



Early View

Original research article

One-year mortality in lung cancer in France according to key time points of care pathways

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Title

One-year mortality in lung cancer in France according to key time points of care pathways

Running title

Treatment initiation and survival in lung cancer

Authors

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“Take-home” message:

Lung cancer care pathways were described using medico-administrative and clinical databases. In unresectable cases, rapid care access was not associated with better survival. Additional time for molecular biology analysis did not impact treatment initiation.

Abstract

Background. It is unclear whether delays in care affect prognosis of patients with lung cancer. The primary objective of this study was to describe the care pathway of patients diagnosed with lung cancer in a French region. Secondary objectives were to identify markers associated with 1) time from imaging to treatment, and 2) with one-year survival.

Methods. In a retrospective cohort study, clinical data from multidisciplinary team meetings (MTM) for all incident lung cancer cases discussed in 2018 in one French region were matched with medico-administrative data from the National Health Insurance Database. Care pathway time intervals were estimated for small cell lung cancer (SCLC), resected non-small cell lung cancer (NSCLC) and unresected NSCLC. Factors associated with delay in the care pathway were identified using linear regression; 1-year survival was analysed using Cox modelling.

Results. A total of 685 patients were included. Median time between imaging and treatment was 49 days (q1-q3: 33-73), and was lower in cases of metastatic disease, SCLC, and private care. At one year, 48% had died (resected NSCLC 12%). In unresected NSCLC, time from diagnostic imaging to first treatment <49 days was associated with a higher risk of death. Time intervals were similar in patients with squamous cell carcinoma vs adenocarcinoma or undifferentiated carcinoma.

Discussion. Time intervals in the care pathways of lung cancer were similar to previous reports, confirming the robustness of retrospective databases. In unresectable NSCLC, rapid care was not associated with better survival.

Key words: care pathway, databases, epidemiology, lung cancer, one-year case fatality

Introduction

Lung cancer is the leading cause of cancer deaths in Europe. In France, lung cancer is the second most frequent cancer in men and the third most frequent cancer in women with 31,231 and 15,132 new cases in 2018, respectively (1). Lung cancer includes non-small cell lung cancer (NSCLC) and small cell lung cancer (SCLC), which differ in their natural history, prognosis and management (2). A fraction of patients with limited NSCLC benefits from surgical resection. In recent years, prognosis of unresectable NSCLC has been improved by the introduction of targeted therapies and immune checkpoint inhibitors in addition to chemotherapy and radiotherapy (3). According to current guidelines on NSCLC care (4), patients with unresectable adenocarcinoma or undifferentiated carcinoma require molecular biological analysis for optimal treatment, as do the rare non-smokers with squamous cell carcinoma. Prognosis of SCLC, mainly treated by chemotherapy and radiotherapy, is poor (5)(6).

Rapid diagnosis and treatment are considered indicators of quality of care in lung cancer, although evidence is lacking to support this concept in patients with unresectable lung cancer (7). While large population-based studies have reported an association between swift surgery and longer survival in patients with resectable NSCLC (8,9), it is unclear whether care pathway delays impact on outcomes in patients with unresectable lung cancer (10). In fact, currently available data indicate either inverse associations between care delays and survival (i.e. longer survival in patients with longer care delays) (11–13) or the absence of any link (14–19). Only one study reported an association between short time to management and increased survival in unresectable NSCLC (8). These data are in opposition with current guidelines of lung cancer care, which set clear objectives in terms of care time intervals for unresectable lung cancer (20–23). Specific to unresectable NSCLC, concern may be raised that the recent need for lengthy molecular analyses, which are required for access to targeted therapies, may lead to increased time intervals between diagnosis and treatment (24).

To better inform public policy into lung cancer care, we aimed to further explore the hypothesis that delays in the care pathway are associated with poor prognosis of lung cancer. The primary goal of the study was to measure time intervals between the major steps of medical care of patients newly diagnosed with lung cancer in a French region. The secondary objectives were to assess factors associated with i) time intervals from imaging to treatment, and ii) with one-year survival. We then explored whether patients with adenocarcinoma or undifferentiated carcinoma, in whom molecular

biology analyses are required, had longer time to treatment in comparison with patients with squamous cell carcinoma.

Methods

Study design and data sources

A multicentre retrospective observational study was conducted in the Centre-Val de Loire region of France by merging two existing databases, the Shared Medical reports in Oncology (*Dossier Communicant en Cancérologie - DCC*) and the French Health Insurance Database (*Système National des Données de Santé – SNDS*). The Centre-Val de Loire region includes six administrative departments that were anonymised (A to F). Patients were identified from DCC data. DCC is a French regional register which gathers data on diagnosis and treatment of all cancer patients. These data were collected during multidisciplinary team meetings (MTMs) in 35 Centre-Val de Loire centres. By law, MTMs are mandatory for all cancer patients in France. Inclusion criteria were i) age over 18 years old and ii) first thoracic oncology MTM in 2018. Diagnoses of lung cancer were checked by reviewing all medical records.

As DCC does not include follow-up data, matching with SNDS was performed to describe the entire care pathway for each patient. SNDS includes all care reimbursement data such as type and date of any diagnostic procedure or treatment, and date of death. Since the two databases have no common patient identification number, matching of DCC and SNDS databases was probabilistic, based on sex, month and year of birth, date of diagnosis and residence postal code. Patients were excluded if i) they had lung cancer in the previous five years, ii) they had pre-existing extra-pulmonary cancer or synchronous extra-pulmonary cancer, iii) review of medical records ruled out lung cancer, or iv) variables required for matching were missing.

Definition of care pathway key points, time intervals and secondary outcomes

The dates of diagnostic imaging, pathological diagnosis, MTM and first treatment were used to calculate pathway-specific time intervals. Date of diagnostic imaging (computed tomography-CT or positron emission tomosintigraphy-PET or chest X-ray), date of cancer pathology diagnosis, date of

treatment (surgical resection, targeted therapy, immunotherapy, chemotherapy, radiotherapy or palliative care), and date of death were obtained from SNDS. Medical procedures were identified using the French medical procedures thesaurus (*Classification Commune des Actes Médicaux- French Classification of procedure terminology CPT*). Date of MTM and lung cancer type (NSCLC including adenocarcinoma, undifferentiated carcinoma and squamous cell carcinoma, or SCLC) were obtained from DCC. The date of pathological diagnosis was the date of the initial pathology report which included the immunohistochemical study (ALK, ROS1, PD-L1), or the sampling date if the pathology report was not available. The date of the final report including molecular biology analyses was not available. Drugs were identified according to the Anatomical Therapeutic Chemical Classification system. Driving times to the first healthcare facility were estimated with the METRIC software from the French Institute of Statistics and Economical Studies (*Institut National de la Statistique et des Etudes Economiques INSEE*).

Patients were allocated to 3 care pathway groups according to tumour type and treatment : “unresected NSCLC”, “SCLC” and “surgically resected NSCLC”. In patients with SCLC and unresectable NSCLC, time intervals from imaging to cancer pathological diagnosis, from pathological diagnosis to MTM, from MTM to treatment initiation and from imaging to treatment were calculated. When the time intervals were negative numbers, these outcomes were excluded from the analyses (for instance when the first treatment occurred before MTM).

In patients with resected NSCLC, only the time intervals from diagnostic imaging to pathological diagnosis and from diagnostic imaging to surgical resection were calculated, MTM was often conducted after surgery, resulting in a negative MTM to treatment interval. The following secondary outcomes were calculated: i) time from diagnostic imaging to first treatment, and ii) death at one year, defined as the frequency of patients deceased one year after diagnostic imaging.

Statistical analyses

Qualitative data were presented as numbers and percentages. Continuous data were presented as medians and interquartile ranges (IQR). Factors associated with time from diagnostic imaging to first treatment were identified by a multiple linear regression model. The type of tumour was used as an adjustment variable. A sensitivity analysis was performed on the subgroup of NSCLC patients. For each care pathway, factors associated with one-year survival were identified by a Cox regression

model giving hazard ratios (HR) along with their 95% confidence intervals (95% CIs). Variables with $p<0.2$ in bivariate analysis were included in the initial multivariate models. A descending stepwise process was then used to select the final models, including all the statistically significant variables at the threshold $p<0.05$. Analyses were performed with SAS Enterprise Guide 7.1 64-bit (SAS Institute Inc., Cary, NC, USA).

The study was authorised by the French data protection board (*Commission Nationale de l'Informatique et des Libertés - CNIL*), allowing access to SNDS data, decision DR-2019-371. Moreover, the study was granted a waiver of the individual information requirement by the French data protection board, as most patients were deceased at the time of the retrospective study and their personal details were not known. As an alternative measure, collective information about the study was made available and required by the protection board on the websites of the regional epidemiology unit and of the regional oncology centre, mentioning the rights of access, rectification and opposition, in accordance with the provisions of article R. 1461-9 of the CSP, as well as the following rights and information, in accordance with the European Data Protection Regulation.

Results

Study population

Screening of the DCC database yielded 962 adult patients. Of these, 102 were excluded due to a previous history of lung cancer ($n=55$), no lung cancer or synchronous extra-pulmonary cancer or pre-existing extra-pulmonary tumours ($n=33$), or missing data ($n=14$). Matching failure between DCC and SNDS data led to 175 additional exclusions (**Figure 1**). Thus, the final sample included 685 patients, corresponding to a 79.7% matching rate.

Median age was 66 years (59-73) (**Table 1**). Most patients were males (72%). Performance status (PS) was ≤ 1 in 82% of patients. One hundred and thirteen patients (16%) had SCLC, 106 (15.5%) had resected NSCLC and 466 (68%) had unresected NSCLC. Among NSCLC cases, the most frequent cancer pathology was adenocarcinoma. The stage of the disease was available for 531 patients (77%); 403 patients had metastatic disease.

Of the 106 patients undergoing resection, surgery was the first treatment in 99 patients. Among patients with unresected NSCLC, the first treatment was chemotherapy in 329 (48%) patients, immunotherapy in 60 patients (9%), targeted therapy in 27 patients (4%), and palliative care or abstention in 136 patients (23%).

A total of 243 patients (35%) were cared for in teaching Hospitals, 195 (28%) in secondary public hospitals, and 220 patients (32%) in private healthcare facilities. One hundred and eighty-eight (28%) patients lived over a 45-minute drive from the healthcare facility.

One year after diagnostic imaging, 331 (48%) patients died, ranging from 12.3% for resected NSCLC to 59.3% for SCLC.

Care pathway key points and time intervals

The diagnostic imaging procedure was CT for 571 patients (83.3%), chest X-ray for 64 patients (9.4%) and PET for 18 patients (2.6%) whereas no imaging was reported in 32 cases (4.7%). The date of pathological diagnosis was the one of the pathology report in 515 patients (75.2%) and the date of sampling in 154 patients (22.5%); no date was found in 16 cases (2.3%). Eventually, 18% of the time intervals were excluded from the analyses.

In the whole study population, time from diagnostic imaging to pathological diagnosis, from pathological diagnosis to MTM and from MTM to first treatment were 12 days (6-30), 14 days (8-22) and 16 days (9-28) respectively. Time from diagnostic imaging to first treatment was 49 days (33-76) (**Figure 2a**). It was shorter for patients with SCLC (**Figure 2b**). Among patients with unresected NSCLC, this time interval was no different in patients with adenocarcinoma or undifferentiated carcinoma vs squamous cell carcinoma (**Figure 2c**). In patients with resected NSCLC, 41% of patients had surgery before MTM. Time from diagnostic imaging to surgery was 59 days (41-91) (**Figure 2d**).

Factors associated with time from diagnostic imaging to first treatment

Multivariate analysis showed that time from diagnostic imaging to first treatment increased in patients with NSCLC (+10.4 days; p<0.001) compared to patients with SCLC (**Table 2**) along with care given in the administrative department C (+14 days; p=0.01) or F (+10 days; p<0.001) compared to D (**Table 2**). Conversely, time from diagnostic imaging to first treatment decreased in patients with metastatic cancer (-16 days, p<0.0001) or in a private healthcare facility (-7 days, p=0.01). No

association was found between time from diagnostic imaging to first treatment, along with driving time to first healthcare facility and either sex or age.

Focusing on patients with unresectable NSCLC, no significant difference was found between cancer requiring molecular biology tests as standard (adenocarcinoma or undifferentiated carcinoma) or not (non-squamous cell carcinoma) (**Table 2**).

Factors associated with death at one year

In patients with unresected NSCLC, factors associated with increased risk of death (**Table 3**) were PS>1, squamous cell carcinoma or undifferentiated carcinoma (vs. adenocarcinoma), metastatic disease, care in the administrative department B or in a public hospital, absence of targeted therapies, radiotherapy or immunotherapy, and time from imaging to treatment under 49 days.

Of the 96 patients with SCLC, 59% died within the year following diagnostic imaging. No association was found between death and the time interval from imaging to first treatment, driving time to the care facility, and type of healthcare facility (Supplementary material – Table S1). No variable was significantly associated with risk of death in patients with resected NSCLC (Supplementary material – Table S2).

Discussion

This study is the first to provide a comprehensive description of care pathways of patients diagnosed with lung cancer in France. Key time intervals of the care pathway were similar to those reported in other European regions, confirming the robustness of retrospective databases, reinforced by a new combined approach. Indeed, completeness and analysis strength were enhanced by matching an exhaustive medico-administrative database with MTM medical records. The exhaustiveness of clinical databases is the key factor for optimizing studies with such a design.

An accurate description of lung cancer care pathways in real-life conditions of patients with new-onset lung cancer was performed in a large French region with more than 2.5 million inhabitants. We could observe that access to treatment initiation reversely impacted 1-year survival in patients with unresected NSCLC, strengthening the notion that rapid care may not translate into better outcomes in patients with unresectable lung cancer. Furthermore, this study suggests that molecular biology analyses such as next-generation sequencing for unresectable NSCLC did not result in extending time

to treatment initiation. It may be speculated that the lack of a difference in the time from imaging to treatment may be related to the fact that, when targeted treatment is prescribed, treatment delivery is quite rapid compared to chemotherapy or immunotherapy.

Although the Centre-Val de Loire region combines multiple factors associated with increased time intervals in care such as low population density, low personal income and educational level, and scarce health resources (25–27), care time intervals were similar to the literature. Hence, although there was a possible selection bias due to possible mismatches between the two databases, the convergence of results with other European studies confirmed the robustness of the design. The median time from diagnostic imaging to treatment initiation was 49 days. The time interval between symptom onset and diagnostic imaging would have been interesting to study, however available databases did not provide information on symptoms. In recent studies, median time from symptom onset to treatment was 130 days in Finland (18) and 123 days in Spain (13). It may be hypothesised that in Centre-Val de Loire, time from symptom onset to diagnosis may be longer due to the low density of general practitioners (25).

Importantly, care intervals in the present study mostly complied with current RAND corporation (22), Ontario Cancer Plan (23), British NHS (21) and BTS (20) guidelines for lung cancer care. At the European level, median time from pathological diagnosis to treatment was 28 days in the Netherlands (28) and 35 days in England (29). In the present study, time from pathological diagnosis to MTM and from MTM to treatment were 14 days and 16 days, respectively. Thus, time from pathological diagnosis to first treatment of lung cancer may actually be rather similar among different Western European countries. Altogether, these data suggest that key time intervals in the lung cancer care pathway are relatively homogeneous across Western European countries, supporting the fact that the present study's results may be useful to other countries with similar healthcare systems.

Shorter time between diagnostic imaging and first treatment was associated with poorer survival in patients with unresected NSCLC, independent of PS and disease stage, as reported in previous studies (11–13), although one previous study reported an inverse association (8). The mechanisms driving the association between shorter care time intervals and worse prognosis are not fully understood. It is questionable whether aiming to reduce unresected NSCLC care time intervals beyond what is currently recommended by RAND corporation (22), Ontario Cancer Plan (23), British NHS (21) and BTS (20) guidelines is of interest in a public health and care quality perspective. The key time

intervals were shorter in patients with metastatic disease and SCLC, in accordance with earlier studies (30,31).

This study presents an evaluation bias, particularly with regard to the definition of the date of pathological diagnosis (absence of date of anatomopathological examination for 24.8% of patients), which led to underestimation of the time from diagnostic imaging to pathological diagnosis and overestimation of time from pathological diagnosis to treatment. Missing data are common in studies of this type. In a previous French national study, the date of pathology report was not available in 19.1% of cases and the time to diagnosis could not be calculated in 37.5% of cases (32).

It is possible that in patients with unresected NSCLC, missing data may have precluded precise estimation of disease severity. In particular, nutritional status, the location of metastases and full documentation of somatic mutations may have been of interest. It is possible that disease severity determines survival to a much larger extent than treatment-related aspects such as achieving reduction in care intervals. Another limitation is that 41% of patients with resectable lung cancer were operated on before MTM, which precluded analysis of the MTM to surgery delay in many patients.

In conclusion, these new results confirm the high interest of retrospective real-life databases and the power of approaches combining medico-administrative and clinical databases, in order to assess healthcare pathways in a cost- and time-saving process, giving trends to help enhance public health policies. Exhaustiveness of clinical databases is a key factor to optimize retrospective studies. It seems useful for each country to develop good databases in order to analyse the key points to improve in order to offer lung cancer patients the best treatment. Key time intervals in the lung cancer care pathway in this large French population-based study were similar to those reported in other European regions. A shorter time from diagnostic imaging to first treatment was associated with increased 1-year mortality in patients with unresectable NSCLC, strengthening the notion that accelerating care beyond what is already achieved may not translate into better outcomes for those patients. Moreover, this study suggests that the introduction of molecular biology analyses did not lead to an increase in management intervals.

Authors' contributions

AR, AIL, EL, CL, TL, EO, DC, LP, LGG conceived the study and drafted the study protocol. CL, EO, AR provided and reviewed clinical data. AIL analysed the data. AR, DC, LP helped to interpret the data. AR, AIL, EL, LGG drafted the manuscript. The final manuscript was read and approved by all authors.

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Declaration of Conflicting Interests

The authors declare that there is no conflict of interest.

Funding information

None

Ethical statement

The study was authorised by the French data protection board (*Commission Nationale de l'Informatique et des Libertés - CNIL*), decision DR-2019-371.

Availability of data and materials

If you require more details concerning data and materials, please contact us by sending an email to:

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Table 1. Patient characteristics – the CAP-Centre study.

	All patients n=685		SCLC n=113 N (%)	Resected NSCLC n=106 N (%)	Unresected NSCLC n=466 N (%)
	TOTAL n=685 N (%)	Death at 1 year n=331 N (%)			
General characteristics					
Male	496 (72.4%)	254 (51.2%)	88 (77.9%)	72 (67.9%)	336 (72.1%)
Age (median [q1-q3])	66.0 [59-73]		65.0 [60-74]	66.0 [60-72]	67.0 [58-74]
Performance Status (PS) score					
0 - 1	565 (82.5%)	233 (41.2%)	90 (79.6%)	104 (98.1%)	371 (79.6%)
2	85 (12.4%)	65 (76.5%)	18 (15.9%)	2 (1.9%)	65 (13.9%)
3 - 4	35 (5.1%)	33 (94.3%)	5 (4.5%)		30 (6.5%)
Histology, n=662					
Adenocarcinoma	311 (47.6%)	138 (44.4%)		54 (56.2%)	257 (57.9%)
Undifferentiated carcinoma	62 (9.5%)	38 (61.3%)		9 (9.4%)	53 (11.9%)
Squamous cell carcinoma	149 (22.8%)	70 (47.0%)		30 (31.2%)	119 (26.8%)
Others	50 (7.3%)	10 (55.5%)		13 (12.2%)	37 (7.9%)
SCLC	113 (17.3%)	67 (59.3%)	113 (100%)		
Disease stage, n=531					
I	29 (5.4%)	2 (6.9%)	1 (1.2%)	26 (38.8%)	2 (0.5%)
II	27 (5.1%)	3 (11.1%)	2 (2.4%)	21 (31.3%)	4 (1.0%)
III	72 (13.5%)	23 (31.9%)	5 (6%)	13 (19.5%)	54 (14.2%)
IV	403 (75.9%)	253 (62.8%)	75 (90.4%)	7 (10.4%)	321 (84.3%)
First healthcare facility					
Administrative department					
A	49 (7.2%)	27 (55.1%)	5 (4.4%)	6 (5.7%)	38 (8.2%)
B	87 (12.7%)	58 (66.7%)	23 (20.4%)	9 (8.5%)	55 (11.8%)
C	38 (5.5%)	21 (55.3%)	9 (8%)	5 (4.7%)	24 (5.2%)
D	265 (38.7%)	111 (41.9%)	36 (31.9%)	45 (42.5%)	184 (39.5%)
E	40 (5.8%)	20 (50.0%)	6 (5.3%)	7 (6.6%)	27 (5.8%)
F	174 (25.4%)	81 (46.5%)	28 (24.8%)	25 (23.6%)	121 (26%)
Outside of CVL region	32 (4.7%)	13 (40.6%)	6 (5.3%)	9 (8.5%)	17 (3.6%)
Nature					
Teaching hospital	203 (29.6%)	96 (47.3%)	29 (25.7%)	31 (29.2%)	143 (30.7%)
Public hospital	223 (32.6%)	135 (60.5%)	49 (43.4%)	24 (22.6%)	150 (32.2%)
Private Hospital	259 (37.8%)	100 (38.6%)	35 (31.0%)	51 (48.1%)	173 (37.1%)
Driving time (minutes)					
[00-15[187 (27.4%)	101 (54.0%)	36 (31.9%)	21 (19.8%)	130 (28.0%)
[15-25[153 (22.5%)	74 (48.4%)	29 (25.7%)	23 (21.7%)	101 (21.8%)
[25-45[153 (22.5%)	71 (46.4%)	22 (19.5%)	24 (22.6%)	107 (23.1%)
≥45	188 (27.6%)	84 (44.7%)	25 (22.1%)	37 (34.9%)	126 (27.1%)
First treatment					
Therapeutic abstention	136 (19.9%)	123 (90.4%)	23 (20.3%)		113 (24.2%)

Resection	99 (14.5%)	12 (12.1%)	3 (2.7%)	96 (90.6%)	
Targeted therapy	27 (3.9%)	10 (37.0%)			27 (5.8%)
Immunotherapy	60 (8.8%)	26 (43.3%)		1 (0.9%)	59 (12.7%)
Chemotherapy	329 (48.0%)	148 (45.0%)	83 (73.5%)	9 (8.5%)	237 (50.9%)
Radiotherapy	30 (4.4%)	11 (36.7%)	3 (2.7%)		27 (5.8%)
Radio-Chemotherapy	4 (0.6%)	1 (25%)	1 (0.9%)		3 (0.6%)
Case-fatality					
Death at one year	331 (48.3%)		67 (59.3%)	13 (12.3%)	251 (53.9%)

Table 2. Factors associated with time from diagnostic imaging to first treatment – the CAP-Centre study.

Variation of time interval between diagnostic imaging to first treatment - Multivariate analysis	All patients (n=520)		patients with unresected NSCLC* (n=305)	
	days (n)	95% confidence interval	days (n)	95% confidence interval
Origin time				
Male	-0.8	[-5.6; 4.1]	1.1	[-5.12; 7.3]
Age \geq 65 years old	0.9	[-3.4; 5.3]	1.4	[-4.2; 7.1]
Metastatic disease	-15.8	[-20.4; -11.2]	-14	[-20.8; -7.3]
NSCLC (as compared to SCLC)	10.4	[4.2; 16.5]		
Adeno/undifferentiated carcinoma (vs squamous cell carcinoma)			2.5	[-4.1; 9.1]
Nature of the first healthcare facility				
Teaching hospital	ref			
Public hospital	-0.7	[-7.8; 6.4]		
Private Hospital	-7.1	[-12.5; -1.7]		
Administrative department of the first healthcare facility				
A	8.7	[-1.2; 18.6]	9.6	[-2.1; 21.3]
B	2.7	[-5.7; 11.1]	-3.6	[-14.1; 7]
C	14.5	[3.4; 25.7]	20.3	[4.4; 36.3]
D	ref		ref	
E	7.6	[-1.8; 17.0]	4.2	[-8.4; 16.8]
F	10.0	[4.5; 15.5]	17.5	[10.6; 24.4]
Outside CVL region	-5.1	[-15.0; 4.8]	0.4	[-15.8; 16.6]
Nature of the first treatment facility**				
Teaching hospital			ref	
Public hospital			10.2	[2.2; 18.2]
Private Hospital			4.3	[-2.9; 11.5]

* excluding patients with histology="others", n=37

**can be different from first healthcare facility

Table 3. Factors associated with death at one year in patients with unresected NSCLC – the CAP-Centre study.

Potential risk factors of death at 1 year - patients with unresected NSCLC (n=360)	Hazard Ratio (HR)	95% confidence interval (95%CI)
General characteristics		
Age ≥65 years old	1.2	[0.8-1.6]
Male	1.0	[0.7-1.5]
PS >1	2.1	[1.5-3.0]
Histology		
Adenocarcinoma	ref	
Squamous cell carcinoma	1.6	[1.1-2.4]
Undifferentiated carcinoma	2.0	[1.3-3.1]
Others	1.4	[0.6-2.9]
Metastatic stage	2.4	[1.6-3.8]
Healthcare pathway		
No immunotherapy	1.8	[1.3-2.7]
No targeted therapy	2.5	[1.3-4.9]
No radiotherapy	2.8	[1.7-4.7]
Administrative department of the first health care facility		
A	1.1	[0.5-2.3]
B	1.9	[1.1-3.3]
C	0.8	[0.4-2.0]
D	ref	
E	1.3	[0.6-2.5]
F	0.8	[0.6-1.3]
Outside of the CVL region	1.7	[0.7-3.8]
Nature of the first health care facility		
Teaching hospital	ref	
Public hospital	0.8	[0.5-1.3]
Private Hospital	0.5	[0.4-0.8]
Time interval between diagnostic imaging and first treatment <49 days	1.5	[1.1-2.1]

PS= performance status

CVL=Centre-Val de Loire

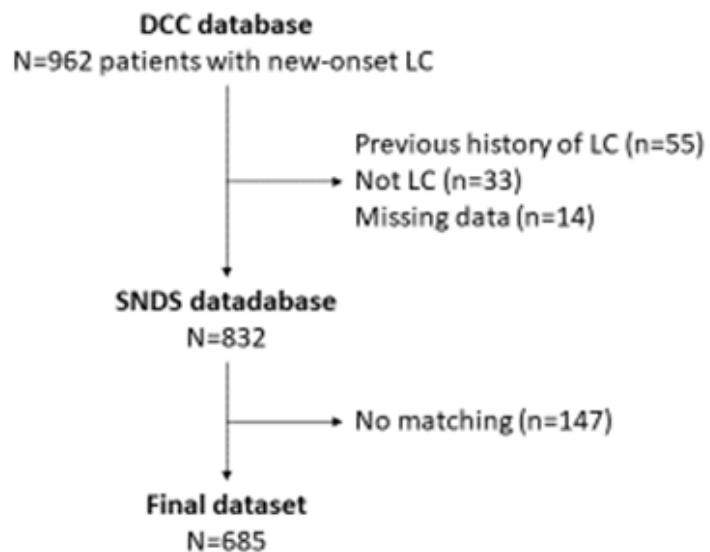
* 49 days corresponded to the median time interval between diagnostic imaging and first treatment

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Figure 1 flow chart – the CAP-Centre study

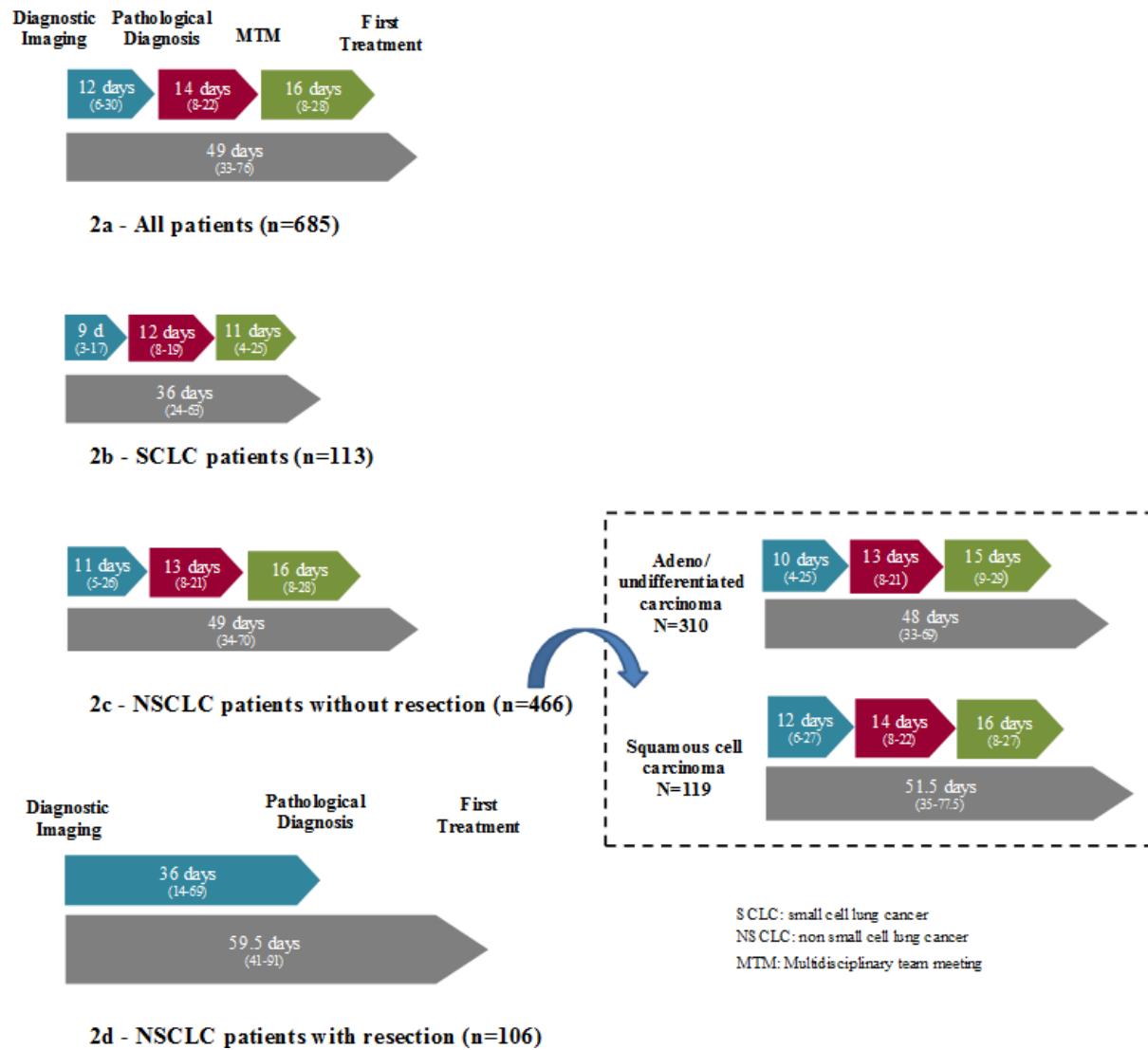


DCC: *Dossier communicant en cancérologie* (regional Shared Files in Cancerology) ;

SNDS: *Système national des données de santé* (French Health Insurance Database) ;

LC : lung cancer

Figure 2 Key time intervals in the care pathway – the CAP-Centre study.



Supplementary material – the CAP-Centre study

Table S1. Factors associated with death at one year in patients with resected NSCLC – the CAP-Centre study.

Potential risk factors of death at 1 year - patients with resected NSCLC (n=96)	Hazard Ratio (HR)	95% confidence interval (95%CI)
General characteristics		
Age ≥ 65 years old	1.6	[0.5-5.6]
Male	1.9	[0.4-9.3]
Metastatic stage	2.1	[0.4-10.8]
Healthcare pathway		
Time interval between imaging diagnosis and first treatment <60 days	1.5	[0.4-5.7]

Table S2. Factors associated with death at one year in patients with SCLC – the CAP-Centre study.

Potential risk factors of death at 1 year - patients with SCLC (n=96)	Hazard Ratio (HR)	95% confidence interval (95%CI)
General characteristics		
Age ≥ 65 years old	2.2	[1.2-3.9]
Male	0.7	[0.3-1.3]
PS >1	2.2	[1.3-3.9]
Metastatic stage	6.2	[2.2-17.4]