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A Rare Complication: Post-Chemotherapy Esophageal Stricture

ABSTRACT

Dysphagia in patients undergoing cancer treatment is often associated with conditions such as reflux esophagitis, infectious esophagitis, malignant infiltration, or as a consequence of radiation therapy, the occurrence of an acute esophageal stricture resulting from chemotherapy is exceedingly rare. In this report, we present a distinctive case of an isolated chemotherapy-induced esophageal stricture in a patient who was undergoing treatment for metastatic osteosarcoma. Notably, this patient had no previous history of gastroesophageal reflux disease, caustic ingestion, or other risk factors commonly linked to the development of esophageal strictures.

Keywords: post chemotherapy ,oesophageal stricture, dysphagia, chemotherapy ,

1. INTRODUCTION

Esophageal strictures are typically classified into two main categories: peptic and non peptic in their aetiology. Peptic strictures arise from prolonged gastroesophageal reflux disease. Non peptic strictures, on the other hand, have diverse causes, including infection, complications following surgery, tumour growth, exposure to toxic substances, or localized radiation exposure.[1] The occurrence of esophageal stricture resulting from chemotherapy alone is an exceedingly rare phenomenon. As far as our knowledge extends, there have been only two reported cases in adult patients [2,3]. In the case of paediatric patients, it has been recognized as an infrequent complication following induction therapy for acute leukaemia [4].

2. CASE PRESENTATION

13 years aged boy was referred for evaluation of progressive dysphagia of 7 months duration. He was diagnosed with osteosarcoma, T1N0M0 of left knee 14 months earlier and started with neoadjuvant chemotherapy. Because of vomiting he was evaluated at 4th month of treatment and found to have mucositis, hepatitis, pancytopenia, COVID 19 positive

40 serology. Chemotherapy stopped and he recovered with conservative management. He underwent above knee
41 amputation a month earlier and referred for grade IV dysphagia. He had hair loss and discolouration of nails. Endoscopy
42 showed narrowing at 25 cms of oesophagus (Fig.1). Barium swallow also showed stricture of oesophagus and chest
43 skiagram(Fig.2) and CT Chest(Fig.3) showed cystic lesions in the both lungs. He underwent graded oesophageal dilation
44 with Savory – Gilliard dilators after discussion with parents (Fig 4). His dysphagia was grade II with periodic dilation. Six
45 months later he failed to follow up and died 1 month later.
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48
49 **FIG 1**

FIG 2

FIG 3

FIG 4

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52 **Fig 1. Endoscopy showed narrowing at 25 cms of oesophagus**

53 **Fig 2. Barium swallow showed stricture of oesophagus and chest skiagram**

54 **Fig 3. CT of Chest**

55 **Fig 4. Graded oesophageal dilation with Savory – Gilliard dilators**
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58 **3. DISCUSSION**

59 Dysphasia in individuals with compromised immune systems is typically attributed to opportunistic infections, often
60 caused by *Candida albicans*, herpes simplex virus, or cytomegalovirus. In both immunocompromised and
61 immunocompetent individuals, dysphagia can also result from conditions such as reflux esophagitis, caustic ingestion,
62 esophageal webs, and foreign body obstruction. However, the patient's medical history did not indicate any such factors
63 contributing to his symptoms, and both endoscopic and histopathologic assessments ruled out these possibilities. While
64 esophageal strictures are a recognized complication following radiation therapy, this patient had no prior history of such
65 treatment. It is important to note that isolated chemotherapy-induced strictures are exceptionally rare.

66 Two extensive retrospective investigations examined the incidence of esophageal strictures in paediatric patients
67 undergoing treatment for cancer. In both studies, all patients were subject to extended follow-up, which encompassed the
68 assessment of delayed complications. [4, 5]

69 In the adult population, only two documented cases exist of patients developing esophageal stricture during systemic
70 chemotherapy treatment [2, 3]. Notably, in the majority of paediatric cases, including our own, symptoms manifested
71 within three weeks of initiating systemic chemotherapy. Our patient underwent 5-fluorouracil (5-FU)-based chemotherapy,
72 and their symptoms initially ameliorated following endoscopic dilation therapy [3]. Subsequent endoscopic examinations
73 with multiple biopsies revealed no signs of esophageal malignancy, and cross-imaging confirmed a benign process. 5-FU-
74 based chemotherapy functions by inhibiting the S phase of the cell cycle, resulting in disruption of DNA replication and
75 substantial mucosal damage across the gastrointestinal tract. Previous studies have also indicated that, in addition to 5-
76 FU, both etoposide and cisplatin can intensify the radiation effects on the esophagus. Given our patient's lack of prior
77 radiation therapy and the rarity of chemotherapy-induced strictures, it remains challenging to pinpoint the specific agent
78 responsible for the adverse outcome.

79 Mucositis and esophagitis are recognized complications associated with various chemotherapy medications. However, the
80 development of strictures is an exceedingly rare occurrence and has been scarcely documented in the existing literature.
81 In this particular case, we suspect that the patient's esophageal stricture resulted from systemic chemotherapy since the
82 patient had no history of long-term acid reflux or known exposure to established non peptic factors contributing to stricture
83 formation. This unexpected adverse event is likely the result of a complex interplay between chemotherapy-induced
84 mucosal damage and the rapid proliferation of cells lining the gastrointestinal tract. Remarkably, this condition presented
85 with considerable severity, even in the absence of concurrent radiation-induced toxicity. These findings should serve as a

86 catalyst for further investigation into the pathways of injury, potentially involved genetic mutations, and the risk factors of
87 such a rare isolated injury.
88

89 **4. CONCLUSION**

90
91 In conclusion, the case presented here underscores the rarity and complexity of chemotherapy-induced esophageal
92 strictures. While such strictures are exceptionally uncommon in the adult population, they have been more extensively
93 documented in pediatric patients undergoing cancer treatment. In our case, the patient's medical history did not reveal any
94 predisposing factors for esophageal stricture, such as prior radiation therapy or chronic acid reflux.
95

96 **COMPETING INTERESTS : NO**

97 **AUTHORS' CONTRIBUTIONS**

98 This work was carried out in collaboration among all authors. All authors read and approved the final manuscript.
99

100 **CONSENT**

101 As per international standard or university standard, parents(s) written consent has been collected and preserved by the
102 author(s).
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105 **ETHICAL APPROVAL**

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107 As per international standard or university standard written ethical approval has been collected and preserved by the
108 author(s).
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