

# The 100 most cited articles on thoracic surgery management of lung cancer

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#### Introduction

Lung cancer is the leading cause of cancer-related deaths worldwide. In 2019, 228,150 new lung cancer cases and 142,670 cancer deaths are projected to occur in the United States (1), and non-small cell lung cancer (NSCLC) accounts for nearly 85% of all diagnosed cases (1,2). As the US Preventative Services Task Force has released recommendations of computed tomography lung cancer screening for long-term smokers, the incidence of newly diagnosed lung cancer will have a marked increase (3).

Over the past two decades, the therapies for lung cancer have evolved into a mature subspecialty, also being the culmination of a progressive research initiative and the continued innovation of interventional techniques. For operable NSCLC, the current guideline states that VATS lobectomy is the standard operation (4,5). In recent years, a great progress in the clinical treatment and basic research of adjuvant therapy for lung cancer has been obtained, such as chemotherapy, radiotherapy, target therapy, and immunotherapy (6-8).

A large amount of literature is available to thoracic medicine specialists who perform thoracic surgery management of lung cancer. It is difficult to recognize articles of significance since the quality of the literature varies substantially and includes low impact studies and open access journals. An understanding of the available literature and the most heavily cited works will allow for a better understanding of the evidence base when discussing and providing these treatments.

A bibliometric analysis is a quantitative method used to examine the knowledge structure and development of a research field and its publications. Among all the bibliometric methods, the citation analysis is the most common which focuses on citation number and number of citations per year (9,10). It evaluates the impact an article has had on a specific field of medicine by measuring the number of citations an article has received.

Since 1971, the global output of academic research into the management of lung cancer has increased. There has been no specific quantitative analysis on the articles which have had the most significant impact on the treatment of lung cancer been performed. The aim of this study was to perform a citation analysis on the most cited articles in the thoracic surgery management for lung cancer and analyze each article individually, collecting the article type, year of publication, topic of interest, citation index, authorship, country of origin, institution, and level of evidence.

#### Methods

#### Identification of the top 100 cited articles

Between July and August 2018, we performed a retrospective bibliometric analysis, identifying articles through the Web of Science (Thomson Reuters, New York, USA) which searches the Science Citation Index Expanded database, allowing access to over 12,000 peer reviewed journals and collected citation data published since 1945.

Using selected key terms ("lung cancer", "surgery", "lobectomy", "segmentectomy", "pneumonectomy", "wedge resection", "video-assisted thoracic surgery", "thoracotomy"), our search performed between July and August 2018 yielded a total of 18,268 articles. The top 100 articles were identified subsequently which were ranked by the number of citations.

#### Analysis of the top 100 cited articles

A further analysis of individual articles was carried out by three reviewers (J Shen, K Jin, and Q Hu) to extract relevant information for the analysis. Individual articles were reviewed to identify the authorship, journal title, journal IF, subject matter, article type (e.g., prospective analysis, retrospective analysis, meta-analysis or systematic review, randomized controlled trial, review article, guideline, technical note, or case report), institution and country of origin, level of evidence, and year of publication. The analysis method used the Australian National Health and Medical Research Council evidence hierarchy (11). Of note, this classification defines clinical guidelines as level I evidence and randomized control trials as level II evidence.

Furthermore, the citation count and citations per year were extracted using the Web of Science. When there was disagreement or discordance between reviewers, the senior author (D Ma) was included to achieve consensus.

#### Results

The top 100 articles regarding the thoracic surgery management of lung cancer according to the most citations received are listed in *Table 1*.

"Cisplatin-based adjuvant chemotherapy in patients with completely resected non-small-cell lung cancer", published in *The New England Journal of Medicine* in 2004, has received totally 1,720 citations, the most citations in the past 14 years. Of note, it also remains the most cited article per year, being cited 122.86 times on average, per year. The article with the lowest number of citations in the top 100 list, "A practical molecular assay to predict survival in resected non-squamous, non-small-cell lung cancer: Development and international validation studies", received 177 citations. Overall, the mean number of citations was 356, with the top 100 articles cited an average of 26 times per year.

Only one of the top 100 articles was published prior to 1980 (*Table 2*), with 63% (n=63) published since the year 2000 and 13% published within the past decade. Eighteen authors contributed four or more articles (*Table 3*). Rusch, Ginsberg, Johnson, and Livingston have contributed the most articles. Rusch and Ginsberg were the first named authors on four articles each. Rusch is the most published author, appearing on a total of ten articles.

A total of 31 journals made contributions to the top 100 articles with 13 journals providing two or more articles. Of note, four journals provided ten or more articles (*Table 4*). The journals included *Journal of Clinical Oncology* (26.303), *Journal of Thoracic and Cardiovascular Surgery* (4.88), *Annals of Thoracic Surgery* (3.779) and *Chest* (IF 7.562).

A total of 14 countries produced the entirety of the top 100 most cited articles (*Figure 1*), with the majority originating in the USA (43%). Japan, the next highest contributor, contributed nine articles. One institution contributed the most to the top 100 (*Table 5*). The Memorial Sloan-Kettering Cancer Center contributed 18 articles while the University of California contributed nine. Overall, 30 institutions contributed three or more articles each.

Topics covered in each article are grouped and outlined in *Table 6*. Subject matter included prognosis (n=21), surgical type (n=20), adjuvant therapy (n=17), perioperative outcome (n=10), and neo-adjuvant therapy studies (n=10).

Table 7 and Figure 2 summarize the type of clinical study and level of evidence within the top 100 articles. Most of the evidence is level III or IV, with 18 original prospective and 40 original retrospective clinical studies. Overall, there were 42 prospective studies which included 24 level II randomized controlled trials. Five commentaries and eight reviews were included in the top 100 cited articles.

#### **Discussion**

In recent decades, the thoracic surgery management for lung cancer has expanded dramatically. This bibliometric analysis emphasized the top 100 articles relating to the

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Table 1 The 100 'citation classics' n	regarding the field of lung cancer surgery and the number of calculated annual c	itations

Rank	Article	Citations	Citations per year	PMID
1	Cisplatin-Based Adjuvant Chemotherapy in Patients with Completely Resected Non-Small-Cell Lung Cancer	1,720	122.86	14736927
2	Randomized trial of lobectomy versus limited resection for T1 N0 non-small cell lung cancer	1,623	70.57	7677489
3	Vinorelbine plus cisplatin vs. observation in resected non-small-cell lung cancer	1,372	105.54	1597286
4	A Randomized Trial Comparing Preoperative Chemotherapy Plus Surgery with Surgery Alone in Patients with Non-Small-Cell Lung Cancer	1,095	45.63	8043059
5	Adjuvant vinorelbine plus cisplatin versus observation in patients with completely resected stage IB-IIIA non-small-cell lung cancer (Adjuvant Navelbine International Trialist Association [ANITA]): a randomised controlled trial	1,014	84.50	1694576
6	A randomized trial comparing perioperative chemotherapy and surgery with surgery alone in resectable stage iiia non-small-cell lung cancer	962	40.08	8156698
7	Incidence of local recurrence and second primary tumors in resected stage I lung cancer	763	33.17	7815787
8	Concurrent cisplatin/etoposide plus chest radiotherapy followed by surgery for stages IIIA(N2) and IIIB non-small-cell lung cancer: Mature results of Southwest Oncology Group Phase II Study 8805	746	32.43	7636530
9	Radiotherapy plus chemotherapy with or without surgical resection for stage III non-small-cell lung cancer: a phase III randomised controlled trial	739	82.11	1963271
10	Prognosis and survival in resected lung carcinoma based on the new international staging system	633	21.10	2842549
11	Lymph node mapping and curability at various levels of metastasis in resected lung cancer	598	14.95	713589
12	Preoperative chemotherapy followed by surgery compared with primary surgery in resectable stage I (except T1N0), II, and IIIa non-small-cell lung cancer	523	32.69	1177317
13	Stereotactic ablative radiotherapy versus lobectomy for operable stage I non-small-cell lung cancer: A pooled analysis of two randomised trials	521	173.67	2598181
14	Mutations in the p53 gene are frequent in primary, resected non-small cell lung cancer	515	18.39	1979160
15	Modern thirty-day operative mortality for surgical resections in lung cancer	501	14.31	6632940
16	The influence of hospital volume on survival after resection for lung cancer	470	27.65	1146301
17	Randomized study of adjuvant chemotherapy for completely resected stage I, II, or IIIA non- small-cell lung cancer	445	29.67	1451975
18	Randomized controlled trial of resection versus radiotherapy after induction chemotherapy in stage IIIA-N2 non-small-cell lung cancer	439	39.91	1776233
19	A randomized trial of postoperative adjuvant therapy in patients with completely resected stage II or IIIA non-small-cell lung cancer	419	23.28	1135558
20	Morbidity and mortality of major pulmonary resections in patients with early-stage lung cancer: Initial results of the randomized, prospective ACOSOG Z0030 trial	418	34.83	1648871
21	Systematic review and meta-analysis of randomized and nonrandomized trials on safety and efficacy of video-assisted thoracic surgery lobectomy for early-stage non-small-cell lung cancer	406	45.11	1928962
22	Radical sublobar resection for small-sized non-small cell lung cancer: A multicenter study	402	33.50	1700028
23	Guidelines on the selection of patients with lung cancer for surgery	394	23.18	1120909
24	Long-term follow-up of patients enrolled in a randomized trial comparing perioperative chemotherapy and surgery with surgery alone in resectable stage IIIA non-small-cell lung cancer	392	19.60	9792048
25	Survival of patients with resected N2 non-small-cell lung cancer: Evidence for a subclassification and implications	384	21.33	1094413

Table 1 (continued)

Rank Article		Citations	Citations per year	PMID
26	ERS/ESTS clinical guidelines on fitness for radical therapy in lung cancer patients (surgery and chemo-radiotherapy)	382	42.44	19567600
27	Effects of Postoperative Mediastinal Radiation on Completely Resected Stage II and Stage III Epidermoid Cancer of the Lung	375	11.72	2877397
28	The role of surgery in N2 lung cancer	357	11.52	3629423
29	Surgery for Early-Stage Non-Small Cell Lung Cancer: A Systematic Review of the Video- Assisted Thoracoscopic Surgery Versus Thoracotomy Approaches to Lobectomy	345	34.50	19022040
30	Physiologic evaluation of the patient with lung cancer being considered for resectional surgery: ACCP evidenced-based clinical practice guidelines (2nd edition)	338	30.73	17873167
31	Stereotactic body radiotherapy (SBRT) for operable Stage i non-small-cell lung cancer: Can SBRT be comparable to surgery?	328	46.86	20638194
32	Increased MET gene copy number negatively affects survival of surgically resected non-small- cell lung cancer patients	320	35.56	19255323
33	The American Association for Thoracic Surgery guidelines for lung cancer screening using low- dose computed tomography scans for lung cancer survivors and other high-risk groups	308	51.33	22710039
34	Preoperative chemotherapy followed by concurrent chemoradiation therapy based on hyperfractionated accelerated radiotherapy and definitive surgery in locally advanced non- small-cell lung cancer: Mature results of a phase II trial	307	15.35	9469351
35	Neutrophil/lymphocyte ratio and its association with survival after complete resection in non- small cell lung cancer	302	33.56	19185164
36	Wedge resection versus lobectomy for stage I (T1 N0 M0) non-small-cell lung cancer	299	14.24	9104978
37	The influence of hospital and surgeon volume on in-hospital mortality for colectomy, gastrectomy, and lung lobectomy in patients with cancer	296	18.50	11812957
38	Preoperative F-18 fluorodeoxyglucose-positron emission tomography maximal standardized uptake value predicts survival after lung cancer resection	292	20.86	15310769
39	Results of resection in non-oat cell carcinoma of the lung with mediastinal lymph node metastases	289	8.26	6615059
40	Outcomes after stereotactic lung radiotherapy or wedge resection for stage I non-small-cell lung cancer	283	35.38	20065181
41	A Japanese lung cancer registry study: Prognosis of 13,010 resected lung cancers	280	28.00	18166840
42	Prognostic Significance of p53 Mutations and 3p Deletions in Primary Resected Non-Small Cell Lung Cancer	280	11.20	8380124
43	Adjuvant chemotherapy after complete resection in non-small-cell lung cancer	275	12.50	23931022
44	Video-assisted thoracoscopic lobectomy achieves a satisfactory long-term prognosis in patients with clinical stage IA lung cancer	266	14.78	10594199
45	Prognostic and predictive gene signature for adjuvant chemotherapy in resected non-small- cell lung cancer	264	33.00	20823422
46	Potential impact on survival of improved tumor downstaging and resection rate by preoperative twice-daily radiation and concurrent chemotherapy in stage IIIA non-small-cell lung cancer	263	12.52	20823422
47	Risk Factors for Acute Lung Injury after Thoracic Surgery for Lung Cancer	261	17.40	14633519
48	Evaluation of epidermal growth factor-related growth factors and receptors and of neoangiogenesis in completely resected stage I-IIIA non- small-cell lung cancer: Amphiregulin and microvessel count are independent prognostic indicators of survival	256	12.80	9516978
49	Induction chemoradiation and surgical resection for superior sulcus non-small-cell lung carcinomas: Long-term results of Southwest Oncology Group trial 9416 (Intergroup trial 0160)	254	23.09	17235046
50	Sites of recurrence in resected stage I non-small-cell lung cancer: A guide for future studies	251	7.38	6512581

Table 1 (continued)

Table 1 (continued)

Rank Article		Citations	Citations per year	PMID
51	Lobectomy by video-assisted thoracic surgery (VATS) versus thoracotomy for lung cancer	246	27.33	19577048
52	Intentional limited resection for selected patients with T1 N0 M0 non- small-cell lung cancer: A single-institution study	244	11.62	9305186
53	The role of extrapleural pneumonectomy in malignant pleural mesothelioma. A lung cancer study group trial	243	9.00	2072706
54	Patient and hospital characteristics related to in-hospital mortality after lung cancer resection	240	9.23	1582293
55	Morbidity and mortality after neoadjuvant therapy for lung cancer: The risks of right pneumonectomy	238	14.00	11603428
56	ERCC1 expression is a predictor of survival in resected patients with non-small cell lung cancer	238	18.31	15764785
57	Effect of tumor size on prognosis in patients with non-small cell lung cancer: The role of segmentectomy as a type of lesser resection	236	18.15	15632829
58	Association of PD-L1 overexpression with activating EGFR mutations in surgically resected nonsmall-cell lung cancer	236	59.00	25009014
59	miR-34a as a prognostic marker of relapse in surgically resected non-small-cell lung cancer	234	26.00	19736307
60	Segmental resection spares pulmonary function in patients with stage I lung cancer	233	16.64	15223434
61	18F-FDG uptake as a biologic prognostic factor for recurrence in patients with surgically resected non-small cell lung cancer	232	14.50	11801701
62	The Importance of Surgery to Non-Small Cell Carcinoma of Lung with Mediastinal Lymph Node Metastasis	231	7.70	2848463
63	The physiologic evaluation of patients with lung cancer being considered for resectional surgery	229	15.27	12527570
64	Induction chemotherapy before surgery for early-stage lung cancer: A novel approach	229	12.72	10694600
65	Outcomes of Sublobar Resection Versus Lobectomy for Stage I Non-Small Cell Lung Cancer: A 13-Year Analysis	226	18.83	16863738
66	Randomized phase III trial of vinorelbine plus cisplatin compared with observation in completely resected stage IB and II non-small-cell lung cancer: Updated survival analysis of JBR-10	226	28.25	19933915
67	Intentional limited pulmonary resection for peripheral T1 N0 M0 small-sized lung cancer	226	15.07	12698157
68	Long-term results of the international adjuvant lung cancer trial evaluating adjuvant cisplatin- based chemotherapy in resected lung cancer	223	27.88	19933916
69	A Phase III randomized trial of lobectomy versus limited resection for small-sized peripheral non-small cell lung cancer (JCOG0802/WJOG4607L)	222	24.67	19933688
70	Similar long-term survival of elderly patients with non-small cell lung cancer treated with lobectomy or wedge resection within the surveillance, epidemiology, and end results database	222	17.08	16002941
71	Physiologic evaluation of the patient with lung cancer being considered for resectional surgery: Diagnosis and management of lung cancer, 3rd ed: American college of chest physicians evidence-based clinical practice guidelines	221	44.20	23649437
72	Lymph node involvement, recurrence, and prognosis in resected small, peripheral, non-small- cell lung carcinomas: Are these carcinomas candidates for video-assisted lobectomy?	220	10.00	8642812
73	Induction chemoradiation and surgical resection for non-small cell lung carcinomas of the superior sulcus: Initial results of southwest oncology group trial 9416 (Intergroup Trial 0160)	217	12.76	11241082
74	A controlled study of postoperative radiotherapy for patients with completely resected non- small cell lung carcinoma	214	11.26	10994397
75	Is segmentectomy with lymph node assessment an alternative to lobectomy for non-small cell lung cancer of 2 cm or smaller?	212	12.47	11269480

Table 1 (continued)

Table 1 (continued)

Rank	Article	Citations	Citations per year	PMID
76	A prospective randomized trial to determine the benefit of surgical resection of residual disease following response of small cell lung cancer to combination chemotherapy	212	8.83	7988254
77	Neoadjuvant chemotherapy and radiotherapy followed by surgery in stage IIIA non-small-cell carcinoma of the lung: Report of a cancer and leukemia group B phase II study	212	8.15	1321893
78	Prognostic factors in patients with resected stage I non-small cell lung cancer. A report from the Lung Cancer Study Group	211	6.21	6478416
79	What happens to patients undergoing lung cancer surgery? Outcomes and quality of life before and after surgery	211	13.19	1211433
80	Thoracoscopic Lobectomy Facilitates the Delivery of Chemotherapy after Resection for Lung Cancer	206	18.73	1738332
81	Prognostic significance of CCND1 (cyclin D1) overexpression in primary resected non-small- cell lung cancer	205	9.32	8562333
82	Prognosis of 6644 resected non-small cell lung cancers in Japan: A Japanese lung cancer registry study	203	15.62	1606130
83	Complete resection in lung cancer surgery: Proposed definition	202	15.54	1594958
84	Prognostic assessment of 2,361 patients who underwent pulmonary resection for non-small cell lung cancer, stage I, II, and IIIA	201	11.17	1066967
35	Evaluation of microRNA expression profiles that may predict recurrence of localized stage I non-small cell lung cancer after surgical resection	200	25.00	2002885
36	Epidermal growth factor receptor overexpression correlates with a poor prognosis in completely resected non-small-cell lung cancer	199	14.21	1467911
37	Preoperative risk evaluation for lung cancer resection: Predicted postoperative product as a predictor of surgical mortality	198	8.25	792146
38	Adjuvant MAGE-A3 immunotherapy in resected non-small-cell lung cancer: Phase II randomized study results	197	39.40	2371556
39	Video-assisted thoracic surgery versus open lobectomy for lung cancer: A secondary analysis of data from the American College of Surgeons Oncology Group Z0030 randomized clinical trial	196	24.50	2017253
90	Impact of Postoperative Radiation Therapy on Survival in Patients With Complete Resection and Stage I, II, or IIIA Non-Small-Cell Lung Cancer Treated With Adjuvant Chemotherapy: The Adjuvant Navelbine International Trialist Association (ANITA) Randomized Trial	196	19.60	1843976
91	Video-Assisted Thoracoscopic Surgery is More Favorable Than Thoracotomy for Resection of Clinical Stage I Non-Small Cell Lung Cancer	195	17.73	1753237
92	Resection rates and postoperative mortality in 7,899 patients with lung cancer	194	8.82	883432
93	Complications of surgery in the treatment of carcinoma of the lung	191	5.31	708393
94	Radiotherapy as an alternative to surgery in elderly patients with resectable lung cancer	185	6.17	284914
95	Postoperative radiation therapy in lung cancer: A controlled trial after resection of curative design	183	4.82	699893
96	BRCA1: A novel prognostic factor in resected non-small-cell lung cancer	182	16.55	1798711
97	Surgical resection of stage IIIA and stage IIIB non-small-cell lung cancer after concurrent induction chemoradiotherapy: A Southwest Oncology Group trial	182	7.28	838047
98	Global histone modifications predict prognosis of resected non-small-cell lung cancer	181	16.45	1790620
99	Cancer recurrence after resection: T1 N0 non-small cell lung cancer	181	6.46	215495
100	A practical molecular assay to predict survival in resected non-squamous, non-small-cell lung cancer: Development and international validation studies	177	29.50	2228505

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management of NSCLC in the areas of prognosis, surgical type, adjuvant and neoadjuvant therapy, perioperative outcome, recurrence, and second primary tumors. It highlights the high impact literature available for reference in practicing lung cancer therapy. Furthermore, our list of the top 100 articles includes selections which have been

 Table 2 The number of articles in the top 100 most cited articles by decade of publication

Decade	No. of article	
1960–1969	0	
1970–1979	1	
1980–1989	11	
1990–1999	25	
2000–2009	50	
2010–2018	13	

published across the spectrum of thoracic surgery, including anesthesia, nuclear medicine, radiology, and respiratory medicine, highlighting the multidisciplinary approach to lung cancer.

A prospective multi-institutional trial by Ginsberg *et al.* compared lobectomy with sublobectomy, and showed that locoregional recurrence was lower in the lobectomy group (12). It has remained highly cited (Web of Science 1623) but also historically relevant (70.23 citations per year). This work provided a guide for the choice of surgery type in clinical practice. Whitson *et al.* in 2007 demonstrated that VATS lobectomy for patients with clinical stage I NSCLC appeared to be a less morbid operation (Web of Science 195, 17.73 citations per year) (13). Moreover, a meaningful meta-analysis suggests that VATS lobectomy is an appropriate procedure for selected patients with early-stage NSCLC when compared with open surgery (Web of Science 406, 45.11 citations per year) (14).

Table 3 The authors that contributed four or more articles to the 100 most cited articles in lung cancer surgery

Author	No. of articles	Position on author list
Rusch VW	10	First [4], second [1], third [1], sixth [1], ninth [1], tenth [1], twelfth [1]
Ginsberg RJ	5	First [2], second [2], seventh [1]
Johnson DH	5	Sixth [1], seventh [2], ninth [1], thirteenth [1]
Livingston RB	5	Seventh [1], eighth [2], eleventh [1], fifteenth [1]
Tsuchiya R	5	Third [2], fourth [1], sixth [1], eighth [1]
Asamura H	4	First [2], second [1], fifth [1]
Downey RJ	4	First [1], fourth [1], fifth [1], eighth [1]
Gandara D	4	Sixth [1], twelfth [1], fourteenth [1], eighteenth [1]
Giaccone G	4	Second [1], fifth [1], sixth [1], twelfth [1]
Goya T	4	First [1], second [3]
Jassem J	4	Ninth [1], fifteenth [1], eighteenth [1], thirty-fifth [1]
Komaki R	4	Third [1], third [1], fifth [1], twenty-second [1]
Landreneau RJ	4	First [1], second [1], sixth [1], ninth [1]
Le Chevalier T	4	Fourth [1], tenth [1], twelfth [1], thirteenth [1]
Martini N	4	First [3], third [1]
Naruke T	4	First [3], sixth [1]
Okada M	4	First [3], fourth [1]
Rosell R	4	First [2], second [2]

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Table 4 The number of articles each journal contributed to the top 100 articles

Journal	No. of 100 top articles	Impact factor (2017–2018)
Journal of Clinical Oncology	18	26.303
Journal of Thoracic and Cardiovascular Surgery	17	4.88
Annals of Thoracic Surgery	11	3.779
Chest	10	7.652
New England Journal of Medicine	6	79.258
International Journal of Radiation Oncology Biology Physics	3	5.554
Journal of the National Cancer Institute	3	11.238
Lung Cancer	3	4.486
Annals of Oncology	2	13.926
Cancer	2	6.537
Cancer Research	2	9.13
European Respiratory Journal	2	12.242
Lancet	2	53.254
American Journal of Respiratory and Critical Care Medicine	1	15.239
Anesthesia and Analgesia	1	3.463
Annals of Surgery	1	9.203
British Journal of Cancer	1	5.922
Carcinogenesis	1	5.072
Clinical Cancer Research	1	10.199
Japanese Journal of Clinical Oncology	1	2.37
Journal of Nuclear Medicine	1	7.439
Journal of Thoracic and Cardiovascular Surgery	1	4.88
Journal of Thoracic Oncology	1	10.336
Lancet Oncology	1	36.418
Oncogene	1	6.854
PLoS ONE	1	2.766
Radiotherapy and Oncology	1	4.942
Surgery	1	3.574
Surgical Clinics of North America	1	1.947
Thorax	1	9.655
World Journal of Surgery	1	2.766

Regarding neoadjuvant and adjuvant chemotherapy, notably in 1994, Rosell *et al.* described how the preoperative chemotherapy increases the median survival in patients with stage IIIA NSCLC (Web of Science 1095, 45.63 citations per year) (15). In 2004, the International Adjuvant Lung Cancer Trial Collaborative Group concluded that cisplatinbased adjuvant chemotherapy improves survival among patients with completely resected non-small cell lung



**Figure 1** A total of 14 countries produced the entirety of the top 100 most cited articles, with the majority originating in the USA (43%); Japan, the next highest contributor, contributed nine articles; Australia, Belgium, Switzerland, and Hungary all contributed a single article each.

**Table 5** The institutions that contributed the most articles to 100most cited articles in the lung cancer surgery

Institution	No. of articles
Cornell University Medical Center	21
University of California	12
University of Toronto	17
National Cancer Center Hospital, Japan	7
University of Texas M.D. Anderson Cancer Center	7
University of Washington	6
Mayo Clinic	6
Fred Hutchinson Cancer Research Center	5
Brigham and Women's Hospital	4
Dana-Farber Cancer Institute	4
Vanderbilt University	4
Evanston Hospital	4
Allegheny General Hospital	3
Fox Chase Cancer Center	3
Hospital Germans Trias i Pujol	3
Hyogo Medical Center for Adults	3
Medical University of South Carolina	3
Mount Sinai Hospital	3
National Cancer Institute, Bethesda	3
Swedish Medical Center	3
University of Barcelona	3
University of North Carolina	3
Veterans Administration Medical Center	3
Institut Gustave Roussy	3
H. Lee Moffitt Cancer Center & Research Institute	3

cancer (Web of Science 1720, 122.86 citations per year) (16). In 2006, Douillard *et al.* compared the effect of adjuvant vinorelbine plus cisplatin versus observation on survival in patients with completely resected NSCLC [Adjuvant Navelbine International Trialist Association (ANITA)], better defining indication of adjuvant chemotherapy (Web of Science 1014, 84.50 citations per year) (17).

A bibliographic analysis aiming to identify and review the 100 most-cited papers in general thoracic surgery was published in 2018 (18). Ding et al. focused on surgical treatment of diseases in the lung, esophagus, trachea, pleura, chest wall and mediastinum and offered insights into the development and advances of general thoracic surgery which could help us understand the evidential basis of clinical decision-making today in this area. However, to our knowledge, there is no other bibliographic analysis of the literature surrounding the thoracic surgery management of lung cancer using Web of Science results. Our analysis had some strengths, including the focused analysis of the thoracic surgery management spanning a range of medical disciplines, including many topics not discussed in prior analyses. The analysis contained a high proportion of original articles, including a higher proportion of randomized controlled trials compared with other similar bibliographic analyses.

However, there are several limitations which should be considered in this bibliometric analysis. While providing a literature overview, the analysis does not provide or allow for any conclusions as to the quality or merit of individual articles. Older articles are more likely to be cited regardless of their impact or relevance accumulating more citations over time. Importantly, many important discoveries do not

**Table 6** The main topics discussed within the 100 most cited articles in lung cancer surgery

Subject	No. of articles
Prognosis	21
Surgical type	20
Adjuvant therapy	17
Perioperative outcome	10
Neoadjuvant therapy	10
Radiotherapy vs. surgery	6
Recurrence and second primary tumors	5
Guidelines	5
Other	6

 Table 7 Levels of evidence and article type composing the 100 most cited articles in lung cancer surgery

Level of evidence	No. of articles	
I	Guidelines	5
Ш	Randomized controlled trial	24
Ш	Original prospective	18
IV	Original retrospective	40
V	Commentary	5
	Review	8

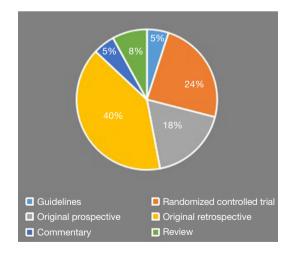


Figure 2 A summary of the levels of evidence composing the 100 most cited articles in the thoracic surgery management of lung cancer.

make the top 100 list as their findings are often accepted as general knowledge by the scientific community and subsequently not heavily cited. Furthermore, there is often an influence on citation patterns due to citation bias and social relations within the scientific community (including self-citations), influence by publication structures, and low rates of non-English language citations.

#### **Questions to be further considered:**

### What are the future directions of surgical treatment of lung cancer?

#### Expert opinion 1: Dr. Michael K. Hsin

There are many trends which suggest the likely pathway of the future of surgical treatment of lung cancer. I will address a few of such trends which I believe will be particularly significant.

The increasingly common findings of subcentimeter lesions and GGOs, as a result of lung cancer screening, or just incidental findings as CT scans are performed with increasing frequency.

This will lead to the following issues:

- (I) While the findings of the two large multicenter RCTS that address the issue of sublobar resection versus lobectomy—the Japanese RCT: JCOG0802/ WJOG4607L, and the North American RCT CALGB 140503—are eagerly anticipated, we are already witnessing an increasingly common use of sublobar resection in the thoracic surgical community for the treatment of small lesions, including GGOs.
- (II) Following on the previous point, there will be increasing effort to develop sophisticated and reliable means for the accurate localization of these small lung lesions, such as the routine use of 3 D CT reconstruction in the preoperative planning, electromagnetic navigational bronchoscopy with or without intraoperative real time CT imaging (19).
- (III) The incorporation of ablative therapy into the armamentarium of treatment for small lung cancers in appropriately selected patients. These include cryotherapy, microwave ablation and radiofrequency ablation, which may be administered percutaneously, via endobronchial approach, or minimally invasive surgery.
- (IV) The advent of immunotherapy has significantly altered the landscape of advanced stage lung cancer treatment.

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There are already a sleuth of clinical trials looking is at the role of neoadjuvant immunotherapy in resectable lung cancer. For example, the NEOMUN trial looks at neoadjvant Pembrolizumab as part of a multimodality treatment in resectable Stage II/IIIA NSCLC (20). The role of neoadjuvant PD-1 blockade in resectable early stage lung cancer has been reported in pilot studies (21). However, the role of immunotherapy for early stage resectable lung de

cancer will be better defined by the results of phase 3 studies with long term outcomes by ongoing studies such as Checkmate 816, IMpower030, and KEYNOTE-671.

#### Expert opinion 2: Dr. Carmelina C. Zirafa

Since the introduction of thoracoscopy, at the beginning of the 20th century, the surgical treatment of lung cancer has been characterized by a reduction of invasiveness, in terms of surgical approach and impact on the patient's life. Thanks to the improvement of technology, currently, it is possible to perform also complex procedures by minimally invasive surgery, represented by video-assisted thoracic surgery (VATS) and robotic surgery. In VATS the introduction of even more peculiar and technologically advanced devices, like 3D high definition view and energy devices, enables to do major surgical procedure routinely (22). Moreover, robotic surgery, that is considered the evolution of VATS in terms of technology, permits to approach also complex surgical operations in a safe and comfortable way. In fact, chest wall or vascular invasion, neoadjuvant therapy or prior thoracic surgery are no longer considered contraindications to robotic surgery, as described by various Authors (23).

Furthermore, integration of advanced technological devices will be able to support challenging operations, like electromagnetic-navigational bronchoscopy or fluorescent imaging system (24). In addition, the recent introduction in the practical field of the 3D printing, a promising radiologic model, could become a consistent support for surgeons to plan the surgical procedure tailored to the patient's case, to reduce time of surgery and be that extra aid that might significantly reduce intraoperative complications (25). In future, 3D printing could represent a stimulating basis for educational program and as well as the new frontier for biomaterial implant (26).

The development of minimally invasive technique is one of the main elements of the new approach to surgery, in which the fulcrum is the quality of life of patients. Consequently, diffusion of programs such as ERAS and the reduction of invasiveness during the anesthesiologic phase, in particular intubation and analgesia management, is fundamental (27,28).

Considering this background, the future of surgery in lung cancer appears to be exponentially less traumatic and technologically advanced.

#### Expert opinion 3: Dr. Nuria M. Novoa

Probably, I would highlight three main lines in the future development around surgical treatment of lung cancer.

The first is related to the controversy: lobectomy versus sublobar resection. Currently, we have sound retrospective data supporting lobectomy over any other sublobar resection whenever the patient can tolerate the resection and even for 1cm tumors scenario (29). However, prospective data are mandatory to determine the role of each technique and the best strategy in different groups of patients. Two randomized control ongoing studies had been designed to assess this problem the CALGB 140503 and the Japanese study (JCOG0802/ WJOG4607L) (30). The first study is currently reporting initial results on morbimortality (31) and discussing the need of a preoperative diagnosis for a correct enrollment and assignment (32) but no data on overall survival and local relapse, the most relevant issues, had been published yet. The second is still on going and no initial results had been reported. I look forward with great interest to the results of these studies.

The second pertinent line is related to the impact of the developing immunotherapy on surgical treatments. Again, very promising results had been reported (21) but further information is needed. And, finally, lung cancer screening. Although the results of the NLST (33) were very relevant and induced the introduction of the screening in the USA at a national scale, the raised a lot of unsolved questions that other posterior trials are trying to clarify (34).

#### Expert opinion 4: Dr. Stefano Bongiolatti

Nowadays the treatment options for early stage or metastatic stage NSCLC are well-established while some controversial issue are present regarding the multimodal approach on locally advanced stage (IIIA). From an historical point of view, the thoracic surgeon evolved from thoracotomic lobectomy to minimally invasive anatomical pulmonary resection (VATS, RATS, uniportal, subxyphoidal etc.) while the oncological treatment has evolved to a more targeted therapy. In my opinion, these targeted strategies (EGFR-TKI, ALK-inhibitors, ROS1-TKI, immunotherapy) associated with local treatment will be the new direction in the multimodal approach of locally advanced NSCLC. However, some randomized controlled trial and multi-institutional studies are necessary to demonstrate the safety, feasibility and benefits of this intention treatment approach.

#### Expert opinion 5: Dr. Robert J. Cerfolio

Almost all of lung cancer surgery will be done minimally invasively over the next few years. In our own practice, we have already essentially achieved a 100% robotic approach for all lung cancer, including patients that had chemo and radiation, patients who require a sleeve of the artery or the bronchus and even patients who have tumors greater than 10 cm. We make it a daily work now with zero conversions in our last 200–300 robotic procedures. In the next 5–7 future years, all T1A and T1b tumors will all be treated with bronchoscopic intervention either using heat or thermal or other ablative forms of treatment via the bronchoscope and/or using the injection of immunotherapy.

## Should guidelines be established for early-stage lung cancer, especially for GGO?

#### Expert opinion 1: Dr. Michael K. Hsin

There are two aspects to this question:

- (I) The management of a GGO where there is no histological diagnosis.
- (II) The surgical treatment, if deemed appropriate, of GGO that has been proven to be malignant.

There are already several well-established guidelines on the management of GGO without histological diagnosis, such as the Fleischner society and NCCN guidelines, and no doubt as we gain more experience, new guidelines will be written to reflect the accumulated knowledge.

The management of GGOs known to have malignant potential is fraught with difficulty. What is missing is not so much guideline but good evidence. The main issue centers around whether to perform a lobectomy or a sublobar resection; and for sublobar resection, then which typesegmentectomy or wedge resection?

The decision leading to the appropriate management of GGOs should include the following considerations:

- Consolidation-to-tumour ratio.
- Impact of spread through air space.
- Intraoperative lymph node sampling.
- The place of intraoperative frozen section for resection margins and lymph nodes.
- Solitary GGO vs. multiple GGOs vs. GGO with

concurrent proven lung cancer in a different lobe (same side/different side).

#### Expert opinion 2: Dr. Carmelina C. Zirafa

The increasingly frequent detection of small pulmonary lesions, due to latest enhancement of screening programs or to incidental detection, has encouraged the diagnosis and treatment of earlier stages of lung cancer. The recurring undefined characteristics of these small lesions, among which the GGO have taken on more relevance in the recent years, have made it necessary to define some guidelines. Several national and international guidelines have been published in the last years, albeit sometimes, disagreeing on the management of small or stable GGOs (35).

The optimisation and homogeneity of indications of different guidelines are essential to limit surgical resection in benign lesions or to avoid delay in diagnosis in case of malignancy. In addition, there are still some confounding elements in the evaluation of indeterminate small lesions, like the presence of multiple lesions, the indication to nonsurgical biopsy and which is better surgical strategy to be applied in order to avoid overtreatment (36).

Specifically, guidelines are lacking in surgical indications. Minimally invasive surgery should be recommended for resection, although depending on the dimensions of these lesions the support of intraoperative localization techniques (e.g., radio-guided resection, near-infrared fluorescence localization, hook wire placement) is required. Currently, some important points of discussion, not yet fixed, in GGO are the kind of pulmonary resection to be preferred (wedge resection vs. anatomical resection) and the role of lymphadenectomy (37). According to the proposal of JCOG (Japan Clinical Oncology Group) a wedge resection should be favoured for lesions smaller than 2 cm, whereas larger lesions require anatomical resection (segmentectomy, lobectomy). Whilst waiting for the results of randomized trials and updated guidelines, the treatment of undefined lesion should be always evaluated by a multidisciplinary oncology board.

#### Expert opinion 3: Dr. Nuria M. Novoa

There is no doubt that GGO are a special subgroup of lung cancer patients that need specific considerations. For that reason, a guideline is needed to pool all the current information allowing to design a clear management strategy for these patients. The European Society of Thoracic Surgery (ESTS) is currently collaborating in the development of this guideline.

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#### Expert opinion 4: Dr. Stefano Bongiolatti

The guidelines for every stage NSCLC are a continuous and strenuous process to define the clinical and pathological stage in order to find a right treatment choice. In the last years, with the diffusion of CT-screening programs, new radiological findings (for example ground glass opacities, part-solid nodules, solid nodules associated with cystic airspaces) were frequently identified and are associated with specific subtypes of NSCLC, in particular adenocarcinoma. Furthermore, these adenocarcinoma subtypes are correlated with particular survival patterns. In my opinion, the current international guidelines must consider these features, but it is necessary to clear some aspects of multiple or multifocal ground glass opacity that could be locally treated as primary cancer, instead of a metastatic disease.

#### Expert opinion 5: Dr. Robert J. Cerfolio

Yes, there is a plethora of patient with GGO, especially of non-smoking women, and most of them have several in their lungs. Although we only focus on the invasive or solid component of these tumors, but certainly overtime we are going to get more and more information about how to optimize therapy.

#### The role of surgical treatment in stage IV lung cancer.

#### Expert opinion 1: Dr. Michael K. Hsin

Survival for patients with stage IV NSCLC is poor. Fiveyear survival is 1% (38). Surgery can be offered for patients with stage IV lung cancer for either palliation or in highly selected patients with the goal of improving prognosis.

The indications for palliation are well established, and include drainage and pleurodesis for malignant pleural effusion, pericardial window for pericardial effusion, airway intervention such as disobliteration/dilatation and stenting for central airway obstruction.

In terms of surgery for improving prognosis in stage IV lung cancer, it remains controversial. There is limited amount of published evidence most of which are from small single center series and focus on patients with oligometastasis. The number of patients with oligometasis remains low and is estimated to be 2-7%(39,40). In general, the fewer the metastasis, the better the prognosis. In patients with solitary metastasis, 5-year survival close to 50% has been achieved after surgery, whereas when there are five metastases, this dropped to 17% after surgery (41). The presence or absence of nodal disease can affect survival significantly. Tamura reported that for Stage IV N0 NSCLC patients, median survival was 18 months after surgery. However, when there is nodal metastasis as well, the median survival after surgery was 11 months (42).

Most of the publications on surgery in Stage IV NSCLC patients pertain to brain or adrenal metastasis, and it should be emphasised that these represent highly selected patient populations. The best results of aggressive treatment in oligometastasis to the brain is probably by Daniels, who reported that after lung resection and resection of the brain metastasis, 5-year survival of 60% was achieved (43). Lanuti reviewed published series where there was synchronous oligometastasis NSCLC to the brain, and concluded that 5-year survival averages 21% (range 1–35) (44).

For NSCLC with oligometastasis to the adrenal, in carefully selected patients, Raz et al. reported a 5-year overall survival of 34a% for patients who underwent resection of adrenal metastasis, and 0% for those managed non-surgically. Worse survival was seen in patients with contralateral adrenal metastasis or mediastinal lymph node involvement (45). In a systematic review by Tanvetvanon et al. where they performed a pooled analysis of 114 patients from 10 published series, they showed that median overall survival was shorter for patients with synchronous metastasis than those with metachronous metastasis (12 months vs. 31 months). However, the 5-year survival estimates were equivalent at 26% and 25%, respectively (46). The ACCP evidence-based guideline on oligometastatic adrenal metastases recommend that in patients with a synchronous resectable N0, N1 primary NSCLC and an isolated adrenal metastasis with no other sites of metastases, resection of the primary tumor and the adrenal metastasis is recommended. In patients who have undergone a curative resection of an isolated adrenal metastasis, adjuvant chemotherapy is suggested (47).

The role of surgery in extracranial, extra-adrenal NSCLC oligometastasis is less clear. Salah performed a systematic review on patients who underwent resection of the metastasis and definitive treatment of the lung primary (n=62). The 5-year survival rate was 50% for the entire cohort. Mediastinal lymph node involvement was independently predictive of inferior outcome. Similarly, patients with intra-thoracic stage III disease had an inferior outcome compared to patients with stage II and stage I disease. Other factors including site of metastasis have no effect on outcome (48).

#### NSCLC in contralateral lung

According to the 8<sup>th</sup> edition of the IASLC classification, NSCLC in the contralateral lung is grouped under M1a. It is not always possible to differentiate between a metastatic disease, or multifocal localized disease, which have significantly different prognostic implications. Bertolaccini advocate that in a fit patient with a full preoperative staging including also a negative video-mediastinoscopy, aggressive therapy should be encouraged. On the other hand, if there is regional nodal involvement (pN1/N2), then the benefit of aggressive surgery should be carefully reconsidered (49). All oligometastatic patients should always be discussed in a multidisciplinary meeting. Surgical options include lobectomy for the dominant lesion and sublobar resection for the lesser lesion. Alternatively, in some patients, sublobar resection may be done on one side and stereotactic ablation may be offered for the contralateral side.

A recent multicenter phase II randomized controlled trial from MD Anderson/London Ontario/Colorado compared local consolidative treatment, i.e., definitive radiotherapy or surgery to all sites of known disease, to maintenance treatment/observation. They showed that in patients with oligometastasis (3 or fewer metastases), following frontline systemic therapy and who showed no disease progression at 3 months or more, the local consolidative therapy group is superior for both progression free survival (median 14.2 vs. 4.4 months, P=0.022), and overall survival (median 41.2 vs. 17.0 months, P=0.017), with no differences in grade 3 or greater toxicities between the two groups. This exciting work is practicing changing and will for sure generate significant impetus for more aggressive treatment of appropriately selected patients with oligometastases (50).

#### Expert opinion 2: Dr. Carmelina C. Zirafa

Despite the progress in the treatment of lung cancer, unfortunately the prognosis of clinical IV stage NSCLC is still negative. Hence the surgical treatment should be considered only for selected cases of advanced stage of lung cancer, considering also the promising outcomes obtained with new chemotherapy strategies. In selected IV stage cases, the survival in patients treated with resection of the primary lesion has been proven to be better than in non-surgical subjects, rising a 5-year-survival of about 20–40% (51). According to NCCN (National Comprehensive Cancer Network) guidelines, the satisfying prognosis following the radical surgery in selected cases encourages the surgical option. In the analysis of prognostic factors after surgery in patients with advanced disease, the nodal involvement appears to be an independent predictor of a worse prognosis, being N2 positivity associated with a reduction in survival when compared to N0 and N1 (52). In IV stage, other negative prognostic factors after surgery are: higher T status, presence of malignant pleural effusion, more of 2 metastases and surgical resection different from lobectomy (wedge resection, segmentectomy, pneumonectomy) (53).

Accurate staging remains fundamental to select limitedsite oligometastatic N0-1 patients with good performance status for surgical treatment, to reach the best therapeutic option and to improve oncologic outcomes.

#### Expert opinion 3: Dr. Nuria M. Novoa

Current data supports the utility of surgery in stage IV whenever a proper selection of the patients is done (54). Probably a better knowledge of the biological stress test is needed in order to better define the population that will really benefit from undergoing surgery. This is an interesting study line. As I mentioned before, the introduction of the immunotherapy and the use of target therapies probably will increase the number of patients with oligo-recurrences or oligo-progression and this is a population that clearly benefit from the resection strategy. Still a lot of new achievements should come, and we need to be ready to adapt to the new circumstances. I am clearly optimist about the future of stage IV patients.

#### Expert opinion 4: Dr. Stefano Bongiolatti

Published data regarding the surgical/local treatment of metastatic lung cancer are controversies because, in my opinion, there is a lack in the definition of the "oligometastatic state" as affirmed by Hellman and Weichselbaum in 1995. Several progresses have been made in the diagnostic, staging and treatment fields, but the scientific community needs a universally accepted definition of oligometastatic state. Patients with oligometastatic NSCLC demonstrated favourable overall survival when treated with an intention to cure multidisciplinary plan, but some clinical conditions are mandatory although not universally stated. Several questions can arise from this situation. First of all, the number of metastases which can be locally treated. Second, the site and the treatment modality (surgery or radiation therapy) are not welldefined. Third, the treatment should be the same for patient with synchronous or metachronous metastases? Despite all these questions, the most important criterion that every clinician should follow in the decision-making

process for NSCLC in stage IV, is an accurate and invasive staging and consequently absence of lymph nodal hilar or mediastinal metastasis (N0 disease). Some robust data are necessary to well define this controversial topic and perhaps the definition of metastatic or oligometastatic disease may change thanks to the future developments and clinical applications of new biomarkers such as circulating tumour cells (CTCs) and circulating cell-free DNA or RNA.

#### Expert opinion 5: Dr. Robert J. Cerfolio

Interestingly as surgeons, although we would have a shrinking or decreasing role in the treatment of T1a and T1b tumors, we would concomitantly play a more important role in the treatment of patients with stage III and stage IV cancers.

We have done for years with patients of adenocarcinoma who have solitary metastatic lesions of the brain or adrenal, plural-based disease with multiple lesions, or even those with hepatic disease or nodules in other parts of the lung.

Nowadays even patients with multiple plural based modules would receive resection of the primary tumor two or three years later when they were diagnosed with stage IV cancer. A rogue metastasis is resistant to the targeted therapy.

# How should we think of citation ranking of articles in the field of lung cancer surgery?

#### Expert opinion 1: Dr. Carmelina C. Zirafa

In the last decades, treatment and prognosis of lung cancer have been in continuous evolution and several papers have been published to report these changes. The analysis of the top 100 cited articles on lung cancer has underlined the heterogeneity of the evaluated aspects in literature. The principal elements studied by the Authors was the impact of neoadjuvant and adjuvant chemotherapy and radiotherapy in patients undergoing surgery for lung cancer, the extension of surgical resection, the post-operative oncologic outcomes, the application and the results of minimally invasive surgery and the investigation of prognostic factors for resected NSCLC (e.g., clinical characteristics, gene mutations). Usually, the higher citation ranking corresponds to a relevant impact of the paper on the clinical and scientific field, yet the reverse might not always be true. In fact, some outstanding article not have a high citation count, due to the fact that the number of citations can be influenced not only by the quality of the research but also by other factors, such as the author's relevance in the

scientific community or to self-citations.

#### Expert opinion 2: Stefano Bongiolatti

In my opinion, it could be difficult to change some universally accepted ranking methods as the use h-index. For example, this score should be improved removing selfcitation or inserting a function that analyse the type of study performed and the author position in the authors' list. In this way, more information and more precise details will be included into the score.

#### Expert opinion 3: Dr. Nuria M. Novoa

The only way to know the relevance of any work is looking at its repercussion in the rest of the scientific community. How to measure it? Using a ranking system. The current metrics is based on the number of citations of every published paper. Based on it you can calculate the author's Impact Factor and the h-index among others or you can measure the citations of each individual paper. No index is perfect. For instance, the IF is a good system although refinements are necessary. For example, not all the papers published in one journal received the same number of citations, but all received the same IF based on the IF of the journal. Moreover, it does not distinguish between the type of manuscript. Therefore, manuscript from originals to letters to the editor or review papers have the same IF. One interesting problem is self-citation. Some journals and some disciplines favor self-citation meanwhile others penalized it. In my opinion, unless you are working in a very specific area, self-citation is not a good option. Nowadays knowledge grows thanks to little steps made by different members of the community and all them deserve the benefit of their contribution. I am sure that refinements will take place in the current metric system although I consider it a good one. The relevance of this paper relays in the fact that reviews the greatest contributions in the knowledge of lung cancer, the major topic of our specialty.

#### Expert opinion 4: Dr. Robert J. Cerfolio

I'm not quite sure how to give a clear objective answer to this question. My first question is why is it important? Why do we have to rank the articles of lung cancer from the most important to the least important? Why should we strive to do this thing?

It is like ranking which movie, which book or which painting is the best. Is there any inherent subjectivity and why does it matter?

Currently the only metric is the impact factor of the

journal in which the article published. But the number of times that an article is referenced is heavily biased. I think if we ask ourselves what is the most important article in the field of lung cancer surgery, it should be the one which has the largest influence or one that improves the therapy effect for most patients, which means that it has the greatest value.

#### Conclusions

This study provides an important and comprehensive overview of the most cited articles in the field of the thoracic surgery management of lung cancer over the past 40 years. It recognizes the important contributions made by key authors and institutions to the field and provides an important framework to a better understanding of the evidence base behind the multidisciplinary treatment of lung cancer.

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#### Footnote

*Conflicts of Interest:* The authors have no conflicts of interest to declare.

*Ethical Statement:* The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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