| First author, year (reference) | Study characteristics | | | | Adjustment for confounding | | | | | | | | | | | | | | |
|-----------------------------------|-----------------------|-------|--------------------|--------------------|---|----------|----------|-----------|-----------|------------------------|------------------------|-----------------------|-------------------------|--|------------------------|------|---|------------------------|------|
| | | | | | gr F rbid ge span ing jery | | | | ing | jery mor | ds ds | or/ ts | Confid RE Tmt effect | Adjusted % 5 yr OS W <i>vs.</i> Seg | | | Adjusted % 5 yr LCSS W <i>vs</i> . Seg | | |
| | Source | Yrs | Ν | Stage ^a | Demogr F CoMorbid | CoMorbid | Hi stage | Time span | Q setting | Q surgery Fav tumor | Statistical methods | # adj for/ Subsets | Con Tmt | W | Seg | HR | W | Seg | HR |
| Wedge resection vs. s | segmentector | iy | | | | | | | | | | | • | | - A- | | | | |
| Smith ⁿ 2013 (63) | SEER | 98-06 | 3,525 ⁿ | clA1,2 | | | | | | | PA, PQ, PM | 7/2 | М | - | - | 1.19 | - | - | 1.22 |
| Smith ⁿ 2013 (63) | SEER | 98-06 | 3,525 | clA | | | | | | | PA, PQ, PM | 7/2 | М | - | - | 1.23 | - | - | 1.32 |
| Koike 2013 (64) | Japan ×1 | 98-09 | 328 | clA | | | | | | | MV | 15 | М | - | - | - | 68 ^d | 91 ^d | 3.18 |
| Cao 2018 (36) | SEER | 04-13 | 252 ^b | cIA1 | | | | | | | PM | 11 | L | 76 | 74 | 1.05 | 83 | 91 | .75 |
| Cao 2018 (36) | SEER | 04-13 | 852 ^b | cIA2 | | | | | | | PM | 11 | L | 64 | 72 | 1.34 | 75 | 85 | 1.65 |
| Cao 2018 (36) | SEER | 04-13 | 440 ^b | cIA3 | | | | | | | PM | 11 | L | 48 | 53 | 1.17 | 62 | 69 | 1.25 |
| Zhang ° 2016 (65) | SEER | 98-12 | 3,391 | clA | | | | | | | PA | 8/2 | L | - | - | 1.15 | - | - | 1.09 |
| Zhang ^p 2016 (65) | SEER | 98-12 | 1,949 | clA | | | | | | | PA | 8/2 | L | - | - | 1 | - | - | .92 |
| Fan 2020 (47) | SEER | 04-15 | 1,026 | clA1 | | | | | | | MV | 5 | VL | 71 ^d | 76 ^d | 1.42 | - | - | - |
| Dai 2016 (48) | SEER | 00-12 | 981 | clA1 | | | | | | | MV | 6 | VL | 68 ^d | 71 ^d | 1.08 | 83 ^d | 81 ^d | .93 |
| Dai 2016 (48) | SEER | 00-12 | 3,104 | cIA2 | | | | | | | MV | 8 | VL | 62 ^d | 67 ^d | 1.36 | 73 ^d | 82 ^d | 1.42 |
| Zhao 2019 (66) | SEER | 04-15 | 1,372 ^b | clA | | | | | | | MV, PM | 10/3 | VL | 39 | 68 | 1.29 | 77 | 78 | - |
| Dziedzic 2017 (50) | Polish Reg | 07-13 | 462 ^b | cl-IIA | | | | | | | PM | 5 | VL | 54 | 79 | 1.49 | - | - | - |

Table 3 Long-term outcomes in generally healthy patients: wedge resection vs. segmentectomy Ordered by resection extent, degree of confidence that results reflect the effect of the treatment, stage

Inclusion criteria: studies with multivariable or propensity adjustment of wedge resection vs. segmentectomy, 2000–21, with >50 pts per arm in generally healthy patients with generally solid tumors; excluding studies that accrued most patients before 2000. The HR reference is segmentectomy, i.e., HR >1 reflects worse outcome compared with segmentectomy. Bold highlights better outcome (>2-point difference); Light green shading highlights statistically significant difference (lighter shade = univariable; darker = multivariable).

Legend (Tables 1-3):

^a, 8th edition stage classification (reported stage is translated into current 8th edition nomenclature for the sake of uniformity and contemporary application); ^b, propensity matched pairs (total); ^c, all solid tumors (GGN excluded); ^d, unadjusted results; ^e, 3-year survival (in brackets because not comparable to 5-year OS); ^f, All resected by VATS; ^g, 30–50% were "lobe-like" segments (lingula-sparing Left Upper Lobectomy, lingulectomy or basilar quadri-segmentectomy); ^h, cN0 but pN1 (OS in brackets because not comparable to unselected cN0 cohorts; ⁱ, all with visceral pleural invasion (technically stage IB but ≤ 2 cm); ^k, predominantly wedge (\geq 80%); ⁱ, ACS special study (involving enhanced chart abstraction of clinical factors); ^m, lepidic adenocarcinoma; ⁿ, for entire study, not this specific cohort; ^o, adenocarcinoma; ^p, squamous carcinoma.

HR, hazard ratio; LCSS, lung cancer specific survival; Lobe, lobectomy; NCDB, US national cancer database; NS, not statistically significant; OS, overall survival; Reg, registry; SEER, Surveillance, Epidemiology, and End Results database; Seg, segmentectomy; SL, sublobar resection (segmentectomy or wedge); STS-MC, Society of thoracic Surgeons Database, linked to Medicare; VATS, video-assisted thoracic surgery; W, wedge; Yrs, years (of patient accrual).

Adjustment for Confounding: Demogr F, demographic factors (age, sex, socioeconomic); CoMorbid, comorbidities; Hi stage, occult stage inaccuracy due to differences in extent of assessment; Time span, adjustment for changes during the study period or differential use of the interventions; Q settings, discrepancy in the facilities or settings performing the interventions; Q treatmt, quality of the treatment (e.g., margin distance, adjuvant therapy); Fav tumor, selection of less aggressive tumors for an intervention; Statistical methods, methods used to adjust for confounding; Subset, additional subset or sensitivity analyses; # adj for, number of factors adjusted for; Conf RE tmt effect, Confidence that results reflect the effect of the treatment vs. confounding factors. MV, Multivariable model (e.g., Cox regression); PA, propensity score adjustment; PM, propensity matching; PQ, analysis of propensity score quintiles.

| Color | Categories of confounding | Addressed | Neutral (likely little effect) | Limited concern | Moderate concern | High concern | Clearly confounded |
|-------|--------------------------------|--------------|-----------------------------------|-----------------|------------------|--------------|-----------------------|
| code: | Confidence RE treatment effect | VH-very high | H-high | M-moderate | L-low | VL-very low | confidence |