

# Cerebral carbon dioxide embolism after kidney cancer laparoscopic surgery with full neurological recovery

## A case report

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### Abstract

**Rationale:** Cerebral carbon dioxide embolism (CCDE) is a rare cause of stroke and is a recognized life-threatening complication. CCDE may result from direct intravascular CO<sub>2</sub> insufflation during surgery. Due to the lack of typical clinical manifestations, the disease is often missed or mistaken for another condition. The clinical signs and symptoms depend on the speed and volume of embolized gas entering the blood and the patient's condition. In particular, patent foramen ovale has been found to be associated, in rare cases, with the intraoperative entry of gas into the arterial system.

**Patient concerns:** In this report, we present the case of a 35-year-old woman with kidney cancer who underwent laparoscopic right partial nephrectomy.

**Diagnosis:** After the laparoscopic surgery, the patient was initially diagnosed with acute cerebral infarction.

**Interventions:** The patient was treated according to the standard method for treatment of acute cerebrovascular disease.

**Outcomes:** Three days after the laparoscopic procedure, the patient gained consciousness, and she was discharged without any neurologic sequelae on postoperative day 12.

**Lessons subsections as per style:** Due to the low incidence and sudden occurrence of CCDE, there is a strong likelihood of missed diagnosis or misdiagnosis, and it is; therefore, important to be aware of the risk. The findings from this report would be highly useful as a reference to clinicians in the future.

**Abbreviations:** BP = blood pressure, CCDE = cerebral carbon dioxide embolism, CO<sub>2</sub> = carbon dioxide.

**Keywords:** cerebral carbon dioxide embolism, laparoscopic surgery, neurological recovery, treatment options

### 1. Introduction

Laparoscopic surgery is a common technique in abdominal surgery, and its application in clinical practice is on the rise. In laparoscopic procedures, gas insufflation is usually used for

accurate surgical visualization and manipulation. Carbon dioxide (CO<sub>2</sub>) is the most commonly used gas for insufflation because it is colorless, inexpensive, and nonflammable.<sup>[1]</sup> Further, it has higher blood solubility than air, and is; therefore,

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YL and EZ contributed equally to this paper.

As it is a case report, ethics approval is not necessary after consulting the Ethics Committee of Peking University Third Hospital.

Written informed consent was obtained from the patient for publication of this case report and any accompanying images.

All data generated or analyzed during this study are included in this article.

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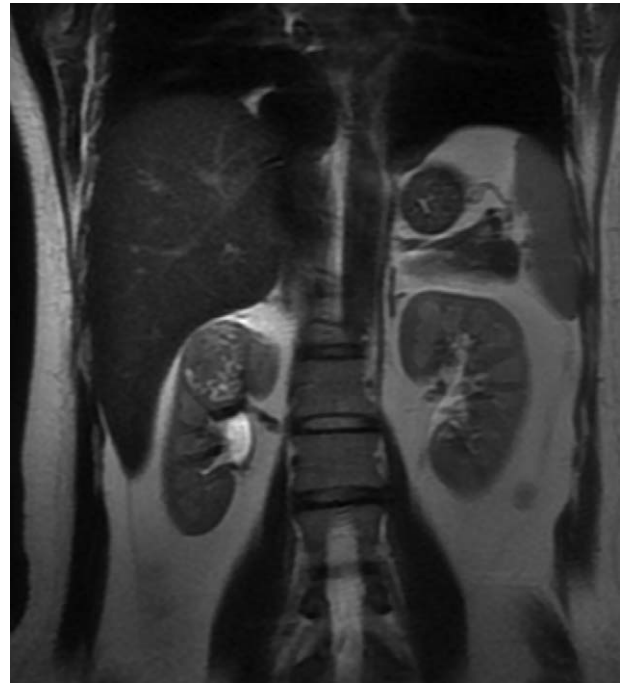
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associated with a reduced risk of complications in case of venous embolism. Although laparoscopic surgery with CO<sub>2</sub> insufflation is generally a beneficial procedure, it does have the disadvantage of potentially causing complications such as subcutaneous emphysema, hypercapnia, respiratory acidosis, cerebral edema, and CO<sub>2</sub> embolism.<sup>[2,3]</sup> Clinically significant CO<sub>2</sub> embolism is rare and unnoticeable, but it is potentially fatal.<sup>[4]</sup> The clinical presentation of CO<sub>2</sub> embolism ranges from asymptomatic to neurologic injury, cardiovascular collapse or even death, and is dependent on the speed and volume of embolized gas entering the blood and the condition of the patient. CO<sub>2</sub> embolism is typically venous and may also cause pulmonary embolism and acute myocardial infarction. More rarely, the gas is able to cross from venous to arterial circulation, and this is referred to as a “paradoxical embolus.”<sup>[5]</sup> In paradoxical CO<sub>2</sub> embolism, CO<sub>2</sub> from the right heart flows into the left heart through the intracardiac or extracardiac shunt, and patients with patent foramen ovale are likely to develop cerebral carbon dioxide embolism (CCDE), which is a rare, but serious neurological deficit with fatal consequences.<sup>[6]</sup> Published articles have reported that most patients with CCDE exhibit a transitional stage of delayed recovery from general anesthesia, temporary hemiplegia, and disturbance of consciousness.<sup>[7]</sup> Here, we present a well-documented case of a patient with patent foramen ovale who had a stroke associated with CCDE that developed after laparoscopic surgery for kidney cancer. The patient was later discharged as she did not exhibit any neurologic sequelae.

## 2. Case presentation

A 35-year-old female patient (height, 165 cm; weight, 62 kg) presented to our hospital with pain in the right back for more than 1 month. On physical examination, her abdomen appeared soft and nontender, with no palpable masses. Abdominal computed tomography (CT) and magnetic resonance imaging (MRI) revealed a 4.2 × 3.2 × 2.6 cm solid mass in the upper pole of the right kidney (Fig. 1). Renal cell carcinoma was suspected, and the patient was scheduled for a laparoscopic right partial nephrectomy. According to her medical history, she had undergone post-laparoscopic cholecystectomy and surgery for ectopic pregnancy, and was allergic to penicillin. She did not have a family history of renal cell carcinoma or genetic abnormalities.

In preparation for the procedure, general anesthesia was induced and maintained according to the standard protocol. The preoperative heart rate was 88 beats/min; systolic blood pressure (BP), 138/68 mm Hg; and oxygen saturation, 99% to 100%. The patient was placed in the left lateral decubitus position. CO<sub>2</sub> insufflation was used to establish pneumoperitoneum and maintain the intra-abdominal pressure at 12 mm Hg. Extraperitoneal fat from the surface of the lateral vertebral fascia was removed, and the perirenal fascia was opened along the leading edge of the psoas major. The peripheral lymphatic vessels of the renal pedicle were cut off with an ultrasonic knife, and the renal artery was detached from the dorsal side of the hilum. The kidney fat capsule was opened and attached to the surface of the kidney to dissociate the kidney. Titanium clamps were used to trap small peripheral blood vessels and bleeding points. Laparoscopic arteriotomy forceps were used to block the renal artery. The tumor and some kidney tissue located 0.5 to 1.0 cm away from the tumor edge were removed. The tumor was removed through the expanded camera port incision and placed in an organ bag. The renal pelvis and parenchyma were joined by continuous



**Figure 1.** MRI demonstrating a lesion consistent with a primary renal neoplasm. MRI = magnetic resonance imaging.

suture. The renal artery was opened, and any oozing blood was wiped off the wound surface. The bleeding area was pressed down with a hemostatic gauze. The procedure time and warm ischemia time were 186 minutes and 25 minutes, respectively. The estimated blood loss volume during the procedure was 50 mL, and the patient did not require blood transfusion. No vein or artery was injured, and no complications occurred during the surgery. The patient remained hemodynamically stable.

After surgery, the patient was moved to the anesthesia recovery room, and her vital signs were found to be stable after extubation (body temperature, 37.1°C; heart rate, 86/min; BP, 112/63 mm Hg; breathing rate, 20 breathes/min; and oxygen saturation, 92%–95%). The patient was breathing spontaneously, but was still deeply sedated and vomiting frequently. She was also responsive to orbital pressure and pain stimulation. Her clinical symptoms included left hemiparesis, positive left-sided Babinski sign, suspicious positive left-sided Hoffman sign, and increased right facial muscle tension. The left and right pupils were of the same size (about 4 mm in diameter), and the patient was positive for the doll’s eye test. In both eyes, the stare was fixed to the top left area of the field of vision. Wrinkling and teeth motion on the right side of the face were observed on pain stimulation. Pain stimulation in the right limb induced some activity, but pain stimulation in the left upper limb and left lower limb resulted in little and no obvious activity, respectively.

An urgent consultation was arranged with the department of neurology, in order to examine the patient for cerebral infarction and cerebral hemorrhage. An emergency CT scan (taken approximately 3 hours after surgery) showed no bleeding foci or occupying lesions (Fig. 2). This was indicative of cerebral infarction, but not intracerebral hemorrhage, so the latter was excluded. The patient received oxygen ventilation via a mask at a rate of 2 L/min, and was moved to the intensive care unit and connected to a mechanical ventilator. In order to assess for the

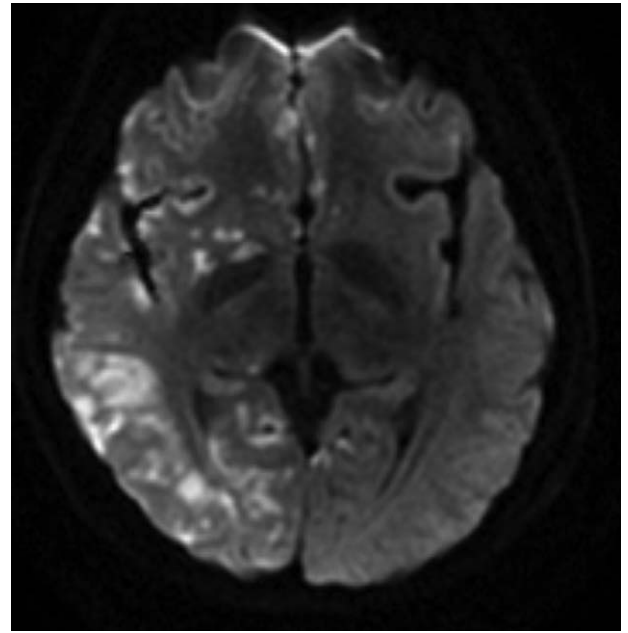


**Figure 2.** Cerebral CT scan taken 3 h after the surgical procedure showing no obvious abnormalities. CT = computed tomography.

presence of acute cerebrovascular diseases, emergency brain MRI and cerebrovascular angiography were recommended. The patient was administered midazolam, butorphanol, norepinephrine, and vinpocetine. However, no obvious abnormalities were observed in cerebral blood vessels of the head and neck in the angiography findings. The patient had no previous history of



**Figure 3.** Follow-up CT scan taken 2 d later showing a hypodense lesion in the right hemisphere. CT = computed tomography.



**Figure 4.** Brain MRI showing paradoxical CO<sub>2</sub> embolism. The diffusion-weighted imaging (DWI) image revealed multiple cerebral infarction, predominantly on the right side of the cerebral cortex on day 5 after the procedure. MRI = magnetic resonance imaging.

hypertension, diabetes, hyperlipidemia, or abnormality in a local echocardiograph. Plavix and aspirin were prescribed, and treatment for maintaining BP was administered by the neurologists at the department. A repeat CT scan taken 2 days later showed that there was a hypodense lesion in the right hemisphere, but there was no sign of cerebrovascular gas (Fig. 3). Mannitol was administered for intracranial dehydration; edaravone, to eliminate free radicals present in the brain; and dexamethasone, to relieve edema. Apart from symptomatic treatment, plavix and aspirin were administered as antiplatelet therapy. On day 4, the patient regained consciousness but had slow reaction times. On day 5 after the surgery, brain MRI showed multiple cerebral infarction located predominantly on the right side of the cerebral cortex (Fig. 4). Contrast-enhanced echocardiography of the right heart was performed to determine if she had right-to-left systemic shunt in the lung or heart. The result was indicative of patent foramen ovale. Based on the results of brain CT, MRI, cerebrovascular angiography, and contrast echocardiography of the right heart, the patient was eventually diagnosed with CCDE. On day 9, the patient had generally recovered and was discharged with completely normal neurological functions on day 12.

### 3. Discussion and conclusion

The high blood solubility of CO<sub>2</sub> in plasma can reduce the risk of CO<sub>2</sub> embolization during laparoscopic surgery via rapid absorption and elimination of CO<sub>2</sub> through the lungs. However, if a large amount of CO<sub>2</sub> enters into the vascular system, it can lead to venous CO<sub>2</sub> embolism, which is rarely seen in humans.<sup>[6]</sup> The incidence rate of gas embolism during laparoscopic surgery is approximately 0.15%, and the mortality rate can be as high as 30%.<sup>[8]</sup> CO<sub>2</sub> embolism has been reported in several laparoscopic procedures, including laparoscopic liver resection, laparoscopic

**Table 1****Previous reports on cases of CO<sub>2</sub> embolism as a complication of laparoscopic radical nephrectomy or partial nephrectomy.**

Author	Patient	Procedure	Year
Bruyère M <sup>[10]</sup>	A 70-yr-old woman	Retroperitoneal laparoscopic right radical nephrectomy	2001
Umegaki T <sup>[11]</sup>	A 76-yr-old man	Laparoscopic partial nephrectomy	2007
Huang YY <sup>[12]</sup>	A 76-yr-old woman	Laparoscopic nephrectomy	2008
Yu X <sup>[13]</sup>	A 52-yr-old woman	Laparoscopic nephrectomy converted to open procedure	2014

cholecystectomy, laparoscopic appendectomy, and gynecological laparoscopy.<sup>[9]</sup> The presence of a gas in venous circulation may obstruct pulmonary circulation, and clinically significant CO<sub>2</sub> embolism can be asymptomatic or even present with cardiac symptoms, including cardiovascular collapse and neurological deficits, and even death.<sup>[9]</sup> In patients with patent foramen ovale, CO<sub>2</sub> embolism may take the form of a very rare but fatal paradoxical gas embolism that leads to cerebral gas embolism.

In the present case report, a 35-year-old female patient with kidney cancer who underwent laparoscopic right partial nephrectomy. Although the surgical procedure was successfully completed with perioperative vein or artery injury, cerebral infarction induced by a gas embolism still occurred. To our knowledge, this is the first report of cerebral paradoxical gas embolism with complete neurological recovery after laparoscopic nephrectomy.

During laparoscopic surgery, it is important to be aware of the possibility of CO<sub>2</sub> embolism and the potential risk factors, including venous bleeding, high ventilatory pressure, patent foramen ovale, and patient positioning. Prompt and accurate recognition and management can limit the clinical impact of such events. The symptoms of gas embolism include systemic hypotension, tachypnea, dyspnea, cyanosis, tachycardia or bradycardia, arrhythmia, and asystole. Paradoxical embolism may also be associated with altered mental status, focal neurological deficits, and loss of consciousness. In our case, a right-to-left shunt caused paradoxical gas embolism when CO<sub>2</sub> entered through the vein. No obvious vascular injury was observed during the operation (probably because the patient was young), and no obvious bubbles were found in the brain CT scan. This indicates that the embolism was caused by a very small volume of CO<sub>2</sub> gas. Further, the patient remained hemodynamically stable during the surgery, and this might explain her quick recovery.

We searched PubMed for previously reported cases of complications caused by CO<sub>2</sub> embolism associated with laparoscopic radical nephrectomy or partial nephrectomy. The following keywords were used: “carbon dioxide embolism,” “nephrectomy,” and “laparoscopy,” as shown in Table 1.<sup>[10–13]</sup> The findings of the identified studies indicate that CO<sub>2</sub> embolism is extremely rare in cases of laparoscopic nephrectomy. We also researched the incidence of CCDE during laparoscopic procedures by searching PubMed with the keywords “cerebral gas embolism,” “carbon dioxide,” and “laparoscopy,” and have summarized the patients’ backgrounds, treatments administered, and outcomes in Table 2.<sup>[7,14–20]</sup> Overall, the findings indicate that clinically significant CO<sub>2</sub> embolism may be fatal.

Preoperative medical evaluation should be conducted to understand the cardiopulmonary risks of a procedure and anticipate possible complications. When gas embolism is suspected, the pneumoperitoneum must be released and the patient must be placed in the Trendelenburg position. Further, if a gas embolism is suspected at any point of time, the surgeon should be notified immediately and insufflation should be discontinued. In order to reduce ventilation-perfusion mismatch and improve hypoxemia, CO<sub>2</sub> should be washed out with 100% oxygen ventilation.<sup>[21]</sup> In addition to symptomatic treatment, prevention of cerebral edema, and hyperbaric oxygen therapy, the amount of gas entering the brain could be reduced by keeping the head in a lower position.<sup>[22]</sup> For patients with severe cardiovascular collapse, supportive treatment with fluid, vasopressors, and cardiopulmonary bypass may be necessary. Currently, there is little literature on standard treatment for cerebral CO<sub>2</sub> embolism, so the prevention of this condition is very important.

Due to the low incidence and sudden occurrence of CCDE, the possibility of missed diagnosis or misdiagnosis exists. Therefore, it is important to be aware of the risk of this condition, and the

**Table 2****Previous reports on the incidence of cerebral carbon dioxide embolism (CCDE) during laparoscopic procedures.**

Author	Surgery	Treatments	Outcomes
Root B <sup>[14]</sup>	Diagnostic laparoscopy and partial ovarian cystectomy	Closed chest massage and administration of tropine, sodium bicarbonate, isoproterenol, metaraminol, and calcium chloride	Sudden death at about 1 h after instillation of CO <sub>2</sub>
Gomar C <sup>[15]</sup>	Laparoscopy and hysterectomy	Administration of tropine, adrenaline, sodium bicarbonate and calcium chloride	Irreversible brain damage and death 1 wk later
Schindler E <sup>[16]</sup>	Laparoscopic cholecystectomy	Intravenous administration of epinephrine (5 µg), corticosteroids and mannitol	Discharge without any neurologic sequelae
Ghimouz A <sup>[17]</sup>	Hysteroscopy-curettage and laparoscopy	Cardiopulmonary resuscitation and intravenous administration of lignocaine	Blindness for 3 h
Benitez PO <sup>[18]</sup>	Laparoscopic cholecystectomy	Advanced cardiopulmonary resuscitation	Sequelae and a prolonged hospital stay
Van Gorp V <sup>[19]</sup>	Laparoscopic partial hepatectomy	Administration of 1 unit of packed cells, and 100% oxygen ventilation with positive-end expiratory pressure	Complete recovery
Kawahara T <sup>[20]</sup>	Laparoscopic liver resection	Cooling of the head and administration of thiamylal and botox	Coma and quadriplegia

findings from this report might prove useful to clinicians in the future.

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### Author contributions

YL and ELZ designed the idea and wrote the manuscript. YL and ELZ collected the data. HSY edited the manuscript. All authors read and approved the manuscript.

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