

Robotic-assisted bronchoscopy: a narrative review of systems

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Background and Objective: Robotic-assisted bronchoscopy (RAB) has emerged as an advanced technology for lung cancer diagnosis. This review explores the three approved robotic bronchoscopy systems: Ion[™] Endoluminal (Intuitive Surgical, Sunnyvale, CA, USA), Monarch[™] (Johnson & Johnson, Redwood City, CA, USA), and Galaxy System[™] (Noah Medical, San Carlos, CA, USA), and their different operational systems. This narrative review aims to summarize their findings and outcomes for sampling peripheral pulmonary lesions (PPL) suspected of lung cancer.

Methods: A search in PubMed and Google Scholar databases was conducted for articles and abstracts published between January 2018 to May 2024 using the terms "robotic bronchoscopy" or "robotic-assisted bronchoscopy" for biopsy of PPL.

Key Content and Findings: Lung cancer is the leading cause of cancer-related mortality. The introduction of RAB aims to improve the feasibility and safety of sampling PPL. Current literature describes high diagnostic yields with low risk of complications, allowing concurrent hilar and mediastinal staging within the same procedure. RAB can potentially improve early diagnosis and treatment of pulmonary malignancies and survival rate in long term, while progressing towards therapeutic applications in the near future.

Conclusions: As RAB evolves, its potential as a "one-stop shop" for diagnosis, staging, and treatment can positively impact lung cancer detection, focusing on improved patient-centered outcomes and reducing multiple diagnostic and therapeutic procedures.

Keywords: Robotic-assisted bronchoscopy (RAB); shape-sensing technology; electromagnetic navigation (EMN); lung cancer

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Introduction

Lung cancer stands as the leading cause of cancer-related deaths in the United States, surpassing breast, prostate, and colon cancers in mortality rates (1). The severe impact of this malignancy demands a constant search for effective diagnostic and treatment approaches, for which timely screening and detection are crucial to reduce mortality.

New guidelines from the American Cancer Society now recommend annual lung cancer screening for healthy individuals aged 50 to 80 years with a \geq 20 pack-year smoking history, regardless of the time since quitting. This expanded eligibility includes an additional 5 million people, potentially preventing more lung cancer deaths (2). Screening has played a vital role, reducing mortality by 16– 24% through the detection of asymptomatic malignancies amenable to curative treatment in high-risk individuals (3,4). However, uptake remains low, with approximately only 6% adherence to screening guideline criteria among the eligible population in 2020 (5).

In diagnostic efforts, robotic-assisted bronchoscopy (RAB) has emerged as an advanced technology for early lung cancer detection. Aside from cost-related barriers, including platform expenses and tools, RAB is showing promise in improving diagnostic yield for pulmonary peripheral lesions (PPL). Recent studies highlight RAB's maneuverability, farther reach, and navigational and diagnostic capabilities compared to conventional methods. When available, it is becoming the diagnostic modality of choice. This narrative review aims to summarize the findings and outcomes of the three available robotic platforms, including their different operational systems and characteristics. We present this

 Table 1 The search strategy summary

article in accordance with the Narrative Review reporting checklist (available at https://jtd.amegroups.com/article/ view/10.21037/jtd-24-456/rc).

Methods

For this narrative review, we conducted a search in the PubMed and Google Scholar databases for articles and abstracts published between January 2018 to May 2024 using the terms "robotic bronchoscopy" or "robotic-assisted bronchoscopy" for biopsy of PPL. The search strategy is detailed in *Table 1*.

Review of systems

Ion[™] Robotic Endoluminal System

The IonTM Robotic Endoluminal System, developed by Intuitive Surgical (Sunnyvale, CA, USA), received approval from the US Food and Drug Administration in February 2019. This system comprises a flexible robotic catheter equipped with shape-sensing fibers along its entire length, a removable video scope for direct visualization during navigation, a planning station, a robot cart with display screens, and a controller featuring a trackball and a scroll wheel (*Figure 1*). The articulating catheter has a 3.5-mm outer diameter (OD) and a 2.0-mm inner diameter (ID) working channel. For direct visualization, a 1.7-mm OD vision probe is inserted via the working channel, although the IonTM video scope must be removed for biopsying after navigation, precluding live imaging during sampling.

Pre-procedure planning for IonTM involves a contiguous

Table 1 The search strategy summary	
Items	Specification
Date of search	10/01/2023 to 1/5/2024
Databases and other sources searched	PubMed and Google Scholar
Search terms used	"Robotic bronchoscopy" or "robotic-assisted bronchoscopy"
Timeframe	January 2018 to May 2024
Inclusion and exclusion criteria	Inclusion criteria: original articles including retrospective and prospective studies, meta- analysis, and conference abstracts) for human and animal studies, in the English language
	Exclusion criteria: not in the English language
Selection process	The selection process was conducted by A.K. and N.C.C. independently, and later approved by two board-certified interventional pulmonologists. Duplicate results were eliminated. Consideration for additional studies and review of the final references were performed by all authors



Figure 1 Ion[™] Robotic Endoluminal System bronchoscopy. (A) Robotic arm, platform, and screen and trackball controllers (courtesy of Intuitive Surgical). (B) Trackball controllers and navigation screen. (C) Navigation display with 3-dimensional airway pathway in the upper screen, and mobile cone-beam computed tomography images in the axial, coronal, and sagittal view showing biopsy tool-in-lesion in the lower screen.

thin-cut (1.5 mm or less) chest computed tomography (CT) scan. The Ion[™] PlanPoint[™] software generates a three-dimensional (3D) virtual airway reconstruction, automatically identifying potential pathways towards the target, which can also be done manually if necessary. Once the robotic arm is docked and the catheter introduced, registration begins by aligning the distance and rotation of the main carina between the virtual and live bronchoscopic images. The bronchoscope is then driven into each of the main bronchi, and into the individual lobar and distal airway segments.

The Ion[™] platform employs proprietary shape-sensing technology with a fiber embedded along the robotic catheter, providing real-time shape and location feedback. This data enables the catheter tip to correlate with the CTderived airway map, determining the catheter tip position within the lung. Real-time information on the direction, distance to the target, and distance from the nearest pleura is constantly provided by the system to achieve successful navigation. Fluoroscopy, radial endobronchial ultrasound (rEBUS), and cone-beam CT (CBCT), mobile CBCT (mCBCT) or O-arm CT are used as supportive devices for secondary confirmation of target reach and tool-inlesion. The catheter is then locked in position, and the vision probe removed to facilitate the use of biopsy tools. A comprehensive comparative description of shape-sensing robotic-assisted bronchoscopy (ssRAB) to the other RAB systems is presented in *Table 2*.

Evidence

In the initial feasibility human study of the IonTM Endoluminal System, conducted by Fielding *et al.*, 29 consecutive cases were analyzed in a single center (6). The mean lesion size was 12 mm (range, 10–30 mm), with 58.6% of lesions exhibiting a positive bronchus sign. Successful navigation, assisted by rEBUS, was achieved in 96.6% (28/29) of cases. Diagnostic yield was 79.3%, with a follow-up period of 6 months. Subsequently, in the first real-world prospective study, 59 nodules from 52 patients were biopsied, using fine needle aspiration and forceps, and

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	Monarch™ Robotic Endoscopy System	The Ion™ Robotic Endoluminal System	Galaxy System™
FDA approval	March 2018	February 2019	March 2023
Bronchoscope	4.2 mm inner bronchoscope,6 mm outer sheath	3.5 mm outer diameter fully articulating catheter with a thin 1.8 mm removable visual probe	4.0 mm outer diameter
Working channel	2.1 mm	2 mm	2.1 mm
Navigation	Electromagnetic navigation along with peripheral vision and real time input from the micro-camera at the tip of the bronchoscope	Fiberoptic shape-sensing and peripheral vision	Electromagnetic navigation with digital tomosynthesis Tool-in-Lesion+ Technology™
Scope reprocessing	Yes	Yes	No (single use disposable scope)
Vision during biopsy	Yes	No	Yes

Table 2 Comparison of the three robotic bronchoscopic platforms

FDA, United States Food and Drug Administration.

standard CBCT for secondary confirmation of successful navigation (7). The mean nodule size was <20 mm, achieving a 100% target reach rate, 83% overall diagnostic yield and 84% sensitivity for malignancy. Reported complications were two pneumothoraxes, one requiring chest tube insertion.

At Memorial Sloan Kettering Cancer Center, a retrospective single-center trial was conducted, using ssRAB with mCBCT and rEBUS. Encompassing 130 patients with 159 lesions with a median lesion size of 18 mm [interquartile range (IQR), 13-27 mm], successful navigation was achieved in 98.7% (157/159) of cases, resulting in an overall diagnostic yield of 81.7% (8). This yield was stratified by rEBUS view, 93% for concentric and 78.8% for eccentric views. The reported complication rate was 3% (4/130), including 2 patients developing post-procedural pneumothorax requiring tube thoracostomy. Finally, in a large single-center study, Styrvoky et al., performed 200 ssRAB, combined with CBCT and rEBUS, to sample 209 nodules with a mean size of 22.6 mm (±13.3) (9). Diagnostic yield was 91.4% and sensitivity for malignancy was 87.3%. Procedure-related complications were two pneumothoraxes, only one resulting in chest tube insertion.

For PPL under 2 cm, suboptimal diagnostic yields at 66% for <1 cm and 70.4% for 1–2 cm have been reported (8). A retrospective multicenter study led by Abia-Trujillo *et al.* assessed the diagnostic accuracy for small PPL under 2 cm using ssRAB integrated with mCBCT (10). The study included 192 nodules from 173 patients, with 117 nodules utilizing mCBCT and standard C-arm for the remaining ones. Diagnostic yield was 83.8% and 88% for subgroups

with and without mCBCT, respectively, highlighting the improved diagnostic performance of ssRAB for small lung nodules regardless of the type of intraprocedural imaging. Likewise, the capability of ssRAB to accurately biopsy ground glass nodules (GGNs) and subsolid nodules (SSNs) with a solid component less than 6 mm, in combination with mCBCT and rEBUS, was evaluated in a cohort study comprising 12 SSNs and 11 pure GGNs (11). ssRAB demonstrated an overall diagnostic yield of 87%, with a sensitivity for malignancy reaching 88.9%. Importantly, no procedure-related complications were observed. Lastly, the same authors recently presented their findings utilizing ssRAB with mCBCT and rEBUS to biopsy multiple nodules within the same procedure, including unilateral and bilateral lesions (12). A total of 46 nodules less than 20 mm in median size were sampled from 22 patients, achieving a diagnostic yield of 84.8% with two complications comprising one grade 2 bleeding in the Nashville Scale and one pneumothorax requiring chest tube. A comprehensive summary of findings for IonTM Endoluminal is described in Table 3.

Monarch[™] Robotic Endoscopy System

The Monarch[™] Robotic Endoscopy System, developed by Auris Robotics (now Johnson & Johnson) in Redwood City, CA, USA, was the first RAB platform to be approved by the US Food and Drug Administration in March 2018. Comprising a bronchoscope system, cart, and tower, the Monarch[™] system employs an articulating bronchoscope within an articulating sheath, each controlled by independent

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Author, year	Type of study	Patients (n)	Mean/median diameter (mm)	SD/IQR or range	Nodules sampled (n)	Diagnostic yield (%)	
Ion™ Robotic Endoluminal Syst	em						
Fernandez-Bussy 2024, (12)	Retrospective	22	14	IQR, 1–2	46	86.9	
Fernandez-Bussy 2024, (13)	Retrospective	27	16.6	IQR, 13.6–19.9	28	85.7	
Abia-Trujillo 2024, (10)	Retrospective	173	12	IQR, 10–14	192	77.4	
Abia-Trujillo 2023, (11)	Retrospective	22	18	IQR, 14–20	23	88.9	
Gupta 2023, (14)	Abstract prospective	240	16	IQR, 1.1–25	265	76	
Hammad Altaq 2023, (15)	Retrospective	42	12	IQR, 10–18	42	88.1	
Low 2023, (16)	Retrospective	133	19	IQR, 14–28	143	77	
Chambers 2023, (17)	Retrospective	75	20	IQR, 13–35	79	77.2	
Styrvoky 2022, (9)	Retrospective	198	22.6	SD: 13.3	209	91.4	
Oberg 2022, (18)	Retrospective	112	22	IQR, 13–34.3	120	90	
Yu Lee-Mateus 2023, (19)	Retrospective	113	18	IQR, 13–27	113	87.6	
Reisenauer 2022, (20)	Prospective	30	17.5	SD: 6.8	30	93.3	
Tavakoli 2022, (21)	Abstract retrospective	65	21.2	SD: NA	65	86.2	
Kalchiem-Dekel 2022, (8)	Prospective	130	18	IQR, 13–27	159	81.7	
Ost 2021(22)	Abstract prospective	155	17	SD: 5.5	155	83	
Benn 2021, (7)	Prospective	52	19.6	SD: 10.9	59	83	
Ross 2021, (23)	Abstract prospective	45	14	Range, 5-44	58	89	
Bawek 2021, (24)	Abstract retrospective	76	17	Range, 6–70	76	92	
Ghosh 2021, (25)	Abstract retrospective	95	19	Range, 7–69	103	79.6	
Pritchett 2021, (26)	Abstract retrospective	192	15	IQR, 10–21	230	92.2	
Folch 2020, (27)	Abstract prospective	129	18.4	SD: 5.44	129	80.6	
Fielding 2019, (6)	Prospective	29	12.2	SD: 4.2	29	79.3	
Monarch™ Robotic Endoscopy	System						
Agrawal 2023, (28)	Retrospective	124	24	IQR, 13–30	124	77	
Khan 2023, (29)	Retrospective	264	19.3	Range, 32–72.5	264	85.2	
Manley 2023, (30)	Prospective	20	14.5	Range, 8–28	20	80	
Hedstrom 2022, (31)	Abstract retrospective	45	16.9	Range, 4-35	45	91	
Cumbo-Nacheli 2022, (32)	Retrospective	20	22	SD: 7	20	90	
Chen 2021, (33)	Prospective	54	23.2	SD: 10.8	54	82.5	
Ekeke 2021, (34)	Retrospective	25	NA	NA	25	96	
Chaddha 2019, (35)	Retrospective	165	25	SD: 15	167	77	
Rojas-Solano 2018, (36)	Prospective	15	26	Range, 10-63	15	86.7	
Galaxy™ Robotic Endoscopy S	ystem						
Saghaie 2023, (37)	Abstract Prospective	14	20.5	Range, 10–34	15	93	

Table 3 Review of human studies using robotic bronchoscopy platforms

SD, standard deviation; IQR, interquartile range; NA, not available.



Figure 2 Monarch[™] Robotic Endoscopy System bronchoscopy. (A) Tower and cart with robotic arms. (B) Controllers (courtesy of Johnson & Johnson).

robotic arms mounted on the RAB cart (*Figure 2A*). Connecting to the RAB cart, a separate tower houses a monitor screen and a controller (*Figure 2B*). With a 6.0-mm OD for the outer sheath and a 4.2-mm OD with a 2.1-mm ID working channel for the inner bronchoscope, the system prioritizes stability through the outer sheath and enhanced maneuverability and articulation via the flexible inner bronchoscope.

Pre-procedure planning involves a contiguous thincut chest CT scan, enabling the creation of a 3D virtual lung reconstruction. The planning software automatically generates pathways to the target, with manual planning available as an alternative. Subsequent registration aligns the robotic bronchoscope's position with the virtual pathway. The Monarch[™] platform's registration process entails moving the bronchoscope to the main carina and then retracting it, followed by a prompt to advance and retract the bronchoscope in the contralateral main bronchus, ensuring smooth movement within the airways.

The proprietary navigation algorithm of the Monarch[™] system combines optical pattern recognition, electromagnetic navigation (EMN) positioning, and robotic insertion data to confirm the scope tip position within the lung. This fusion process corroborates the signals with the CT-derived airway map. Operators control the bronchoscope using a handheld controller resembling a gaming controller. To compensate for the lack of tactile feedback, the system triggers "buckling errors" upon pressure on the airway walls. Fluoroscopy or rEBUS, as well as CBCT and other intraprocedural imaging devices, are used to verify target reach before locking the

entire bronchoscope system in a static position for biopsy acquisition. A comprehensive comparative description of RAB and EMN to the other RAB systems is presented in *Table 2*.

Evidence

In 2018, Rojas-Solano *et al.* conducted the first human safety and feasibility study using the MonarchTM system (36). Out of the 15 lesions studied, 14 (93%) were successfully sampled, without complications. The initial post-market multicenter study by Chaddha *et al.* involved four centers and analyzed 167 PPL in 165 patients (35). With a mean lesion size of 25±15 mm, 71.3% of lesions were \leq 30 mm, and 70.7% were located in the peripheral third of the lung. Successful navigation was achieved in 88.6% of cases, with visual confirmation by rEBUS in 84.4% of lesions (concentric =57.5%; eccentric =42.5%). Diagnostic sensitivity was reported at 69.1% *vs.* 77%, considering inflammation alone as non-diagnostic *vs.* diagnostic, respectively (35).

The first prospective multicenter pilot and feasibility study, known as the BENEFIT study, included 54 patients across five different centers, with a median lesion size of 23 mm (IQR, 15–29 mm) (33). Navigational success, assisted by rEBUS, was achieved in 96% of cases, with an estimated diagnostic yield of 74.1%. Stratification by rEBUS view revealed an accuracy of 80.6% for concentric and 70% for eccentric views. The safety profile mirrored that of prior publications, post-procedural pneumothorax occurred in 3.7% (2/54) of patients, one requiring tube thoracostomy (33).

Agrawal et al. conducted a retrospective study on factors



Figure 3 The Galaxy SystemTM comprising of a single platform, arm, screen, and controller (courtesy of Noah Medical).

influencing diagnostic accuracy using the MonarchTM system and rEBUS (38). Evaluating 124 patients with a median PPL size of 20.5 mm, the study reported an 82% navigational success rate and a diagnostic yield of 77% after a 12-month follow-up period. Similar diagnostic yields were observed for concentric (85%) and eccentric views (84%) on rEBUS. Finally, Khan *et al.* published their findings on a large-scale retrospective real-world study, comprising of 264 patients with a median PPL size of 19.3 mm (range, 3.2–72.5 mm) (29). Diagnostic yield achieved was 85.2%, while adverse event rate was 7.6%, including pneumothorax and bleeding.

Recently completed is the TARGET study, a multicenter prospective post-marketing trial, enrolling 1,200 patients across 30 sites undergoing robotic-assisted transbronchial lung biopsy using the MonarchTM platform over a four-year period. The primary endpoint involved the incidence of device or procedure-related complications, while secondary endpoints assessed diagnostic yield and sensitivity for malignancy, procedural details, and post-intervention adverse events, with patient follow-up extending to 24 months (Clinical Trial ID: NCT04182815). A comprehensive summary of findings for MonarchTM is described in *Table 3*.

Galaxy System[™]

The Galaxy System[™] developed by Noah Medical in San

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Carlos, CA, USA, received United States Food and Drug Administration (FDA) approval in March 2023 (*Figure 3*). It employs a disposable, single-use bronchoscope featuring a 4.0-mm OD, 2.1-mm working channel, and integrated vision capabilities. The navigation system implements EMN in conjunction with integrated tomosynthesis technology and augmented fluoroscopy.

During the navigation process, EMN guides the system to within 2 cm of the target lesion. Subsequently, proprietary digital tomosynthesis, known as Tool in Lesion Technology (TiLT+ TechnologyTM), is employed via a standard fluoroscopy C-arm to confirm the tool's precise placement in relation to the lesion. Digital tomosynthesis, akin to CT scanning, involves capturing a series of X-ray images from various angles to reconstruct a 3D image. Notably, this technique requires fewer images taken from a relatively narrower angle (15–60°), compared to the broader range (180°) employed in CT scans (39). This allows for the use of a standard C-arm instead of a dedicated CT scanner.

Positioning itself as a cost-competitive RAB platform, the Galaxy System[™] incorporates a disposable, single-use scope that eliminates the need for reprocessing the scope and contributes to cost savings and procedural turnaround. The system features native fluoroscopy/CBCT integration, also in line of economic benefits. A comprehensive comparative description of RAB with EMN, digital tomosynthesis and TiLT+ to the other RAB systems is presented in *Table 2*.

Evidence

Bhadra *et al.* published the inaugural animal study, titled "The MATCH Study", which explored the application of the Galaxy System[™] (40). This study involved four operators conducting navigation and target reach on four pigs. Each physician performed 4 to 6 nodule biopsies for a total of 20 simulated PPL. The average lesion size was 16.3±0.97 mm, predominantly located in the lower lobes (65%). Results revealed a 100% success rate in navigation, a 100% diagnostic yield, and a tool-in-lesion rate of 95%.

Currently, an ongoing clinical trial aims to confirm the accuracy of the TiLT+ incorporated in the Galaxy System[™]. The trial, with an estimated enrollment of 25 participants, focuses on the primary outcome of successful tool-in-lesion confirmation via CBCT (Clinical Trial ID: NCT06056128). Preliminary results from 15 sampled PPL with a mean size of 20.5 mm were presented in 2023, showing a promising 100% target reach, 86–93% overall diagnostic yield and 3 reported complications (37).

Discussion

Robotic bronchoscopy aims to enhance diagnostic accuracy and safety for pulmonary diagnostic procedures by implementing advanced planning systems for more accurate navigation and biopsy tool delivery, while overcoming the limitations of standard flexible bronchoscopy and percutaneous transthoracic biopsy.

Since its introduction in 2018, RAB has shown increasing diagnostic yields, with observational studies reporting ranges from 76% up to 93% across published literature (Table 3). In a meta-analysis including 20 RAB studies, 1,779 PPL were evaluated. Reported pooled diagnostic vield was 84.5%, while complication rates were 2.3% for pneumothorax, 1.2% for pneumothorax requiring chest tube, and 0.5% for bleeding (41). Two recent metaanalyses on guided bronchoscopy, including EMN, virtual bronchoscopy, and RAB, showed overall diagnostic accuracies of 69.4% [95% confidence interval (CI): 67-71.1%, 126 studies] and 70.9% (95% CI: 68.4-73.2%, 95 studies) (42,43). For RAB specifically, diagnostic yield was 77.6% (43) and 77.3% when combined with CBCT (42), the highest yield among all procedures in both analyses. Adverse events rates remained at 2% for pneumothorax requiring intervention and for bleeding (42,43).

While these meta-analyses have compared RAB to other bronchoscopic approaches, there is virtually no data directly comparing RAB systems, and scarce analysis between RAB and non-bronchoscopic techniques. Computed tomography-guided transthoracic biopsy (CTTB) has historically been the gold standard for diagnosing PPL, consistently demonstrating yields over 80% (44-46). However, its diagnostic performance is hindered by significant rates of pneumothorax and bleeding, ranging from 6-15% and 18%, respectively (47,48). For RAB, MonarchTM has reported pneumothorax rates from 0–5.7% (29,30,33,35). This pneumothorax occurrence was lower in comparison to other published studies involving PPL sampling through standard flexible bronchoscopy and EMN (49,50). With the Ion[™] system, the pneumothorax rates have ranged from 0-7% (6-9,12,13,19). The incidence of substantial airway bleeding following conventional transbronchial biopsy is estimated to be around 2-3% of cases (49,50). The Monarch[™] Platform has disclosed airway bleeding rates from 0-3.2% (28-30,35). Likewise, the IonTM system has exhibited low airway bleeding occurrence, ranging from 0% to 0.8% (6-8,20,51). To date, there are no published data on procedure-related adverse

events using the Galaxy System[™].

In 2023, a retrospective comparative analysis between ssRAB and CTTB by Yu Lee-Mateus et al. revealed similar diagnostic yields (87.6% and 88.4%, respectively) with significantly lower complication rate for ssRAB (4.4% vs. 17%, P=0.001) (19). The same authors later presented an updated assessment between both procedures focused on subsolid nodules, reconfirming comparable diagnostic performances, 85.7% for ssRAB and 89.5% for CTTB, and fewer complications for the robotic approach (7.4% versus 21.2%, respectively) (13). Compared to CTTB, bronchoscopic procedures do not involve puncturing through the chest wall and lung parenchyma to reach PPL, which inherently increases the risk of pneumothorax and bleeding. In contrast to other bronchoscopic approaches, RAB has the advantages of pre-procedural planning based on contiguous thin-cut CT to create a 3D reconstruction of the airway, navigational plan towards the virtual target, live feedback on distance to target and pleura, and a stable robotic arm with thinner working channel that allows for farther reach and interchangeable biopsy tools. These characteristics enable more accurate sampling and consequently, enhance the safety profile of RAB. As robotic bronchoscopy expands worldwide, further research directly comparing RAB systems and against non-bronchoscopic techniques is needed to ascertain the optimal diagnostic procedure for PPL. An upcoming systematic review and network meta-analysis assessing diagnostic accuracy and safety of navigational guided bronchoscopy, including RAB, and CTTB (PROSPERO ID: CRD42023432829) and two ongoing clinical trials facing RAB to CTTB (Clinical Trial ID: NCT04250194) and to standard EMN (Clinical Trial ID: NCT06308120) aim to answer this question.

One significant challenge impacting precise bronchoscopic navigation and sampling is computed-tomography-to-body divergence (CTBD). Regardless of the robotic platform used, target location and navigational planning relay on pre-procedural CT. These capture lung volumes nearing total lung capacity, at full inspiration on breath hold, which differs from the patient's lung volumes during the procedure under general anesthesia and paralysis, controlled mechanical ventilation, and closer to functional residual capacity (52). Additionally, atelectasis may occur during general anesthesia, particularly in the lower lobes, as well as other factors such as mucus plugging, pleural effusions, bleeding, etc. that can cause structural distortions (39). CTBD originates from the difference between the location of the virtual target from pre-procedural CTs and the real-

time intraprocedural location, which has been reported up to a median of 17 mm (52,53). The three robotic bronchoscopy platforms utilize intraprocedural imaging systems to mitigate CTBD. Ion[™] Endoluminal is supported by standard CBCT and mCBCT with the Cios Spin (Cios Spin, Siemens Healthineers, Siemens Medical Solutions, USA). CBCT uses a cone-shaped X-ray beam projecting onto a flat panel detector that is able to produce 3D images with a reconstruction algorithm, to be reviewed during the procedure for assessment of the robotic tip location in relation to the target lesion for potential readjustment. mCBCT functions in a similar manner, implementing the concepts of X-ray based tomosynthesis, able to produce images similar to conventional fixed CBCT, as well as 2D X-ray fluoroscopic images and 3D reconstructions, although not suitable for augmented fluoroscopy (39). With CBCT, IonTM Endoluminal has reported diagnostic yields of 86% and 91.4% (7,9). In a retrospective study on 30 nodules using ssRAB and mCBCT, divergence, defined as a distance greater than 10 mm between the target location in pre-procedure CT and the time of procedure, was found in 50% of cases (20). With mCBCT, which also allows for correction of the navigation plan to the real-time location (39,54), the virtual pathway and biopsy tool were readjusted when necessary and diagnostic yield achieved was 93% (20). Monarch™ has used standard CBCT as well as LungVision[™] (Body Vision Medical Ltd., Ramat Ha Sharon, Israel), a system that combines augmented fluoroscopy with artificial intelligence to generate a navigational pathway from pre-procedure CT and real-time fluoroscopy (52). With LungVision[™], diagnostic accuracy surpassed 90% in a study with 45 patients (31), while with CBCT, sensitivity for malignancy achieved was 86% in 20 lesions (32). The Galaxy System[™] integrates digital tomosynthesis, which relays on computerbased reconstruction algorithms to produce radiographic images with depth of field from multiple single-plane X-ray images over angles as small as 50° and augmented fluoroscopy (39), using standard C-arm CT. As described previously, an animal study assessing reach and feasibility achieved tool-in-lesion confirmation in 95% of cases, and diagnostic yield of 100% (40). The preliminary results of the first human study reported 100% target reach and approximately 90% of overall diagnostic yield (37).

The reliability and precision of RAB have evolved into potential applications in therapeutic bronchoscopy. RAB has been used for the direct injection of liposomal amphotericin B into a chronic *Aspergillus* lesion, with subsequent radiological and clinical improvement (55). Zhong et al. documented the application of RAB in conjunction with photodynamic therapy for a patient with adenoid cystic carcinoma, involving the entire tracheal wall and resulting in lumen obstruction (56). Exploring the administration of chemotherapy and other agents through direct intratumoral injection is currently under consideration (57). This strategy seeks to boost immunotherapy directly within the local tumor microenvironment. RAB may play a crucial role in facilitating nodule marking during the perioperative phase for minimally invasive thoracic surgery, to improve accuracy in localization and resection with negative margins based on the dyes and dye-soaked coils luminescence (58,59). Currently, active clinical trials are in progress to test direct endobronchial ablation (Clinical Trial ID: NCT05299606; Clinical Trial ID: NCT05890872) for local treatment of lung primary and metastatic malignancies.

Addressing competency and skill acquisition for patient safety in the context of advanced robotic bronchoscopy encounters obstacles related to equipment costs, skill acquisition challenges, and overcoming the learning curve for widespread adoption. Tackling these issues requires the implementation of structured training programs that provide thorough education on both equipment operation and procedural techniques. To mitigate variability in the learning curve, standardized training modules and mentorship programs can assist practitioners in mastering skills consistently. Furthermore, enhancing access requires collaborative efforts among institutions to share expertise and resources, potentially through specialized training centers or virtual platforms. The challenge of conducting a cost-to-benefit analysis remains, as average costs for robotic bronchoscopy plus secondary imaging equipment are over six-figure United States dollars, respectively, in addition to all device-specific tools (39). Comprehensive studies to assess overall cost-effectiveness and clinical impact are crucial for making informed decisions regarding resource allocation and promoting broader accessibility of this technology to enhance patient care.

Strengths and limitations

This narrative review aims to summarize the findings on RAB and present a comprehensive comparison of the three approved systems. We acknowledge the nature of the narrative review may limit an exhaustive systematic review approach and most of the data described comprised

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retrospective or observational prospective studies.

Conclusions

Robotic bronchoscopy, comprised by the IonTM, MonarchTM, and GalaxyTM systems, represents an advancement in lung cancer diagnostics. These technologies demonstrate effective diagnostic performance with low complication rates. As RAB continues to expand worldwide, RAB may soon allow a "one-stop shop" approach with diagnosis, staging, and treatment within the same procedure.

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References

1. Siegel RL, Giaquinto AN, Jemal A. Cancer statistics,

2024. CA Cancer J Clin 2024;74:12-49.

- Wolf AMD, Oeffinger KC, Shih TY, et al. Screening for lung cancer: 2023 guideline update from the American Cancer Society. CA Cancer J Clin 2024;74:50-81.
- de Koning HJ, van der Aalst CM, de Jong PA, et al. Reduced Lung-Cancer Mortality with Volume CT Screening in a Randomized Trial. N Engl J Med 2020;382:503-13.
- Pinsky PF, Church TR, Izmirlian G, et al. The National Lung Screening Trial: results stratified by demographics, smoking history, and lung cancer histology. Cancer 2013;119:3976-83.
- Fedewa SA, Bandi P, Smith RA, et al. Lung Cancer Screening Rates During the COVID-19 Pandemic. Chest 2022;161:586-9.
- Fielding DIK, Bashirzadeh F, Son JH, et al. First Human Use of a New Robotic-Assisted Fiber Optic Sensing Navigation System for Small Peripheral Pulmonary Nodules. Respiration 2019;98:142-50.
- Benn BS, Romero AO, Lum M, et al. Robotic-Assisted Navigation Bronchoscopy as a Paradigm Shift in Peripheral Lung Access. Lung 2021;199:177-86.
- Kalchiem-Dekel O, Connolly JG, Lin IH, et al. Shape-Sensing Robotic-Assisted Bronchoscopy in the Diagnosis of Pulmonary Parenchymal Lesions. Chest 2022;161:572-82.
- Styrvoky K, Schwalk A, Pham D, et al. Shape-Sensing Robotic-Assisted Bronchoscopy with Concurrent use of Radial Endobronchial Ultrasound and Cone Beam Computed Tomography in the Evaluation of Pulmonary Lesions. Lung 2022;200:755-61.
- Abia-Trujillo D, Folch EE, Yu Lee-Mateus A, et al. Mobile cone-beam computed tomography complementing shape-sensing robotic-assisted bronchoscopy in the small pulmonary nodule sampling: A multicentre experience. Respirology 2024;29:324-32.
- Abia-Trujillo D, Chandra NC, Koratala A, et al. Diagnostic Yield of Shape-Sensing Robotic-Assisted Bronchoscopy for Ground-Glass Nodules and Subsolid Nodules with a Solid Component Less than 6 mm. Respiration 2023;102:899-904.
- Fernandez-Bussy S, Garza-Salas A, Barrios-Ruiz A, et al. Shape-Sensing Robotic-Assisted Bronchoscopy in the Multiple Pulmonary Nodule Diagnosis during a Single Anesthetic Event. Respiration 2024;103:397-405.
- Fernandez-Bussy S, Yu Lee-Mateus A, Reisenauer J, et al. Shape-Sensing Robotic-Assisted Bronchoscopy versus Computed Tomography-Guided Transthoracic Biopsy for the Evaluation of Subsolid Pulmonary Nodules.

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5432

Respiration 2024;103:280-8.

- 14. Gupta B, Orr L, Bell J, et al. Enhancing diagnostic yield of peripheral pulmonary nodules: a prospective study on shape-sensing robotic-assisted bronchoscopy with ion endoluminal system. Chest 2023;164:A6511-3.
- Hammad Altaq H, Parmar M, Syed Hussain T, et al. The Use of Robotic-Assisted Bronchoscopy in the Diagnostic Evaluation of Peripheral Pulmonary Lesions: A Paradigm Shift. Diagnostics (Basel) 2023;13:1049.
- 16. Low SW, Lentz RJ, Chen H, et al. Shape-Sensing Robotic-Assisted Bronchoscopy vs Digital Tomosynthesis-Corrected Electromagnetic Navigation Bronchoscopy: A Comparative Cohort Study of Diagnostic Performance. Chest 2023;163:977-84.
- Chambers J, Knox D, Leclair T. O-arm CT for Confirmation of Successful Navigation During Robotic Assisted Bronchoscopy. J Bronchology Interv Pulmonol 2023;30:155-62.
- Oberg CL, Lau RP, Folch EE, et al. Novel Robotic-Assisted Cryobiopsy for Peripheral Pulmonary Lesions. Lung 2022;200:737-45.
- Yu Lee-Mateus A, Reisenauer J, Garcia-Saucedo JC, et al. Robotic-assisted bronchoscopy versus CT-guided transthoracic biopsy for diagnosis of pulmonary nodules. Respirology 2023;28:66-73.
- Reisenauer J, Duke JD, Kern R, et al. Combining Shape-Sensing Robotic Bronchoscopy With Mobile Three-Dimensional Imaging to Verify Tool-in-Lesion and Overcome Divergence: A Pilot Study. Mayo Clin Proc Innov Qual Outcomes 2022;6:177-85.
- 21. Tavakoli H, Tao CT, Chung JK, et al. Robotic Assisted Bronchoscopy in Evaluation of Peripheral Lung Nodules (abstract). Am J Respir Crit Care Med 2022;205:A5553.
- 22. Ost D, Pritchett M, Reisenauer J, et al. Prospective multicenter analysis of shape-sensing robotic-assisted bronchoscopy for the biopsy of pulmonary nodules: results from the PRECISE study. Chest 2021;160:A2531-3.
- 23. Ross P, Skabla P, Meyer T. P02.03 Robotic Bronchoscopy: Navigating the Change in Lung Nodule Management. J Thorac Oncol 2021;16:S977-8.
- 24. Bawek S, Bajwa S, Rathore A. 76 Consecutive Cases of Robotic-Assisted Navigational Bronchoscopy at a Single Center. TP137 TP137 THORACIC ONCOLOGY: DIAGNOSIS AND TREATMENT: IP, SURGERY, AND RADIATION. Am J Respir Crit Care Med 2021;203:A4820.

- 25. Ghosh S, Baltaji S, Weksler B, et al. Robotic bronchoscopy: Our first 100 nodules. Chest 2021;160:A2051.
- 26. Pritchett M, Schirmer C. Shape-sensing robotic assisted bronchoscopy for the diagnosis of peripheral pulmonary lesions. Chest 2021;160:A1631-2.
- 27. Folch EE, Pritchett M, Reisenauer J, et al. A Prospective, Multi-Center Evaluation of the Clinical Utility of the Ion Endoluminal System -Experience Using a Robotic-Assisted Bronchoscope System with Shape-Sensing Technology. A110 ADVANCES IN INTERVENTIONAL PULMONOLOGY. Am J Respir Crit Care Med 2020;201:A2719.
- Agrawal A, Ho E, Chaddha U, et al. Factors Associated With Diagnostic Accuracy of Robotic Bronchoscopy With 12-Month Follow-up. Ann Thorac Surg 2023;115:1361-8.
- Khan F, Seaman J, Hunter TD, et al. Diagnostic outcomes of robotic-assisted bronchoscopy for pulmonary lesions in a real-world multicenter community setting. BMC Pulm Med 2023;23:161.
- Manley CJ, Kramer T, Kumar R, et al. Robotic bronchoscopic needle-based confocal laser endomicroscopy to diagnose peripheral lung nodules. Respirology 2023;28:475-83.
- 31. Hedstrom G, Wagh AA. Combining Real-Time 3-D Imaging and Augmented Fluoroscopy with Robotic Bronchoscopy for the Diagnosis of Peripheral Lung Nodules. Chest 2022;162:A2082.
- 32. Cumbo-Nacheli G, Velagapudi RK, Enter M, et al. Robotic-assisted Bronchoscopy and Cone-beam CT: A Retrospective Series. J Bronchology Interv Pulmonol 2022;29:303-6.
- 33. Chen AC, Pastis NJ Jr, Mahajan AK, et al. Robotic Bronchoscopy for Peripheral Pulmonary Lesions: A Multicenter Pilot and Feasibility Study (BENEFIT). Chest 2021;159:845-52.
- 34. Ekeke CN, Vercauteren M, Istvaniczdravkovic S, et al. Lung Nodule Evaluation Using Robotic-Assisted Bronchoscopy at a Veteran's Affairs Hospital. J Clin Med 2021;10:3671.
- 35. Chaddha U, Kovacs SP, Manley C, et al. Robot-assisted bronchoscopy for pulmonary lesion diagnosis: results from the initial multicenter experience. BMC Pulm Med 2019;19:243.
- Rojas-Solano JR, Ugalde-Gamboa L, Machuzak M. Robotic Bronchoscopy for Diagnosis of Suspected

Journal of Thoracic Disease, Vol 16, No 8 August 2024

Lung Cancer: A Feasibility Study. J Bronchology Interv Pulmonol 2018;25:168-75.

- 37. Saghaie T, Williamson J, Philips M, et al. Preliminary results of the FRONTIER STUDY: First Human Use of a New Robotic Electromagnetic Navigation Bronchoscopy with Integrated Toolin-Lesion Tomosynthesis (TiLT) technology (Galaxy System) for Small Peripheral Pulmonary Nodules. 2023 Abstracts. American Association for Bronchoscopy and Interventional Pulmonology; Chicago, IL, USA: Lippincott; 2023:238.
- Agrawal A, Hogarth DK, Murgu S. Robotic bronchoscopy for pulmonary lesions: a review of existing technologies and clinical data. J Thorac Dis 2020;12:3279-86.
- Jain A, Sarkar A, Husnain SMN, et al. Digital Tomosynthesis: Review of Current Literature and Its Impact on Diagnostic Bronchoscopy. Diagnostics (Basel) 2023;13:2580.
- Bhadra K, Rickman OB, Mahajan AK, et al. "Toolin-lesion" Accuracy of Galaxy System-A Robotic Electromagnetic Navigation BroncHoscopy With Integrated Tool-in-lesion-Tomosynthesis Technology: The MATCH Study. J Bronchology Interv Pulmonol 2024;31:23-9.
- Ali MS, Ghori UK, Wayne MT, et al. Diagnostic Performance and Safety Profile of Robotic-assisted Bronchoscopy: A Systematic Review and Meta-Analysis. Ann Am Thorac Soc 2023;20:1801-12.
- 42. Kops SEP, Heus P, Korevaar DA, et al. Diagnostic yield and safety of navigation bronchoscopy: A systematic review and meta-analysis. Lung Cancer 2023;180:107196.
- Nadig TR, Thomas N, Nietert PJ, et al. Guided Bronchoscopy for the Evaluation of Pulmonary Lesions: An Updated Meta-analysis. Chest 2023;163:1589-98.
- 44. An W, Zhang H, Wang B, et al. Comparison of CT-Guided Core Needle Biopsy in Pulmonary Ground-Glass and Solid Nodules Based on Propensity Score Matching Analysis. Technol Cancer Res Treat 2022;21:15330338221085357.
- 45. Wang B, Zhong F, An W, et al. The diagnostic value of CT-guided percutaneous puncture biopsy of pulmonary ground-glass nodules: a meta-analysis. Acta Radiol 2023;64:1431-8.
- 46. Yun S, Kang H, Park S, et al. Diagnostic accuracy and complications of CT-guided core needle lung

biopsy of solid and part-solid lesions. Br J Radiol 2018;91:20170946.

- Heerink WJ, de Bock GH, de Jonge GJ, et al. Complication rates of CT-guided transthoracic lung biopsy: meta-analysis. Eur Radiol 2017;27:138-48.
- 48. Wiener RS, Schwartz LM, Woloshin S, et al. Population-based risk for complications after transthoracic needle lung biopsy of a pulmonary nodule: an analysis of discharge records. Ann Intern Med 2011;155:137-44.
- 49. Folch EE, Pritchett MA, Nead MA, et al. Electromagnetic Navigation Bronchoscopy for Peripheral Pulmonary Lesions: One-Year Results of the Prospective, Multicenter NAVIGATE Study. J Thorac Oncol 2019;14:445-58.
- 50. Pritchett MA, Schampaert S, de Groot JAH, et al. Cone-Beam CT With Augmented Fluoroscopy Combined With Electromagnetic Navigation Bronchoscopy for Biopsy of Pulmonary Nodules. J Bronchology Interv Pulmonol 2018;25:274-82.
- Reisenauer J, Simoff MJ, Pritchett MA, et al. Ion: Technology and Techniques for Shape-sensing Roboticassisted Bronchoscopy. Ann Thorac Surg 2022;113:308-15.
- Ravikumar N, Ho E, Wagh A, et al. Advanced Imaging for Robotic Bronchoscopy: A Review. Diagnostics (Basel) 2023;13:990.
- 53. Chen A, Pastis N, Furukawa B, et al. The effect of respiratory motion on pulmonary nodule location during electromagnetic navigation bronchoscopy. Chest 2015;147:1275-81.
- 54. Yu Lee-Mateus A, Chandra NC, Johnson MM, et al. Robotic-Assisted Bronchoscopy Integrated with Three-Dimensional Fluoroscopy for Sampling Bilateral Pulmonary Nodules in a Single Procedure. Respiration 2023;102:449-53.
- 55. Bawaadam H, Romero AO, Krishna G. Therapeutic Use of Robotic Bronchoscopy System to Treat Chronic Invasive Pulmonary Aspergillosis. J Bronchology Interv Pulmonol 2022;29:e35-8.
- 56. Zhong C, Li H, Huang J, et al. The Novel Application of Robotic-Assisted Bronchoscopy Combined with Photodynamic Therapy for Adenoid Cystic Carcinoma of the Trachea. Respiration 2023;102:961-8.
- 57. DeMaio A, Sterman D. Bronchoscopic intratumoural therapies for non-small cell lung cancer. Eur Respir Rev 2020;29:200028.
- 58. Bawaadam H, Benn BS, Colwell EM, et al. Lung

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Fernandez-Bussy et al. Robotic-assisted bronchoscopy review

Nodule Marking With ICG Dye–Soaked Coil Facilitates Localization and Delayed Surgical Resection. Ann Thorac Surg Short Rep 2023;1:221-5.

59. Chan JWY, Chang ATC, Yu PSY, et al. Robotic Assisted-

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Bronchoscopy With Cone-Beam CT ICG Dye Marking for Lung Nodule Localization: Experience Beyond USA. Front Surg 2022;9:943531.