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Impact of systematic advanced care planning in lung cancer patients: a prospective study

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ABSTRACT

Background: End-of-life (EOL) communication is crucial, particularly for cancer patients. While advanced care planning is still uncommon, we sought to investigate its impact on care intensity in case of organ failure in lung cancer patients.

Methods: We prospectively included consecutive lung cancer patients hospitalised at the Grenoble University Hospital, France, between January 1, 2014 and March 31, 2016. Patients could be admitted several times and benefited from advanced care planning based on three care intensities: intensive care, maximal medical care, and exclusive palliative care. Patients' wishes were addressed.

Results: Data of 739 hospitalisations concerning 482 patients were studied. During the three first admissions, 173 (25%) patients developed organ failure, with intensive care proposed to 56 (32%), maximal medical care to 104 (60%), and exclusive palliative care to 13 (8%). Median time to organ failure was 9 days [IQR 25%-75%: 3-13]. All patients benefited from care intensity that was either equal to or lower than the care proposed. Specific wishes were recorded for 158 (91%) patients, with a discussion about EOL conditions held in 116 (73%).

Conclusions: In case of organ failure, advanced care planning helps provide reasonable care intensity. The role of the patient's wishes as to the proposed care must be further investigated.

Clinical Trial Registration: The study was registered at www.ClinicalTrials.gov with the identifier NCT02852629.

Keywords: advanced care planning, lung cancer, end-of-life conditions, intensity of care, palliative care.

1. Introduction

Lung cancer patients have the highest rates of intensive care unit (ICU) admission among tumour patients, straight after lower intestinal and genito-urinary cancers [1,2]. In recent decades, there has been a decrease in mortality for these ICU-admitted patients [1]. Despite the improvements in cancer treatments and resuscitation techniques, the mortality rate is still high in the event of organ failure in lung cancer patients. Recently, there has been increasing awareness of the inadequacy of end-of-life (EOL) care, partially because the patients' wishes were not known by their physicians, and due to a lack of communication [3–5], particularly before the patients lose their decision-making capacity. This may result in inappropriate care that the patients would not have chosen if they had been informed properly and involved in the decision-making [6]. Even if cancer patients likely think more about EOL care and advance directives [7,8], this decision-making process proves difficult. Previous studies have reported that patients think they do not have enough medical information about the disease and its treatments, but they likewise experience inner conflicts about quality/quantity of life [9]. Patients and physicians should be better prepared for these decisions, with tools available to assist in the process [10]. The quality of medical care delivered to cancer patients nearing the EOL is of significant concern. The issues that physicians face are how to evaluate the quality of care that patients should receive, as well as its adequacy with the patients' wishes [11]. Advance care planning is a distinct part of EOL care, representing a means to know patients' preferences, but it provides also an opportunity for a multidisciplinary discussion about care limitations in the event of acute disease, in both oncology and emergency units.

We sought to investigate the impact of systematic advance planning of EOL care in lung cancer patients admitted to a thoracic oncology unit.

2. Methods

2.1 Study design

We included all consecutive patients admitted to the thoracic oncology unit (conventional hospital stay) of the Grenoble University Hospital. France, between 28th January 2014 and 31st March 2016, and followed them up until 31st December 2016. Data were collected prospectively. Only patients with thoracic tumours like lung cancer and mesothelioma were studied, regardless of their ECOG-performance status (PS). Exclusion criteria were absence of cytological or histological cancer diagnosis, as well as remission for more than 5 years. At hospital admission, each patient benefited from a standardized advance care planning developed at the Grenoble University Hospital (annex 1). The programme was established based on the legal framework of the Kouchner Act of 4th March 2002, Leonetti Act of 22nd April 2005, and Leonetti-Claeys Act of 2nd February 2016 [12-14]. Depending on the patient's wishes and referent medical team's experience, an appropriate care intensity level was proposed. This was always carried out straight upon the patient's admission to the hospitalisation unit, designed to be just a proposition of care intensity, whereas the physician in charge of the patient at the time of organ failure remained master of the decision-making process. The propositions of care intensity recorded in this tool could be modified any time during the hospitalisation according to the patient's clinical progression and own wishes. Patients were evaluated at each admission, with their wishes as well as discussions with relatives recorded in the electronic medical chart. These discussions could be held before or after the advance care planning, which could be modified accordingly. The discussion was held with at least one physician, one resident and one nurse. The referent oncologist was contacted as soon as possible. The study protocol was published in 2015 [15].

The study primarily sought to evaluate whether the proposed care intensity level in the event of acute organ failure was in accordance with the standardized advance care planning. The secondary objectives were to better determine the conditions surrounding the discussions with patients and relatives, reasons of non-ICU admission in the case of proposed intensive care, as well as variations in the propositions during the first and subsequent hospitalisations. The association between patients' characteristics and care intensity propositions were likewise studied.

2.2 Ethics

Ethical Committee approval was obtained on 5th November 2013 (CECIC Rhône-Alpes-Auvergne, Clermont-Ferrand, IRB 5891). An information letter was handed to the patients, enabling them to refuse study participation. The study was registered at www.ClinicalTrials.gov with the identifier NCT02852629.

2.3 Data collection

The following patients' data were recorded: ECOG-PS, weight loss, Charlson Comorbidity Index (CCI) [16], and cancer history (status, histological type, and previous and current anticancer treatments). Reasons of admission were likewise collected, as were causes of organ failure. The advance care planning performed at admission and upon subsequent hospitalisations was recorded, as was the manner how planning evolved (more or less intensive), along with the respective reasons (annex 1). Care intensity propositions were divided into three categories; 1) intensive care (cardio-pulmonary resuscitation, transfer to the ICU, oro-tracheal intubation or tracheotomy, dialysis, and inotrope administration); 2) maximal medical care (non-invasive ventilation, artificial nutrition, antibiotic treatment, and blood transfusion); 3) exclusive palliative care (peripheral venous catheter, complementary examinations, vital signs, anticoagulation, fibro-aspiration). Reasons for discordance between care intensity proposition and care actually performed in the event of organ failure were likewise recorded, as were discussions with the patients or relatives.

2.4 Statistical analyses

Continuous variables have been expressed as median and interquartile range [IQR 25%-75%] and categorical variables as numbers and percentages (%). Associations between categorical variables were compared using the Chi-squared or Fisher's exact tests, and those between continuous variables using the Wilcoxon test. Patients were followed up until the 31st December 2016, with patients lost to follow-up censored.

Organ failure occurring during the three first hospitalisations was studied. If patients developed more than one organ failure, only the first one was considered. Kaplan-Meier plots of survival curves from organ failure onset were compared between groups using the log-rank test.

All tests were two-sided, with P values <0.05 considered statistically significant. All statistical analyses were performed using SAS 9.4 (SAS Institute, Cary, NC, USA).

3. Results

3.1 Patient characteristics

In total, 739 hospitalisations involving 482 patients were studied (figure 1). Nine patients were excluded and 473 analysed. Overall, 314 patients were not admitted for a second hospitalisation, 93 of whom died during their 1st hospitalisation and 164 thereafter, with 57 still alive. Thus, 159 patients were hospitalised a second time, and 69 a third time.

Patient characteristics at 1^{st} hospitalisation and according to the first planned care intensity level have been provided in table 1. A large majority of patients were metastatic (393, 85% at 1^{st} admission. At 1^{st} admission, 182 (38%) patients were not yet treated and 165 (35%) in cancer progression, whereas at 3^{rd} admission, 52 (75%) were in progressive disease. Reasons of admission at 1^{st} hospitalisation were recorded as acute disease (n=208, 48%), improvement in supportive care (n=167, 35%), scheduled hospitalisation (n=50, 11%), and toxicity of anticancer treatment (n=48, 10%). Among acute diseases, the main cause concerned respiratory disorders (n=119, 57%), followed by neurological disorders (n=41, 20%).

3.2 Planned care intensity

Concerning planned care intensity, at 1^{st} admission, intensive care was proposed for the majority of the patients in the event of acute organ failure (n=331, 70%), whereas it concerned only 77 (48%) at 2^{nd} hospitalisation, and 37 (54%) at 3^{rd} one (table 2). At 1^{st} admission, 62 (13%) patients benefited from a second care proposition. In most cases, the trend was towards more palliative care. Among the 39 patients with at least two propositions at 1^{st} admission and initial intensive care proposition, the 2^{nd} proposition was maximal medical care in 17 (44%) and exclusive palliative care in 2 (5%). Among the 23 patients with initial maximal medical care proposition, the 2^{nd} proposition was exclusive palliative care in 8 (35%) and intensive care in 2 (9%).

Patient characteristics varied according to the planned care intensity (table 1). The ECOG-PS was significantly poorer in the medical or palliative care groups (109, 76% patients with an ECOG-PS of 3 to 4) versus the intensive care group (117, 35% patients with an ECOG-PS of 3 to 4) ($P < 10^{-4}$). In univariate analysis, patient characteristics (age, modified CCI, and weight loss), cancer characteristics (time since cancer diagnosis, metastatic disease, cancer status, and current anticancer treatment) and reason of admission were likewise associated with the planed care intensity.

Median survival from the first hospitalization was 3.5 months [IQR 25%-75%, 1.0-9.7]. Among the 173 patients with organ failure, the in-hospital mortality was 41% (n=7) for patients who benefited from intensive care, 55% (n=27) for those with maximal medical care, and 92% (n=98) for those with exclusive palliative care. From the organ failure, median OS was 2 days [IQR 25%-75%, 0-13].

3.3 Patients with organ failures

At 1st hospitalisation, 127/473 (27%) patients developed organ failure, 34/159 (21%) at 2nd (3 not studied because 2nd organ failure), and 15/69 (22%) at 3rd hospitalisation. These 173 patients have been further described in table 3.

Main causes of organ failure were as follows: respiratory failure (n=68, 39%), coma (n=52, 30%), shock except for septic one (n=19, 11%), and sepsis (n=18, 10%). Median time from admission to organ failure was 9 days [IQR 25%-75%, 3-13].

The characteristics of patients who developed organ failure and care intensity administered are summarised in table 3. The main reason of admission was the improvement in supportive care (n=53, 50%). All patients who received exclusive palliative care suffered from metastatic

disease, their ECOG-PS proved to be worse, and they lost more weight. Most were in disease progression, without any further perspective of anticancer treatment.

3.4 Adequacy between planned and actual care intensity in the event of organ failure

Among the 173 patients who developed organ failure, care intensity was always equal or inferior to the care proposition just before organ failure (table 3). Among patients planned for intensive care (n=56, 32%), 39 (70%) did not benefit from it. In most cases (35/39), the oncologist did not solicit the opinion of the intensivist, reasons for this being a lack of updated care proposition in 18 patients, yet for 12 of them it was clearly stated in the medical records, ability to treat the organ failure in the medical unit in 13 patients, patients considered too frail to benefit from ICU admission in two, unreasonable intensive care proposition regarding patient's general conditions in one patient, with one patient found dead. In 4/39 cases, the intensivist's opinion was sought and the patient was thus not admitted to ICU because he was considered too frail to benefit from ICU admission in three cases, and because no ICU bed was available in the last case. Among patients without initial intensive care proposition, seven cases were discussed with the intensivist, with none admitted to ICU.

3.5 Condition surrounding the discussion with patients and relatives

For each hospitalisation, discussions with patients and relatives for those patients who developed organ failure were held for with most patients or relatives (n= 158, 91%), as summarized in table 4. In some cases (n=17/158, 11%), the discussion took place after organ failure, and the topics discussed were disease, disease severity, and EOL condition. Concerning the latter, discussions were held with patients and relatives for 116/173 (67%) during the hospitalisation, with the care proposition agreed upon in all cases. There was no

divergence between the medical team and patients or relatives point of view regarding the proposition of care at the time of organ failure.

4. Discussion

We have reported herein the data of a 2-year, prospective, hospital-wide study, designed to evaluate advance care planning developed at the Grenoble University Hospital. By means of this tool, no patient underwent more aggressive care than initially planned.

The study's strengths were its prospective design, along with the recording of both the physicians' and patients/relatives' point of view. Most studies published to date concerned only advance care planning according to the physician's decision [17,18], or depending on what the patients chose for their EOL conditions [3,19–21]. This is the first study reporting the use of a formal tool for advance care planning. A large number of patients were included, with an important rate of organ failures (36%). Patient characteristics were in line with real-life setting of hospitalised patients, and similar to those reported by other studies [22,23]. The single-centre patient recruitment, however, does not imply the applicability of our results to other centres. Of particular note is that this study was performed based on the thoracic oncologist's point of view, without systematically calling for the intensivist in the event of an intensive care proposition. Besides, the influence of patients' and relatives' wishes concerning the care intensity proposition was not collected either. These wishes were actually recorded in real-life conditions, but not following a thorough discussion.

In no case was the care intensity administered more aggressive than the one initially proposed. Previous studies [24–26] had already confirmed the crucial place of advance care planning and directives as regards EOL care, which resulted in less patients undergoing

intensive care. Concerning the people participating in the decision-making process, a lack of discussions with intensivists, owing due to an auto-censorship of the thoracic oncologist or physician in charge of the patient upon organ failure, has previously been highlighted [23]. This could be a particularity of our institution where thoracic oncologists are very committed to this topic. Recently, differences in view points between oncologists and intensivists regarding life support measures in critically-ill cancer patients have been reported in two different settings, namely metastatic breast cancer and metastatic pancreatic cancer [27]. It was reported that oncologists tend to focus on the cancer characteristics, but intensivists focus rather on multiple organ failure when confronted with the same decisions. Regular meetings between oncologists and intensivists during the disease course and prior to acute disease onset may reduce possible conflicts in the critical care of cancer patients. Physicians must keep in mind the objective of ICU admission: the patient should leave the ICU alive and still benefit from anticancer treatment, with sufficient quality of life, especially in metastatic cancer.

Regarding discussion with patients and relatives concerning EOL conditions, in 65% of cases, they were held during first hospitalisation. This number reached 87% for the third hospitalisation. Several studies have previously reported how these discussions likely influence the medical decision [4,8,21]. The earlier this discussion takes place, the more the patient's choices are respected. In the Mack et al. report [8], patients who had had EOL discussions with their physicians within the last 30 days of life were less likely to receive aggressive measures at EOL, including chemotherapy. In another study, 120 patients with metastatic cancer were randomized to receive either intervention on end-of-life decision making or treatment as usual [28]. The intervention made up of an informational pamphlet and discussions was associated with earlier exclusive palliative care implementation, along with less likelihood of in-hospital death. In our study, most of the discussions between thoracic oncologist and patients or relatives concerned the cancer and its prognosis,

particularly at EOL. They did, however, not imply a formal discussion to collect EOL wishes and were not meant to write advance directives, as framed by French laws on EOL care [12– 14]. Hospitalisation may occur too late in the patient's history to this end. Given this context, it appears warranted to increase the rate of patients who put down their advance directives at an earlier time in their cancer history. These patients' preferences should be accurately recorded in medical charts, thereby facilitating the decision around care intensity. This is an essential aspect of quality of care for patients with aggressive disease [11].

This advance care planning cannot be self-sufficient. A discussion between the physician in charge of the patient at the time of organ failure (or the intensivist) and the patient (and/or relatives) remains central to the decision-making about care intensity. In cancer cases, conditions of patients may change during the same hospitalisation and differ from others due to efficacy or inefficacy of anti-cancer treatment. In our study, we have noted several times that the deterioration of the patient condition was not properly acknowledged in the medical record. In case of re-hospitalisation, particularly at the emergency unit, we generally prefer the reflection to take into account each prognostic factor rather than a previous advance care planning that may even not be update. For optimal decision-making, emergency physicians, intensivists, oncologists, and the patient and his relatives must communicate in the best possible manner [29].

5. Conclusions

The care intensity propositions recorded in this tool for decision making in advance care planning were followed in terms of maximal care intensity. Such an aid in the decisionmaking appears to be useful for the physicians in charge of the patient, in the event of organ failure, likewise respecting the patients' wishes. It should, however, never replace a thorough discussion with the patient and his relatives, and we should better investigate the precise role of the patients' wishes in terms of the care intensity proposed. This tool must now be validated in a multicentre study involving patients with cancer or other chronic diseases.

Author contributions: A.-C. Toffart and J. –F. Timsit conceived the study. A.-C. Toffart, M. Duruisseaux and L. Sakhri included patients. A.-C. Toffart, N. Denis and J. –F. Timsit were involved in the analysis strategy and drafted the manuscript. All authors participated in the interpretation of the findings, reviewed the manuscript and revised it critically before submission. All authors have seen and approved the final version of the manuscript.

Declarations of interest: none.

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Figure caption

FIGURE 1 Flow chart of the study patients. Quantitative data were reported as median [interquartile range 25%-75%].

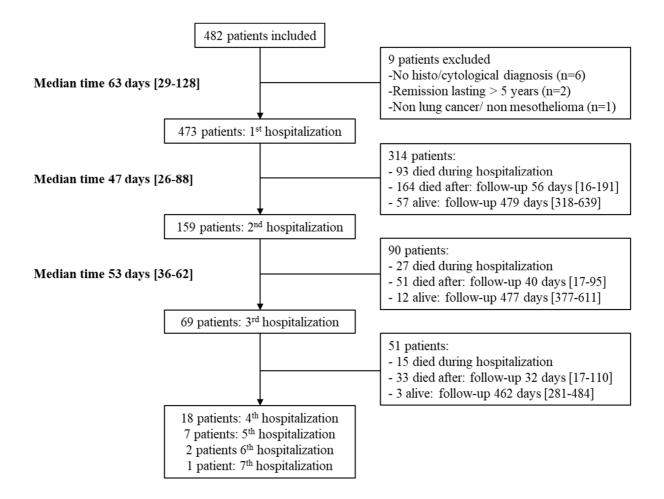


TABLE 1 Patient characteristics at first admission according to first planned care intensity

Variables	1 st admission n=473	Intensive care proposition n=330 (70)	Maximal medical or palliative care proposition n=143 (30)	P-value
Demographics				
Male gender	335 (71)	227 (69)	108 (76)	0.14
Age (years)	67 [60-75]	66 [59-73]	70 [61-77]	8.10-3
Modified CCI (points)	1 [0-3]	1 [0-2]	1 [0-3]	3.10-3
ECOG-PS at admission	- [* •]	- [* -]	- [• •]	<10-4
- 0-1	124 (26)	116 (35)	8 (6)	
- 2	123 (26)	97 (29)	23 (18)	
- 3-4	226 (48)	117 (35)	109 (76)	
Weight loss (%)	8 [1-15]	8 [0-13]	11 [3-19]	2.10^{-3}
	(MD=18)	(MD=11)	(MD=7)	2.10
Cancer history	(1112 10)			
Cancer diagnosis before	400 (85)	270 (82)	130 (91)	0.01
hospitalization	()	()		
Time since cancer diagnosis				
- If diagnosis before	189 [30-505],	146 [24-471],	274 [85-584],	8. 10-4
hospitalization (days)	n=400	n=270	n=130	2. 10
- If diagnosis during	4 [1-8], n=73	4 [1-8], n=60	4 [2-6], n=13	0.91
hospitalization (days)	L J, / J	. [- 0], 00	,,	
Cancer type				0.30
- Adenocarcinoma	264 (56)	195 (59)	69 (48)	
- Squamous cell carcinoma	72 (15)	46 (14)	26 (18)	
- Small cell carcinoma	76 (16)	50 (15)	26 (18)	
- Other NSCLC	49 (10)	31 (9)	18 (13)	
- Mesothelioma	12 (3)	8 (2)	4 (3)	
Metastasis (except mesothelioma)	393 / 461 (85)	261 / 322 (81)	132 / 139 (95)	10-4
Cancer status		(0)		<10-4
- Not yet treated	182 (38)	148 (45)	34 (24)	
- Progression	165 (35)	78 (24)	87 (61)	
- Controlled disease	75 (16)	53 (19)	12 (8)	
- Unknown	51 (11)	41 (12)	10 (7)	
Current anticancer treatment	()	()		<10 ⁻⁴
- Awaiting new treatment	187 (40)	156 (47)	31 (22)	
- Treatment ongoing	148 (31)	98 (30)	50 (35)	
- Included in clinical trial	30 (6)	24 (7)	6 (4)	
- Therapeutic window	53 (11)	40 (12)	13 (9)	
- Palliative care	55 (12)	12 (4)	43 (30)	
Reason of admission	(1-)	(')		6. 10 ⁻⁴
Acute disease	208 (44)	143 (43)	65 (45)	0.10
Respiratory disorder	119/208 (57)	89/143 (62)	30/65 (46)	
- Pleural effusion	33/119 (28)	25/89 (28)	8/30 (27)	
- Pneumonia	27/119 (23)	19/89 (21)	8/30 (27)	
- Tumour compression	25/119 (21)	19/89 (21)	6/30 (20)	
Neurological disorder	41/119 (20)	22/89 (15)	19/30 (29)	
-				
- Coma	7/41 (17)	3/22 (14)	4/19 (21)	
Improvement in supportive care	167 (35)	104 (32)	43 (44)	
Scheduled hospitalisation	50 (11)	46 (14)	4(3)	
Toxicity of anticancer treatment	48 (10)	37 (11)	11 (8)	

Data are expressed as median [interquartile range] for quantitative variables and numbers (%) for qualitative variables. CCI: Charlson Comorbidity Index [16]]; MD: missing data; NSCLC: non-small cell lung cancer, PS: performance status.

TABLE 2 Planned care intensity in the event of organ failure

Variables	1 st admission	2 nd admission	3 rd admission
	n=473 patients	n=159 patients	n=69 patients
1 st proposition	473 (100)	159 (100)	69 (100)
- Intensive care	330 (70)	77 (48)	37 (54)
- Maximal medical care	141 (30)	81 (51)	31 (45)
- Exclusive palliative care	2 (0)	1 (1)	1 (1)
First discussion at the day of	310 (65)	130 (82)	47 (68)
admission			
First discussion within the next days	163 (35)	29 (18)	22 (32)
of admission			
With a median time from admission	2 [1-4]	1 [1-3]	2 [1-3]
(days)			
2 nd proposition	62 (13)	16 (10)	7 (10)
Time between 2 nd and 1 st proposition	9 [4-14]	7 [5-12]	8 [6-9]
(days)			
- Intensive care	22 (35)	2 (13)	3 (43)
- Maximal medical care	30 (48)	11 (69)	2 (29)
- Exclusive palliative care	10 (16)	3 (19)	2 (29)
Sense of the 2 nd proposition			
compared to the first one			
- More palliative	52 (84)	13 (81)	6 (86)
- Equal	3 (5)	1 (6)	1 (14)
- More intensive	7 (11)	2 (13)	
3 rd proposition	16 (3)	2 (1)	•••
Time between 3^{rd} and 2^{nd}	6 [4-10]	6-17	
proposition (days)			
- Intensive care	2 (13)	1 (50)	
- Maximal medical care	8 (50)	1 (50)	
- Exclusive palliative care	6 (38)		

Data are expressed as numbers (%).

TABLE 3 Patients' characteristics according to intensity of care at organ failure

Variables	Organ failure n=173	Intensive care n=17 (10)	Maximal medical care n=49 (28)	Exclusive palliative care n=107 (62)	p-value
Demographics					
Male sex	128 (74)	13 (76)	36 (73)	79 (74)	0.97
Age (years)	67 [59-76]	62 [54-69]	64 [55-74]	69 [61-77]	0.02
Modified CCI (points)	0 [0-2]	1 [0-2]	1 [0-3]	0 [0-2]	0.21
ECOG-PS: 3-4 vs 0-2	134 (77)	11 (65)	35 (71)	88 (82)	0.13*
Weight loss (%)	11 [4-18]	8 [4-12]	9 [2-16]	13 [6-20]	0.02
	(MD=15)	(MD=1)	(MD=5)	(MD=9)	
Cancer history		× ,			
Time since cancer diagnosis (months) if	8.9 [2.6-18.5]	6.4 [1.5-16.4]	12.2 [3.3-25.2]	8.6 [2.7-15.6]	0.32
diagnosis before admission	n=153	n=14	n=42	n=97	
Type of cancer				, ,	0.60*
- Adenocarcinoma	96 (55)	10 (58)	27 (55)	59 (55)	
- Small cell carcinoma	27 (16)	2 (12)	6 (12)	19 (18)	
- Squamous cell carcinoma	24 (14)	4 (24)	5 (10)	15 (14)	
Metastasis (except mesothelioma)	160 / 168 (95)	13 / 17 (76)	44 / 48 (92)	103/103 (100)	<10-4*
Cancer status	100 (100 ()0)	10 / 17 (70)	117 10 (52)	105/105 (100)	0.02*
- Not yet treated	49 (28)	7 (41)	13 (27)	29 (27)	0.02
- Progression	103 (60)	5 (29)	27 (55)	71 (66)	
- Controlled disease	16 (9)	4 (24)	7 (14)	5 (5)	
- Unknown	5 (3)	1 (6)	2(4)	2 (2)	
Current anticancer treatment	5 (5)	1 (0)	2 (4)	2(2)	<10-4*
	47 (27)	9 (17)	17 (35)	22 (21)	N10
88	. ,	8 (47) 6 (25)			
- Treatment ongoing	49 (28)	6 (35) 1 (6)	15 (31)	28 (26)	
- Included in clinical trial	12 (7)	1(6)	7 (14)	4 (4)	
- Therapeutic window	16 (9)	2 (12)	7 (14)	7 (7)	
- Palliative care	49 (28)	•••	3 (6)	46 (43)	0.00*
Reason of admission	04 (54)	10 (7()	22 ((5)	10 (16)	0.02*
Acute disease	94 (54)	13 (76)	32 (65)	49 (46)	
- Respiratory disorder	51 (54)	10 (77)	20 (63)	21 (43)	
- Neurological disorder	18 (19)	•••	3 (9)	15 (31)	
Improvement of supportive care	69 (40)	3 (18)	13 (27)	53 (50)	
Scheduled hospitalisation	2(1)		1 (2)	1(1)	
Toxicity of anticancer treatment	8 (5)	1 (6)	3 (6)	4 (4)	
Planned intensity of care before OF					<10 ⁻⁴ *
- Intensive care	56 (32)	17 (100)	22 (45)	17 (16)	
 Maximal medical care 	104 (60)		27 (55)	77 (72)	
- Exclusive palliative care	13 (8)			13 (12)	
Organ failures					
Time between admission and OF (days)	8 [3-13]	8 [1-10]	4 [1-10]	10 [6-15]	2. 10-4
Respiratory disorder	68 (39)	10 (59)	26 (53)	32 (30)	5.10-3
Coma*	52 (30)	••••	4 (8)	48 (45)	<10 ⁻⁴
Shock (except septic)	19 (11)	2 (12)	6 (12)	11 (10)	0.93*
Sepsis	18 (10)	2 (12)	10 (20)	6 (6)	0.01*
Death	7 (4)	2 (12) 		7 (7)	-

Data are expressed as median (interquartile range) for quantitative variables and numbers (%) for qualitative variables. *Use of Fisher exact. CCI: Charlson Comorbidity Index; MD:

missing data; NSCLC: non-small cell lung cancer; OF: organ failure.

*Reason of coma were: complication of central nervous system metastasis (n=22), drug-related (n=13)

or as a consequence of another organ failure (n=13). No explanation was identified for 4 patients.

Variables	Total of the 3 admissions	1 st admission n=127	2^{nd} admission n=31*	3 rd admission
	n=173		Excluding 3 patients	n=15
Reason of no discussion in the	n=15 (9)	n=13 (10)	n=1 (3)	n=1 (7)
medical chart				
- Patient/ relative unable for	3 (20)	3 (23)	•••	••••
discussion				
- Sudden death	2 (13)	2 (15)		
- Rapidly reversible	2 (13)	1 (1)		1 (100)
- No explanation / missing data	8 (53)	7 (54)	1 (100)	
1 st discussion	158 (91)	n=114 (90)	n=30 (97)	n=14 (93)
- Time from admission (day)	5 [2-9]	5 [2-9]	6 [2-10]	5 [1-10]
- Before organ failure	141 (89)	103 (90)	25 (81)	13 (93)
- End of life condition	73 (46)	49 (43)	14 (47)	10 (71)
- Cancer	61 (39)	47 (41)	11 (37)	3 (21)
- Severity of disease	24 (15)	18 (16)	5 (17)	1 (7)
2 nd discussion	n=86 (50)	n=62 (49)	n=16 (53)	n=8 (53)
- End of life condition	68 (79)	50 (81)	12 (75)	6 (75)
- Cancer	8 (9)	6 (10)	2 (13)	•••
- Severity of disease	10 (6)	6 (10)	2 (13)	2 (25)
End of life discussion during	116 (67)	82 (65)	21 (68)	13 (87)
hospitalisation	~ /		~ /	× /

TABLE 4 Patient and relative information for patients with organ failure (n=173)

Data are expressed as median [interquartile range] for quantitative variables and numbers (%)

for qualitative variables.

*3 patients excluded because of organ failure at 1st hospitalisation.

APPENDIX:

Annex 1. Form for advanced care planning in the thoracic oncology unit

« Advanced care planning	form for patients with advanced dis	ease»	
	part : Legal procedure		
ferring doctor to the patient :			
st meeting date: / /			
gular reassessment is necessary			
- This document refers to the care planning fo	r M./ Ma		
- The goal of this for mis to help decision make	ing in case of acute event, for nurses and on-call physicians.		
 Patient's wishes should be followed. Relative This form is filled under the responsibility of a 		м	n
	a medicine doctor, ey expressed their opinion and comments to enlighten the phy		-
 If the patient cannot express his wishes, a contract of the patient cannot express his wishes, a contract of the patient cannot express his wishes. 	ollegiate procedure should be make*.		
 Propositions of care are regularly reassesse The physician in reaponability of the patient 	d. ient at the time of organ failure remains judge of his decis	lon	
	· · · · · · · · · · · · · · · · · · ·		
AGNOSIS			
 Main comorbidities: 			
SHES	1		
Patient able to express his will	Patient unable to express his v		
tient's wishes :	Collegiate procedure for decision making, written in Patiant's wishes before inability :	the medica	chart
The second se	Was he informed about the diagnosis :	yes 🗆	no 🗆
	Was he informed about the prognosis?	yes 🗆	no 🗆
hat are his wishes about the therapeutic project? :	What were his wishes about the therapeutic project?		
	the theorem is the second seco		
	Are-they advanced directives?	yes 🗆	no 🗆

mself, his will is priority over advanced directives and			••••••••••••••••••••••••••••••••••••••
mself, his will is priority over advanced directives and latives' wishes :	d		
mself, his will is priority over advanced directives and latives' wishes : e-they advanced directives? yes _ no	d Is there a designated proxy?	yes 🗆	no 🗆
mself, his will is priority over advanced directives and latives' wishes : e-they advanced directives? yes _ no	d Is there a designated proxy?		
mself, his will is priority over advanced directives and latives' wishes : e-they advanced directives? yes □ no there a designated proxy? yes □ no Name	d Is there a designated proxy? Name		
mself, his will is priority over advanced directives and latives' wishes : e-they advanced directives? yes □ no there a designated proxy? yes □ no Name	d Is there a designated proxy? Name		
mself, his will is priority over advanced directives and latives' wishes : e-they advanced directives? yes □ no there a designated proxy? yes □ no Name	d Is there a designated proxy? Name Relatives' opinion		
mself, his will is priority over advanced directives and latives' wishes : re-they advanced directives? yes □ no there a designated proxy? yes □ no Name	d Is there a designated proxy? Name		
mself, his will is priority over advanced directives and latives' wishes : re-they advanced directives? yes □ no there a designated proxy? yes □ no Name	d Is there a designated proxy? Name Relatives' opinion Referring doctor's opinion		
mself, his will is priority over advanced directives and latives' wishes : re-they advanced directives? yes □ no there a designated proxy? yes □ no Name	d Is there a designated proxy? Name Relatives' opinion Referring doctor's opinion		
there a designated proxy? yes no	d Is there a designated proxy? Name Relatives' opinion Referring doctor's opinion		
mself, his will is priority over advanced directives and latives' wishes : re-they advanced directives? yes □ no there a designated proxy? yes □ no Name	d Is there a designated proxy? Name Relatives' opinion Referring doctor's opinion		

2nd part : Datas of advanced care planning to help decision making in case of organ failure in a patient with a severe or terminal illness, the physician on duty stays in responsibility of the decision.

Þ				
	1 ^{sr}	2 nd	3rd	4 th
	evaluation	evaluation	evaluation	evaluation
DATE	//	//	//	//
In case of collegiate procedure :				
Name of referring doctor:				
Paramedical team yes no				

	1 ^{sr} evaluation	2 nd evaluation	3 rd evaluation	4 th evaluation
MEDICAL PROPOSITIONS	YES/NO	YES/NO	YES/NO	YES / NO
Blue Code Procedure				
Cardio-pulmonar ressuscitation				
Patient's transfer to intensive care unit (if no, patients stays in medical unit with maximal medical care)				
Oro-tracheal intubation or tracheotomy				
Dialyssis				
Amines				
Non invasive ventilation				
Artificial nutrition				
Start antibiotics				
Transfusion				
Peripheral venous catheter				
Further examination (biology, radiology,)				
Vital signs monitoring				
Anticoagulant				
Fibro-aspiration				
Other : IV fluids, insulin, (precise)				

In every proposition, assessment of patient's confort and treatments: pain killers, anxiolytics,...

The medical team will judge the most appropriated way to inform the patient from the decision taken:

	1sr	2 nd	3rd	4 th
	evaluation	evaluation	evaluation	evaluation
DATE	//	//	//	//
DOCTOR IN RESPONSIBILITY				
&				
Signature				
			1	l

Annex 2. French version of the form for advanced care planning in the thoracic oncology unit

'HU de Grenoble Décembre 2012 - version 6		Etiquette patient :	
	d'aide à la décision		
en cas d'aggravation d'un pat		pathologie avancée»	>
	e : procédure légale		
Nédecin référent du patient : vate de la 1 ^{ère} réunion: / / éévaluation régulière indispensable			
-			
 Ce document synthétise la réflexion de projet de s Il se veut une aide pour les soignants et médecins 			
- La volonté du patient doit être respectée au mieux	L'avis des proches est recueil	i.	
 Ce document est élaboré sous la responsabilité d'i Il fait suite à une réunion au cours de laquelle les s 			lairer au
mieux la décision du praticien.			
 Si le patient n'est pas apte à exprimer sa volonté, Ces propositions sont à réévaluer régulièrement. 	une procedure collegiale s'impo	Se .	
- Le médecin de garde reste juge de ses décision	ns face à la situation qui se p	résente.	
DIAGNOSTIC			
- Pathologie principale			
Stade de la maladie :			
Comorbidité(s) principale(s)			
VIS Patient apte à exprimer sa volonté	Patient non	apte à exprimer sa volonté	
	Procédure collégiale avec dé	cision motivée inscrite dans le dossi	er médical
Avis du patient :	Avis du patient avant l'inap		-
Est-il informé du diagnostic : oui	Etait-il informé du diagnostic Etait-il informé du pronostic		non 🗆
Quelles sont ses volontés concernant le projet	Quel était son avis :		
thérapeutique:			
	Y a-t-il des directives anticipé	es oui 🗆	non 🗆
Tant que le patient est apte à comprendre et à s'exprimer,			
son avis est prioritaire sur les directives anticipées et sur	Y a-t-il une personne de cont		non 🗆
'avis des proches :	Nom		
Y a-t-il des directives anticipées oui 🗆 non 🗆	Avis des proches		·····
Y a-t-il une personne de confiance oui non			
Nom	Avis du médecin consultar	t	
Avis des proches			
	Avis complémentaire éven	tuel	
		de Grenoble.	

2^{ème} partie : Données pour aide à la décision du médecin de garde qui reste juge de ses décisions en cas d'aggravation d'un patient atteint d'une maladie grave ou en fin de vie

	1 ^{ère} évaluation	2 ^{ème} évaluation	3 ^{ème} évaluation	4 ^{ème} évaluation
DATE	//	//	//	//
En cas de décision collégiale :				
Médecin consulté, nom :				
Equipe paramédicale oui 🗆 non 🗆				

	1 ^{ère} évaluation	2 ^{ène} évaluation	3 ^{ème} évaluation	4 ^{ème} évaluation
PROPOSITIONS MEDICALES	OUI / NON	OUI / NON	OUI / NON	OUI / NON
Procédure SIERRA				
Réanimation cardio-pulmonaire				
Transfert du patient en réanimation (Si non, maintien du patient dans le service, avec traitement médical renforcé)				
Intubation ou trachéotomie				
Dialyse				
Amines				
Ventilation non invasive				
Nutrition artificielle				
Débuter ATB				
Transfusion				
Reposer une voie d'abord veineuse				
Examen complémentaire (biologique ou radiologique)				
Prises de constantes				
Anticoagulant				
Fibro-aspiration				
Autre : hydratation, insuline (préciser)		2		

Dans tous les cas, maintien de l'évaluation rapprochée du confort et de ses traitements : antalgie, anxiolyse...

L'équipe médicale jugera de la façon la plus appropriée pour informer le patient des décisions concertées :

	évaluation	évaluation	évaluation	evaluation
ATE	//	//	//	//
ESPONSABLE & nature				
	ESPONSABLE &	ATE// ESPONSABLE	ATE/////////	ATE//// ESPONSABLE &