



Community 365 Roundtable Meeting on Lung Cancer



Dr Matti Aapro

President
European Cancer Organisation

Meeting protocols

- To protect the quality of the audio for everybody **please stay on “Mute” throughout the meeting**
- **We encourage all participants to join the interactive discussion in the Chat box: ask questions, share thoughts and comments**
- **Please note that the meeting will be recorded**



Community 365 Roundtable Meeting on Lung Cancer



Agenda:

15:05-15:20 **Essential Requirements for Quality Cancer Care: Lung Cancer**

Yolande Lievens, Co-Chair of the Quality Cancer Network

15:20-15:30 **Open Discussion**

15:30-15:50 **Early Detection and Screening**

Co-Chair: Françoise Bartoli, VP, Head of Europe and Canada, Oncology Business, AstraZeneca

Presentation: Giorgio Scagliotti, Professor of Oncology, University of Turin and Chief of the Medical Oncology Division at the S. Luigi Hospital

15:50-16:00 **Open Discussion**

Community 365 Roundtable Meeting on Lung Cancer



16:00-16:20 **Molecular Diagnostics in Lung Cancer – Considerations and Relevance for Treatment Selection**

Co-Chair: Geoff Oxnard, Vice President, Global Medical Lead, Liquid Franchise at Foundation Medicine

Presentation: Matthew Krebs, Clinical Senior Lecturer in Experimental Cancer Medicine, University of Manchester and Consultant in Medical Oncology, The Christie NHS Foundation Trust, Manchester, UK

16:20-16:30 **Open Discussion**

16:30-16:50 **The Dutch Lung Cancer Audit: Nationwide Quality of Care Evaluation Using Quality Indicators**

Co-Chair: Ouzna Morsli, EMEAC Oncology Medical Lead at MSD

Presentation: Hans J.M. Smit, MD, PhD, Pulmonologist Rijnstate Hospital, Chairman of the Dutch Lung Cancer Audit, Arnhem, The Netherlands and **Rawa Ismail**, PharmD and PhD Candidate DICA

16:50-17:00 **Open Discussion**

Our Quality Cancer Care Network



Special Network
Impact of COVID-19
on Cancer



Prevention Network



Quality Cancer Care
Network



Survivorship and
Quality of Life
Network



Inequalities Network



Workforce Network



HPV Action Network



Health Systems and
Treatment
Optimisation
Network



Digital Health
Network





Our Quality Cancer Care Network



Community 365 Roundtable on Lung Cancer



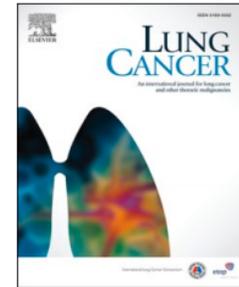
Lung Cancer 150 (2020) 221–239



Contents lists available at [ScienceDirect](#)

Lung Cancer

journal homepage: www.elsevier.com/locate/lungcan



Review

European Cancer Organisation Essential Requirements for Quality Cancer Care (ERQCC): Lung cancer



Thierry Berghmans^{a,1}, Yolande Lievens^{b,1}, Matti Aapro^c, Anne-Marie Baird^d, Marc Beishon^{e,*}, Fiorella Calabrese^f, Csaba Dégi^g, Roberto C. Delgado Bolton^h, Mina Gagaⁱ, József Lövey^j, Andrea Luciani^k, Philippe Pereira^l, Helmut Prosch^m, Marika Saarⁿ, Michael Shackcloth^o, Geertje Tabak-Houwaard^p, Alberto Costa^q, Philip Poortmans^r

europeancancer.org/resources



Community 365 Roundtable on Lung Cancer



Legacy from this meeting will include:

- Action report to be published in early January
- From tomorrow, video and slides on our website europeancancer.org/resources
- Follow up with EU Commission ahead of publication Europe's Beating Cancer Plan
- Next steps on implementation of Essential Requirements in our Quality Cancer Care Network



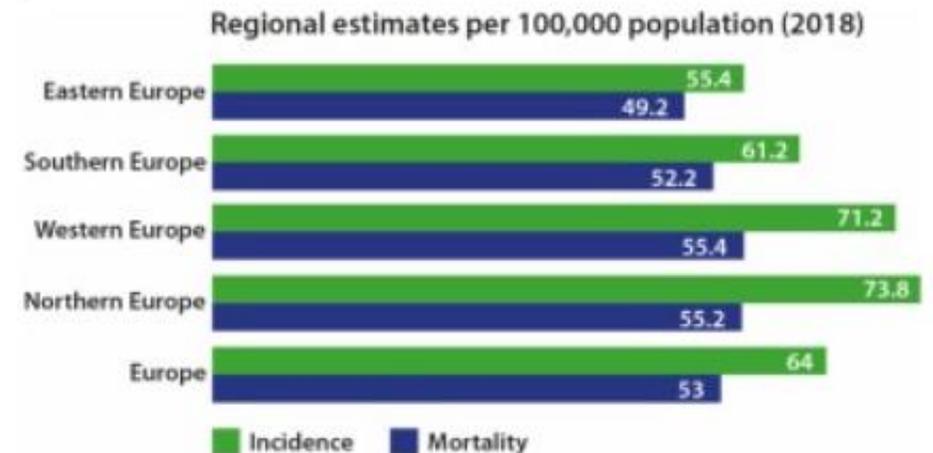
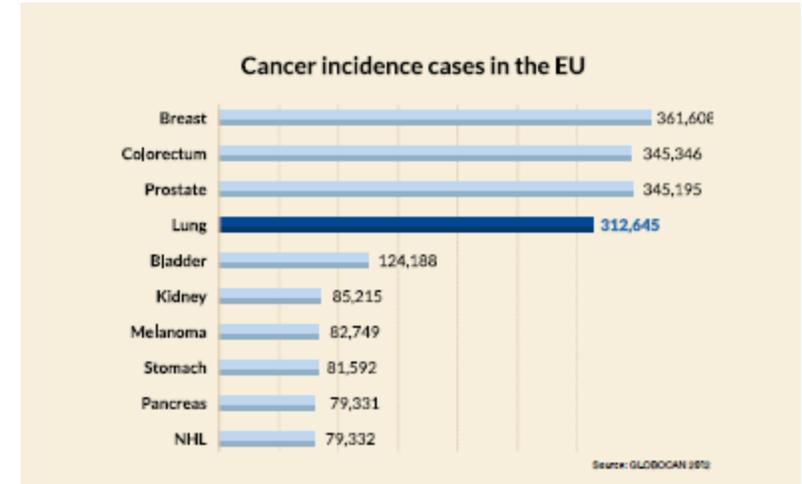
Essential Requirements for Quality Cancer Care: Lung Cancer

Yolande Lievens

Co-Chair of the Essential Requirements for Quality Cancer
Care: Lung Cancer

Lung cancer: So much more to do

- In 2018, the estimated **incidence** of lung cancer in EU countries was around 365,000 with mortality nearly 300,000.
- Lung cancer sadly remains a **poor prognosis** tumour with 5-year survival rates at a very low level. Men represent about two-thirds of mortality – nearly 200,000 were projected to die from lung cancer in 2018.
- **Opportunities for improvement in prevention, care and treatment remain insufficiently exploited.**
- Better **organisation** of care, and ensuring patient **access** to key members of the multidisciplinary and multi-professional **healthcare team**, can drive up both **quality of care and outcomes**.
- **The Essential Requirements for Quality Cancer Care: Lung Cancer** aims to help countries bridge that gap



Essential Requirements for Quality Cancer Care: Lung Cancer

The Lung Cancer manuscript has been produced by the [European Cancer Organisation](#) as part of the Essential Requirements for Quality Cancer Care (ERQCC) programme.

The European Cancer Organisation Essential Requirements for Quality Cancer Care (ERQCC) are written by [experts representing all disciplines involved](#) in cancer care in Europe.

They give patients, health professionals, managers and policymakers a [guide](#) to essential care throughout the [patient journey](#).





Some section highlights

A diagnosis and treatment summary

Recommendations on care pathways and timelines

Treatment centre recommendations

Patient involvement, access to information and transparency

Description and outline of the multidisciplinary team

Performance and quality recommendations

Recommendations on research and education

The image shows the cover of a review article from the European Cancer Organisation Essential Requirements for Quality Cancer Care (ERQCC). The title is "Lung Cancer" and it is from Volume 150, December 2020, pages 221-239. The article is a review by a multidisciplinary team of experts. The Elsevier logo is visible on the left, and a small graphic of a lung is on the right.

Understanding the particularities

The **challenges** section of the ERQCC paper helps audiences understand

- what makes lung cancer particular
- how organisational and health service response must adapt to this

such as

- The **high rate of diagnosis of lung cancer at advanced stages** is a major challenge for improving outcomes, and makes the option of lung cancer screening interesting.
- **Improved strategies at primary care level** are highlighted, such as lowering barriers to access - and to demanding - imaging, and equipping GPs with risk prediction tools.
- **Diagnosing and staging lung cancer is complex.**
It is essential that experienced specialists (*e.g. radiologists, pulmonologists, pathologists, nuclear medicine specialists*) determine results from imaging and pathological samples.
A successful management plan, especially for radical interventions, depends on their input to the MDT.

Understanding the particularities

The **challenges** section of the ERQCC paper helps audiences understand

- what makes lung cancer particular
- how organisational and health service response must adapt to this

such as

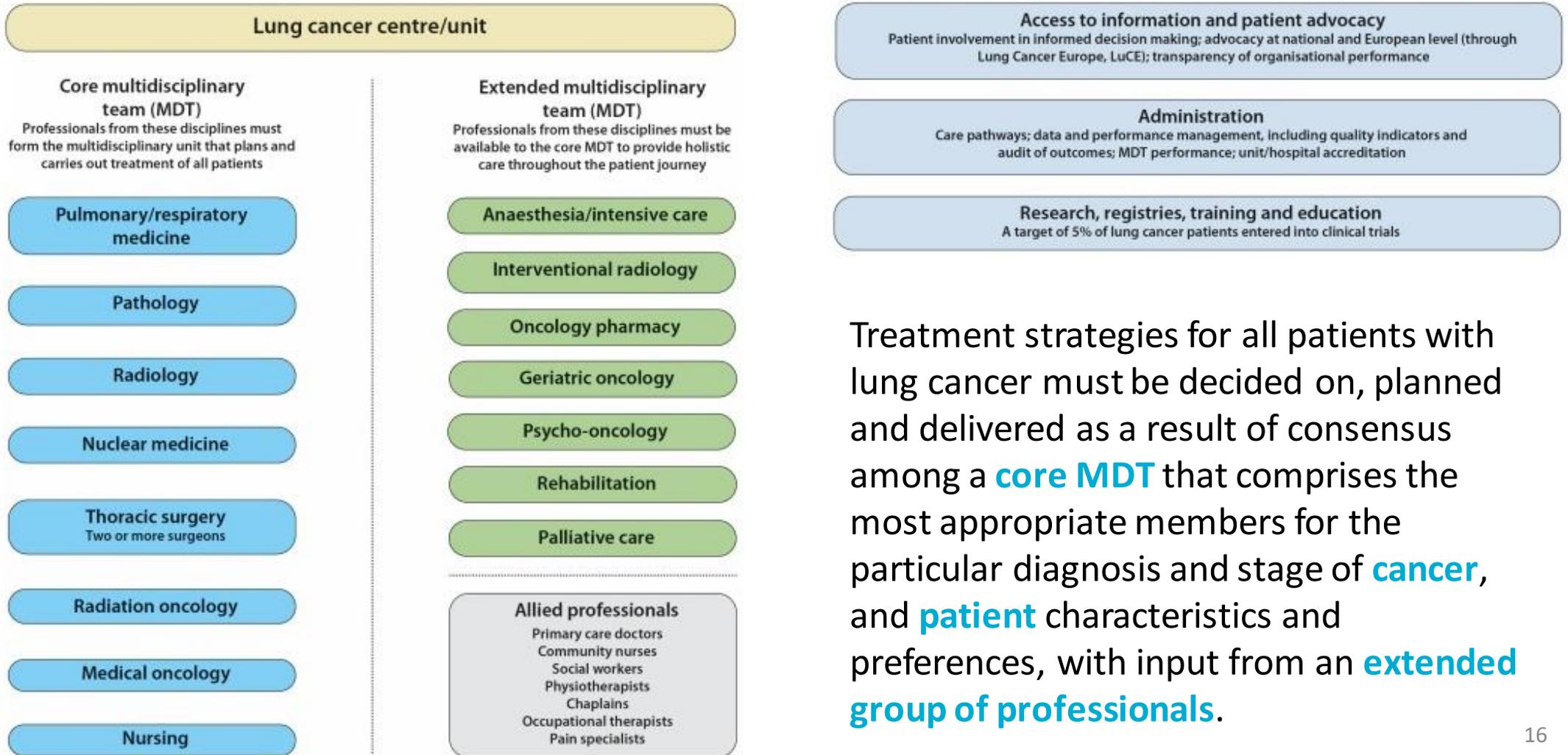
- **Optimal treatment of lung cancer is particularly challenging**, as patients may be older and have important comorbidities, requiring strong multidisciplinary knowledge, commitment and interaction to tailor the treatment to each individual patient.
- **Patients with lung cancer are a neglected population for psychosocial needs** compared with some other cancers, partly owing to the stigma of the disease as being self-inflicted through smoking, and they report increased distress as a result.

Pathway & Organisation: Getting it right!

The ERQCC: *lung cancer*

- provides advice on improving pathway management for lung cancer patients
- highlights good practice on cancer centre organisation
- **After a diagnosis, the patient must know** which professional is responsible for each step in the **treatment pathway** and who is following the patient during the journey (usually called a case manager or patient navigator)
- **Follow-up, support and care for long-term survivorship, as well as palliative care,** must be part of a care pathway.
- **There is direction towards care and treatment for lung cancer performed in higher volume centres.**
European countries taking notable steps in this direction:
 - **Denmark** - now carries out surgery in just 4 centres, and also has fewer locations where lung cancer is diagnosed and evaluated, reduced to 13 sites from about 50 previously
 - **Germany** - the target for a certified lung cancer centre is 200 cases a year (all new presentations)

Defining the core and extended MDT



Treatment strategies for all patients with lung cancer must be decided on, planned and delivered as a result of consensus among a **core MDT** that comprises the most appropriate members for the particular diagnosis and stage of **cancer**, and **patient** characteristics and preferences, with input from an **extended group of professionals**.

Requirements for professions: Examples

Essential requirements: pulmonology

- Pulmonologists must take part in multidisciplinary treatment of lung cancer.
- Pulmonologists must be able to perform bronchoscopy.

Essential requirements: thoracic surgery

- Lung cancer surgery must be carried out by multidisciplinary teams of appropriately trained surgeons.
- There must be at least 2 experienced thoracic surgeons spending a significant amount of their time to ensure sufficient volume of patients to ensure high quality of care.
- Perioperative care for patients undergoing thoracic surgery must be provided by specialist teams (including anaesthetists, theatre and on the wards) and anaesthesia must be provided in an intensive care and high dependency ward also attended by physiotherapists.
- Patients with early stage lung cancer must be offered minimally invasive surgery where appropriate.
- Outcomes of patients undergoing surgery must be audited [117].

Essential requirements: radiation oncology

- Radiation oncology departments treating lung cancer must have access to up-to-date radiotherapy technology and techniques such as IMRT and SBRT, ideally on-site or at a centre through a formal collaborative agreement that includes a common MDT.
- Radiation oncologists must know the indications of radiotherapy for lung cancer, and the place, expected efficacy and potential side-effects of thoracic radiotherapy in multidisciplinary treatment regimens. They must have a special interest and expertise in the multidisciplinary treatment of lung cancer and of other thoracic malignancies to select the optimal treatment for each patient, considering the specific oncologic situation and comorbidities.
- Multimodal imaging including a CT in treatment position and/or a PET/CT scan are mandatory to define the target volume, along with pathological information obtained through mediastinal staging – either EUS-EBUS or mediastinoscopy – in the case of locally advanced disease.
- Radiation oncologists treating lung cancer must have a team of radiation therapists, dosimetrists and medical physicists with expertise in lung cancer and thoracic malignancies.
- Radiation oncologists must be aware of ongoing clinical trials and their methodology performed at their centre or in associated centres.
- The radiation oncology centre must have regularly updated protocols for radiotherapy and concurrent chemoradiotherapy for lung cancer based on international guidelines.
- Image guidance, motion management and adaptive radiotherapy policies and quality assurance guidelines must be clearly described and documented. External quality assurance audits are highly recommended.
- Radiation oncologists must follow up patients to act on early or late toxicity, and in case of relapse.

Essential requirements: nuclear medicine

with management guidelines of pulmonary nodules. Knowledge of the peculiar pattern of lymphatic and haemangio-lymphatic spread of lung cancer (including uncommon sites of metastases) is essential. Found knowledge of the TNM lung cancer staging system and its pitfalls [100,101]. Knowledge of the strengths and limitations of PET/CT and PET/MR with image guided biopsies and minimally invasive procedures (i.e. radiofrequency ablation, cryoablation) and their use with treatment responses to radiotherapy and immunotherapy, and their use in the context of clinical trials. Knowledge of surgical procedures to assist surgery. Knowledge of the use of PET/CT and PET/MR must be available. Knowledge of when to refer a patient to nuclear medicine

Requirements for professions: Examples

Essential requirements: nursing

- Nurses must conduct holistic, personalised and age-appropriate care, ensuring the efficacy throughout the patient's journey and the nature of shared decision-making.
- Nurses must provide support and family-centred care at the point of care.
- Nurses must monitor the patient's trajectory, including psychosocial, nutritional and functional status, and must be updated on advanced

Essential requirements: palliative care

- All older patients should be screened using a validated tool.
- Frail and disabled patients should be identified [147]. The assessment should be an objective assessment of functional status, collaboration with geriatric medicine).
- Cognitive impairment, consent, complex screening using a validated tool, psychiatrist or psychologist, and impaired patient should be present in the team, working with the patient's goals of care.

Essential requirements: palliative care

- The MDT must offer optimal supportive and palliative care at the earliest opportunity.
- There must be access to a dedicated palliative care unit with a specialist team that provides expert outpatient and inpatient care and good knowledge of cancer disease and cancer treatments.
- The palliative care team must include palliative care physicians and specialist nurses, working with an extended team of social workers, psychotherapists, physiotherapists, occupational therapists, dietitians, pain specialists and psycho-oncologists.
- The palliative care team must have experience of taking care of frail older patients and their families.
- To ensure the continuity of care at home, the palliative care team must work with community/primary care providers.
- Palliative care specialists and oncologists must aspire to meet the standards of ESMO Designated Centres of Integrated Oncology and Palliative Care (<http://www.esmo.org/Patients/Designated-Centres-of-Integrated-Oncology-and-Palliative-Care>).

Essential requirements: interventional radiology

Interventions must be performed by an experienced interventional radiologist with access to appropriate interventional CT, MRI, and endoscopy (fluoroscopy and 3D-CT imaging are recommended). Radiologists must be available to the MDT to discuss the use of local ablative techniques for treating lesions not amenable to, or combined with, surgery.

Essential requirements: psycho-oncology/psychosocial care

Psychological assessment by the healthcare team should be provided at all stages of the disease and its impact on the patient and their partners and families. A validated psychological assessment tool (such as a distress thermometer) should be administered at a certain level must be routinely managed; above that level there must be further screening for anxiety and depression, and referral to an appropriate professional, such as a mental health professional.

Guidelines must be based on clinical practice guidelines, such as the ESMO Guidelines for Distress Management (http://www.esmo.org/clinical-advances-and-practice-guidelines/physician_gls/default.aspx).

Measuring what we do

A lung cancer centre must develop:

- **Performance measurement metrics/quality indicators** based on the essential requirements in this paper and on clinical guidelines, in alignment with national requirements and legislation
- **Operational policies** to ensure the full benefits of a coordinated clinical pathway based on published guidelines
- **Accountability** within the governance processes in individual institutions
- **Systems** to ensure safe and high-quality patient care and experience throughout the clinical pathway
- **Effective data management** and **reporting systems**
- **Engagement with patients, their carers and support groups** to ensure reporting of patient outcomes and experience.

Some final words on improving lung care

Lung cancer

- includes **variable disease** entities and patient groups;
- requires a large range of **knowledge and expertise** over the entire care pathway;
- from professionals that work in a **well-structured and organised** manner.

Educational and awareness programs should support that available **research evidence** is accessible, translating into **optimal outcome**.

Initiatives such as **ERQCC: lung cancer should not remain a static endeavour** but should continuously integrate **optimised** treatment and health system approaches and be **informed** by data collected in clinical trials as in real-life practice

Thank you to the paper contributors!





Open Discussion

Please use the Chat feature to ask questions and make comments



Redefining the future for lung cancer patients in Europe

Co-Chair:

Françoise Bartoli, VP, Head of Europe and Canada,
Oncology Business, AstraZeneca



Accelerating advances for lung cancer patients through collaboration

Giorgio Scagliotti, Professor of Oncology, University of Turin and Chief of the Medical Oncology Division, S. Luigi Hospital

Eight pillars of the current IASLC strategic plan

Education

Research

Membership

Int'l Development

Governance

Policy

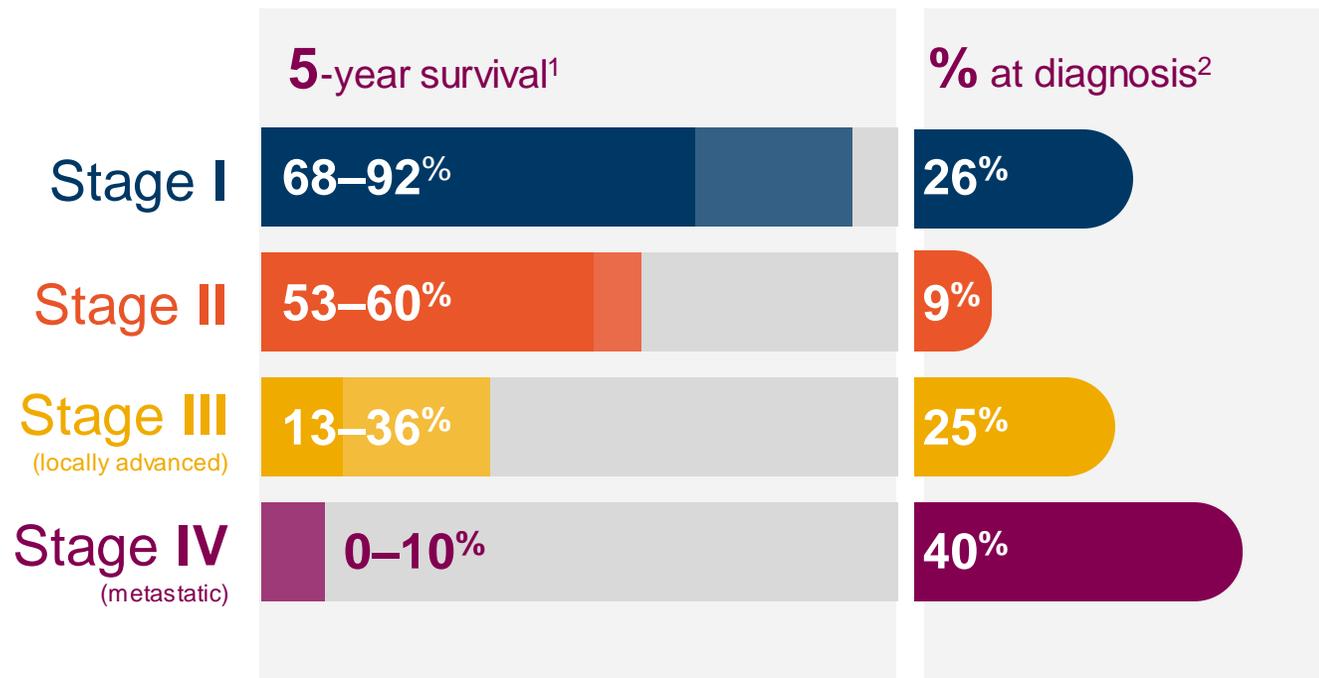
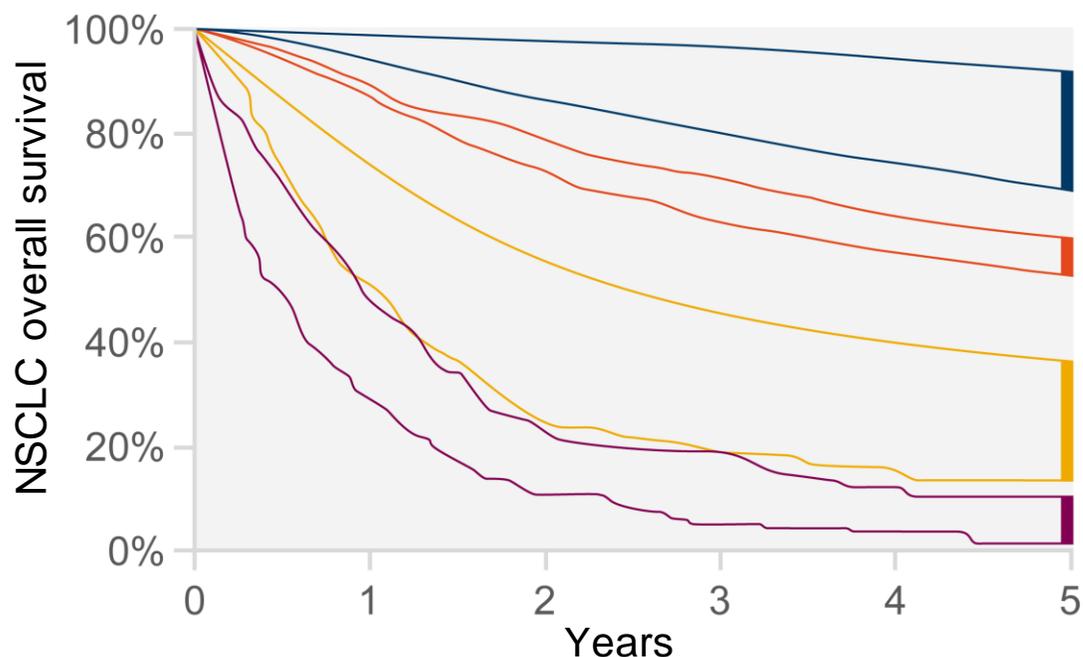
Funding

Strategic Partnerships



The Challenge

Globally lung cancer is the leading cause of cancer death with 1.8 million lives lost annually



¹Goldstraw *et al. J Thorac Oncol.* 2016; 11(1):39–51; ²EpiCast Report: NSCLC Epidemiology Forecast to 2025. GlobalData. 2016.

Staging: Improving Lung Cancer Staging through International Collaboration

- For more than 20 years, the IASLC has provided valuable recommendations for the TNM classification of lung cancer
- IASLC is currently gathering data to inform the 9th edition of the TNM Staging System, which for the first time, will include new data elements such as gene mutations, fusions, and copy number alterations and protein expression level.
- The goal of these additional tumor characteristics is to significantly enhance the accuracy of the Staging System, leading to more precise treatment regimens and increased patient survival.
- As of November 2020, the following number of cases have been collected from over 20 countries: Lung Cancer: 54,789, Mesothelioma: 436, Thymic malignancies: 7,183

We are at a critical moment in the fight against lung cancer

In 2018 alone,
1.8 million people
died from lung
cancer¹



Every 18 seconds,
a life is lost to
lung cancer¹



40% are diagnosed
after the disease
has spread
beyond the
lung, worsening
prognosis²



Only 1 in 5 patients
are alive 5 years
after diagnosis³



The Time For Change Is Now

Diverse perspectives enable our four founding partners to identify needs more precisely, while our depth of resources help us to amplify impact as we work to **eliminate lung cancer as a cause of death**

Founding Partners



Project partners :Merck - BMS - Genentech - Lilly - Novartis

Lung Ambition - The Mission

Vision

What does the coalition aim to achieve?

As global partners with a common and enduring commitment, the coalition aims to one day **eliminate lung cancer as a cause of death**

Mission

Why should the coalition exist?

As a coalition, we can **accelerate progress** by amplifying the multi-disciplinary expertise of our partners

Goals

What do we want to do?

Together, we can shape the environment to improve outcomes for patients with lung cancer. As a first goal we will use the evidence, advance the science and motivate the community to **double 5-year survival in lung cancer by 2025**

Strategic Focus

How will we achieve it?

We will deliver on the mission through 3 key areas of focus:

- A commitment to **early screening & diagnosis**
- A promise to deliver **precision, curative treatments**
- A passion for ensuring **quality care** for patients

Focusing on Today and Future Potential



Screen & Diagnose Early



Deliver Innovative Medicine



Enhance the Quality of Care

- 1 Screening Access & Policy
- 2 Screening Rates
- 3 New Technology (ctDNA)

- 1 Early-Stage Disease
- 2 Resistance Mechanisms
- 3 Clinical Endpoints

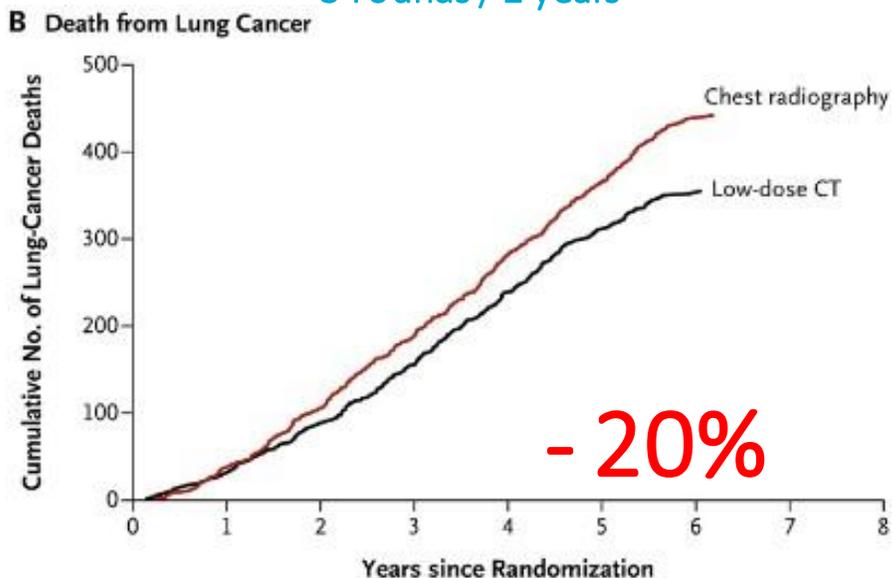
- 1 Long-term Survivorship
- 2 Access
- 3 Standards of Optimal Care



LDCT screening trials results: lung cancer mortality can be reduced

large size trial (> 50,000)
diameter > 3 mm no
risk modulation

3 rounds / 2 years

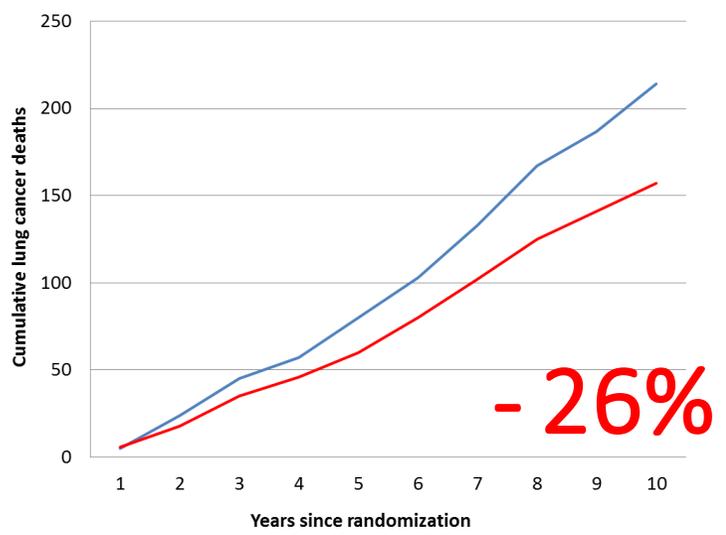


Aberle DR, N Engl J Med. 2011; 365:395

NLST

medium size trial (> 15,000)
volume > 50 mm³+ VDT no risk
modulation

4 rounds / 6.5 years

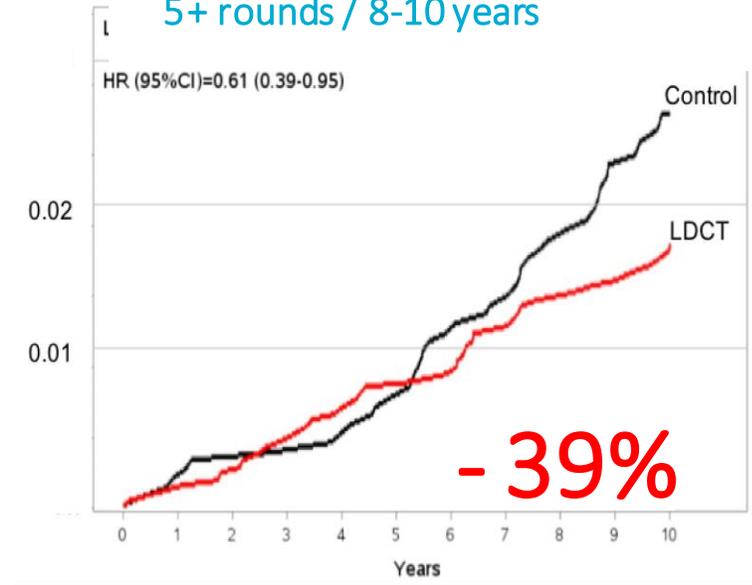


deKoning H, WCLC 2018; Toronto

NELSON

small size trial (> 4,000)
volume > 60 mm³+ VDT +
PET LDCT risk modulation

5+ rounds / 8-10 years



Pastorino U, Ann Oncol 2019 doi: 10.1093

MILD

Screening of Lung Cancer: key issues

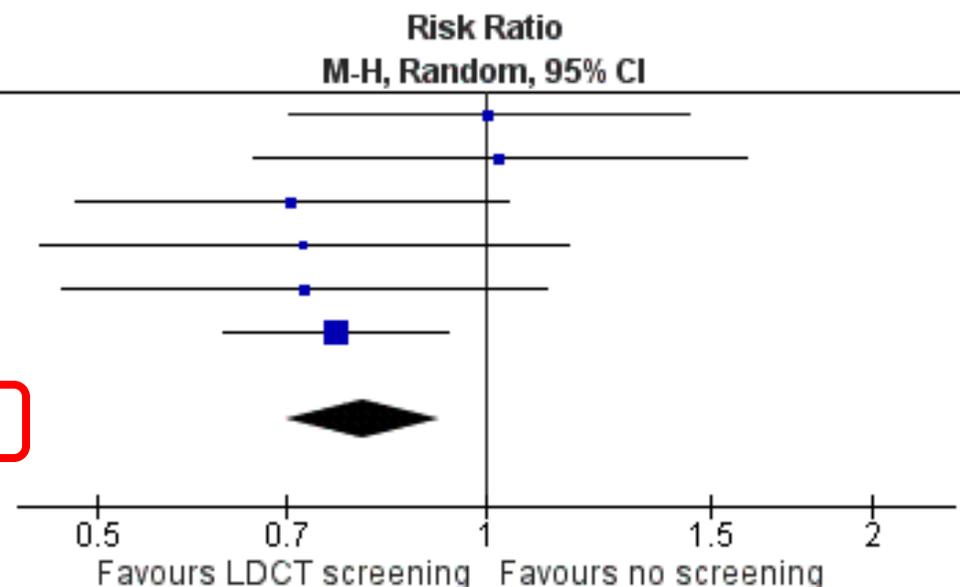
- Evidence from RCTs
- LDCT technology and risk
- Overtreatment & surveillance
- Benefits beyond LC detection
- Liquid biopsy & biomarkers
- Boosting prevention with LDCT
- Personalized strategy
- Implementation challenges
- Screening in the covid-19 time



Benefits and Harms of Lung Cancer Screening by Low-Dose Computed Tomography: a systematic review and meta-analysis

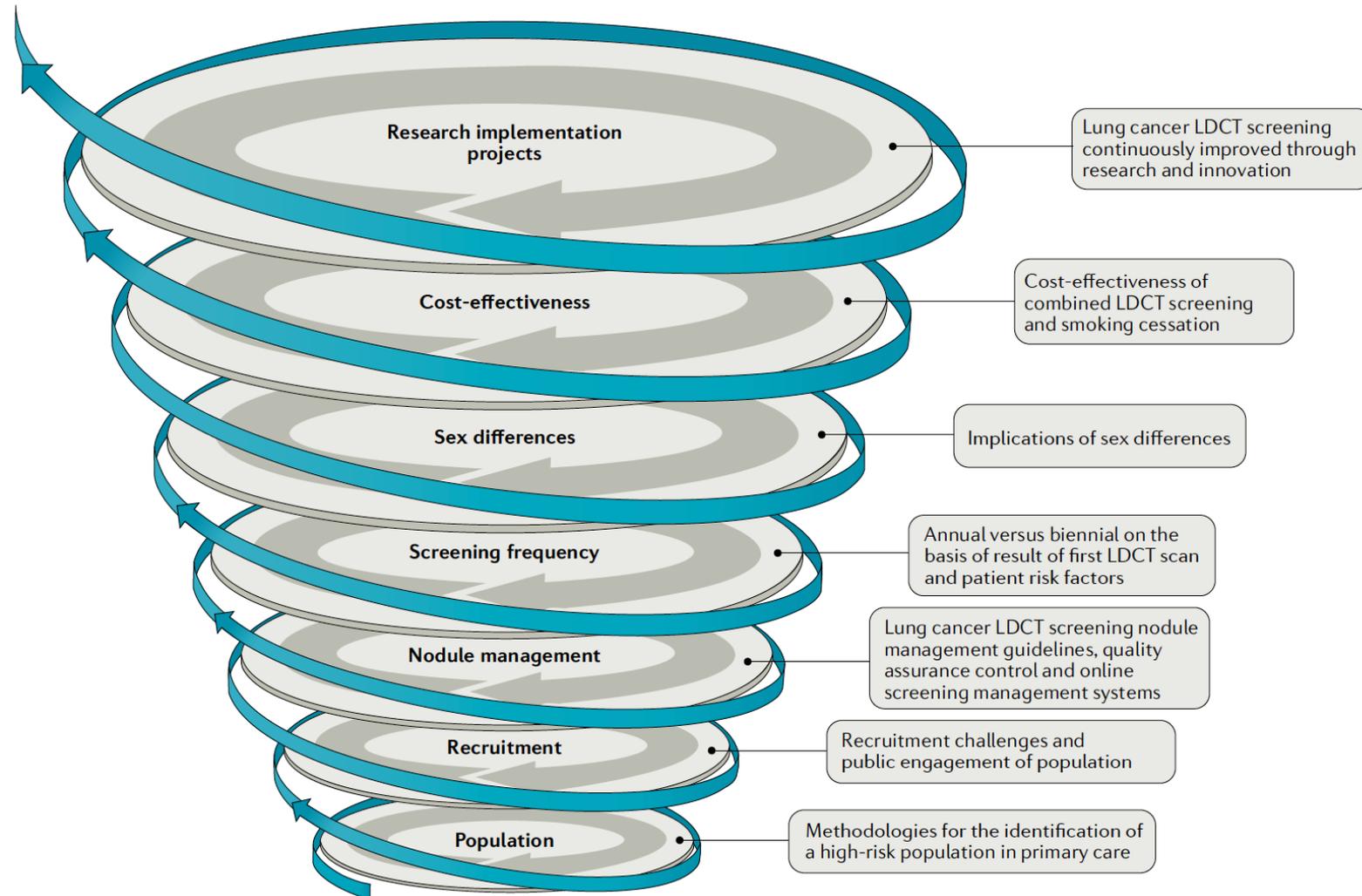
Study or Subgroup	LDCT screening		no screening		Weight	Risk Ratio	
	Events	Total	Events	Total		M-H, Random, 95% CI	
DANTE	59	1264	55	1186	14.5%	1.01	[0.70, 1.44]
DLCST	39	2052	38	2052	9.5%	1.03	[0.66, 1.60]
ITALUNG	43	1613	60	1593	12.5%	0.71	[0.48, 1.04]
LUSI	29	2029	40	2023	8.3%	0.72	[0.45, 1.16]
MILD	40	2376	40	1723	9.9%	0.73	[0.47, 1.12]
NELSON	160	6583	210	6612	45.3%	0.77	[0.62, 0.94]
Total (95% CI)		15917		15189	100.0%	0.80	[0.70, 0.92]

Total events 370 443
 Heterogeneity: $\tau^2 = 0.00$; $\chi^2 = 3.73$, $df = 5$ ($P = 0.59$); $I^2 = 0\%$
 Test for overall effect: $Z = 3.16$ ($P = 0.002$)



SPIRAL

Screening planning and implementation

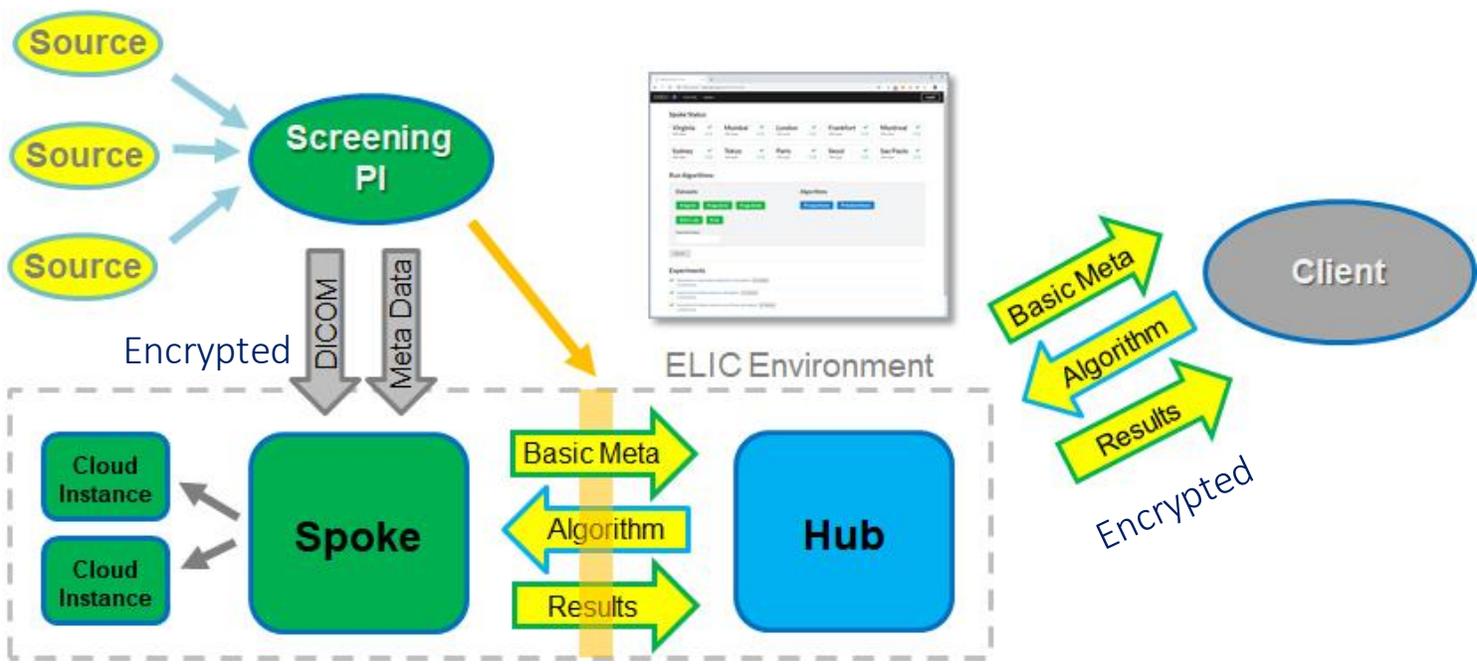




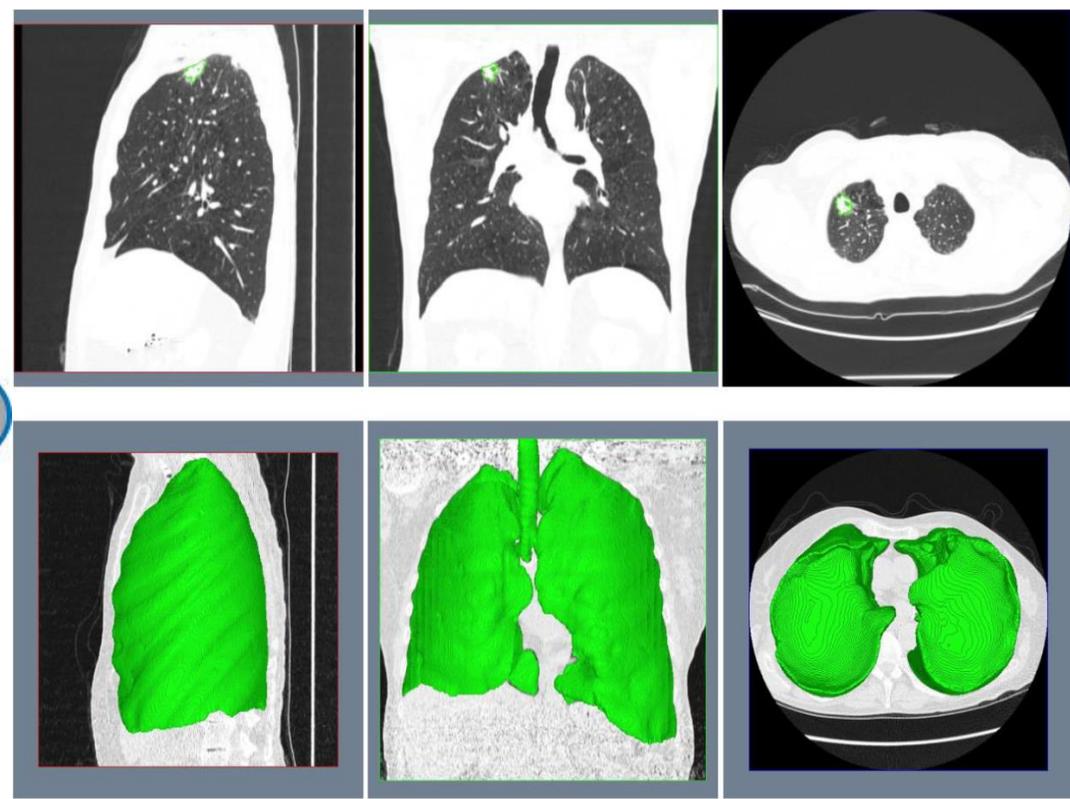
Early Lung Imaging Confederation (ELIC)

Lung cancer imaging database and computational analysis environment designed to enable the study of extremely large collections of quality-controlled internationally assembled CT chest images and associated biomedical data.

ELIC Hub and Spoke Model



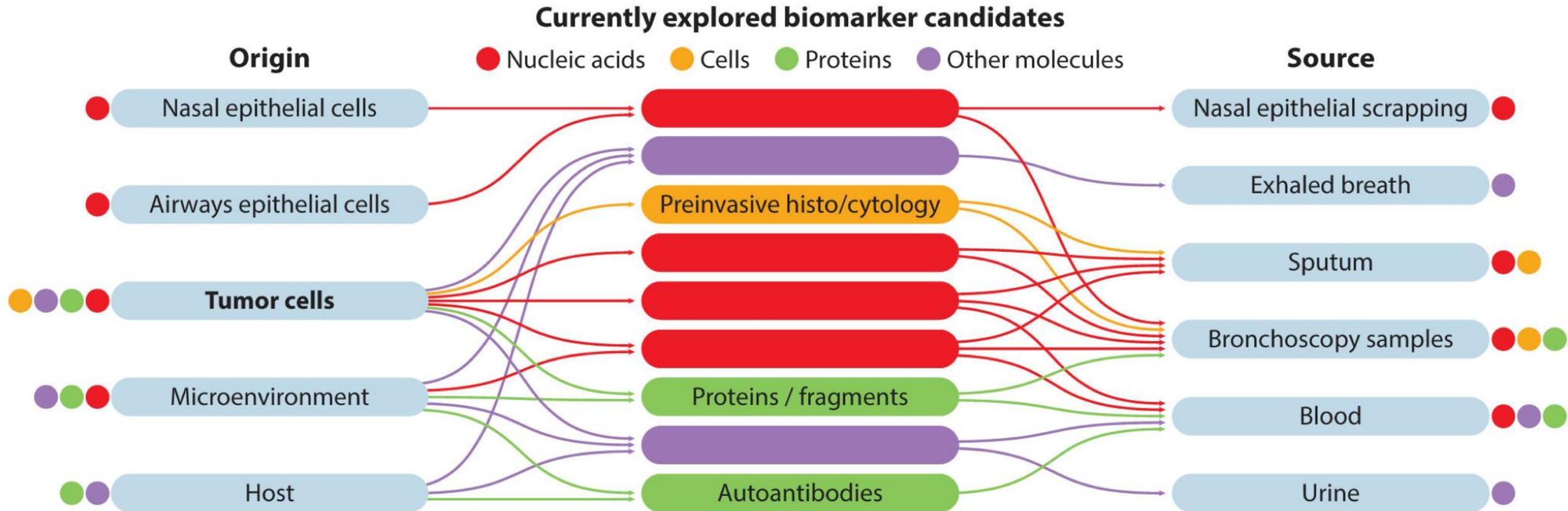
Nodule and Lung Volumes



ELIC Moving Forward

- ✓ **'Overall uptake of low-dose CT-screening needs to increase, if we are going to make a further dent in the lung-cancer mortality'**
 - Awareness Campaigns
 - Educational Efforts
 - Videos/Webinars
 - Podcasts with Multidisciplinary Participation
 - Workshops
- ✓ **Stratification of the indeterminate nodule on the screening scans**
- ✓ **Detection of squamous-cell and small-cell lung cancer earlier will be of importance**
- ✓ **Application of Artificial Intelligence, Machine and Deep learning will pick up relevant images for further invasive procedures and non-invasive bio-marker studies**
 - Precise volumetric segmentation of lung nodules automatically -- both detected and user found (Deep Learning will reduce variability)
 - Benign vs. Malignant

Biomarkers in lung cancer screening: achievements, promises and challenges

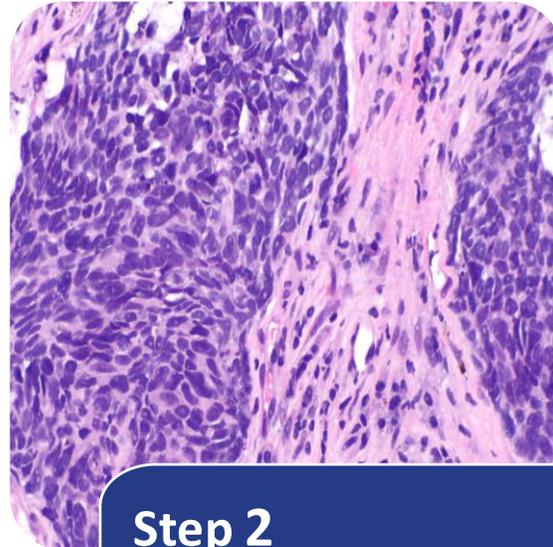


Major Pathologic Response Project



Step 1

MPR multidisciplinary recommendations paper for standardized tissue processing and pathologic assessment



Step 2

Conduct interobserver reproducibility study based on recommendations for assessment



Step 3

Collect data to compare pathologic response and other biomarkers to survival – ultimate goal is surrogate marker for outcome

ILC2 :Initiatives in Lung Cancer Care

ILC² invites local PAGs around the world to develop and submit projects with a potential to transform care and improve survival of lung cancer pts.

Awardees by Region



60 applications received

30 applications were reviewed

14 were selected by the committee for a total of \$1,035,812 to be distributed during this application cycle

Round 2 is Now Open!

To complete your application please click here [↗](#)

ILC² grant applications are reviewed and selected by the four founding partners of the Lung Ambition Alliance: the Global Lung Cancer Coalition, Guardant Health, the International Association for the Study of Lung Cancer and AstraZeneca. **Funding for the ILC² grant is provided by AstraZeneca.**

All applications must be completed by 31 December 2020.

Organisation	Project		
Hope Foundation for Cancer Care	Patient Centered Quality Care for Lung Cancer: Research Based De	United States	\$ 100,000
Swedish Medical Center Foundation	Native American CommunityLung Cancer Screening Road Map	United Kingdom	\$ 98,918
LUNGeivity Foundation	Model for improved patient participation in shared decision- making t biomarker testing report results	KENYA	\$ 95,335
Lung Health Foundation Canada	Bridging the Gaps in Lung Cancer Care	Egypt	\$ 93,728
GO2 Foundation for Lung Cancer	Lung Cancer Advocate Media Training & Network	Australia	\$ 90,831
Roy Castle Lung Cancer Foundation	L-Can Connect: lung cancer advocacy network	Canada	\$ 58,000
KENYA HOSPICES AND PALLIATIVE CARE ASSOCIATION	Collective Hope and Quality Life for Lung Cancer Patients	United States	\$ 25,000
Mersal Civil Society Foundation for charities and catering	LUNG Cancer Advanced patient Support (LUCAS) Program	Rwanda	\$ 25,000
Lung Foundation Australia	Australian Lung Cancer Nurse Telehealth service	Costa Rica	\$ 25,000
Lung Cancer Canada	Airways of Hope	Argentina	\$ 24,000
Dusty Joy Foundation	New Lung Cancer Support Group Chapters		
RWANDA PALLIATIVE CARE AND HOSPICE ORGANIZATION	Support and advocacy for lung cancer patients in Rwanda.		
Asociación Tour Rosa de Costa Rica	Lung Cancer in Costa Rica: Current Situation and Challenges for a Person-Centered Approach		
Fundacion Pacientes de Cancer de Pulmón	Learning with the patient		
		Grand Total	\$ 1,035,812

LAA: Other Funded Projects

COVID-19 Research Grants

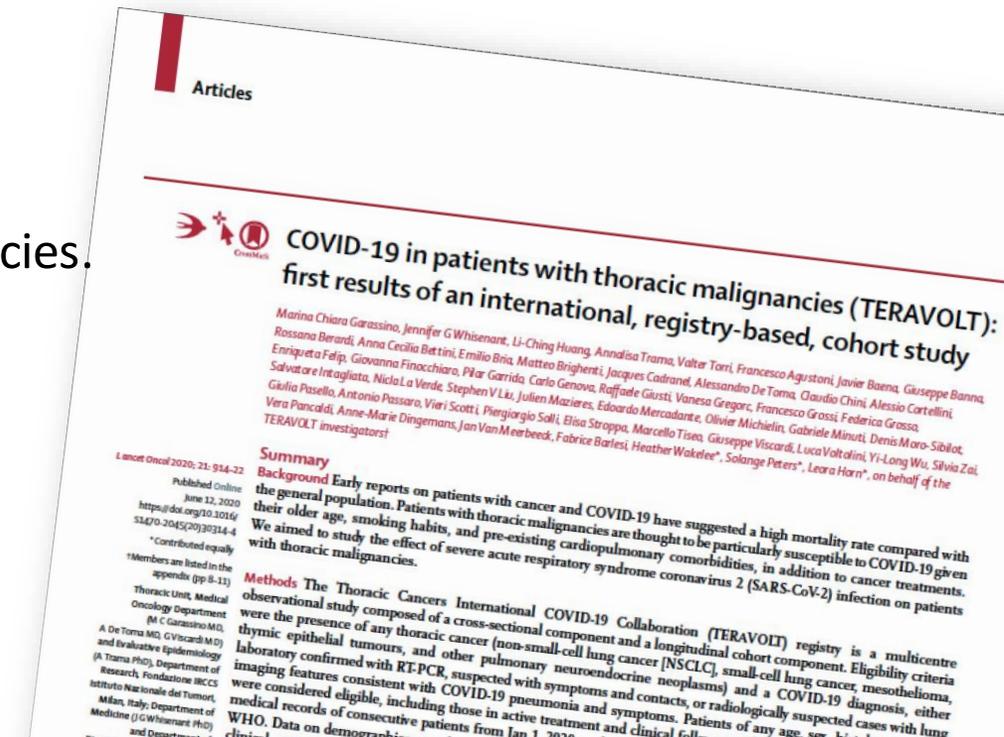
Junior Faculty Research Grants for the Study of Lung Cancer and COVID-19

3 Grants for \$50,000 each

Goal: Obtain preliminary data to enable successful competition for national peer-reviewed grants

With this effort Novartis and Eli Lilly became Project Partners of LAA

- TERAVOLT Project
- The IASLC, through the LAA, funded TERAVOLT.
- Goal: Understanding the impact of COVID-19 on thoracic malignancies.
 - Risk factors associated with morbidity and mortality
 - Therapies that may impact survival
 - Guidance on the management of patients



Developing Partnerships through Data Sharing with Academic Institutions, Societies & Industry

- Enhance and improve DATA SHARING
- Recognizing Patterns in Large Volume of Data
- Identifying Characteristics that cannot be perceived by the human brain, e.g., mutations with non-invasive techniques
- Ultimately improve the survival of lung cancer patients

The
LungAmbition
Alliance



Accelerating advances
for people with lung cancer.



Thank you!



Giorgio V. Scagliotti
giorgio.scagliotti@unito.it



Open Discussion

Please use the Chat feature to ask questions and make comments



Final Remarks

Co-Chair:

Françoise Bartoli, VP, Head of Europe and Canada,
Oncology Business, AstraZeneca



Molecular Diagnostics in Lung Cancer – Considerations and Relevance for Treatment Selection

Co-Chair:

Geoff Oxnard, Vice President, Global Medical Lead,
Liquid Franchise at Foundation Medicine

Precision cancer care & precision cancer diagnostics

EMPIRIC CANCER CARE

- Treatment based on tumor location verses genomic drivers of disease
- Potential for more toxicity with less reliable treatment outcomes

PRECISION CANCER CARE

- Finding the right treatment for the right patient
- Potential for less toxicity with more reliable treatment outcomes

Precision cancer care & precision cancer diagnostics

EMPIRIC CANCER CARE

- Treatment based on tumor location verses genomic drivers of disease
- Potential for more toxicity with less reliable treatment outcomes

Precision cancer diagnostics

Targeted therapy biomarkers

Kinase activation
(EGFR, ALK, NTRK, etc.)

HRD biomarkers
(BRCA1/2, ATM, etc.)

Immunotherapy biomarkers

PDL-1 IHC

Microsatellite instability

Tumor mutational burden

PRECISION CANCER CARE

- Finding the right treatment for the right patient
- Potential for less toxicity with more reliable treatment outcomes

Precision cancer care & precision cancer diagnostics

EMPIRIC CANCER CARE

- Treatment based on tumor location verses genomic drivers of disease
- Potential for more toxicity with less reliable treatment outcomes

Precision cancer diagnostics

Targeted therapy biomarkers

Kinase activation
(EGFR, ALK, NTRK, etc.)

HRD biomarkers
(BRCA1/2, ATM, etc.)

Immunotherapy biomarkers

PDL-1 IHC

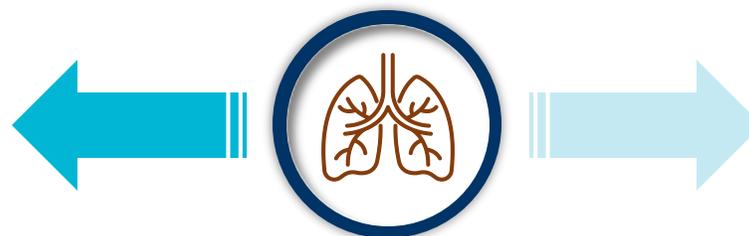
Microsatellite instability

Tumor mutational burden

PRECISION CANCER CARE

- Finding the right treatment for the right patient
- Potential for less toxicity with more reliable treatment outcomes

Multi-drug chemo-immunotherapy



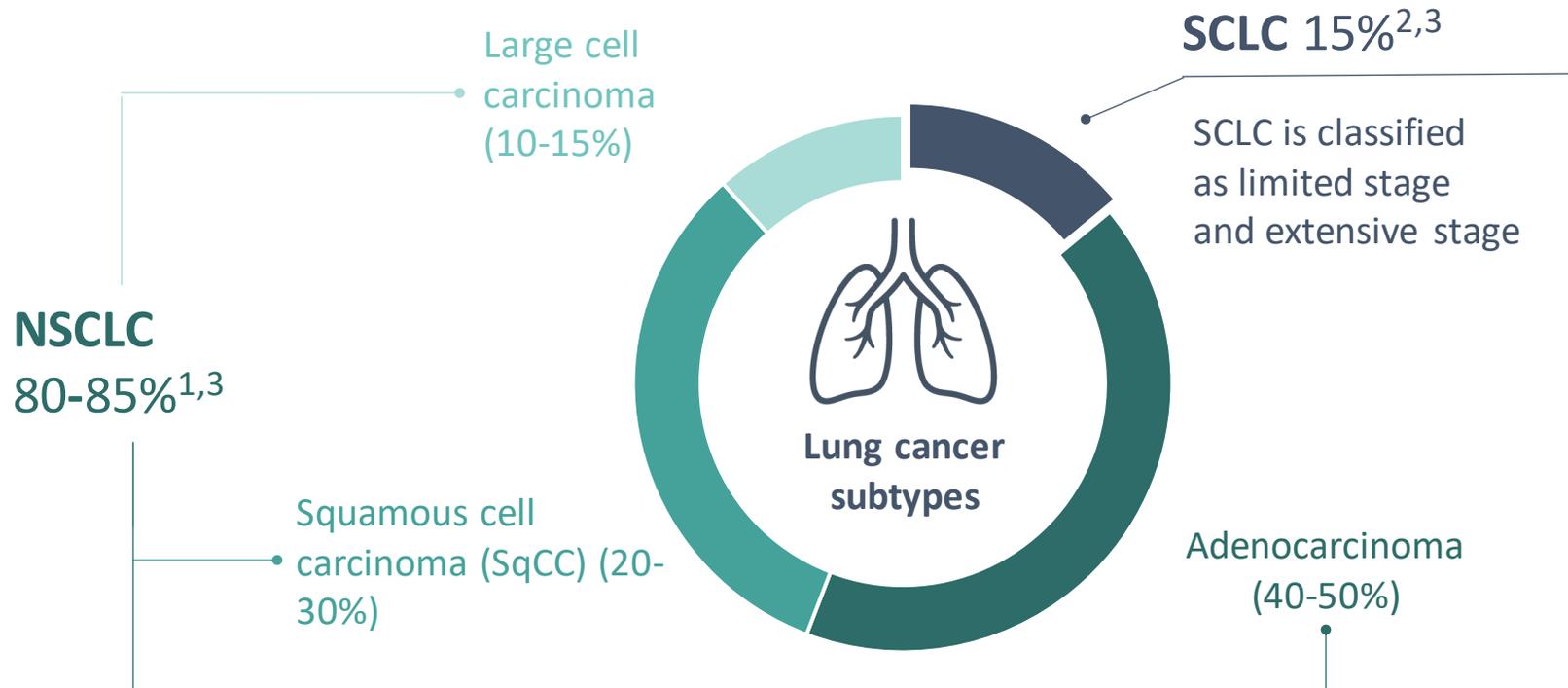
Single-agent precision therapies



Molecular Diagnostics in Lung Cancer – Considerations and Relevance for Treatment Selection

Matthew Krebs, Clinical Senior Lecturer in Experimental Cancer Medicine, University of Manchester and Consultant in Medical Oncology, The Christie NHS Foundation Trust, Manchester, UK

Traditionally lung cancer divided into histological sub-types



Pathologists are required to categorise lung cancer into adenocarcinoma and SqCC, because the same drugs used to treat adenocarcinoma can be inappropriate for treatment of SqCCs due to side effects, e.g. in the case of bevacizumab^{3,4}

NSCLC: non-small cell lung cancer; SCLC, small cell lung cancer; SqCC, squamous cell carcinoma.

1. Osmani, L., et al. (2018) *Semin Cancer Biol* 52:103–109; 2. SCLC NCCN Guidelines Version 2.2020;

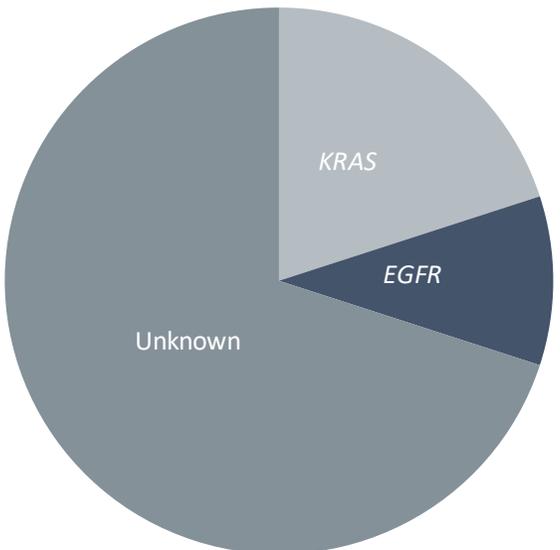
3. Inamura, K., (2017) *Front Oncol* 7:193; 4. Reck, M., et al. (2012) *Ann Oncol* 23:1111-1120.



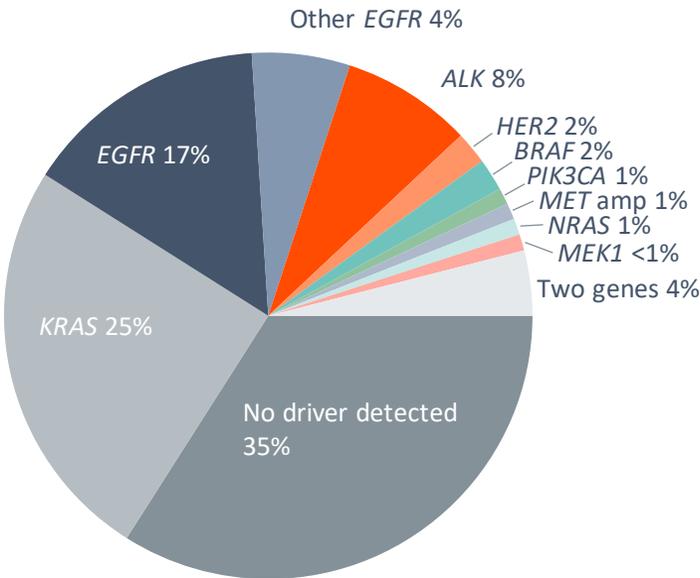
Understanding of genomic landscape of lung cancer has dramatically evolved in recent years

Adenocarcinoma

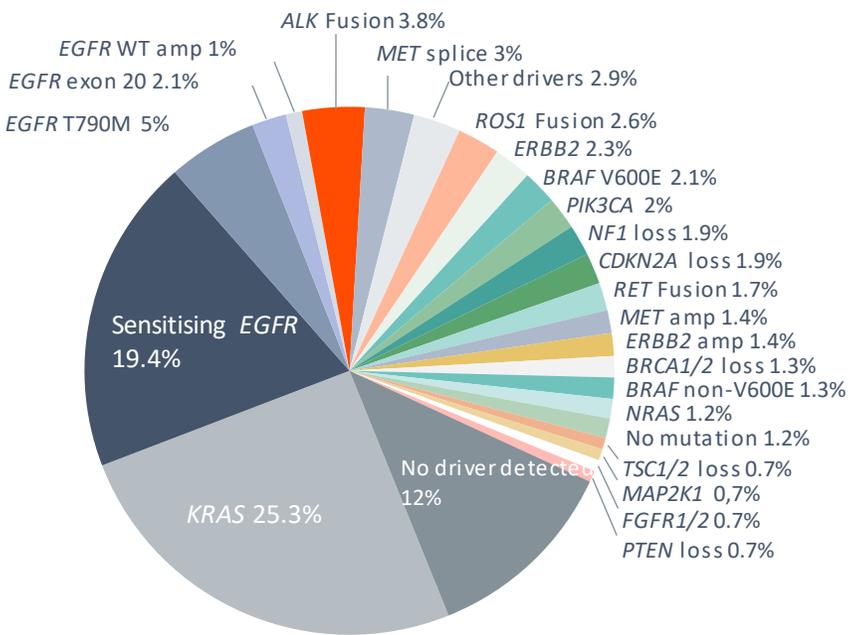
2004¹



2014²

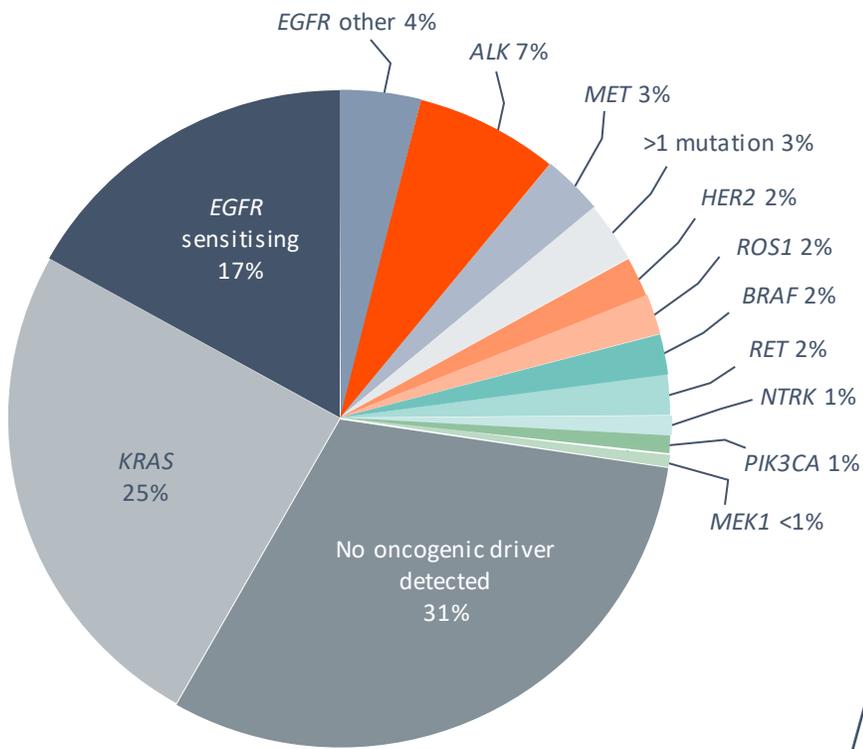


2017³



Advanced diagnostics inform therapy selection in lung cancer

Targetable mutations in lung cancer¹



Approved drugs
Investigational drugs



Identifying actionable mutations with broad genomic profiling²

Gene	Approved drugs	Investigational drugs
EGFR	<ul style="list-style-type: none"> Afatinib Dacomitinib ▼ Erlotinib (± anti-VEGF / VEGFR) Gefitinib Necitumumab ▼¹ Osimertinib ▼ 	<ul style="list-style-type: none"> JNJ-372³ Poziotinib⁶ TAK-788⁵ U3-1402⁴
ALK	<ul style="list-style-type: none"> Alectinib ▼ Brigatinib ▼ Ceritinib ▼ Crizotinib Lorlatinib ▼ 	<ul style="list-style-type: none"> Ensartinib⁸ Repotrectinib⁷
MET	<ul style="list-style-type: none"> Cabozantinib ▼¹ Crizotinib Capmatinib⁹ Savolitinib¹⁰ Tepotinib¹¹ 	
HER2	<ul style="list-style-type: none"> Afatinib¹ Dacomitinib ▼¹ Pertuzumab + trastuzumab ▼¹³ Trastuzumab emtansine¹ / deruxtecan¹⁴ 	<ul style="list-style-type: none"> Poziotinib⁶ TAK-788⁵
RET	<ul style="list-style-type: none"> Apatinib¹ Cabozantinib ▼ Lenvatinib ▼¹ Ponatinib ▼¹ Vandetanib ▼ 	<ul style="list-style-type: none"> Selpercatinib¹⁵ Pralsetinib¹⁶
BRAF	<ul style="list-style-type: none"> Dabrafenib (± trametinib) Vemurafenib 	
NTRK	<ul style="list-style-type: none"> Cabozantinib ▼¹ Entrectinib Larotrectinib ▼ Repotrectinib⁷ Selitrectinib¹⁷ 	
ROS1	<ul style="list-style-type: none"> Ceritinib ▼ Crizotinib Lorlatinib ▼ Repotrectinib⁷ 	<ul style="list-style-type: none"> DS-6051b¹
PIK3CA	<ul style="list-style-type: none"> Copanlisib¹² 	
MEK1	<ul style="list-style-type: none"> Cobimetinib ▼¹ Selumetinib¹ Trametinib¹ 	

All drugs listed are included in NSCLC NCCN Guidelines unless otherwise indicated.

Some drugs are investigational and not approved in any indication. Some non-investigational drugs are only approved for use in specific indications in Europe and / or USA and / or Japan. Therapies marked with ▼ are subject to additional monitoring. Reporting suspected adverse reactions after authorisation of the medicinal product is important. Adverse events should be reported to your respective local office. Amgen Europe B.V.: Trastuzumab (Kanjinti); AstraZeneca AB: Osimertinib; Bayer AG: Larotrectinib; Celltrion Healthcare Hungary Kft.: Trastuzumab (Herzuma); Eli Lilly Nederland B.V.: Necitumumab; Eisai Europe Limited: Lenvatinib; Genzyme Europe B.V.: Vandetanib; Incyte Biosciences Distribution B.V.: Ponatinib; Ipsen Pharma: Cabozantinib; Mylan S.A.S.:

Trastuzumab (Oqivri); Novartis Europharm Limited: Ceritinib; Pfizer Europa MA EEG: Trastuzumab (Trazimera); Pfizer Europe MA EEIG: Dacomitinib, Lorlatinib; Roche Registration GmbH: Alectinib, Cobimetinib; Samsung Bioepis UK Limited: Trastuzumab

(Ontruza); Takeda Pharma A/S: Brigatinib. 1. Adapted from Tsao, A.S., et al. (2016) *J Thorac Oncol* 11:613-38; 2. NSCLC NCCN Guidelines Version 2.2020; 3. NCT02609776; 4. NCT03260491; 5. NCT02716116; 6. NCT03318939; 7. NCT03093116; 8. NCT02767804; 9.

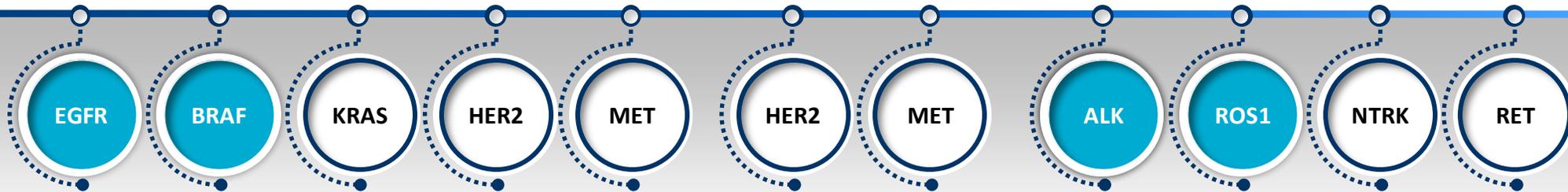
NCT03693339; 10. NCT03778229; 11. NCT02864992; 12. NCT02465060; 13. NCT03845270; 14. NCT03505710; 15. NCT04268550; 16. NCT04204928; 17. NCT03206931.

Clinical guidelines do not always match clinical practice



Biomarker testing in non-small cell lung cancer

● Key recommended tests ○ Recommended to be assessed if NGS is used for broader testing



Molecular testing method	Point mutations and small indels	Copy number alterations	Rearrangements
PCR and conventional sequencing	✓		
FISH			✓
IHC		✓*	✓*
NGS (amplicon-based)	✓		
NGS (hybrid capture-based)	✓	✓	✓

CAP / IASLC / AMP¹ / ESMO²

Molecular testing guideline

“In general, capture-based [NGS] methods may be preferable for initial testing of lung cancer samples in order to detect rearrangements... as well as a broader range of potential genetic markers”¹

“If available, multiplex platforms (NGS) for molecular testing are preferable”²

ASCO Educational Book³

Biomarker testing for advanced NSCLC

“For very limited samples... for which multiple tests cannot be performed, [hybrid capture-based] assays are preferable for upfront comprehensive assessment”

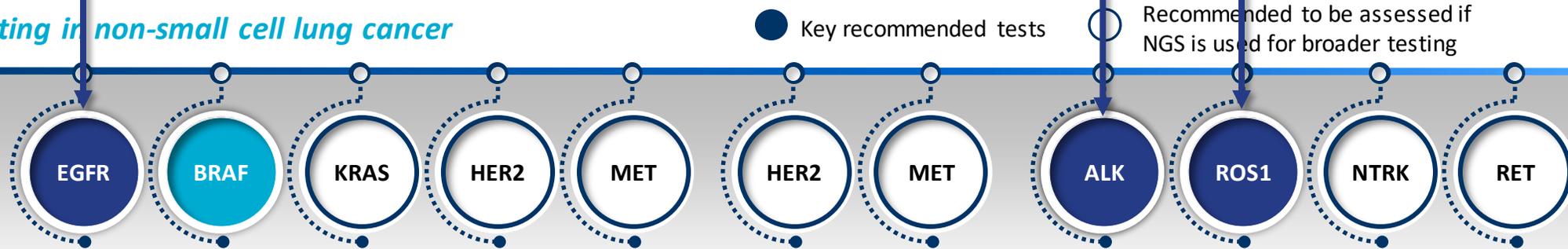
* IHC is used to detect *MET* overexpression and *ALK* translocations respectively.

AMP: Association for Molecular Pathology; ASCO: American Society of Clinical Oncology; CAP: College of American Pathologists; ESMO: European Society for medical oncology; FISH: fluorescence in situ hybridisation; IASLC: International Association for the Study of Lung Cancer; NSCLC: non-small cell lung cancer; PCR: polymerase chain reaction. Table adapted from Pennell, N.A., et al. (2019). 1. Lindeman LI, et al. *J Mol Diagn* 2018; 20:129–159; 3. Pennell NA, et al. *ASCO Educational Book* 2019; 39:531–542; 4. Domagala-Kulawik J, et al. *Front Med (Lausanne)* 2019; 6:284.

Clinical guidelines do not always match clinical practice

Biomarker testing in non-small cell lung cancer

Usual current clinical practice



Molecular testing method	Point mutations and small indels	Copy number alterations	Rearrangements
PCR and conventional sequencing	✓		
FISH			✓
IHC		✓*	✓*
NGS (amplicon-based)	✓		
NGS (hybrid capture-based)	✓	✓	✓

CAP / IASLC / AMP¹ / ESMO²

Molecular testing guideline

“In general, capture-based [NGS] methods may be preferable for initial testing of lung cancer samples in order to detect rearrangements... as well as a broader range of potential genetic markers”¹

“If available, multiplex platforms (NGS) for molecular testing are preferable”²

ASCO Educational Book³

Biomarker testing for advanced NSCLC

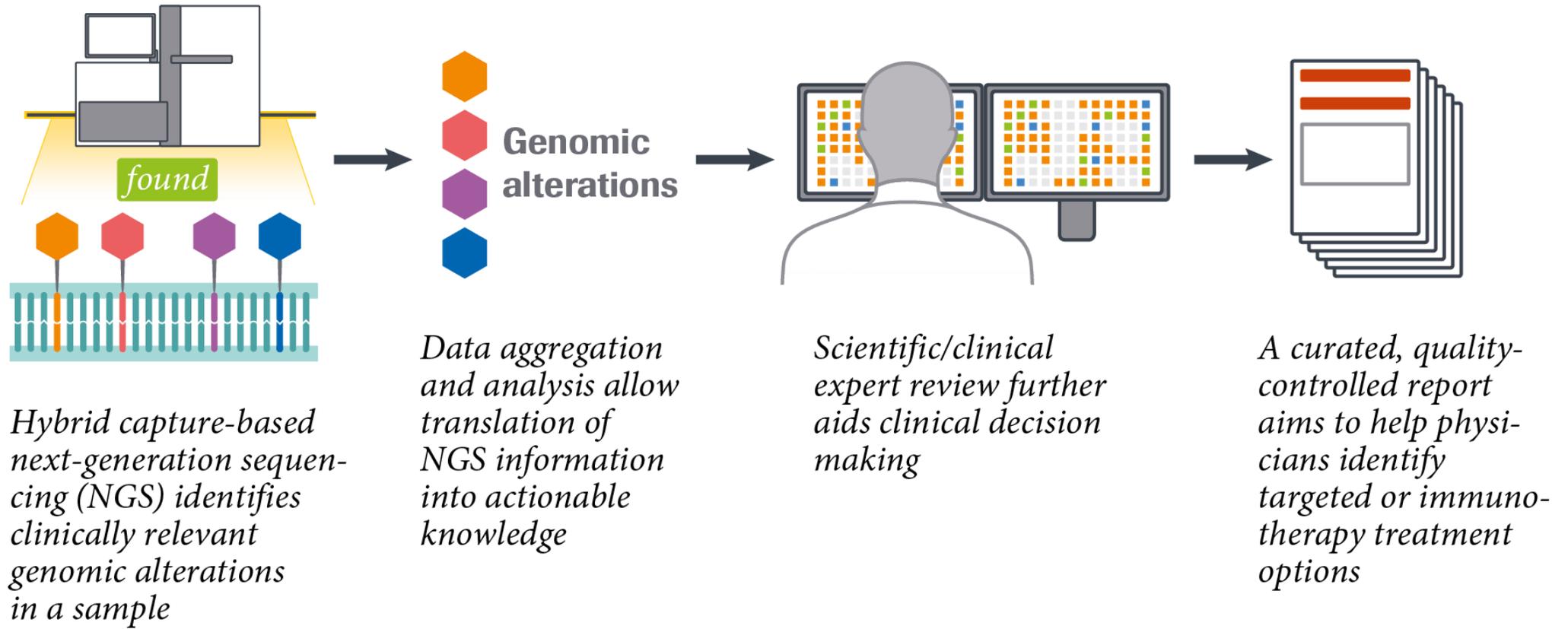
“For very limited samples... for which multiple tests cannot be performed, [hybrid capture-based] assays are preferable for upfront comprehensive assessment”

* IHC is used to detect *MET* overexpression and *ALK* translocations respectively.

AMP: Association for Molecular Pathology; ASCO: American Society of Clinical Oncology; CAP: College of American Pathologists; ESMO: European Society for medical oncology; FISH: fluorescence in situ hybridisation; IASLC: International Association for the Study of Lung Cancer; NSCLC: non-small cell lung cancer; PCR: polymerase chain reaction. Table adapted from Pennell, N.A., et al. (2019). 1. Lindeman LI, et al. *J Mol Diagn* 2018; 20:129–159; 3. Pennell NA, et al. *ASCO Educational Book* 2019; 39:531–542;

4. Domagala-Kulawik J, et al. *Front Med (Lausanne)* 2019; 6:284.

Next generation sequencing – analysis *and* bioinformatics



Tissue versus liquid biopsy testing for genomic alterations

Tissue based genomic profiling remains standard of care with advantages such as higher sensitivity for certain types of alterations¹³ but suffers from limitations impacting patient care.



solid biopsy is **difficult** or poses a **high risk**¹²



Does not capture tumour **heterogeneity**⁶



Robust and highly specific tests intensively validated



capture tumour **heterogeneity**⁶



tissue samples may be **insufficient**, inadequate or exhausted⁵⁻⁷



Enable **timely** personalised treatment decisions by shortening time to results



lower cost, ease of collection^{7,9}



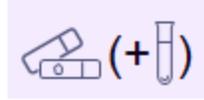
monitoring disease progression/recurrence^{3,4}

Patient care plan may suffer if timely testing not available, leading to reduced therapy options with potential higher adverse events

The potential clinical applications of liquid biopsy are wide-ranging



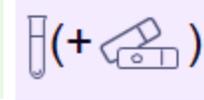
Early cancer detection



MRD



Recurrence surveillance



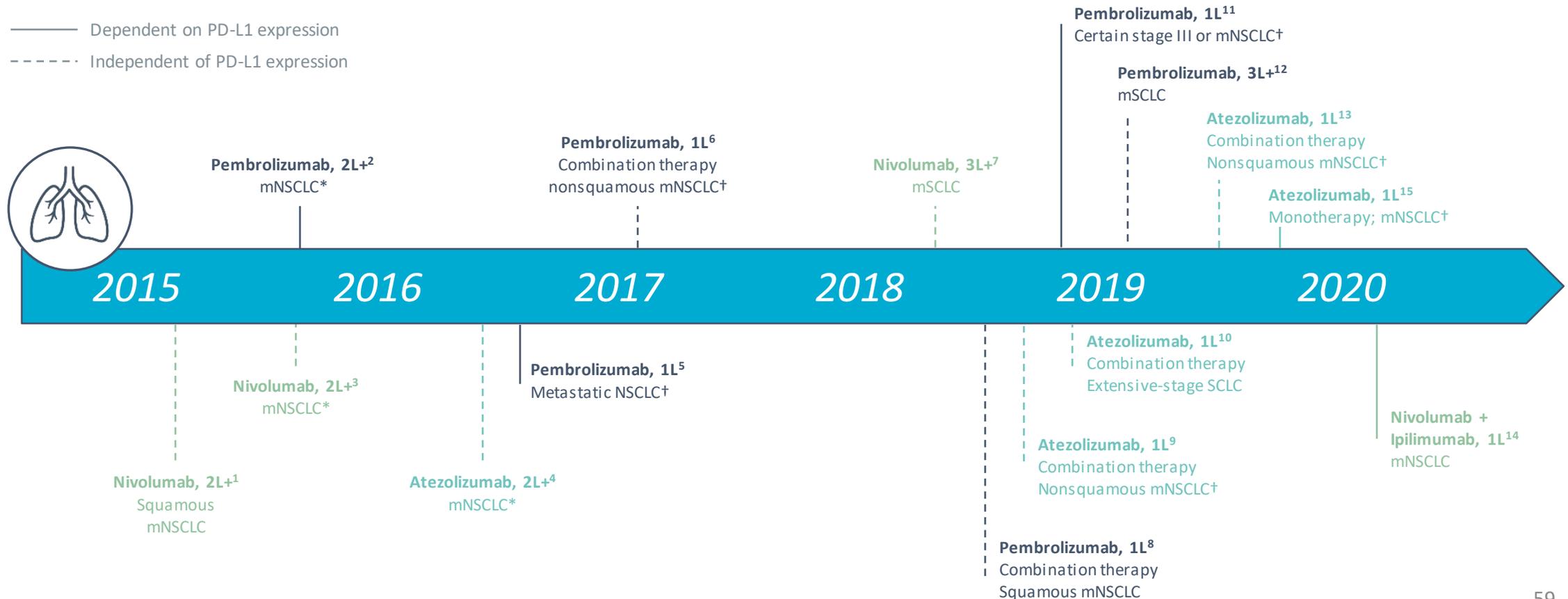
Tx selection (CGP)

Tx response monitoring

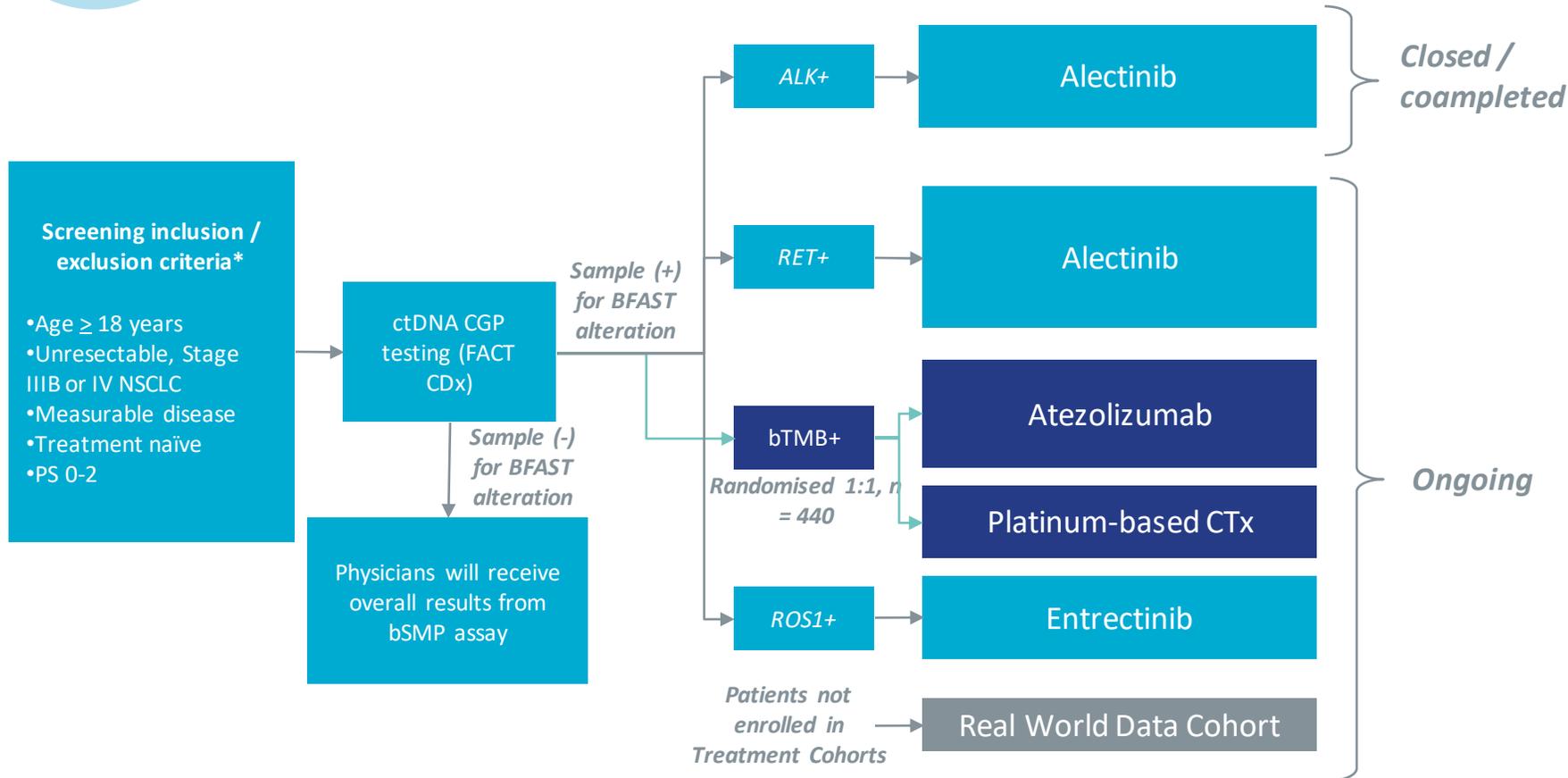


Immunotherapy is well established in advanced lung cancer

Timeline of developments for immunotherapies in lung cancer



BFAST: blood-first assay screening trial measures bTMB to inform clinical decision making in NSCLC



Phase II/III BFAST trial¹

- Treatment-naïve, aNSCLC patients screened using blood-based NGS assays and enrolled into targeted treatment / immunotherapy cohorts
- Median bTMB at baseline was 2 mutations
- 3/87 (3.4%) had bTMB \geq 16 mutations
- LBx identified a similar proportion of patients with ALK mutations (5.4%) to that typically seen with traditional biopsy (5%)
- 12-month DoR was 75.9%

*All cohorts have additional, treatment-specific inclusion / exclusion criteria

aNSCLC: advanced non-small cell lung cancer; BFAST: blood-first assay screening trial; BID: twice daily; bSMP: blood somatic mutation profiling; bTMB: blood based tumour mutational burden; ctDNA: circulating tumour DNA; CTx: chemotherapy; DoR: duration of response; FMI: Foundation Medicine Inc.; IV: intravenously; LBx: liquid biopsy; NSCLC: non-small cell lung cancer; PD: progressive disease; PO: orally; PS: performance status; Q3W: every 3 weeks. Gadgeel, S., et al. presented at ESMO 2019, abstract LBA81_PR.

Conclusions

- Lung cancer is the poster boy for precision medicine with effective targeted treatments licensed and in development
- Liquid biopsy is convenient, efficient and technologies have evolved to perform broad panel testing from ctDNA
- Tissue testing for NGS still has its place and will likely be complementary with liquid-based testing
- Serial sampling provides important insights into potential resistance mechanisms and will guide next generation of targeted therapies.
- Challenges will be funding of NGS in different health-care systems and reimbursement of therapies

How to integrate these advanced diagnostics tools into clinical routine?



Open Discussion

Please use the Chat feature to ask questions and make comments



Final Remarks

Co-Chair:

Geoff Oxnard, Vice President, Global Medical Lead,
Liquid Franchise at Foundation Medicine



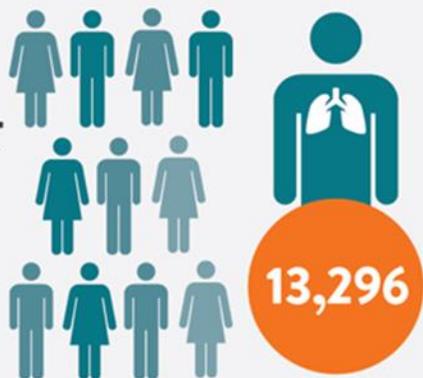
Removing health system delays in lung cancer

Co-Chair:

Ouzna Morsli, EMEAC Oncology Medical Lead, MSD

Time is of the essence for people living with lung cancer: delays in diagnostic testing and referral for treatment must be eliminated

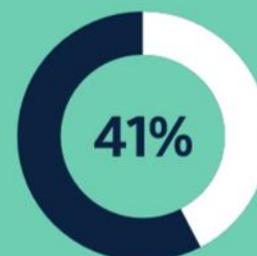
**13,296 MORE LUNG
CANCER PATIENTS
ARE ALIVE FOR AT LEAST
FIVE YEARS FOLLOWING
A DIAGNOSIS IN 2014,
COMPARED TO THOSE
DIAGNOSED IN 2004**



Lung cancer incidence and survival data taken Hofmarcher, T. et al. (2019) *Comparator Report on Cancer in Europe 2019*, IHE Report 2019:7. IHE: Lund, Sweden



In the lung cancer clinical guidelines of the countries studied:



Do not include fast-tracking people suspected of having lung cancer for diagnostic testing.

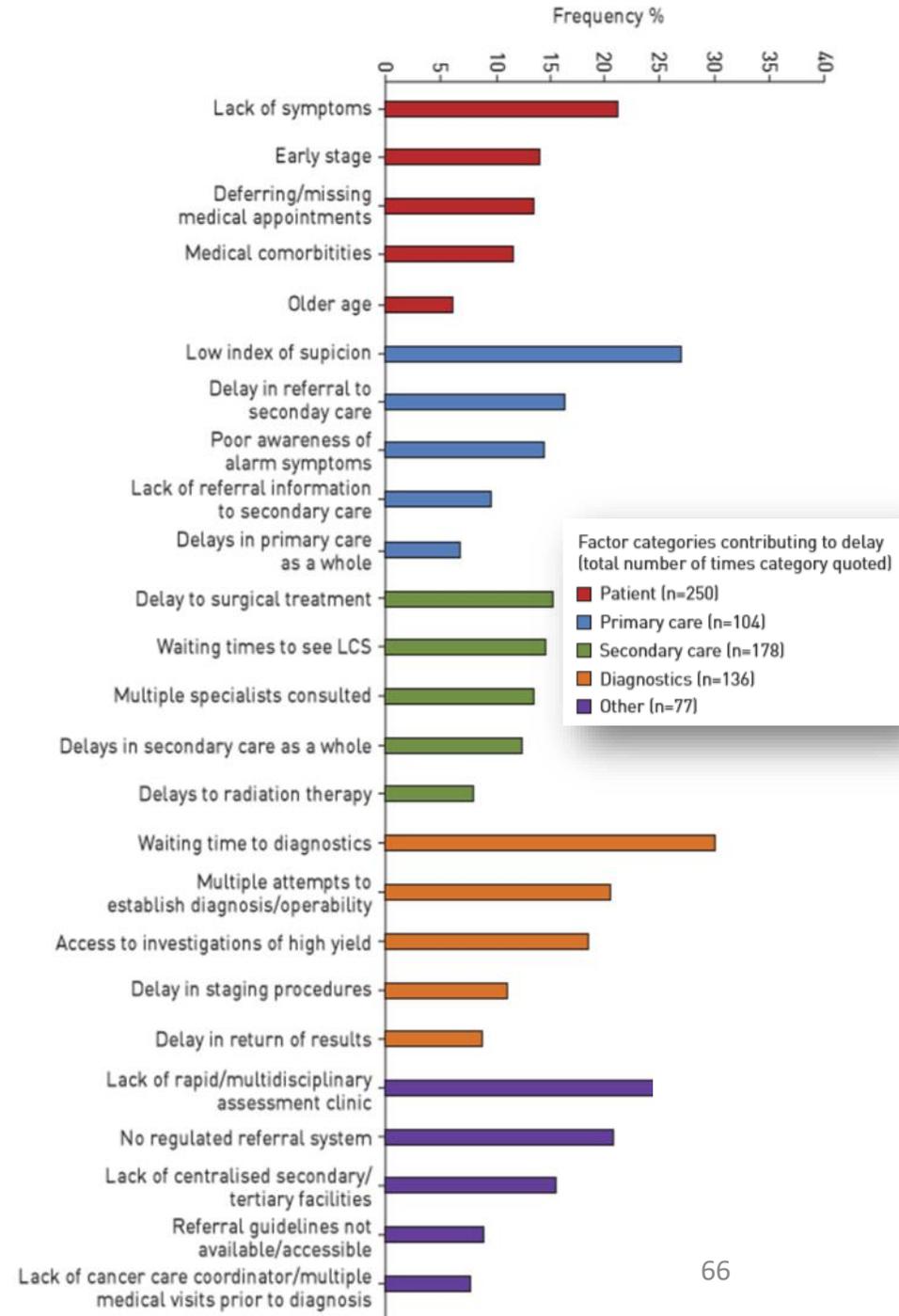
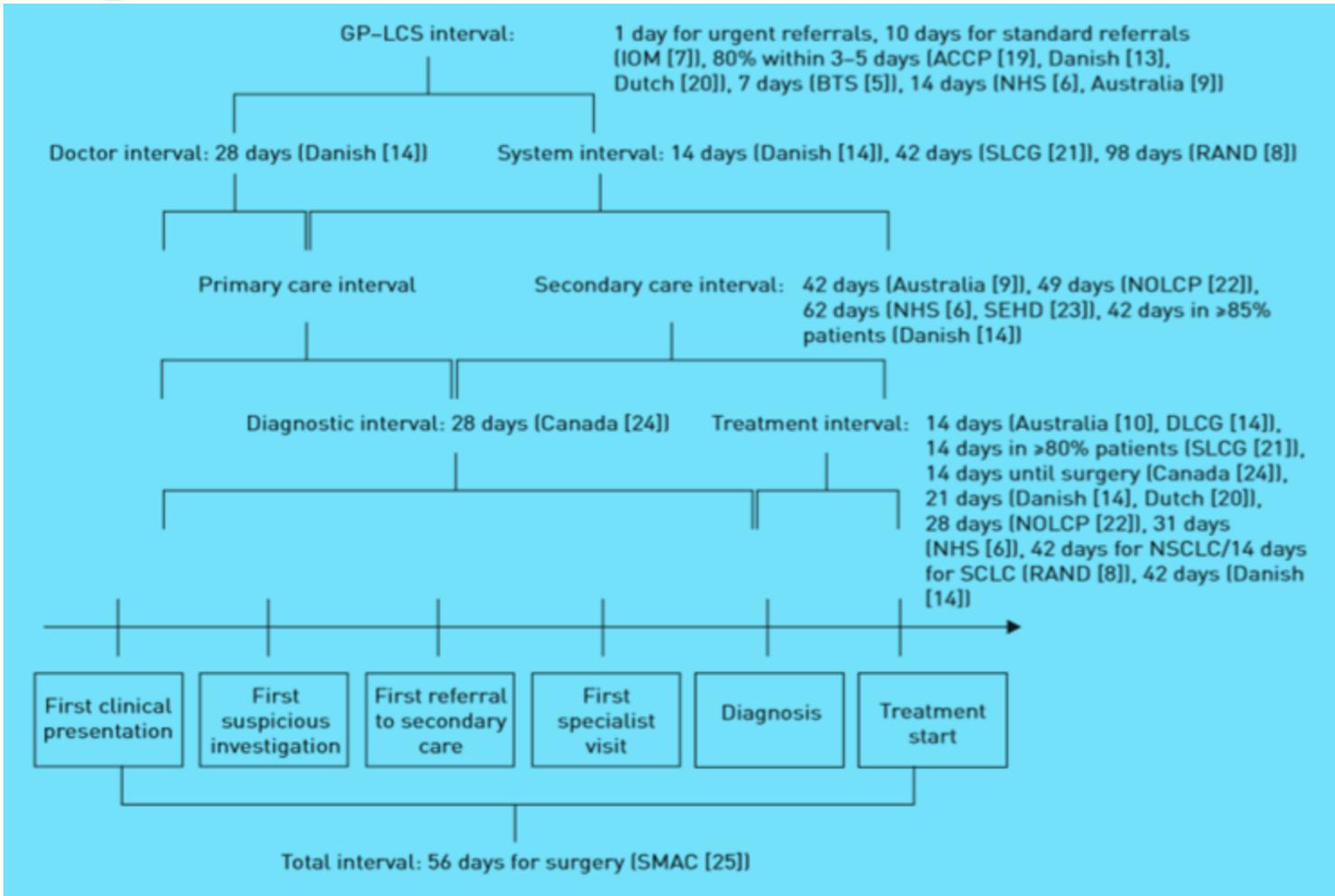


Do not include a specific timeframe for obtaining diagnostic testing.



Do not include rapid referral for newly diagnosed patients to obtain treatment.

Removing health system delays in lung cancer





The Dutch Lung Cancer Audit: Nationwide Quality of Care Evaluation Using Quality Indicators

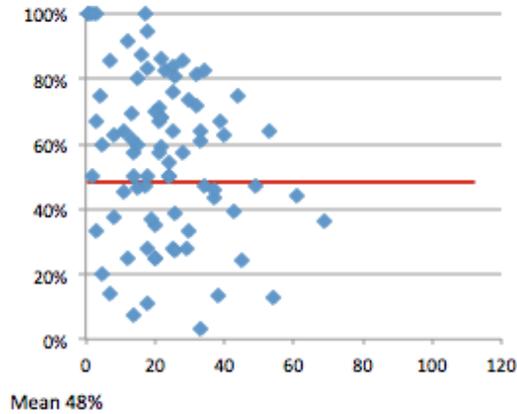
Hans J.M. Smit, Pulmonologist, Rijnstate Hospital, Chairman
of the Dutch Lung Cancer Audit, Arnhem, The Netherlands

Rawa Ismail, PharmD and PhD Candidate DICA

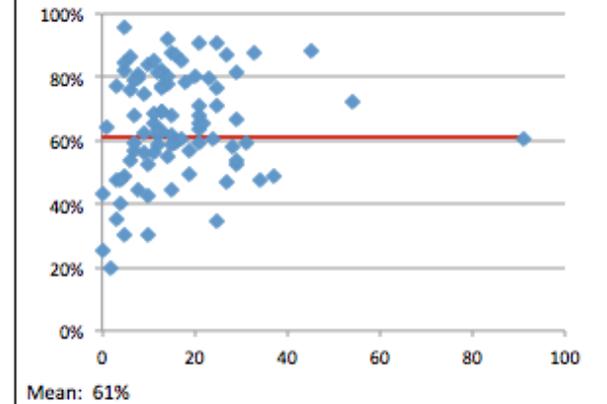
What to achieve

- Improving quality by bench marking
- Preferably in a safe setting (anonymous)

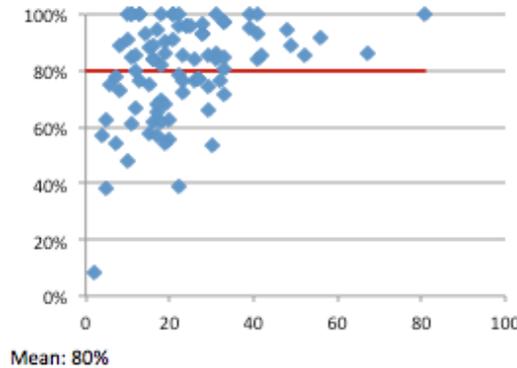
2009



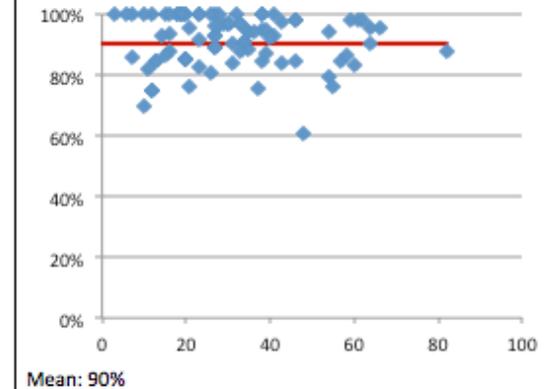
2010



2011



2012



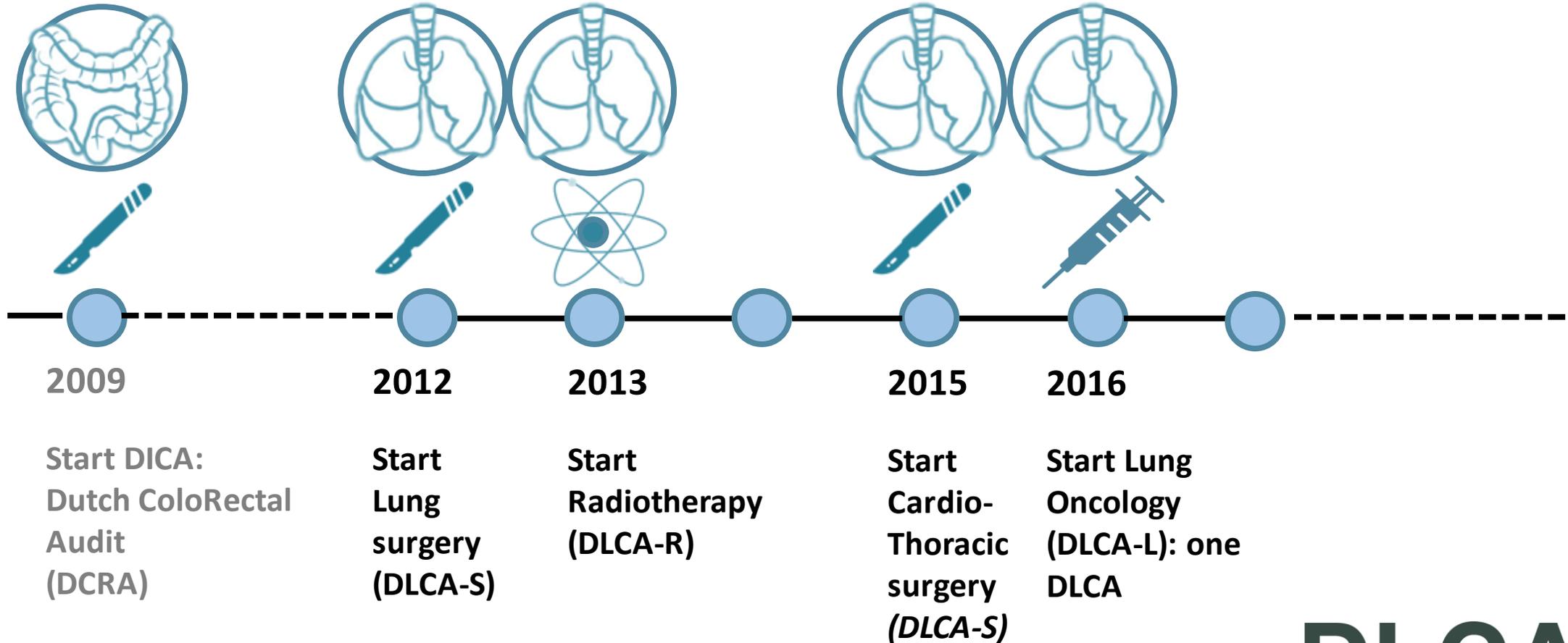
Key messages

- Quality registries play an important role in quality care improvement
 - It takes time to initiate a nation-wide registry
 - Insight into own data leads to improved care
 - Benchmarking with other hospitals can lead to discussions and best-practice examples
- Quality indicators are important to measure the quality of care: start “simple”, improve over time
 - Be aware of the registration burden for hospitals
 - Indicators on processes in the hospital can lead indirectly to better care
 - Outcome indicators are of high value and should be measured when registry data are rich and trustworthy

Content

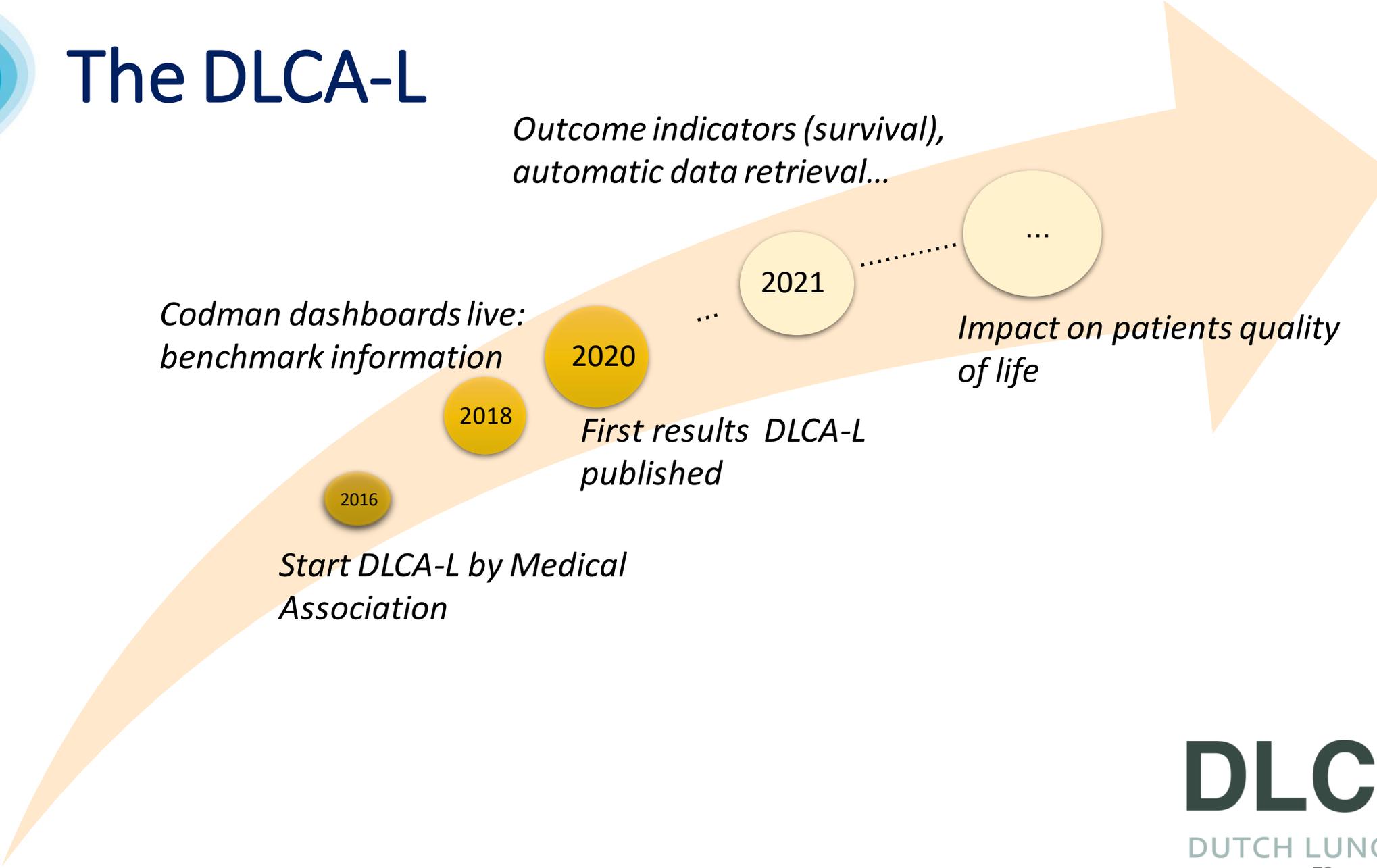
- What is DLCA?
- How was DLCA initiated?
- What is measured?
- What has been achieved?
- How to use it in clinical practice?

Multidisciplinary DLCA within DICA





The DLCA-L





Finance by production payment

- Hospital 1
- Hospital 2
- Hospital 3
- Hospital 4
- Hospital 5

Finance by government

DICA through MRDM

DICA Scientific buro

DLCA board

Mandated from Specialists Association and Patient Representatives

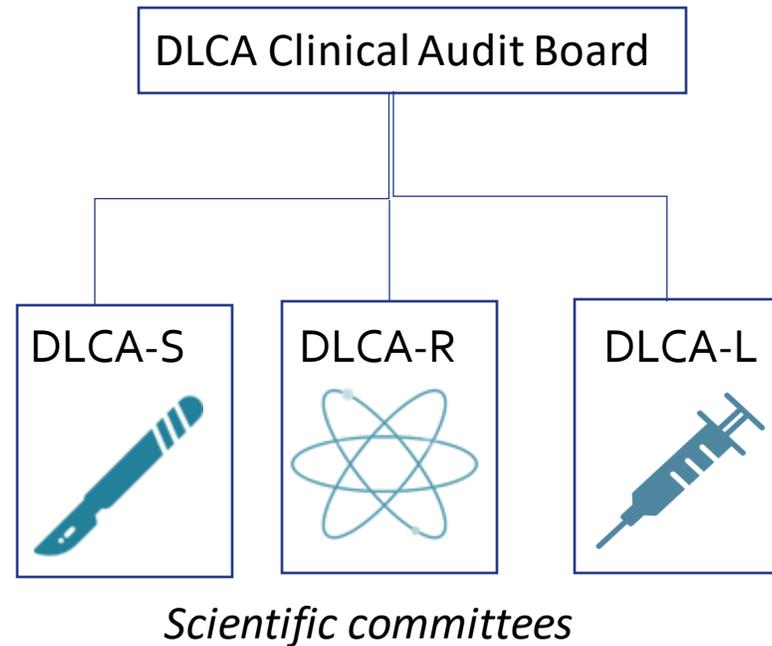
- Open character
- Publications
- Develop Indicators:
 - Based on guidelines*
 - Based on science*
 - Based on patient needs*

DICA supports law, ICT, epidemiology

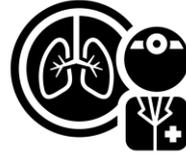
Scientific board

Initiation of DLCA

- Initiated by professional association of chest physicians (NVALT)
- Facilitated by DICA
- Subregistries with own scientific committees and clinical audit board



DLCA-L



>40.000 patients
73 hospitals

DLCA-S



>42.000 patients
43 hospitals

DLCA-R



>18.000 patients
19 hospitals

Initiation of the DLCA

Why?

Insights into quality of care of lung cancer patients

by focusing on

- Diagnostics
- Time to Diagnosis and Therapy
- Monitoring of in-hospital times,
- Outcomes of systemic therapy
- Including Best Supportive Care = complete

Potential problems

Registration = time consuming

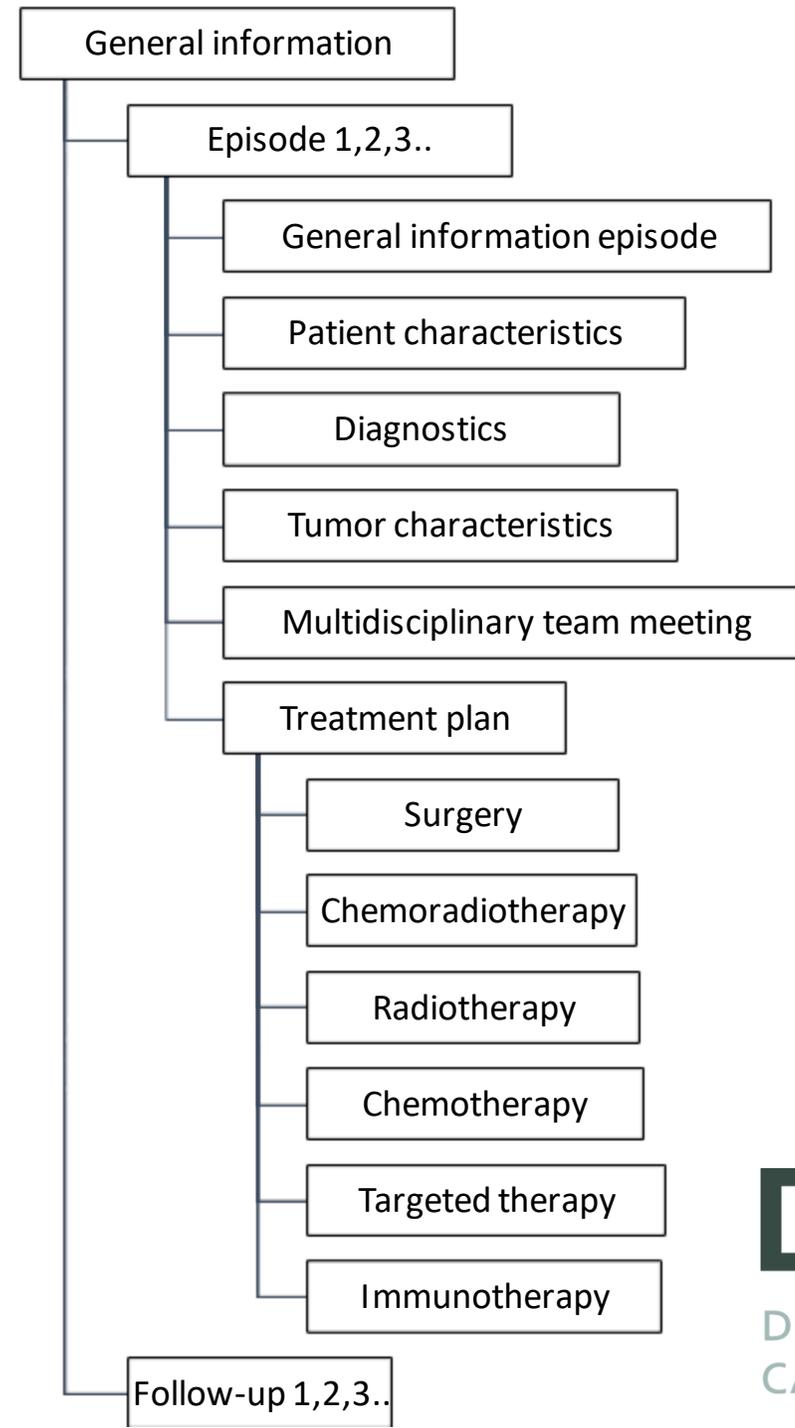
Time = competitive with patient care

Automatic subtraction = less control

Privacy law = reduction of possibilities and time consuming

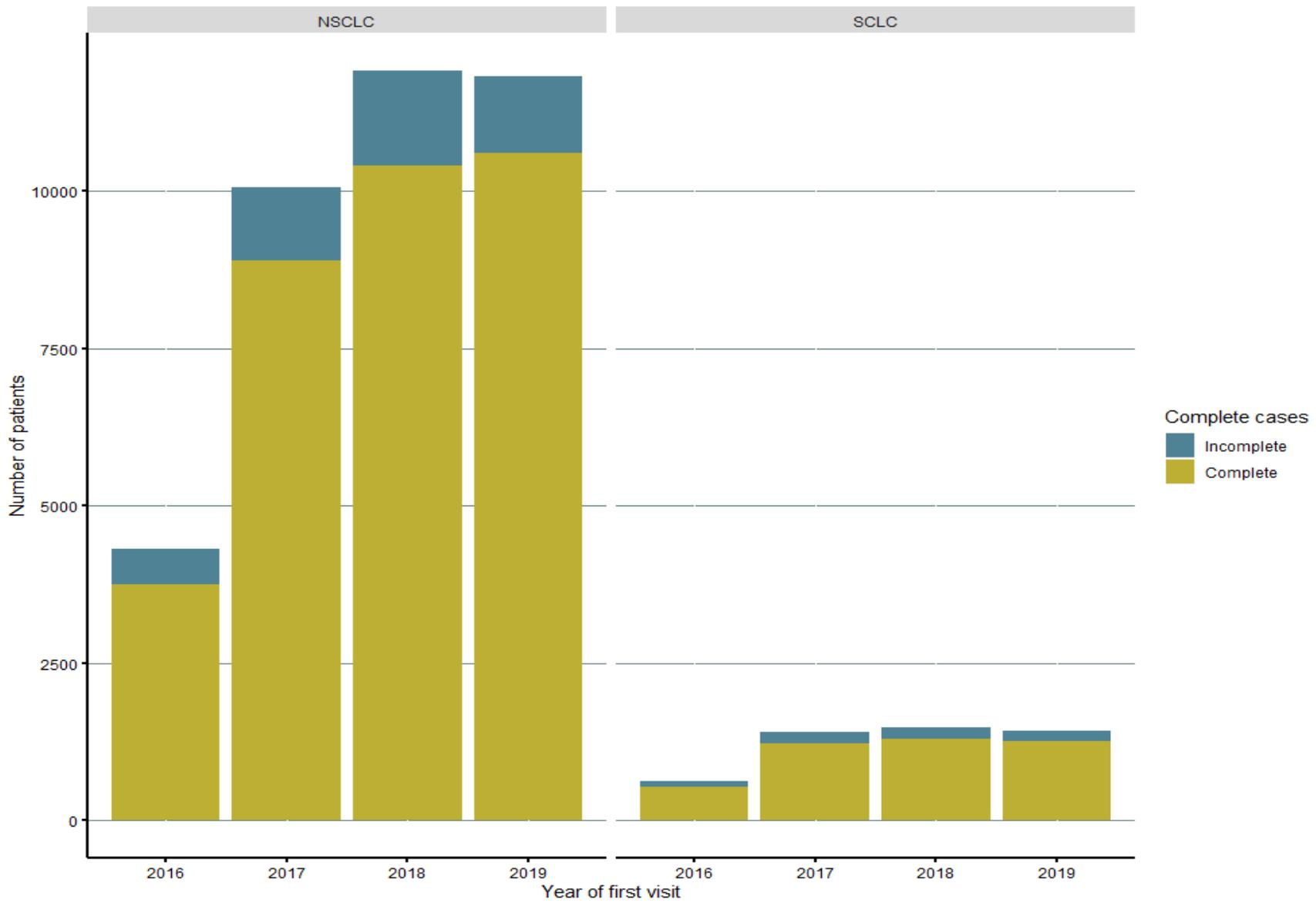
The DLCA dataset

- Inclusion: all patients with lung carcinoma
- Including clinically suspected
- 153 variables
- >40% mandatory items



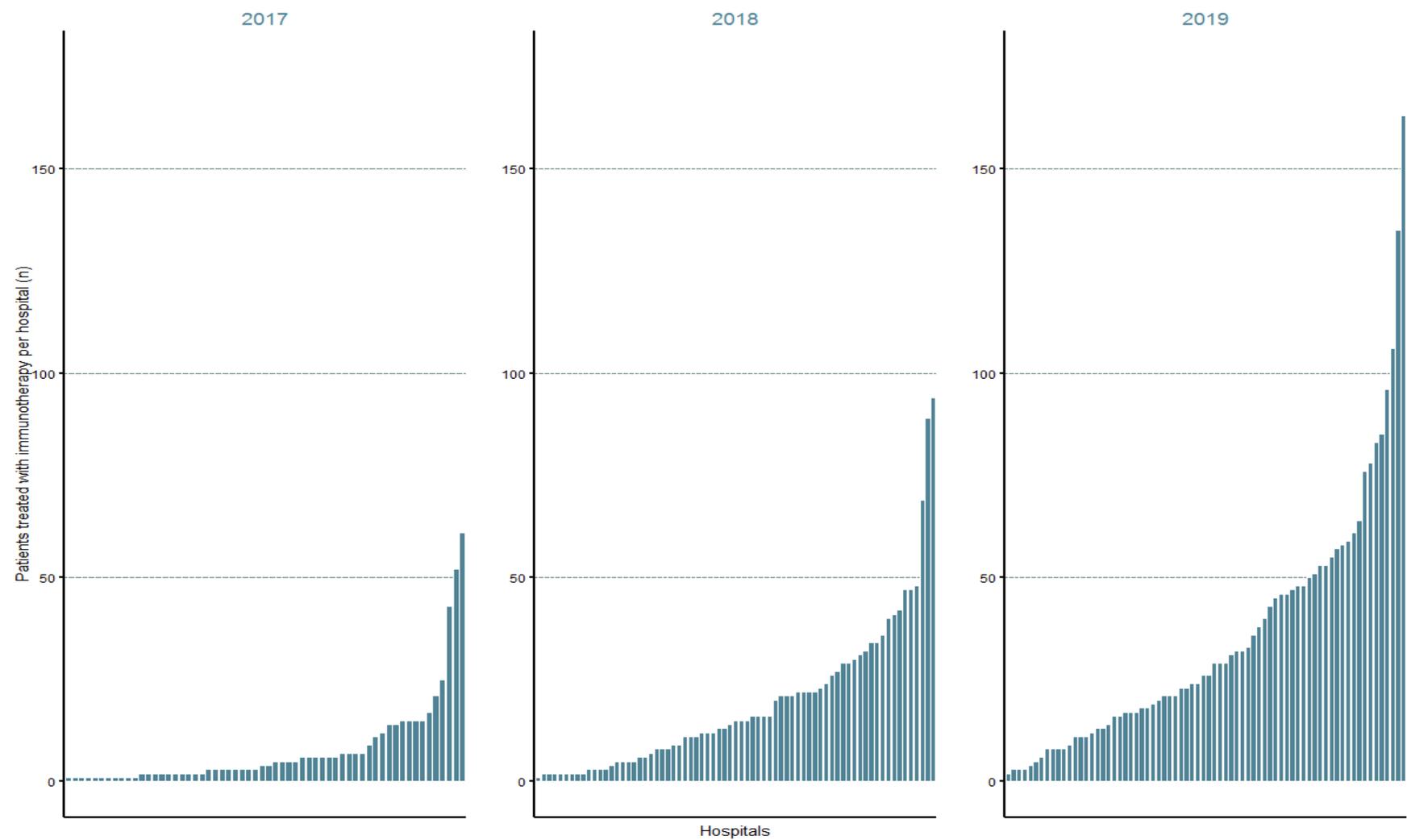


Results: Data set completeness



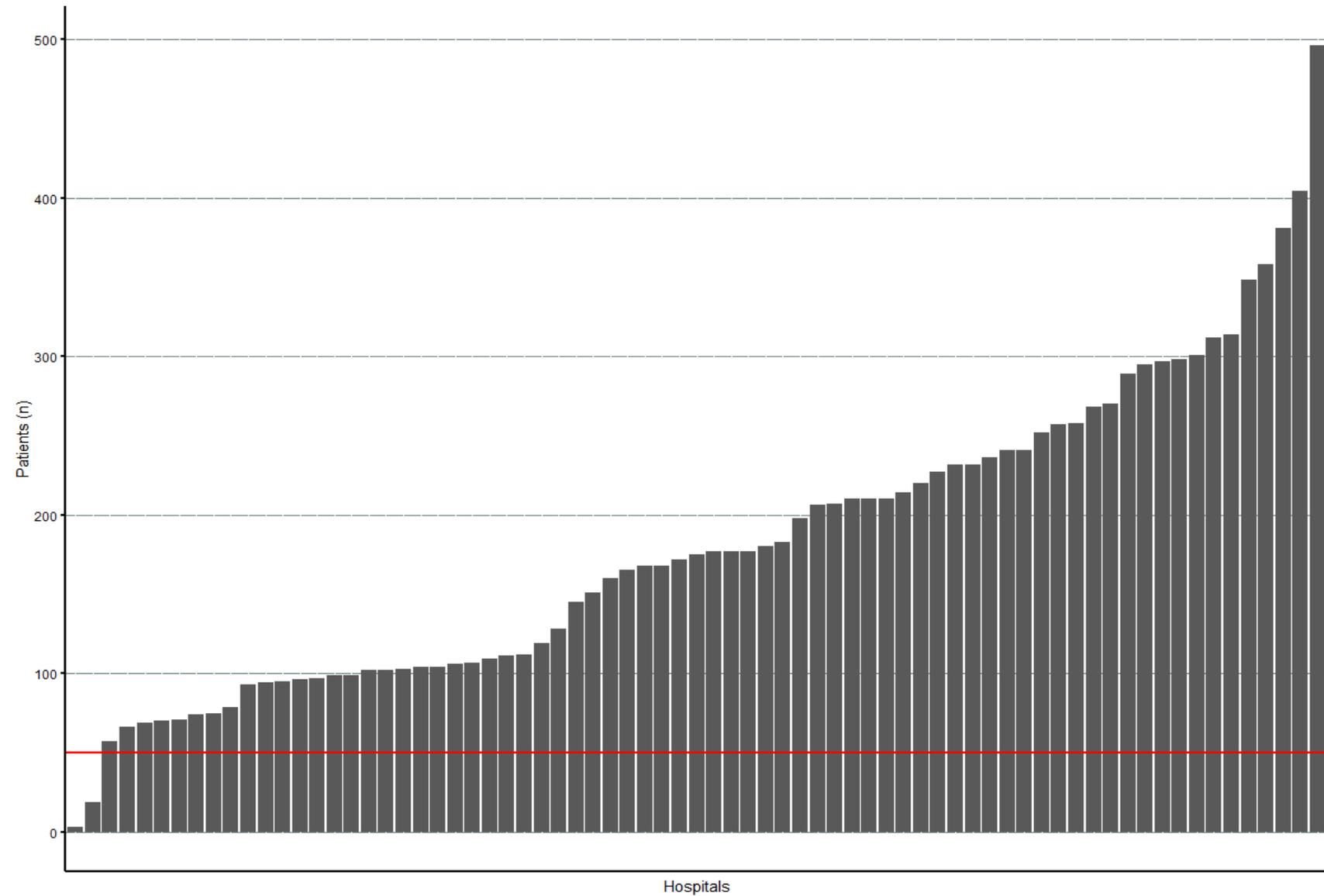


Results: Hospitals and immunotherapy





Results: Numbers of patients per hospital



DLCA
DUTCH LUNG
CANCER AUDIT



Results: Feedback process



Ernest Amory Codman, 1910

Results: Quality indicators

1) Structure quality indicators

- *Number of patients*
- *Completeness of registration*

2) Process quality indicators

- *Brain imaging*
- *Molecular diagnostics*
- *Multidisciplinary consultation*
- *Duration diagnostic trajectory*
- *First-line treatments NSCLC/SCLC*
- *Use of treatments in elderly*

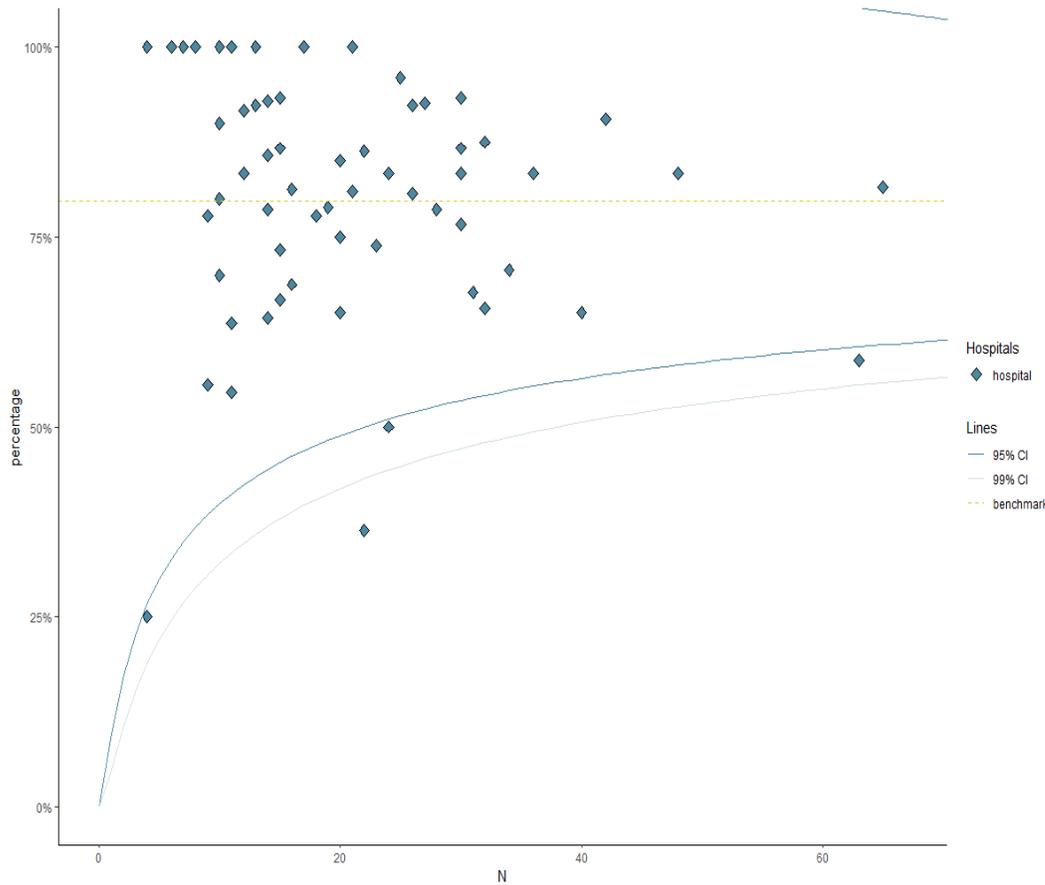
3) Outcome quality indicators

- *Grade 3/4 toxicities related to systemic treatment*

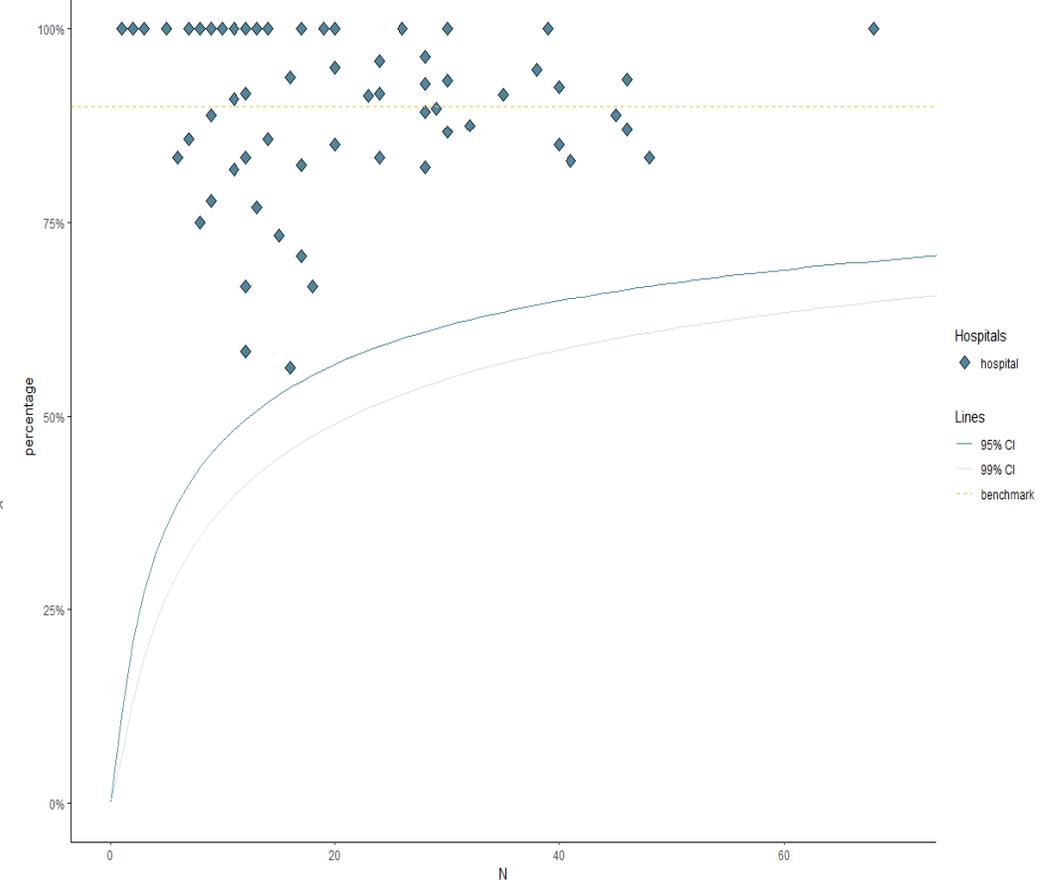


Results: Improvement in brain imaging

2017



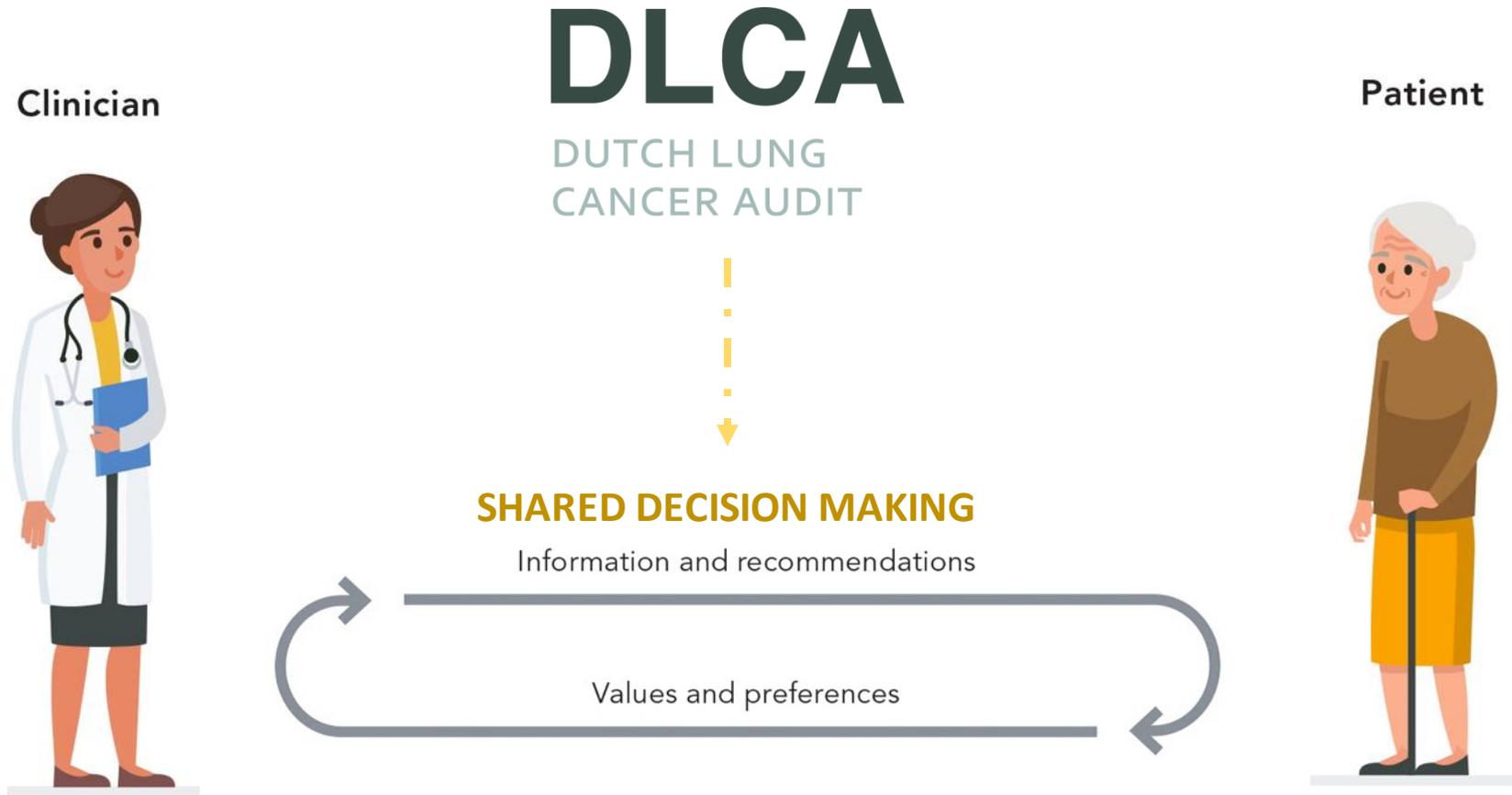
2019



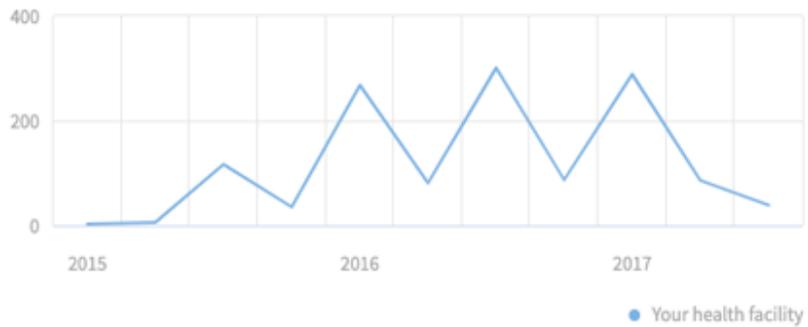
Results: Other examples of QIs

- Stage III NSCLC patients undergoing brain imaging:
 - **82%** in 2017 → **90%** in 2019
- Stage IV adenocarcinoma patients undergoing molecular diagnostics:
 - **89%** in 2017 → **93%** in 2019
- Time from diagnosis to start treatment
 - Without invasive mediastinal diagnostics (<21 days): **62%**
 - With EUS/EBUS (<21 days): **46%**
 - With mediastinoscopy (<35 days): **59%**

Use in clinical practice



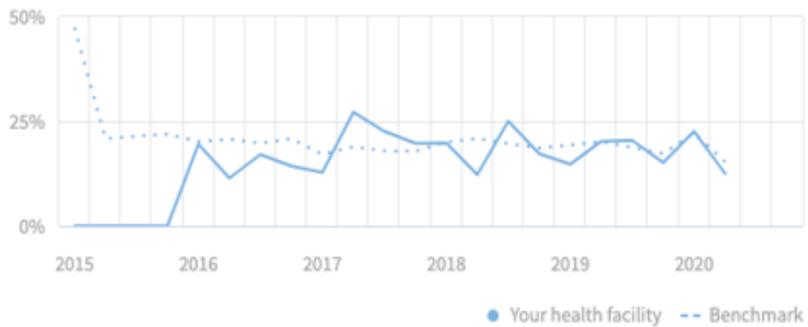
Trend number of patients



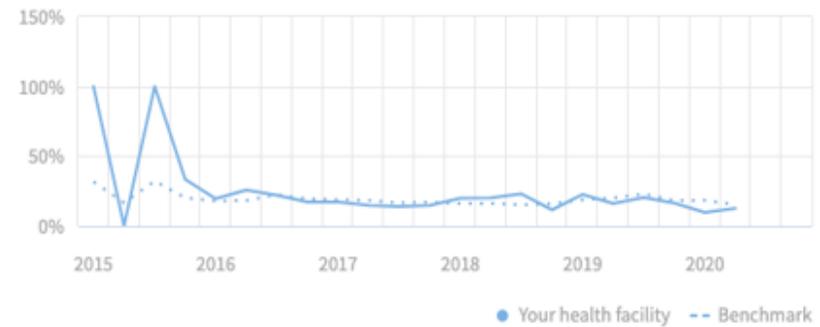
Immunotherapy ▾

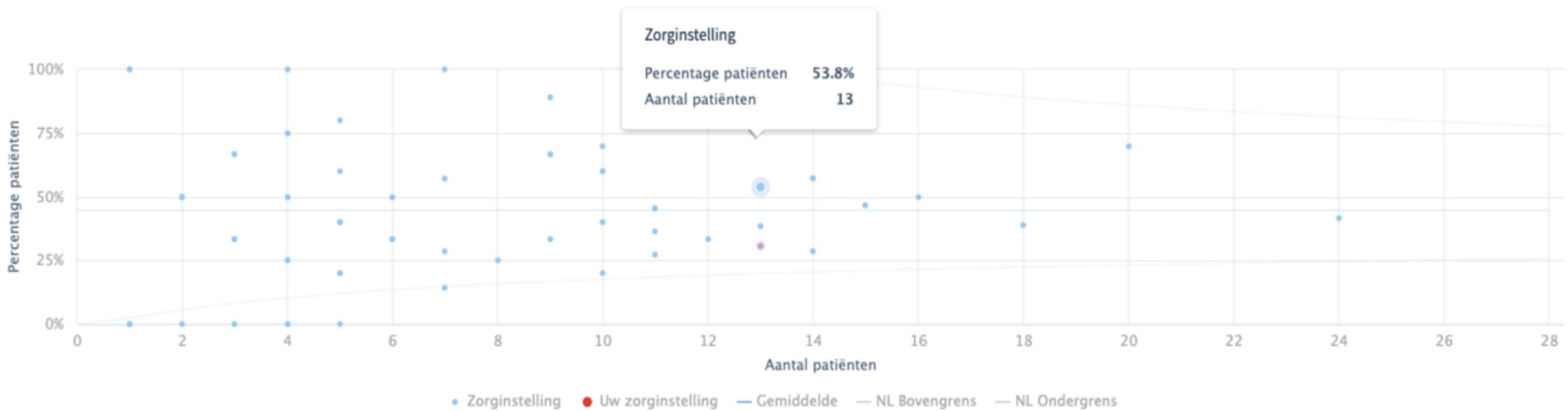


Radiotherapy ▾



Chemotherapy ▾





	Amount	Your health facility	Your health facility SD	Benchmark NL	Benchmark NL SD
Leeftijd ten tijde van diagnose	61	67.8	10.6	69.5	10.1
	Amount	Your health facility	Benchmark NL		
Leeftijd ten tijde van diagnose					
20-59 jaar	14	22.6 %	13.6 %		
60-69 jaar	40	64.5 %	57.7 %		
70-79 jaar	7	11.3 %	14.1 %		
> 80 jaar	1	1.6 %	14.6 %		
Geslacht					
Man	33	53.2 %	53.8 %		
Vrouw	29	46.8 %	44.9 %		
Onbekend / niet ingevuld	–	–	1.4 %		
ECOG score					
ECOG 0	22	35.5 %	24.2 %		
ECOG 1	25	40.3 %	38.9 %		
ECOG 2	8	12.9 %	14.1 %		
ECOG 3	4	6.5 %	7.8 %		
ECOG 4	–	–	1.4 %		
Onbekend / niet ingevuld	3	4.8 %	13.5 %		

Key messages

- Quality registries play an important role in quality care improvement
 - It takes time to initiate a nation-wide registry
 - Insight into own data leads to improved care
 - Benchmarking with other hospitals can lead to discussions and best-practice examples
- Quality indicators are important to measure the quality of care: start “simple”, improve over time
 - Be aware of the registration burden for hospitals
 - Indicators on processes in the hospital can lead indirectly to better care
 - Outcome indicators are of high value and should be measured when registry data are rich and trustworthy



Thank you!



Rawa Ismail
r.ismail@dica.nl



Open Discussion

Please use the Chat feature to ask questions and make comments



Final Remarks

Co-Chair:

Ouzna Morsli, EMEAC Oncology Medical Lead, MSD



Community 365 Roundtable Meeting on Lung Cancer



Legacy from this meeting will include:

- Action report to be published in early January
- From tomorrow, video and slides on our website: europeancancer.org/resources
- Follow up with EU Commission ahead of publication Europe's Beating Cancer Plan
- Next steps on implementation of Essential Requirements in our Quality Cancer Care Network



European Cancer Summit 2020



All the sessions from this year's European Cancer Summit are now available free of charge on wondrmedical.net/ch/european-cancer-organisation

- Home
- Videos
- Topics
- Publishers
- Events

Terms · Privacy Policy · Cookie Policy
Contact Us

© 2020 All rights reserved



01:11:34

EUROPEAN CANCER

How to implement Quality Cancer Care in Europe: From good intentions to...



01:10:11

EUROPEAN CANCER

Inequalities: Disparities and Discrimination in Cancer Care



01:11:38

EUROPEAN CANCER

Early Detection Saves Lives: Taking European Cooperation on Cancer Screening...



57:05

EUROPEAN CANCER

Europe's Beating Cancer Plan

Quality Cancer Care Catalogue



- The European Quality Cancer Care Catalogue aims to provide a central repository to signpost individuals to the tools they will find helpful in improving the quality of cancer care
- The Catalogue is a continually evolving home for societies and other entities to profile and disseminate their work to broad audiences likely to have interest in their initiatives

Research Projects

Education

Standards & Accreditation

Journals

Patient Tools