

World Journal of *Clinical Cases*

World J Clin Cases 2022 September 6; 10(25): 8808-9179



MINIREVIEWS

- 8808** Ear, nose, and throat manifestations of COVID-19 and its vaccines
Al-Ani RM
- 8816** Potential influences of religiosity and religious coping strategies on people with diabetes
Onyishi CN, Eseadi C, Itechukwu LC, Okoro KN, Okolie CN, Egbule E, Asogwa E

ORIGINAL ARTICLE**Case Control Study**

- 8827** Effectiveness of six-step complex decongestive therapy for treating upper limb lymphedema after breast cancer surgery
Zhang HZ, Zhong QL, Zhang HT, Luo QH, Tang HL, Zhang LJ

Retrospective Study

- 8837** Hospital admissions from alcohol-related acute pancreatitis during the COVID-19 pandemic: A single-centre study
Mak WK, Di Mauro D, Pearce E, Karran L, Myintmo A, Duckworth J, Orabi A, Lane R, Holloway S, Manzelli A, Mossadegh S

- 8844** Indocyanine green plasma clearance rate and ^{99m}Tc-galactosyl human serum albumin single-photon emission computed tomography evaluated preoperative remnant liver
Iwaki K, Kaihara S, Kita R, Kitamura K, Hashida H, Uryuhara K

- 8854** Arthroscopy with subscapularis upper one-third tenodesis for treatment of recurrent anterior shoulder instability independent of glenoid bone loss
An BJ, Wang FL, Wang YT, Zhao Z, Wang MX, Xing GY

- 8863** Evaluation of the prognostic nutritional index for the prognosis of Chinese patients with high/extremely high-risk prostate cancer after radical prostatectomy
Yang F, Pan M, Nie J, Xiao F, Zhang Y

Observational Study

- 8872** Chlorine poisoning caused by improper mixing of household disinfectants during the COVID-19 pandemic: Case series
Lin GD, Wu JY, Peng XB, Lu XX, Liu ZY, Pan ZG, Qiu ZW, Dong JG

- 8880** Mental health of the Slovak population during COVID-19 pandemic: A cross-sectional survey
Kralova M, Brazinova A, Sivcova V, Izakova L

Prospective Study

- 8893** Arthroscopic anatomical reconstruction of lateral collateral ligaments with ligament advanced reinforcement system artificial ligament for chronic ankle instability
Wang Y, Zhu JX

SYSTEMATIC REVIEWS

- 8906** How to select the quantitative magnetic resonance technique for subjects with fatty liver: A systematic review
Li YW, Jiao Y, Chen N, Gao Q, Chen YK, Zhang YF, Wen QP, Zhang ZM
- 8922** Lymphocytic choriomeningitis virus: An under-recognized congenital teratogen
Ferenc T, Vujica M, Mrzljak A, Vilibic-Cavlek T

CASE REPORT

- 8932** Alagille syndrome associated with total anomalous pulmonary venous connection and severe xanthomas: A case report
Zeng HS, Zhang ZH, Hu Y, Zheng GL, Wang J, Zhang JW, Guo YX
- 8939** Colo-colonic intussusception with post-polypectomy electrocoagulation syndrome: A case report
Moon JY, Lee MR, Yim SK, Ha GW
- 8945** Portal vein gas combined with pneumatosis intestinalis and emphysematous cystitis: A case report and literature review
Hu SF, Liu HB, Hao YY
- 8954** Quadricuspid aortic valve and right ventricular type of myocardial bridging in an asymptomatic middle-aged woman: A case report
Sopek Merkaš I, Lakušić N, Paar MH
- 8962** Treatment of gastric carcinoma with lymphoid stroma by immunotherapy: A case report
Cui YJ, Ren YY, Zhang HZ
- 8968** Gallstone associated celiac trunk thromboembolisms complicated with splenic infarction: A case report
Wu CY, Su CC, Huang HH, Wang YT, Wang CC
- 8974** Extracorporeal membrane oxygenation for lung cancer-related life-threatening hypoxia: A case report
Yoo SS, Lee SY, Choi SH
- 8980** Multi-disciplinary treatment of maxillofacial skeletal deformities by orthognathic surgery combined with periodontal phenotype modification: A case report
Liu JY, Li GF, Tang Y, Yan FH, Tan BC
- 8990** X-linked recessive Kallmann syndrome: A case report
Zhang P, Fu JY
- 8998** Delayed complications of intradural cement leakage after percutaneous vertebroplasty: A case report
Ma QH, Liu GP, Sun Q, Li JG

- 9004** Coexistent Kaposi sarcoma and post-transplant lymphoproliferative disorder in the same lymph nodes after pediatric liver transplantation: A case report
Zhang SH, Chen GY, Zhu ZJ, Wei L, Liu Y, Liu JY
- 9012** Misdiagnosis of pancreatic metastasis from renal cell carcinoma: A case report
Liang XK, Li LJ, He YM, Xu ZF
- 9020** Discoid medial meniscus of both knees: A case report
Zheng ZR, Ma H, Yang F, Yuan L, Wang GD, Zhao XW, Ma LF
- 9028** Simultaneous laparoscopic and arthroscopic excision of a huge juxta-articular ganglionic cyst compressing the sciatic nerve: A case report
Choi WK, Oh JS, Yoon SJ
- 9036** One-stage revision arthroplasty in a patient with ochronotic arthropathy accompanied by joint infection: A case report
Wang XC, Zhang XM, Cai WL, Li Z, Ma C, Liu YH, He QL, Yan TS, Cao XW
- 9044** Bladder paraganglioma after kidney transplantation: A case report
Wang L, Zhang YN, Chen GY
- 9050** Total spinal anesthesia caused by lidocaine during unilateral percutaneous vertebroplasty performed under local anesthesia: A case report
Wang YF, Bian ZY, Li XX, Hu YX, Jiang L
- 9057** Ruptured splenic artery aneurysms in pregnancy and usefulness of endovascular treatment in selective patients: A case report and review of literature
Lee SH, Yang S, Park I, Im YC, Kim GY
- 9064** Gastrointestinal metastasis secondary to invasive lobular carcinoma of the breast: A case report
Li LX, Zhang D, Ma F
- 9071** Post-bulbar duodenal ulcer with anterior perforation with kissing ulcer and duodenocaval fistula: A case report and review of literature
Alzerwi N
- 9078** Modified orthodontic treatment of substitution of canines by first premolars: A case report
Li FF, Li M, Li M, Yang X
- 9087** Renal cell carcinoma presented with a rare case of icteric Stauffer syndrome: A case report
Popov DR, Antonov KA, Atanasova EG, Pentchev CP, Milatchkov LM, Petkova MD, Neykov KG, Nikolov RK
- 9096** Successful resection of a huge retroperitoneal venous hemangioma: A case report
Qin Y, Qiao P, Guan X, Zeng S, Hu XP, Wang B
- 9104** Malignant transformation of biliary adenofibroma combined with benign lymphadenopathy mimicking advanced liver carcinoma: A case report
Wang SC, Chen YY, Cheng F, Wang HY, Wu FS, Teng LS

- 9112 Congenital hepatic cyst: Eleven case reports
Du CX, Lu CG, Li W, Tang WB
- 9121 Endovascular treatment of a ruptured pseudoaneurysm of the internal carotid artery in a patient with nasopharyngeal cancer: A case report
Park JS, Jang HG
- 9127 Varicella-zoster virus meningitis after spinal anesthesia: A case report
Lee YW, Yoo B, Lim YH
- 9132 Chondrosarcoma of the toe: A case report and literature review
Zhou LB, Zhang HC, Dong ZG, Wang CC
- 9142 Tamsulosin-induced life-threatening hypotension in a patient with spinal cord injury: A case report
Lee JY, Lee HS, Park SB, Lee KH
- 9148 CCNO mutation as a cause of primary ciliary dyskinesia: A case report
Zhang YY, Lou Y, Yan H, Tang H
- 9156 Repeated bacteremia and hepatic cyst infection lasting 3 years following pancreatoduodenectomy: A case report
Zhang K, Zhang HL, Guo JQ, Tu CY, Lv XL, Zhu JD
- 9162 Idiopathic cholesterol crystal embolism with atheroembolic renal disease and blue toes syndrome: A case report
Cheng DJ, Li L, Zheng XY, Tang SF
- 9168 Systemic lupus erythematosus with visceral varicella: A case report
Zhao J, Tian M

LETTER TO THE EDITOR

- 9176 Imaging of fibroadenoma: Be careful with imaging follow-up
Ece B, Aydin S

ABOUT COVER

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The primary aim of *World Journal of Clinical Cases* (*WJCC*, *World J Clin Cases*) is to provide scholars and readers from various fields of clinical medicine with a platform to publish high-quality clinical research articles and communicate their research findings online.

WJCC mainly publishes articles reporting research results and findings obtained in the field of clinical medicine and covering a wide range of topics, including case control studies, retrospective cohort studies, retrospective studies, clinical trials studies, observational studies, prospective studies, randomized controlled trials, randomized clinical trials, systematic reviews, meta-analysis, and case reports.

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Extracorporeal membrane oxygenation for lung cancer-related life-threatening hypoxia: A case report

Seung Soo Yoo, Shin Yup Lee, Sun Ha Choi

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Abstract

BACKGROUND

Life-threatening hypoxia can occur in patients with lung cancer due to bronchial obstruction. Extracorporeal membrane oxygenation (ECMO) can be used as a bridge therapy for patients with severe hypoxia not relieved by conventional mechanical treatment. However, the usefulness of chemotherapy in patients with lung cancer receiving ECMO therapy is not well known.

CASE SUMMARY

A 53-year-old man visited the emergency room with worsening dyspnea for 1 mo. A series of imaging and diagnostic tests were performed, and stage IIIB (cT4N2M0) lung cancer was eventually diagnosed. On hospital day 3, he experienced dyspnea and hypoxia that was not relieved with oxygen support *via* a high-flow nasal cannula. ECMO was initiated because his respiratory condition did not improve even with mechanical ventilation. The patient then underwent gemcitabine/cisplatin chemotherapy without dose reduction while on ECMO. After two cycles of chemotherapy, there was a decrease in the size of the primary tumor in the right main bronchus. After the completion of concurrent chemoradiotherapy, a computed tomography scan revealed further improvement in the right main bronchus narrowing. Eight months after a lung cancer diagnosis, the patient did well without any dyspnea.

CONCLUSION

ECMO is a potential bridge therapy for respiratory failure in patients with central airway obstruction secondary to lung cancer.

Key Words: Extracorporeal membrane oxygen; Lung cancer; Hypoxia; Respiratory failure; Chemoradiotherapy; Case report

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Core Tip: Extracorporeal membrane oxygenation can be selected as an important salvage treatment for patients with severe cardiopulmonary dysfunction caused by lung cancer and other malignant tumors until the patients are stabilized or even cured.

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INTRODUCTION

Lung cancer is one of the most commonly diagnosed malignancies and is the leading cause of cancer-related deaths worldwide, with an average 5-year survival rate of 19% [1]. The clinical presentation of lung cancer varies among individuals, resulting in a range of symptoms that patients may find distressing [2]. Several common signs and symptoms associated with lung cancer can be classified as a result of the primary tumor, intrathoracic spread, distant metastases, paraneoplastic syndromes, or nonspecific symptoms [2]. Airway obstruction, superior vena cava syndrome, and massive pleural effusion secondary to lung cancer may cause dyspnea. Moreover, sudden airway obstruction can cause ventilation/perfusion mismatch or shunt, leading to life-threatening respiratory failure.

Extracorporeal membrane oxygenation (ECMO) is the only way to overcome hypoxia that persists even with conventional mechanical ventilator treatment. Patients with hematological malignancies, such as leukemia or lymphoma, may develop respiratory failure related to disease or chemotherapy, and in these cases, ECMO may be required for bridge treatment [3,4]. However, there are only a few reports on the use of chemotherapy during ECMO support as a treatment for respiratory failure in lung cancer, which has a low response rate and delayed response time compared with hematologic malignancy. Herein, we report the case of a patient treated with chemotherapy using ECMO for acute respiratory failure due to lung cancer.

CASE PRESENTATION

Chief complaints

In March 2021, a 53-year-old man visited the emergency room with worsening dyspnea.

History of present illness

The patient's dyspnea started 1 mo ago and has progressed since then.

History of past illness

The patient is a 32 pack-year smoker.

Personal and family history

The patient had a free personal and family history.

Physical examination

On admission, his vital signs were as follows: Blood pressure, 126/80 mmHg; body temperature, 36.5 °C; heart rate, 112 bpm; and respiratory rate, 20 breaths/min.

Laboratory examinations

Atrial blood gas analysis (ABGA) showed a pH of 7.47, PCO₂ of 27 mmHg, PaO₂ of 67.7 mmHg, HCO₃ of 19.8 mmHg, and SpO₂ of 94.8% on room air.

Imaging examinations

Chest radiography revealed a suprahilar mass (Figure 1A), and a chest computed tomography (CT) scan showed a 4.9-cm mass in the right upper lobe (RUL). The mass on the CT scan appeared to cause narrowing of the right main bronchus and pulmonary artery (Figure 1B). He was admitted for evaluation of the lung mass. On day 2 of hospitalization, flexible bronchoscopy revealed an obstructing

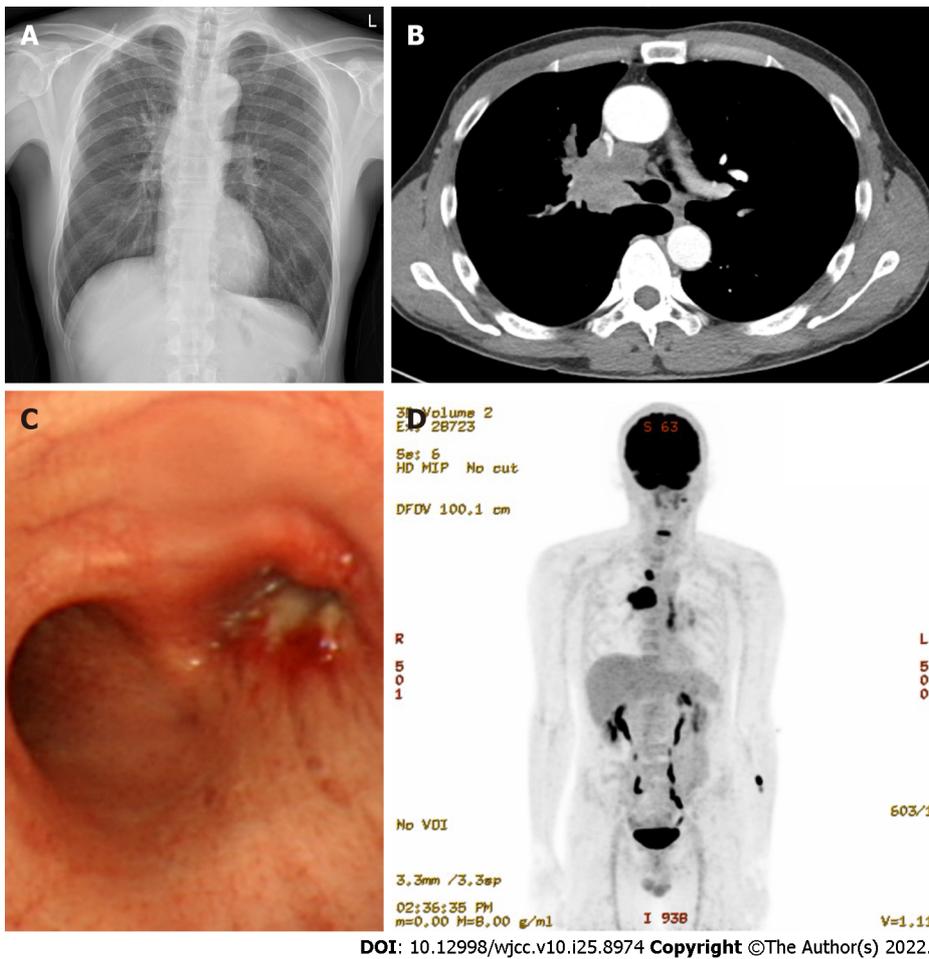


Figure 1 Initial chest radiography, chest computed tomography scan, bronchoscopy, and positron emission tomography-computed tomography. A: Chest radiography reveals a mass-like shadow in the right supra-hilar area; B: Chest computed tomography (CT) showing narrowing of the right main bronchus secondary to a mass; C: Bronchoscopy confirms that the right bronchus is almost completely obstructed by a mass; D: Positron emission tomography-CT showing a 5-cm hypermetabolic mass (SUVmax 20.8) in the right upper lobe and hypermetabolic 4R and 2R lymph nodes.

mass in the right main bronchus (Figure 1C). Bronchoscopic biopsy and endobronchial ultrasound-guided transbronchial needle aspiration (EUBS-TBNA) were performed for the 4R lymph nodes. Upon suspicion of lung cancer, positron emission tomography (PET)-CT was performed (Figure 1D).

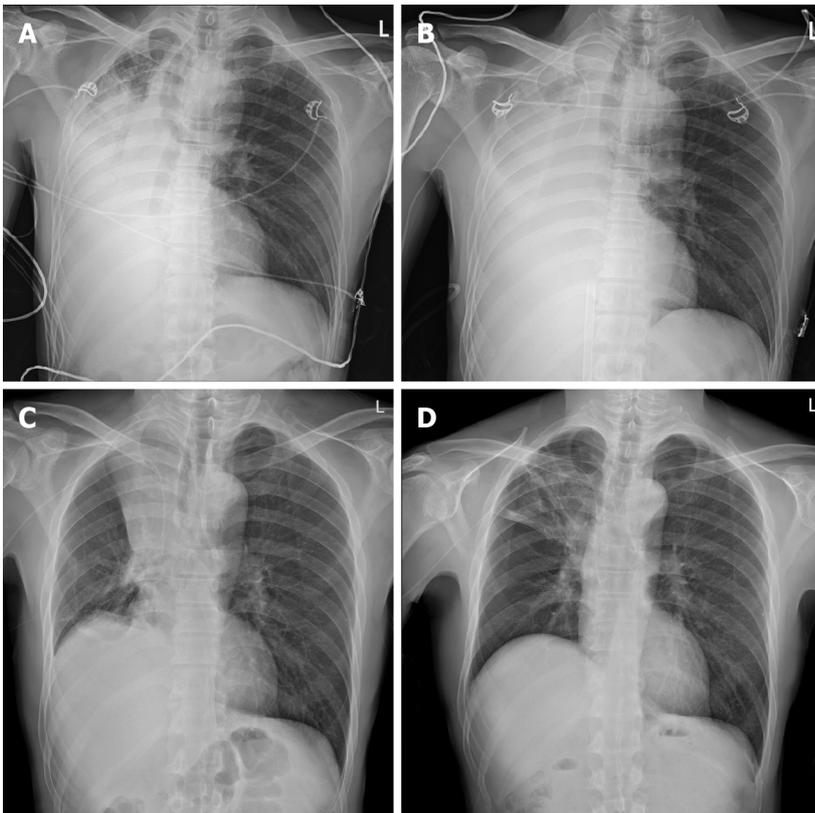
FINAL DIAGNOSIS

The final diagnosis of the presented case is stage IIIB (cT4N2M0) lung cancer, squamous cell carcinoma.

TREATMENT

On hospital day 3, the dyspnea suddenly worsened. The O₂ saturation in room air decreased from 98% to 71%. Chest radiography revealed a nearly complete collapse of the right lung (Figure 2A). A reserve mask with an oxygen flow rate of 15 L/min was used, which improved the O₂ saturation to 88%-90%. A high-flow nasal cannula (HFNC) was administered, and the patient was then transferred to the intensive care unit.

On hospital day 4, ABGA showed a pH of 7.301, PCO₂ of 39 mmHg, PaO₂ of 44.4 mmHg, HCO₃⁻ of 18.4 mmHg, and SpO₂ of 75.9% on oxygen supply *via* an HFNC with FiO₂ of 80% and flow rate of 60 L/min. Mechanical ventilation support was started with lung-protective mechanical ventilation and pronation, according to the current protocols. However, the respiratory condition did not improve, and the patient eventually presented with refractory hypoxemia with a PaO₂/FiO₂ ratio of 46.7, FiO₂ of 100%, and positive end-expiratory pressure (PEEP) of 12 cmH₂O. Accordingly, an urgent decision was made to initiate venovenous ECMO (CardioHelp; Getinge Critical Care, Gothenburg, Sweden) by percutaneous cannulation of the right femoral vein (25-Fr draining cannula) and right jugular vein (17-Fr restitution



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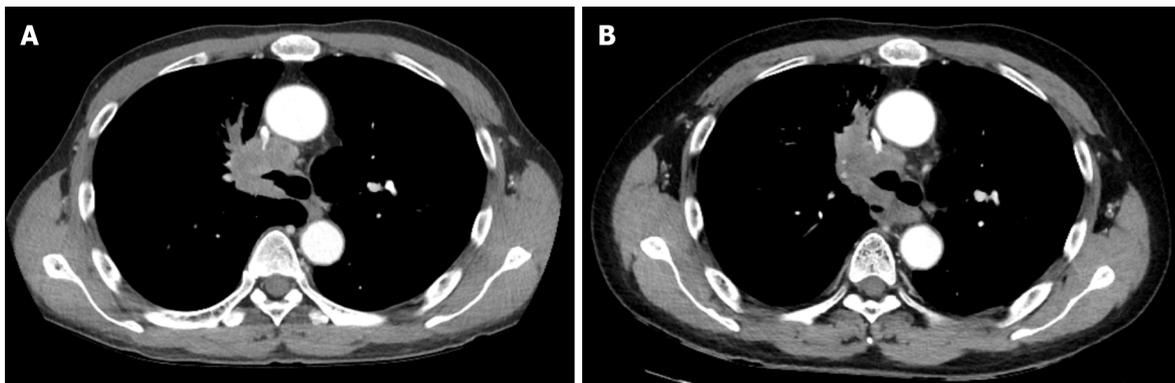
Figure 2 Changes on chest radiography. A: On hospital day 3, chest radiography shows near total atelectasis of the right lung; B: On hospital day 5, atelectasis in the right lung is aggravated; C: On hospital day 16, the near total atelectasis of the right lung has improved, although right upper lobe atelectasis remains; D: On hospital day 19, right upper lobe atelectasis has improved.

cannula). At the start of the ECMO run, blood flow was set at 4 L/min, sweep gas flow was set at 2 L/min, and unfractionated heparin was infused to match a partial thromboplastin time ratio of 1.5- to 2-fold that of the baseline. Ultra-protective mechanical ventilation was instituted in pressure-controlled ventilation mode with a driving pressure of 8 cmH₂O, PEEP of 12 cmH₂O, respiratory rate of 8 breaths/min, and FiO₂ of 60%. Chest radiography revealed an aggravated collapse of the right lung (Figure 2B). ABGA showed pH of 7.462, PCO₂ of 25.9 mmHg, PaO₂ of 373.1 mmHg, HCO₃ of 20.9 mmHg, and SpO₂ of 99.8% on ECMO and ventilator support.

Biopsy of the mass in the right main bronchus and 4R lymph nodes revealed squamous cell carcinoma. The patient was diagnosed with stage IIIB (cT4N2M0) lung cancer. On hospital day 6, the patient was extubated and in spontaneous breathing supported by HFNC oxygen therapy at 35 L/min with a FiO₂ of 45% (awake ECMO). On hospital day 7, gemcitabine/cisplatin chemotherapy without dose reduction was initiated while the patient was on ECMO support. The doses of gemcitabine and cisplatin were 1250 mg per body surface area (BSA) and 70 mg/BSA, respectively. ECMO therapy was discontinued on day 12 of hospitalization, while HFNC was continued. On day 15 of hospitalization, the HFNC was discontinued. On hospital day 16, the total atelectasis of the right lung had improved, although RUL atelectasis remained (Figure 2C). On hospital day 20, the RUL atelectasis improved (Figure 2D). The patient was discharged on hospital day 26 without dyspnea.

OUTCOME AND FOLLOW-UP

After two cycles of gemcitabine/cisplatin chemotherapy, a CT scan showed a decrease in the primary tumor size in the right main bronchus (Figure 3A). The patient received concurrent chemoradiotherapy with a weekly paclitaxel/cisplatin regimen. After the completion of concurrent chemoradiotherapy, the CT scan revealed further improvement in the right main bronchus narrowing due to the mass (Figure 3B). Eight months after lung cancer diagnosis, the patient did well without any dyspnea.



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Figure 3 Changes on chest computed tomography scan. A: After two cycles of gemcitabine/cisplatin chemotherapy, the size of the right upper lobe mass has decreased; B: After concurrent chemoradiotherapy, the right main bronchus, narrowed by the tumor, has further improved.

DISCUSSION

Lung cancer can cause atelectasis, which can lead to shortness of breath. In this case, sudden obstruction of the right main bronchus caused severe hypoxia, which was not relieved with oxygen supply. The mechanism of refractory hypoxia can be explained by a shunt, implying perfusion to the non-ventilated lung. Radiotherapy or endobronchial treatment with a stent is an option for intraluminal obstruction of the main bronchus. However, the patient in the current case did not have the time to receive such treatment because of life-threatening hypoxia.

ECMO can provide cardiac and pulmonary support in critically ill patients with cardiogenic shock or acute respiratory failure[5]. The use of ECMO in cancer patients who develop severe acute respiratory distress syndrome (ARDS) remains controversial[6]. However, several studies have reported the successful use of ECMO in patients who develop respiratory failure after chemotherapy for hematologic malignancies[3,4,6,7]. Chung *et al*[8] reported that under ECMO support, the patient underwent bronchial stent implantation and was successfully weaned off ECMO. Airway obstructions may qualify for intensive respiratory support compared with diffuse parenchymal malignancy because bronchial intervention procedures enable quick recovery from respiratory failure if airway stenosis is relieved[7, 8]. In the case of solid tumors, respiratory failure may occur either during the disease itself or during the treatment course; however, only a few studies have been conducted on ECMO use in these patients. Because of the adverse effects of ECMO, specifically, infections, bleeding, and thrombocytopenia, the administration of chemotherapy is not routinely advised in patients suffering from malignancies[9]. Our experience, in this case, indicates that ARDS caused by shunting in patients with central airway lung cancer should not be considered a contraindication to ECMO treatment. With our successful treatment outcome with chemotherapy in ECMO, we suggest that chemotherapy during ECMO in solid tumors should be considered in the presence of severe hemodynamic instability or respiratory failure refractory to standard medical support.

The pharmacokinetics of chemotherapy with ECMO is not yet fully understood. Several factors may affect pharmacokinetics, including the circuit of ECMO, the changes in the volume distribution, and the alteration in drug clearance. Sherwin *et al*[9] reported pharmacokinetic parameters and dose recommendations for anti-infective drugs, including antibiotics, under ECMO treatment, primarily in neonates. However, information on the pharmacokinetics of anti-cancer drugs during ECMO is still lacking. The current case showed that standard doses of gemcitabine and cisplatin were effective in ECMO treatment, although further studies on chemotherapy with ECMO in solid tumors are needed. Numerous clinical studies will be necessary to determine the appropriate indications for ECMO.

CONCLUSION

In conclusion, ECMO may play an important role as a bridge therapy for respiratory failure due to ARDS in patients with central airway obstruction secondary to lung cancer. As such, this treatment approach must be considered in these patients.

FOOTNOTES

Author contributions: Yoo SS and Choi SH conceived and designed the experiments; Yoo SS and Choi SH performed

the experiments; Yoo SS, Lee SY, and Choi SH acquired clinical data; Yoo SS, Lee SY, and Choi SH wrote the main manuscript text; all authors reviewed the manuscript.

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