# **BMJ Open** Defining timeliness in care for patients with lung cancer: a scoping review

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### **ABSTRACT**

**Objectives** Early diagnosis and reducing the time taken to achieve each step of lung cancer care is essential. This scoping review aimed to examine time points and intervals used to measure timeliness and to critically assess how they are defined by existing studies of the care seeking pathway for lung cancer.

Methods This scoping review was guided by the methodological framework for scoping reviews by Arksey and O'Malley. MEDLINE, EMBASE, CINAHL and PsycINFO electronic databases were searched for articles published between 1999 and 2019. After duplicate removal, all publications went through title and abstract screening followed by full text review and inclusion of articles in the review against the selection criteria. A narrative synthesis describes the time points, intervals and measurement guidelines used by the included articles.

Results A total of 2113 articles were identified from the initial search. Finally, 68 articles were included for data charting process. Eight time points and 14 intervals were identified as the most common events researched by the articles. Eighteen different lung cancer care guidelines were used to benchmark intervals in the included articles; all were developed in Western countries. The British Thoracic Society guideline was the most frequently used guideline (20%). Western guidelines were used by the studies in Asian countries despite differences in the health system structure.

**Conclusion** This review identified substantial variations in definitions of some of the intervals used to describe timeliness of care for lung cancer. The differences in healthcare delivery systems of Asian and Western countries, and between high-income countries and low-income-middle-income countries may suggest different sets of time points and intervals need to be developed.

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# BACKGROUND

Lung cancer is the most common cancer, with an incidence of 2.1 million globally during 2018, and is the most frequent cause of deaths in both sexes in 14 regions of the world. Incidence and mortality vary across countries due to differences in smoking prevalence and other risk factors, but overall survival rates are low globally (5 year survival of 10%–20% in most countries) with most patients diagnosed at an advanced stage. I

Timely diagnosis and access to effective treatment are important determinants of

# Strengths and limitations of this study

- ▶ This scoping review documented the commonly studied time points in the lung cancer care pathway and the heterogeneity in naming the intervals and, guidelines adopted in the disease care pathway for lung cancer across different studies.
- Arksey and O'Malley's five-stage scoping review framework and Preferred Reporting Items for Systematic Reviews and Meta-Analyses checklist was followed for this scoping review.
- ➤ This study was informed by a previously published protocol which dictated a transparent and rigorous search strategy for four databases.
- Quality of studies was not assessed.
- Only studies published in English were included in the review, which may miss potential literature in other languages.

outcome in patients with cancer.<sup>2</sup> Higher cancer survival rates are evident in high performing healthcare systems. For example, patients with lung cancer in Japan (33%), Israel (27%) and Korea (25%) have a much higher 5-year survival rate than their counterparts in India, Thailand, Brazil and Bulgaria (all less than 10%).<sup>3</sup> Early diagnosis can improve survival and reduce lung cancer mortality through timely initiation of treatment.<sup>4</sup>

Numerous studies have been conducted to assess timeliness of initiation and completion of cancer treatment. However, the pathway to cancer diagnosis and treatment is complex. The patient journey from onset of symptoms to initiation of treatment involves multiple stages, which vary significantly across different health systems, with different health systems having different 'bottlenecks' in the patient journey.

The patient journey can be categorised into different care time points. Time points are the landmarks or events that take place in a patient journey to healthcare, for example, onset of symptom(s), contact with a healthcare provider, referral, diagnosis, initiation of treatment, and so on. Depending on the



outcome of interest of a research or intervention, intervals are defined by calculating the time between two agreed time points. Timeliness can be defined as reaching different time points of care in a way that supports the best patient outcomes. It usually starts from the date of onset of symptoms and ends at the date of initiation of treatment. Guidelines can be defined as a set of agreed recommendation that aim to streamline the process in each step of the disease care pathway to set routine or standard clinical practice. In some countries, clinical guidelines have been developed to establish a maximal length requirement for the intervals between different time points to ensure optimal patient care outcomes. These have enabled measurement of delay. However, studies describing time intervals often mislabeled these intervals as 'delays' despite a lack of benchmarking, creating confusion among readers. There are also marked variations in the definitions of these intervals across studies, and in how the data were obtained, measured and presented. This ambiguity leads readers to make assumptions about the interpretation of the terms and findings. Moreover, due to differences in health systems, studies are seldom comparable across countries. 6 Referral pathways vary between countries. For example, in some developing countries, all the diagnostic tests required to diagnose a cancer are completed before a patient is referred to a specialist, thus contributing to variation in the definition and length of the diagnostic segment in the care pathway between such developing countries and the developed country which was the source of the guidance.

Existing guidelines for lung cancer care vary in the benchmarks or cut-off values used to describe acceptable limits of time for each step in the disease care pathway. As a result, definitions and measures of 'timeliness of care' vary across countries. Furthermore, the majority of guidelines were developed in Western countries, considering country-specific resources and healthcare mechanisms, and associated with effective referral systems governed by policies. It is unlikely that guidelines developed for Western health systems can be fully effective in poorly resourced health systems, which require different definitions, measurements and guidelines for timely care compatible with their available resources and the strength of their health systems.

Several models were proposed in an attempt to improve consistency in the definition, classification and measurement of timeliness of care, but the models are not devoid of limitations. These include the Andersen model of total patient delay, <sup>11</sup> the model of pathways to treatment <sup>12</sup> and the Aarhus statement. <sup>6</sup> Andersen's model can capture the decisional and behavioural processes that occur before the initiation of treatment, but is limited in its capacity to address the complex and dynamic journey into and through the healthcare system. <sup>12</sup> The subsequently proposed 'Model of pathways to treatment' is a descriptive framework which can encompass the psychological theories with a focus on patient factors in the appraisal and help-seeking intervals. The most recent and widely

accepted framework, 'The Aarhus Statement,' proposes a universal framework to incorporate the issue of lack of consensus in definitions and methods across studies conducted on timeliness of cancer care. It defines four important time points that links different interval durations with patient outcomes to determine targets and guidelines (date of first symptom, date of first presentation to a general practitioner (GP), date of referral and date of diagnosis). It also provides guidance on how to design research with greater precision and transparency. All these models provide an overarching framework that can be adapted to different system contexts. This scoping review aimed to examine time points and intervals used to measure timeliness and to critically assess and compare how they are defined by existing studies of the care seeking pathway for lung cancer.

# **METHODS**

This scoping review followed the methodological framework for scoping reviews by Arksey and O'Malley<sup>14</sup> which was further enhanced by Levac *et al*<sup>15</sup> and the Joanna Briggs Institute.<sup>16</sup> Stages of the scoping review framework included (1) Identifying the research question, (2) Identifying relevant studies, (3) Study selection, (4) Charting the data and (5) Collating, summarising, and reporting the results. The University of York Centre for Reviews and Dissemination guidance for undertaking reviews in health care<sup>17</sup> and the Preferred Reporting Items for Systematic Reviews and Meta-Analyses checklist<sup>18</sup> were followed to ensure the comprehensiveness of the review. This scoping review categorised available definitions and terminologies relating to timeliness in the disease care pathway, without an intention of achieving consensus.

### **Identifying the research guestion**

To address the aim of assessing definitions describing timeliness of seeking and receiving care in patients with lung cancer in published articles, the following research questions were posed:

- 1. What are the time points and intervals commonly identified in the care pathway for lung cancer in the existing literature?
- 2. How is timeliness of seeking and receiving care for lung cancer described and related to guidelines in the existing literature?
- 3. Are there differences in definitions, measurements and benchmarking of timeliness used in Western and Asian countries?

# **Identifying relevant studies**

The study population of included literature was patients with diagnosed lung cancer, irrespective of histological type and disease stage. Studies were identified through the keywords that were used to describe timeliness of seeking care, time points in seeking care and intervals between time points in the disease care pathway. Studies were excluded if timeliness of care or time points and intervals

in the care pathway were ambiguous, were not specific for lung cancer, if the primary focus of the article was not timeliness of care, if the articles were not published in English, or if studies were published only as abstracts. This scoping review included all studies, irrespective of study methodology, quality and publication type to gain a better understanding of how researchers have operationalised and measured timeliness of seeking and receiving care for lung cancer in various study settings between May 1999 and May 2019.

The text contained in the titles and abstracts of the papers from the initial search and the keywords used to describe those articles were used to formulate the search strategies specific to the selected databases. MEDLINE, EMBASE, PsycINFO and CINAHL were searched for published articles. An academic health sciences librarian was consulted on selecting the appropriate keywords and the most appropriate MeSH terms and filters to maximise inclusion of articles within the search, and how to modify them for selected bibliographic databases (full search strategy in online supplemental file 1). Reference lists were screened for relevant articles. Search results were imported into EndNote (V.X9) to organise search results specific to each database and later used to generate the reference list for the review. References were imported to Covidence, which was used for documenting the process including duplicate identification and removal, title and abstract screening, and full-text review for included articles. Detailed keywords mapping and database specific search strategies were published in the protocol of this scoping review. 19

# **Study selection**

Selection of publications involved two stages. First, title and abstract were screened against the inclusion criteria, and second, the potentially relevant papers went through full-text review. To increase the reliability of the decision process all selected papers were independently assessed by at least two researchers. Due to the exploratory nature of this scoping review, a detailed methodological quality assessment was not required. One author (AA) performed a search of the electronic database for literature. Two authors (AA and AR) independently reviewed and screened the abstracts of the searched articles for inclusion. The other two authors (VL and CFM) reviewed the disagreements and resolved by discussion with all the authors.

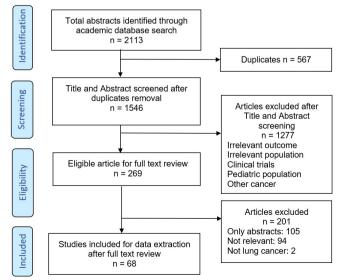
# Data charting, collating and summarising

A data extraction chart was used to capture the data from selected articles (online supplemental file 2), which was recorded on Microsoft Excel 365. Data were extracted by AA independently and examined by authors (VL, CL, CFM and AR).

Initially a coding tree was constructed which had three levels: time points as the first level, time intervals (with starting and ending time point) as the second level, and timeliness (with a definition or benchmarking) as the third level. The initial coding tree was further expanded and divided when new categories emerged from data. An exhaustive list of time points related to seeking or receiving care on the patient care journey was extracted through comparing and merging similar terminologies. The sequence of the time points was determined as follows, (1) patient recalled onset of symptoms, (2) first contact with a healthcare provider, (3) diagnosis, (4) referral to a specialist, (5) first visit to a specialist/ hospital admission, (6) patient informed about diagnosis, (7) pre-initiation of treatment, and (8) initiation of treatment. Afterwards, we summarised and charted the type of intervals examined in the included studies. Intervals in the lung cancer patient care pathway considered the duration between one time point and another time point. Relevant definitions or measurements in relation to the three level coding themes (time points, intervals and timeliness) were also extracted with or without further verification from the cited guidelines. The data on definition of interval or delay were extracted when an article explicitly mentioned the guiding principle (cancer care guideline or self-definition) which included researcher/ study constructed definitions as well. Comparisons between Asian and Western countries were based on the similarities or differences in using time points, intervals and measurement of timelines for intervals.

### **RESULTS**

A total of 2113 articles were identified from the initial search. After duplicates removal, 1546 articles were screened for eligibility and 269 articles were selected for full-text review. Two hundred and one articles were excluded because they were not relevant, only published as abstract or not related to lung cancer. Finally, 68 articles were included for the data charting process (figure 1).



**Figure 1** PRISMA flow chart. PRISMA, Preferred Reporting Items for Systematic Reviews and Meta-Analyses.



Table 1 Characteri	stics of included articles	
N=68	Characteristics of included articles	N (%)
Year of publication	2001–2010 2011–2018	25 (37) 43 (63)
Study setting*	North America (USA, Canada)	21 (30.88)
	UK (England, Scotland, Wales and Northern Ireland)	15 (22.06)
	Europe (Denmark, Netherlands, Norway, Spain, Italy, Sweden, France, Poland, Finland)	13 (19.12)
	Asia (Turkey, India, Mainland China, Taiwan, Nepal)	9 (13.24)
	Australia and New Zealand	8 (11.76)
Study design	Cross-sectional Other study designs Cohort Case control Systematic review Scoping review	41 (60.83) 13 (19.1) 9 (13.2) 3 (4.4) 1 (1.5) 1 (1.5)
Sample size	Range All studies total	12–171208 280591

<sup>\*</sup>Review papers not counted in study settings and sample size.

Characteristics of the included articles are given in table 1 (review articles were excluded).

### **Time points**

Based on the selected articles, time points were classified and the sequence was determined into eight categories (table 2). Commonly mentioned time points included onset of symptom(s), first contact with healthcare provider, diagnosis/first suspicious investigation result, referral/receipt of referral by a specialist (at secondary care), first visit to a specialist/hospital admission, patient informed of lung cancer diagnosis and initiation of treatment.

### **Intervals**

Fourteen different intervals, from onset of symptom(s) to initiation of treatment were identified in this scoping review (table 3): (1) From onset of symptoms to first contact with healthcare provider, (2) From first contact with general healthcare provider to first contact with specialist healthcare provider, (3) From first contact with secondary/tertiary healthcare provider to diagnosis, (4) From first contact with healthcare provider to diagnosis, (5) From diagnosis to contact with secondary/tertiary healthcare provider, (6) From onset of symptoms to contact with secondary/tertiary healthcare provider, (7) From contact with secondary/tertiary healthcare

provider to initiation of treatment, (8) From onset of symptom(s) to referral to a specialist/receipt of referral by a specialist or thoracic department, (9) From referral to a specialist/receipt of referral by a specialist or thoracic department to diagnosis, (10) From onset of symptom to diagnosis, (11) From referral to a specialist/receipt of referral by a specialist or thoracic department to treatment, (12) From first contact with healthcare provider to treatment, (13) From diagnosis to initiation of treatment and (14) From onset of symptom to Initiation of treatment. Intervals were not measured as completion of treatment or death.

Some articles used different terminologies to label the same intervals; and similarly, the same terminology was used to label different intervals in different articles.

- 1. From onset of symptoms to first contact with health-care provider interval: patient delay <sup>21–26</sup> and patient's application interval. <sup>27 28</sup>
- 2. Duration from first contact with healthcare provider to first contact with specialist at secondary care or next level: GP delay, <sup>21</sup> <sup>23–25</sup> GP interval, <sup>29</sup> primary care interval, <sup>30</sup> referral delay <sup>21</sup> <sup>23</sup> and referral interval. <sup>27</sup> <sup>28</sup>
- 3. From first contact with secondary or tertiary health-care provider to diagnosis interval: specialist interval, <sup>29</sup> specialist's delay (second doctor's delay), <sup>21 24 25</sup> diagnosis delay<sup>31</sup> and diagnosis interval. <sup>28</sup>
- 4. From first contact with healthcare provider to diagnosis: diagnostic interval<sup>29 30 32 33</sup> and delay in diagnosis.<sup>34</sup>
- 5. From diagnosis to contact with secondary/tertiary healthcare provider: referral interval in one study.<sup>35</sup>
- 6. Interval between onset of symptom to contact with secondary/tertiary healthcare provider: patient delay.<sup>36</sup>
- 7. Interval between contact with secondary/tertiary healthcare provider and initiation of treatment: hospital delay<sup>25 31</sup> and treatment interval.<sup>35</sup>
- 8. From onset of symptoms to referral to a specialist thoracic department: referral delay,<sup>37</sup> specialist delay.<sup>31</sup>
- 9. From referral to a specialist or receipt of referral by a specialist or thoracic department to diagnosis: referral interval.<sup>30</sup>
- 10. Interval between onset of symptom to diagnosis: total diagnostic delay<sup>29</sup> and time to diagnosis.<sup>38</sup>
- 11. From referral to a specialist/receipt of referral by a specialist or thoracic department to treatment interval: time to treatment (hospital delay)<sup>39</sup> and delay in secondary healthcare.<sup>22</sup>
- 12. Interval between first contact with healthcare provider to treatment: healthcare interval, <sup>30</sup> system delay<sup>22</sup> and doctor's interval. <sup>27 28</sup>
- 13. From diagnosis to initiation of treatment: therapeutic delay, <sup>23</sup> treatment delay, <sup>22 31</sup> treatment interval, <sup>30 33</sup> system interval, <sup>40</sup> pretreatment interval, <sup>32</sup> diagnosisto-treatment delay <sup>41</sup> and diagnosis-to-treatment interval. <sup>42</sup>



Time points	Articles	Definition of time point	Settings
Onset of symptoms	Baughan <i>et al</i> UK <sup>80</sup>	Date patient first noticed symptoms	UK
Shoot of Cymptolic	Corner et al UK <sup>94</sup>	The date, week, or month when a symptom or health change was recalled, and actions taken as a result by the patient were recorded as well as a description of the health change or symptom	
	Dobson et al UK <sup>95</sup>	The date of symptom onset was defined as the first symptom reported	
	Melling et al UK <sup>84</sup>	First symptom reported by the patients to their GPs	
	Neal et al UK <sup>96</sup>	Onset of first symptom	
	Smith et al Scotland <sup>97</sup>	The date participant defined first symptom	
	Salomaa et al Finland <sup>21</sup>	The dates of onset of symptoms	Europe
	Yang et al Mainland China98	First symptom	Asia
	Yilmaz et al Turkey <sup>27</sup>	Date of initial symptoms	
	Özlü <i>et al</i> Turkey <sup>69</sup>	Onset of symptoms	
irst contact with ealthcare provider	Baughan <i>et al</i> UK <sup>80</sup>	Date patient of first presentation with a GP	UK
	Corner et al UK94	Timing of first visit to the GP	
	Dobson et al UK <sup>95</sup>	Date on which person consulted a GP about their symptoms.	
	Smith et al Scotland <sup>97</sup>	Date of presentation to a medical practitioner	
	Melling et al UK <sup>84</sup>	Presentation of the first cancer symptom to the GP	
	Neal et al UK <sup>96</sup>	First presentation (Face-to-face consultations, nurse consultations, telephone consultations) to primary care	
	Vidaver et al USA <sup>68</sup>	First visit to primary healthcare provider	North Americ
	Helsper <i>et al</i> 2017 Netherlands <sup>30</sup>	First contact (physical or telephone) with the GP for suspected cancer-related signs or symptoms	Europe
	Salomaa et al Finland <sup>21</sup>	First visit to a doctor, who was in general, a GP	
	Rankin et al Australia32	First consultation with primary healthcare provider	Australia and
	Largey <i>et al</i> Australia <sup>99</sup>	Dates of first presentation as the time point the clinician started investigation or referral for possible investigation	New Zealand
	Yang et al Mainland China <sup>98</sup>	First contact with local doctor	Asia
	Yilmaz et al Turkey <sup>27</sup>	Date of first doctor visit	
	Özlü <i>et al</i> Turkey <sup>69</sup>	First presentation to a physician	
viagnosis/first uspicious investigation esult	Corner et al UK <sup>94</sup>	Date of diagnosis (the investigation procedure was not specified)	UK
	Neal et al UK96	Date of diagnosis (CT/PET scan, a tissue diagnosis)	
	Melling et al UK <sup>84</sup>	Date of Diagnosis (bronchoscopy, mediastionsocopy, CT scan, bone scan, plural cytology)	
	Vidaver et al USA <sup>68</sup>	First imaging result with a lung abnormality	North Americ
	Singh et al USA <sup>65</sup>	Earliest date that a diagnostic clue could have been recognised by a care provider	
	Li et al Canada <sup>100</sup>	Date of diagnosis	
	Maiga et al USA <sup>42</sup>	Date of pathology diagnosis	
	Schultz et al USA <sup>70</sup>	Date when a pathologic diagnosis of lung cancer was confirmed	
	Grunfeld et al Canada <sup>83</sup>	Date of confirmed diagnosis (date of the pathology or radiology report)	
	Helsper et al Netherlands <sup>30</sup>	Date of the histological confirmation of the primary tumour	Europe
	Rankin et al Australia <sup>32</sup>	Time of the formal cancer diagnosis being made	Australia and New Zealand
	Largey et al Australia99	Date of histological diagnosis	
	Malalasekera <i>et al</i> 2018 Australia <sup>33</sup>	First suspicious investigation report (the investigation procedure was not specified)	
	Özlü <i>et al</i> Turkey <sup>69</sup>	Date of histopathological diagnosis	Asia
	Yang et al Mainland China <sup>22</sup>	Date of diagnosis (CT scan and biopsy)	
	Yilmaz et al Turkey <sup>27</sup>	Date of diagnosis	
Referral to a specialist/ eceipt of referral by a pecialist or thoracic department	Baughan et al UK <sup>80</sup>	Date of decision to refer by primary care	UK

Continued

Time points	Articles	Definition of time point	Settings
	Melling et al UK <sup>84</sup>	Date of referral to secondary care	
	Neal et al UK <sup>96</sup>	Date of GP referral to specialist or admission to hospital	
	Grunfeld et al Canada <sup>83</sup>	Referral for diagnostic assessment was received by the consultant	North Americ
	Vidaver et al USA <sup>68</sup>	Date of referral to a specialist	
	Helsper et al Netherlands <sup>30</sup>	The time point when the responsibility for the patient was transferred from a GP to secondary care	Europe
	Salomaa et al Finland <sup>21</sup>	The date of the writing of the referral requesting consultation from a specialist	
	Stokstad et al Norway <sup>87</sup>	A referral letter for suspected lung cancer was received by the Department of Thoracic Medicine	
	Largey et al Australia99	Date of referral by primary healthcare provider	Australia and
	Malalasekera et al Australia <sup>33</sup>	Date of first referral to secondary care	New Zealand
	Yang et al Mainland China <sup>22</sup>	Date of referral to hospital from primary physician	Asia
First visit to a specialist/ Hospital admission	Baughan et al UK <sup>80</sup>	Date patient first seen by specialist	UK
	Vidaver et al USA <sup>68</sup>	First visit to a specialist	North Americ
	Salomaa et al Finland <sup>21</sup>	The first appointment with the specialist	Europe
	Largey et al Australia <sup>99</sup>	First specialist visit	Australia and New Zealand
	Malalasekera et al 2018 Australia <sup>33</sup>	First specialist visit	
	Alexander et al 2016 Australia <sup>76</sup>	Date of first medical oncology or haematology review for patients with an urgent presentation	
	Yilmaz et al 2008 Turkey <sup>27</sup>	Date of admission to pneumology department	Asia
Patient informed of the cancer diagnosis	Baughan et al 2009 UK <sup>80</sup>	Date patient told the diagnosis	UK
	Grunfeld et al 2009 Canada <sup>83</sup>	Date patient informed of diagnosis	North Americ
	Vidaver et al 2016 USA <sup>68</sup>	Date patient informed of the biopsy result	
Pre-initiation of creatment	Maiga et al USA <sup>42</sup>	<ul> <li>Date of lung nodule identification on CT imaging according to the medical record</li> <li>Date when a lung nodule originally less than 10 mm in size was documented as having new growth on CT imaging.</li> </ul>	North America
nitiation of treatment	Melling et al UK <sup>84</sup>	Date treatment started (surgery, radical radiotherapy with chemotherapy).	UK
	Li et al Canada <sup>100</sup>	Date of first treatment, surgery and adjuvant treatment	North Americ
	Shugarman et al USA <sup>66</sup>	First date recorded for treatment (surgery, radiation, or chemotherapy)	
	Vidaver et al USA <sup>68</sup>	First treatment date	
	Grunfeld et al Canada <sup>83</sup>	Date of initiation of neoadjuvant chemotherapy, surgery if no preoperative treatment was required, chemotherapy, radiotherapy, or a decision not to treat.	
	Maiga et al USA <sup>42</sup>	Time of resection.	
	Stokstad et al Norway <sup>87</sup>	The time for treatment decision as the date when such a decision was documented in the Electronic Medical Record	Europe
	Helsper et al Netherlands <sup>30</sup>	Date of start of therapy as registered in the Network of Cancer Registries	
	lachina et al Denmark <sup>85</sup>	First day of treatment is defined as the date of initiation of surgical, oncological, or radiological treatment, whichever comes first	
	Alexander et al Australia <sup>76</sup>	Time to chemotherapy should be measured from the date that chemotherapy treatment was decided. For adjuvant chemotherapy, time to chemotherapy should be measured from the date of surgery.	Australia and New Zealand
	Evans et al Australia <sup>77</sup>	Date of initial definitive management	
	Malalasekera et al Australia <sup>33</sup>	Treatment start date	
	Rankin et al Australia <sup>32</sup>	Start of treatment	
	Özlü <i>et al</i> Turkey <sup>69</sup>	Start of treatment	Asia
	Yang et al Mainland China <sup>22</sup>	Initiation of treatment date	
	Yilmaz et al Turkey <sup>27</sup>	Date of thoracotomy	

GP, general practitioner.



Table 3 Intervals in	the lung cancer care p	oathway
Intervals	Articles	Study setting
From onset of symptoms	Baughan et al UK80	UK
To First contact with healthcare provider		
	Corner et al UK94	
	Neal et al UK96	
	Smith et al Scotland <sup>97</sup>	
	Brocken <i>et al</i> Netherlands <sup>23</sup>	Europe
	Helsper et al Netherlands <sup>30</sup>	
	Koyi et al Sweden <sup>24</sup>	
	Salomaa et al Finland <sup>21</sup>	
	Sawicki et al Poland <sup>101</sup>	
	Rolke et al Norway <sup>25</sup>	
	Ezer et al Canada <sup>81</sup>	North America
	Ellis and Vandermeer Canada <sup>43</sup>	
	Verma et al Australia <sup>102</sup>	Australia and New Zealand
	Thapa et al Nepal <sup>26</sup>	Asia
	Yang <i>et al</i> Mainland China <sup>41</sup>	
	Yilmaz et al Turkey <sup>27</sup>	
	Özlü et al Turkey <sup>69</sup>	
	Sulu et al Turkey <sup>28</sup>	
From first contact with general healthcare provider To First contact with specialist healthcare provider	Forrest et al UK <sup>78</sup>	UK
	Baughan et al UK80	
	Barrett and Hamilton 2008 UK <sup>103</sup>	
	Devbhandari et al UK <sup>71</sup>	
	Melling et al UK <sup>84</sup>	
	Girolamo et al UK <sup>79</sup>	
	Rolke et al Norway <sup>25</sup>	Europe
	Hueto Pérez De Heredia et al Spain <sup>72</sup>	
	Koyi et al Sweden <sup>24</sup>	
	Helsper et al Netherlands <sup>30</sup>	
	Salomaa et al Finland <sup>21</sup>	
	Brocken <i>et al</i> Netherlands <sup>23</sup>	
	Vidaver et al USA <sup>68</sup>	North America
	Olsson et al USA <sup>104</sup>	
	Ellis and Vandermeer Canada <sup>43</sup>	
	Grunfeld et al Canada <sup>83</sup>	
		Continued

Table 3   Continued		
Intervals	Articles	Study setting
	Verma et al Australia 102	Australia and New
	Emery et al Australia <sup>29</sup>	Zealand
	Sood <i>et al</i> New Zealand <sup>73</sup>	
	Yilmaz et al Turkey <sup>27</sup>	Asia
	Thapa et al Nepal <sup>26</sup>	
	Sulu et al Turkey <sup>28</sup>	
From first contact with secondary/tertiary healthcare provider To diagnosis	Salomaa et al Finland <sup>21</sup>	Europe
	Rolke et al Norway <sup>25</sup>	
	Koyi et al Sweden <sup>24</sup>	
	Gozalez et al Spain <sup>31</sup>	
	Ellis and Vandermeer Canada <sup>43</sup>	North America
	Emery et al Australia <sup>29</sup>	Australia and New Zealand
	Sulu et al Turkey <sup>28</sup>	Asia
	Özlü <i>et al</i> Turkey <sup>69</sup>	
From first contact with healthcare provider To diagnosis	Barrett and Hamilton UK <sup>103</sup>	UK
	Corner et al UK94	
	Devbhandari et al UK <sup>71</sup>	
	Forrest et al UK <sup>78</sup>	
	Neal et al UK <sup>96</sup>	
	Helsper et al Netherlands <sup>30</sup>	Europe
	Ezer et al Canada <sup>81</sup>	North America
	Vidaver et al USA <sup>68</sup>	
	Emery et al Australia <sup>29</sup>	Australia and New Zealand
	Rankin et al Australia <sup>32</sup>	
	Özlü <i>et al</i> Turkey <sup>69</sup>	Asia
	Hsieh et al Taiwan34	
From diagnosis to contact with secondary/ tertiary healthcare provider	Kanarek et al USA <sup>35</sup>	North America
	Wai et al Canada <sup>105</sup>	
	Winget et al Canada <sup>106</sup>	
	Zullig et al USA <sup>107</sup>	
From onset of symptoms To contact with secondary/ tertiary healthcare provider	Bjerager <i>et al</i> Denmark <sup>108</sup>	Europe
	Ampil et al USA <sup>36</sup>	North America
	Thapa et al Nepal <sup>26</sup>	Asia

rinued Continued

Intervals	Articles	Study setting
From contact with secondary/tertiary healthcare provider To initiation of treatment	Devbhandari et al UK <sup>86</sup>	UK
	Girolamo et al UK <sup>79</sup>	
	Gozalez et al Spain <sup>31</sup>	Europe
	Rolke et al Norway <sup>25</sup>	
	Hueto Pérez De Heredia et al Spain <sup>72</sup>	
	Hubert et al Canada <sup>109</sup> Kanarek et al USA <sup>35</sup>	North America
	Winget et al Canada <sup>106</sup>	
	Vidaver et al USA <sup>68</sup>	
	Ellis and Vandermeer Canada <sup>43</sup>	
	Ampil et al USA <sup>36</sup>	
	Olsson et al USA <sup>104</sup>	
	Wai et al Canada <sup>105</sup>	
	Verma et al Australia <sup>102</sup>	Australia and New Zealand
From onset of symptoms to referral to specialist/ receipt of referral by a specialist or thoracic department	Lee et al UK <sup>74</sup>	UK
	Gozalez et al Spain <sup>31</sup> Buccheri and Ferrigno Italy <sup>37</sup>	Europe
From referral to a specialist/ receipt of referral by a specialist or thoracic department to diagnosis	Barrett and Hamilton UK <sup>103</sup>	UK
	Smith et al Scotland <sup>97</sup>	
	Helsper et al Netherlands <sup>30</sup>	Europe
	Grunfeld et al Canada <sup>83</sup>	North America
	Evans et al Australia <sup>77</sup>	Australia and New Zealand
	Largey et al Australia <sup>67</sup>	Zealaliu
	Sood <i>et al</i> New Zealand <sup>73</sup>	
From onset of symptoms to		UK
diagnosis	Lee et al UK <sup>74</sup>	
	Walter et al UK <sup>38</sup>	
	Koyi et al Sweden <sup>24</sup>	Europe
	Wai <i>et al</i> Canada <sup>105</sup>	North America
	Emery et al Australia <sup>29</sup>	Australia and New Zealand
	Sachdeva et al India <sup>88</sup>	Asia
	Chandra et al India <sup>41</sup>	
	Dubey et al India89	

Table 3 Continued		
Intervals	Articles	Study setting
From referral to a specialist/ receipt of referral by a specialist or thoracic department to treatment	Devbhandari et al UK <sup>71</sup>	UK
	Smith et al Scotland <sup>97</sup>	
	Forrest et al UK <sup>78</sup>	
	Bozcuk and Martin UK <sup>39</sup>	
	lachina et al Denmark <sup>85</sup>	Europe
	Olsson et al USA <sup>104</sup>	North America
	Grunfeld et al Canada <sup>83</sup>	
	Ampil et al USA <sup>36</sup>	
	Evans et al Australia <sup>77</sup>	Australia and New Zealand
	Largey et al Australia <sup>67</sup>	Zealand
	Sood <i>et al</i> New Zealand <sup>73</sup>	
	Yang et al Mainland China <sup>22</sup>	Asia
From first contact with healthcare provider to treatment	Melling et al UK <sup>84</sup>	UK
	Helsper et al	Europe
	Netherlands <sup>30</sup>	
	Sawicki et al Poland <sup>101</sup>	
	Vidaver et al USA <sup>68</sup>	North America
	Ezer et al Canada <sup>81</sup>	
	Yang et al Mainland China <sup>22</sup>	Asia
	Yilmaz et al Turkey <sup>27</sup>	
	Özlü <i>et al</i> Turkey <sup>69</sup>	
	Sulu et al Turkey <sup>28</sup>	
From diagnosis to	Forrest et al. 2014 UK <sup>78</sup>	UK
initiation of treatment	Brocken <i>et al</i> Netherlands <sup>23</sup>	Europe
	Gozalez et al Spain31	
	Salomaa et al Finland <sup>21</sup>	
	Helsper <i>et al</i> Netherlands <sup>30</sup>	
	lachina et al Denmark <sup>85</sup>	
	Schultz et al USA <sup>70</sup>	North America
	Kanarek et al USA <sup>35</sup>	
	Grunfeld et al Canada <sup>83</sup>	
	Borrayo et al USA <sup>110</sup>	
	Kim et al Canada <sup>40</sup>	
	Olsson et al USA <sup>104</sup>	
	Ost et al USA <sup>75</sup>	
	Yorio et al USA <sup>111</sup>	
	Zullig et al USA <sup>107</sup>	
	Li et al Canada <sup>100</sup> Maiga et al USA <sup>42</sup>	
	Vidaver et al USA <sup>68</sup>	
		Continued

Continued Continued



initiation of treatment Koyi et al Sweden<sup>24</sup> Rolke et al Norway<sup>25</sup> Sawicki et al Poland<sup>10</sup> Ellis and Vandermeer North America Canada<sup>43</sup> Olsson et al USA<sup>104</sup> Verma et al Australia Australia and New Zealand

India41

Europe

Asia

From onset of symptoms Salomaa et al Finland<sup>21</sup>

14. From onset of symptom(s) to initiation of treatment: global delay, 43 total delay 25 and symptom to treatment

Yilmaz et al Turkey<sup>27</sup>

Özlü et al Turkey69

Sulu et al Turkev<sup>28</sup>

Chandra et al India<sup>41</sup>

Table 4 presents the time intervals commonly studied in the included articles. The most frequently studied interval was 'diagnosis to initiation of treatment', followed by 'first contact with healthcare provider to specialist'

and 'symptom onset to first contact'. Both 'diagnosis to specialist' and 'specialist to diagnosis' paths were studied. Very few studies have researched onset of symptom to referral and specialist consultation. The time point 'patient informed of diagnosis' and intervals involving this time point were rarely studied.

# **Timeliness measures**

The review identified 30 articles which conceptualised delay in the care pathway by adapting benchmarks from established guidelines to set cut-off values. The benchmarks were guided by British Thoracic Society (BTS) recommendations on organising the care of patients with lung cancer, 44 National Institute for Clinical Excellence (NICE) guideline, 45 46 UK National Cancer Plan (UKNCP), 47 UK National Health Service (UKNHS) guideline, 48 49 UK Department of Health guideline, 50 Research and Development (RAND) Corporation guideline,<sup>51</sup> Canadian Strategy for Cancer Control, 52 Canadian guidelines, 53 Standing Medical Advisory Committee (SMAC), 54 Cancer Council Australia and Cancer Australia, 55 Danish Lung Cancer Group and Registry,<sup>56</sup> Swedish Lung Cancer Group<sup>57</sup> and Scottish Executive Health Department (SEHD), 58 59 Institute of Medicine, 60 Dutch Association of Physicians for Pulmonary Disease and Tuberculosis, 61 Joint Council for Clinical Radiology, 62 American College of Chest Physicians, <sup>63</sup> and Norwegian National

Six articles referenced cut-off values from other articles to compare timeliness<sup>24</sup> 35 41 65-67 and one article proposed a benchmark cut-off value based on their findings. 68 Fifteen articles used single guidelines and fifteen articles used more than one guideline to conceptualise timeliness measures. Out of 30 articles, BTS was adopted by 14 articles. 23 25 27 28 33 41 65 69-75 UKNHS was used seven times, <sup>33</sup> 67 72 76-79 NICE guideline by four articles, <sup>71</sup> 73 80 81 RAND corporation guideline by four articles 33 70 75 82 and Canadian guidelines by four articles, 27 28 41 83 SEHD guidelines by three articles, <sup>33</sup> 80 84 Danish Lung Cancer Group guidelines by three articles, 33 67 85 UKNCP guidelines by two articles, <sup>71</sup> <sup>86</sup> SMAC guideline by two articles, <sup>33</sup> <sup>84</sup>

Table 4 Time intervals commonly studied—dark blue >10 (most commonly), light blue >7 (commonly), lighter blue >3 (occasionally), white=none

	Ending point					
Starting point	First contact with healthcare provider	Referral	Specialist consultation	Diagnosis	Patient informed of diagnosis	Initiation of treatment
Onset of symptom	18	3	3	9	-	11
First contact with healthcare provider	X	-	22	12	-	9
Referral		X	-	7	-	12
Specialist consultation			Χ	7	-	14
Diagnosis			4	X	3	28
Patient informed of diagnosis					X	3

Norwegian National Guidelines by two articles<sup>25</sup> 87 and Swedish Lung Cancer Group guidelines by two articles.<sup>28</sup> 33 Online supplemental file 3 describes the 'measures of timeliness'/'benchmark for intervals' with cut-off values adopted from different guidelines. Table 5 presents the timeliness measures according to study settings.

BTS guidelines were those most frequently cited in the included studies (20%). Studies guided by the BTS guidelines adapted the definition of intervals and measurement of timeliness depending on the interval of interest. Common timeliness measures adapted from BTS included the length of time that should elapse from initial GP referral of suspected lung cancer to evaluation/ respiratory assessment (≤1 week), primary care referral to receiving diagnostic tests (bronchoscopy/histology/ cytology) (≤2 weeks), presentation of symptom to diagnosis (≤8 weeks), diagnosis to initiation of treatment (≤6 weeks), GP referral to specialist consultation (≤1 week), GP referral and initiation of any type of treatment (≤62 days), specialist consultation and surgery (thoracotomy) (≤8 weeks), surgical waiting list and thoracotomy (4 weeks), referral to surgeons (≤4 weeks), oncology referral to commencement of radiotherapy or chemotherapy (≤2 weeks), decision-to-treat to initiation of treatment (31 days).

Table 6 presents the frequently used intervals and guidelines to measure timeliness in the included articles.

### **Differences between Asian and Western countries**

There were nine studies from five Asian countries/territories included in the scoping review. There were no differences in the terminology for labelling time points and intervals in the lung cancer care pathway between studies from Asian and Western countries. Studies from Asian countries/territories adapted timeline for intervals from Western guidelines in many instances. One study from India<sup>41</sup> and several Turkish<sup>27</sup> 28 69 studies measured timeliness by adapting guidelines from the BTS, Canada and Sweden. The reporting of timeliness was not described as being guided by any specific guideline in studies from mainland China, <sup>41</sup> Nepal, <sup>26</sup> Taiwan<sup>34</sup> and two other studies from India. <sup>88</sup> 89

# **DISCUSSION**

The lung cancer care journey is not linier. Eight time points found to be most frequently used time points in the included studies, which leads to variations in selection of time points and measurements of intervals (determined by the context) in different studies. Which introduces challenges in assessing timeliness due to lack of appropriate benchmarking, in particular in Asian countries. Moreover, different time points and intervals were defined, and different guidelines were used depending on the interest of the study objectives. This also makes comparisons across studies difficult.

# **Time points**

Different time points were studied depending on the objective of the research in the included studies. 'Onset

 Table 5
 Most frequently cited guidelines used to measure timeliness across settings

unit	Guidelines	Articles included	Settings
1.	British Thoracic Society	Lee et al UK <sup>74</sup>	UK
١.	British Phoracic Godlety	Forrest et al UK <sup>78</sup> Singh et al USA <sup>65</sup> Schultz	North America
		et al USA <sup>70</sup> Olsson et al USA <sup>104</sup> Ost et al USA <sup>75</sup>	Norm America
		Brocken et al Netherlands <sup>23</sup> Rolke <i>et al</i> Norway <sup>25</sup>	Europe
		Malalasekera et al Australia <sup>33</sup> Sood et al New Zealand <sup>73</sup>	Australia and New Zealand
		Özlü et al Turkey <sup>69</sup> Yilmaz et al Turkey <sup>27</sup> Sulu et al Turkey <sup>28</sup> Chandra et al Indian <sup>41</sup>	Asia
2.	UK National Health Service	Barrett and Hamilton 2008 UK <sup>103</sup>	UK
		Hueto Pérez De Heredia et al Spain <sup>72</sup>	Europe
		Malalasekera et al Australia <sup>33</sup> Alexander et al Australia <sup>76</sup> Evans et al Australia <sup>77</sup> Sood et al New Zealand <sup>73</sup> Largey et al Australia <sup>67</sup>	Australia and New Zealand
3.	National Institute for Clinical Excellence guideline	Baughan et al UK <sup>80</sup> Forrest et al UK <sup>78</sup>	UK
		Olsson et al USA <sup>104</sup>	North America
		Verma et al Australia <sup>102</sup>	Australia and New Zealand
4.	RAND corporation	Schultz et al USA <sup>70</sup> Ost et al USA <sup>75</sup> Bullard et al USA <sup>82</sup>	North America
		Malalasekera et al Australia <sup>33</sup>	Australia and New Zealand
5.	Canadian guidelines	Grunfeld <i>et al</i> et al. 2009 Canada <sup>83</sup>	North America
		Yilmaz et al Turkey <sup>27</sup> Sulu et al Turkey <sup>28</sup> Chandra et al India <sup>41</sup>	Asia
6.	Scottish Executive Health Department	Baughan <i>et al</i> UK <sup>80</sup> Melling <i>et al</i> UK <sup>84</sup>	UK
		Malalasekera et al Australia <sup>33</sup>	Australia and New Zealand
7.	Danish Lung Cancer Group	lachina et al Denmark <sup>85</sup>	Europe
		Malalasekera et al Australia <sup>33</sup> Largey et al Australia <sup>67</sup>	Australia and New Zealand
8.	UK National Cancer Plan	Forrest <i>et al</i> UK <sup>78</sup> Devbhandari <i>et al</i> UK <sup>86</sup>	UK
9.	Standing Medical Advisory Committee	Melling et al UK <sup>84</sup>	UK
		Malalasekera et al Australia <sup>33</sup>	Australia and New Zealand
10.	Norwegian National Guidelines	Stokstad <i>et al</i> Norway <sup>87</sup> Rolke <i>et al</i> Norway <sup>25</sup>	Europe
11.	Swedish Lung Cancer Group	Malalasekera et al Australia <sup>33</sup>	Australia and New Zealand
		Sulu et al Turkey <sup>28</sup>	Asia

Continued



Tab	ole 5 Continued		
	Guidelines	Articles included	Settings
12.	Cut-off values referenced from other articles	Singh <i>et al</i> USA <sup>65</sup> Shugarman <i>et al</i> USA <sup>66</sup> Kanarek <i>et al</i> USA <sup>35</sup>	North America
		Koyi et al\ Sweden <sup>24</sup>	Europe
		Largey et al Australia <sup>67</sup>	Australia and New Zealand
		Chandra et al India <sup>41</sup>	Asia

**BAND** Research and Development

of symptoms', 'first contact with a healthcare provider, 'specialist consultation', 'diagnosis' and 'initiation of treatment' were the most frequently studied time points . The first event in any health-seeking behaviour relates to the first health changes or the onset of symptom(s). It is difficult to capture the exact time point of onset of symptom(s) except by asking respondents directly. It may also be difficult to establish a link between onset of symptoms and health-seeking behaviour relating to the diagnosis of lung cancer as similar symptoms are shared by other respiratory diseases. Included studies obtained data from a variety of sources including cancer registries, longitudinal surveillance data, insurance claims data, and hospital records. Not all the studies included the time point 'onset of symptoms' because of the differences in the interval of interest or objective of the study. The relevance and importance of the first time point to understanding the overall patient care pathway is likely to vary across countries with different health systems and resources. In contrast, clinical processes post diagnosis are highly standardised. As a result, research about timeliness in healthcare is focused primarily on the time points prior to diagnosis.

After onset of symptom(s) the next time point in the care seeking pathway is first contact with any healthcare provider. The studies included in this review reported only contact with formal healthcare providers. This may have been because of the difficulty involved in capturing reliable information on seeking healthcare from informal healthcare providers in the absence of any specific record management system and because of the potential for recall bias associated with self-report. Nonetheless, informal healthcare providers (including provision of over-the-counter medicines from unregulated pharmacies, village doctors and traditional or herbal remedies) are predominant in developing countries where, sometimes, informal healthcare is the only available healthcare option accessible. 90 It was evident from the included studies that patients' movement across different tiers of the health system is dynamic and complex. These different tiers within the systems are often not interlinked and using different medical record systems. However, the studies do not necessarily interpret or present this information in a way that makes it easy to understand why the time points are not consistently recorded.

After first contact with any healthcare provider the next time point in the lung cancer care pathway is diagnosis or referral to the next level of healthcare for evaluation of the disease. The way this occurs will depend on the characteristics of the healthcare system and patient behaviour. In some settings, there may be multiple contacts with different providers and the diagnosis could be made at any point, not just as an 'endpoint' before hospital admission. Furthermore, the way patients move across different sectors and services will vary across health systems but may not be described clearly in studies. Patients do not necessarily move through time points in sequential order. In some systems, patients may bypass certain time points. Most included studies were conducted in countries with a 'gate keeper' system consisting of GPs as the first point of contact for healthcare. However, this pathway is not common to all healthcare systems, and was generally not seen in studies from Asian countries. In these countries, confirmatory investigation requisition can be initiated before the referral to a specialist. For instance, a request for a CT and fine needle aspiration cytology can be initiated by a primary care physician and hence, a patient can be diagnosed with lung cancer by a GP before referral to secondary healthcare. Some of the studies included a time point reflecting hospital admission or first specialist visit date. Inclusion of referral time and hospital admission time or first specialist consultation time helped to measure the time elapsed from date of referral to consultation with a specialist or hospital admission. The date when a patient was informed of his/her diagnosis was mentioned by three studies. The last time point in the disease care pathway is the date of initiation of any oncological treatment.

## **Intervals**

Studies have segmented the lung cancer care pathway into different intervals depending on the objectives of those studies and sources of data. 'Onset of symptom' to 'first contact with any healthcare provider', 'first contact with any healthcare provider to 'specialist consultation', 'first contact with any healthcare provider to 'diagnosis' and 'diagnosis' to 'initiation of treatment' were the most commonly used intervals in the included articles. However, there were marked differences in how the intervals were named and this heterogeneity in typologies can be misleading as the same name is used for different intervals. For instance, the 'patient's application interval' and 'the time between onset of symptoms to first contact with primary healthcare provider' were descriptions of the same interval in two studies<sup>27</sup> <sup>28</sup> while the term 'patient delay' was used to measure both 'onset of symptom to primary healthcare provider' and 'onset of symptom to secondary healthcare provider, intervals. 'Patient delay' may not be entirely related to patient factors as lack of health resources can influence the time lapse from onset of symptom to contact with a healthcare provider.

Similarly, the interval 'first contact with a primary healthcare provider to secondary healthcare provider'

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Table 6 Guidelines and interval benchmarks referenced in included articles	erval ber	chmarks	referenced in in	cluded artic	sels										
BTS	NICE	UKNCP	UKNHS UKDOH	H RAND	cscc	SMAC	SEHD	SIGN	NOLCP	CCA	SLCG	DLCG	DAPPDT NNG	ACCP	IOM
Onset of symptoms to first doctor visit															
First clinical presentation to first suspicious investigation												_			
First abnormal investigation (Chest X-Ray) to confirmation of diagnosis/ specialist visit				-											
GP to Specialist	-		-		-	-		_		_	-	_	-	-	-
Primary care to initiation of treatment		_	-	-	-						-	-	•		
Referral to secondary care to diagnosis			-		-							_			
First referral to secondary care to treatment start	-		-				-		_	_		_			
First clinical presentation to Diagnosis				-	-										
First investigation to treatment												-			
Diagnostic investigation to patient informed of diagnosis															
Diagnosis to Treatment start			•	-	-				_	_	-	_	_		
First clinical presentation to treatment start			-			-		_							
Decision to treatment to initiation of treatment		_	_												
Surgery to chemotherapy (Adjuvant chemotherapy)			-												
Referral receipt to specialist consultation	-		-				_								
Oncology referral to radiotherapy/ chemotherapy	-														
Specialist consultation to surgery	-														
Surgeon consultation/ Surgical waiting list to surgery	-				-										
Onset of symptoms to treatment					-										
														Ö	Continued

Table 6 Continued																
BTS	NICE	UKNCP	BTS NICE UKNCP UKNHS UKDoH	RAND	RAND CSCC SMAC SEHD	SMAC		SIGN	NOLCP CCA	CCA	SLCG	DLCG	DAPPDT NNG ACCP IOM	NNG	ACCP	MOI
Primary care referral to first diagnostic evaluation of symptom																
Primary care referral to completion of evaluation at referral centre																

Dutch Association of Physicians for Pulmonary Disease and Tuberculosis; DLCG National Optimal Lung

was measured to reflect 'referral delay' 21 23 25 in some studies<sup>35</sup> and 'diagnosis to secondary/tertiary healthcare provider' and 'referral or receipt of referral by a specialist to diagnosis'30 in others. There were also differences in defining diagnostic intervals including 'from first contact with the secondary healthcare provider to diagnosis', 28 31 'from first contact with primary healthcare provider to diagnosis', 29 30 32-34 and 'from onset of symptom to diagnosis'. 29 38 The interval between 'first contact with primary healthcare provider' and 'treatment initiation' was labelled as 'system delay'22 and 'system interval' and was also described as the 'diagnosis to initiation of treatment' interval. 40 'Treatment delay' was measured using the intervals 'diagnosis to initiation of treatment', 22 and 'onset of symptoms to initiation of treatment'. 41 Use of different terminology for the same intervals and use of the same terminology to label different intervals is confusing and can lead to difficulties in interpretating results. Standardised typology would be helpful in order to streamline consistency and enable comparability across studies.

# **Timeliness**

The terms 'delay' and 'interval' were both used in studies to describe timeliness. The term 'delay' conveys a negative connotation, despite most articles using the term in the absence of benchmarking. It would seem more appropriate to use the term 'time interval' rather than 'delay' as this may imply, inaccurately, that the patient has not sought help promptly. Therefore, several articles suggested using the term 'time interval' as a neutral alternative to 'delay'. 11 12 91 In contrast, other researchers have argued that the term 'time interval' should not be replaced by 'delay' unless the results are compared with others or against benchmarks.

There are some differences in the recommended timeframes for each interval between the guidelines. There were similarities in timeliness measures between the BTS guidelines and most of the European guidelines, with some differences compared with the North American guidelines.

More than half of the included studies (38) did not quantify upper limits for intervals based on existing guidelines. Studies which did not compare their results to any guideline generally compared their results with other timeliness of lung cancer treatment related studies and among the subgroups of patients within the study. Studies also have used different time intervals with different time points. As a result, they were not always comparable between studies. The comparison and interpretation of the results were difficult and created confusion when the studies were not from similar context and health system strength.

# **Asian and Western country differences**

There were no differences between Asian and Western countries in the way they defined timeliness of care. Among 68 studies included in this review, nine studies were from Asian countries and/or territories. 22 26-28 34 41 69 88 89 Four



of nine Asian studies used Western lung cancer guidelines to measure timeliness<sup>27</sup> 28 41 69 and the other five studies did not use a guideline. It remains unclear how effective and relevant Western guidelines are for Asian countries, especially those with low and middle income. The lack of qualified providers, low availability of surgery and radiotherapy services, and poor access to and affordability of up-to-date treatments remain a prevailing concern for lung cancer care in low-income and middleincome countries (LMICs) compared with high-income countries (HICs).<sup>8 9</sup> Moreover, universal healthcare and health insurance mechanisms are still in the development phase in many Asian countries and LMICs. Western guidelines were developed in a context where such health system factors contribute to the effectiveness of guidelines. Using a guideline meant for highly resourced health systems in a resource-constrained country may not accurately reflect expectations and goals for timeliness of lung cancer care; culturally sensitive and resource-sensitive guidelines are likely required.<sup>8</sup> As most of the existing guidelines do not account for diversity in health resources, economic disparities or healthcare infrastructure, their applicability could be limited. 92 93 The articles included from Asian countries/territories did not discuss the compatibility of Western guidelines in terms of relevance and appropriateness of recommended time limits for intervals in the disease care pathway in their context. Although the use of Western guidelines for LMICs with different health systems may not be appropriate, there is currently no guideline for lung cancer care which dictates standard time limits that considers the limitations of weaker health systems. The Asian Oncology Summit 2009 proposed a resource-stratified management guideline for non-small cell lung cancer treatment; however, it does not provide benchmarking for intervals in the care pathway, which need to be developed by respective countries adapting this guideline. <sup>10</sup> Informal healthcare is a unique feature of the diverse healthcare system in Asian countries and LMICs, whereas Western guidelines do not have to consider the inclusion of informal healthcare in the care pathway for lung cancer. Considering inclusion of a time point related to informal healthcare seeking and a measure of the number of times patients sought care from informal healthcare providers could be useful for Asian countries and LMIC settings.

This scoping review is not devoid of limitations. The broad search strategy enabled inclusion of different study designs. This scoping review used a robust and established method guided by a published protocol. Independent screening and assessment of articles against inclusion and exclusion criteria by authors ensured minimisation of selection bias. As this review followed a scoping review methodology, it did not assess the quality of the included articles. Excluding Arksey and O'Malley's optional stage of conducting stakeholder consultation might have limited this scoping review from reaching a consensus, however, the authors intended to undertake stakeholder consultation in the next phase of the research project based on the availability of funding. The majority of

the included studies were from HICs, thus limiting the generalisability for low-income countries. Only studies published in English were included in the review, which could have missed potentially relevant literature in other languages. The search strategy used the most widely used databases; however, articles which were not identified through those databases could have been missed. Although we used common search terms for our search, missing a pertinent term could have limited the search results. Other potential limitations were limiting the search and inclusion of articles published in the last 20 years.

# **CONCLUSION**

Although this review identified similarities in most of the time points and intervals of the included studies, there were substantial variations in selection and interpretation of the meaning of intervals. This lack of consistency creates a challenge for researchers who are trying to undertake research about timeliness of care for lung cancer. As timeliness of care studies are mostly carried out in Western countries and guidelines appear unsuited to weaker healthcare delivery systems, there is a need to revisit existing definitions to conduct timeliness of care related studies and a unified set of definitions needs to be set which can accommodate different structures and characteristics of health systems. The differences in healthcare delivery systems of Asian and Western countries, and between HICs and LMICs may suggest different sets of time points and intervals that reflect resources and feasibility need to be developed. The lack of data capture points in weaker resource-poor health systems and the presence of unregulated and untrained healthcare providers in LMICs make it difficult to conduct research on timeliness of lung cancer care. Differences in the structure and strength of health systems create challenges when comparing results of health service research in lung cancer between HICs and LMICs. Existing frameworks for understanding healthcare pathways such as The Aarhus Statement and Andersen's model of health service utilisation could support synthesis of research but would need to be revisited and modified to be applicable to LMIC-specific contexts.

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# ORCID iDs

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Search strategy for different database

Database	Search strategy
Medline	exp Lung Neoplasms/ OR exp Carcinoma, Non-Small-Cell Lung/ OR exp Carcinoma,
	Small Cell/ OR adenocarcinoma/ OR exp adenocarcinoma, bronchiolo-alveolar/ OR exp
	pulmonary adenomatosis, ovine/ AND General Practitioners/ OR Family Practice/ OR
	General Practice/ OR Primary Health Care/ OR Secondary healthcare.mp. OR Patient
	Admission/ OR exp Tertiary Healthcare/ OR Hospitals, Public/ OR Hospitals, Private/ OR
	Hospitals, Special/ OR Palliative Care/ OR exp Pulmonologists/ OR exp Oncologists/ OR
	exp surgical oncology/ OR exp thoracic surgery/ OR "Referral and Consultation"/ AND
	Diagnostic timelines.mp. OR Delay.mp. OR exp "Early Detection of Cancer"/ OR Primary
	delay.mp. OR Secondary delay.mp. OR Tertiary delay.mp. OR Health system delay.mp.
	OR Timeliness.mp. OR Interval.mp. OR Patient interval.mp. OR Patient delay.mp. OR
	Clinician delay.mp. OR Physician delay.mp. OR *"Referral and Consultation"/ OR Referral
	delay.mp. OR exp *Delayed Diagnosis/ OR Diagnosis delay.mp. OR Diagnostic
	evaluation.mp. OR exp *Time-to-Treatment/ OR Treatment initiation.mp. OR Treatment
	initiation.mp. OR Treatment delay.mp OR exp *Waiting Lists/ OR Wait time.mp. OR exp *"Appointments and Schedules"/ OR Wait time intervals.mp. OR Help seeking
	intervals.mp. OR *Prognosis/ OR Lung cancer Survival.mp. OR Prognostic implication.mp.
	AND limit 43 to (English language and humans and last 20 years)
Embase	exp lung tumor/ OR exp non-small cell lung cancer/ OR exp small cell lung cancer/ OR
	expluing adenocarcinoma/ AND General Practitioners.mp. or exp general practitioner/ OR
	exp primary health care/ OR exp secondary health care/ OR exp tertiary health care/ OR
	exp public hospital/ OR exp private hospital/ OR exp cancer center/ OR exp palliative
	therapy/ OR exp pulmonologist/ OR exp thoracotomy/ OR exp lung lobectomy/ OR exp
	*patient referral/ OR exp consultation/ AND exp delayed diagnosis/ OR Primary delay.mp.
	OR Secondary delay.mp. OR tertiary delay.mp. OR health care system/ OR health care
	system delay.mp. OR timeliness.mp. OR Patient interval.mp. OR Patient delay.mp. OR
	Clinician delay.mp. OR Physician delay.mp. OR delayed lung cancer diagnosis.mp. OR
	time to diagnosis.mp. OR time to treatment.mp. or *time to treatment/ OR Treatment
	initiation.mp. OR treatment delay.mp. OR *hospital admission/ OR Help seeking
	intervals.mp. OR Lung cancer Survival.mp. OR lung cancer prognosis.mp. AND limit 41 to
D 11/50	(human and English language and last 20 years)
PsycINFO	exp neoplasm/ OR (Lung Neoplasms or (lung adj3 neoplasm)).mp. [mp=title, abstract,
	heading word, table of contents, key concepts, original title, tests & measures] OR (lung
	cancer or (lung adj3 cancer)).mp. [mp=title, abstract, heading word, table of contents, key concepts, original title, tests & measures] OR Respiratory tract cancer.mp. OR
	Bronchogenic carcinoma.mp. OR Non-Small-Cell Lung Cancer.mp. OR Non-Small-Cell
	Lung Carcinoma.mp. OR Small Cell lung Cancer.mp. OR Small Cell lung Carcinoma.mp.
	OR (Lung cancer symptom* or (lung cancer adj3 symptom*)).mp. [mp=title, abstract,
	heading word, table of contents, key concepts, original title, tests & measures] AND
	physicians/ or exp family physicians/ or exp general practitioners/ OR (General
	Practitioner* or General practice or Family Practice or Family Physician*).mp. OR (Primary
	healthcare or Secondary healthcare or Tertiary healthcare).mp. OR (Public hospital* or
	Private hospital* or Special hospital* or Cancer hospital* or Cancer Center* or cancer
	centre*).mp. [mp=title, abstract, heading word, table of contents, key concepts, original
	title, tests & measures] OR exp palliative care/ OR Cancer Palliative care.mp. OR
	(Pulmonologist* or oncologist* or thoracic surger*).mp. [mp=title, abstract, heading word,
	table of contents, key concepts, original title, tests & measures] OR (Thoracotom* or Lung
	lobectom* or Pneumonectom*).mp. [mp=title, abstract, heading word, table of contents,
	key concepts, original title, tests & measures] OR (Cancer surgical resection* or Surgical
	resection*).mp. OR (Referral or consultation).mp. OR ((Healthcare adj2 delivery) or patient
	admission).mp. [mp=title, abstract, heading word, table of contents, key concepts, original
	title, tests & measures] AND (Diagnostic timeline* or Timeliness).mp. OR (((early detection adj3 cancer) or delay* detection) adj5 cancer).mp. [mp=title, abstract, heading word, table
	of contents, key concepts, original title, tests & measures] OR (Primary delay* or
	or contente, ney concepts, original title, tests a measures of the finally delay of

Database	Search strategy
	Secondary delay* or Tertiary delay* or Health system delay*).mp. OR (Patient interval* or Patient delay* or Clinician delay* or Physician delay*).mp. OR Referral delay*.mp. OR ((diagnos* adj3 delay*) or diagnostic evaluation).mp. [mp=title, abstract, heading word, table of contents, key concepts, original title, tests & measures] OR ((time adj3 treatment) or treatment initiation).mp. [mp=title, abstract, heading word, table of contents, key concepts, original title, tests & measures] OR Treatment delay*.mp. OR (wait* time* or wait* time* interval or wait* list* or appointment).mp. [mp=title, abstract, heading word, table of contents, key concepts, original title, tests & measures] OR Health service accessibility.mp. OR Help seeking intervals.mp. OR (Prognostic implication* or Lung cancer Survival*).mp. AND limit 38 to (human and English language and last 20 years)
CINAHL	(MH "Respiratory Tract Neoplasms+") OR (MH "Lung Neoplasms+") OR (MH "Carcinoma, Non-Small-Cell Lung/DI/DT/EP/HI/MO/PR/RA/RT/RH/SU/SS/TH") OR (MH "Carcinoma, Small Cell/DI/DT/EP/HI/MO/PR/RA/RT/SU/SS/TH") OR "carcinoma, non-small-cell lung OR Carcinoma, Small Cell lung" OR "lung adenocarcinoma" AND (MH "Physicians, Family") OR (MH "Primary Health Care") OR (MH "Family Practice") OR "general practitioner or gp or family doctor or primary care" OR (MH "Secondary Health Care") OR (MH "Multidisciplinary Care Team") OR (MH "Tertiary Health Care") OR (MH "Hospitals, Public") OR (MH "Hospitals, Private") OR (MH "Hospitals, Veterans") OR (MH "Hospitals, Military") OR (MH "Hospitals, Special") OR (MH "Hospitals, Urban") OR (MH "Hospitals, Rural") OR (MH "Cancer Care Facilities") OR (MH "Oncologic Care+") OR (MH "Pulmonologists") OR (MH "Oncologists") OR "pulmonologist OR oncologist" OR (MH "Surgery, Lung+") OR (MH "Thoracic Surgery+") OR (MH "Pneumonectomy") OR (MH "Referral and Consultation+") OR (MH "Patient Admission") AND "Diagnostic timelines" OR (MH "Early Detection of Cancer") OR "early detection of cancer" OR (MH "Diagnosis, Delayed") OR "delayed diagnosis of cancer" OR "health system delay" OR "timeliness" OR "timeliness in healthcare" OR "timeliness of care" OR "patient delay" OR "patient interval" OR "Physician delay" OR (MH "Treatment Delay") OR "diagnostic delay" OR "diagnostic evaluation" OR "time to treatment" OR "treatment initiation" OR (MH "Waiting Lists") OR "wait* times" OR (MM "Appointments and Schedules") OR "prognostic implication" OR "lung cancer survival" Limiters - English Language; Published Date: 19990101-20190528; Human

Table 1: Timeliness definition and timepoints identified

#	Author, pub date and country	Type/ design of study	Aim of study	Definition/ concept of timeliness in seeking care	Onset of symptom	First visit to healthcare provider	First imaging result with suspicion/ diagnosis	Referral to a specialist	First visit to a specialist	Invasive diagnostic test (e.g. FNAC, biopsy)	Patient informed of the biopsy result	Referral for treatment	Initiation of treatment
1	Alexander et al 2016 Australia	Position paper	Recommendations for the timely triage, review and treatment of cancer patients receiving systemic chemotherapy for six priority cancer groups (breast cancer, colorectal cancer, lung cancer (non-small-cell and small cell), ovarian cancer, lymphoma and myeloma)						The first medical oncology or haematology review for patients with an urgent presentation (Category 1) should occur immediately, within no longer than 48 h of referral receipt. Patients with suspected cancer, not classed as Category 1 or 2 (Category 3), should be seen in a medical oncology or haematology clinic within 14 days of referral receipt as recommended by existing local and international guidelines.			When chemotherapy is the first anti-cancer treatment for a patient, time to chemotherapy should be measured from the date that chemotherapy treatment was decided and the patient was prepared to receive chemotherapy (ready for care) to the date when chemotherapy was first administered (chemotherapy start date). However, in the setting of adjuvant chemotherapy, time to chemotherapy should be measured from the date of surgery.	
2	Ampil et al 2014 USA	Cross sectional	Evaluating the types of delay in the management of people with SVCO-L Ca and the impact of palliative thoracic radiotherapy (PTR) delay on patient outcomes.										
3	Barrett & Hamilton 2008 UK	Nested retrospective case-control study	Aimed at identifying and quantifying clinical features of lung cancer										
4	Baughan et al 2009 UK	Cross sectional	The aim of this study is to gain a better understanding of how quickly patients with cancer initially present to their GP, and how they are then referred to secondary care for further investigation and treatment.		Date patient first noticed symptoms	Date patient first reported symptoms to primary care		Date of decision to refer	Date patient first seen by specialist		Date patient told the diagnosis		
5	Bjerager et al 2006 Denmark	Population based observational case series		Delay in general practice: the time from the patient's presentation of the first symptoms or signs that could be related to the lung cancer until referral to hospital. Delay in general practice was subdivided into: doctor delay: time elapsed without investigation of cancer-related symptoms and signs. System delay: time elapsed due to waiting times related to investigation of cancer-related symptoms and administration.									
6	Borrayo et al 2016 USA	Mixed Method	To better understand the institution- and the patient-level determinants associated with the timely initiation of cancer treatment among underserved Hispanic patients diagnosed with lung and head and neck cancers.										
7	Bozcuk & Martin 2001 UK	Retrospective medical record review	to analyse survival in relation both to time to treatment (hospital delay) and other known prognosticators, in a cohort of NSCLC patients presenting in 1 year in a UK Hospital with thoracic surgery and clinical oncology departments.										

#	Author, pub date and country	Type/ design of study	Aim of study	Definition/ concept of timeliness in seeking care	Onset of symptom	First visit to healthcare provider	First imaging result with suspicion/ diagnosis	Referral to a specialist	First visit to a specialist	Invasive diagnostic test (e.g. FNAC, biopsy)	Patient informed of the biopsy result	Referral for treatment	Initiation of treatment
8	Brocken et al 2012 Netherlands	Retrospective medical record review	To compare various delays in a rapid outpatient diagnostic program (RODP) for suspected lung cancer patients with those described in literature and with guideline recommendations, to investigate the effects of referral route and symptoms on delays, and to establish whether delays were related to disease stage and outcome.	Timeliness of lung cancer care starts with timely recognition of symptoms by patients themselves, which is often inadequate or delayed									
9	Buccheri & Ferrigno 2004 Italy	Retrospective medical record review	provide a more recent profile of the clinical manifestations of lung cancer; 2) evaluate possible time-related changes in the occurrence of symptoms; and 3) explore the possible relationship between symptoms and time to specialist referral.										
10	Bullard et al 2017 USA	Retrospective medical record review	To evaluate the impact that the initiation of timely treatment has on patient survival among a cohort of privately insured patients with NSCLC in South Carolina	Analysis of treatment timeliness was informed by the Andersen and Cacioppo model of delays in seeking cancer care.16 Delay in seeking cancer care is defined as the number of days from the identification of the first symptom to visiting a physician, being diagnosed as having a condition, or beginning a regimen for treating the condition. The model interprets delay as an aggregate of underlying decision-making processes imposed by the patient. Treatment delay is the time between receiving medical attention and when care or treatment is initiated. Timely care was defined according to the RAND Corporation as a maximal time limit of 6 weeks (≤42 days) from diagnosis to treatment.									
11	Corner et al 2004 UK	Exploratory study	To explore the pathway to diagnosis among a group of patients recently diagnosed with lung cancer.		Symptoms were recalled as having started between 4 months and more than 2 years	timing of their visits to the GP	Date of diagnosis						
12	Devbhandari et al 2007 UK	Prospective Cohort	To compare our waiting times with national recommendations										
13	Devbhandari et al 2008 UK	Prospective Cohort	To ascertain the causes of delays in treatment to all patients presenting to our centre with a working diagnosis of lung cancer										
14	Dobson et al 2017 UK	Qualitative study	to explore the patient intervals of people with symptoms of lung or colorectal cancer, considering how symptom appraisal and help-seeking experiences were influenced by the wider context of people's lives, such as family and work.		The date of symptom onset was defined as the first symptom reported	The end of the patient interval was defined as the date on which they consulted about their symptoms.							
15	Ellis & Vandermeer 2011 Canada	Cross sectional	Our objective was to establish the time delays in each phase to help inform strategies to reduce overall diagnostic delays.										

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16	Emery et al 2013 Australia	Mixed methods study	The overall objective of this study was to identify the major subcomponents of the diagnostic interval for rural cancer patients in WA to inform the design of an intervention aimed at reducing time to diagnosis.										
17	Evans et al 2016 Australia	Retrospective cohort study	To assess factors associated with second-line delays in the management of patients diagnosed with lung cancer										
18	Ezer et al 2017 Canada	Cross sectional	The aim of the study was to assess the impact of this model of care (Rapid Investigation Clinic) on timeliness of lung cancer diagnosis, staging and treatment.										
19	Forrest et al 2014 UK	Population-based, data- linkage study	To investigate the factors (socioeconomic position (SEP), age, sex, histology, comorbidity, year of diagnosis, stage and performance status (PS)) that may influence the likelihood of post-primary care referral, diagnosis and treatment within target times.										
20	Kanarek et al 2014 USA	Retrospective cohort	Evaluated the hypothesis that delay to first surgery and other time-related factors reduce survival after treatment (surgery). Then assessed the hypothesis that age, race, gender, place of residence, tumor characteristics, and morbidity confound the relationship between these factors and survival.										
21	Kim et al 2016 Canada	Retrospective medical record review	The aim of this study was to quantify the time intervals that NSCLC patients in Alberta with stage lellI disease spend waiting for diagnosis (diagnostic interval), treatment (treatment interval) and their sum (system interval) and to determine which factors are associated with delays.										
22	Koyi et al 2001 Sweden	Cross sectional	The aim of the present study was to prospectively investigate a material of lung cancer patients in order to measure the delays, both by the patient and by the doctors.										
23	Kudjawu et al 2016 France	Retrospective medical record review	To describe time delays in each phase of lung cancer treatment after bronchoscopy.										
24	Largey et al 2015 Australia	Pilot study.	The audit was conducted as part of routine cancer quality improvement activities at Southern Metropolitan Integrative Cancer Services.			Dates of first presentation as the time point the clinician started investigation or referral for possible investigation		Referral	First specialist appointment	Diagnosis		Referral.	

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	Largey et al 2016 Australia	Retrospective medical record audit	(1) examine the current interval times for lung cancer patients from the point of initial referral to the start of first treatment at three large public principal referral hospitals in Victoria; (2) assess the effects difference treatment type (surgery, radiotherapy and chemotherapy) and health service had on interval times across the selected components of the lung cancer pathway; and (3) compare interval times and identify the proportion of patients who met the established target measures.										
	Lee et,al. 2002 UK	Retrospective medical record audit	assessed the delays in their care against BTS guidelines.										
	Li et al 2012 Canada	Retrospective medical record review	The purpose of this study was to assess the value in measuring specific time intervals across cancer sites to identify potentially important variation in the timeliness of cancer care that may inform needed changes and/or improvements incoordination of care.							dates of diagnosis			first treatment, surgery and adjuvant treatment.
28	Maiga et al 2017 USA	Retrospective cohort study	Investigation of the reasons for delays in treatment and the impact these delays have on tumor-stage progression.										
	Malalasekera et al 2018 Australia	Scoping review	1) synthesise health system related waiting times to milestones of lung cancer care using standardised definitions; 2) benchmark measures of performance against relevant guidelines for timeframes; 3) supplement quantitative findings with barriers to timely care described in the literature; and 4) explore the impact of facilitators such as fast-track referral systems on waiting times.			First clinical presentation	First suspicious investigation	First referral to secondary care	First specialist visit	Diagnosis			Treatment start
	Melling et al 2002 UK	Cross sectional	The purpose of this study was to find out what proportion of patients are referred as lung cancer guidelines assume, whether different referral pathways result in different management and what proportion of patients are seen within recommended time intervals between referral and treatment.	Definitive treatment was defined as surgery (pneumonectomy or lobectomy), radical radiotherapy (radiotherapy directed at treating lung cancer itself) and chemotherapy. Palliative treatment recorded was palliative radiotherapy (for symptom control only), palliative surgery or best supportive care.	Symptom	Presentation	Diagnosis	referral					treatment
	Neal et al 2015 UK	Mixed method	aims to provide a detailed analysis of the diagnostic process of lung cancer from a primary-care perspective.		Onset of first symptom	face-to-face consultations, nurse consultations, telephone consultations, out of hours, home visits before initial referral or investigation request First presentation to primary care	Date of diagnosis  CXR requested  CXR report received  Diagnosis	Referal or admission					

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32	Girolamo et,al. 2018 England	Retrospective medical record review	To assess the association between meeting waiting time targets, as currently available to the policymakers, and individual patients' cancer survival, and measure the time to different types of treatments.	Maximum two-week wait (TWW) between an urgent referral for a suspicion of cancer from a general practitioner (GP) to being seen by a specialist, a maximum 62 days from the referral to the start of the first treatment, and a maximum 31 days from the decision taken to treat a patient to the start of the first treatment, irrespective of the route to diagnosis the patient went through .									
33	Gozalez et,al. 2014, Spain	Retrospective medical record audit	To analyse the delays in the diagnosis and treatment of LC and the factors associated with the timeliness of care and their possible relationship with the survival of these patients										
34	Grunfeld et al 2009 Canada	Cross sectional	To prospectively measure peri- diagnostic and surgical time intervals for patients with suspected colorectal, lung, or prostate cancer				date of the pathology or radiology report	the date the referral for diagnostic assessment was received by the consultant		date of first relevant investigation initiated by consultant, whichever came first; relevant investigations included biopsy, bronchoscopy, chest X-ray, colonoscopy, sigmoidoscopy, CT scan, MRI, PSA, pulmonary function test, transrectal ultrasound, and other	date patient informed of diagnosis		date of initiation of first treatment (first treatment was definedas neoadjuvant chemotherapy, surgery if no preoperativetreatment was required, chemotherapy, radiotherapy, or a decisionfor no treatment
35	Helsper et al. 2017 Netherlands	Retrospective medical record review	To chart the diagnostic pathway for the five most common cancers in the Netherlands			The date of the first cancer-related GP consultation was defined as the first contact (physical or telephone) with the GP for suspected cancer-related signs or symptoms		The date of referral was defined as the moment when the responsibility for the patient was transferred from a GP to secondary care			the date of diagnosis was the date of the histological confirmation of the primary tumour.		The date of treatment initiation denotes the date of start of therapy as registered in the NCR
36	Hsieh et al 2012 Taiwan	Retrospective medical record review	To understand the delay in the diagnosis of lung cancer under the healthcare system in Taiwan, and to identify the factors associated with it										
37	Hubert et al 2018 Canada	Retrospective medical record review	To measure the timeliness of care with a standardized Rapid diagnostic assessment programs (DAP) in patients with early-stage non-small cell lung cancer (NSCLC) and to evaluate the impact of an ERP (enhanced recovery protocols) in these patients.										

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38	Heredia et al 2012 Spain	Cross sectional	To analyze the results obtained in a lung cancer (LC) screening program since its inception five years ago regarding correct referrals, diagnostic and therapeutic delay times and days of hospitalization. To compare the diagnostic—therapeutic delays and hospital stays with those obtained in patients evaluated with the standard system										
39	lachina et al 2017 Denmark	Retrospective cohort study	To investigate the significance of primary investigation and treatment at two or more hospitals on the delay in Danish patients with Non-Small Cell Lung Cancer (NSCLC).	** Time from referral (time of diagnosis) to end of primary investigation = 28 days **Time from referral (time of diagnosis) to first day of treatment = 42 days End of primary investigation is defined as the date of decision on treatment. Referral is defined as the date where the investigating department receives the referral.									First day of treatment is defined as the date of initiation of surgical, oncological, or radiological treatment, whichever comes first
40	Ju et al 2017 USA	Computer process modelling	To evaluate delays in care delivery, in order to identify potential 'bottlenecks' in waiting time, the reduction of whichcould produce greater care efficiency.										
41	Olsson et al 2009 USA	Systematic review	To summarise all recently published studies that described the timeliness of care in patients with lung cancer, identified factors that were associated with more or less timely care, or examined the association between the timeliness of care and lung cancer outcomes, including stage distribution and survival. In addition, we aimed to identify studies that evaluated interventions to improve the timeliness of care for patients with lung cancer.										
42	Ost et al 2013 USA	Guideline/review	This guideline is intended to provide an evidence-based approach to the initial evaluation of patients with known or suspected lung cancer. It also includes an assessment of the impact of timeliness of care and multidisciplinary teams on outcome.										
43	Özlü et al 2004 Turkey	Retrospective medical record review	To determine the delay between the onset and the diagnosis and treatment of patients with lung cancer in two cancer centres in the Eastern Black Sea Region of Turkey.		onset of symptoms	first presentation to a physician				histopathological diagnosis			start of treatment
44	Rankin et al 2017 Australia	Qualitative study	To describe the lung cancer diagnostic pathway, focusing on the perspective of patients and general practitioners about diagnostic and pretreatment intervals			first consultation with HCP	diagnosis						start of treatment

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45	Rolke et al 2006 Norway	Cross sectional	to evaluate the delays in the diagnostic pathways for primary lung cancer in Southern Norway, and to compare results with recommendations from the British Thoracic Society (BTS) and the Swedish Lung Cancer Group (SLCG).	Patients referred by general practitioners, who have obvious clinical evidence of lung cancer, should be seen within 1 week of referral receipt in a respiratory physician's clinic, i.e. Referral delay.  The results of bronchoscopy or any other similar diagnostic test, including the histological or cytological result, should be available and communicated to the patient within 2 weeks of a decision to do it, i.e. Informed diagnostic delay. Suspected lung cancer should wait no more than 1 week before they are investigated by a specialist, i.e. Referral delay. Diagnosed lung cancer should wait no more than 3 weeks since first specialist investigation to a treatment decision was made and no more than 10 days from a treatment decision was made until start of treatment, summarised as Hospital delay.							Tesuit		
46	Thapa et al 2014 Nepal	Cross sectional, prospective observational study.	To identify the steps through which the patients passed before he/she finally arrived to specialist care at Manmohan Cardiothoracic Vascular and Transplant Center (MCVTC) and also determine the time lost in each step.										
47	Verma et al 2018 Australia	Cross sectional	to identify any differences in time delays in lung cancer referral pathways between rural and urban patients and explore patients' perceived barriers to timely lung cancer diagnosis and management.										
48	Vidaver et al 2017 USA	Mixed method	This study explored when and why delays occur in lung cancer care and compared timeliness between two states with divergent disease incidence.	The RAND Corporation suggested that the diagnosis of lung cancer should be established within 2 months of abnormal radiography, and treatment should begin within 6 weeks of diagnosis.  British Thoracic Society recommended that patients with suspected lung cancer be seen by a respiratory specialist within 7 days of referral; a specialist visit should occur within 2 weeks of an abnormal radiograph, and surgery should be within 8 weeks of a visit to a respiratory specialist.		A—first visit to health care provider with symptoms	B— first imaging result with a lung abnormality	C— referral to a specialist	D— first visit to a specialist	E— first diagnostic test F— last diagnostic test	G— patient informed of the biopsy result	H— first referral to treatment	I— first treatment
49	Wai et al 2012 Canada	A case-control study	The primary goal of this study is to investigate if delays in care may decrease the curability of patients with stage III NSCLC.  The secondary goal is to describe the patterns of staging and diagnostic evaluation for palliatively and radically treated patients with stage III NSCLC in British Columbia.										

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50	Walter et al 2015 UK	Prospective cohort study	To investigate the symptoms and other clinical and sociodemographic factors associated with lung cancer diagnosis, time to diagnosis and stage at diagnosis.	The total diagnostic interval (TDI), or 'time to diagnosis', defined as the time from the first symptom/s to the date of diagnosis.									
51	Wilcock et al 2016 UK	Mixed-methods	to identify areas where there may be potential to improve the care provided so as to inform the need for further focused research.										
52	Winget et al 2007 Canada	Stakeholders workshop	1) identify a set of criteria and variables needed to create comparable measures of important time-to-cancer-care intervals that could be applied across provinces and 2) use the measures to compare time-to-care across participating provinces for lung cancer patients diagnosed in 2004.										
53	Yang et al 2015 China	Case control	In this study, we determined the total time from the first symptoms to the initial treatment for lung cancer patients at the Department of Respiratory Disease of Zhongshan Hospital (Fudan University, Shanghai, China), a tertiary health care medical center	In China, a diagnosis delay for lung cancer has been defined as more than 1 month between the first symptom or radiological change and the clinical diagnosis or suspicion for lung cancer.	First symptom	First contact with local doctor		Referral to hospital		Diagnosis/ referral to treatment			Initiation of treatment
54	Yilmaz et al 2009 Turkey	Cross sectional	The aims of this study were to investigate the delays in patients with lung cancer from the first symptom to thoracotomy and to examine whether the delays affect the stage of lung cancer at the time of thoracotomy.	The application interval that exceeded 30 days was considered indicative of a patient's delay.  The interval that exceeded 14 days was considered indicative of a referral delay.  The diagnosis interval that exceeded 14 days was considered as indicative of a delayed diagnosis.  The interval that exceeded 14 days was considered as indicative of a delayed treatment.  The interval that exceeding 6 weeks was considered as indicative of a doctor's delay.  If exceeding 72 days it was considered indicative of a total delay	date of initial symptoms	date of first doctor visit			date of admission to pneumology department of our hospital	date of diagnosis			date of thoracotomy
55	Yorio et al 2009 USA	Cross sectional	to examine the predictors and impact of the timing of lung cancer care in this context, we examined diagnostic and treatment intervals at a large American medical center providing care to a diverse patient population within two different hospital systems.	Date of tissue diagnosis was defined as the date of final pathology report.  Date of treatment was defined as the date of surgery, initial date of chemotherapy, or initial date of radiation therapy, whichever occurred first.									
56	Zullig et al 2013 USA	Cross sectional	Aim 3: Examine patient-level factors associated with (a) receipt of timely lung cancer care and (b) subsequent health outcomes										
57	Sachdeva et al 2017 India	Cross sectional	To determine time delay from the onset of initial symptoms to diagnosis of primary lung cancer.										

#	Author, pub date and country	Type/ design of study	Aim of study	Definition/ concept of timeliness in seeking care	Onset of symptom	First visit to healthcare provider	First imaging result with suspicion/ diagnosis	Referral to a specialist	First visit to a specialist	Invasive diagnostic test (e.g. FNAC, biopsy)	Patient informed of the biopsy result	Referral for treatment	Initiation of treatment
58	Salomaa et al 2001 Finland	Retrospective medical record review	To measure delays of diagnosis and to assess the causes for those delays in patients with lung cancer. To evaluate whether the lengths of the delays were acceptable according to the British recommendations, and To examine the relations between delays and survival			the first symptoms until the first visit to a doctor, who was in general, a GP		the date the consultation request for a specialist was written	the first appointment with the specialist				
59	Sawicki et al 2013 Poland	Cross sectional	To compare the differences in the periods of time and reasons for delay in diagnosisand initiation of treatment of lung cancer among patients who are inhabitants of the rural and urban regions of LublinVoivodeship, and who were consulted in Thoracic Surgery Department										
60	Schultz et al 2009 USA	Cross sectional	To evaluate timeliness of lung cancer care and identify institutional characteristics associated with timely care within the Veterans Affairs (VA) health care system	British Thoracic Society guidelines) *Specialist visit within 2 wk of abnormal CXR *Surgery within 8 wk of specialist visit RAND guidelines *Diagnosis within 8 wk of abnormal CXR *Treatment within 6 wk of diagnosis							Time to diagnosis is the time from the first suspicious chest x-ray or CT scan to the date when a pathologic diagnosis of lung cancer was confirmed		
61	Shugarman et al 2009 USA	Cohort study	To evaluate the relationship of sex and race with the receipt of timely and clinically appropriate NSCLC treatment for each stage of diagnosis	Timely treatment as a 6-week timeframe from the date diagnosis to receipt of treatment (surgery, chemotherapy or radiation therapy)									
62	Singh et al 2010 USA	Cohort study	To evaluate characteristics and predictors of missed opportunities for earlier diagnosis of lung cancer in a health care system with an advanced integrated EHR		the first appearance of a diagnostic clue as the earliest date that the clue could have been recognized by the care providers, regardless of when the patient first started experiencing symptoms								
63	Smith et al 2009 Scotland	Cross sectional	To determine what factors are associated with the time people take to consult with symptoms of lung cancer, with a focus on those from rural and socially deprived areas		the date participant defined first symptom	date of presentation to a medical practitioner							
64	Sood et al 2009 NZ	Retrospective medical record review	To determine the patient characteristics, referral patterns and delays in assessment and treatment of patients with primary lung cancer in South Auckland, New Zealand and compare with international standards										

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65	Stokstad et al 2017 Norway	Retrospective medical record review	To quantify the proportion of patients who started treatment within the recommended timeframes; and to assess the proportion of non-complex patients for which there were no good reasons for delays.	For suspected lung cancer, the first hospital appointment should be offered within seven calendar days of receiving a referral letter; a treatment decision should be made within 28 calendar days; systemic therapy should start within 35 calendar days, and surgery or radiotherapy within 42 calendar days. According to Norwegian recommendations, start of treatment within 42 days (surgery or radiotherapy) or 35 days (systemic therapy) was considered "timely treatment"				start time as the date when a referral letter for suspected lung cancer was received by the Department of Thoracic Medicine – or the date when the decision was made to start diagnostic workup in patients with a known single pulmonary nodule (SPN)					the time for treatment decision as the date when such a decision was documented in the EMR
66	Sulu et al 2011 Turkey	Cross sectional	To investigate patterns of delays among patients with non-small-cell lung cancer and to identify reasons for the delays.	**An application interval that exceeded 30 days was considered indicative of a patient's delay. **The referral interval that exceeded 14 days was considered indicative of a referral delay. **A diagnosis interval that exceeded 14 days was considered as indicative of a delayed diagnosis. **A treatment interval that exceeded 14 days was considered as indicative of a delayed treatment **Doctor's interval that exceeded 6 weeks was considered as indicative of a doctor's delay. ** Total interval exceeded 72 days was considered indicative of a total delay									
67	Chandra et al 2009 India	Retrospective review	To determine the average time period required at various steps for diagnosing lung cancer from the onset of symptoms at a tertiary referral centre in Northern India										
68	Dubey et al 2015 India	Cross sectional	The aim was also to study the time duration for confirming the diagnosis, the relative yield of the investigations in diagnosis of lung cancer and the lung cancer stage in which patients are presenting.										

# Table 2: Intervals identified

#	Author, pub date and country	Symptom to doctor/ GP	GP to LCS/ Chest clinic/ referral/G P to first hospital appointm ent/ admission	Referral to first attendan ce to specialis t	Chest clinic to referral for Chest Physicia n	Chest Physician/ hospital appointment to Diagnosis	GP to diagnosi s	Diagno sis to referral to LCS/ or hospita	Sympto m to hospital admissi on	LCS to treatment	Hospitalizat ion to treatment referral	Diagnostic intervals (imaging/ biopsy)	Referral for treatmen t to initiation of treatmen t	Sympto m to 'referral for diagnosi s'	Sympto m to referral to LCS	Referral for diagnosis' to diagnosis	Sympto m to diagnosi s	Sympto m to referral (by GP or chest physicia n to next Mx)	Symptom to secondary care	Referral to treatment	GP to treatment	Diagnosis to initiation of treatment	Outpatie nt to decision to treat	Decision to treat/ specialist consultatio n to treatment	Symptom to initiation of treatment
·	et al 2016 Australia																								
2	Ampil et al 2014 USA								Patient delay was inferred from the duration of presenti ng sympto ms until hospital admissi on		In-hospital delay was defined as the interval from the date of hospitalizati on to the date of referral for therapy		Professio nal delay was defined as the interval from the date of referral to first treatment												
3	Barrett & Hamilton 2008 UK						First symptom presented to primarly care to diagnosis							Interval between first presentat ion to primary care with a symptom of lung cancer and referral		Interval from referral to diagnosis	The intervals between first symptom presentati on and diagnosis								
4	Baughan et al 2009 UK	time from patient first noticing symptoms to first presentati on with a GP																Time from first presentat ion to time of referral							
5	Bjerager et al 2006 Denmark																		First symptom until referral to secondary care						
6	Borrayo et al 2016 USA																					Diagnosis to treatmentinitiati on			

#	Author, pub date and country	Symptom to doctor/ GP	GP to LCS/ Chest clinic/ referral/G P to first hospital appointm ent/ admission	Referral to first attendan ce to specialis t	Chest clinic to referral for Chest Physicia n	Chest Physician/ hospital appointment to Diagnosis	GP to diagnosi s	Diagno sis to referral to LCS/ or hospita	Sympto m to hospital admissi on	LCS to treatment	Hospitalizat ion to treatment referral	Diagnostic intervals (imaging/ biopsy)	Referral for treatmen t to initiation of treatmen t	Sympto m to 'referral for diagnosi s'	Sympto m to referral to LCS	Referral for diagnosis' to diagnosis	Sympto m to diagnosi s	Sympto m to referral (by GP or chest physicia n to next Mx)	Symptom to secondary care	Referral to treatment	GP to treatment	Diagnosis to initiation of treatment	Outpatie nt to decision to treat	Decision to treat/ specialist consultatio n to treatment	Symptom to initiation of treatment
7	Bozcuk & Martin 2001 UK												Time to treatment (measure of hospital delay): time from receipt of referral letter from GP /referring physician to first treatment .  Referral time (measure of referral delay): time from receipt of GP /referring physician referral letter to first appointm ent in Norfolk & Norwich Hospital. It actually is a componen to f time to treatment												
8	Brocken et al 2012 Netherlan ds	Patient delay as the time from first symptom until the first visit to a GP	GP delay as the time between first GP visit and referral to a chest physician		referral delay as the time between referral (written or by phone) and first rapid outpatient diagnostic program (RODP) day	Diagnostic delay as the time between first RODP day and date of final (accurate) diagnosis																Therapeutic delay as the time between diagnosis and start of treatment.			
9	Buccheri & Ferrigno 2004 Italy				ucy										Referral delay was defined as the time interval between the occurren ce of the first sympto m of alarm (as reported by the patients and confirme d by their relatives) and the date of the first specialis t referral made to the study group). (normall y made to the										

#	Author, pub date and country	Symptom to doctor/ GP	GP to LCS/ Chest clinic/ referral/G P to first hospital appointm ent/ admission	Referral to first attendan ce to specialis t	Chest clinic to referral for Chest Physicia n	Chest Physician/ hospital appointment to Diagnosis	GP to diagnosi s	Diagno sis to referral to LCS/ or hospita	Sympto m to hospital admissi on	LCS to treatment	Hospitalizat ion to treatment referral	Diagnostic intervals (imaging/ biopsy)	Referral for treatmen t to initiation of treatmen t	Sympto m to 'referral for diagnosi s'	Sympto m to referral to LCS	Referral for diagnosis' to diagnosis	Sympto m to diagnosi s	Sympto m to referral (by GP or chest physicia n to next Mx)	Symptom to secondary care	Referral to treatment	GP to treatment	Diagnosis to initiation of treatment	Outpatie nt to decision to treat	Decision to treat/ specialist consultatio n to treatment	Symptom to initiation of treatment
10	Bullard et														study group).										
11	al 2017 USA Corner et	Time					Visit to										Time								
	al 2004 UK	between first change in health status and onset of symptom that prompted patient to visit GP or other service Time between onset of symptom prompting patient to visit GP and date of visit to GP or other service					GP or other service and date of diagnosis										between first recalled change in health status and date of diagnosis								
12	ari et al 2007 UK		Urgent GP referral to date first seen in outpatient clinics was calculated by subtracting the date of urgent referral from the date first seen in chest outpatient clinics													Intervals for investigati ons such as bronchosc opy were calculated by subtracting the date of urgent GP referral from the date of investigati on				GP referral to date of first definitive treatment was calculated by subtracting the date of urgent GP referral from the date of commence ment of the first definitive treatment.					
13	Devbhand ari et al 2008 UK																						The intervals from outpatient to decision-to-treat	Decision-to- treat to treatment	
14	Dobson et al 2017 UK																								

#	Author, pub date and country	Symptom to doctor/ GP	GP to LCS/ Chest clinic/ referral/G P to first hospital appointm ent/ admission	Referral to first attendan ce to specialis t	Chest clinic to referral for Chest Physicia n	Chest Physician/ hospital appointment to Diagnosis	GP to diagnosi s	Diagno sis to referral to LCS/ or hospita	Sympto m to hospital admissi on	LCS to treatment	Hospitalizat ion to treatment referral	Diagnostic intervals (imaging/ biopsy)	Referral for treatmen t to initiation of treatmen t	Sympto m to 'referral for diagnosi s'	Sympto m to referral to LCS	Referral for diagnosis' to diagnosis	Sympto m to diagnosi s	Sympto m to referral (by GP or chest physicia n to next Mx)	Symptom to secondary care	Referral to treatment	GP to treatment	Diagnosis to initiation of treatment	Outpatie nt to decision to treat	Decision to treat/ specialist consultatio n to treatment	Symptom to initiation of treatment
15	Ellis & Vanderme er 2011 Canada	T1: time from initial symptoms to first presentati on to a family doctor or emergenc y departme nt	T3: time from initial presentation to the first appointme nt with a specialist, either directly to the JCC or to a respirologist or thoracic surgeon		T5. Time from JCC referral to initial consultati on	T4: time between the initial appointment with the specialist and the last date of additional diagnostic testing	T2: time from initial presentati on to the last date of diagnostic testing ordered by the family physician			T6: time from initial contact with a medical or radiation oncologist to the starting date of treatment, defined as chemothera py, radiation therapy, or the decision not to pursue treatment															T7: Overall time from onset of symptoms to commence ment of defiitive therapy was also calculated as a global delay
16	Emery et al 2013 Australia		Fist presentation in general practice to referral (GP interval)	From date of referral to fist attendan ce at specialist (specialis t access interval)		Time from fist attendance at the specialist to date of diagnosis (specialist interval)	The diagnostic interval is the time from fist presentati on until cancer diagnosis										Total diagnostic interval was defied as the time from fist symptom to diagnosis.								
17	Evans et al 2016 Australia															Referral to diagnosis				Referral to initial definitive managemen t		Diagnosis to initial definitive management			
18	Ezer et al 2017 Canada	time interval (in days) between first contact with a local physician for suspected lung cancer (T0)					time interval (in days) between first contact with a local physician to date of tissue diagnosis														Time interval (in days) between first contact with a local physician to date of first treatment				
19	Forrest et al 2014 UK		GP referral date to first hospital appointme nt date			First hospital appointment date to diagnosis date	GP referral date to diagnosis date														GP referral date to first treatment date	Diagnosis date to first treatment date			
20	Kanarek et al 2014 USA							Time from diagnosi s to first contact at SKCCC was defined as the referral interval.					Time from first contact at SKCCC to first surgery is defined as the treatment interval									Diagnosis to first surgery interval			

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21	Kim et al 2016 Canada											Diagnostic imaging interval: From Date of the chest X-ray which preceded the last computed tomography scan prior to the first diagnostic biopsy attempt to Date of the last computed tomography scan prior to the first diagnostic biopsy attemptDiagn ostic biopsy attemptDiagn ostic biopsy interval: From Date of the last computed tomography scan prior to the first diagnostic biopsy interval: biopsy interval: biopsy interval: biopsy attemptDiagn ostic biopsy interval: biopsy scan prior to the first diagnostic biopsy attempt to Date of the diagnostic biopsy procedure whichprovided pathological diagnosis										System interval: From Date of the chest X-ray which preceded the last computed tomography scan prior to the first diagnostic biopsy attempt to First day of treatmentTreat ment interval: From Date of diagnostic biopsy procedure which provided pathological diagnosis to First day of treatment			
22	Koyi et al 2001 Sweden	the patient's delay is the time from the first symptom(s) until the date he /she visits the doctor, in general the GP	GP delay, from the time a visit was arranged with the GP until the patient was referred to the specialist			specialist's delay (Second doctor's delay) is the time from when the lung specialist received the referral papers until the diagnosis was made.											Time symptom- diagnosis								Time symptom- treatment
23	Kudjawu et al 2016 France																								
24	Largey et al 2015 Australia																								
25	Largey et al 2016 Australia															Referral to- diagnosis				Referral-to- treatment		Diagnosis-to- treatment			
26	Lee et,al. 2002 UK																Onset of symptom s and their first chest radiograp h	Onset of symptom s and referral to a surgeon by a chest							
27	Li et al 2012 Canada																	physician				Time from diagnosis to first treatment			

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28	Maiga et al 2017 USA																					The interval between T2 and T3 is the diagnosistotreatment interval for patients with a tissue diagnosis before resection.			
29	Malalasek era et al 2018 Australia		Primary care interval				Diagnosti c interval													Secondary care interval		Treatment interval			
30				Referral by GP to first seen by specialist				1 week of a CXR request to first hospital visit													First visit to any treatment				
31	Neal et al 2015 UK	'Patient interval' (time from symptom onset to presentati on)					Date of request of first GP- initiated chest X- ray and date report received																		
32	Girolamo et,al. 2018 England			urgent referral for a suspicion of cancer from a general practition er (GP) to being seen by a specialist																			The decision taken to treat a patient to the start of the first treatment		
33	Gozalez et,al. 2014, Spain	from the first symptom to the first specialist consultati on (specialist delay)				from the first specialist consultation until confirmation of the diagnosis (diagnosis delay)														From the first specialist consultation until the start of treatment (hospital delay)	From the confirmation of the diagnosis up to the start of the first treatment (treatment delay)				

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3	Grunfeld et al 2009 Canada		admission	Date of referral to date of first diagnosti c consultati on											Date of referral to date of confirme d diagnosi s				Date of referral to date of initiation of first treatment (first tx was defined as neoadjuvan t chemother apy, surgery if no preoperative treatment was required, chemother apy, radiotherap y, or a decision for no tx						**Date the referral for diagnostic assessment was received by the consultant ('date of referral') to date patient informed of diagnosis ** Date of first diagnostic consultation to date patient informed of diagnosis **Date of referral to date of surgery or decision for no surgery **Date of confirmed diagnosis to date of surgery or decision for no surgery **Date of referral to date of surgery or decision for no surgery **Date of referral to date of surgery to date of first oncology consultation or decision for no consultation
3	Helsper et al. 2017 Netherlan ds		the time between the first cancer symptom related contact with the general practitioner (GP) and its correspond ing referral to secondary care (Primary care interval (ICP)				the time from the first presentati on to the GP to diagnosis (diagnosti c interval (ID)								The time from referral to histologi cal diagnosi s (refferal interval (IR)					The time from the first presentation to the GP to initial treatment (health care interval (IHC)	The time from diagnosis to initiation of the treatment (Treatmnet interval (IT)				
3	Hsieh et al 2012 Taiwan																								Delay in diagnosis' has been defined as the period from a patient's initial medical visit to any hospital to his/her confirmed diagnosis of lung cancer

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37	Hubert et al 2018 Canada																								**The first one was the interval between the moment that the green file was opened until all lung cancer staging and clinical tests were performed, and patient was referred for surgery after discussion with the respirologist .**The second interval was the time between the referral to the thoracic surgery department the consult with the surgeon ** The last interval was from the surgical
38	al 2012																								consult to the date of surgery
39	Spain  lachina et al 2017 Denmark																						Time from end of primary investigat ion to first dayof treatment = 14 days		
	Ju et al 2017 USA																								
41	Olsson et al 2009 USA			from referral to first respirator y specialist visit																GP referral to initial treatment		from diagnosis to treatment		specialist consultation to surgery	symptom onset to initial treatment
42	Ost et al 2013 USA																					Diagnosis to treatment			

#	Author, pub date and country	Symptom to doctor/ GP	GP to LCS/ Chest clinic/ referral/G P to first hospital appointm ent/	Referral to first attendan ce to specialis t	Chest clinic to referral for Chest Physicia n	Chest Physician/ hospital appointment to Diagnosis	GP to diagnosi s	Diagno sis to referral to LCS/ or hospita	Sympto m to hospital admissi on	LCS to treatment	Hospitalizat ion to treatment referral	Diagnostic intervals (imaging/ biopsy)	Referral for treatmen t to initiation of treatmen t	Sympto m to 'referral for diagnosi s'	Sympto m to referral to LCS	Referral for diagnosis' to diagnosis	Sympto m to diagnosi s	Sympto m to referral (by GP or chest physicia n to next Mx)	Symptom to secondary care	Referral to treatment	GP to treatment	Diagnosis to initiation of treatment	Outpatie nt to decision to treat	Decision to treat/ specialist consultatio n to treatment	Symptom to initiation of treatment
43	Özlü et al 2004 Turkey	From first symptom to presentati on	admission			admission and tissue diagnosis	From presentati on to tissue diagnosis														From presentatio n to first treatment	From diagnosis to treatment			From symptoms to treatment
44	Rankin et al 2017 Australia						The diagnostic interval is defined as "the time between first appointm ent with a health-care provider (HCP) and the formal cancer diagnosis being made."															The pretreatment interval is defined as "the time between formal cancer diagnosis and initiation of treatment"			
45	Rolke et al 2006 Norway	Patient delay: Time from first symptom to first personal contact with doctor	GP delay: Time from first contact with general practitioner (GP) to date on written referral.	Referral delay: Time from dated referral receipt to first contact with pulmonar y consulta nt.		Specialist delay: Time from first contact with pulmonary consultant to dated diagnostic histology/cyto logy																		Hospital delay: Time from first contact with pulmonary consultant to start of treatment.	Total delay: Time from first symptom to start of treatment.
46	Thapa et al 2014 Nepal	D1=Time from onset of symptoms to fist contact with a doctor (T1-T2) or patient delay						D 2=Time from fist contact with doctor to referral to MCVTC (T2-T3) or doctor delay																	
47	Verma et al 2018 Australia	T2: Time between fi rst symptoms to fi rst GP consultati on	T3: Time between GP and specialist consultatio n					coay		T4: Time between specialist consultation and commence ment of treatment.															T1: Time from first symptoms to commence ment of treatment.
48	Vidaver et al 2017 USA		Initial presentatio n-specialist referral	Specialis t referral- specialist consultati on			Initial presentati on-confirmed diagnosis			Specialist consultation -treatment											Initial presentatio n-treatment	Abnormal radiograph-treatment  Confirmed diagnosis-treatment		Treatment consultation-treatment	

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49	Wai et al 2012 Canada							Diagnos is to cancer centre referral Diagnos is to radiatio n oncolog y									First symptom to diagnosis							Radiation oncology consult to start of radiation treatment	
	Walter et al 2015 UK							consult									'time to diagnosis' , defined as the time from the first symptom/ sto the date of diagnosis								
51	Wilcock et al 2016 UK																							time from lung cancer MDT treatment recommenda tion to commencem ent of an 'active' oncological treatment	
52	Winget et al 2007 Canada																					diagnosis to first treatment in a cancer facility (that is, radiation or chemotherapy)		3) first consult with an oncologist to first treatment in a cancer facility.	
53	Yang et al 2015 China	Patient delay: First symptom to first contact with a local doctor	Delay in primary care: first contact with a local doctor to referral to hospital													Diagnostic delay in secondary healthcare: referral to hospital to diagnosis				Delay in secondary health care: referral to hospital to initiation of treatment	System delay: First contact with a local doctor to initiation of treatment	Treatment delay: Diagnosis to initiation of treatment			
54	Yilmaz et al 2009 Turkey	patient's application interval was defined as the time passed between the onset of symptoms and the first doctor visit.	The referral interval was defined as the time from the first doctor visit to admission to one of the pneumolog y departmen ts of our hospital for the further investigation																		Doctor's interval was defined as the time from the first doctor visit to thoracotom y	The treatment interval was the time passed from the diagnosis to thoracotomy			The total interval was the time between the onset of symptoms and thoracotomy
55	Yorio et al 2009 USA		п																			diagnosis to treatment.			

#	Author, pub date and country	Symptom to doctor/ GP	GP to LCS/ Chest clinic/ referral/G P to first hospital appointm ent/ admission	Referral to first attendan ce to specialis t	Chest clinic to referral for Chest Physicia n	Chest Physician/ hospital appointment to Diagnosis	GP to diagnosi s	Diagno sis to referral to LCS/ or hospita	Sympto m to hospital admissi on	LCS to treatment	Hospitalizat ion to treatment referral	Diagnostic intervals (imaging/ biopsy)	Referral for treatmen t to initiation of treatmen t	Sympto m to 'referral for diagnosi s'	Sympto m to referral to LCS	Referral for diagnosis' to diagnosis	Sympto m to diagnosi s	Sympto m to referral (by GP or chest physicia n to next Mx)	Symptom to secondary care	Referral to treatment	GP to treatment	Diagnosis to initiation of treatment	Outpatie nt to decision to treat	Decision to treat/ specialist consultatio n to treatment	Symptom to initiation of treatment
56	Zullig et al 2013 USA		dumsson					Days from diagnosi s to referral to palliativ e care or hospice														Days from diagnosis to initiation of treatment			
57	Sachdeva et al 2017 India																Delay in diagnosis from the onset of initial symptom s to histologic al confirmati								
58	Salomaa et al 2001 Finland		Patient's delay is the time from the first symptoms until the first visit to a doctor, who was in general, a GP	GP delay, which is the time from the date the patient visited the first doctor until the date the consultati on request for a specialist was written	The referral delay is the time between the writing of the referral and the first appointm ent with the specialist		The specialist's delay is the time from the first appointm ent until the diagnosis was made										on					The treatment delay is the time from the diagnosis until the treatment began			symptom-to- treatment delay
59	Sawicki et al 2013 Poland	Time from the first signs of the disease to the first medical examinati on																			the time from the first visit to a doctor to the start of treatment, or disqualifica tion from the causative treatment				
	Schultz et al 2009 USA	Time to treatment was the time from the first suspiciou s radiograp h to the date on which any treatment was first initiated ** In patients who refused treatment, we used the date of refusal as the endpoint for time to treatment																							
	Shugarma n et al 2009 USA Singh et al 2010 USA	first date recorded for treatment																							

#	Author, pub date and country	Symptom to doctor/ GP	GP to LCS/ Chest clinic/ referral/G P to first hospital appointm ent/ admission	Referral to first attendan ce to specialis t	Chest clinic to referral for Chest Physicia n	Chest Physician/ hospital appointment to Diagnosis	GP to diagnosi s	Diagno sis to referral to LCS/ or hospita	Sympto m to hospital admissi on	LCS to treatment	Hospitalizat ion to treatment referral	Diagnostic intervals (imaging/ biopsy)	Referral for treatmen t to initiation of treatmen t	Sympto m to 'referral for diagnosi s'	Sympto m to referral to LCS	Referral for diagnosis' to diagnosis	Sympto m to diagnosi s	Sympto m to referral (by GP or chest physicia n to next Mx)	Symptom to secondary care	Referral to treatment	GP to treatment	Diagnosis to initiation of treatment	Outpatie nt to decision to treat	Decision to treat/ specialist consultatio n to treatment	Symptom to initiation of treatment
63	Smith et al 2009 Scotland	The number of days from date of first symptom defined by the participant until date of presentati on of symptoms to a medical practitione r																							
64	Sood et al 2009 NZ	-																							
65	Stokstad et al 2017 Norway																								
66	Sulu et al 2011 Turkey		Patient's application interval was defined as the time elapsed from the onset of symptoms to the first doctor's visit		The referral interval was defined as the time from the first doctor's visit to admission to our hospital for the further investigati on.		The diagnosis interval was regarded as the time elapsed from admission to our hospital to the pathologic al diagnosis.														Doctor's interval was defined as the time elapsed the first doctor's visit to treatment	The treatment interval was the time elapsed from the diagnosis to treatment			The total interval was the time elapsed from the onset of symptoms to treatment
67	Chandra et al 2009 India																symptom- to- diagnosis delay, between the onset of symptom s to confirmed diagnosis					diagnosis-to- treatment delay, between diagnosis and treatment started			symptom-to- treatment delay, between onset of symptoms and treatment
68	Dubey et al 2015 India																The onset of symptom s to the confirmati on of diagnosis								

# Table 3: Other uncommon timepoint and intervals

#	Author, pub date and country	Other time point or Intervals		
1	Alexander et al 2016 Australia	NSCLC: Where systemic chemotherapy is the first anti-cancer treatment modality, in either definitive or palliative treatment settings, chemotherapy should commence within 3 weeks of the ready for care date (level III, grade C †). Adjuvant chemotherapy should commence as soon as the patient is medically fit following surgery and within 8 weeks of the date of surgery (level III, grade C †).  SCLC: Patients with severe or life-threatening symptoms should be regarded as a medical emergency and chemotherapy initiated immediately, within no longer than 48 h ‡ of the ready for care date – hospitalisation may be required (good practice point †). All other patients should commence chemotherapy within 2 weeks of the ready for care date (good practice point †)		
12	Devbhandari et al 2007 UK	GP referral to chest outpatient GP referral to decision to treat GP referral to treatment GP referral to treatment Oncology referral to chemotherapy Waiting on surgical waiting list Oncology referral to radiotherapy		
23	Kudjawu et al 2016 France	1) from bronchoscopy to: (a) first neo-adjuvant chemotherapy, (b) first combined neo-adjuvant radiotherapy (c) surgery, (d) first chemotherapy (in patients who underwent chemotherapy (in patients who underwent radiotherapy only), (f) first treatment (irrespective of treatment type); 2) from last neo-adjuvant chemotherapy to surgery; 3) from last combined neo-adjuvant radiotherapy chemotherapy surgery; 4) from surgery to: a) first chemotherapy, and b) first radiotherapy. 1- Patients with surgical pathway Time from bronchoscopy to surgery, Time from last neo-adjuvant chemotherapy, Time from bronchoscopy to surgery as first treatment, Time from bronchoscopy to first radiotherapy as only treatment, Time from bronchoscopy to surgery as first treatment, Time from bronchoscopy to surgery as only treatment, Time from bronchoscopy to surgery to first chemotherapy, Surgery followed by radiotherapy, Time from bronchoscopy to surgery, Time from surgery to first chemotherapy to surgery and chemotherapy, Time from bronchoscopy to surgery, Time from surgery to first chemotherapy to surgery to first chemotherapy to surgery to first chemotherapy to surgery to first chemotherapy, Time from bronchoscopy to surgery, Time from bronchoscopy to surgery to first chemotherapy to surgery to first chemotherapy to surgery to first chemotherapy. Time from bronchoscopy to surgery to first chemotherapy and radiotherapy. Time from bronchoscopy to surgery to first chemotherapy and radiotherapy Time from bronchoscopy to surgery to first chemotherapy. Time from bronchoscopy to surgery, Time from bronchoscopy to surgery to first radiotherapy.		
26	Lee et,al. 2002 UK	interval between referral by a respiratory physician and surgical out-patient attendance between referral by a respiratory physician and the surgical procedure time from surgical out-patient attendance to the surgical procedure		
27	Li et al 2012 Canada	Time from surgery to post-surgical treatment. Time from surgery to consultation with an oncologist.		
28	Maiga et al 2017 USA	Timepoints:Time zero (T0) is the date of lung nodule identification on computed tomography (CT) imaging according to the medical record; T1 is the date when a lung nodule originally less than 10 mm in size was documented as having new growth on CT imaging. T2 is the date of pathology diagnosis. T3 is time of resection and final pathology diagnosis.Intervals:Date of lung nodule identification on CT (T0) or date when a lung nodule originally less than 10 mm (T1) to time of resection and final pathology diagnosis (T3) is the time-totreatment interval.		
29	Malalasekera et al 2018 Australia	Doctor interval: First clinical presentation to First suspicious investigation System interval: First suspicious investigation to Treatment start		
38	Heredia et al 2012 Spain	**Interval in days between the 1st evaluation and staging **Interval in days between the first evaluation and the start of treatment **Interval in days between the referral date and staging **Interval in days between the staging date of the tumor and the start of treatment **Therapeutic delays in days since the first evaluation: Interval until surgical treatment, Interval until the start date of oncologic treatment, Interval until the start date of palliative treatment		
39	lachina et al 2017 Denmark	** Time from referral (time of diagnosis) to end of primary investigation = 28 days  **Time from referral (time of diagnosis) to first day of treatment = 42 days		
40	Ju et al 2017 USA	**End of primary investigation is defined as the date of decision on treatment. Referral is defined as the date where the investigating department receives the referral.  1. initial radiologic lesion detection by chest x-ray or CT scan (Step 1) tp diagnostic biopsy (Step 2),  2. diagnostic biopsy (Step 2) to radiologic staging (Step 3),  3. radiologic staging (Step 3) to invasive staging (Step 4),  4. invasive staging (Step 4) to surgery (Step 5).  5. initial radiologic lesion detection by chest x-ray or CT scan (Step 1) to radiologic staging (Step 3)  6. initial radiologic lesion detection by chest x-ray or CT scan (Step 1) to surgery (Step 4)  7. initial radiologic lesion detection by chest x-ray or CT scan (Step 1) to surgery (Step 5)		
41	Olsson et al 2009 USA	Waiting list for surgery Decision-to-treat to treatment other than surgery		
42	Ost et al 2013 USA	Suspicion to treatment		
45	Rolke et al 2006 Norway	Informed diagnostic delay: Time from decision of doing a diagnostic procedure to informing patient of diagnosis.		
46	Thapa et al 2014 Nepal	T1=Time since the onset of symptoms to assessment at hospital (MCVTC) T2=Time since fist contact with a doctor to assessment at Hospital T 3=Time since referral to MCVTC with suspicion of Lung Cancer		
48	Vidaver et al 2017 USA	First diagnostic test-last test		
49	Wai et al 2012 Canada	Driving times to the nearest cancer center at the time of diagnosis First symptom to first abnormal test First abnormal test to diagnosis		
51	Wilcock et al 2016 UK	From emergency admission to diagnosis From emergency admission to discussion at the lung cancer MDT		
52	Winget et al 2007 Canada	2) diagnosis to first consult with an oncologist		
54	Yilmaz et al 2009 Turkey	The diagnosis interval was regarded as the time passed between the admission to our hospital and the pathological diagnosis was made.		
55	Yorio et al 2009 USA	Survival time was defined as the interval between the date of treatment and the date of death or censoring. The intervals included in this analysis were image to diagnosis. Image to treatment		
56	Zullig et al 2013 USA	Days from diagnosis to death		
62	Singh et al 2010 USA	Two types of missed opportunities that could result in diagnostic delays: (1) type I missed opportunities, defined as episodes of care in which there was failure to recognize a predefined clinical clue (ie, no required action or work-up was initiated within 7 days of clue appearance); appropriate decisions to watch and wait were not considered missed opportunities; and (2) type II missed opportunities, defined as episodes of care in which there was failure to complete within 30 days a diagnostic procedure, consultation, or other requested follow-up action in response to a predefined clue.		
63	Smith et al 2009 Scotland	Two definitions of first symptom were used—participant-defined and health professional defined—using a checklist of symptoms compiled from CancerResearch UK lung cancer symptoms and SIGN guidelines.  **the number of days from date of earliest symptom checklist until date of presentation of symptoms to a medical practitioner		

# Author, pub date and country	Other time point or Intervals
Sood et al 2009 NZ	** postal delay (time taken to receive the referral at the outpatient clinic from the referrer)
	**grading delay (time taken to grade the referral)  **clinic delay (interval between date of receiving referral and to date of patient assessment)
	**interval from initial chest physician assessment to bronchoscopy
	**interval from initial respiratory assessment to CT chest
64	**interval from initial CT chest to CT-guided fine needle aspiration (CT FNA)  ** First respiratory assessment to final diagnosis
04	**Date referral received to diagnosis achieved
	**Date of GP referral to first respiratory assessment
	**First respiratory assessment to surgery
	**Date referred to surgeons to surgery  **Date of oncology referral to commencement of radiotherapy
	**Date of oncology referral to commencement of chemotherapy
Stokstad et al 2017	Timepoint:
Norway	Start of treatment as date of surgery, first fraction of radiotherapy, first day of intra-venous chemotherapy, or date of prescription of oral cancer therapy.
65	Time to start of treatment was defined as the number of calendar days from start time until start of treatment
	** time to treatment decision: start time to the date when such a decision was documented in the EMR

Table: Measures of timeliness with cutoff values from different guidelines

Interval	Cutoff value	Guidelines	Naming of interval
Onset of symptoms to first doctor visit <sup>28 51</sup>	30 days	BTS	Patient's Application interval <sup>28 51</sup>
First clinical presentation to first suspicious investigation <sup>35 80</sup>	28 days	DLCG	
First abnormal investigation	14 days	BTS	
(CXR) to confirmation of diagnosis/specialist visit <sup>41</sup>	56 days	RAND	<del>-</del>
GP to Specialist <sup>24</sup> <sup>28</sup> <sup>35-37</sup> <sup>42</sup> <sup>49</sup> <sup>51</sup> <sup>61</sup> <sup>69</sup> <sup>70</sup> <sup>84</sup>	1 day for urgent referrals, 10 days for	IOM	Referral delay <sup>49</sup> or
	standard referrals		Referral Interval <sup>28 51</sup>
	80% within 3-5 days	ACCP, DLCG, DAPPDT	_
	7 days	BTS, NICE, NNG	_
	14 days	UKNHS, Australian, UKDoH, SIGN, SMAC, CSCC, SLCG	
Primary care to initiation of	14 days	DLCG	System interval <sup>35</sup> or
treatment <sup>28 35 42 51 63 67 68 77</sup>	42 days	SLCG, CSCC	Doctor's interval <sup>28 51</sup>
	62 days	UKNHS, UKNCP, BTS, Joint Council for Clinical Radiology	_
	98 days	RAND	
	28 days for treatment decision, 35 days for systemic therapy 42 days for surgery or radiotherapy	Norwegian National Guidelines	_
Referral to secondary care to Diagnosis <sup>28</sup> <sup>36</sup> <sup>45</sup> <sup>51</sup> <sup>61</sup> <sup>84</sup>	28 days	UKDoH, CSCC, DLCG	Diagnosis Interval <sup>28 51</sup>
Diagnosis	14 days	BTS	_
First referral to secondary care	42 days	Australian	Secondary care interval 35
to treatment start 21 35 44 69-71 80	49 days	NOLCP	
	62 days	UKNHS, SEHD, NICE, BTS	
	42 days in ≥85% patients	DLCG	
First clinical presentation to	28 days	CSCC	_ Diagnostic interval <sup>35</sup>
Diagnosis <sup>35 84</sup>	60 days	RAND	
First investigation to treatment <sup>45</sup>	14 days	DLCG	
Diagnostic investigation to patient informed of diagnosis <sup>49</sup>	7 days	BTS	Informed diagnostic delay <sup>49</sup>
Diagnosis to Treatment start <sup>28 35</sup>	14 days	Australian, DLCG	Treatment interval 28 35
41 45-47 51 55 68 80 84 110	14 days in ≥80%	SLCG, DAPPDT	51 55 68
	patients, 35 days if mediastinoscopy		or Therapeutic delay <sup>47</sup>
	14 days until surgery	CSCC	_
	21 days	DLCG, DAPPDT	_
	28 days	NOLCP	_
	31 days 42 days for NSCLC/14 days for SCLC	UKNHS RAND	_
	42 days	DLCG, *Other study	_

Interval	Cutoff value	Guidelines	Naming of interval
First clinical presentation to	56 days for surgery	SMAC, UKDoH, SIGN,	Total interval <sup>35</sup>
treatment start <sup>24 34 35</sup>	52 days	Cutoff value proposed by authors	-
Decision to treatment to initiation	21 days	UKNHS	
of treatment 43 67 71 77	31 days (28 days for	UKNCP, BTS, Joint	
	surgery & radiotherapy,	Council for Clinical	
	7 days for chemotherapy)	Radiology	
Surgery to chemotherapy (Adjuvant chemotherapy) <sup>43</sup>	48 days	UKNHS	
Referral receipt to specialist consultation <sup>21 43</sup>	14 days	UKNHS, SEHD, NICE	_
Oncology referral to radiotherapy/ chemotherapy <sup>70</sup>	14 days	BTS, NICE	
Specialist consultation to surgery <sup>41</sup> 69 70 79	56 days	BTS, NICE	_
Surgeon consultation/Surgical	28 days	BTS, NICE	
waiting list to surgery 61 70 79	14 days	CSCC, *Other study	-
Onset of symptoms to treatment <sup>28 51</sup>	72 days	BTS, Canadian guidelines	Total interval <sup>28 51</sup>
Primary care referral to first diagnostic evaluation of symptom <sup>37</sup>	7 days	BTS	Type I missed opportunity (No evaluation or work-up was initiated within 7 days of appearance of a predefined clinical clue) <sup>37</sup>
Primary care referral to completion of evaluation at referral center <sup>37</sup> *Cutoff value adapted from other	30 days	BTS, *Other article	Type II missed opportunity (Failure to complete within 30 days a diagnostic procedure or consultation or the follow-up action requested in response to a predefined clue) <sup>37</sup>

\*Cutoff value adapted from other studies. IOM: Institute of Medicine, CSCC: Canadian Strategy for Cancer Control, NHMRC: National Health and Medical Research Council, ACCP: American College of Chest Physicians, BTS: British Thoracic Society, UKDoH: United Kingdom Department of Health, UKNHS: United Kingdom National Health Service, NICE: National Institute for Health and Care Excellence, UKNCP: United Kingdom National Cancer Plan, SLCG: Swedish Lung Cancer Group, RAND: Research and Development USA, NOLCP: National Optimal Lung Cancer Pathway, SEHD: Scottish Executive Health Department, DLCG: Danish Lung Cancer Group, SMAC: Standing Medical Advisory Committee, SIGN: Scottish Intercollegiate Guideline Network, CCA: Cancer Council Australia, DAPPDT: Dutch Association of Physicians for Pulmonary Disease and Tuberculosis, NNG: Norwegian National Guidelines.