

Image-guided thoracoscopic surgery with dye localization in a hybrid operating room

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Background: The rate of detection of small pulmonary nodules (SPNs) has increased. Thoracoscopic resection following image-guided localization had been a reliable alternative in their treatment. We present our experience with image-guided dye localization using robotic C-arm computed tomography (CT) followed by immediate video-assisted thoracoscopic surgery (VATS) for SPNs in a hybrid operating room (OR).

Methods: From July 2015 to July 2016, 25 consecutive patients with SPNs smaller than 2 cm underwent robotic C-arm CT-guided blue dye tattooing followed by immediate VATS in a hybrid OR. Their medical records were retrospectively reviewed to evaluate the feasibility and safety of this novel procedure.

Results: Robotic C-arm CT-guided dye localization was successfully performed in 23 patients (92%). Wound extension was required for nodule identification in the remaining two patients. The median size of the nodules was 1.0 cm (range, 0.6–2.0 cm). The median needle localization time and surgery time were 46 and 109 min, respectively. All 25 patients had successful resection of their lesions. The pathological diagnoses were primary lung adenocarcinoma in 18 (72%), benign tumors in 5 (20%), and metastatic lesions in 2 (8%). There was no operative mortality. The median length of the postoperative stay was 3 days (range, 2–8 days). Complications were noted in two patients (8%). One patient had a penetrating injury of the diaphragm during needle localization. The other had pneumonia postoperatively. Both patients were managed conservatively.

Conclusions: Our experience showed that robotic C-arm CT-guided dye localization followed by immediate thoracoscopic surgery in a hybrid OR is safe and feasible. It may become an effective and attractive alternative in managing SPNs.

Keywords: Computed tomography (CT); hybrid operating room; localization; pulmonary nodules; video-assisted thoracoscopic surgery (VATS)

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Introduction

As the National Lung Screening Trial demonstrating screening for lung cancer with chest computed tomography (CT) has reduced mortality from lung cancer (1,2), the use of low-dose CT screening is increasing and has increased the rate of detection of small lung nodules in asymptomatic patients. Most of these CT-detected lesions are indeterminate (3,4). A histopathological diagnosis is recommended when the nodule persists and has a solid component larger than 5 mm, or has enlarged on follow-up CT scans (5). However, an accurate diagnosis using CT-guided percutaneous biopsy is challenging, especially for lesions smaller than 2 cm with a predominately ground glass opacity (GGO) component (6,7). Transbronchial or transthoracic needle biopsy and positron emission tomography scans do not always allow definitive exclusion of malignancy in these lesions (8,9). Thus, patients with small indeterminate lung nodules and suspicious lung cancer lesions are often referred for surgical excision for both diagnostic and curative management.

Small lung lesions which do not cause surface abnormalities may not be identifiable under thoracoscopy. These lesions may require preoperative image-guided localization or wound extension for direct finger palpation, especially for GGOs or small lung nodules. In recent years, uniportal video-assisted thoracoscopic surgery (VATS) has grown in popularity because there is less postoperative pain and better patient satisfaction (10,11). However, finger palpation is not easy to perform under thoracoscopy through only one small incision, which makes preoperative localization even more essential for uniportal thoracoscopic surgery.

Our previous experience demonstrated the efficacy and safety of CT-guided dye localization at CT room followed by uniportal thoracoscopic surgery for resection of lung lesions (12,13). However, postprocedure complications such as pneumothorax, pulmonary hemorrhage, and hemoptysis increased the risks and the burden of nursing care for patients awaiting surgery, even though most of them received conservative management (14,15). By utilizing a hybrid operating room (OR), small lung lesions can be localized and resected precisely as a single procedure in the same place (16-19), which helps avoid the inherent risks of transfer after a transthoracic puncture, and reduces smudging of the injected dye.

Our aim is to provide a new workflow for image-guided dye localization using robotic C-arm CT before a

thoracoscopic procedure in a hybrid OR, and evaluate its safety and feasibility.

Methods

Study design and patients

From July 2015 to July 2016, the medical records of patients who underwent image-guided dye localization of small pulmonary nodules (SPNs) using robotic C-arm CT immediately followed by thoracoscopic surgery in a hybrid OR were reviewed. The selection criteria for this procedure were patients with small peripheral lung nodules sized between 0.5 and 2.0 cm which would be difficult to identify during thoracoscopic surgery (pure or part-solid GGO lesions ≤ 2 cm, or solid nodules ≤ 1 cm). Patients requiring lobectomy or with deep lung nodules with depth > 3 cm were excluded. This study was reviewed and approved by the National Taiwan University Hospital Research Ethics Committee (approval number: 201608041RIND).

Workflow

Hybrid OR setting

The hybrid OR is equipped with a robotic C-arm CT (Artis zeego, Siemens Healthcare GmbH, Forchheim, Germany). The robotic C-arm CT system is able to adapt to almost any position of the integrated free-floating surgical table such as tilted and cradled. The robotic C-arm CT system can be controlled near the surgical table or outside the hybrid OR.

Anesthesia and patient positioning

General anesthesia was performed on the free-floating table and the patients were intubated with either a double-lumen or single-lumen endotracheal tube. According to the location of the lung lesion, the patient was positioned in the supine or lateral decubitus position for the optimal access route for needle insertion and then fixed on the table with positioning aids (*Figure 1A*). During C-arm CT docking, all pipelines from the anesthesia side were gathered to the rostral axis and the C-arm was rotated laterally to check for any entanglement with the patient or lines.

Localization with needle procedure

One of two radiologists (YC Chang, WC Ko) planned the needle routes and performed the dye-localization procedure. After making sure the C-arm could move

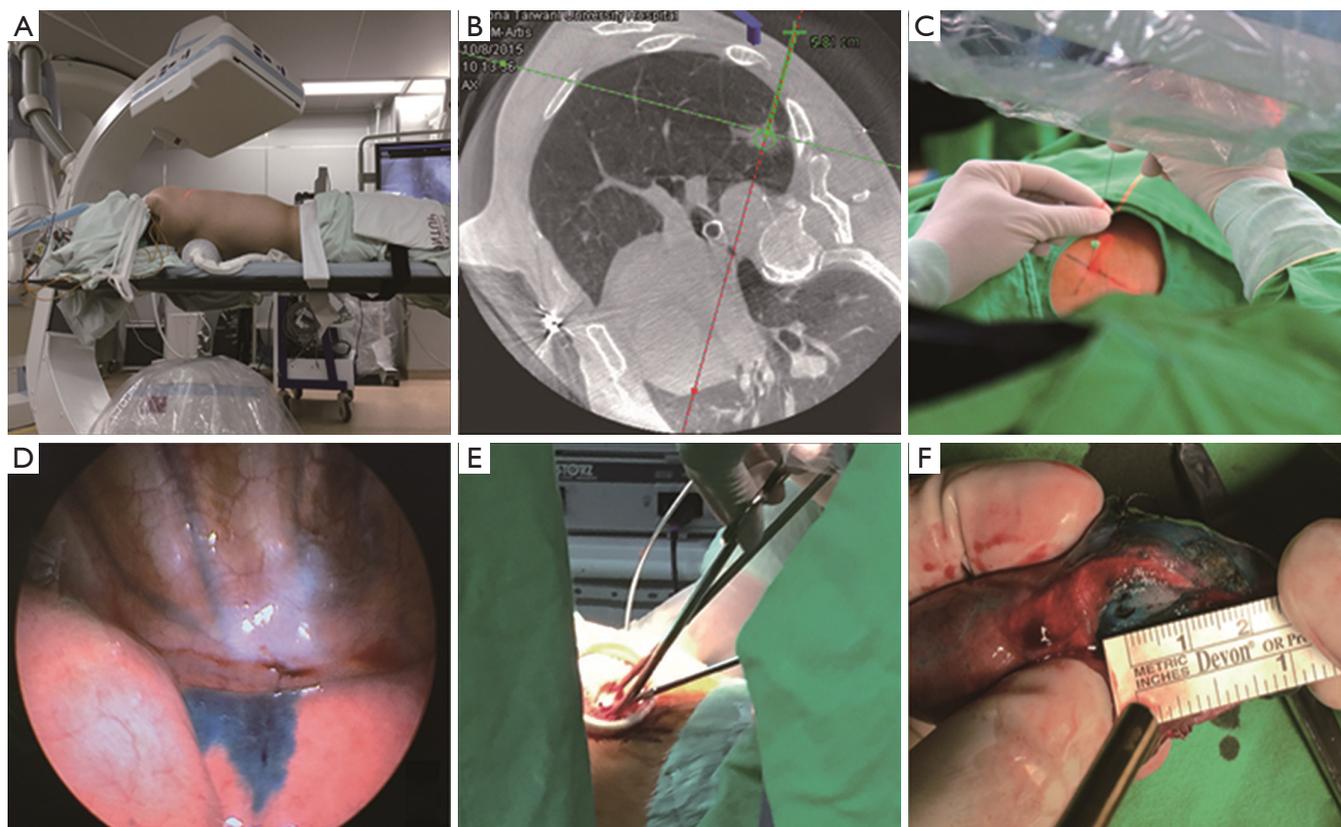


Figure 1 Steps of image-guided thoroscopic surgery. (A) The patient is placed in the lateral decubitus position on the floatable angiography table with an Artis zeego C-arm computed tomography (CT); (B) image-guided needle pathway planning on the C-arm CT image; (C) needle localization through the planned pathway provided by laser beam projection; (D) patent blue vital dye around the target lesion under thoracoscopy; (E) thoracoscopic surgery through a uniportal approach following dye localization; (F) resected lung specimen with a whitish lesion inside and surrounding blue dye.

freely around the patient, an initial C-arm CT volume was acquired (*syngo* DynaCT) for planning the intervention using a 6 sec scan protocol with $0.36 \mu\text{Gy}/\text{projection}$ and 397 frames acquired over 200° during an end-inspiratory hold phase. This was initiated by clamping the endotracheal tube using a hemostat once the ventilator bellow provided the targeted tidal volume. After the protocol was completed, the hemostat was released and the patient resumed ventilation immediately. The access path was laid out in the isotropic data set using the *syngo* Needle Guidance of *syngo* X-Workplace (Siemens Healthcare GmbH, Forchheim, Germany). According to the initially defined paths, the skin entry point and target lesion were selected in *syngo* Needle Guidance, which then calculated the needle path (*Figure 1B*). During the needle procedure, the end-inspiratory hold

phase was created and apnea was sustained 1 minute or longer according to the patient's tolerance to secure the target lesion at the location shown on the last CT image. Before the needle procedure, the planned needle path was projected with a laser beam, making it possible to correct the position and direction of the needle. Under the end-inspiratory hold phase, a 22-gauge Chiba needle was gradually advanced to the planned depth under laser-supported guidance (*Figure 1C*). Thereafter, a second, control C-arm CT volume was acquired to evaluate the needle placement using an identical 6 sec scan protocol before cessation of the end-inspiratory hold phase. After confirmation of the optimal needle location, approximately 0.2 mL of PBV dye (patent blue V 2.5%; Guerbet, Aulnay-sous-Bois, France) was injected into the nodule. The last



Figure 2 This video presented the workflow of image-guide VATS in hybrid OR, which include the following steps (20). Step 1: identify lesion with dynamic CT; step 2: needle procedure planning; step 3: needle placement and dye marking; step 4: dye enabled VATS. VATS, video-assisted thoracoscopic surgery; OR, operating room.

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C-arm CT volume was acquired under the end-inspiratory hold phase to confirm immersion of the injected dye around the target. In some patients with deeply localized nodules, the subpleural lung parenchyma of the localization route was also localized with approximately 0.2 mL of PBV dye after additional C-arm CT confirmation.

Thoracoscopic surgery

After localization was completed, the anesthesiologist placed an endobronchial blocker or utilized a double lumen endotracheal tube to create one lung ventilation. The operation site was sterilized and draped for thoracoscopic surgery. Uniportal or multiportal thoracoscopic wedge resection was performed with the guidance of the blue dye (*Figure 1D,E,F*). If a frozen section of the lesion showed primary lung cancer, mediastinal lymphadenectomy was performed. An additional pulmonary resection was taken if the section margin was less than 2 cm. All incisions were closed, the chest was drained and the patient awakened, extubated, and transferred to the recovery room. The video of this section was shown in *Figure 2*.

Data collection and analysis

The demographic data, CT-localization results, anesthesia results, operative findings, and complications were retrospectively collected by chart review. Descriptive statistics are reported as median (range) for continuous data and as number (%) for categorical data.

Results

From July 2015 to July 2016, 25 consecutive patients with SPNs smaller than 2 cm underwent robotic C-arm CT-guided blue dye tattooing followed by immediate VATS in a hybrid OR. The demographics and baseline characteristics are reported in *Table 1*. Their median age was 59 years and 20 (80%) were women. Their median body mass index was 23.3 kg/m². The median tumor size was 10 mm. The pulmonary function of the patients was generally good, with a median forced expiratory volume in 1 second of 113% of predicted. Coexisting medical diseases were present in 17 patients. At the time of the procedure, the American Society of Anesthesiologists Physical Status Classification was class 1 in 5 patients, class 2 in 7, and class 3 in 13.

Twenty patients (80%) received single lumen endotracheal tube intubation with an endobronchial blocker and the others received a double lumen for general anesthesia and lung separation. Twenty-one patients (84%) were positioned in the lateral decubitus (or semi-prone) position and 4 were supine for the localization procedure. Twenty-three patients (92%) had successful intraoperative dye injection to localize the SPN. *Figure 3* demonstrates images of C-arm CT-guided dye localization. The blue dye immersed around the lesions in these patients was visible on the pleural surface of the lung at thoracoscopy (*Figure 1D*). Two patients (8%) had unexpected events during the needle localization procedure, and both of them received thoracoscopic surgery without adequate localization. One patient with a lesion over the right middle lobe had a puncture through the diaphragm into the liver parenchyma. The other patient with two lesions over the left lower lobe had a large pneumothorax after dye injection for the first lesion. The CT-guided localization procedures were discontinued when these events were noted and thoracoscopic surgery was performed. The lung nodules were identified with the conventional palpation method after extension of the wounds. One patient required addition pulmonary resection because the safety margin of wedge resection was less than 2 cm.

Table 2 showed the results of localization and operation. The median length of the global operation time, defined as the whole duration of the patient in the OR, was 182 (range, 123–300) min. The median duration of anesthesia induction before localization was 10 (range, 5–25) min. The median localization time measured from patient positioning with C-arm docking to the end of the needle procedure was 46 (range, 27–69) min. The median duration

Table 1 Patient demographics and baseline characteristics

Variable	Values (N=25)
Age, years	59 [28–76]
Sex (female)	20 (80.0%)
Height, cm	159.3 (147.1–176.1)
Weight, kg	57.9 (47.8–78.9)
Body mass index, kg/m ²	23.3 (18.5–28.0)
Smoking	2 (8.0%)
FVC (% of prediction)	112.6 (68.4–137.8)
FEV _{1.0} (% of prediction)	113.3 (73.5–155.3)
Nodule size, cm	1.0 (0.6–2.0)
Nodule depth, cm	0.7 (0.4–2.8)
Nodule location (lobe)	
Right upper	3 (12.0%)
Right middle	3 (12.0%)
Right lower	3 (12.0%)
Left upper	7 (28.0%)
Left lower	9 (36.0%)
Malignancy	20 (80.0%)
Comorbidity	
Prior cancer history	8 (32.0%)
Hypertension	8 (32.0%)
Diabetes mellitus	2 (8.0%)
Cardiac diseases	4 (16.0%)
ASA class	
I	5 (20.0%)
II	7 (28.0%)
III	13 (52.0%)

Continuous data are shown as median (range) and categorical data as number (%). FVC, forced vital capacity; FEV_{1.0}, forced expiratory volume in 1 second; ASA, American Society of Anesthesiologists.

of thoracoscopic surgery was 109 (range, 31–185) min. *Syngo* DynaCT was performed intraoperatively an average of 3.6 times (median: 3 times), and the median durations of postoperative chest tube drainage and hospital stay were 1 and 3 days, respectively. *Table 3* provides the pathological diagnoses of the 25 nodules resected. Eighteen (72%)

were primary lung adenocarcinomas, all of which were stage Ia. Two (8%) metastatic lesions were reported and benign disease was diagnosed in 5 (20%) lesions with two organizing pneumonitis, two intrapulmonary lymph nodes, and one atypical adenomatous hyperplasia. *Table 4* provides the surgical and pathological outcomes of the primary lung cancers in this cohort.

Discussion

In recent years, robotic C-arm CT-guided interventions have been used more frequently because of the improving image quality of flat-panel CT and the availability of three-dimensional mapping and navigation applications. A robotic C-arm CT interventional suite can provide real-time visualization for transthoracic needle biopsy (TNB) and more flexibility of the detector system around the patient compared with conventional gantry-based, multidetector CT (MDCT) (21–24). Furthermore, the patient has not to be repositioned and the system complies even with orthopedic hygienic standards. *Rotolo et al.* (21) evaluated 324 TNBs using either C-arm CT or MDCT and demonstrated that C-arm CT-guided TNB provided high and comparable diagnostic accuracy (96% *vs.* 94%) with a similar radiation dose. However, MDCT guidance is only possible in an axial plane, thus making off-plane interventions more difficult and time consuming, whereas, the C-arm CT system is not restricted to a single plane (22). Thus, the C-arm CT guidance enables better angle flexibility for nodule localization, especially in decubitus position.

Several preoperative localization methods, including dye injection (14,25,26), hookwire placement (27–30), and coil labeling (31), have been described for SPNs, and these methods have their advantages and disadvantages. Because hookwire dislodgment and coil migration may occur during patient transfer, we prefer to use dye injection routinely. We initially used methylene blue dye for preoperative CT-guided localization, however, the dye rapidly diffused into the surrounding lung parenchyma in some patients. Therefore, we began using PBV dye for pulmonary nodule localization in 2013, and the localization success rate in our institution was 100% in patients with a single nodule and 97.2% in those with synchronous multiple nodules (25). In our experience, preoperative CT-guided PBV dye localization also contributes to a satisfactory operative outcome for SPNs managed with uniportal VATS (12,13), which comprised 80% of cases in our cohort. In this study, the success rates of localization using PBV injection under C-arm CT guidance

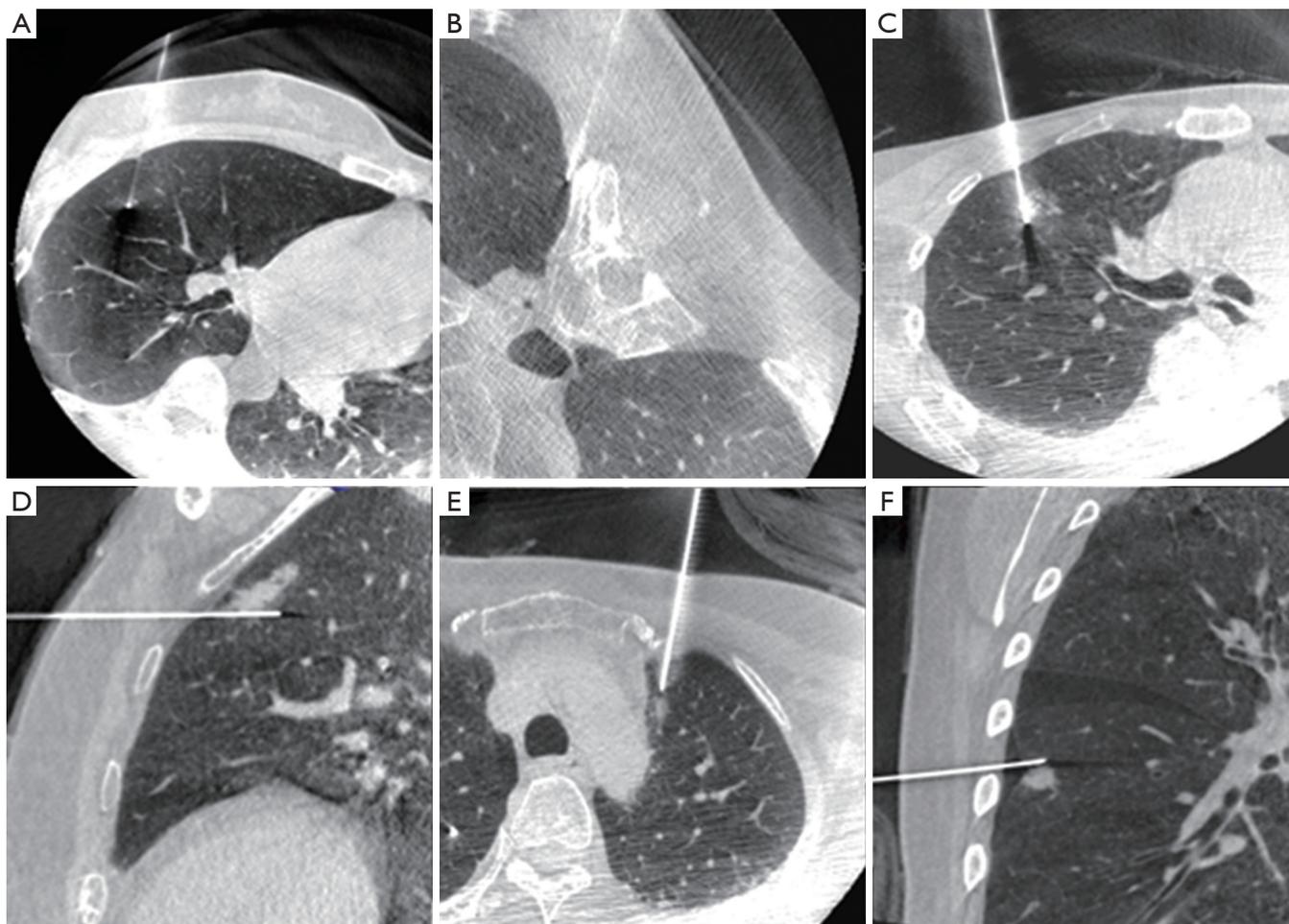


Figure 3 Series needle procedure under C-arm CT guidance. (A) A 6 mm small ground glass opacity in the right lower lung lobe; (B) one small metastatic lesion located in the left upper lobe near the spine; (C) a right upper lobe lung lesion approached from the supine position; (D) localization planning using sagittal CT image; (E) one small left upper lobe lesion near the aortic arch is localized successfully using C-arm CT; (F) small pneumothorax detected during the needle localization procedure.

was 92% which is slightly lower than MDCT guidance (25). We speculate that the learning curve using C-arm CT guidance and limited case number may contribute the difference. Also the C-arm CT patients were intubated with positive ventilation, which may lead to a higher risk of a large pneumothorax during a needle procedure.

Several groups have demonstrated successful combination of intraoperative CT-guided localization and immediate thoracoscopic surgery for SPNs (16-18). Narayanam *et al.* (16) described a series of 31 pediatric patients with a total of 34 SPNs receiving lung tattooing with immediate video-assisted thoracoscopic resection performed as a single procedure in a hybrid room. Lung tattooing was performed under conventional gantry-based MDCT. The technical

success rate was 91.1% and the diagnostic yield was 100%. Ohtaka *et al.* (17) used mobile CT for intraoperative pulmonary nodule localization. Ten patients with small lung nodules underwent VATS using an intraoperative imaging device with a flat-panel detector that provides CT imaging. The positional relationship between the needle marking and the tumor was recognized intraoperatively. In 9 patients (90%), the tumor could be seen on intraoperative CT images using their mobile CT.

Another group from Harvard Medical School (18) developed a prospective phase I-II clinical trial to facilitate localization and resection of small lung nodules. In their study, an intraoperative C-arm CT scan was utilized for guidance of percutaneous marking with T-bars followed

Table 2 Localization and operation results

Variable	Values (N=25)
DynaCT for localization (times)	3 [2–6]
Global OR time (min)	182 [123–300]
Anesthesia induction time (min)	10 [5–25]
Needle localization time (min)	46 [27–69]
Surgery time (min)	109 [31–185]
Successful localization	23 (92%)
Chest tube drainage (days)	1 (0–2)
Post-op hospital stay (days)	3 [2–8]
Complications	2 (8%)
Diaphragm injury during localization	1 (4%)
Pneumonia	1 (4%)

Continuous data are shown as median (range) and categorical data as number (%). OR, operating room.

Table 3 Diagnoses of the nodules

Diagnoses	Number [%] (N=25)
Primary lung adenocarcinoma	18 [72]
Metastatic cancer from	2 [8]
Thyroid cancer	1 [4]
Sarcoma	1 [4]
Benign lung tumors	5 [20]
Organizing pneumonitis	2 [8]
Intrapulmonary lymph node	2 [8]
Atypical adenomatous hyperplasia	1 [4]

Table 4 Surgical and pathological results of primary lung cancer

Variable	Number (%) (N=18)
Adenocarcinoma	18 (100.0)
Stage Ia	18 (100.0)
R0 resection	18 (100.0)
Post-op hospital stay [range] days	3 [2–8]
Post-op complications	
Pneumonia	1 (5.5)

Continuous data are shown as median [range] and categorical data as number (%).

by VATS resection of the tumor. All 23 patients underwent complete resection of their lesions and CT imaging of the resected specimens confirmed the removal of the T-bars and the nodule. The median time required for placement of the two T-bars and for the VATS were 39 and 67 min, respectively, which saved a lot of time compared with the first MDCT-based series and also our results (average localization procedure: 33 min; VATS: 109 min).

Selection of a hybrid OR table depends on the primary use of the system. Surgeons, particularly orthopedic, general, and neurosurgeons, usually expect a segmented tabletop for flexible patient positioning (32,33). Thoracic surgeons also require a flexible table to help bridging, which is opening up the intercostal spaces (ICS) when the patient is in the lateral decubitus position. However, most hybrid ORs were originally set up for cardiac and vascular procedures and are usually equipped with floatable angiography tables. Ng *et al.* (19) described their substitution for bridging by placing a rolled up pillow under the patient's mid thorax to open up the ICS when doing an image-guided localization uniportal lobectomy in a hybrid OR. In our series, the patients were positioned laterally without trying to open up the ICS. Patients with anticipated anatomical pulmonary resections were excluded from our project, because those procedures are thought to be more difficult and require standard surgical tables. Measures must be taken to avoid touching the patient and entangling accessory lines and tubing during C-arm rotation before acquiring *syngo* DynaCT images. Ng *et al.* (19) suggested that the anesthesia equipment be placed further from the patient than usual to ensure collision-free C-arm rotation. This may require extension tubing. In our practice, the anesthesia equipment is located as usual, and all tubes and lines are secured together extending from the patient to the machine through the axis of the table, preventing entanglement with the rotating C-arm.

In our series, one patient had a failed localization due to large pneumothorax noted on real-time imaging after the needle puncture. Under positive pressure ventilation, the incidence of a large pneumothorax after thoracic needle procedures may rise compared with procedures done under spontaneous breathing. In future practice, we will use catheter drainage or create a thoracic incision as salvage for a large pneumothorax during needle localization to complete the procedure. Another patient had a diaphragm puncture into the liver parenchyma, which is usually encountered with lesions located at the basal portion close to the diaphragm. With lesions at the basal portion, care

should be taken before the needle enters the pleural cavity. *Syngo DynaCT* should be performed to prevent unexpected diaphragm puncture when the needle tip is still in the chest wall. Alternative localization methods without puncture into the pleural cavity have been described in Japanese centers (34,35). These can also be considered to avoid complications.

The success of this program depends on close cooperation between different specialties, including anesthesiologists, intervention radiologists, technicians, and surgeons. To gather these experts at a scheduled time to go through a time-consuming process in the hybrid OR, which was in paucity and usually controlled by cardiovascular specialist, is the most challenging part. For the practitioners wishing to start such a program in their own institution, we suggest starting for peripheral and probably palpable lesions in case of unsuccessful localization, and to avoid lesions near the diaphragm, at the base of lower lobe, or area covered by scapula. A special consideration should be focused on the general anesthesia during needle localization. Although some of the center prefer to maintain contralateral lung ventilation during needle localization, our workflow suggest totally apnea with inflated lung for better image acquisition and accuracy of localization.

Conclusions

In conclusion, we presented a new workflow for image-guided dye localization with an immediate thoracoscopic procedure for SPNs in a hybrid OR. Through this multidisciplinary approach, patients can be safely managed with successful resection for both diagnostic and therapeutic needs in one stop under a single anesthetic.

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Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

Ethical Statement: This study was reviewed and approved by

the National Taiwan University Hospital Research Ethics Committee (approval number: 201608041RIND).

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