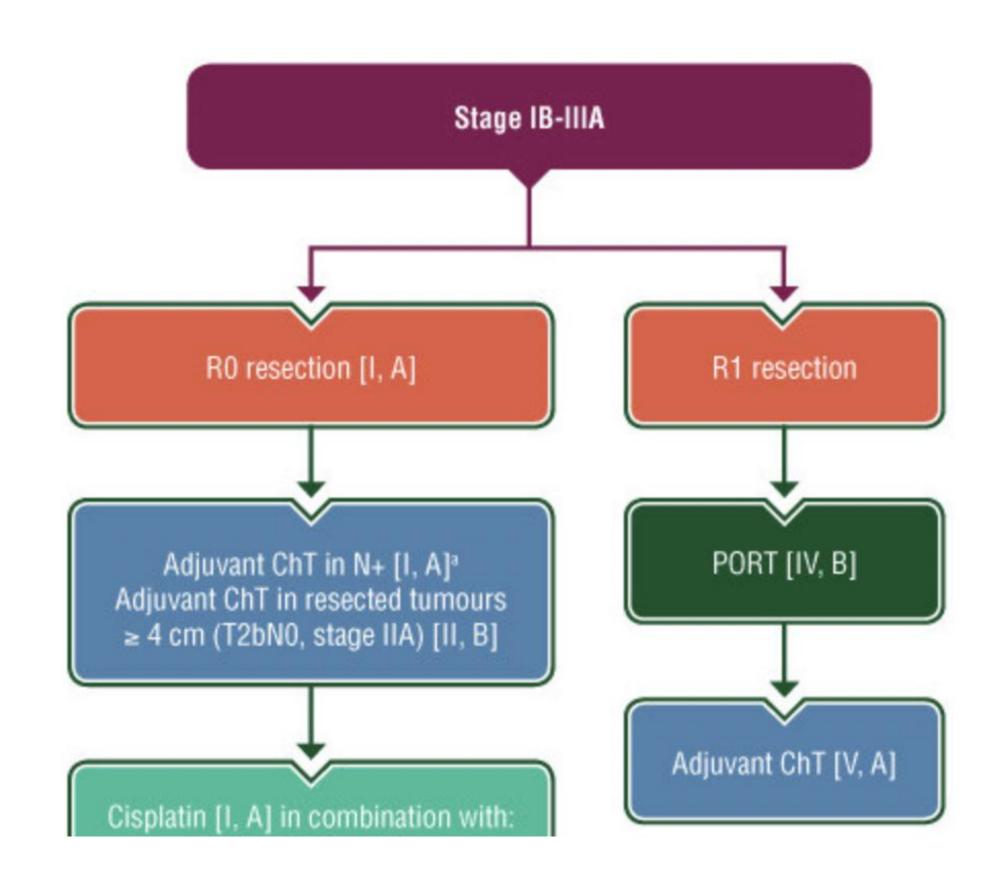
Recommendation

NCCN

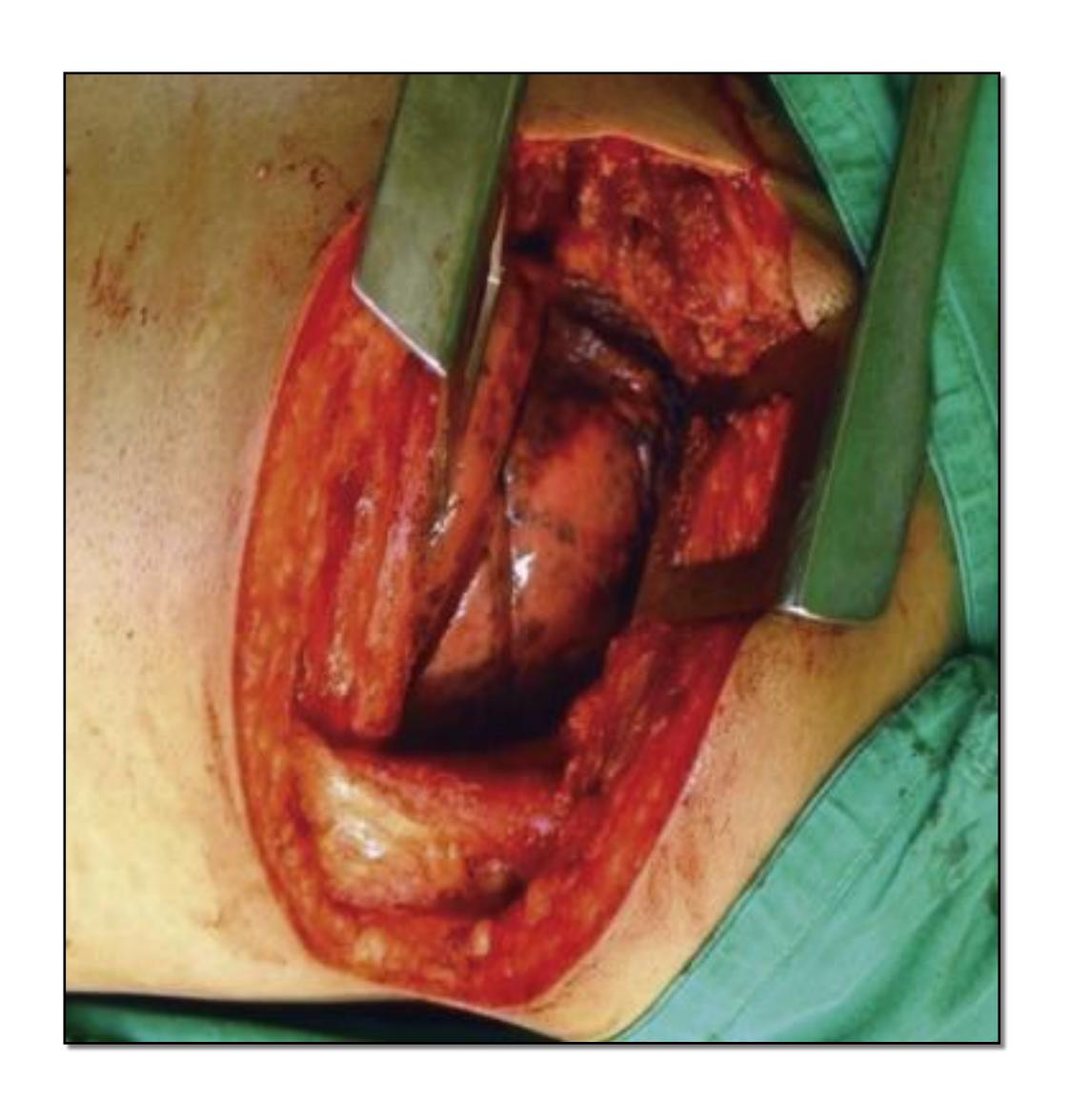


Recommendation

ESMO



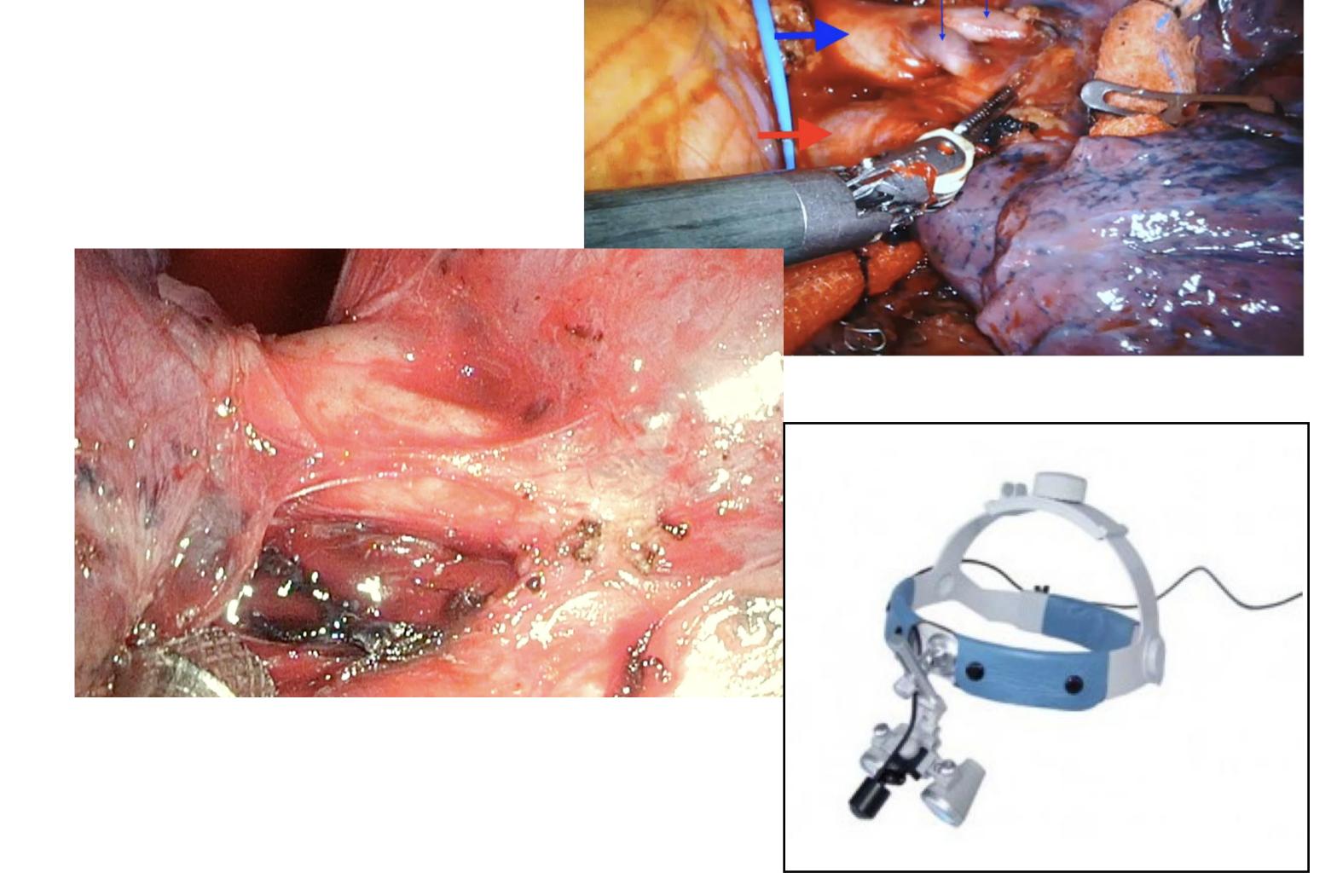
Which type- VATS versus Open





Purported Benefits

- Less pain, faster recovery
- Less complications
- Better quality of life
- Overall reduced cost
- Oncological equivalence



VATS v/s Open: Meta-analyses

	N	Mortality	Morbidity	Oncological outcomes
C Cao, 2013	3634			
Cai Y-x, 2013, China				
Zhang Z, 2013, China	5389			
Wang Z, 2019	1514			

Randomised Evidence

Bendixen et al. 2016

N = 206, single centre

- •Decreased procedure time, blood loss
- •Less epidural analgesia duration
- Cost effective
- •Similar complications

Long et al. 2018

N = 425, multi centre

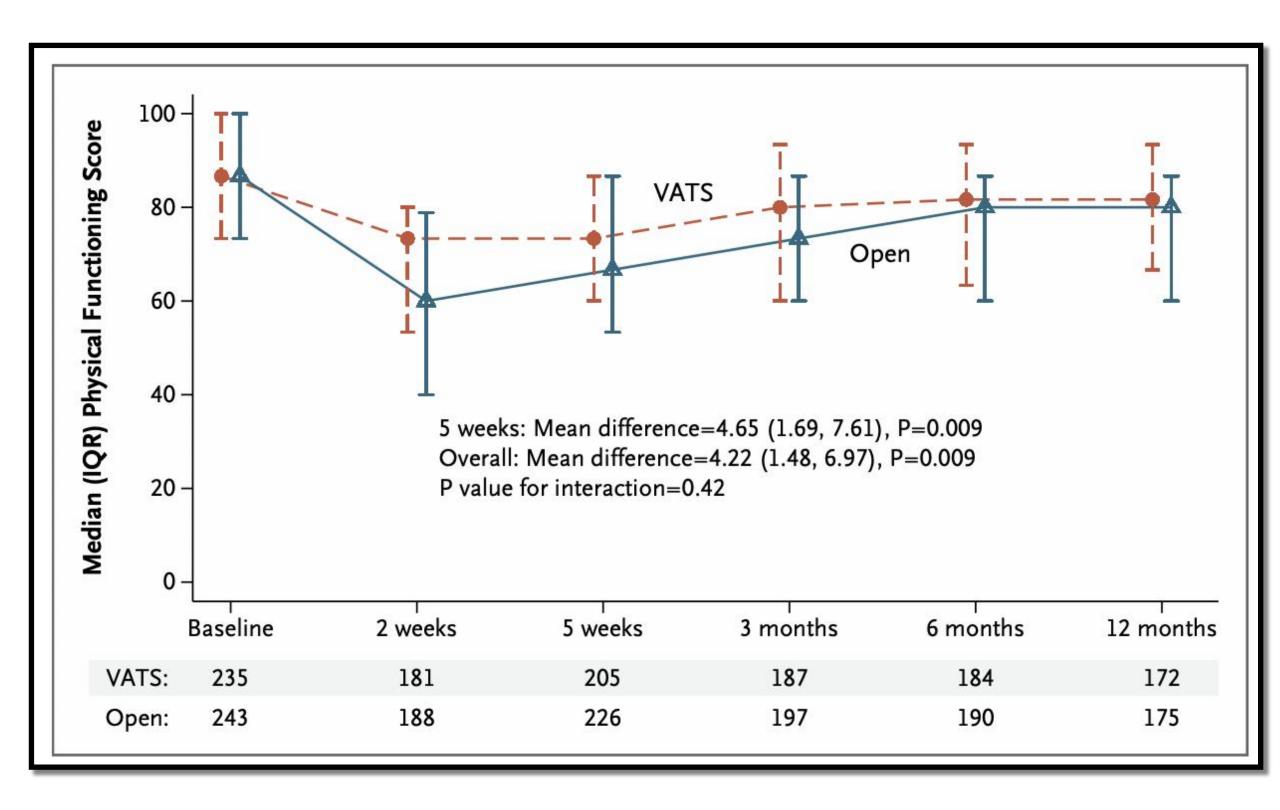
- •Decreased procedure time, blood loss
- •Equivalent nodal yield, R0 rates
- •Similar complications

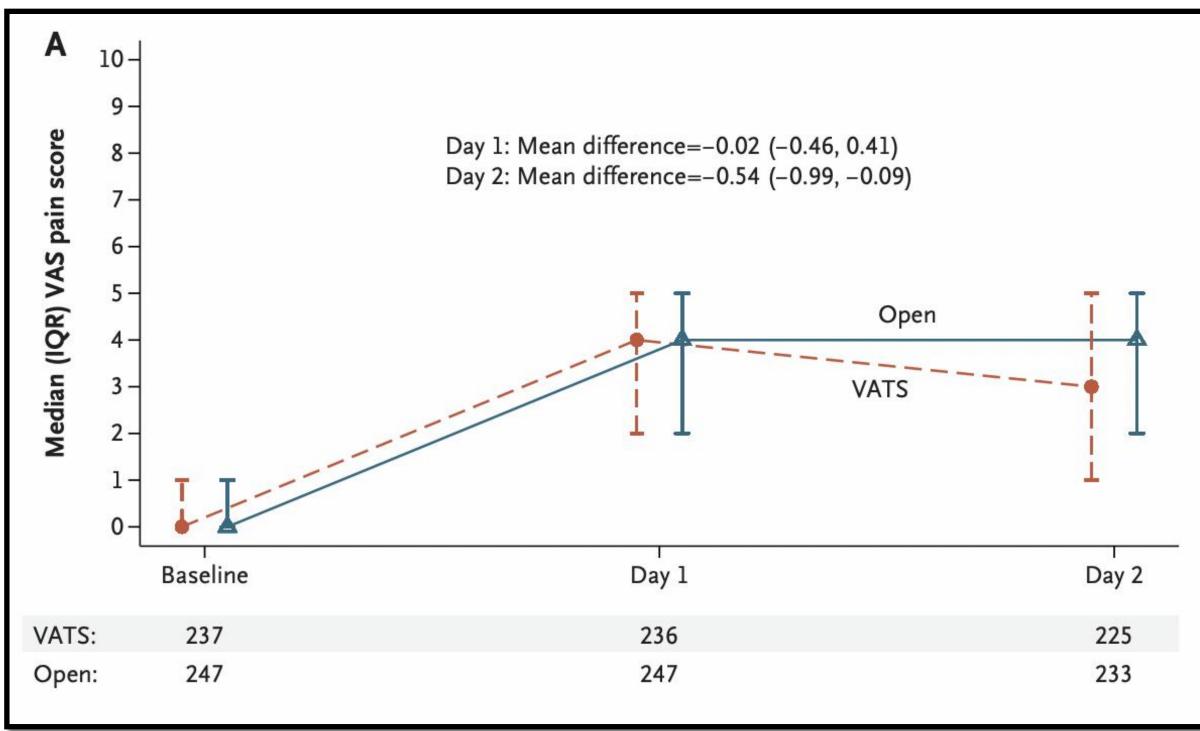
- Poor capture of QoL data
- •No oncological outcomes reported

- No comment on post op pain
- Survival results awaited

VIOLET- NEJM Jan 2022

Results





Global Trends

STS Lung Cancer Resection Risk Model: Higher Quality Data and Superior Outcomes

Felix G. Fernandez, MD MSc,¹ Andrzej S. Kosinski, PhD,² William Burfeind, MD,³ Bernard Park, MD,⁴ Malcolm M. DeCamp, MD,⁵ Christopher Seder, MD,⁶ Blair Marshall, MD,⁷ Mitchell J. Magee, MD,⁸ Cameron D. Wright, MD,⁹ and Benjamin D Kozower, MD MPH¹⁰

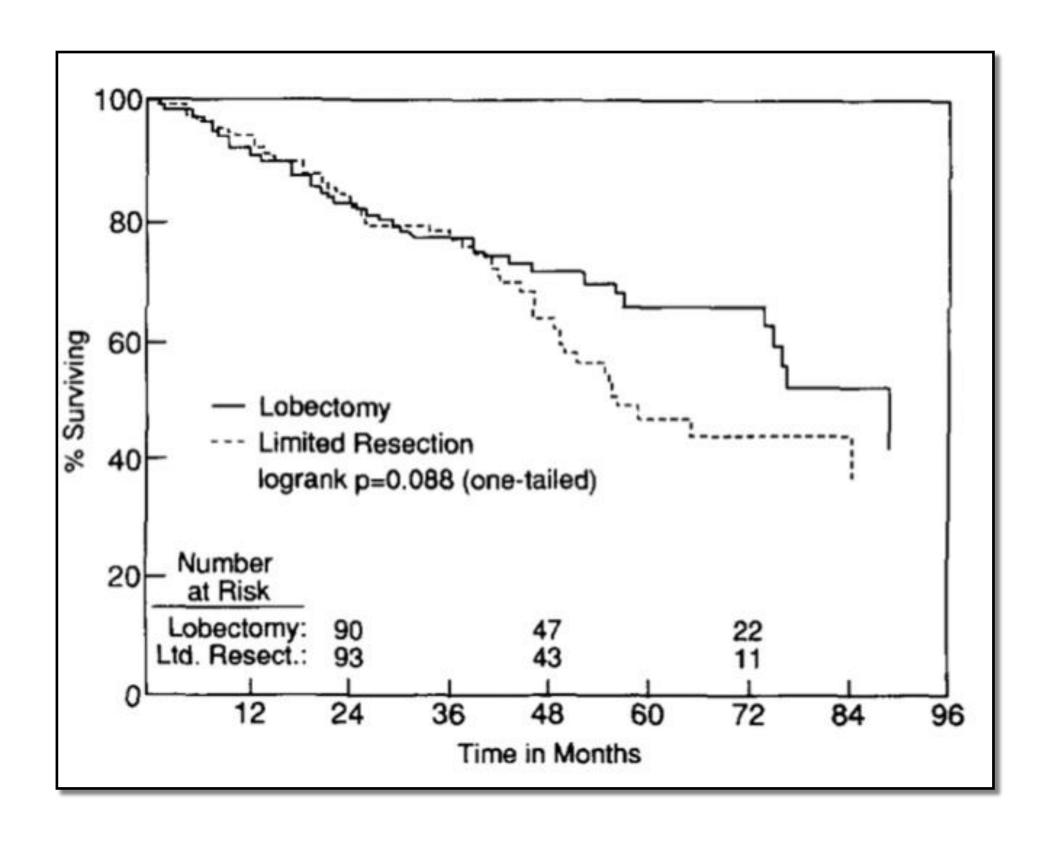
- January 2012 to December 2014
- 27,844 resections for lung cancer
- 62% VATS resections
- Significantly lower morbidity and mortality in MIS group

VATS preferred..

What is adequate for local tumour

Lobectomy versus Segmentectomy

- Lobectomy- standard of treatment
- Randomised trial Ginsberg et. Al., 1995, LCSG
- T1N0, 276 patients
- Patients undergoing limited resections had
 - •75% increased recurrence
 - •Local recurrence tripled
 - •30% increased overall deaths
 - •50% increased cancer deaths



Lobectomy versus sublobar resection

RCT by Nasser K Altorki, Lancet 2018

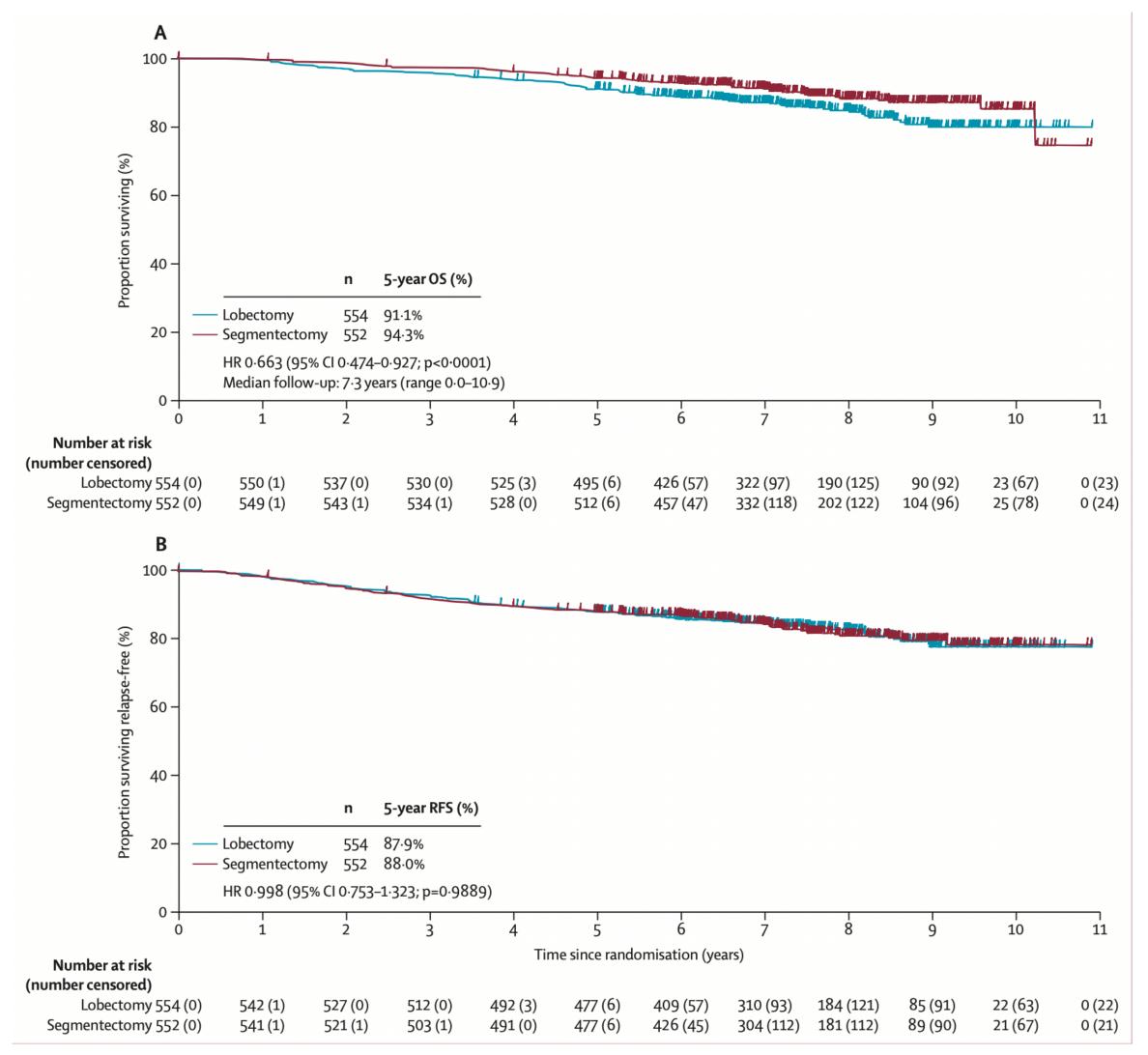
• 697 patients randomised

• Grade 3/4/5 AEs- slightly more frequent with segmentectomy (18.6%), compared to lobectomy (15.6%) and wedge resections (11%)- but no statistical significant difference

• More of air leak in segmentectomy group

Lobectomy versus Segmentectomy

JCOG 0802, Lancet, April 2022



- •Improved survival in segmentectomy group
- •At 1 year follow up- PFT improvement of 3.5%, statistically significant but less than expected (10%)
 - •Local recurrence double in segmentectomy group (11% versus 5%)

Lobectomy versus Segmentectomy

• Evidence is so far only for tumours ≤ 2 cm, N0

• All segmentectomies are not the same- technically challenging

More air leak

Still Lobectomy is standard..

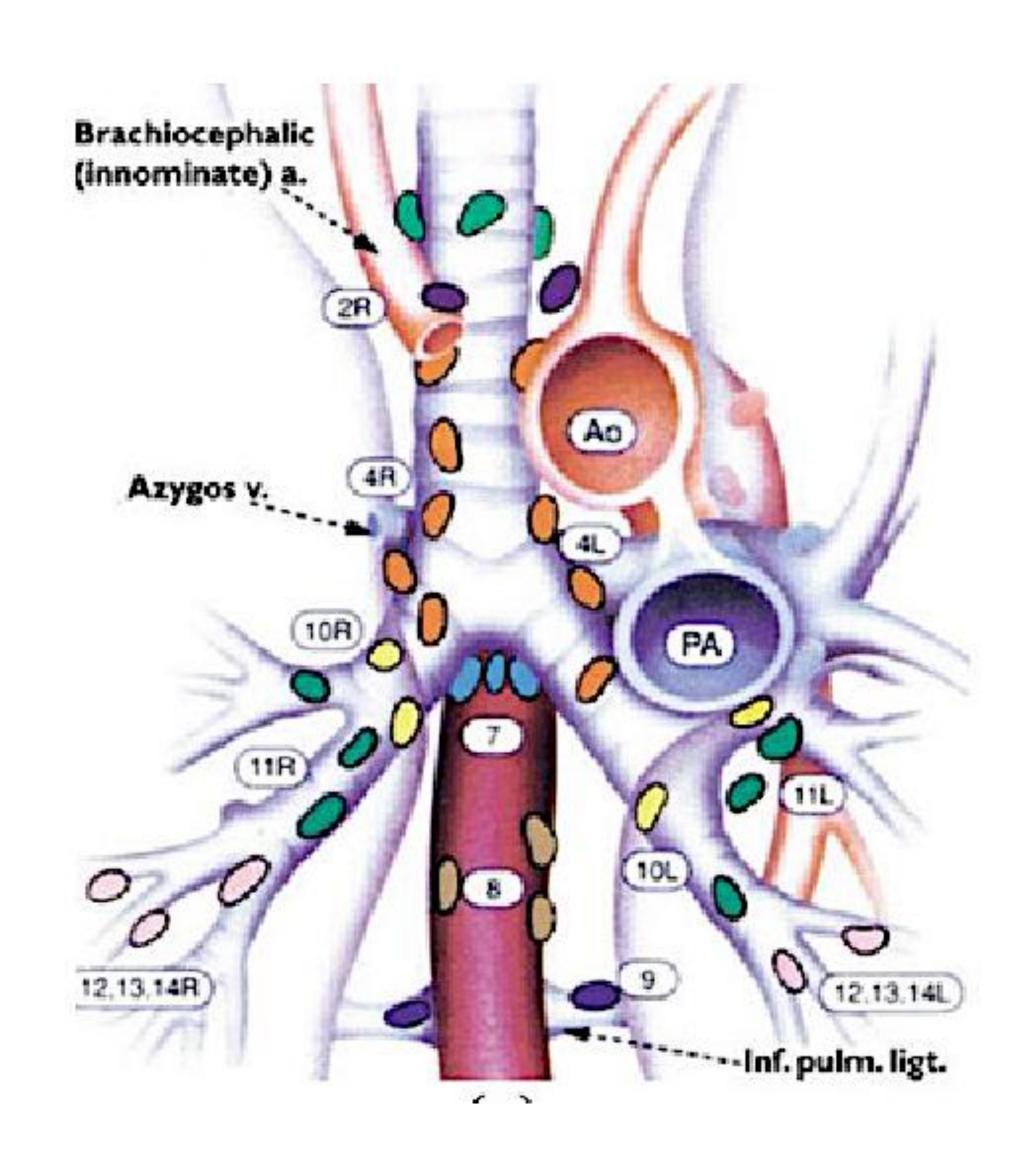
What is adequate for nodal burden

SMLND versus SMLNS

Systematic lymphadenectomy versus sampling of ipsilateral mediastinal lymph-nodes during lobectomy for non-small-cell lung cancer: a systematic review of randomized trials and a meta-analysis

١	MLN		MLN		. 2227271270	Odds Ratio			Odds	Ratio	
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI	Year		M-H, Rand	lom, 95% CI	
Sugi et al	0	59	0	56	7-02-000	Not estimable	1998				
Izbicki et al	2	76	4	93	23.6%	0.60 [0.11, 3.38]	1998		-	_	
Wu et al	1	240	0	231	6.8%	2.90 [0.12, 71.55]	2002		_	-	_
ACOSOG Z0031 trial	4	525	10	498	51.6%	0.37 [0.12, 1.20]	2011			+	
Zhang et al	2	95	2	107	17.9%	1.13 [0.16, 8.18]	2013			-	
Total (95% CI)		995		985	100.0%	0.59 [0.25, 1.36]			•	-	
Total events	9		16								
Heterogeneity: Tau ² =	0.00; Chi ²	= 1.94,	df = 3 (P	= 0.58); I ² = 0%			-		1 10	400
Test for overall effect: 2			The state of the s		100			0.01	0.1 Favours MLND	1 10 Favours MLNS	100

Mokhles et.al. EJCTS, 2017 SMLND- no increase in operative mortality Better clearance and prognostication



Postoperative outcome- mortality and morbidity

- ACOSOG Z0030, 2011
- More than 1000 patientsopen lobectomy (93%) with SMLND/SMLNS
- Mortality- 1.37%
- Morbidity- 38%

- JCOG0802, 2022
- More than 1000 patients,
 VATS (90%)
 lobectomy/segmentectomy
- Mortality 0%
- Morbidity- 26%

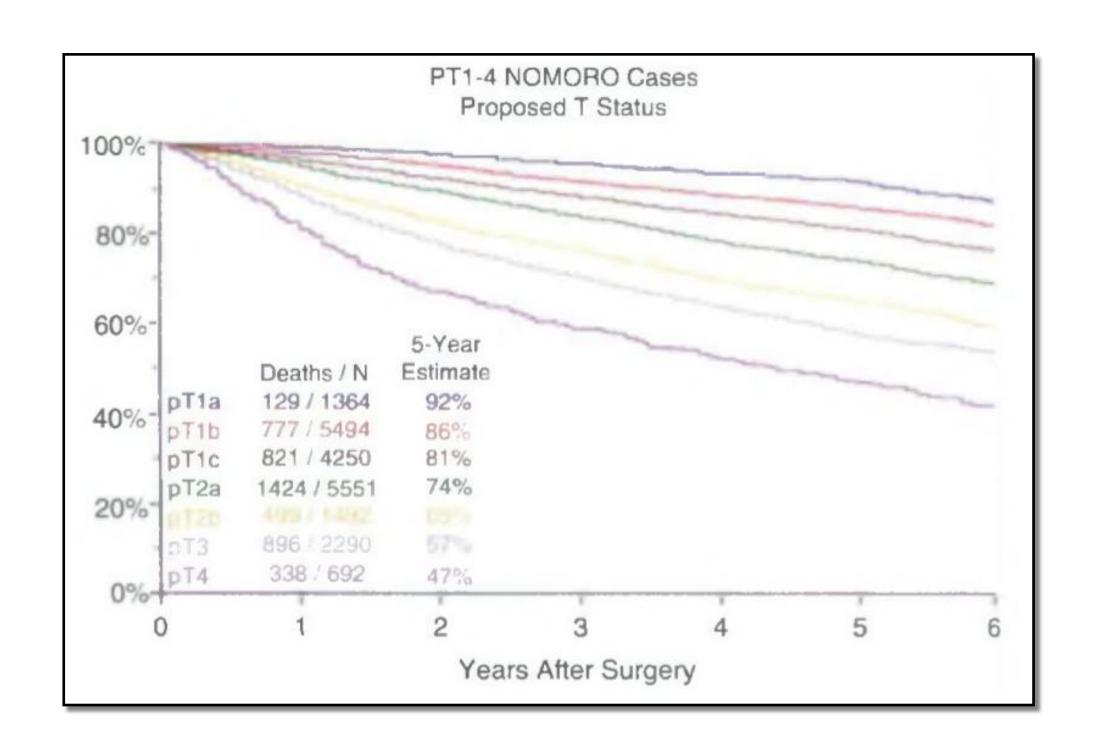
TMH data

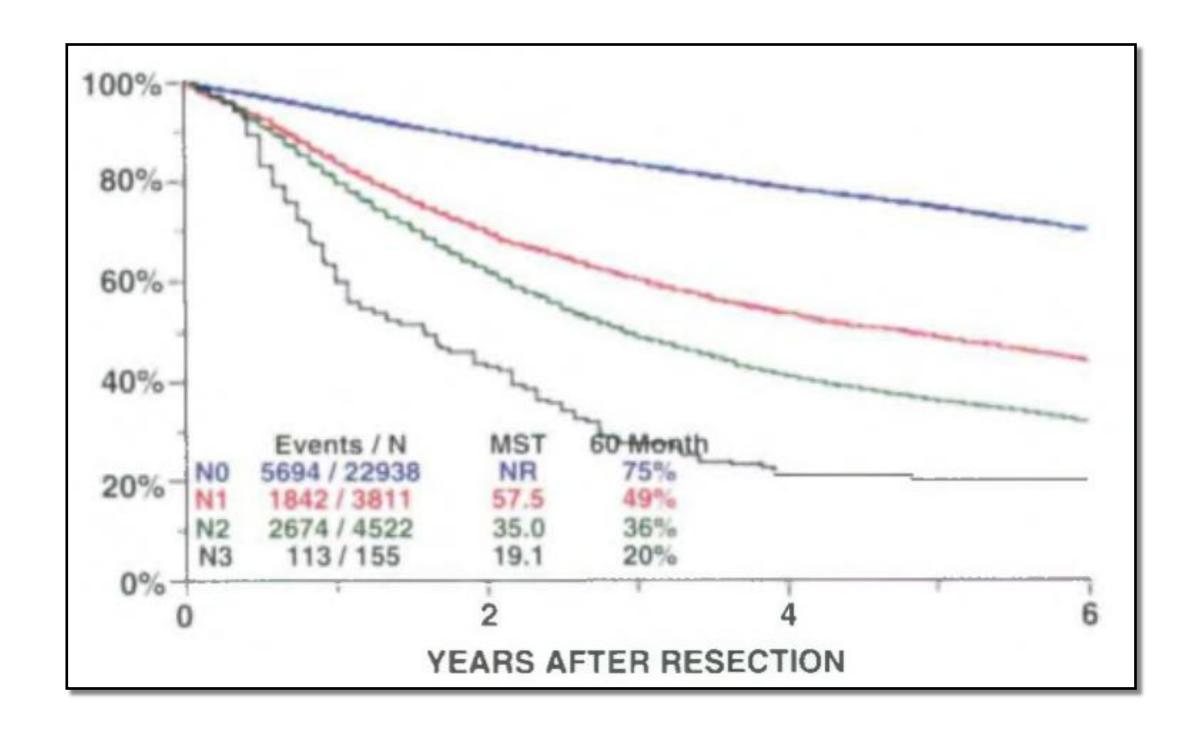
uVATS anatomical lung resection

- May 2017 to Feb 2021, n=106
- Median length of hospital stay- 4 days
- Post operative mortality- 0.9%
- Morbidity (CD IIIa and above)- 13.2%

Results of Surgery

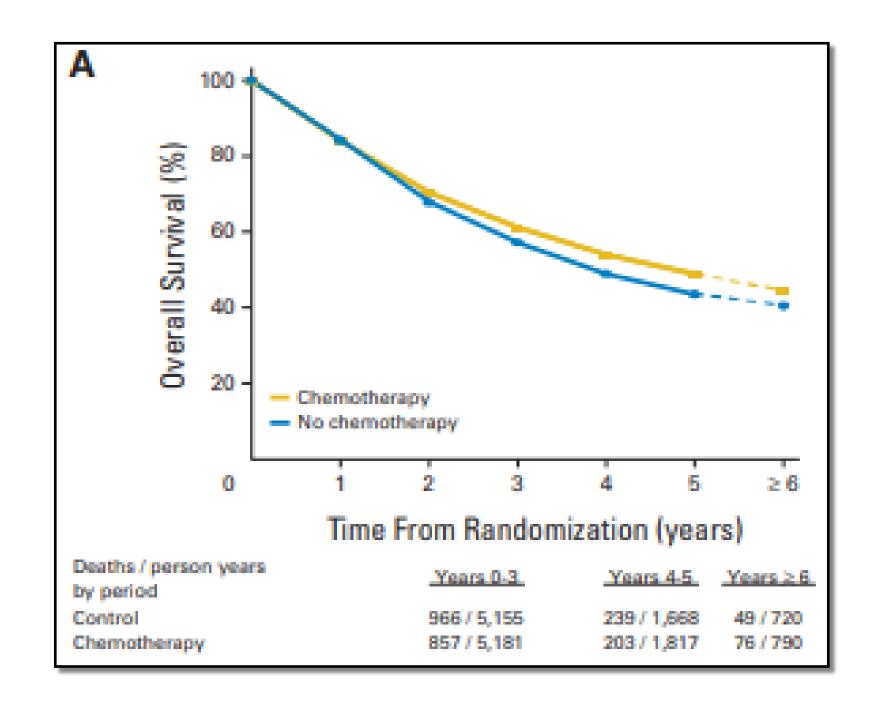
• Outcome worsens- as the stage advances





Role of systemic therapy

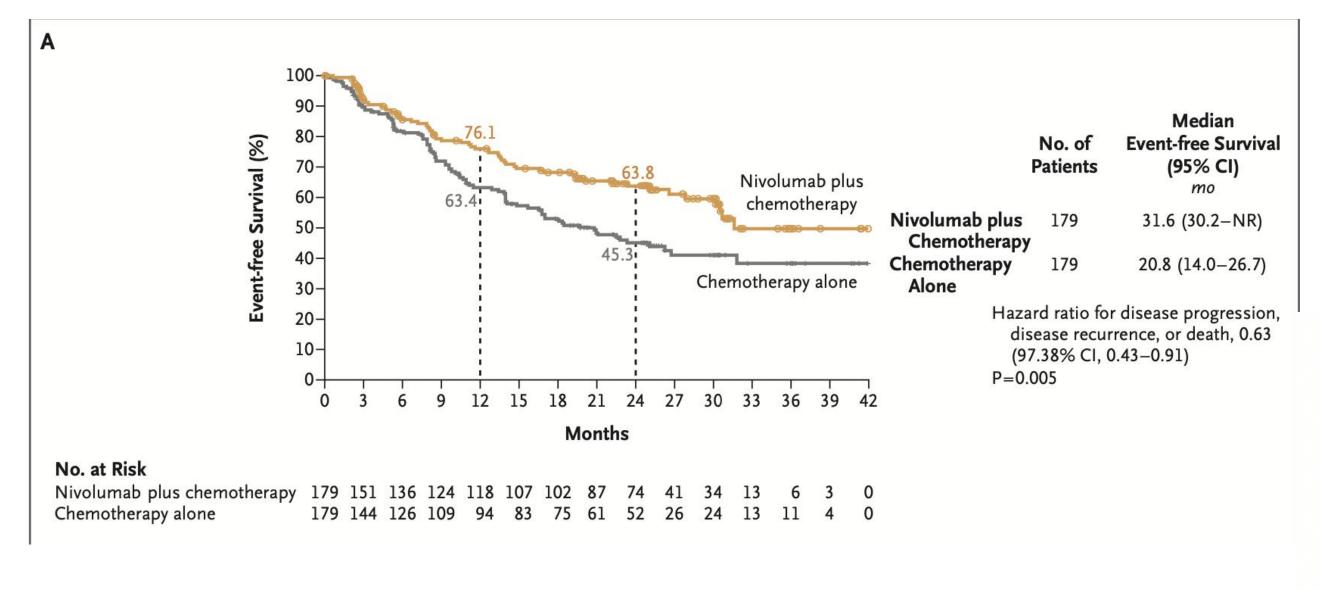
- Need for systemic therapy to improve outcome
- LACE meta-analysis-
 - Overall 5.4%, 5 year survival benefit

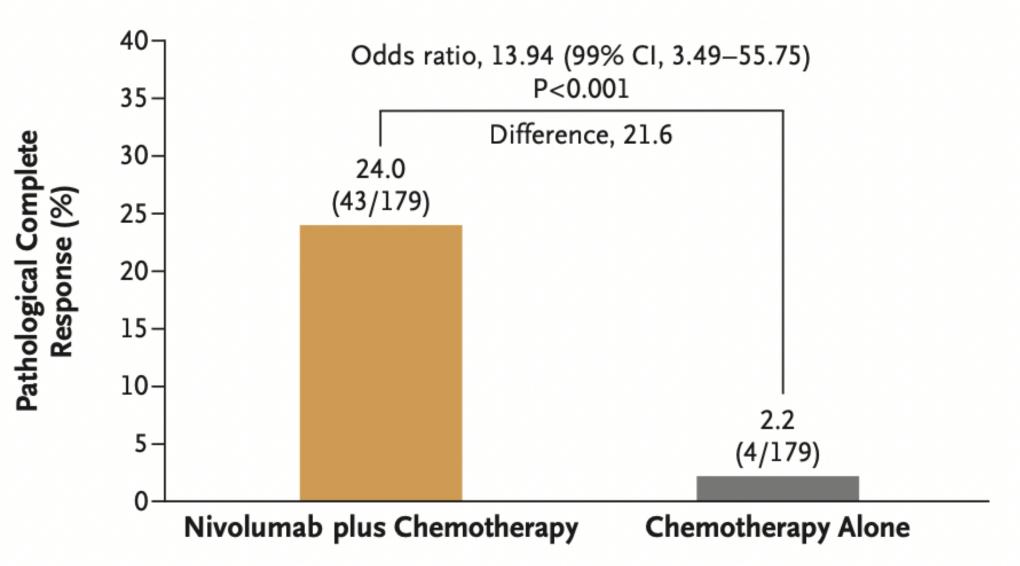


Stage	HR				
IA	1.40				
IB	0.93				
	0.83				
	0.83				

Unmet need

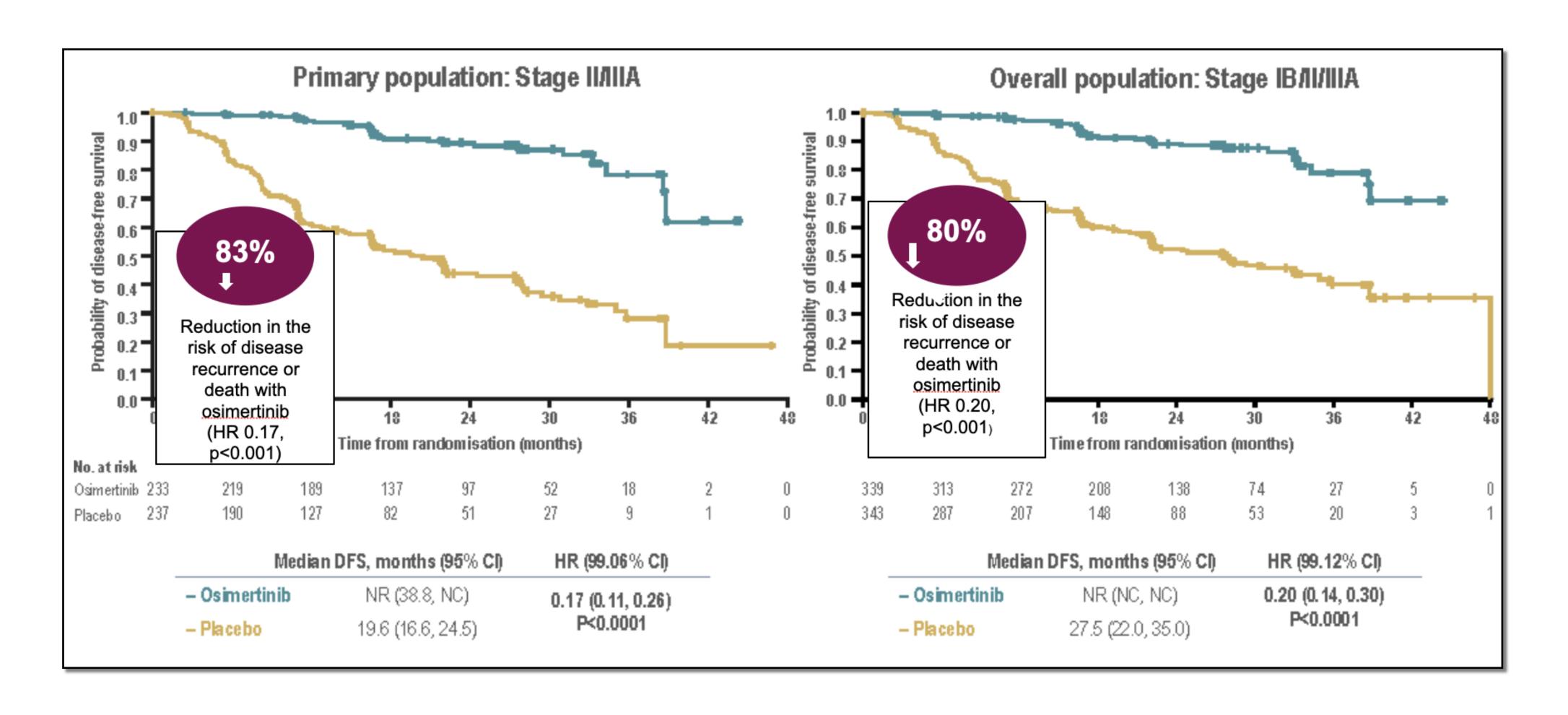
Checkmate 816





Unmet need

ADAURA



Conclusion

• Surgery- Standard of care in early stage NSCLC

MIS- VATS preferred

• Lobectomy + SMLND

• Unmet need persists- multiple ongoing trials

Thank You