# The Relationship Between Volume or Surgeon Specialty and Outcome in the Surgical Treatment of Lung Cancer A Systematic Review and Meta-Analysis

Erik M. von Meyenfeldt, MD,\* Gea A. Gooiker, MD,† Willem van Gijn, MD,† Piet N. Post, PhD,‡ Cornelis J. H. van de Velde, MD, PhD,† Rob A. E. M. Tollenaar, MD, PhD,† Houke M. Klomp, MD, PhD,\* and Michel W. J. M. Wouters, MD,\*†

**Background:** Whether improvement of quality of surgical cancer care can be achieved by centralizing care in high-volume specialized centers is a subject of ongoing debate. We have conducted a meta-analysis of the literature on the effect of procedural volume or surgeon specialty on outcome of lung resections for cancer.

**Methods:** A systematic search of articles published between January 1, 1990 and January 20, 2011 on the effects of surgeon specialty and hospital or surgeon volume of lung resections on mortality and survival was conducted. After strict inclusion, meta-analysis assuming a random-effects model was performed. Meta-regression was used to identify volume cutoff values. Heterogeneity and the risk of publication bias were evaluated.

**Results:** Nineteen relevant studies were found. Studies were heterogeneous, especially in defining volume categories. The pooled estimated effect size was significant in favor of high-volume hospitals regarding postoperative mortality (odds ratio [OR] 0.71; confidence interval 0.62–0.81), but not for survival (OR 0.93; confidence interval 0.84–1.03). Surgeon volume showed no significant effect on outcome. General surgeons had significantly higher mortality risks than general thoracic (OR 0.78; 0.70–0.88) or cardiothoracic surgeons (OR 0.82; 0.69–0.96). A minimal annual volume of resections for lung cancer could not be identified.

**Conclusions:** Hospital volume and surgeon specialty are important determinants of outcome in lung cancer resections, but evidence-based minimal-volume standards are lacking. Evaluation of individual institutions in a national audit program might help elucidate the influence of individual quality-of-care parameters, including hospital volume, on outcome.

**Key Words:** Lung cancer, Quality improvement, Thoracic surgery, Procedural volume, Surgeon specialty.

(J Thorac Oncol. 2012;7: 1170-1178)

Disclosure: The authors declare no conflict of interest.

Copyright @ 2012 by the International Association for the Study of Lung Cancer ISSN: 1556-0864/12/0707-1170

mprovement of safety, quality, and increased cost-effectiveness of cancer care are a subject of ongoing debate in the professional, public, and political arenas in many countries. The outcome of high-risk, low-volume procedures, such as esophageal and pancreatic resections, is especially thought to improve when centralized in high-volume specialized centers.<sup>1,2</sup> One of the parameters that is frequently suggested in guiding this centralization process is procedural volume. Introduction of minimal volume standards to impose centralization might be appealing to policy makers, because it is a clear and easily enforceable parameter. Because many factors-for example, patient selection, perioperative care protocols, and dedicated nursing staff-influence the outcome of low-volume, high-risk surgical cancer care, the emphasis of governing bodies on hospital-volume criteria seems too limited when the quality of research supporting these criteria is substandard.

Even though the annual case-load is larger than that of pancreatic or esophageal cancer, lung cancer surgery is also deemed to be a low-volume, high-risk surgery. There is, however, limited evidence to support volume standards in lung cancer surgery.

The methodological quality of volume-outcome studies is sometimes disputable and no meta-analysis is yet available. The vast majority of studies is observational and based on administrative data collected for other purposes. Differences in case mix are not always accounted for, and postoperative mortality is often presented as the sole outcome measure. These inadequate reports only feed an endless debate and hamper the introduction of minimal-volume standards in several countries, including The Netherlands.<sup>3</sup>

The aim of this study was to contribute to the ongoing debate by conducting a systematic review and meta-analysis of methodologically high-quality studies on the relationship between hospital volume, surgeon volume, and surgeon specialization on mortality and survival after lung resections.

#### METHODS AND MATERIALS

#### Systematic Search Strategy

We performed a systematic search to identify articles published after January 1, 1990, which described the

<sup>\*</sup>Department of Surgical Oncology, Netherlands Cancer Institute—Antoni van Leeuwenhoek hospital, Amsterdam, The Netherlands; †Department of Surgery, Leiden University Medical Centre, Leiden, The Netherlands; and ‡Post Voor Zorg, Delft, The Netherlands.

Address for correspondence: Erik M. von Meyenfeldt, MD, NKI-AVL, Department of Surgical Oncology, PO Box 90203, 1006 BE Amsterdam, The Netherlands. E-mail: emvonm@gmail.com

TABLE 1.	Search Terms Used;	Last Search	January	/ 20,	2011
----------	--------------------	-------------	---------	-------	------

Medline (PubMed)

(("Pneumonectomy "[majr] OR " Lung Neoplasms /surgery "[MAJR] OR ("Surgical Procedures, Operative"[MAJR:NoExp] AND "Neoplasms"[MAJR:NoExp])) AND ("hospital volume" OR "surgeon volume" OR "surgeon specialty" OR "provider volume" OR "Outcome Assessment (Health Care)"[MAJR] OR regionalization[ti] OR regionalization[ti] OR "Health Facility Size"[majr] OR "Workload"[majr] OR (outcome\*[ti] AND volume\*[ti]) OR (outcome\*[ti] AND complication\*[ti])))

association between surgeon specialty, surgeon volume or hospital volume of lung resections and postoperative mortality and survival. The search was conducted in the Medline database (PubMed) and Cochrane library with a combination of MESH terms and text words (Table 1). Because volume is not well indexed in electronic databases, we formulated the search terms to be as sensitive as possible to ensure no publications were missed. The last search was conducted on January 20, 2011.

#### **Study Selection**

Two reviewers (G.G. and E.v.M.) independently screened titles and abstracts of all retrieved articles. Studies were selected using the following inclusion criteria:

- Written in English.
- Primary data.
- Subject: surgical treatment of lung cancer.
- Comparisons between providers (hospitals or surgeons).
- No single-hospital nor single-surgeon studies.
- Postoperative mortality or survival as outcome parameters.
- Distinct cutoff value for procedural volume or clearly defined specialty.

After this first selection on the basis of titles and abstracts, the remaining articles were obtained in full text and were further selected by the same reviewers. Any discrepancies regarding inclusion of a study were solved by consensus after discussion with a third author (M.W. or P.P.). Reference lists of relevant articles and recent reviews were hand-searched to identify

additional articles. We also used the "related articles" function in PubMed.

The following information was retrieved from the selected articles and entered in our database: year of publication, country, data source, study period, number of patients, hospitals and surgeons, surgeon specialty, case-mix adjustments, volume categories for hospitals, volume categories for surgeons, outcome parameters (mortality and survival), and results regarding these outcome parameters (statistically significant or nonsignificant).

#### Assessment of Study Quality

Two authors (E.v.M. and G.G.) critically appraised each study in the review on methodological quality and multiple use of the same database. Data were gathered in a data-extraction form, which was based on the Strengthening the Reporting of Observational studies in Epidemiology criteria (www.strobestatement.org).

Study inclusion criteria were checked for probability of selection bias. When multiple use of the same database was established, the study with the largest patient cohort was used. Cutoff values for high- and low-volume were noted per volume group. The study results were recorded separately for each unit (surgeon, surgeon specialty, or hospital) and for each outcome parameter (postoperative mortality and long-term survival). Subsequently, we noted the estimated effect size after adjustment comparing the highest volume group with the lowest, expressed as odds ratios (OR), hazard ratios (HR), or risk rates (RR) with confidence intervals (CI). If necessary, we converted the effect sizes so that the lowest-volume group was the reference in all studies. Studies without a multivariate analysis and/or no reporting of OR, HR, or RR were excluded from the meta-analysis.

#### Synthesis of the Data for Meta-Analysis

A meta-analysis was performed for the relationship between hospital volume, surgeon volume, and surgeon specialty on the one hand and postoperative mortality and long-term survival on the other. The random-effect model, which accounts for the expected heterogeneity in pooling observational studies,



FIGURE 1. Selection process of reviewed studies. OR, odds ratio.

Copyright © 2012 by the International Association for the Study of Lung Cancer

**Case-Mix Adjustment** 

**TABLE 2.** Studies Included in the Systematic Review of the Literature on the Relationship Between Volume and Outcome of Lung Resections for Cancer

Study	Country	Data	Patients	Hospitals	Surgeons	Demographic	Comorbidities	Urgency	Tumor	Treatment
Birkmeyer et al. <sup>1</sup>	USA	Adm	19410	1877	n.r.	•	•	•	_	•
Birkmeyer et al. <sup>2</sup>	USA	Adm	24092	n.r.	4178	•	•	•	—	—
Sioris et al.13	Finland	Adm	5339	26	n.r.	•	_	—	•	•
Bilimoria et al. <sup>14</sup>	USA	Adm	40754	1528	n.r.	•	٠	—	•	•
Urbach et al.15	Can	Adm	5156	54	n.r.	•	•			
Freixinet et al. <sup>16</sup>	Spain	Clin	2994	19	n.r.	•	•	—	•	•
Romano and Mark <sup>17</sup>	USA	Adm	12439	389	n.r.	•	•	—	•	•
Simunovic et al. <sup>18</sup>	Canada	Adm	2698	67	n.r.	•	•	—	•	_
Finlayson et al. <sup>19</sup>	USA	Adm	21890	674	n.r.	•	•	•	—	•
Osada and Yamakoshi <sup>20</sup>	Japan	Clin	3233	91	n.r.	•	—	—	•	•
Osada and Yamakoshi <sup>20</sup>	Japan	Clin	18055	540	n.r.	•	—	—	—	•
Hollenbeck et al. <sup>21</sup>	USA	Adm	90088	n.r.	n.r.	•	•	•	—	—
Kim et al. <sup>22</sup>	South Korea	Adm	987	n.r.	n.r.	•	•	•	—	—
Li et al.23	Netherlands	Clin	1097	20	n.r.	•	_	_	•	_
Birkmeyer et al. <sup>24</sup>	USA	Adm	12967	517	n.r.	•	•	•	•	•
Cheung et al <sup>25</sup>	USA	Adm	13469	90	n.r.	•	•	_	•	•
Lien et al.26	Taiwan	Adm	4841	79	377	•	•		•	•
Goodney et al. <sup>27</sup>	USA	Adm	25545	n.r.	n.a.	•	•	—		•
Schipper et al. <sup>28</sup>	USA	Adm	86538	n.r.	n.a.	•		—	—	•
Farjah et al. <sup>29</sup>	USA	Adm	19745	n.r.	n.a.	•	٠	—	•	•

Adm, based on administrative data; Clin, based on clinical data; n.r., not reported; n.a., not applicable; Demographic, adjusted for demographic data (e.g., patient age, sex, race, and income); Comorbidities, adjusted for comorbidities (including American Society of Anesthesiologists classification); Urgency, adjusted for urgency of the operation; Tumor, adjusted for tumor characteristics (e.g., stage, grade, and location); Treatment, adjusted for treatment differences (e.g., surgical approach; [neo]adjuvant treatments); Mortality, s urvival analysis adjusted for postoperative mortality; Hospital, adjusted for other hospital characteristics (e.g., teaching or academic status); Low: <10, low-volume group < 10, High: >20, high-volume > 20 resections a year; S, statistically significant; NS, statistically nonsignificant; L, lesser lung resections; P, pneumonectomies; TS, general thoracic surgeon; CTS, cardio thoracic surgeon; GS, general surgeon.

was used.<sup>4</sup> Heterogeneity was quantified by the *P* test. An *P* less than 40 was considered homogeneous; between 40 and 60, moderately heterogeneous; and greater than 60 very heterogeneous.<sup>5</sup> We conducted a sensitivity analysis to assess the impact of subgroups in case of heterogeneity. Publication bias was assessed with an Egger's regression intercept.<sup>6</sup> Meta-regression analysis was done for cutoff values of hospital volume, using a fixed-effects regression test. The meta-analysis was conducted with Comprehensive Meta Analysis, professional version 2.2 (Biostat Inc., Englewood NJ).

#### RESULTS

#### Search Results

Our initial search identified 95 potentially relevant articles. After the first screening, 17 articles were retrieved for more detailed evaluation. From these, six articles were excluded.<sup>7–12</sup> The remaining 11 articles were selected. After this first selection, the related-articles feature in PubMed was used, and the reference lists of retrieved articles were hand-searched to identify eight additional articles, which met the

			Hospi	tal Volume			Surg	geon Volume		Sur	geon Specialty	7
		Volu Categ	ime ories			Vol Cate	ume gories			Surgoon		
Mortality	Hospital	Low	High	Mortality	Survival	Low	High	Mortality	Survival	Categories	Mortality	Survival
		<9	>46	S(L) & (P)	_		_			_	_	
—	•			—	—	<7	>17	NS	—	—	—	—
_	_	<5	>20	NS	S	_			_	_	_	_
—	—	<21	>83	S	S			—	—	—	—	—
_	_	<18.2	>129,4	NS	_	_			_	_	_	_
—	—	<44	>54	NS	NS				—	—	—	—
—	•	<9	>24	S (L), NS (P)	—	—	—	—	—	—	—	—
_	•	<10.7	>43,7	NS	S		—	_	_	—	—	_
_	—	<19	>37	NS(L) &(P)	_	_	—	—	—	_		_
_	_	<20	>79		S	_	_		_	—	—	
_	_	<25. quartiles	>99	NS	_	_	—	—	_	_		_
_	_	<3.6	>116,3	S	_	_	_		_	—	—	
—	—	<5	>20	NS	_	_	_	_	_	—	—	_
	•	<40	>60	_	NS	_	_	_	_	CTS vs. GS	_	NS
•	—	<11.4	>25,2	—	S	—	—	_	—	—	—	
_	•	<60	>60	_	S	_	_	_	_	_	_	_
		_	_			<11.5	>33	S				_
—	•	—	—	—	—	—	—	—	—	663 TS, 1516 CTS, 2614 GS	S	—
_	•	—	_	_	_	—	_	_		331 TS, 3401 CTS, 9194 GS	S (L), NS (P)	—
—	—	—	—	—	—	—	—	—	—	388 TS, 684 CTS, 776 GS	NS	S

predefined inclusion criteria. Nineteen eligible articles were analyzed (Fig. 1).

#### **Study Characteristics**

Table 2 shows the characteristics of the 19 studies included in the review. Most studies are from the United States and Canada, and the remaining studies are from Asia and Europe. Study data have been obtained frequently from the databases of insurance companies or national cancer registries (Medicare, National Inpatient Sample). The number of patients, hospitals, and surgeons varied widely among the included studies. In all selected studies, the results were adjusted for differences in case mix, but the parameters used for the adjustments differed largely. For hospital volume, cutoff values of the highest volume strata varied between 20 and 129.4 procedures annually, and the lowest hospital-volume strata cutoff varied between 3.6 and 60. The rationale for the cutoff values used varied and was not always explained in the sections on methodology in the articles. **FIGURE 2.** Forest plot of the included studies using OR in the meta-analysis on hospital volume and postoperative mortality for lung resections for cancer. Separate analysis of pneumonectomies and lobectomies in case of sufficient data. OR, odds ratio; Study name, name of first author (resection type if specific odds ratios available); Year, year of publication; Low, highest annual volume of low-volume category; High, lowest annual volume of high-volume category; CI, confidence interval.

# Hospital mortality



Meta Analysis

### **Hospital Volume**

Postoperative mortality was used as an outcome parameter in 11 studies; in five of these studies, a significant inverse relationship between hospital volume and 30-day or in-hospital mortality was found.<sup>1,13-22</sup>

Ten studies were used for the meta-analysis, as one did not provide an OR.<sup>13</sup> All of these studies had an observational design, and only two studies were based on clinical data; the remaining eight were based on administrative data. In all but one, the results were adjusted not only for age and sex, but also for other parameters. (Table 2)

Figure 2 shows the forest plot regarding hospital volume and postoperative mortality using ORs. The effect was significant in favor of high-volume hospitals (OR 0.71; CI 0.62–0.81). One study used HRs and one study used relative risk, both favoring high hospital volume, with an HR of 0.76 (CI 0.66–0.88) and RR of 0.83 (CI 0.59–1.25), respectively.<sup>14,15</sup> There was moderate heterogeneity between the two studies ( $I^2 = 53$ ). Subgroup analysis of the studies using ORs did not reveal any parameter showing significant influence on the pooled estimate effect size. (Table 3) No publication bias was detected (p = 0.36).

Regression analysis did not identify a cutoff value for volume of resections for lung cancer to differentiate hospitals with high versus low postoperative mortality rates. The p value for the slope is 0.73 (Fig. 3).

Differences in survival between high- and low-volume hospitals were evaluated in eight studies, five of which showed a significant inverse relationship between hospital volume and survival and one of which showed a significant benefit for the lower-volume category.<sup>13,14,16,18,20,23–25</sup>

Seven studies met the criteria for the meta-analysis, as one did not provide an HR.<sup>16</sup> All studies were observational; two studies used clinical instead of administrative data. In all studies, the results were adjusted not only for age and sex, but also for other parameters (Table 2).

Figure 4 shows the forest plot on hospital volume and survival. The effect was in favor of high-volume hospitals but it did not reach statistical significance (HR 0.93; CI 0.84–1.03). This result was very heterogeneous ( $I^2 = 85$ ). In

**TABLE 3.** Sensitivity Analysis of the 11 Included Series onHospital Volume and Postoperative Mortality with OddsRatios as Effect Size

Factor	Subgroup	Ν	OR	CI	p Value
Datasource					0.367
	Administrative	9	0.69	0.60-0.79	
	Clinical	2	0.79	0.60-1.04	
Comorbidity					0.198
	Not adjusted	1	0.55	0.36-0.83	
	Adjusted	10	0.73	0.64-0.83	
Urgency					0.810
	Not adjusted	5	0.69	0.55-0.86	
	Adjusted	6	0.72	0.60-0.85	
Tumor stage					0.367
_	Not adjusted	9	0.69	0.60-0.79	
	Adjusted	2	0.79	0.60-1.04	

This (two-sided) p value is the result of the mixed-effects analysis and tells us whether there is a statistically significant difference in the effect between subgroups. N, number of studies; OR, odds ratio; CI, confidence interval; P, result of I square test on heterogeneity of study results.<sup>8</sup>



# **FIGURE 3.** Regression analysis of the effect of high procedural hospital volume on hospital mortality.

Copyright © 2012 by the International Association for the Study of Lung Cancer

## **Hospital survival**

<u>Mode</u> l	<u>el Study nam¥ear Low High</u>			St	atistic	s for e	ach stud	study Hazard ratio and 95%					
				I	Hazard ratio	Lower limit	Upper limit	Z-Value	o-Value				
	Bilimoria	2008	20	84	0.917	0.876	0.961	-3.680	0.000				
	Birkmeye	r 2007	11	25	0.830	0.777	0.887	-5.536	0.000		-		
	Cheung	2009	10	61	0.930	0.875	0.988	-2.352	0.019				
	Li	2008	40	60	1.180	0.875	1.591	1.085	0.278		_		
	Osada	2007	19	80	0.641	0.413	0.996	-1.976	0.048	← ■			
	Simunovi	c2006	11	44	0.769	0.638	0.928	-2.745	0.006				
	Sioris	2008	4	21	1.250	1.102	1.417	3.481	0.001				
Random					0.931	0.843	1.029	-1.404	0.160		-		
										0.5	1		2
										Favours hi	gh volume	Favours low vol	ume
Meta A	nalysis												

**FIGURE 4.** Forest plot of the included studies in the meta-analysis on hospital volume and survival of lung resections for cancer. Study name, name first author (resection type if specific odds ratios available); Year, year of publication; Low, highest annual volume of low-volume category; High, lowest annual volume of high-volume category; CI, confidence interval.

**TABLE 4.** Sensitivity Analysis of the Included Studies on Hospital Volume and Survival with Hazard Ratios as Effect Size

Factor	Subgroup	Ν	OR	CI	<i>p</i> Value
Datasource					0.961
	Administrative	3	0.93	0.79-1.10	
	Clinical	4	0.93	0.79 - 1.09	
Comorbidity					0.003
	Not adjusted	3	1.15	0.98-1.34	
	Adjusted	4	0.88	0.81-0.95	
Urgency					0.472
	Not adjusted	5	0.96	0.83-1.12	
	Adjusted	2	0.88	0.72 - 1.07	
Tumor stage					0.127
	Not adjusted	1	0.64	0.39-1.05	
	Adjusted	6	0.95	0.86-1.05	

This (two-sided) p value is the result of the mixed-effects analysis and tells us whether there is a statistically significant difference in the effect between subgroups. Bold indicates statistical significance.

N, number of studies; OR, odds ratio; CI, confidence interval; I, result of I square test on heterogeneity of study results.<sup>8</sup>

subgroup analysis, a significant effect of hospital volume on survival was observed after adjustment for comorbidities (OR 0.88; CI 0.81–0.95) (p = 0.003, Table 4). No publication bias was detected (p = 0.82).

#### **Surgeon Volume**

In two studies, the relationship between surgeon volume and outcome was investigated.<sup>2,26</sup> Both studies used postoperative mortality as an outcome parameter; one showed a significant result favoring high volume. Both were used for the meta-analysis. Age, sex, and comorbidities were included in the multivariate analysis in both studies. One adjusted for tumor characteristics and treatment differences in its outcome analyses, the other for urgency of operation and other hospital characteristics.

Figure 5 shows the forest plot of the two studies. In the meta-analysis, a pooled estimated effect size was detected in favor of high-volume surgeons, but this effect did not reach statistical significance (OR 0.68; CI 0.42–1.08) and was very heterogeneous ( $I^2 = 66$ ). The number of studies is

too small to properly assess publication bias or perform subgroup analysis.

#### **Surgeon Specialty**

In three studies, the relationship between surgeon specialty and postoperative mortality was investigated.<sup>27-29</sup> These studies compared three provider groups: general surgeons (GS), cardiothoracic surgeons (CTS), and general thoracic surgeons (TS). One study showed a significant difference; in another, the difference was significant in lobectomies, not in pneumonectomies. The third study did not show a significant difference. All three studies met the inclusion criteria for the meta-analysis. In all, age, sex, comorbidities, and treatment differences were included in the multivariate analysis. Two adjusted for other hospital characteristics and one adjusted for tumor characteristics in their survival analyses. Figures 6A and B show the forest plots of the included studies. There was a significant pooled estimated effect size in favor of TS over GS (OR 0.78; 0.70-0.88), and of CTS over GS (OR 0.82; CI 0.69-0.96). As a result of the lack of ORs comparing TS with CTS, no pooled estimate could be calculated. The number of studies, again, is too small to properly assess publication bias or perform subgroup analysis.

Differences in survival between different specialties were evaluated in two studies, one of which showed a significant difference both when comparing GS with CTS and when comparing GS with TS.<sup>23,29</sup> Both studies met the inclusion criteria for the meta-analysis. Both adjusted for age, sex, and tumor characteristics; one also adjusted for comorbidity and treatment differences, the other, for other hospital characteristics.

Figure 7 shows the forest plot of the included studies. In the meta-analysis, a pooled estimated effect size was detected in favor of TS over GS, but this effect did not reach statistical significance (HR 0.86; CI 0.74–1.00). This result was very homogeneous ( $l^2 = 28$ ). The number of studies is too small to properly assess publication bias or perform subgroup analysis.

#### DISCUSSION

This study is the first meta-analysis on the effect of surgeon specialty and of procedural volume on postoperative

Surgeon mortality

#### Statistics for each study Odds ratio and 95% CI Model Study nam Year Low High Odds Lower Upper ratio limit limit Z-Valuep-Value FIGURE 5. Forest plot of the Birkmeyer 2003 6 18 0.806 0.698 0.931 -2.931 0.003 included studies in the meta-analysis 2007 9 26 0.490 0.283 0.849 -2.545 0.011 Lien on surgeon volume and postopera-0 677 0 424 1 079 -1 642 0 101 Random tive mortality of lung resections for 0.5 2 cancer. Year, year of publication; Low, highest annual volume of low-vol-Favours high volume Favours low volume ume category; High, lowest annual volume of high-volume category; Cl, Meta Analysis confidence interval. Specialisation GS vs TS Α Model Study nameYear Statistics for each study Odds ratio and 95% CI Odds Lower Upper limit Z-Value p-Value ratio limit 2009 0.700 1.051 -1.719 0.086 Schipper 0 466 Goodney 2005 0.770 0.881 -3.807 0.000 0.673 Farjah 2009 0.920 0.679 1.247 -0.537 0.591 Random 0.784 0.697 0.883 -4.038 0.000 0.5 2 Favours high volume Favours low volume

Meta Analysis

**FIGURE 6.** *A* and *B*, Forest plot of the included studies in the metaanalysis on surgeon specialty and postoperative mortality of lung resections for cancer. *A*, General surgeon versus general thoracic surgeon. *B*, General surgeon versus cardiothoracic surgeon. GS, general surgeon; TS, general thoracic surgeon; CTS, cardiothoracic surgeon; Year, year of publication; Low, highest volume of low-volume category; High, lowest annual volume of high -volume category; CI, confidence interval.

### B Specialisation GS vs CTS

Model	<u>Study nameYear</u>			Year Statistics for each study					O <u>dds ratio and 95% C</u> I			
			Odds ratio	Lower limit	Upper limit	Z-Value	p-Value					
	Schipper	2009	0.880	0.722	1.073	-1.265	0.206		∎∔			
	Goodney	2005	0.720	0.636	0.815	-5.196	0.000	-	∎-			
	Farjah	2009	0.920	0.730	1.160	-0.706	0.480					
Random			0.818	0.694	0.964	-2.403	0.016	•				
								0.5	1	2		
								Favours hi	gh volume Favours	low volume		

Meta Analysis

mortality and long-term survival after lung resections for cancer. Only a minority of the increasing number of published studies met our strict inclusion criteria for methodological quality. We found that hospital volume has a significant inverse relationship with postoperative mortality, although moderate heterogeneity among studies exists. A cutoff value for hospital volume could not be identified.

The effect of hospital volume on survival only reached significance in the subgroup of these very heterogeneous studies, which adjusted for comorbidity (Table 4). This suggests that high-volume hospitals tend to treat more patients with multiple comorbidities, which influences the overall survival in these hospitals. The pooled-effect estimate of surgeon volume on mortality did not reach significance (very heterogeneous), suggesting that outcome depends more on hospital setting and team performance than on volume of individual surgeons. Specific thoracic training, however, is of significant importance. Postoperative mortality and survival rates are significantly in favor of CTS and TS when compared with GS (very homogeneous).

The available evidence has limitations that cannot be ignored. Our review confirms that most studies are observational, retrospective, and based on administrative data collected for other purposes, instead of carefully designed comparative studies (Table 2). However, we did not observe

# Specialisation GS vs TS survival

Model	Study name	Year	Statistics for each study				,	H <u>azard ratio and 95% C</u> I			
			Hazard ratio	Lower limit	Upper limit	Z-Value	o-Value				
	Li	2008	0.730	0.529	1.008	-1.913	0.056		■		
	Farjah	2009	0.890	0.835	0.949	-3.574	0.000				
Random			0.861	0.744	0.996	-2.020	0.043				
								0.5	1	2	
								Favours hi	igh volume Fa	vours low volume	
Meta Ar	nalvsis										

**FIGURE 7.** Forest plot of the included studies in the meta-analysis on surgeon specialty and survival of lung resections for cancer. General surgeon versus general thoracic surgeon. GS, general surgeon; TS, general thoracic surgeon; Year, year of publication; Low, highest annual volume of low-volume category; High, lowest annual volume of high-volume category; CI, confidence interval.

significantly different effect sizes among studies using clinical data and those using administrative data. Large variation was observed in demographical, geographical, and epidemiological factors and in standards of care and definitions of surgeon specialties. Awareness of these differences in training and certification of "specialized" surgeons is important in interpreting these results. In the Dutch situation, for instance, a growing number of GSs have had specialist training in lung resections, which should qualify them as general TSs.<sup>23</sup>

Substantial heterogeneity was identified in evaluating studies' methodological quality. The choice of volume categories was extremely diverse but seldom explained. The variety in cutoff values made it impossible to identify a minimal hospital volume with regression analysis, despite looking for the biggest difference by comparing the highest- and lowestvolume category from each article.

Apart from the methodological shortcomings mentioned above, volume-outcome studies have other important limitations. In only few studies, data have been corrected for (other) provider characteristics, such as the available infrastructure, teaching or academic status, experience with other high-risk operations, expertise in multimodality cancer treatments, a hospital's budget, focus and/or referral bias.<sup>17,18</sup> Also, in almost all studies, only the resected patients were analyzed. Essential data on patient selection and resection rates are missing; therefore, the effect of procedural volume or surgeon specialty on the ability to provide appropriate surgical care for more challenging patients remains unclear. Lowvolume centers might predominantly select a more favorable patient group (lower stage and less comorbidity), thus influencing their outcome. The question of whether this clinically more favorable patient group would also benefit from referral to high-volume centers could not be assessed in this metaanalysis. These factors are often related to, but not identical with, procedural volume.

In the Netherlands the Quality of Cancer Care task force of the Dutch Cancer Society has recently proposed to concentrate specific cancer treatments in hospitals that meet a set of criteria.<sup>30</sup> The Dutch Surgical Society set the criteria for lung cancer surgery.<sup>31</sup> These criteria focus not only on procedural volume, but also on the available infrastructure, specialization, and outcome measures, which should be reported by individual institutions.<sup>32</sup> Careful analysis of audit data retrieved from different hospitals that vary in patterns of care and outcomes might identify ways to improve lung cancer treatment more than merely enforcing volume criteria.<sup>33–35</sup> In our opinion, policy makers should bear this in mind when efforts are made to centralize complex high-risk surgical procedures.

In conclusion, this meta-analysis has shown that higher procedural volume is associated with better outcomes in lung cancer surgery. Annual hospital volume seems more important than the volume of individual surgeons. Surgeon specialty is also of significant importance. Concentration of lung cancer surgery could lead to a decrease in postoperative mortality and a better overall survival, although there is no evidence for a specific annual volume cutoff. Evaluation of individual institutions in a national audit program might help elucidate the influence of individual quality-of-care parameters, including hospital volume, on outcome, and provide better parameters to help improve surgical cancer care.

#### REFERENCES

- Birkmeyer JD, Siewers AE, Finlayson EV, et al. Hospital volume and surgical mortality in the United States. N Engl J Med 2002;346:1128–1137.
- Birkmeyer JD, Stukel TA, Siewers AE, Goodney PP, Wennberg DE, Lucas FL. Surgeon volume and operative mortality in the United States. *N Engl J Med* 2003;349:2117–2127.
- van Heek NT, Kuhlmann KF, Scholten RJ, et al. Hospital volume and mortality after pancreatic resection: a systematic review and an evaluation of intervention in the Netherlands. *Ann Surg* 2005;242:781–788, discussion 788.
- Mahid SS, Hornung CA, Minor KS, Turina M, Galandiuk S. Systematic reviews and meta-analysis for the surgeon scientist. *Br J Surg* 2006;93:1315–1324.
- Higgins JP, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analyses. *BMJ* 2003;327:557–560.
- Egger M, Davey Smith G, Schneider M, Minder C. Bias in meta-analysis detected by a simple, graphical test. *BMJ* 1997;315:629–634.
- Finley CJ, Bendzsak A, Tomlinson G, Keshavjee S, Urbach DR, Darling GE. The effect of regionalization on outcome in pulmonary lobectomy: a Canadian national study. *J Thorac Cardiovasc Surg* 2010;140:757–763.
- Ho V, Heslin MJ, Yun H, Howard L. Trends in hospital and surgeon volume and operative mortality for cancer surgery. *Ann Surg Oncol* 2006;13:851–858.
- Meguid RA, Brooke BS, Chang DC, Sherwood JT, Brock MV, Yang SC. Are surgical outcomes for lung cancer resections improved at teaching hospitals? *Ann Thorac Surg* 2008;85:1015–1024; discussion 1024.
- Hannan EL, Radzyner M, Rubin D, Dougherty J, Brennan MF. The influence of hospital and surgeon volume on in-hospital mortality for colectomy, gastrectomy, and lung lobectomy in patients with cancer. *Surgery* 2002;131:6–15.

- Bach PB, Cramer LD, Schrag D, Downey RJ, Gelfand SE, Begg CB. The influence of hospital volume on survival after resection for lung cancer. N Engl J Med 2001;345:181–188.
- Begg CB, Cramer LD, Hoskins WJ, Brennan MF. Impact of hospital volume on operative mortality for major cancer surgery. *JAMA* 1998;280:1747–1751.
- Sioris T, Sihvo E, Sankila R, Salo J. Effect of surgical volume and hospital type on outcome in non-small cell lung cancer surgery: a Finnish population-based study. *Lung Cancer* 2008;59:119–125.
- Bilimoria KY, Bentrem DJ, Feinglass JM, et al. Directing surgical quality improvement initiatives: comparison of perioperative mortality and longterm survival for cancer surgery. J Clin Oncol 2008;26:4626–4633.
- Urbach DR, Bell CM, Austin PC. Differences in operative mortality between high- and low-volume hospitals in Ontario for 5 major surgical procedures: estimating the number of lives potentially saved through regionalization. *CMAJ* 2003;168:1409–1414.
- 16. Freixinet JL, Julià-Serdà G, Rodríguez PM, et al. Bronchogenic Carcinoma Cooperative Group of the Spanish Society of Pneumology and Thoracic Surgery GCCB-S. Hospital volume: operative morbidity, mortality and survival in thoracotomy for lung cancer. A Spanish multicenter study of 2994 cases. *Eur J Cardiothorac Surg* 2006;29:20–25.
- Romano PS Mark DH. Patient and hospital characteristics related to inhospital mortality after lung cancer resection. *Chest* 1992;101:1332–1337.
- Simunovic M, Rempel E, Thériault ME, et al. Influence of hospital characteristics on operative death and survival of patients after major cancer surgery in Ontario. *Can J Surg* 2006;49:251–258.
- Finlayson EV, Goodney PP, Birkmeyer JD. Hospital volume and operative mortality in cancer surgery: a national study. *Arch Surg* 2003;138:721–5; discussion 726.
- Osada H Yamakoshi E. Hospital volume and surgical outcomes of lung cancer in Japan. *Gen Thorac Cardiovasc Surg* 2007;55:360–365.
- Hollenbeck BK, Dunn RL, Miller DC, Daignault S, Taub DA, Wei JT. Volume-based referral for cancer surgery: informing the debate. *J Clin* Oncol 2007;25:91–96.
- Kim SY, Park JH, Kim SG, et al. Disparities in utilization of high-volume hospitals for cancer surgery: results of a Korean population-based study. *Ann Surg Oncol* 2010;17:2806–2815.

- 23. Li WW, Visser O, Ubbink DT, Klomp HM, Kloek JJ, de Mol BA. The influence of provider characteristics on resection rates and survival in patients with localized non-small cell lung cancer. *Lung Cancer* 2008;60:441–451.
- Birkmeyer JD, Sun Y, Wong SL, Stukel TA. Hospital volume and late survival after cancer surgery. *Ann Surg* 2007;245:777–783.
- Cheung MC, Hamilton K, Sherman R, et al. Impact of teaching facility status and high-volume centers on outcomes for lung cancer resection: an examination of 13,469 surgical patients. *Ann Surg Oncol* 2009;16:3–13.
- Lien YC, Huang MT, Lin HC. Association between surgeon and hospital volume and in-hospital fatalities after lung cancer resections: the experience of an Asian country. *Ann Thorac Surg* 2007;83:1837–1843.
- Goodney PP, Lucas FL, Stukel TA, Birkmeyer JD. Surgeon specialty and operative mortality with lung resection. *Ann Surg* 2005;241:179–184.
- Schipper PH, Diggs BS, Ungerleider RM, Welke KF. The influence of surgeon specialty on outcomes in general thoracic surgery: a national sample 1996 to 2005. *Ann Thorac Surg* 2009;88:1566–1572; discussion 1572.
- Farjah F, Flum DR, Varghese TK Jr, Symons RG, Wood DE. Surgeon specialty and long-term survival after pulmonary resection for lung cancer. *Ann Thorac Surg* 2009;87:995–1004; discussion 1005.
- Quality of Cancer Care Taskforce. *Quality of Cancer Care in The Netherlands [Dutch]*. Oisterwijk, the Netherlands: Dutch Cancer Society, 2010.
- Dutch Surgical Society. Minimal Criteria for Surgical Treatment [Dutch]. [Dutch Surgical Society (NVvH) web site] January 2011. Available at: http://www.heelkunde.nl/uploads/50/5H/505H5htx15bBtDFNcBE mkw/nvvh-normen-januari-2011.pdf. Accessed September 27, 2011.
- Wouters MW, Jansen-Landheer ML, van de Velde CJ. The Quality of Cancer Care initiative in the Netherlands. *Eur J Surg Oncol* 2010;36 Suppl 1:S3–S13.
- Birkmeyer JD, Dimick JB, Birkmeyer NJ. Measuring the quality of surgical care: structure, process, or outcomes? JAm Coll Surg 2004;198:626–632.
- van Gijn W, van de Velde CJ. members of the EURECCA consortium. Improving quality of cancer care through surgical audit. *Eur J Surg Oncol* 2010;36 Suppl 1:S23–S26.
- 35. Wouters M. Outcome-based referral to improve quality of care in upper GI surgery. *J Surg Oncol* 2008;98:307–308.