INTRODUCTION

The simultaneous occurrence of achalasia and esophageal diverticula is rare. There have been few reports of esophageal cancer in achalasia and esophageal diverticula. Although most patients with only esophageal diverticula are asymptomatic, patients with both achalasia and esophageal diverticula may present with dysphagia or regurgitation of food. Additionally, the risk of esophageal cancer is increased in patients with both achalasia and esophageal diverticula. Similar to the pathogenesis of esophageal achalasia, chronic inflammation caused by repeated injury and chronic irritation by stagnated food are thought to result in esophageal carcinogenesis. There have been a few reports of esophageal cancer in achalasia with multiple esophageal diverticula. Our group published one such case report in 2009 regarding a case of achalasia associated with multiple large esophageal diverticula. This is a follow-up of the aforementioned case, in which esophageal carcinoma was subsequently diagnosed in the diverticula that occurred simultaneously with esophageal achalasia.

CASE REPORT

A 68-year-old man was diagnosed with achalasia associated with multiple esophageal diverticula. He had symptoms of dysphagia and regurgitation of food for 2 years. Thus, barium esophagography, esophageal manometry, and esophagogastroduodenoscopy (EGD) were performed. The endoscopic findings showed multifocal lugol-voiding lesions identified as moderate dysplasia. We decided to use photodynamic therapy to treat the multifocal dysplastic lesions. At follow-up EGD 2 months after photodynamic therapy, more lugol-voiding lesions representing a squamous cell carcinoma in situ were found. The patient ultimately underwent surgery for the treatment of recurrent esophageal multifocal neoplasia. After a follow-up period of 3 years, the patient showed a good outcome without symptoms. To manage premalignant lesions such as achalasia with esophageal diverticula, clinicians should be cautious, but have an aggressive approach regarding endoscopic surveillance.

Key Words: Esophageal achalasia; Diverticulum; Esophageal; Esophageal; Neoplasms
of achalasia associated with lower pressures. His symptoms improved after botulinum toxin injection. However, dysphagia, regurgitation of food, and epigastric discomfort recurred 5 years later. A follow-up EGD showed a slight increase in the size of the previous diverticula, mild erythematous mucosal changes, and multifocal lugol-voided lesions (Fig. 3). The biopsy result showed a moderate degree of epithelial dysplasia. Because the lesions were located in a large and tortuous diverticulum, performing endoscopic submucosal dissection (ESD) was considered too difficult and dangerous. Photodynamic therapy (PDT) was therefore considered a better option and was administered. Two months after PDT, EGD was repeated to evaluate the response to PDT (Fig. 4). Mucosal nodularity was observed at a nontreated diverticulum, which on biopsy was revealed to be squamous cell carcinoma in situ (Fig. 5). The patient underwent an Ivor-Lewis esophagectomy with regional lymph node dissection for recurrent multifocal esophageal neoplasia, and the final pathological staging was TisN0M0 (stage 0). The patient showed no evidence of disease at the 3-year follow-up.

**DISCUSSION**

Patients with achalasia can develop a variety of complications. Among them, the most serious may be esophageal cancer. The overall prevalence of esophageal squamous cell
carcinoma in patients with achalasia has been estimated to be 3%, accounting for a 50-fold increase in cancer risk. In a recent, large prospective cohort study with long-term (mean, 15 years) follow-up data of 448 achalasia patients, the relative hazard ratio for the development of esophageal cancer in achalasia was 28. Although many cases of esophageal carcinoma in patients with diverticula have been reported, the precise level of cancer risk in esophageal diverticulum remains to be elucidated.

In both achalasia and esophageal diverticulum, swallowed foods stagnate easily. This food and fluid stasis leads to repeated injury of the esophageal epithelium and chronically irritates the mucosal surface. In addition, bacterial overgrowth and impaired clearance of regurgitated gastric acid contents occur during persistent esophageal distention. Due to these diverse factors, chronic inflammation may occur in the esophageal mucosa, which is presumed to develop epithelial hyperplasia, multifocal dysplasia, and finally esophageal cancer.

Before a diagnosis of cancer, the duration of symptoms is usually at least 15 years in patients with achalasia and at least 10 years in patients with esophageal diverticulum. A large prospective cohort study was conducted with a mean follow-up period of 15 years after the onset of achalasia symptoms. Elderly patients have an increased risk of carcinoma. A diverticulum of less than 2 cm rarely develops into esophageal cancer.

Treatment for esophageal dysplasia or carcinoma arising from achalasia or esophageal diverticulum follows the same principles as esophageal dysplasia or carcinoma without underlying esophageal etiology. Esophageal dysplasia or early esophageal cancer can be treated by endoscopic resection, classic endoscopic mucosal resection (EMR), and ESD, which is widely used. In the present case, however, the diverticula were much too large and tortuous, making EMR or ESD too risky; thus, we performed an operation.

The optimal time and interval for the surveillance of patients with achalasia and/or diverticulum is open-ended. If a patient presents with newly developed symptoms such as weight loss, melena, or hematemesis, endoscopic evaluation should be performed. If not, there is no strong recommendation for endoscopic surveillance. Regarding achalasia, the current version of the American Society of Gastrointestinal Endoscopy guidelines recommends that surveillance should be initiated 15 years after the onset of achalasia symptoms, although the subsequent surveillance interval is not defined. As in the present case, however, patients with both achalasia and diverticula should be considered for endoscopic surveillance earlier than 15 years, as the earlier development of symptoms necessitates shorter surveillance intervals. Moreover, clinicians should consider chromoendoscopy or narrow-band image endoscopy as well as white light endoscopy for surveillance of these patients.

Although the simultaneous occurrence of achalasia and esophageal diverticulum is unusual, this case emphasizes the high risk of esophageal carcinoma arising from both etiologies, as well as the importance of early detection by aggressive endoscopic surveillance.

Conflicts of Interest

The authors have no financial conflicts of interest.

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