

RECENT DEVELOPMENTS IN ADVANCED

DIAGNOSTIC BRONCHOSCOPY

MINIMIZING INVASIVENESS WHILE MAXIMIZING ACCURACY

150 years, Since Gustav Killian the “father of bronchoscopy”, bronchoscopy, especially flexible one, has been used as a way for airway inspection for diagnosing and therapeutic purposes .

Challenges to the wider use of flexible bronchoscopy have included difficulty in navigating to the lung periphery due to the complex nature of vasculature structures when performing diagnostic biopsies, and the ability to biopsy a lesion unconfined to the airway lumen.

The last 10–15 years have seen major advances in thoracic imaging, navigational platforms to direct the bronchoscopist to lung lesions, and the ability to visualize lesions during biopsy.

key advancements in diagnostic bronchoscopy aim to enhance imaging and provide bronchoscopy on to lung lesions are worth mentioning

Techniques : convex EBUS, radial EBUS [rEBUS], fluoroscopy

Additional Techniques: Virtual bronchoscopic navigation , electromagnetic navigational bronchoscopy, percutaneous endobronchial nodule access, CT bronchoscopy.

Advanced bronchoscopic local imaging techniques: Optical coherence tomography (OCT), Confocal laser endomicroscopy, Thin convex probe EBUS.

Technological Changes in the Bronchoscope : Ultrathin bronchoscopy , Robotic bronchoscopy.

ULTRASOUND

central airway radial ultrasound probes are used to detail imaging of the airway wall and surrounding structures

(e.g. 10 MHz radial probe EBUS fitted with a catheter that has a water-inflatable balloon at the tip), peripheral

allow for visualisation and subsequent sampling of peripheral intrapulmonary lesions

(e.g. 10 MHz ultra-miniature radial probe can be extended into subsegmental bronchi housed in a guide sheath)

In a feasibility study for the sampling of peripheral pulmonary lesions tested a prototype of a

visualisation device integrating r-EBUS and biopsy needle into a single device to sample lesions in real time

EBUS-TBNA

OLYMPUS



CP probe (CP)-endobronchial ultrasound

EBUS-TBNA has emerged as a technique that combined the high yield of mediastinoscopy (up to 90 %) with the minimally-invasiveness of TBNA, has the ability to locate lymph nodes and obtain the sample under direct visualization.

EBUS is integrated with a convex-shaped ultrasonic transducer at the tip of the bronchoscope. The physician can visualize the airway walls and surrounding structures by placing the end of the bronchoscope directly against the bronchial wall or by inflating the balloon with saline solution. It can be observed in real-time that the needle passes through the bronchial wall and the lesion of interest, it is possible to avoid blood vessels through the power Doppler mode. The procedure is installed with ultrasound guidance.

Complication rates in EBUS-TBNA were low (1% to 5%) such as bronchogenic cyst infection, mediastinal abscess, emphysema, pneumothorax, tracheobronchovascular fistula, tracheitis, pericarditis, and sepsis have been reported.

Convex probe (CP)-endobronchial ultrasound is the “gold standard” for lung cancer staging except for Hodgkin lymphoma remains lower at ~70% .

Excellent diagnostic accuracy for infectious aetiologies such as mediastinal tuberculosis (TB) lymphadenopathy.

EBUS has proven utility in the evaluation of sarcoidosis.

Cyst drainage and therapeutic drug injections

Concurrent EBUS and endoscopic ultrasound (EBUS is limited to the anterosuperior mediastinum, and EUS is limited to sample the posteroinferior mediastinum, which justifies the case for combining EBUS with EUS for thorough and systematic mediastinal staging)

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Video source: AstraZen

st recent advances in EBUS are focused on involving the **ultrasonographic characteristics of the LN**
r patterns and elastography

est study on this topic to date was published by Fujiwara and coworkers :Round shape, distinct margin
eneous echogenicity, presence of coagulation necrosis were found to be independently predictive
nancy. When all four factors were absent, 96% of the lymph nodes were benign.

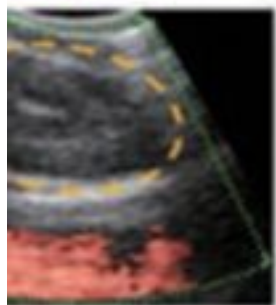
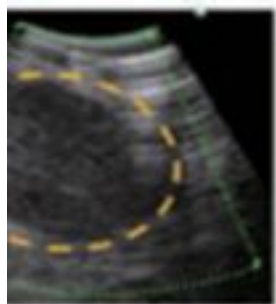
Chest 2010;13

et al. found that lymph nodes **measuring >10 mm** were associated with malignancy.

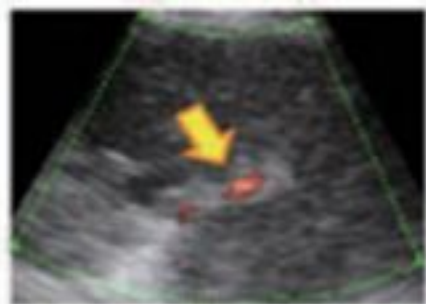
Chest 2011;14

jima et al. ,a classification system was developed based on the **pattern and number of vessels** in the L
aded from 0–III .When **grade 0 (no or minimal flow)** and **I (few main vessels running towards center of**
ined as “benign” and **grade II (few punctiform or rod-shaped flow signals)** and **III (rich flow with more t**
with different diameter and a helical flow signal) as “malignant”, they found that the sensitivity and diagn
y rate were 87.7% and 78% respectively. They also described with color-doppler imaging the “inflow si
ng of **blood arising in bronchial artery and flowing towards the LN (away from the probe)** resultin
nal . The accuracy of predicting metastasis solely from a positive BA inflow sign was 80.3%.

J Thorac Oncol 2012;7:



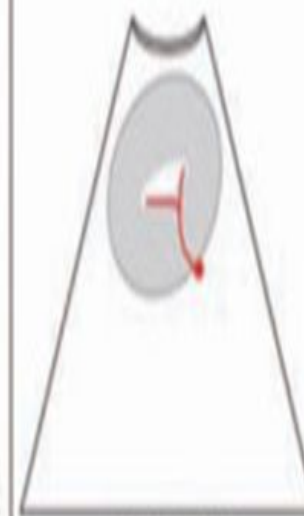
Grade 0



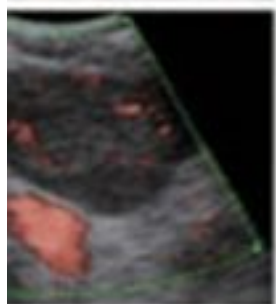
(B) Grade I



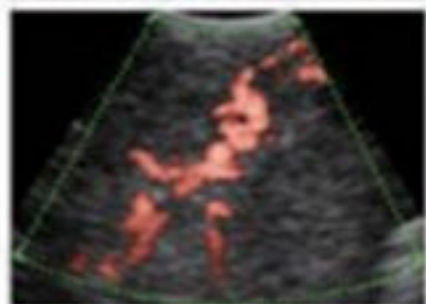
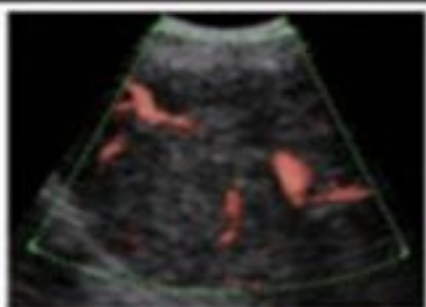
(A) Grade 0



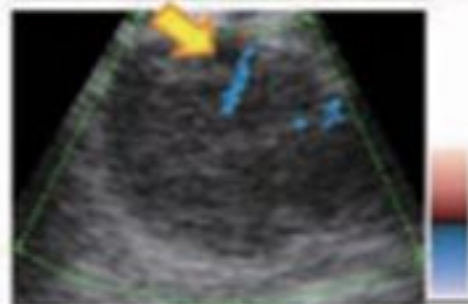
(B) Grade I



Grade II



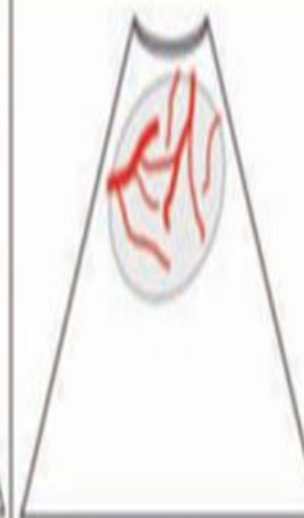
(D) Grade III



(E) BA inflow sign



(C) Grade II



(D) Grade III



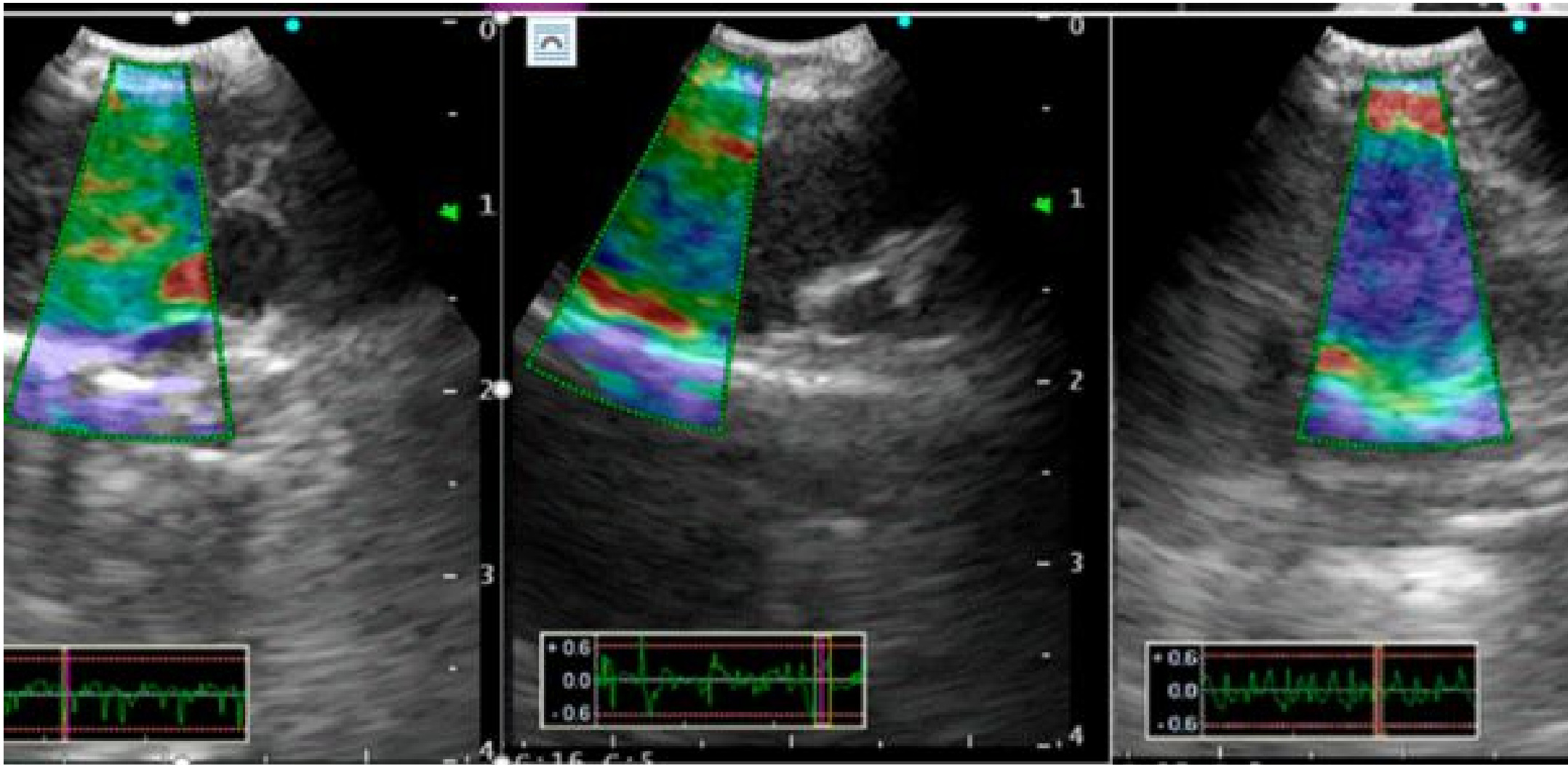
(E) BA

graphy

, pathological processes such as malignancy makes tissues less compressible.

, a new EBUS processor has been equipped with elastography which allows measurement of tissue **compressibility** (the elasticity of the tissue within the scanned area is compared with the surrounding tissue, and to a color signal that is superimposed on the B-mode image). **Colors** associated with **hard, intermediate** tissues **are blue, green and yellow/red** respectively.

et al. published the first retrospective data on this modality, and classified lymph nodes into 3 types (Type 1-predominantly non-blue, Type 2-part blue, and Type 3-predominantly blue), in the order of stiffness of the lymph node. They found 94.6% of type 3 lymph nodes to be positive for lymph node metastasis



B

C

1 elastography pattern (homogenous green) in a patient with tuberculosis. (B) Type 2 elastography pattern (heterogeneous color pattern) in a patient with sarcoidosis. (C) Type 3 elastography pattern (homogenous blue) in a patient with carcinoma.

Neither the ultrasound **characteristics** nor the **elastographic** appearance are likely to **replace** the need for **biopsy** of the LN.

But, When performing EBUS for mediastinal staging, we often find several LN in a given nodal station and it is not always feasible to sample all of these. These ultrasound/ elastographic characteristics may help us determine our best targets.

tional systems can be virtual (virtual bronchoscopic navigation, usually non-contrast enhanced magnetic (ENB)

bronchoscopic navigation encompasses multi-row detector CT-derived images to create three-dimensional renderings of the bronchial tree, thereby mimicking the view of a real flexible bronchoscope, but virtual bronchoscopy does not provide real-time positional guidance .

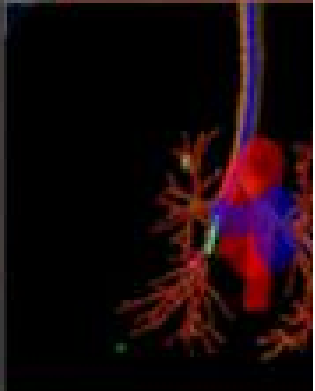
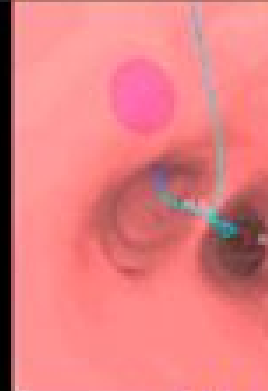
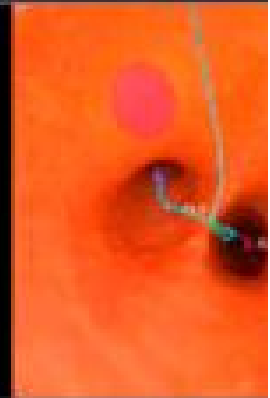
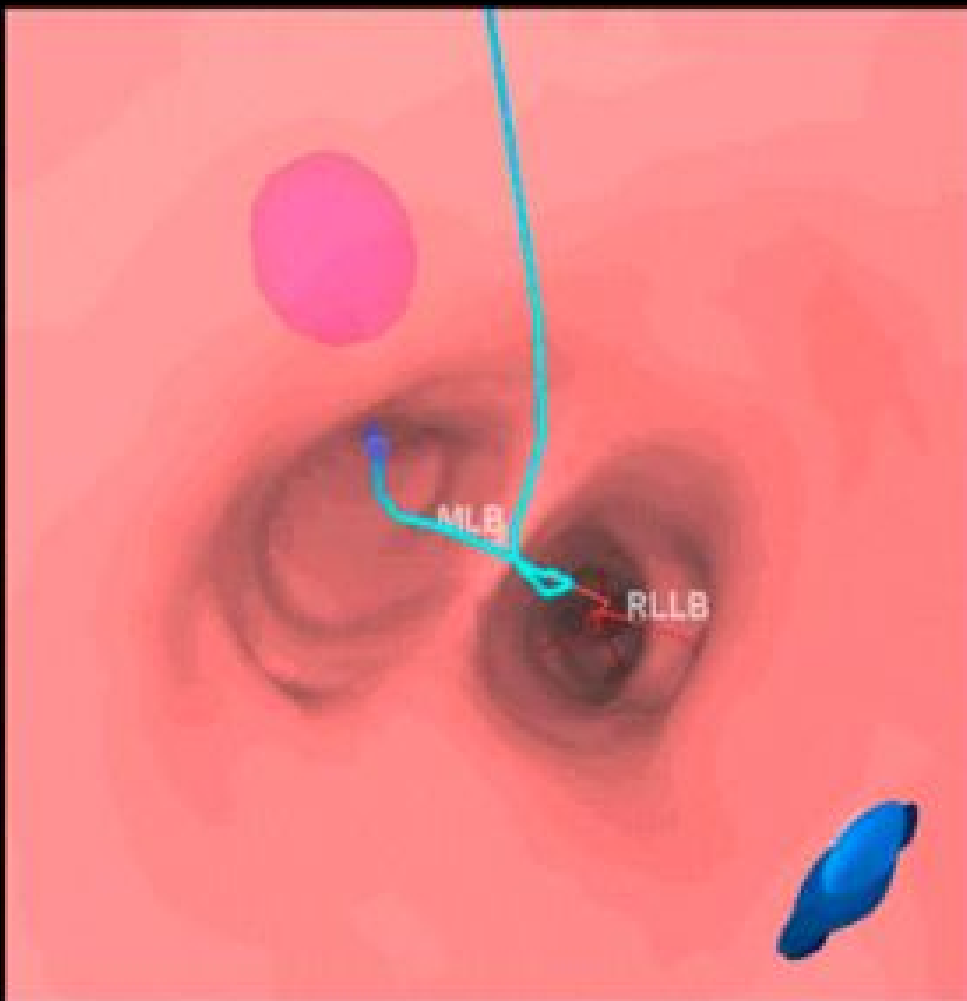
magnetic navigation (EMN) bronchoscopy relies on a pre-procedural CT of the chest to create a three-dimensional virtual airway map, which is then linked to an electromagnetic field to provide spatial feedback in electromagnetic navigation bronchoscopy (ENB) was cleared for use in the United States via FDA in 20



[superDimension™ Navigation System
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scopic transparenchymal nodule access (BTPNA):

nodules lack a bronchus sign and are so distant from a bronchus that bronchoscopic sampling techniques are not feasible. In these situations, transparenchymal nodule access was developed.

To access the nodule by creating a direct pathway that starts at the airway, goes through lung parenchyma, and reaches the lesion.

Using CT scan data, the procedure plan was uploaded to a virtual bronchoscopic navigation system that guided the bronchoscopist to the point-of-entry area. There, the airway wall was pierced by an 18-gauge needle and the opening was sealed by a small balloon catheter. Next, a 2.0 mm working channel sheath was inserted into the opening and held in place by a 15-gauge stylet and advanced together to the target lung lesion under fused CT scan-fluoroscopy guidance. The 2.0 mm sheath was then kept in place to allow various instruments to be used to sample the lesion.

ROBOTIC ASSISTED BRONCHOSCOPY

robotic system to be introduced in the field of bronchoscopy was the [Monarch™ platform](#) by Auris Health, which received FDA approval in March [2018](#). Subsequently another robotic bronchoscopy platform, [Ion™ Endoluminal Navigation System](#) developed by Intuitive Surgical received FDA approval in February [2019](#).

These systems use a [small endoscope controlled by robotic steering devices under direct visualization](#) by the operator. The system still requires thin slice CT scan data to plan the pathway and navigate to the desired target.

Key advantages of such technology include continuous endobronchial visualization and greater maneuverability of the bronchoscope with the ability to lock into a desired position.



ULTRATHIN BRONCHOSCOPY

The small size of the peripheral airways limits the ability of conventional bronchoscopes to navigate to peripheral lesions. The channel of conventional pediatric bronchoscopes limits the size of the tools needed to diagnose peripheral nodules. The introduction of ultrathin bronchoscopes (with 2.8–3.5 mm outer diameter) allows for greater maneuverability to traverse small airways. Although no strict definition of “ultrathin” exists, most have outer dimensions of 3.2 mm or less. They can be guided deep into the lung to a median of the sixth generation bronchi (range, fourth- to ninth-generation bronchi), allowing more distal diagnosis.



Fluorescence bronchoscopy (AFB):

uses green- and red-spectrum light to detect mucosal alterations. Normal mucosa presents green color, while precancerous and cancerous lesions absorb the green spectrum and turn magenta.

High sensitivity but low specificity except when used in follow up of surgical margins after curative surgery or to detect extension of lung cancer.

One of the open issues on AFB is question of its usability in bronchoscopic lung cancer screening. Results of previous studies did not support the general use of AFB as a screening tool for lung cancer.

AFI

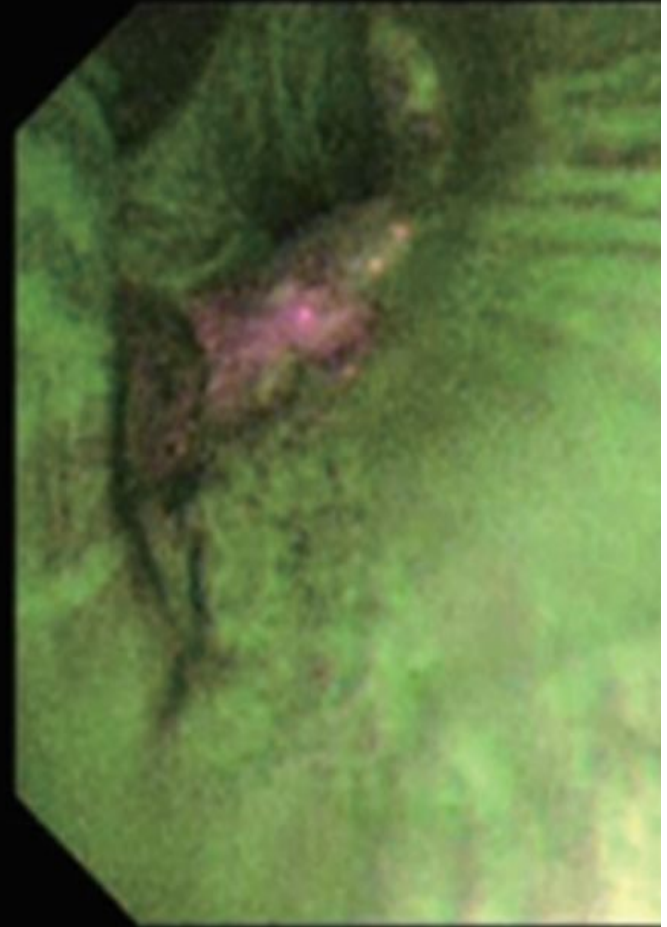
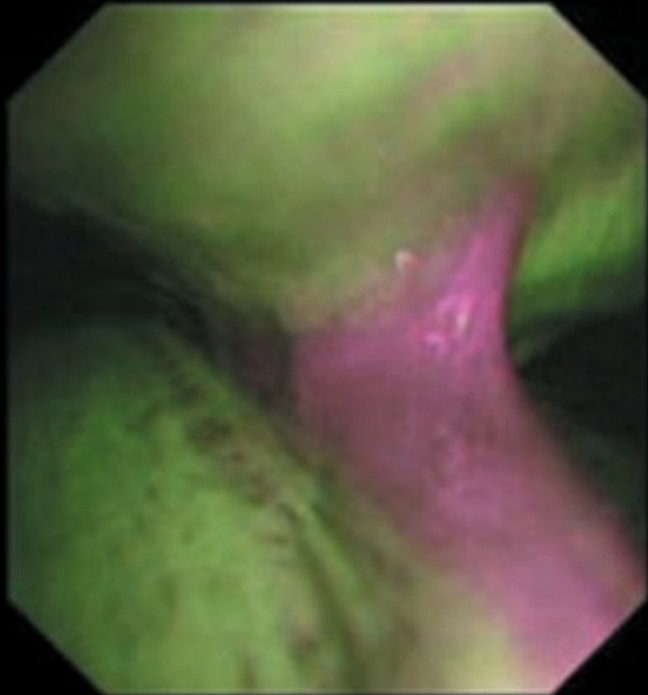
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V BAND IMAGING

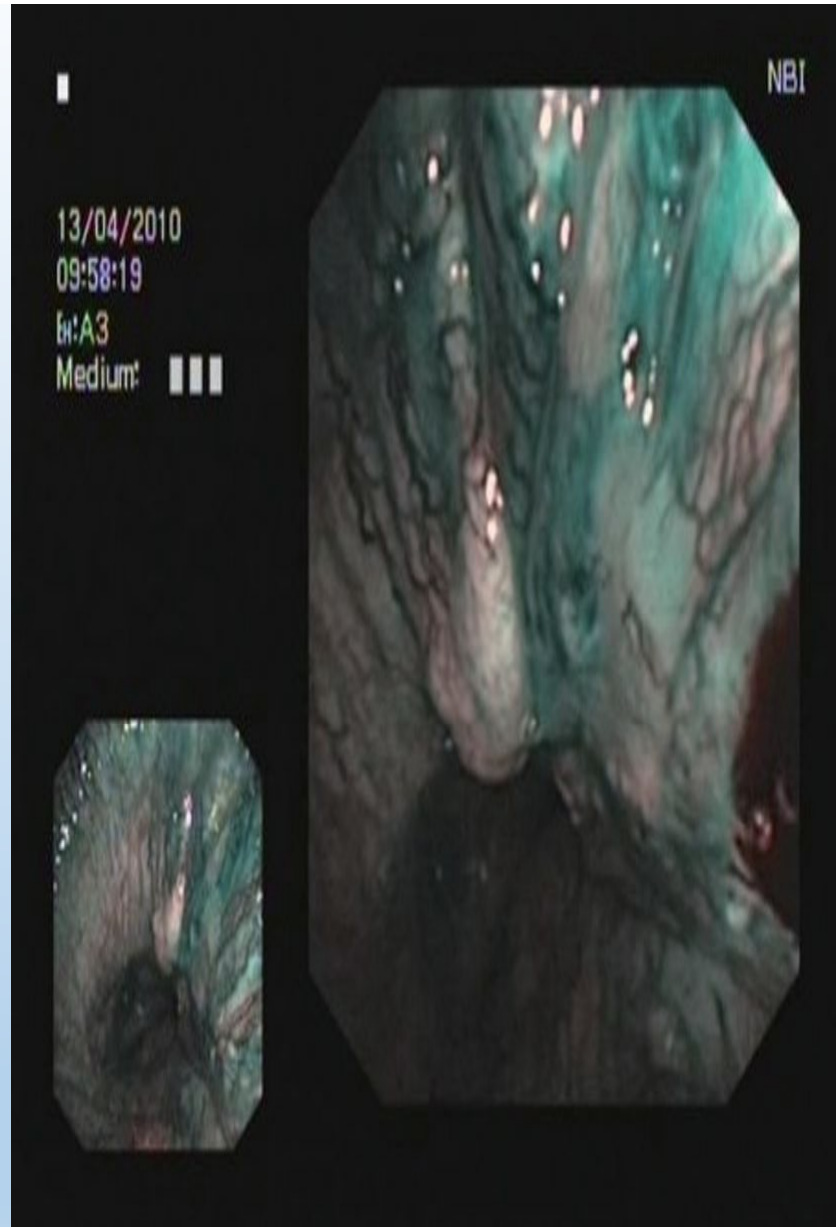
and imaging removes all wavelengths except two that are **absorbed by hemoglobin**, thereby creating the **vasculature** (cyan) and surrounding mucosa (brown)

et al. first described the pathological patterns on bronchial mucosa that are known as Shibuya's description (**tortuous and abrupt ending vessels**)

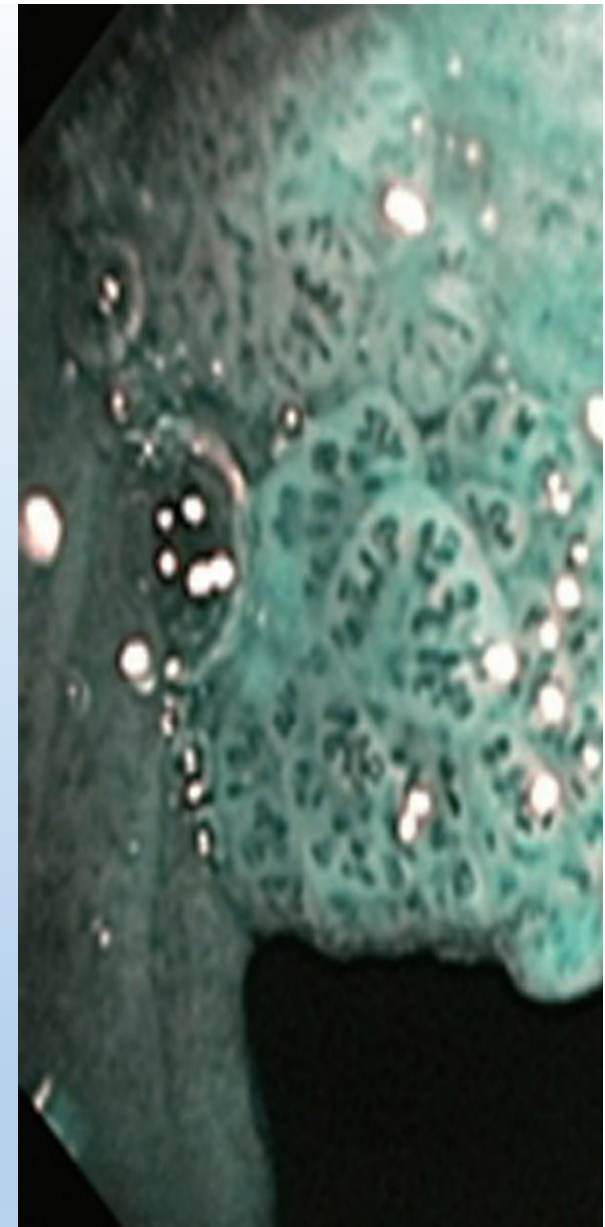
, this technology could be successfully used for evaluation of tumor margins, follow up after curative surgery



tortuous blood vessels-
squamous cell lung cancer



abrupt ending blood vessels-
squamous cell lung cancer



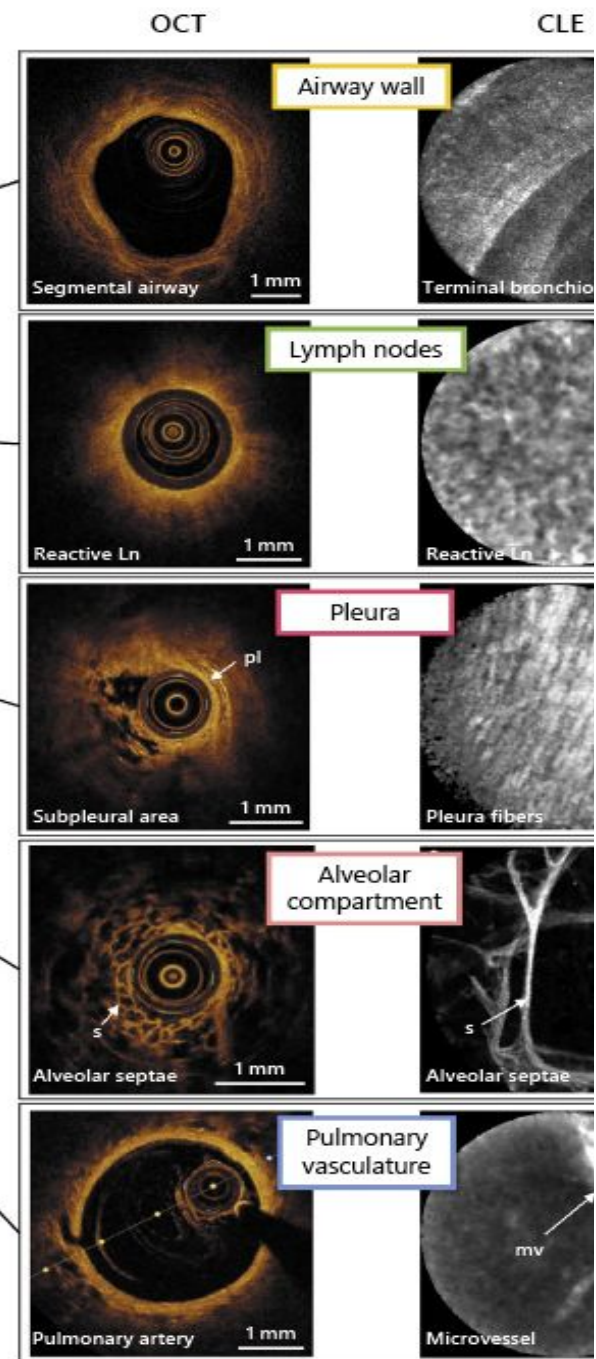
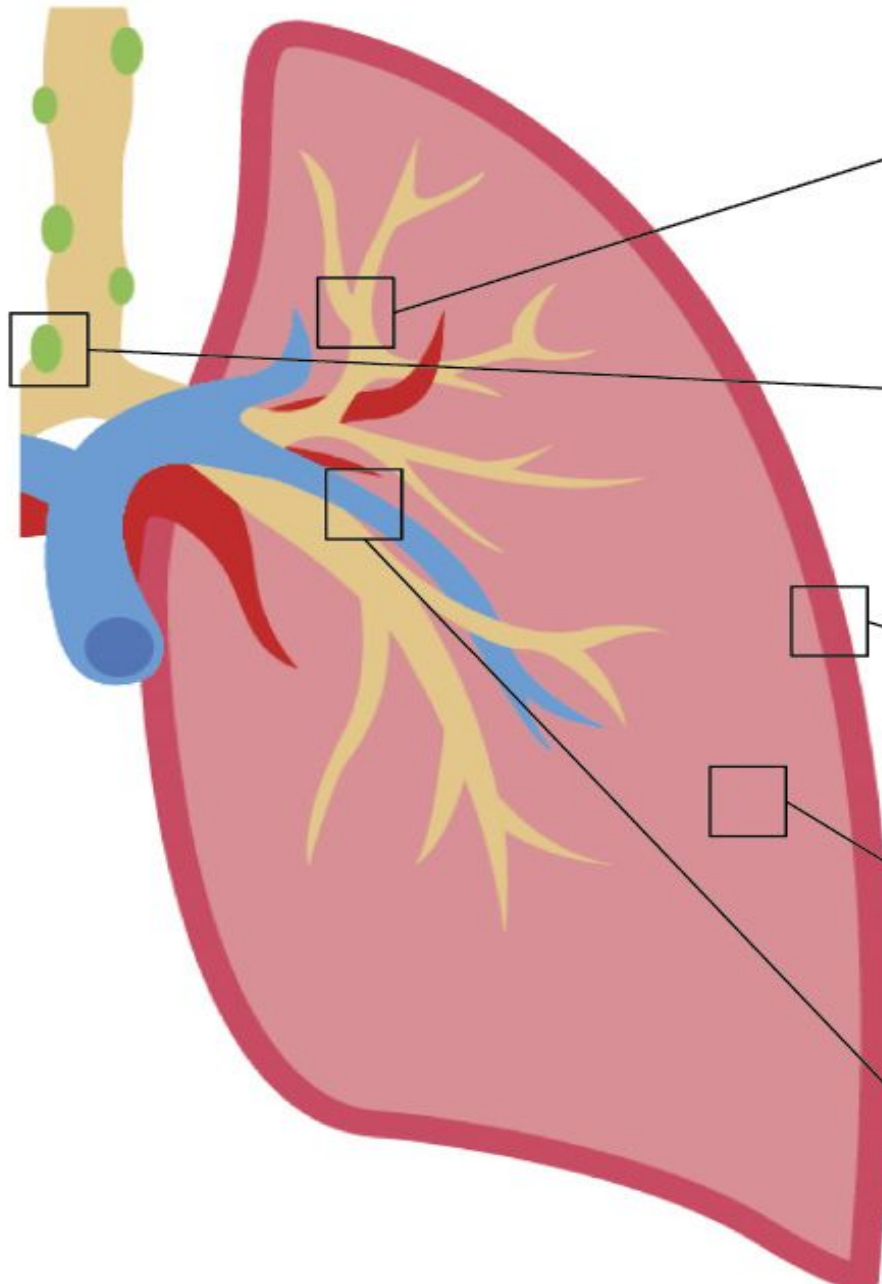
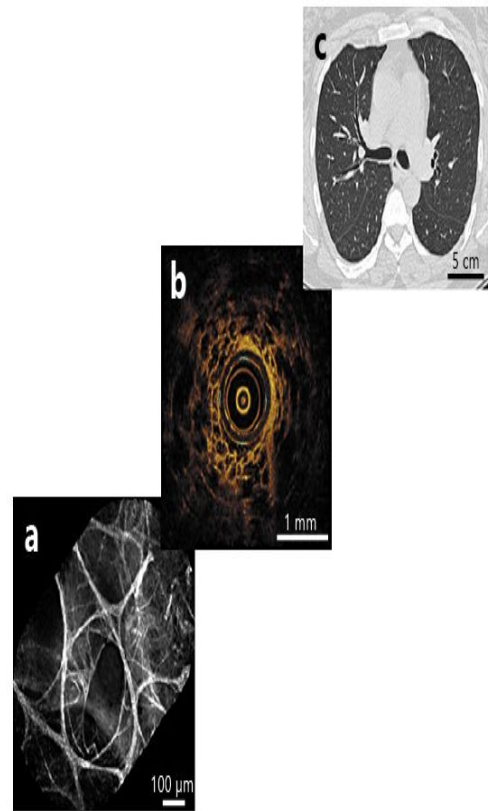
dotted vascular p
adenocarcinoma of th

coherence tomography (OCT) :

n imaging technique using near-infrared light to generate high-resolution images of tissue structures with a resolution of $\pm 10\text{--}15\ \mu\text{m}$ and depth of $2\text{--}3\ \text{mm}$. The conceptual idea of OCT is comparable to ultrasound, but instead of the reflection of acoustic waves, OCT uses the scattering of near-infrared light to generate images. In OCT, an optical beam generates near-infrared light and focuses on the tissue.

PCLE BASED CONFOCAL LASER ENDOMICROSCOPY “PCLE”

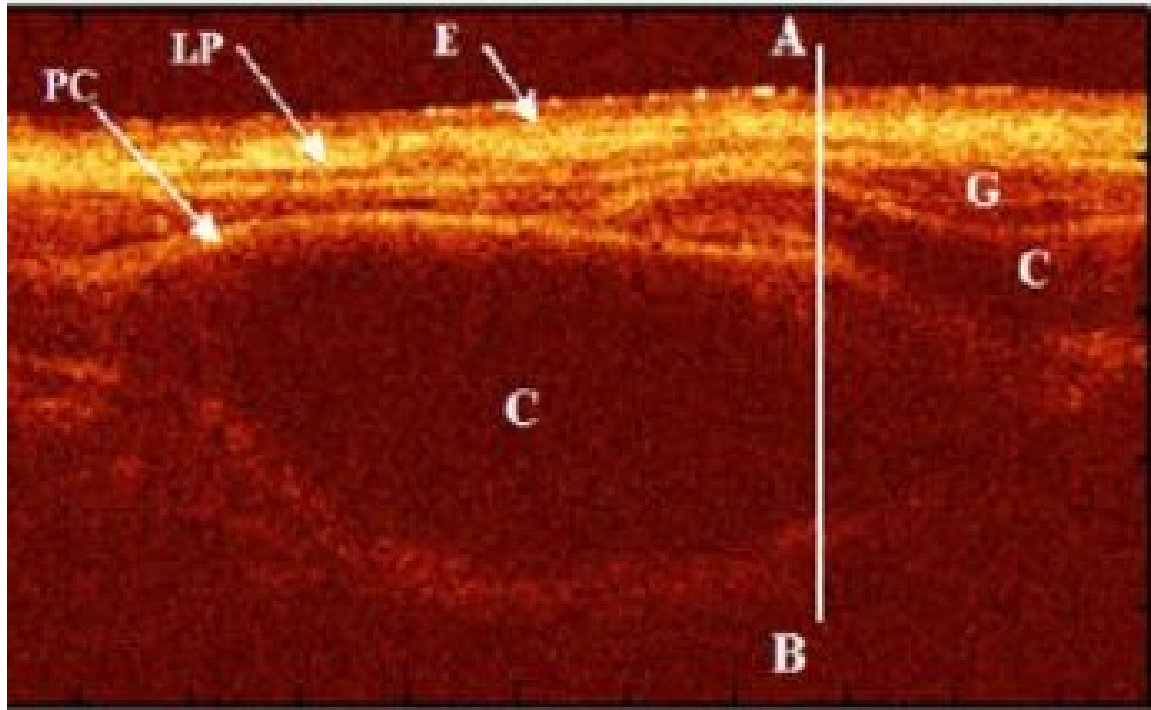
real-time images of the airways, alveoli, lung tumours, pleura and lymph nodes with a resolution up to $10\ \mu\text{m}$ and a maximum depth of $70\ \mu\text{m}$ and a maximum field of view of 600° . A fibre-optic probe is advanced through the working channel of a bronchoscope directed to the area of interest where it illuminates tissue with laser light (usually used $488\ \text{nm}$). Reflected light is redirected back through a pinhole.



$\pm 100 \mu\text{m}$ $\pm 2 \text{ mm}$ $\pm 0.5 \text{ m}$

Imaging depth \longrightarrow

KK '19



0.5 1 1.5 2 2.5 3 3.5 4 4.5 5

Transverse Length (mm)

E: epithelium
LP: lamina propia
G: glandules
PC: pericondrium
C: cartilage
A-B: 500 μ m

ymal lung disease

Transbronchial cryobiopsy (TBCB) is gaining popularity in the diagnostic approach to diffuse
ymal lung diseases.

performing via bronchoscopic placement of a flexible cryoprobe inside the lung parenchyma, freezing
and shearing out the frozen lung tissue, providing larger specimens and more alveolar tissue (tend to b

mal positioning of the cryoprobe with a distance of 1 cm from the pleura is recommended to minimize
ations)

