

New techniques for Diagnosis & Treatment in pulmonology

Kyungjong Lee

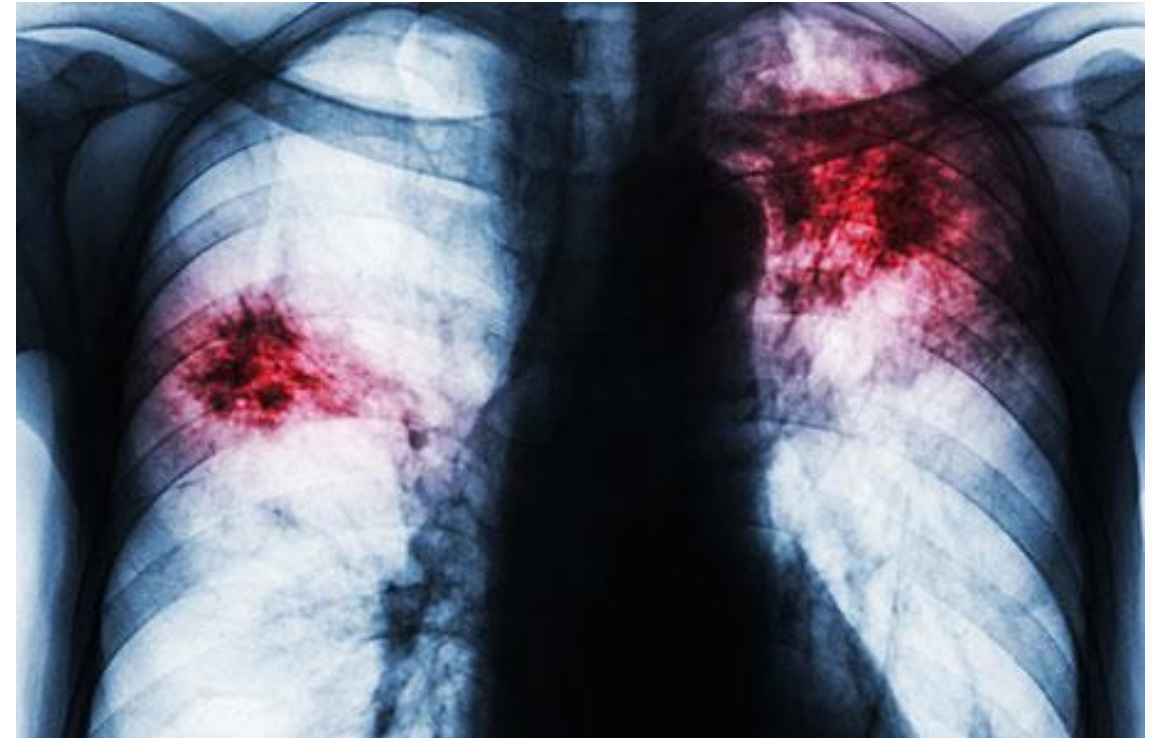
Division of Pulmonary and Critical Care Medicine, Department of Medicine,
Samsung Medical Center, Sungkyunkwan University School of Medicine



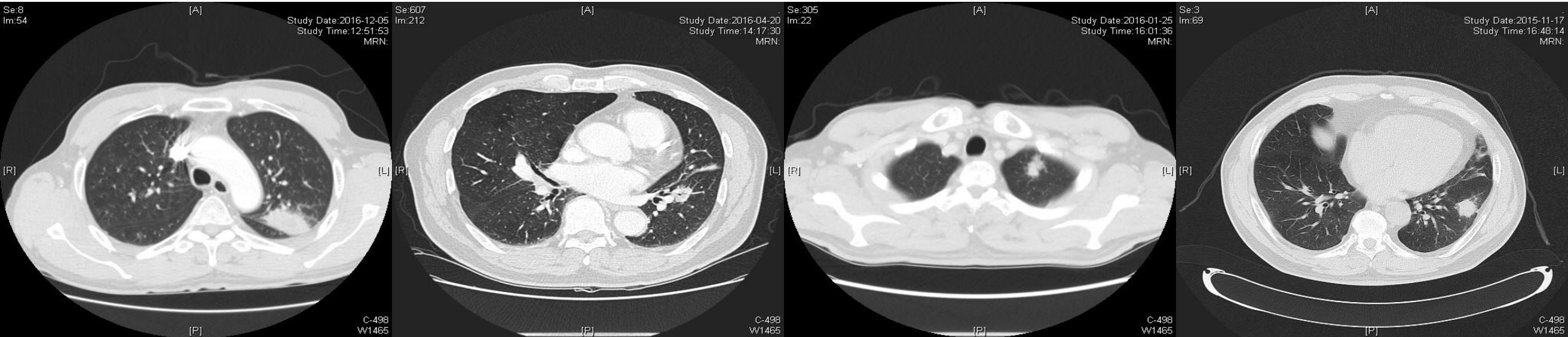
I

Diagnosis of lung nodule

Human Lung

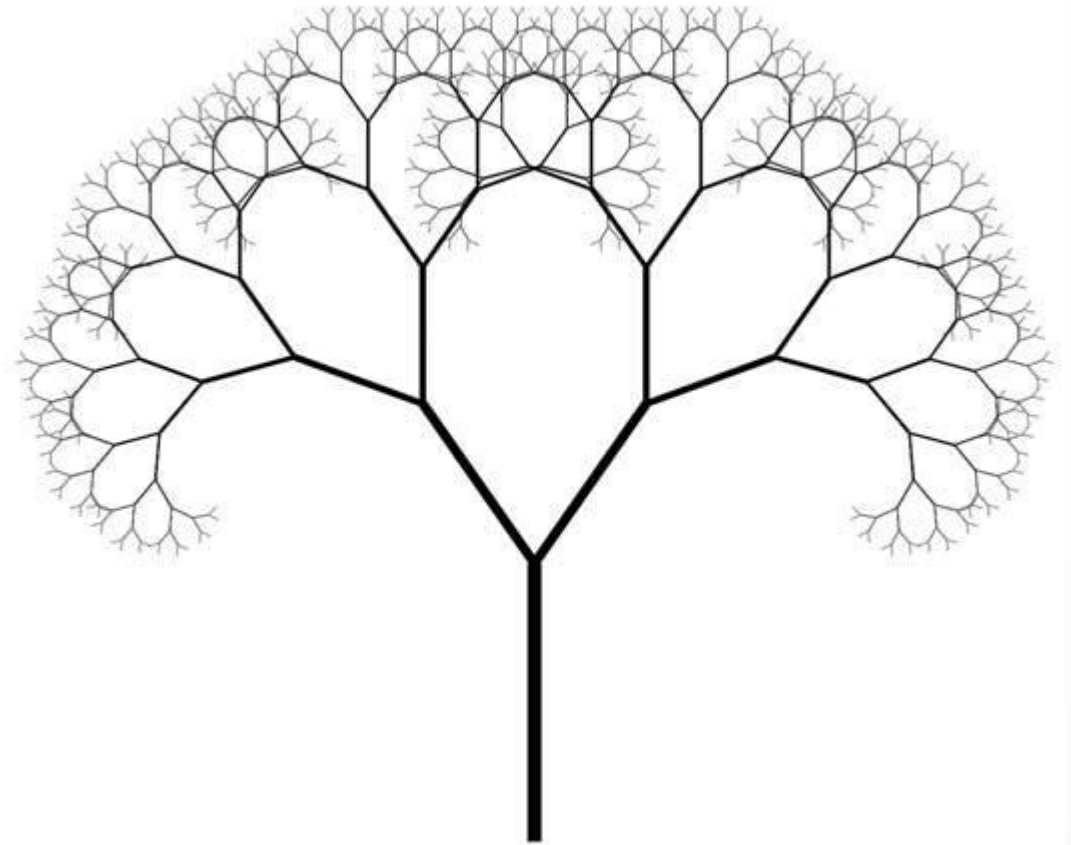
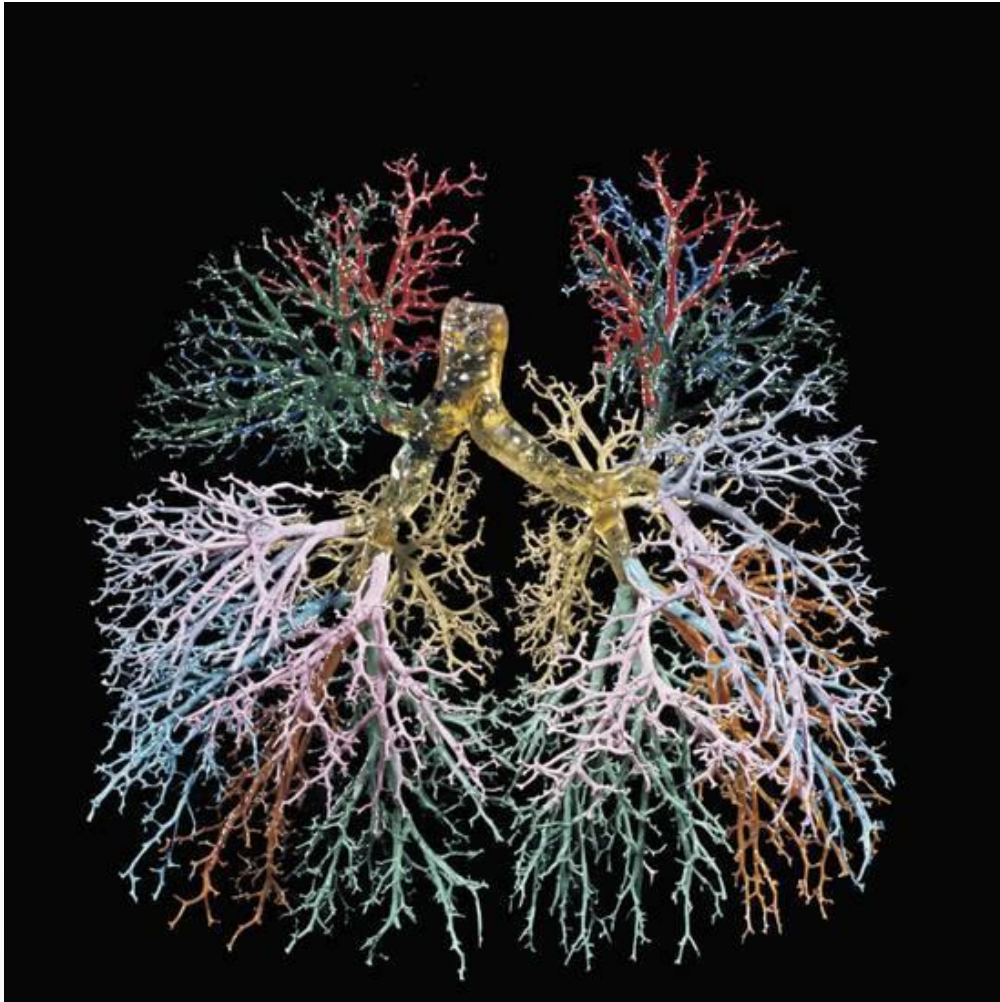


Human Lung



Chest CT에서 폐암 의심 소견을 주었으나 양성 폐질환으로 판명됨

Tissue sampling from the lung



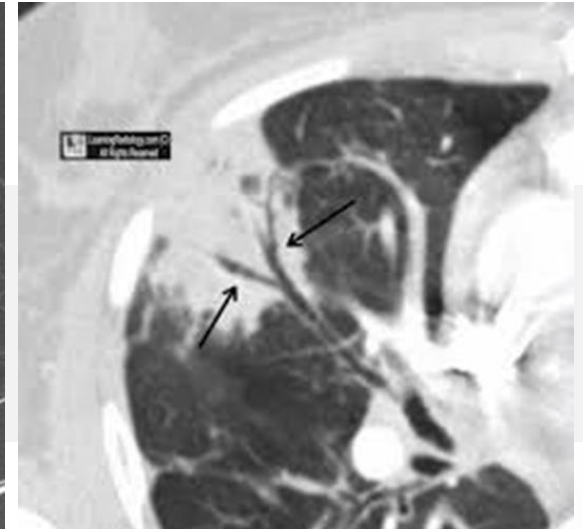
The fractal tree

(Photo: [Manuel Noah Angeja](#))

In the past...

» Percutaneous needle aspiration

- Risk of pneumothorax
- Hemoptysis
- Air embolism



» Surgical resection

- Morbidity: 4.53%
- Mortality : 1.21%

» Conventional bronchoscopy +/- fluoroscopy guided TBLB

Low diagnostic yields : 14-62%

Guided Bronchoscopy

» Radial endobronchial ultrasound

- Distance measure
- Guide sheath + Fluoroscopy

» Virtual Navigation Bronchoscopy

- Lung point (Bronchus)
- Virtual bronchoscopy navigation (Olympus)

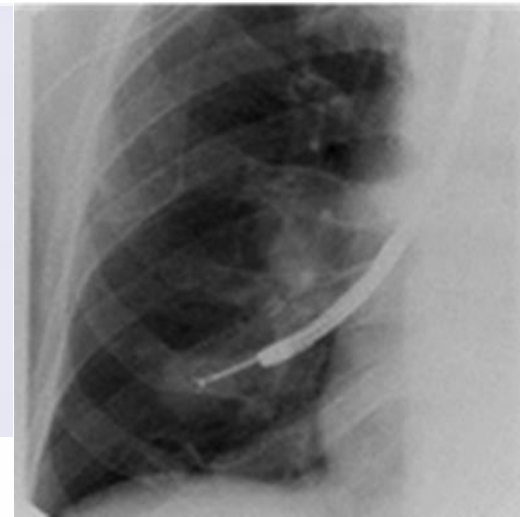
» Electromagnetic Navigation Bronchoscopy

- Super Dimension (Medtronic)
- Spin Drive (Veran Medical)

Guided Bronchoscopy

» Radial endobronchial ultrasound

Guide sheath + Fluoroscopy



Guided Bronchoscopy

» Radial endobronchial ultrasound

Distance measure method at SMC first

어떻게 하면 최소한의 장비를 가지고 폐 결절의 조직검사를 할 수 있을까?



Can Computed Tomography Characteristics Predict Outcomes in Patients Undergoing Radial Endobronchial Ultrasound-Guided Biopsy of Peripheral Lung Lesions?

Matthew Evison,*† Philip A.J. Crosbie,*† Julie Morris,‡ Julie Martin,* Philip V. Barber,* and Richard Booton*†

Introduction: Percutaneous computed tomography (CT)-guided lung biopsy is a standard minimally invasive technique for sampling peripheral lung lesions. Radial endobronchial ultrasound (EBUS) offers an alternative approach but it has yet to be defined which patients are most suited to this procedure. The primary aim of this study was to investigate whether CT characteristics could predict the success of radial EBUS-guided sampling.

Methods: The University Hospital South Manchester provides a radial EBUS service, under conscious sedation without fluoroscopy, double-hinged cassettes, or guide sheaths, to a large cancer Network in the United Kingdom. This retrospective analysis of a prospectively maintained database included all patients undergoing radial EBUS from January 2011 to June 2013. Lesion size, structure, location, and presence of a bronchus sign on thoracic CT were analyzed against predefined outcomes using multivariate analysis.

Results: One-hundred and seventeen patients underwent radial EBUS in the study period (mean age 69.5, mean lesion size 36.6 mm). The presence of a bronchus sign on CT was the only independent predictor of all predefined outcomes: (1) lesion identification with radial EBUS, (2) positioning of probe within the center of the lesion, and (3) accurate pathological diagnosis; odds ratio (OR) 31.1 (7.8–123.9, $p < 0.0001$), OR 44.8 (5.6–354.9, $p < 0.0001$) and OR 46.6 (11.1–195.3, $p < 0.0001$) respectively. The sensitivity and diag-

Key Words: Endobronchial ultrasound, Bronchoscopy, Computed tomography, Lung cancer, EBUS, EBUS-TBB, Radial probe

(*J Thorac Oncol.* 2014;9: 1393–1397)

Radial endobronchial ultrasound (EBUS), performed during flexible bronchoscopy, has gained increasing acceptance as a feasible method of locating and facilitating sampling of peripheral lung lesions. The associated literature supported the publication of an Interventional Procedure Guidance by the National Institute for Clinical Excellence in 2010 (N2124) and a meta-analysis of 1420 patients from 16 studies in 2011.¹ The favorable safety profile of radial EBUS-guided sampling is a consistently reported advantage over percutaneous computed tomography (CT)-guided biopsy where the risk of pneumothorax is up to 25%.^{2–4} The complication rate from radial EBUS is considerably lower including a risk of pneumothorax of approximately 1%.^{5–10} In contrast, meta-analysis report a higher pooled sensitivity for percutaneous CT-guided biopsy compared with radial EBUS; 90% versus 73%, respectively.^{1,11}

Patient selection and the place of radial EBUS in the diagnostic pathway have not been clearly defined. The techniques and environment used by expert groups around the

anesthetic support. Endobronchial disease is excluded using a standard diagnostic bronchoscope (Olympus BFF260 or BF6C260) before the selective cannulation of the target lobe (and subsegmental airways) using a 1.7 mm 20 MHz radial ultrasound probe. If the target lesion is identified, the distance from the lesion to the bronchoscope is measured using the same method previously described by Fuso et al.⁹ The radial probe is removed and biopsy forceps introduced into the same subsegmental bronchi and biopsies taken at the appropriate distance from the bronchoscope. Fluoroscopy, guide sheaths, and double-hinged cassettes are not used. Rapid on-site evaluation of specimens is unavailable. A minimum of five biopsies per lesion are taken assuming safety, patient comfort and tolerance. A post procedure chest x-ray is only performed if: (1) a patient reports any symptoms suggestive of a pneumothorax including breath-

this study was to prospectively evaluate which factors, based on pre-procedure CT imaging, were associated with: (1) lesion identification, (2) location of the probe within the center of the lesion and (3) increased diagnostic accuracy in a population of patients undergoing radial EBUS without any adjuncts (fluoroscopy, cassettes, guide sheaths or general anesthesia).

MATERIALS AND METHODS

The University Hospital of South Manchester provides the only radial EBUS service within Manchester Cancer, a large cancer Network in the United Kingdom that includes a regional cancer center. Respiratory physicians, thoracic surgeons, medical and clinical oncologists can all refer patients for radial EBUS. The accepted indication for radial EBUS is a target pulmonary lesion, located distal to the central airways unlikely to be visible with standard diagnostic bronchoscopy.



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Lung Cancer

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



Role of ultrasound-guided transbronchial biopsy in the diagnosis of peripheral pulmonary lesions^{☆,☆☆}

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Guide sheath

ABSTRACT

Introduction: Endobronchial ultrasound (EBUS) can be used as an alternative to fluoroscopy to visualize a peripheral pulmonary lesion (PPL) and to provide an image guidance for transbronchial biopsy (TBB). The aim of this study was to verify the accuracy of EBUS-guided TBB in the diagnosis of PPLs.

Methods: All the patients with CT-scan evidence of PPL who underwent bronchoscopy with EBUS in the period between 2008 and 2011 were retrospectively evaluated. EBUS was performed using a radial-type miniature ultrasound probe. Once obtained an EBUS image of the PPL, we measured the distance of the PPL from the outer orifice of the working channel of the bronchoscope in order to perform TBB at PPL site.

Results: A total of 662 patients were examined. The mean diameter of lesions was 36 ± 20 mm. PPLs were visualized in 494 patients (75%) and the TBB was performed in 479 patients. Thirty-two patients were lost in follow-up and data from 447 patients were analyzed. TBB results were 255 cancers and 192 non-malignant lesions. The final diagnosis reported was 359 cases of cancer and 88 of benign lesion. EBUS-guided TBB had a sensitivity of 71% for the diagnosis of cancer, a negative predictive value of 46% and an overall diagnostic accuracy of 77%.

Conclusions: These data obtained from a large series of patients and using an original method show that EBUS represents a valid support to bronchoscopy and that the EBUS-guided TBB has a high diagnostic yield in the diagnosis of PPLs.

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1. Introduction

A peripheral pulmonary lesion (PPL) can represent a real diagnostic challenge. Solitary nodules are visualized in the lung parenchyma of asymptomatic subjects with increasing frequency because of a larger use of imaging procedures such as CT-scan. Although lung cancer might be suspected on some radiological features of the lesion, a histological diagnosis is often mandatory

[☆] Dr. Fuso is the guarantor of the paper; he had full access to all of the data in the study and takes responsibility for the integrity of the work and the accuracy of the data analysis.

^{☆☆} Ongoing early results of this work have been presented as poster at ERS congress, Amsterdam, 27th September 2011.

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<http://dx.doi.org/10.1016/j.lungcan.2013.04.004>

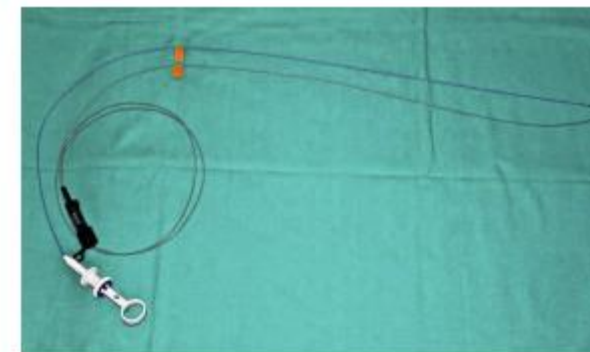


Fig. 1. Once the lesion was identified by EBUS, the distance measured on the probe was computed on the biopsy forceps.

Guided Bronchoscopy

» Radial endobronchial ultrasound

Distance measure method at SMC first
No guide sheath Kit
No fluoroscopy
Radial ebus + disposable forcep

Easy and convenient method !!



Guided Bronchoscopy

» Radial endobronchial ultrasound with distant measure



Thin bronchoscopy approach with DM

» Thin bronchoscopy

4.0 mm outer diameter

보통 4차 기관지 까지 접근이 가능

GS가 없이 targeting하기 위해서는 최대한 병변 가까이 접근해서 bronchus tip wedging을 시도해야 함.

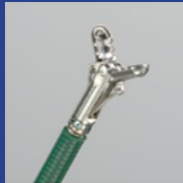
장점

방사선에 노출을 피하기 위해 납가운을 입을 필요가 없어서 몸이 매우 가볍다.

시술을 준비하는 시간이 안걸린다.

Radial probe handling이 매우 쉽다.

1.8 mm disposable forcep 사용
매우 큰 조직 채취가 가능

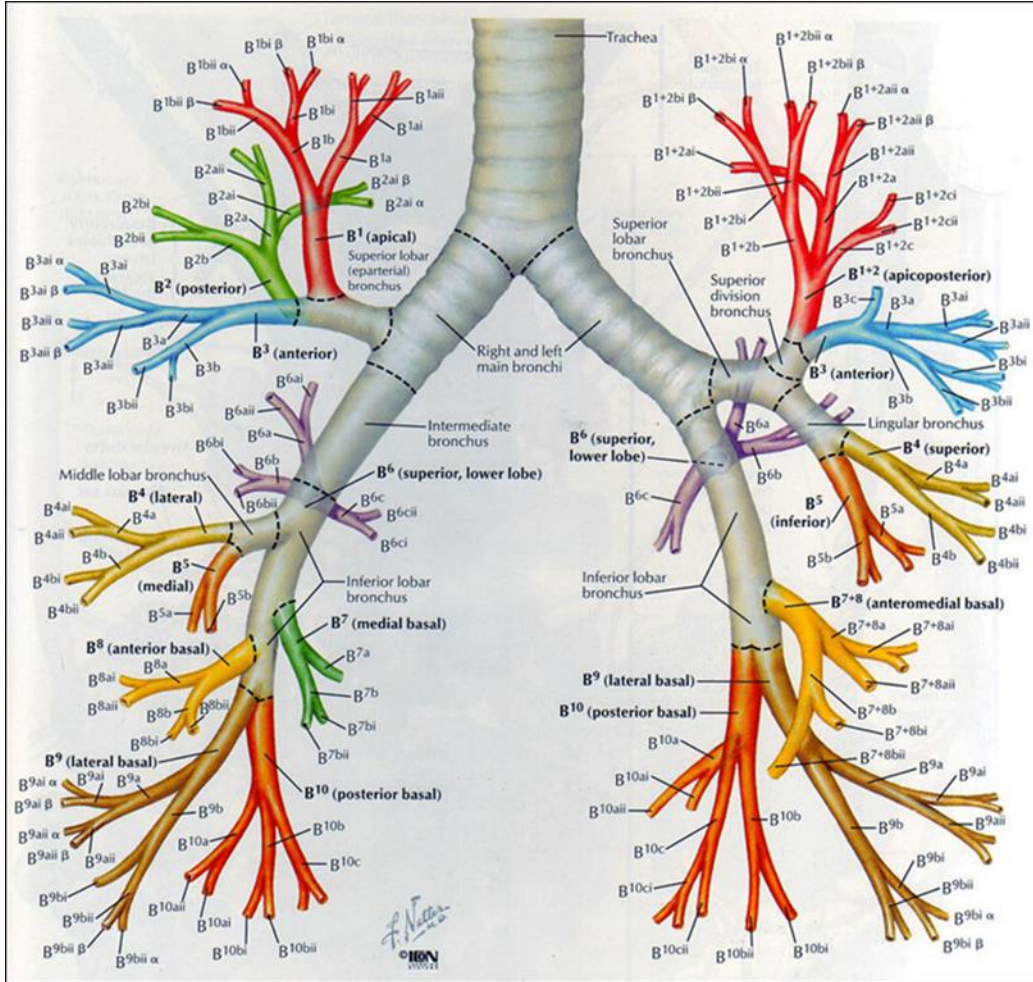


단점

조직검사후 출혈로 시야 확보가 어려움

기침을 하는 경우 내시경 wedging이 안되어 쉽게 target bronchus를 이탈함.

Mapping of Bronchus



1968년 일본 기관식도 학회

II차 기관지 : 1, 2, 3

III차 기관지 : a, b, c

IV차 기관지 : i, ii

V차 기관지 : α, β, γ

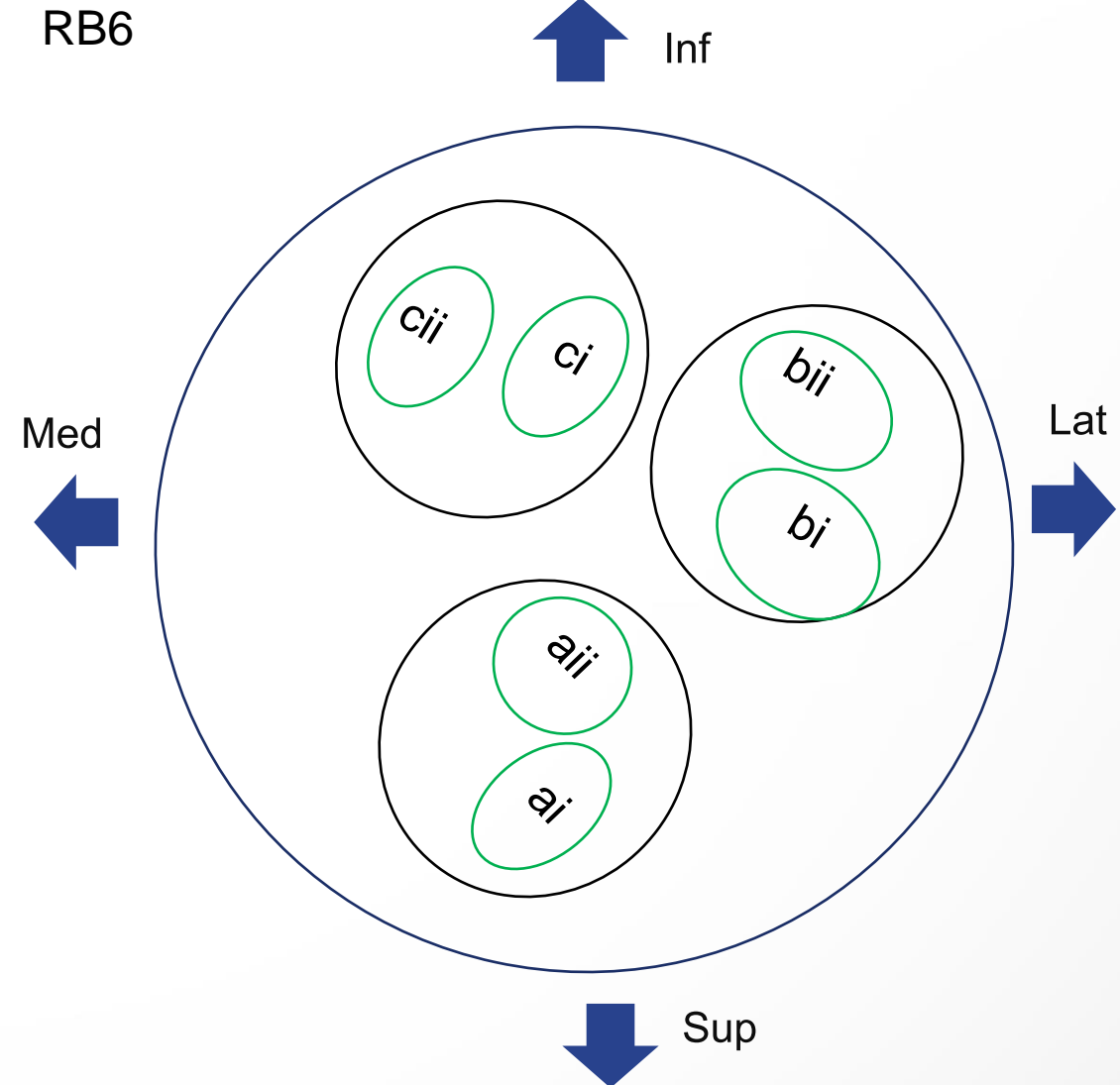
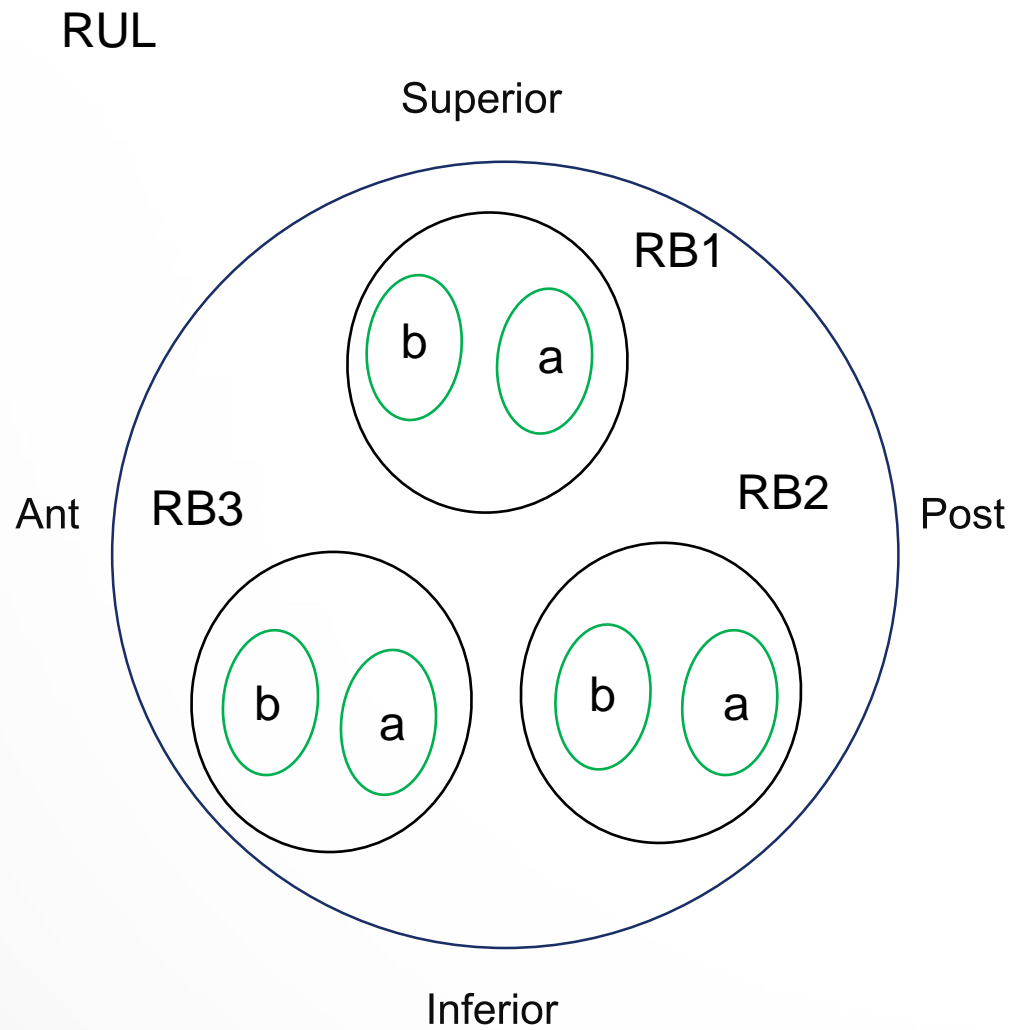
Lateral → Medial

Superior → Inferior

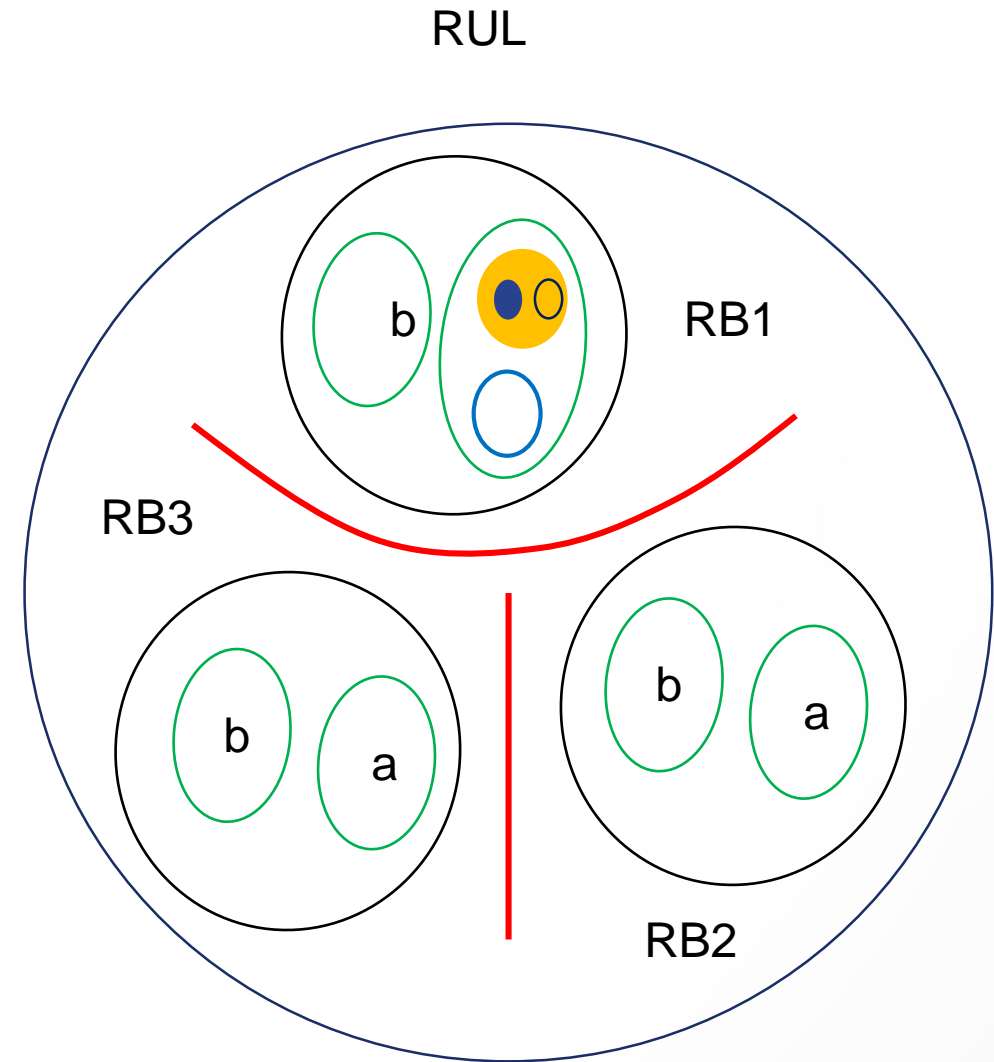
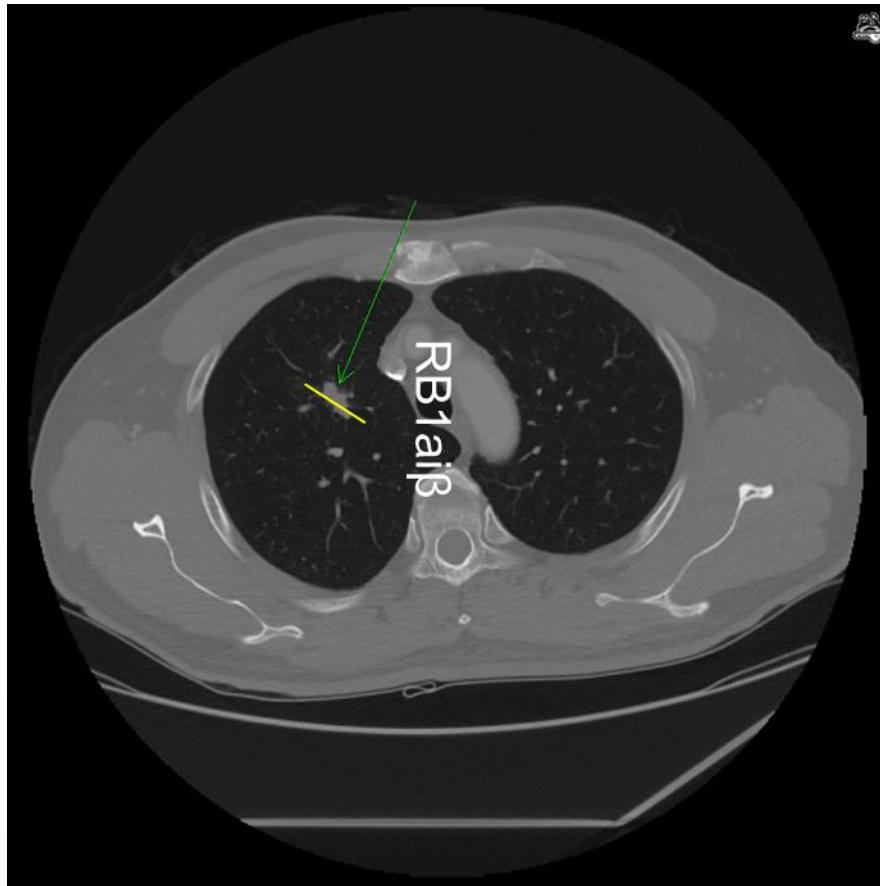
Posterior → Anterior

순으로 명명을 함

Mapping of Bronchus



Mapping of Bronchus



Case

46/F

러시아 병원에서 검사후 내원

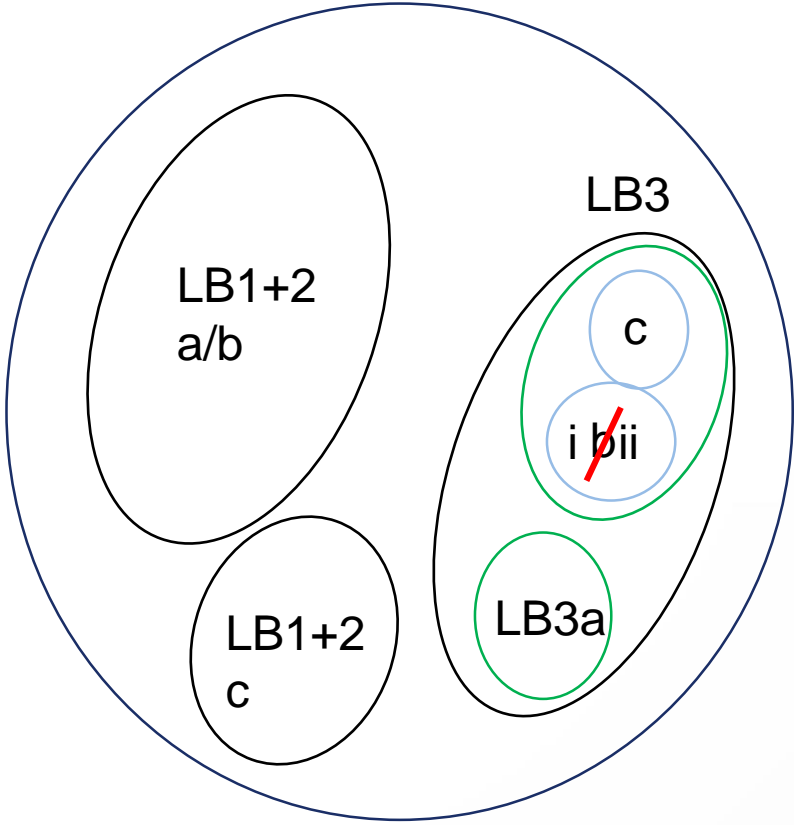
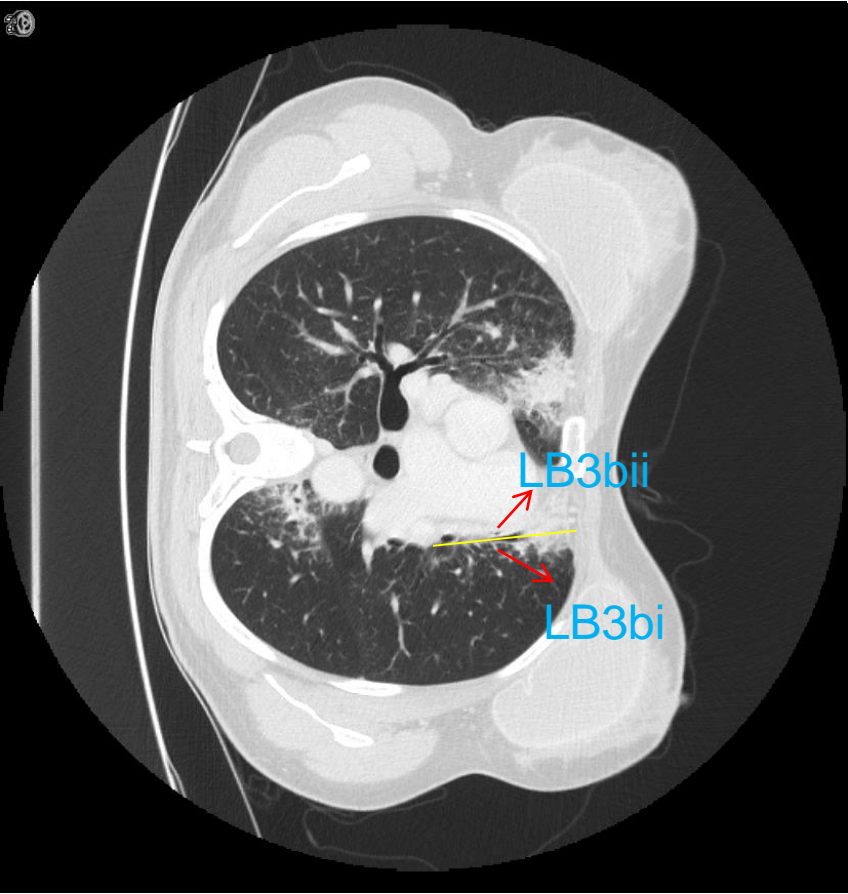
러시아 병원 진료 기록

Patch lung consolidation

TBLB : sclerotic cell



Navigation



LB1-3

Radial EBUS guided TBLB



Radial EBUS guided TBLB

» Pathology Report

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【검사명】생검 1~3개 [BP1A101] 【구분】입원
【처방일】2017-08-10 【접수일】2017-08-10 16:19 【검사일】2017-08-10 16:19 【보고일】2017-08-14 11:39
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**** 【최종보고】 ****

【판독의1】 고영혜
【검체】 Lung 【병리번호】 S1731940
■ 결론 및 진단

(PP4489)

1. Lung, LB3biibeta.#1, EBUS-transbronchial biopsy :

. Extranodal marginal zone lymphoma

<< Result of immunohistochemistry (I1719187 - 단) 2017-08-14 >>

- . Ki-67 : Positive in 5% of tumor cells
- . CD 3 T-CELL : Negative in tumor cells
- . CD 20 L26 : Positive in tumor cells

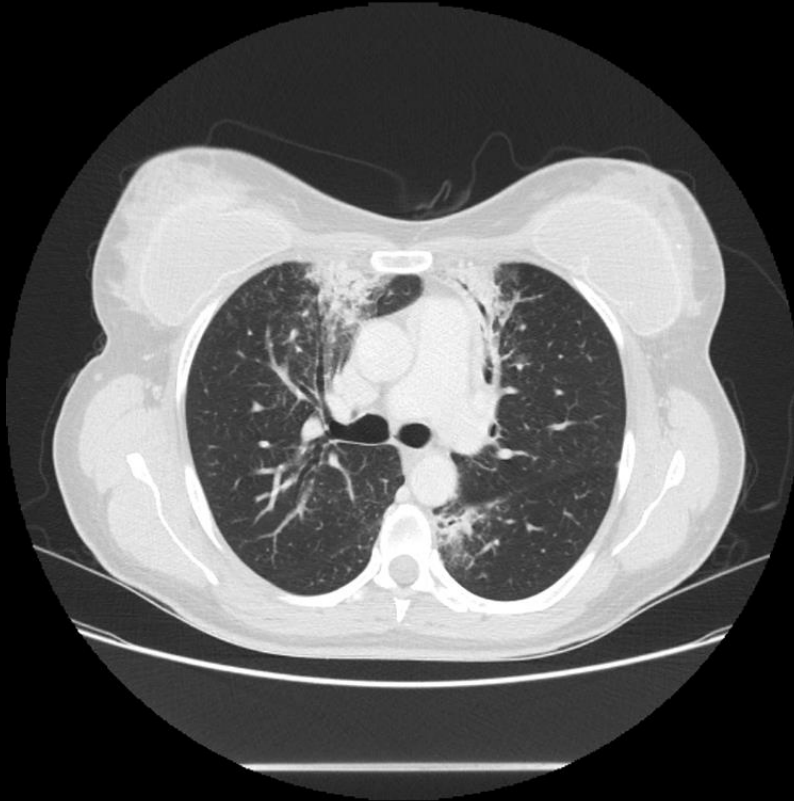
<< Result of immunohistochemistry (I1719236 - 단) 2017-08-14 >>

- . CD 5 (Leu-1) : Negative in tumor cells
- . Lambda light chain :
- . Kappa light chain :
- . Cyclin D-1 : Negative in tumor cells

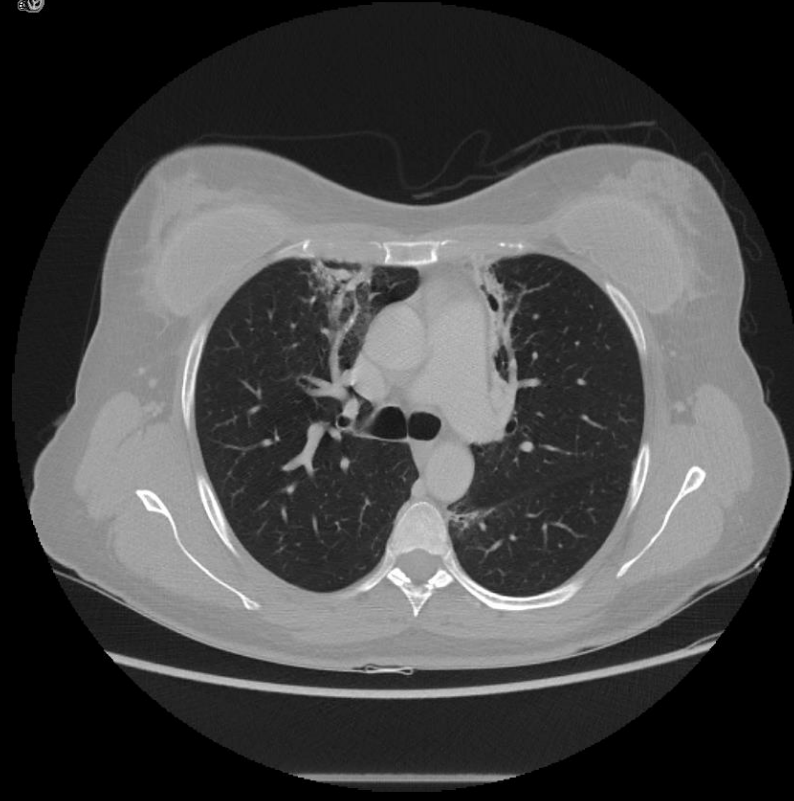
- . Kappa:Lambda=1:2, polyclonal pattern

R-CHOP#4 Chemotherapy

Before



After



Guided Bronchoscopy

» Radial endobronchial ultrasound

Distance measure

Guide sheath + Fluoroscopy

» Virtual Navigation Bronchoscopy

- Lung point (Bronchus)
- Virtual bronchoscopy navigation (Olympus)

» Electromagnetic Navigation Bronchoscopy

Super Dimension (Medtronic)

Spin Drive (Veran Medical)

Naviation Bronchoscopy



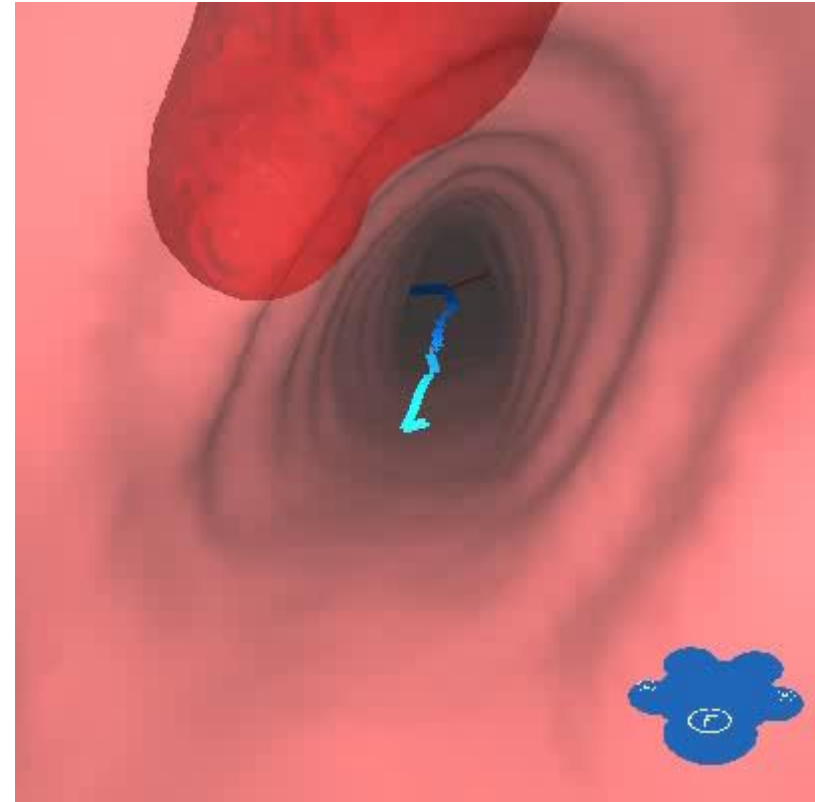
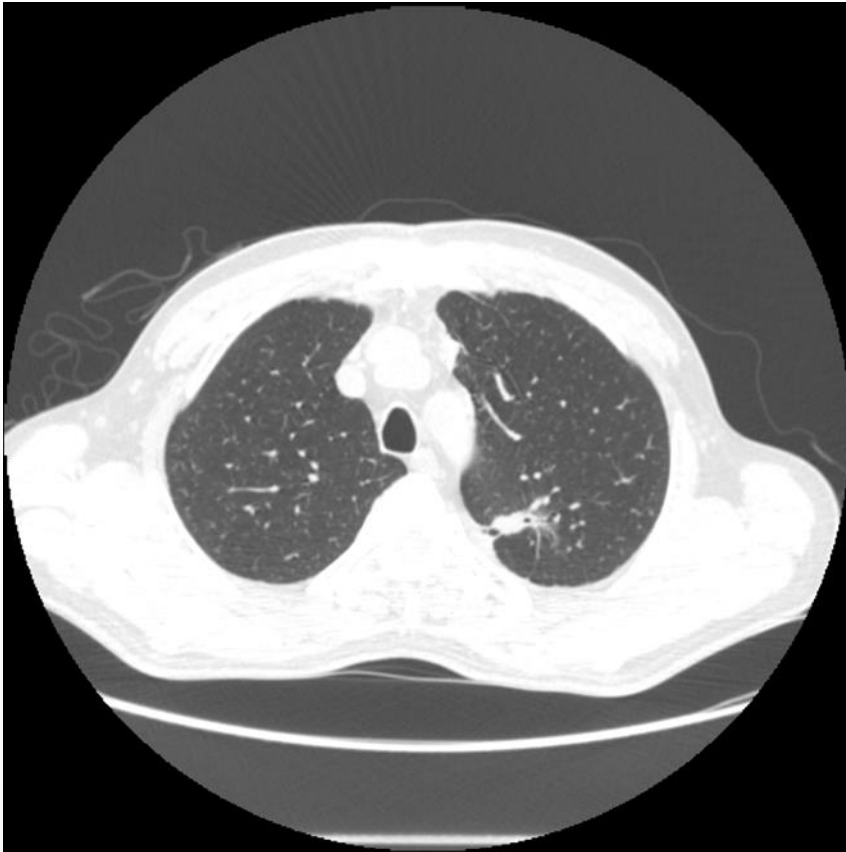
Get faster route suggestions with lifetime traffic



Virtual Navigation Bronchoscopy

» Lung Point (Bronchus)

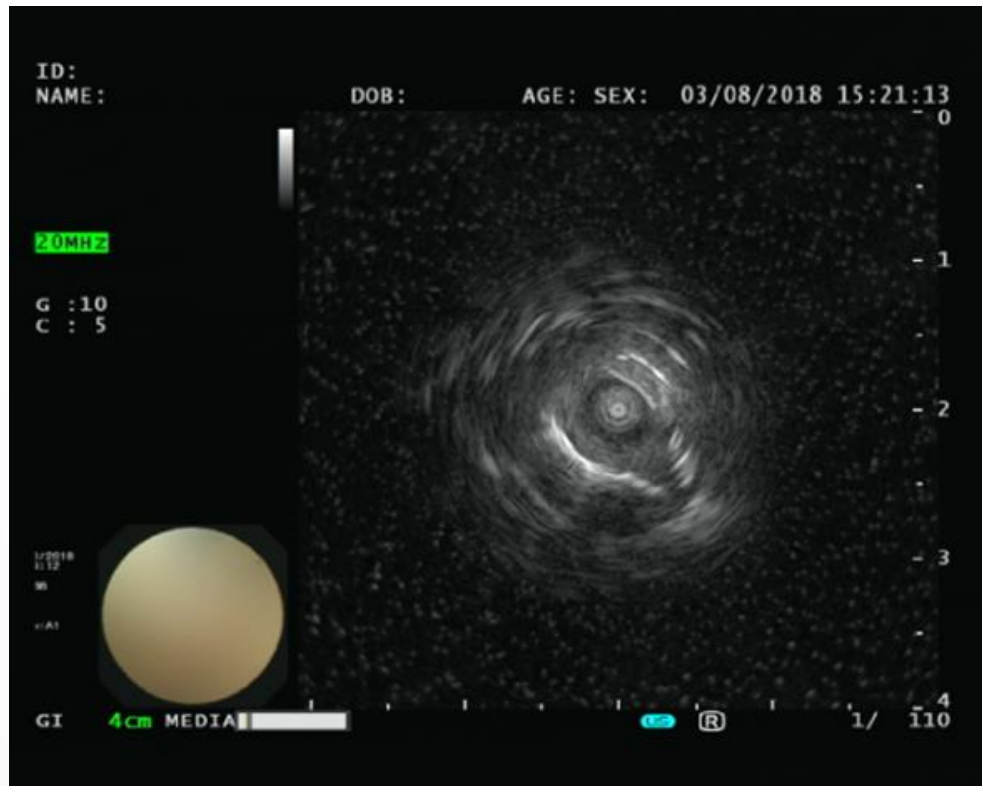
67/M 흥00



Virtual Navigation Bronchoscopy

» Lung Point (Bronchus)

67/M 홍00



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【검사명】 생검 4~6개 [BP1A102] 【구분】 입원
【처방일】 2018-03-08 【접수일】 2018-03-08 17:44 【검사일】 2018-03-08 17:44 【보고일】
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**** 【최종보고】 ****
【판독의1】 조수연
【검체】 Lung 【병리번호】 S1809978
☐ 결론 및 진단

(507011)

Lung, LUL, EBUS-transbronchial biopsy :

. ADENOCARCINOMA
```

Guided Bronchoscopy

» Radial endobronchial ultrasound

Distance measure

Guide sheath + Fluoroscopy

» Virtual Navigation Bronchoscopy

Lung point (Bronchus)

BF navigation (Olympus)

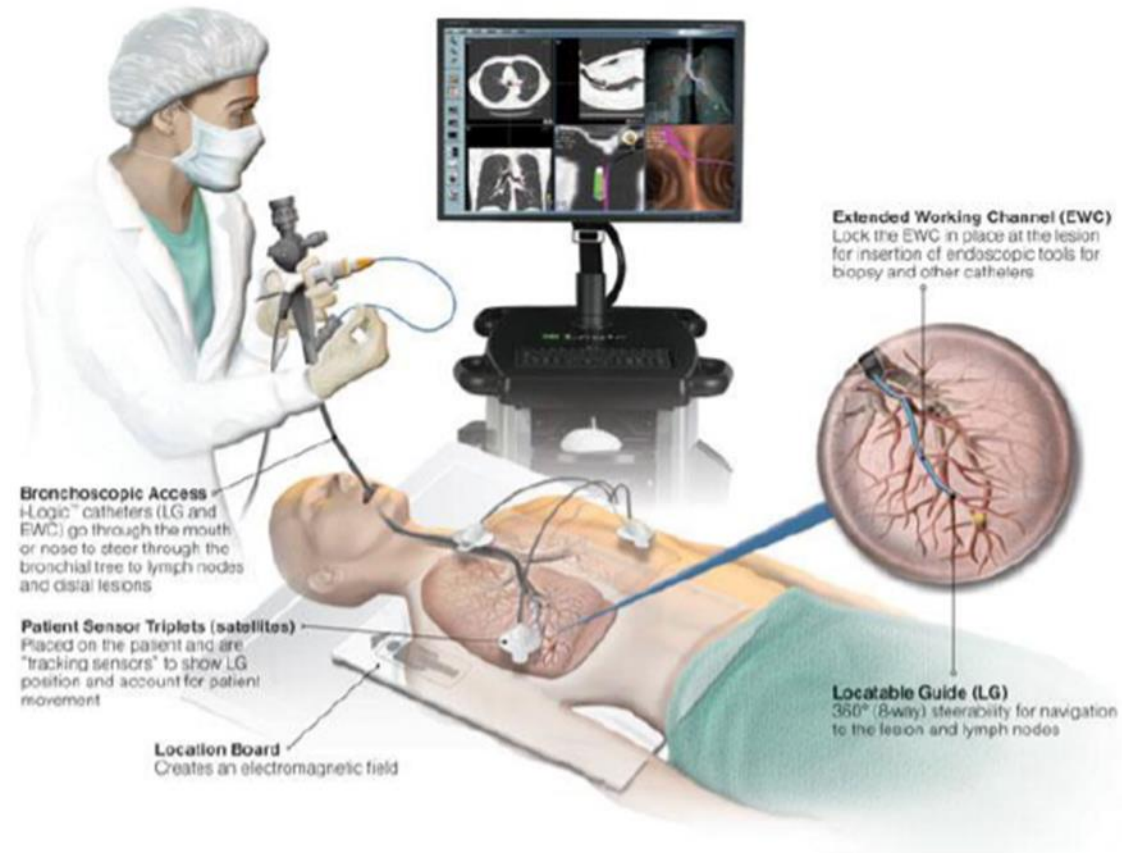
» Electromagnetic Navigation Bronchoscopy

- Super Dimension (Medtronic)

- Spin Drive (Veran Medical)

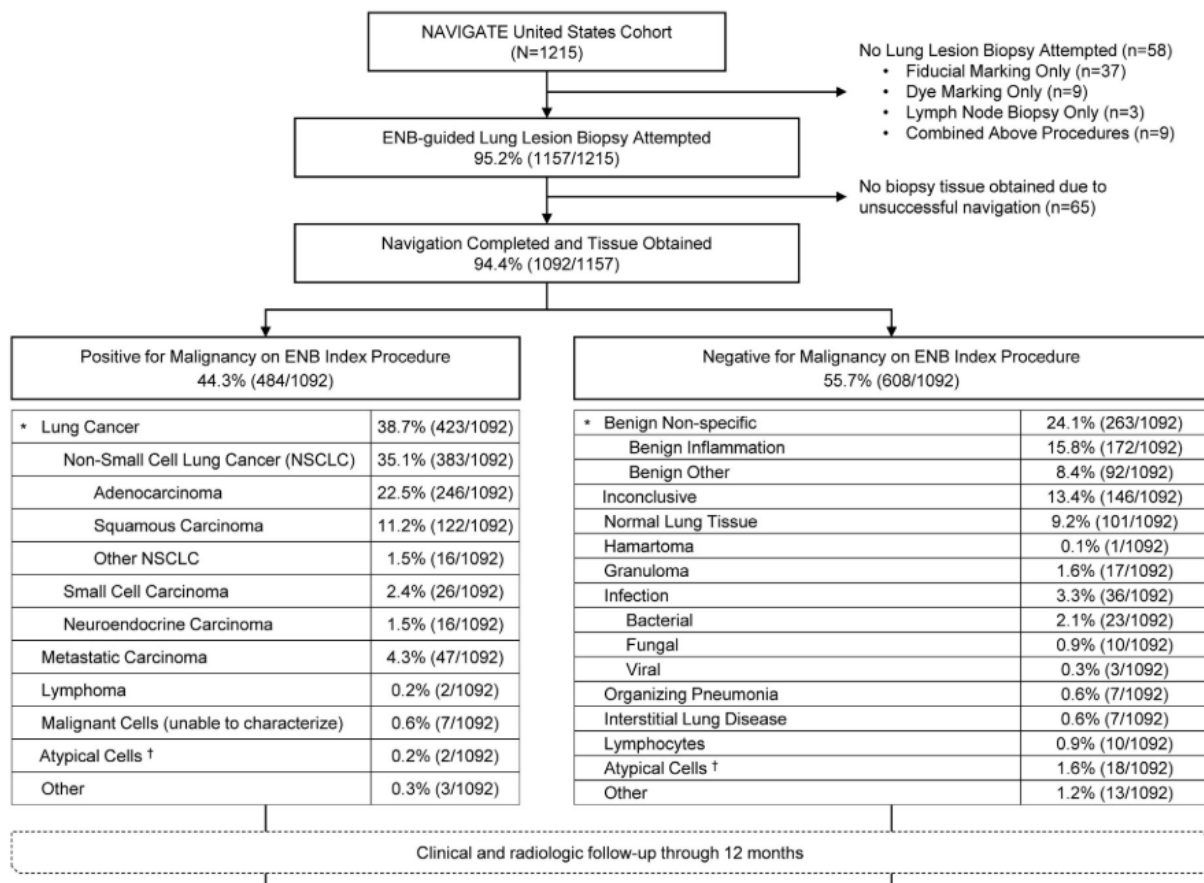
Electromagnetic Navigation Bronchoscopy

» SuperDimension, Covidien



Prospective, Multicenter NAVIGATE Study

» SuperDimension, Covidien

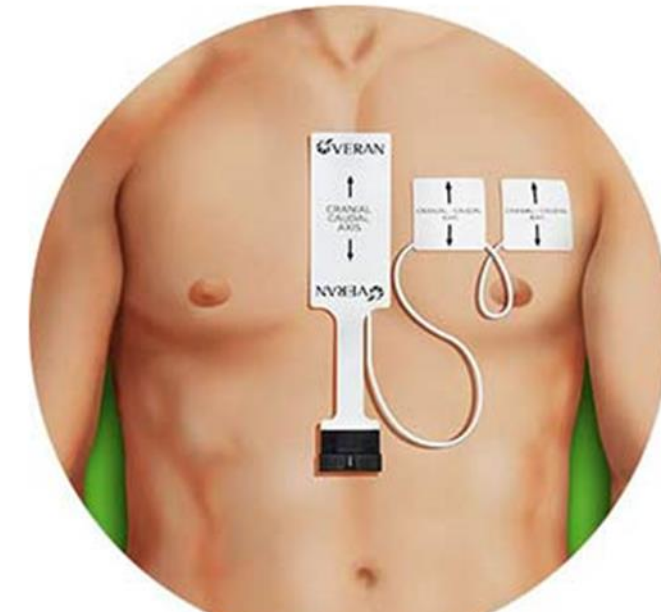
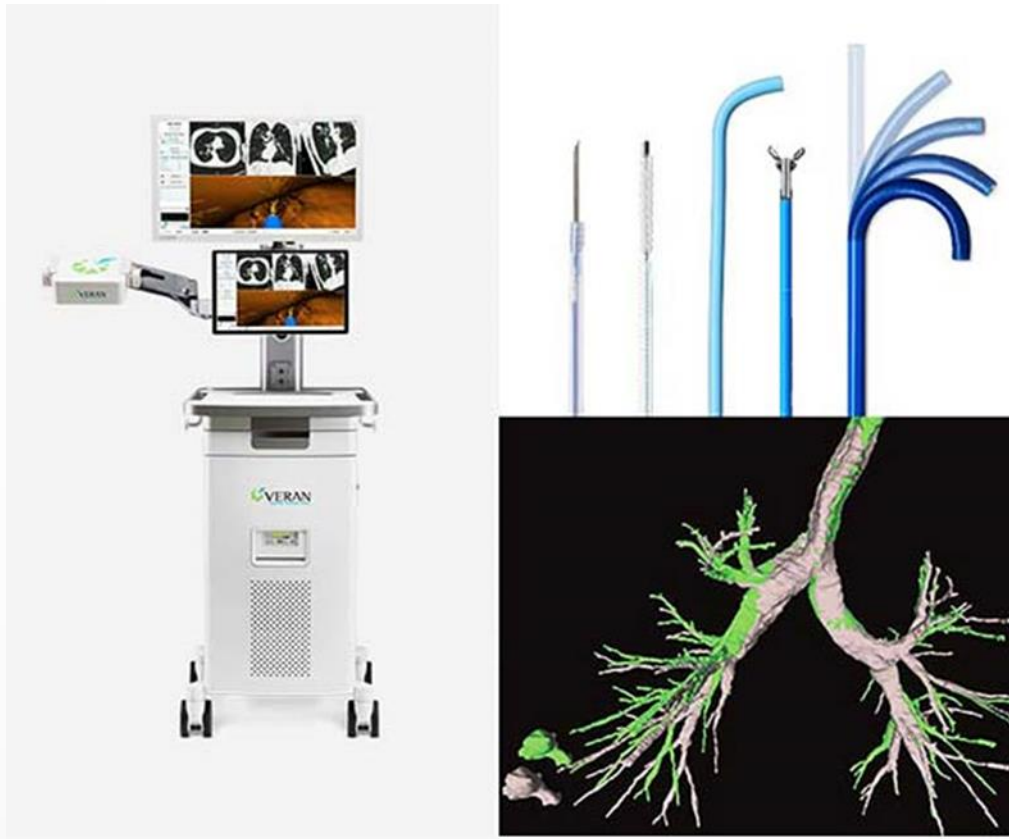


	SuperD
TP	484
FP	0
TN	284
FN	220

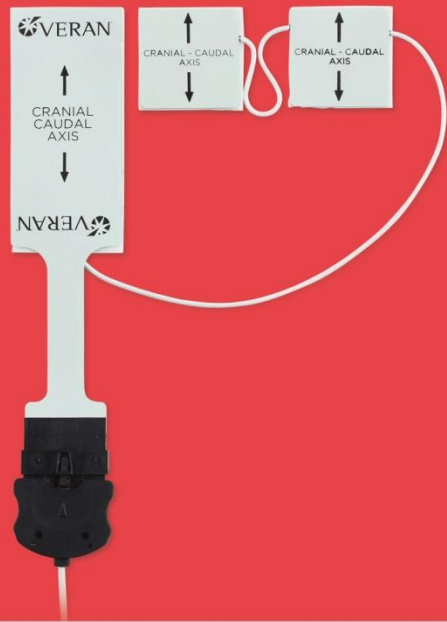
12 month diagnostic yields : **72.9%**

Electromagnetic Navigation Bronchoscopy

» Spin Drive (Veran Medical)



1: Scan



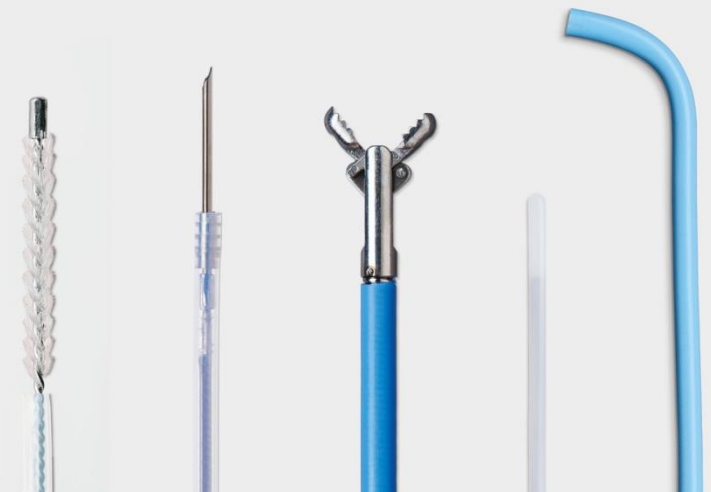
2: Plan



3: Access

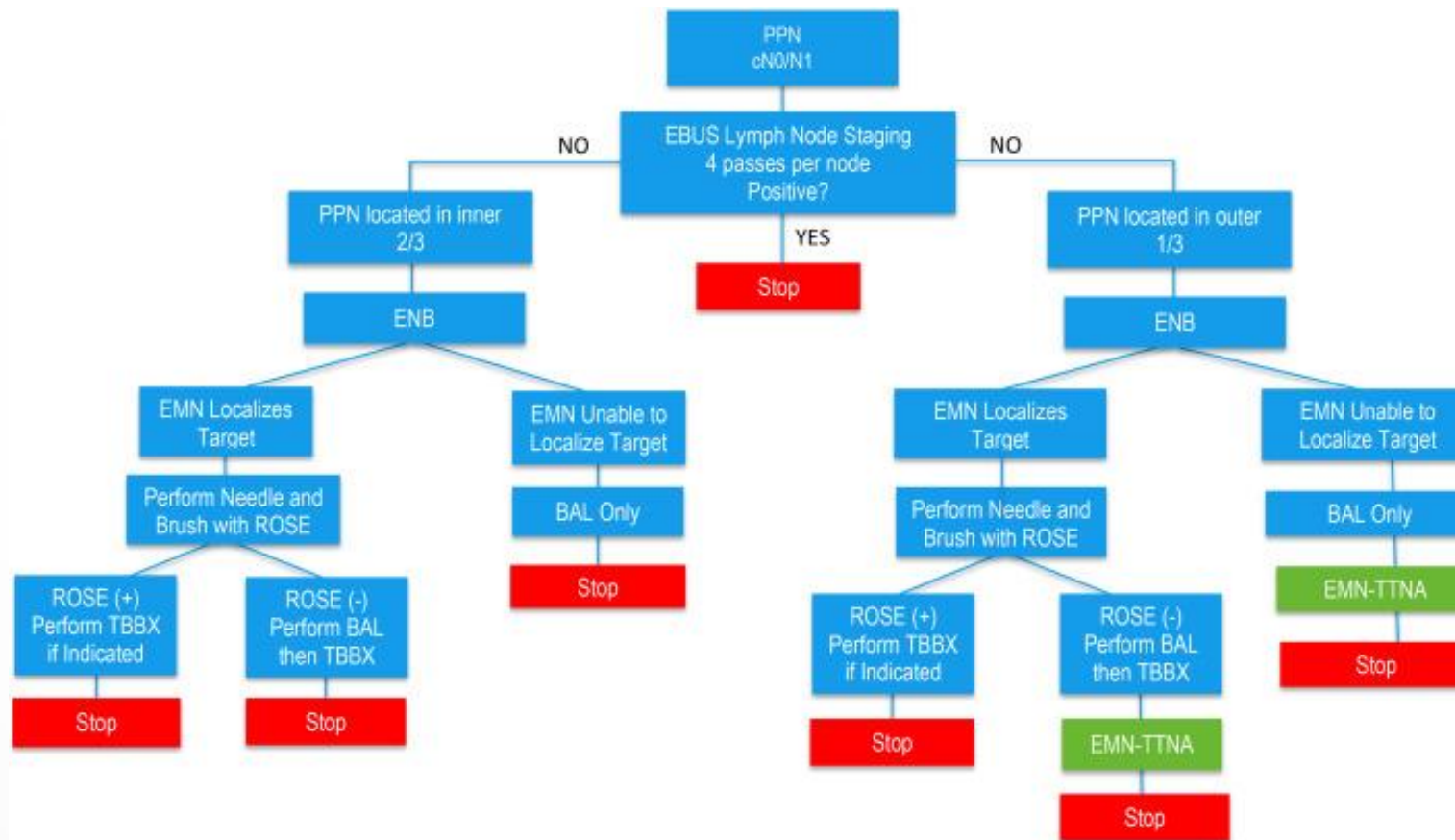


4: Biopsy and Guide Treatment



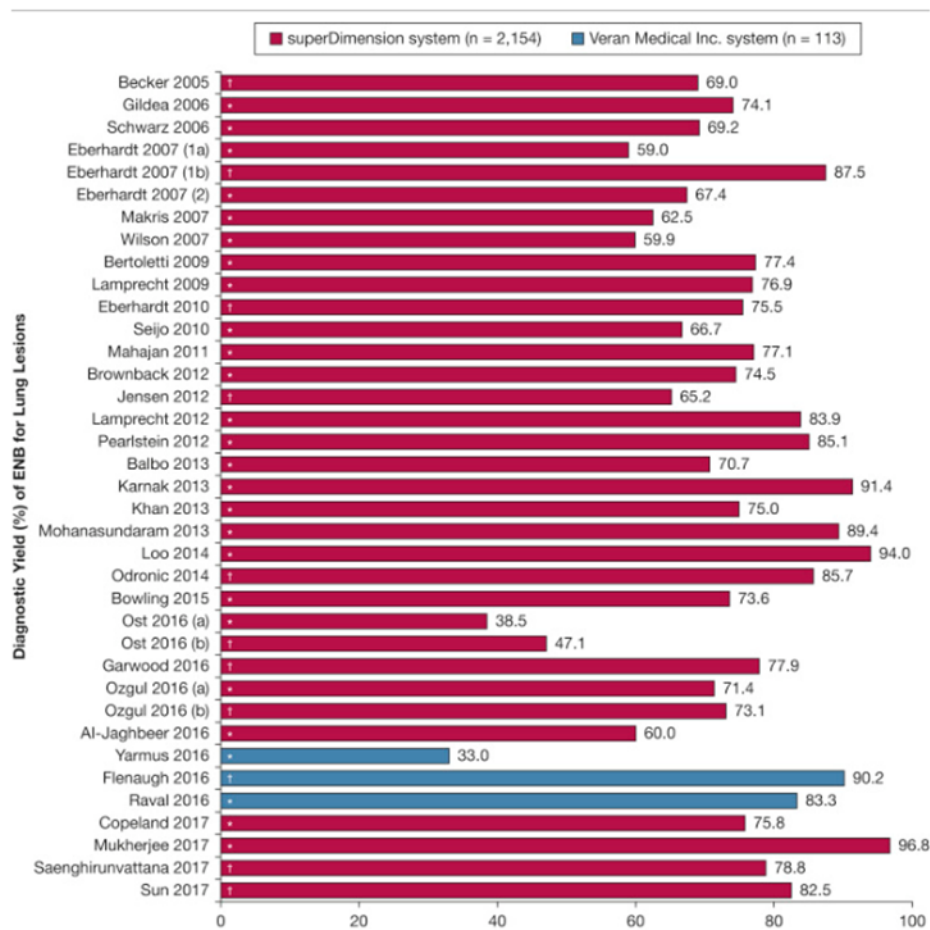
Multicenter, Prospective ALL IN ONE trial

» Spin Drive (Veran Medical)



Electromagnetic Navigation Bronchoscopy

» Diagnostic yields



	SuperD	Veran
High	96.8%	90.2%
Low	38.5%	33.0%

Guided Bronchoscopy

» Meta-analysis of Diagnostic yields

Table 2—Inverse Weighted Diagnostic Yield Overall and by Modality

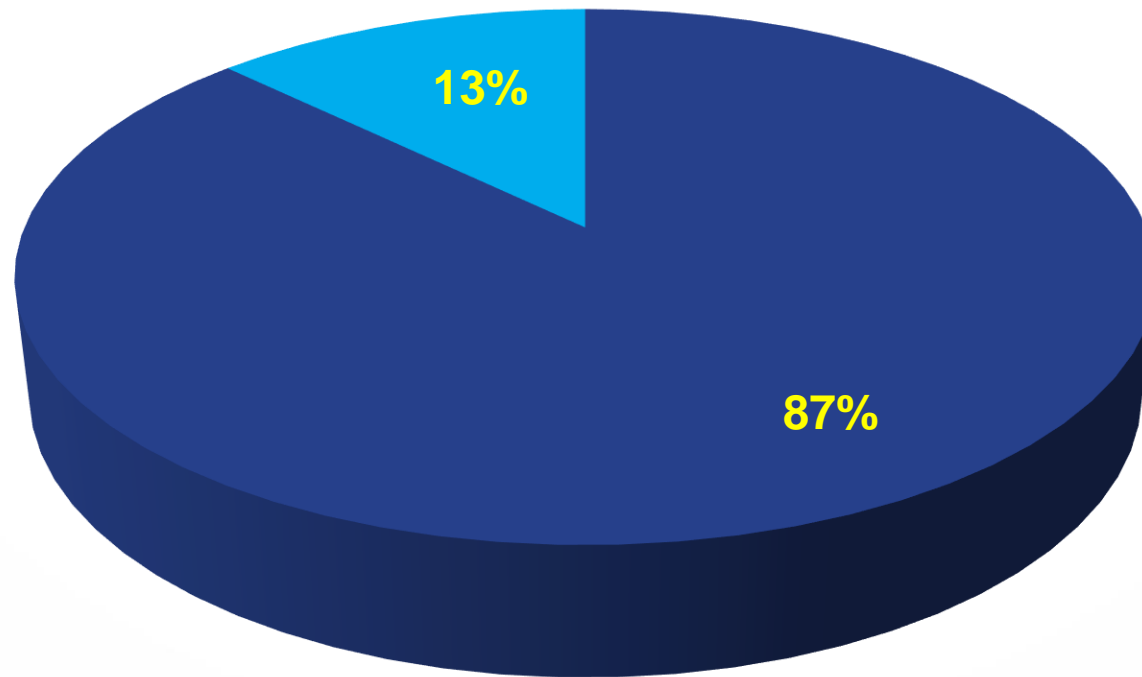
Technology	Studies, No.	Weighted Proportion, %	95% CI	Q Statistic	Q P Value
VB	10	72.0	(65.7-78.4)	21.0	.01
ENB	11	67.0	(62.6-71.4)	13.3	.21
GS	10	73.2	(64.4-81.9)	63.8	<.0001
U	11	70.0	(65.0-75.1)	15.2	.12
R-EBUS	20	71.1	(66.5-75.7)	84.2	<.0001
All	39	70.0	(67.1-72.9)	119.4	<.0001

See Table 1 legend for expansion of abbreviations.

Radial EBUS at SMC

» 2016.06 ~ 2017.05 211 cases

■ 접근 성공 ■ 접근 실패



Radial EBUS at SMC (접근 가능한 case)

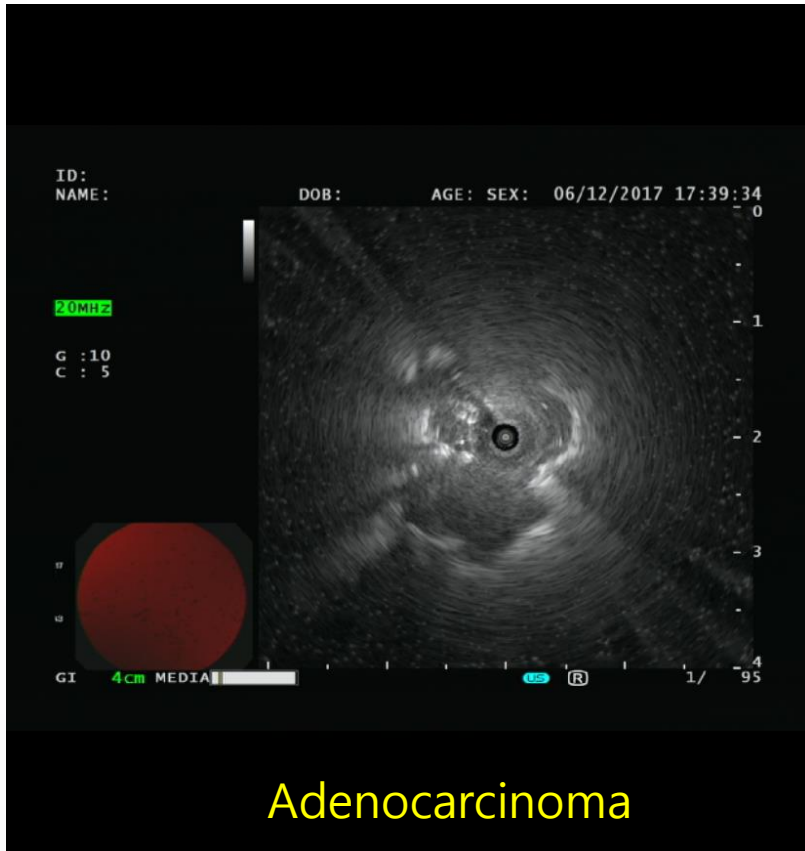
» 2016.06 ~ 2017.05 211 cases

	Final diagnosis: malignant	Final diagnosis: benign	Total	
R-EBUS: Positive for malignancy	109 True positive	0 False positive	109	Diagnostic accuracy: 80%
R-EBUS: Negative for malignancy	34 False negative	27 True negative	61	Sensitivity 76%
				NPV 44%

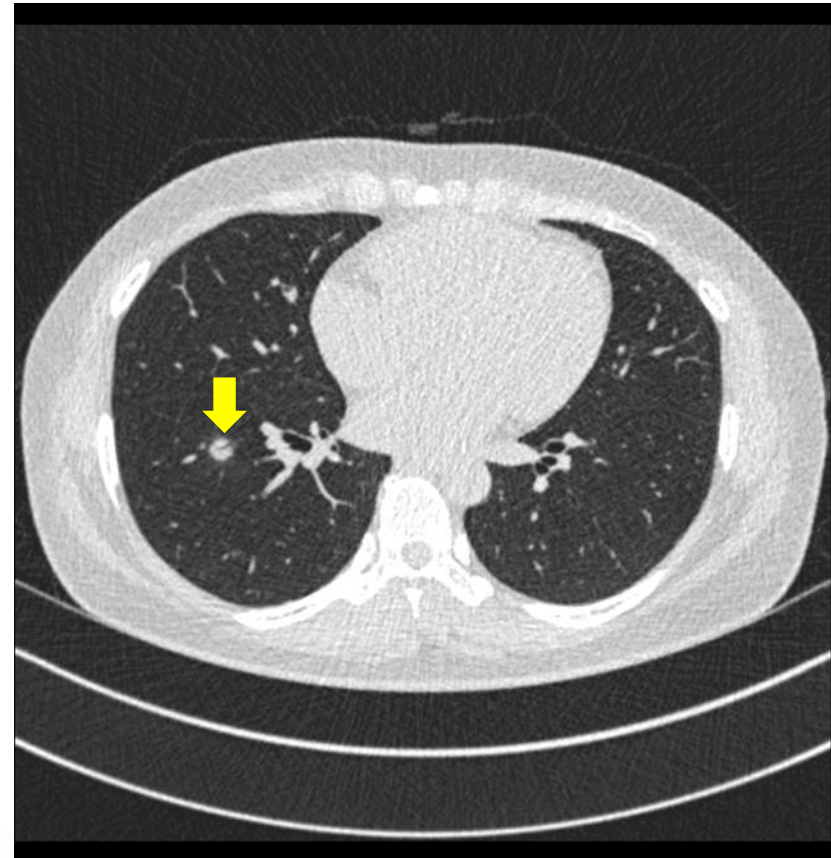
Guided Bronchoscopy

» Thin slice Chest CT의 필요성

Thin slice Chest CT가 필요함 (적어도 1mm cut)



5 mm

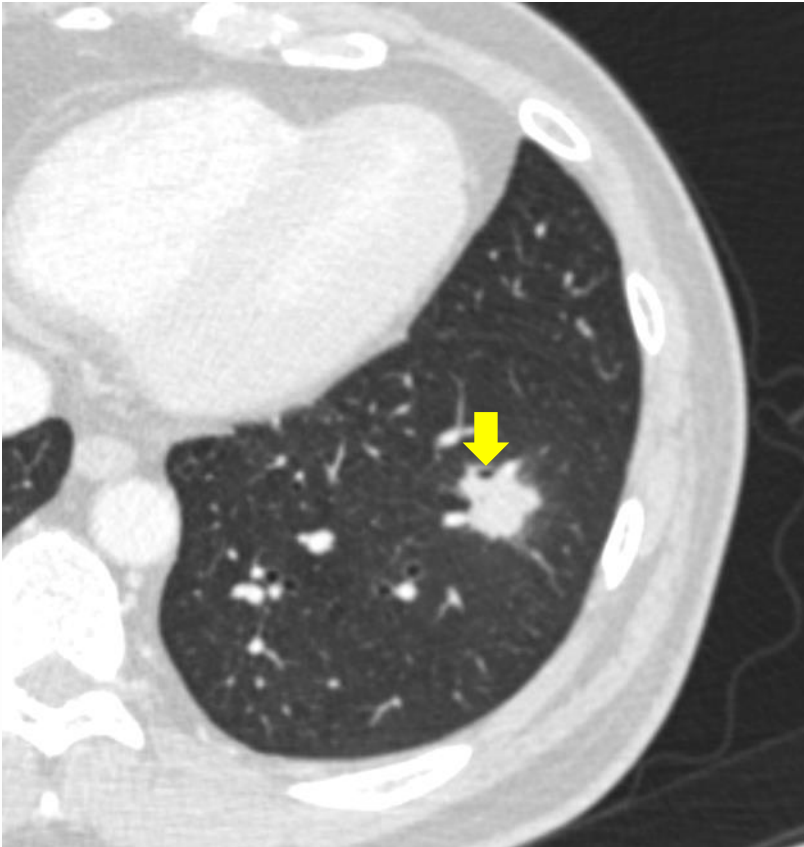


0.6 mm

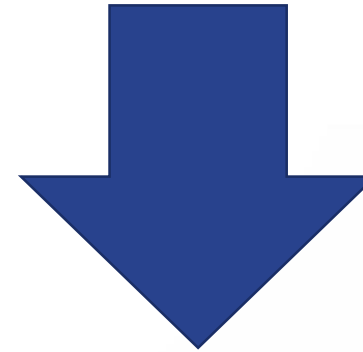
Guided Bronchoscopy

» 어떤 경우에 의뢰 되어야 하는가?

전제 조건 : 병변이 기관지 안에 있어야 함, 기관지로 접근이 가능 해야 함



진단을



Extrabronchial location하는 경우는?

» Bronchoscopic TransParenchyma Nodule Access (BTPNA)

ARCHIMEDES - TOTAL LUNG ACCESS PLATFORM



Extrabronchial location하는 경우는?

» Bronchoscopic TransParenchyma Nodule Access (BTPNA)

Lung cancer



ORIGINAL ARTICLE

Bronchoscopic transparenchymal nodule access (BTPNA): first in human trial of a novel procedure for sampling solitary pulmonary nodules

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¹Department of Pneumology and Respiratory Care Medicine, Thoraxklinik, University of Heidelberg, Heidelberg, Germany

²Translational Lung Research Center, Heidelberg, Germany

³Department of Pulmonary Medicine, University of Pennsylvania Medical Center, Philadelphia, Pennsylvania, USA

⁴Department of Pulmonary Medicine, Medical University of South Carolina, Charleston, South Carolina, USA

⁵Department of Thoracic Surgery, Thoraxklinik, University of Heidelberg, Heidelberg, Germany

⁶The NIHR Respiratory Biomedical Research Unit at Royal Brompton and Harefield NHS Foundation Trust and Imperial College, London, UK

⁷Chelsea & Westminster Hospital, London, UK

⁸National Heart & Lung Institute, Imperial College,

ABSTRACT

Introduction The promise of benefits from lung cancer screening is tempered by the false positive rate and the need to perform diagnostic procedures to determine the aetiology of the solitary pulmonary nodules (SPN) identified. We have developed a novel procedure that allows sampling of SPNs via a transparenchymal approach, and report the results from this as a first human trial.

Methods This study was a prospective single-arm interventional study. We recruited patients with a nodule detected on CT imaging, which was suspicious for cancer, who were suitable for surgical resection. On the subject's CT, an optimal airway wall point of entry (POE), and an avascular path through lung tissue from the POE to the SPN was calculated. A tunnel was created from the POE to the nodule using a set of catheter-based tools under fused fluoroscopy guidance. The patients proceeded to surgical resection immediately after the biopsy. The participants were followed-up for 6 months after the procedure. The primary endpoint of the study was to evaluate the feasibility to access and biopsy the nodule.

Results Twelve patients were recruited, and a tunnel

Key messages

What is the key question?

12 patients recruited



10 patients Success

that it is a safe approach that is technically feasible with a high success rate.



1st case in China



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阿基米德导航实例操作

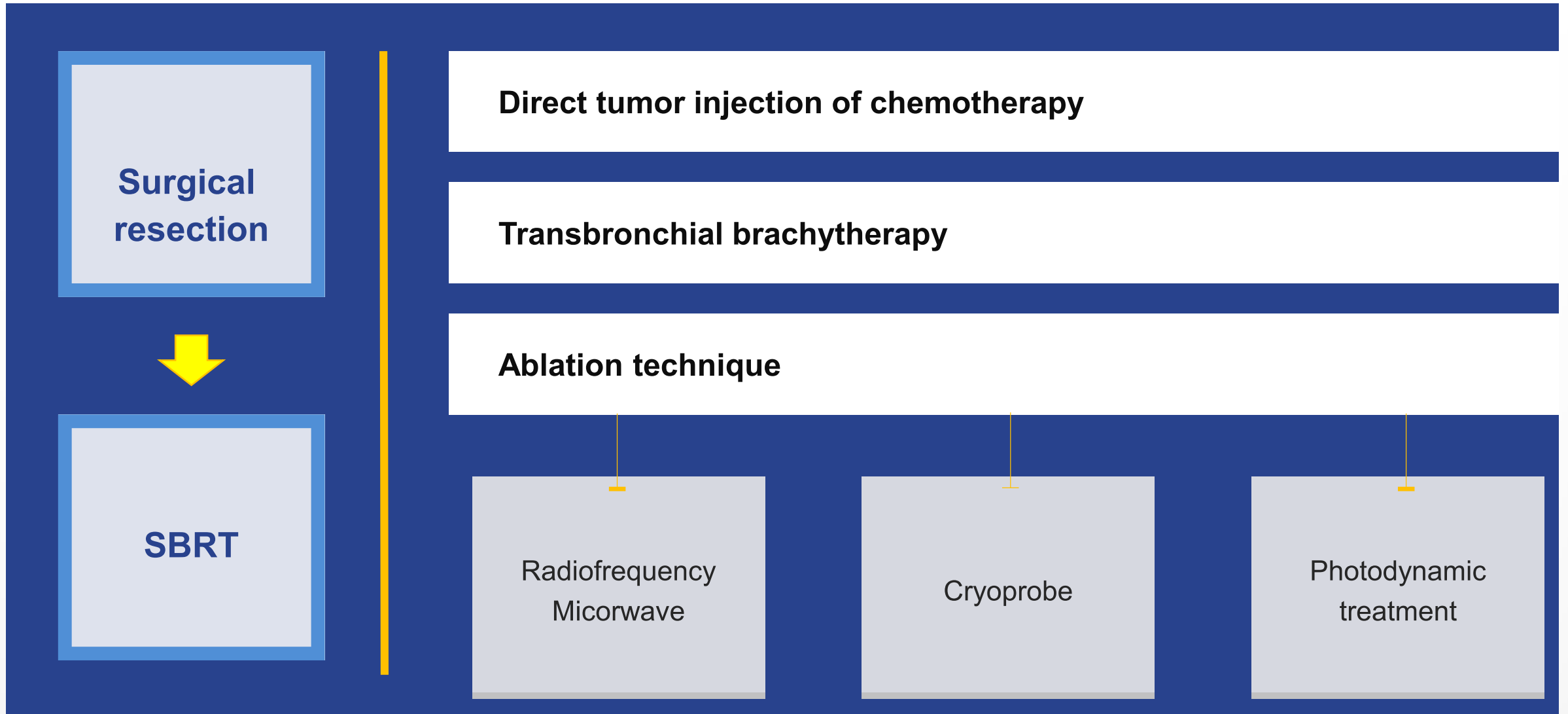
Diagnostic Bronchoscopy

	Radial ebus	Radial ebus+GS	Lung point navigation	EMN	TBA
Cost	+	++	+++	++++	NA
Pure GGO	-	-	-/+	++	-
Navigation	+	+	+++	++	++++
TBLB	+++	++	++	+ ~ +++	++
Fluoroscopy	-	++	++	-	+++
Maintenance	+	+++	NA	++	NA
Outside the bronchus	-	-	-	++	+++
Sedation	Conscious	Conscious	Conscious	Conscious/General	General

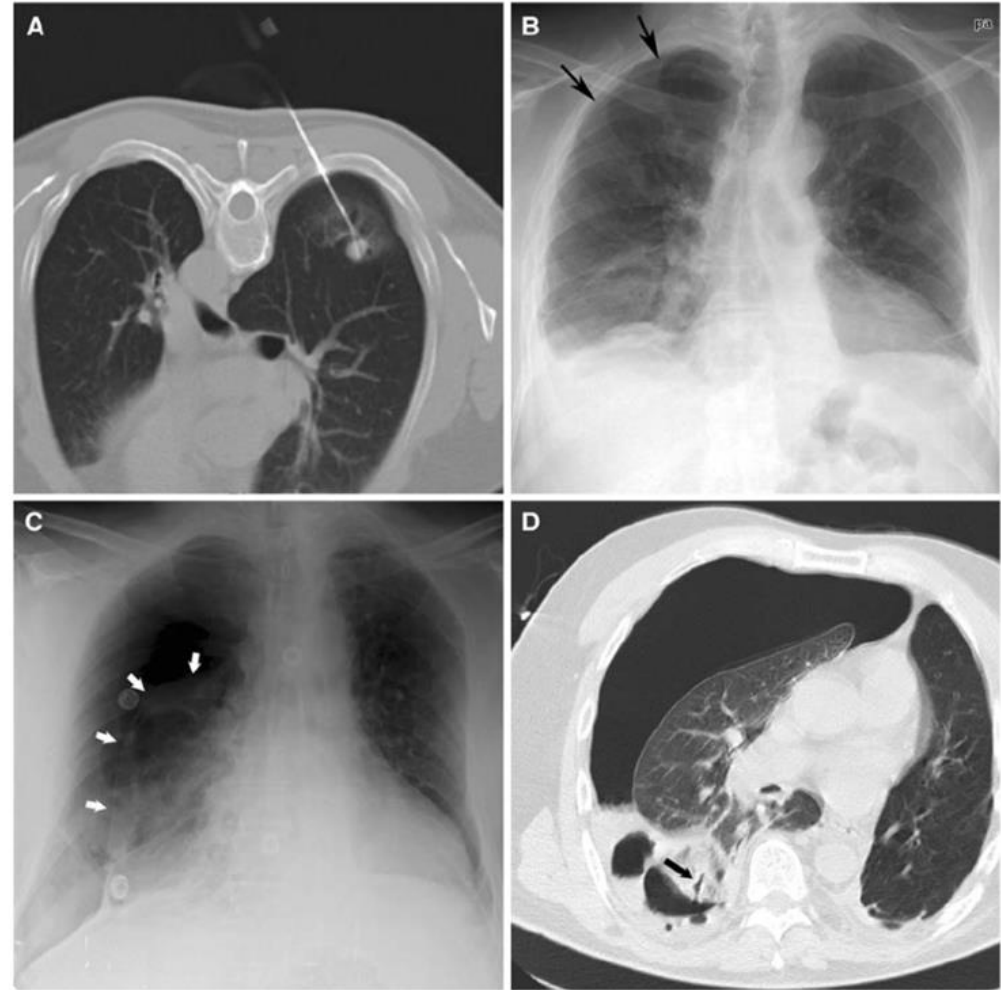
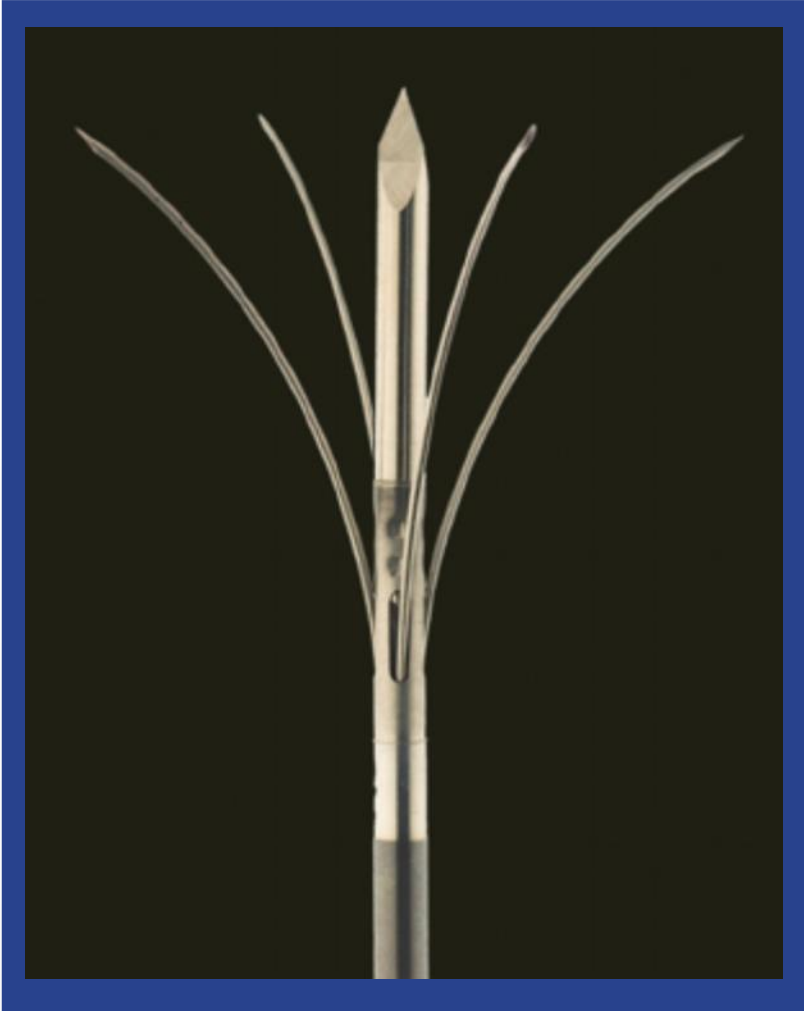
A white Roman numeral "II" centered within a white square frame.

Bronchoscopic treatment of lung cancer

Treatment of peripheral lung cancer (cT1 No Mo)



Percutaneous ablation of peripheral lung cancer



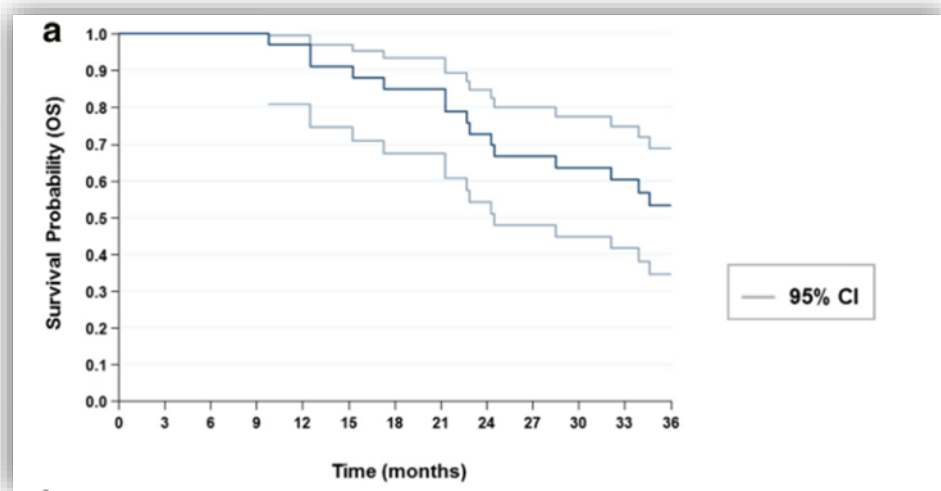
RF ablation of stage IA NSLCL

» Ineligible for surgery

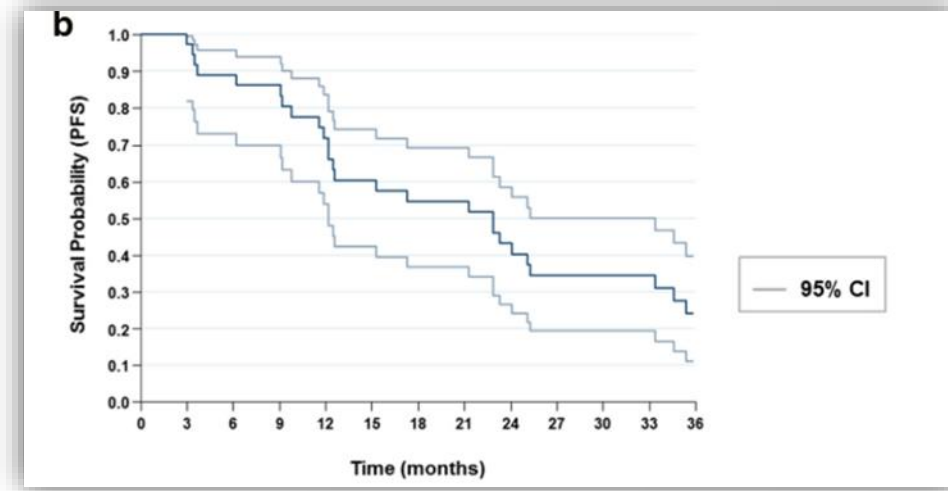
Prospective multicenter phase II trial

biopsy-proven stage IA NSCLC with a maximum tumor diameter ≤ 3 cm

Under CT-guided percutaneous ablation



1yr OS: 91.67%, 3yr OS: 58.3%



1yr PFS: 71.76%, 3yr PFS: 25%

Question for bronchoscopy guided ablation

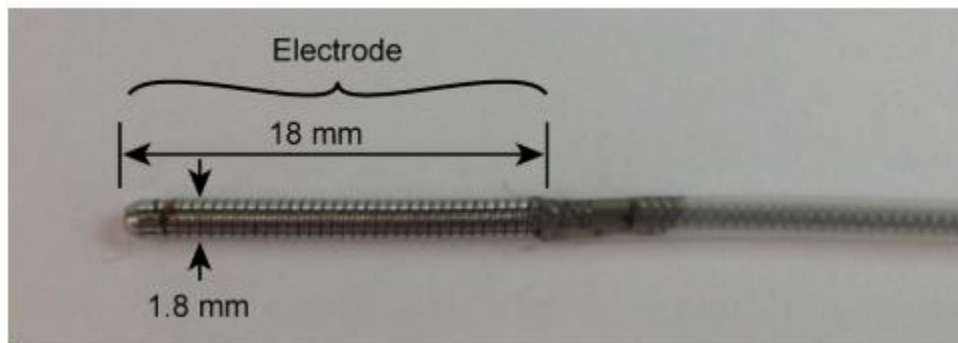
» Biopsy and ablation simultaneously

Recent advancer of bronchosocpy technique for peripheral lung nodule

To access peripheral lung nodules in combination with the simultaneous bronchoscopic ablation

→ Least invasive treatment

» New ablation device for bronchoscoy



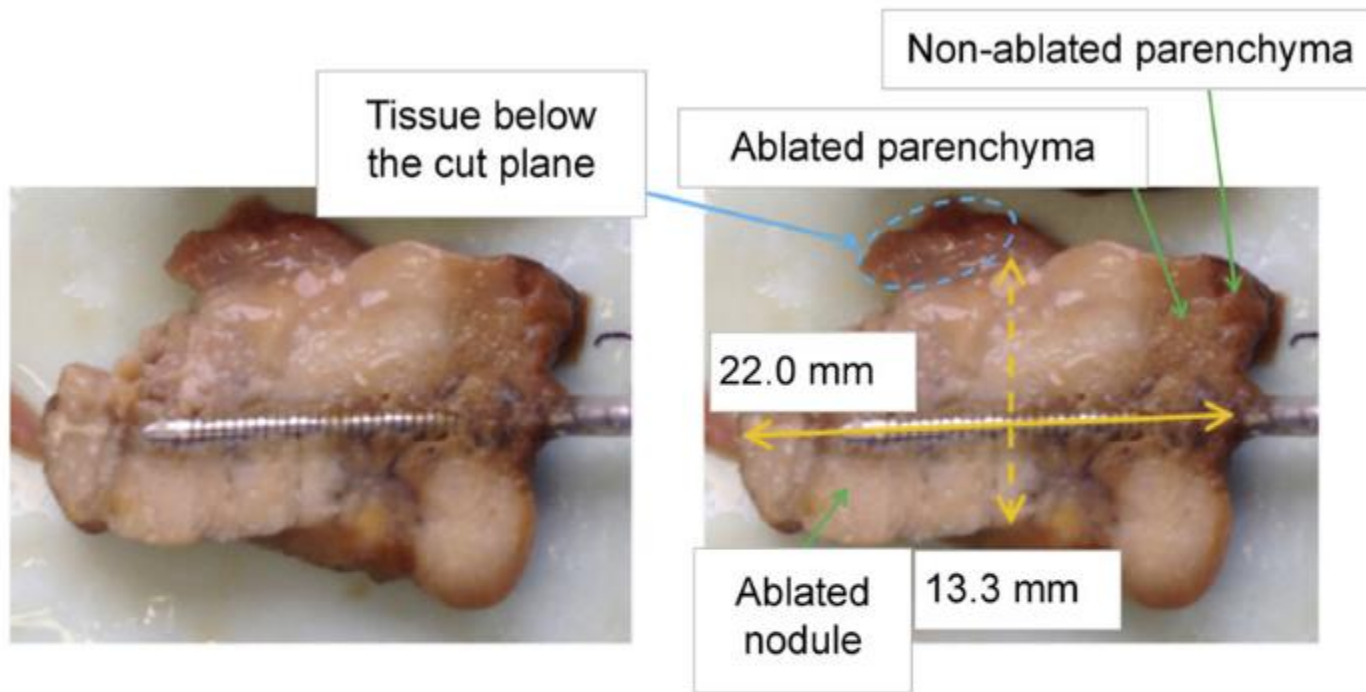
Monopolar RF catheter

18 mm length

1.8 mm diameter

good flexibility → Upper lobar 접근 가능

New ablation device in the resected lung tumor



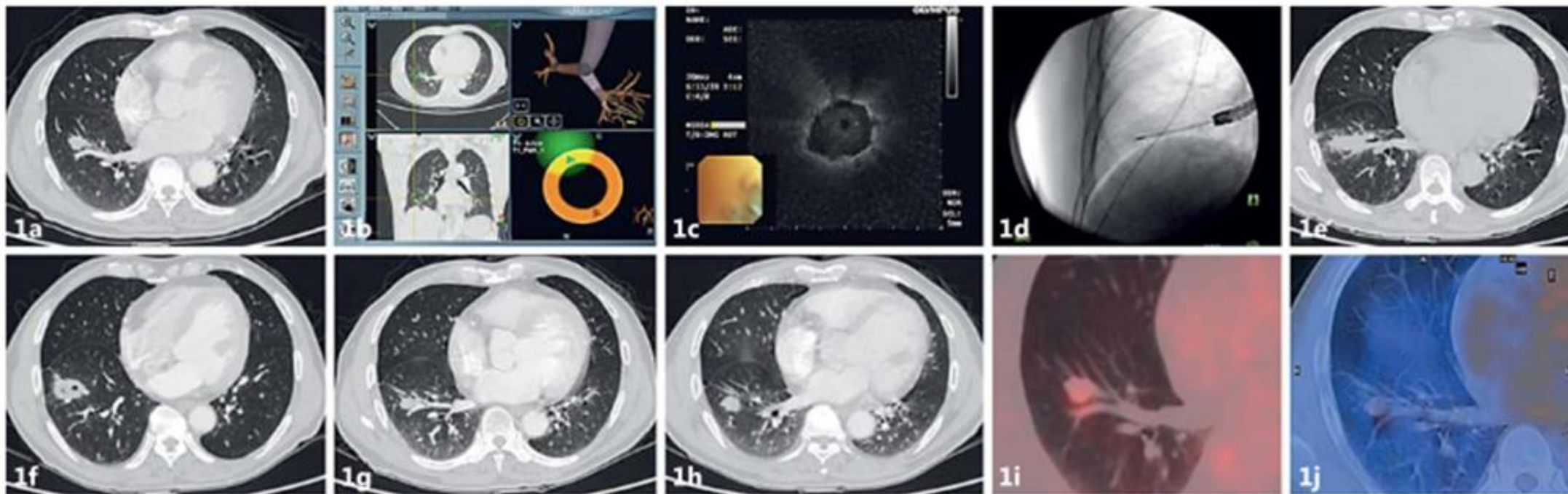
Five resected lung tumor

	Ablation zone
Major axis length	18.9 ~ 22.8 mm
Minor axis length	13.3 ~ 18.0 mm

- Both radial ablation distances reached the edge of tissue

Navigation guided RF ablation

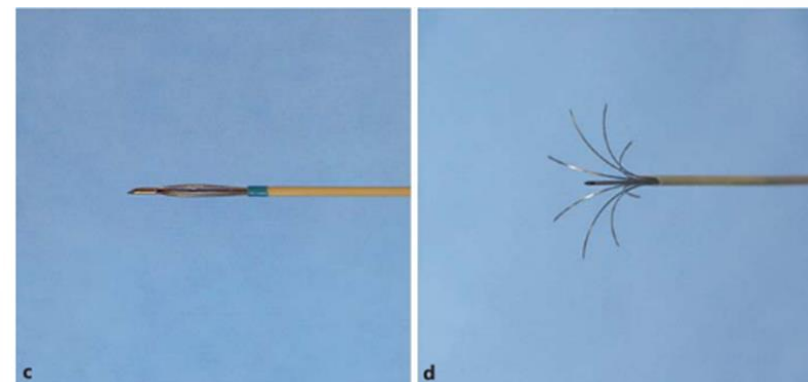
Case 1



13X11 mm sized lung nodule in the RLL

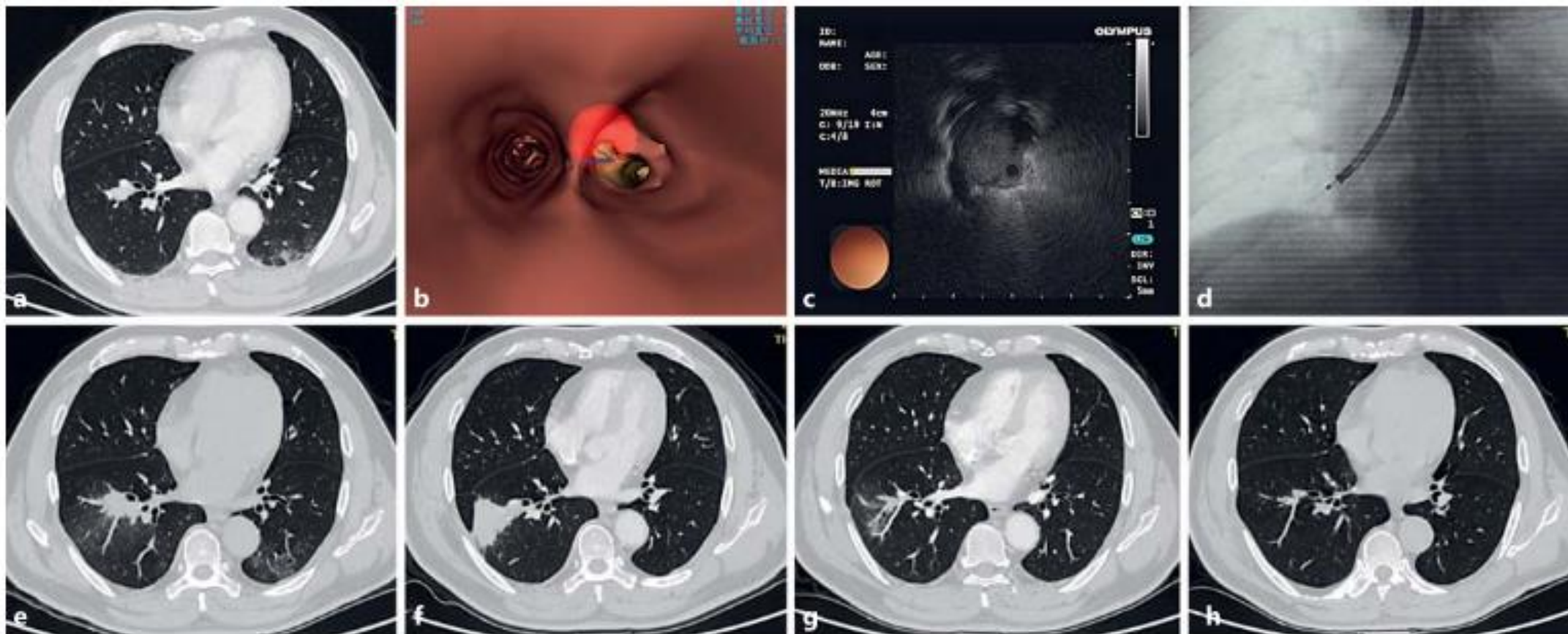
TBLB: SQC

EMB guided bronchoscopic RFA



Navigation guided RF ablation

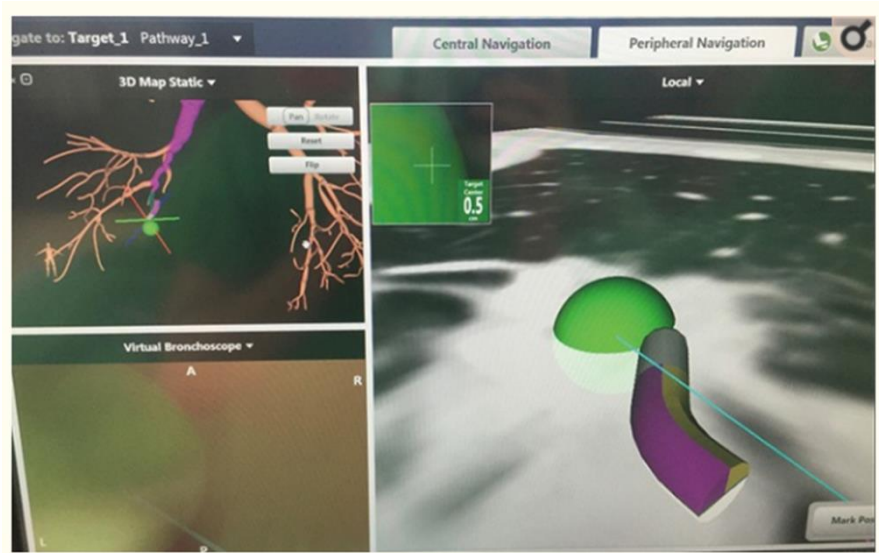
Case 3



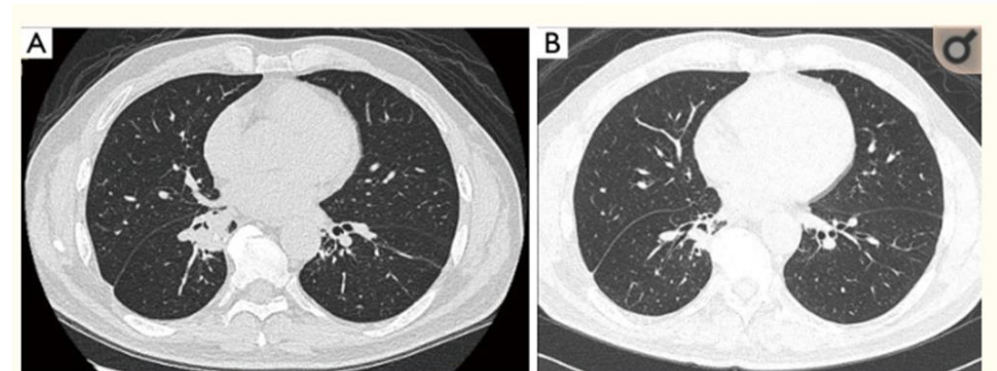
21.8X11.9 mm sized lung nodule in the RLL

Lung metastasis of RCC

Navigation guided PDT for lung nodule



No	Age/sex	Diagnosis	Location	Size (mm)	Admission day	Follow time (months)	Progression-free survival	Complications
1	75/M	Lung metastasis from colon cancer	RLL	36	5	16	+(PR)	-
2	57/F	Lung metastasis from colon cancer	RUL	20	4	14	+(PR)	-
3	65/F	Lung adenocarcinoma	RLL	8	3	12	+(CR)	Skin hypersensitivity, grade I

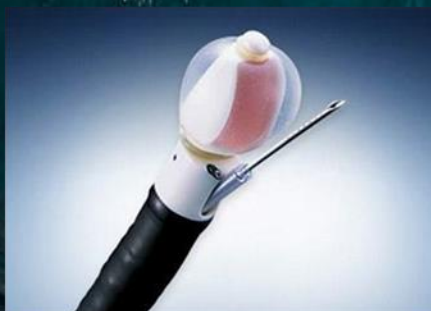




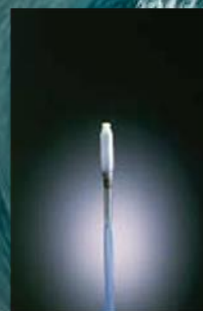
Gustav Killian
The first bronchoscopy in 1897



Shigeto Ikeda
fiberoptic bronchoscopy Invented in 1966



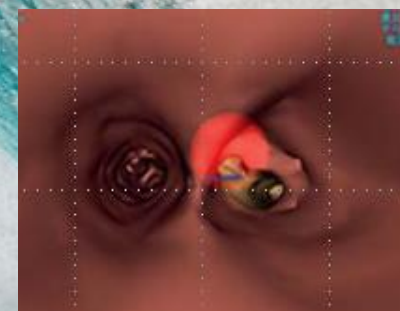
Convex probe EBUS



Radial probe EBUS



Radial balloon EBUS



Future of the bronchoscopy

» Diagnostic bronchoscopy

Development of transbronchial biopsy technique

- needle aspiration under real time ultrasound guidance

Robotic bronchoscopy

» Limitation of bronchoscopy guided treatment

SBRT, generally acceptable in case of surgical contraindication

- SBRT vs Guided therapy

Confirmation of RF zone

» Advantage of bronchoscopy guided treatment

Less complication than percutaneous approach

Repeat treatment

- Progression after SBRT
- Progression after previous RF ablation



SAMSUNG MEDICAL CENTER

Thank you
for your attention