Original Article

Management of infected post-pancreatic resection fluid collections under endoscopic ultrasound guidance using lumen apposing metal stent: A case series and review of the literature

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Backgrounds/Aims: Post-operative pancreatic fistulas (POPF) and fluid collections (POPFC) remain significant sources of morbidity and mortality after pancreatic resections. There remains a paucity of literature describing endoscopic ultrasound (EUS) guided drainage of POPFC using a Hot AXIOS™ lumen apposing metal stent (LAMS).

Methods: We conducted a retrospective study, encompassing all consecutive patients with POPFC managed using Hot AXIOS™ LAMS at our institution between January 2017 and December 2019. Primary outcome measures were technical and clinical success. Secondary outcome measures were adverse events and recurrence rates.

Results: Five patients underwent EUS guided drainage using Hot AXIOS™ LAMS during the study period. Mean age of patients was 67.8 ± 2.16 years. The majority (60.0%) of patients were males. Median duration of symptom onset after surgery was 9 days. All patients presented with abdominal pain. Median size of the collection measured on computed tomography was 91 mm. Median interval time between symptom onset and EUS drainage was 30 days. Two patients required percutaneous drainage prior to EUS guided drainage. Technical and clinical success were achieved for all patients. No adverse events were observed. Median duration of follow-up was 90 days. No recurrence of collection occurred during the follow-up period.

Conclusions: EUS guided drainage of POPFC using Hot AXIOS™ LAMS is a safe and effective treatment modality with technical and clinical success rates of 100% in our experience.

Key Words: Pancreatic fistula; Endosonography

INTRODUCTION

Infected post-operative pancreatic fistulas (POPF) and fluid collections (POPFC) are major sources of morbidity and mortality following pancreatic resections. The majority of collections that occur following distal pancreatectomies are due to a persistent leak from the pancreatic stump. The existing literature demonstrates that 30%–40% of patients may suffer from a clinically significant POPF leading to a collection following a distal pancreatectomy [1-3]. A recent systematic review and meta-analysis has shown that the incidence of POPF is higher after a central pancreatectomy than that after a distal pancreatectomy [4]. In those undergoing a distal pancreatectomy, predisposing factors for POPF include technique of pancreatic transection and stump closure, site of resection, and additional splenectomy [5-8]. POPFC has three grades based on clinical significance and the need for intervention. Grade A has been recently reassigned the term of biochemical leak.
as it does not affect clinical outcome of the patient. Grades B and C are clinically relevant that require further intervention [9]. Grade C is the most severe one. It may be fatal in 25% of patients [10]. The majority of POPFC are asymptomatic. Symptomatic collections may present with pain, fever, gastric outlet obstruction, fistulae, intra-abdominal sepsis, and multi-organ failure. Conservative management of patients with POPFC includes octreotide infusion, intravenous antibiotics, and total parenteral nutrition. Patients with Grade B and Grade C fistula require interventional therapy in the form of percutaneous or endoscopic drainage. Percutaneous drainage (PCD) is widely used because of its higher rate of technical success, widespread availability, safety, and low rate of adverse event or complications. PCD has its own limitations. It can leave an indwelling external catheter which may affect the overall quality of life. It also needs monitoring of drain output and frequent flushing. In addition, it requires trained health care professionals. Furthermore, serial upsizing of the drain may be required in addition to elective drain exchange [11-14]. Endoscopic management of pancreatic leak following pancreatic resection involves pancreatic sphincterotomy with or without trans-papillary stenting and endoscopic ultrasound (EUS) guided transmural drainage of POPFCs. Multiple case series have described the role of trans-papillary pancreatic duct stenting for POPFC as a safe and effective technique for management of POPF [15,16]. However, complications associated with endoscopic retrograde cholangiopancreatography (ERCP), especially post ERCP pancreatitis, lack of benefit in larger and infected collections, and collections not communicating with the pancreatic duct, are associated with a trans-papillary stenting.

There is significant evidence in the literature that supports the use of EUS guided transmural drainage of pancreatic fluid collections (PFCs) following pancreatitis [17]. EUS has revolutionised the management of PFCs. It has almost replaced other treatment modalities as the initial procedure of choice for drainage of PFCs following pancreatitis [18]. The role of EUS guided PFC drainage has evolved from plastic stents to fully covered self-expandable metal stents (fcSEMS) and most recently lumen apposing metal stents (LAMS), including Hot AXIOS™ stent (Boston Scientific, Marlborough, MA, USA), Nagi stent (Taewoong Medical Co., Goyang, Korea), and Niti-S SPAXUS stent (Taewoong Medical Co.) [19-24]. One advantage of the Hot AXIOS™ LAMS system is that it eliminates multiple steps during EUS guided drainage of PFC. Consequently, it is safer and more time-efficient than other LAMS and fcSEMS. Previous studies have evaluated the role of EUS guided drainage in POPFCs. However, the experience with Hot AXIOS™ for drainage of POPFCs has been limited [25,26]. In this service evaluation project, we report the safety and efficacy of LAMS Hot AXIOS™ for management of POPFCs based on our experience.

**MATERIALS AND METHODS**

This was a service evaluation project involving retrospective analysis of routinely collected clinical data. Consecutive cases of POPF at our institution presenting between January 2017 and December 2019 were retrospectively identified from the database. Patients with POPFC grades B and C who underwent EUS guided drainage using Hot AXIOS™ LAMS were included in the analysis. EUS guided drainage was performed after multidisciplinary discussion between surgeons, interventional radiologists, and endoscopists. Symptomatic patients with well-defined walled off collections having close adherence to stomach or duodenum were chosen for EUS guided drainage.

All patients underwent contrast enhanced computed tomography (CT) before drainage per our institute’s policy (Fig. 1–3). Informed consent was obtained from all patients prior to drainage. Patients less than 18 years of age, pregnant patients, patients with collections due to causes other than pancreatic resection, and patients with collections not amenable to EUS guided drainage were excluded from this study.

**Data collection**

Data were collected using institutional electronic patient record and entered into a pre-designed excel spreadsheet. Demographic details including age, sex, nature of underlying disease and surgery, time interval between surgery and first symptom, time interval between onset of symptom and EUS drainage, imaging results (single vs. multiple collections, size of the largest collection), laboratory results (white blood cell count, C reactive protein at the time of admission when POPFC was

![Fig. 1. Pancreatic fluid collection at the resection site extending up to the under surface of the greater curvature of the stomach. Contrast enhanced computed tomography of abdomen (pre-endoscopic ultrasound drainage).](https://www.ahbps.org)
first detected), and follow-up duration were recorded. Regarding EUS guided drainage, size of the collection, calibre of Hot AXIOS™ LAMS inserted, and LAMS indwelling time before removal were recorded. In addition, whether PCD was performed prior to EUS drainage and whether abdominal drain was removed before EUS drainage were also noted. Follow-up assessment was conducted either in the hospital’s outpatient clinic or by telephone.

**Procedures**

EUS guided drainage using LAMS Hot AXIOS™ was performed by experienced endosonographers and team under propofol sedation or general anaesthesia. All procedures were carried out using a Pentax linear echo-endoscope (EG-3870-UTK; PENTAX House, Slough, United Kingdom). LAMS (Hot AXIOS™ stent and electrocautery-enhanced delivery system; Boston Scientific) used in this study were through-the-scope fcSEMS of 15 or 20 mm in diameter and 10 mm in length. After direct puncture of the wall of the collection using the electrocautery tip (without using a guidewire to assist in stent insertion), the delivery catheter was advanced into the fluid collection and the distal flange was deployed under EUS guidance (Fig. 4–7). The proximal flange was released under EUS or endoscopic view. A contrast enhanced CT of the abdomen and pelvis was performed for all patients before removing the LAMS (Fig. 8, 9). The LAMS was removed within 6 weeks for all patients following repeat cross-sectional imaging at 4–5 weeks (Fig. 10). It is our routine practice to subsequently replace the LAMS with two double pigtail plastic stents using the same tract.
Outcomes

Primary outcome
Primary outcomes were technical and clinical success. Technical success was defined as successful placement of Hot AXIOS™ LAMS. Clinical success was defined as resolution of POPFC as confirmed on CT before removing Hot AXIOS™ LAMS.

Secondary outcome
Secondary outcomes were defined as adverse events related to Hot AXIOS™ LAMS such as bleeding, non-target perforation, stent migration, or any other unexpected events attributable to Hot AXIOS™ LAMS. Recurrence was defined as persistence of PFC following removal of the LAMS.

Ethical clearance
This was a service evaluation using retrospectively collected clinical data. Only the clinical team responsible for managing patients had access to the data. The clinical team conducted all analyses and drafted the manuscript.

Statistical analysis
Descriptive statistics was used to describe continuous vari-
ables. Results are presented as mean or median with standard deviations or interquartile range. Microsoft Excel ver. 2010 (Microsoft, Redmond, WA, USA) was used for data set compilation and statistical analysis.

RESULTS

Demographic details
A total of 434 patients underwent pancreatic and or splenic resections between January 2017 and December 2019. Twenty-three patients had POPFC and 18 (78.2%) patients had resolution of POPFC with conservative management or PCD. One patient required re-laparotomy and wash out. Five patients underwent EUS guided drainage using Hot AXIOS™ LAMS.

Demographic details including underlying surgery details are summarized in Table 1. In the EUS drainage group, there were two patients who underwent distal pancreatectomy and splenectomy for neuroendocrine tumour of the pancreas, two who underwent splenectomy for B cell lymphoma and traumatic splenic rupture, respectively, and one who underwent subtotal pancreatectomy and splenectomy for pancreatic ductal adenocarcinoma.

Clinical details including radiology and inflammatory markers
Clinical details are summarised in Table 2. All patients presented with abdominal pain. In addition, two (40.0%) patients had clinical features of sepsis. All patients had raised C reactive protein levels. All but one patient had raised total leukocyte count at the time of initial presentation with POPFC. The size of the smallest collection was 83 mm (patient 2) and the largest collection had a size of 135 mm (patient 5). Multiple communicating collections were observed in 3 (60.0%) patients.

| Table 1. Baseline demographic details and characteristics of five POPFC patients |
|-----------------------------|-------------------|
| Patients detail             | Value             |
| Age (yr)                    | 67.8 ± 2.16       |
| Sex (male : female)         | 3 : 2             |
| Reasons for surgery         |                   |
| Pancreatic malignancies     | 3                 |
| Splenectomy for trauma      | 1                 |
| Splenectomy for B cell lymphoma | 1          |
| Histopathology              |                   |
| Neuroendocrine tumour       | 2                 |
| Pancreatic ductal adenocarcinoma | 1          |
| Others                      | 2                 |
| Type of surgery             |                   |
| Distal pancreatectomy       | 2                 |
| Splenectomy                 | 2                 |
| Subtotal pancreatic head excision + splenectomy | 1          |
| Values are presented as mean ± standard deviation or number only. POPFC, post-operative pancreatic fluid collections. |

| Table 2. Clinico-radiological profile and outcome measures of five POPFC patients |
|-----------------------------|-------------------|
| Variable                    | Value             |
| Presenting symptoms         |                   |
| Pain abdomen                | 5                 |
| Fever                       | 2                 |
| Pain and fever both         | 2                 |
| Inflammatory markers        |                   |
| White blood cell (mm$^3$)   | 16,000 (6,200–28,200) |
| C reactive protein (mg/dL)  | 209 (7–233)       |
| Maximum size of the collection on CT (mm) | 91 (83–135)    |
| Multiple collections on CT  | 3                 |
| Interval between surgery and first symptom pertaining to POPFC (days) | 9 (17–50)       |
| Interval between symptom onset and EUS drainage (days) | 30 (21–77)   |
| PCD before EUS guided drainage | 2             |
| Size of LAMS used (mm)      |                   |
| 15 × 10                     | 2                 |
| 20 × 10                     | 3                 |
| Technical success           | 5                 |
| Clinical success            | 5                 |
| Adverse events              | 0                 |
| Median follow up after LAMS removal (days) | 90 (30–364)  |
| Values are presented as number only or median (range). POPFC, post-operative pancreatic fluid collections; CT, computed tomography; EUS, endoscopic ultrasound; PCD, percutaneous drainage; LAMS, lumen apposing metal stent. |

Drainage
The median interval between the onset of symptom due to POPFC and EUS drainage was 30 days. In one patient, it was
performed at 3 weeks after the onset of symptoms. In all patients, the abdominal drain inserted intraoperatively had been removed before the onset of symptoms. Two of five patients underwent PCD before the EUS guided drainage. Two patients had LAMS of 15 mm in diameter inserted and the remainder had 20 mm in diameter. Technical success and clinical success rates were 100% in all patients. LAMS was removed after successful resolution of the collection by 6 weeks in all patients. No major or minor adverse events were encountered. The median follow-up duration was 90 days (range, 30–364 days). No recurrence was observed in the study population in the follow-up period.

Microbiological profiles of fluids aspirated from collections during EUS guided drainage are summarised in Table 3. Fluids from patient 1 and 5 had growth of commensal flora while patients 2, 3, and 4 showed growths of pathogenic microorganisms in addition to commensal flora. Patient-wise demographic details are summarised in Table 4. Three of our patients had a large hiatus hernia. If post-pancreatectomy collection extends superiorly, it may present a technical challenge. Thus, it is important to avoid deploying the LAMS within a hiatus pouch.

**DISCUSSION**

The Hot AXIOS™ stent system purports to offer a larger diameter stent, superior anchorage, ease of use, limited procedural time, and minimal complications. Our findings are concordant with the previously published case reports and series in this particular clinical context. Li et al. [25] have described the use of Hot AXIOS™ in a 59-year-old patient with pseudocyst at 4 months following distal pancreatectomy and splenectomy. A 15 × 10 mm stent was used for drainage which was removed 60 days later. No adverse event or recurrence was observed. Li et al. [25] removed the LAMS after 60 days while in our study we removed the LAMS by 6 weeks as there was evidence that removal of LAMS after 6 weeks could increase the risk of complications, especially bleeding. To date, Mudireddy et al. [26] have published the largest case series on the use of LAMS for management of post-surgical collections. However, the population in their multi-centric cohort was quite heterogeneous. Of 47 patients, 26 (55.3%) had fluid collections after pancreatic resections. The authors used both electrocautery enhanced (hot) LAMS and non-electrocautery enhanced LAMS. The authors used LAMS of 15 mm and 10 mm in size in 70% and 30% patients, respectively, while in our study, 15 mm and 20 mm LAMS were used in 40% and 60% of patients, respectively. Fourty percent of our patients had attempted PCD prior to EUS guided drainage in the present study. However, in the study of Mudireddy et al. [26], 30.7% of 26 patients had PCD. The authors repeated the imaging after drainage at 1–8 weeks at the discretion of the endoscopist while we routinely performed a repeat imaging at 4–5 weeks after EUS drainage. The authors also described through scope dilatation of LAMS immediately after placement which was not a strategy utilised by our unit. Despite all these differences in protocols, they reported technical and clinical success rates of 100% and 96%, respectively.

Priyanka et al. [27] have described three cases of post-surgical PFCs successfully managed with EUS guided LAMS drain-

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**Table 3.** Microbiological profiles of drained collections of five POPFC patients

| Patient | Microorganism |
|---------|---------------|
| 1       | Neisseria subflava + Streptococcus anginosus + Streptococcus mitis |
| 2       | Coagulase-negative staphylococci + Enterococcus faecalis + Escherichia coli + Pseudomonas aeruginosa |
| 3       | E. fecalis + S. mitis + Yeast |
| 4       | Streptococcus oralis + P. aeruginosa + Candida albicans + Veillonella atypica |
| 5       | Nakaseomyces glabrata |

POPFC, post-operative pancreatic fluid collections.

**Table 4.** Patient-wise demographic, clinical, and drainage details of five POPFC patients

| No. | Age (yr)/Sex | Surgery | Interval between surgery and first symptom (day) | Symptom | PCD before EUS drainage | Collection size on CT (mm) | Multiple collection | Size of LAMS (mm) | Follow up after LAMS removal (day) | Adverse event |
|-----|-------------|---------|-------------------------------------------------|---------|-------------------------|---------------------------|---------------------|-----------------|-----------------------------------|--------------|
| 1   | 67/M        | Splenectomy | 40 | Pain | N | 93 | Y | 15 × 10 | 364 | None |
| 2   | 70/F        | Lap distal pancreatectomy and splenectomy | 70 | Pain + fever | Y | 83 | N | 20 × 10 | 30 | None |
| 3   | 70/M        | Lap distal pancreatectomy and splenectomy | 9 | Pain | Y | 135 | Y | 20 × 10 | 45 | None |
| 4   | 65/M        | Splenectomy | 3 | Pain + fever | N | 91 | N | 20 × 10 | 90 | None |
| 5   | 67/F        | Subtotal excision head of pancreas and splenectomy | 7 | Pain | N | 90 | Y | 15 × 10 | 150 | None |

POPFC, post-operative pancreatic fluid collections; PCD, percutaneous drainage; EUS, endoscopic ultrasound; CT, computed tomography; LAMS, lumen apposing metal stent; M, male; F, female; Y, yes; N, no.
age. The first patient underwent Whipple procedure for pancreatic cancer and developed symptomatic PFC at five months after the surgery. An EUS guided drainage of the collection was performed using 10 mm LAMS. Unfortunately, the patient succumbed to the illness. The second patient had a pancreatic leak from the tail following neuroendocrine tumour surgery and underwent ERCP guided trans-papillary stenting of the pancreatic duct. However, the PFC persisted. It was drained using 15 mm calibre LAMS. The patient eventually died following a neurological event. The third patient presented with multi-organ injury following a road traffic accident and underwent splenectomy and hepatectomy. He developed a pancreatic leak one week later. He was initially managed with trans-papillary pancreatic duct stenting. Subsequently he presented with sepsis and a PFC. Following a failed PCD, EUS guided LAMS drainage of the collection was performed using 15 mm LAMS. The patient had an uneventful recovery. The stent was removed at eight weeks after the insertion. Technical and clinical success were achieved for all three patients in the series reported by Priyanka et al. [27]. Two of three patients underwent ERCP guided pancreatic stenting before EUS guided drainage. Unfortunately, two of the three patients died from events unrelated to the drainage [27].

However, none of the aforementioned series was focussed on challenges of EUS guided Hot AXIOS™ drainage of post-surgical pancreatic collections in comparison with drainage of pancreatitis related collections. As described in our results, POPFC are associated with a technical challenge of stent placement. These collections are often lying in a superior position in relation to the fundus of the stomach compared to collections seen following pancreatitis which are more commonly lying inferiorly. This poses a risk of deployment of the proximal flange of the stent in the esophagus, especially if patients have hiatus hernia as evident in three patients in our study. Another challenge in these collections is recurrent obstruction of the stent lumen by food debris as stents are often deployed immediately inferior to the gastroesophageal junction due to anatomical limitations. In addition, there is an increased potential risk of diaphragmatic injury while puncturing the collection again due to a close anatomical relationship with these collections.

Denzer et al. [28] have conducted a retrospective study on the role of EUS guided drainage using plastic stents for PFCs following pancreatic surgery. Twenty patients underwent 24 procedures and drainage was performed within a median duration of 30 days. Mean size of the collection was 72.5 mm and PCD was done in four patients before EUS guided drainage. Technical success rate was 100% and clinical success rate was 90%. Only one patient had a recurrence. This patient was subsequently referred for surgical management. Varadarajulu et al. [29] have published a case series on the management of EUS guided drainage of POPFC using double pigtail plastic stents with or without the use of naso-cystic drains. Out of 10 patients, successful drainage was achieved in nine patients. Only one patient had to undergo surgery for persistent symptoms. One patient in this series had stent migration. No other complications were observed. Three out of the ten patients had positive bacterial or fungal culture from aspirated fluid. Overall, authors concluded that EUS guided drainage was a safe and effective modality for managing POPFC.

Prior to introduction of EUS in the management of POPFC, PCD with or without trans-papillary pancreatic duct drainage was the modality of choice for the management of POPFC associated with pancreatic fistulas following surgery. Few studies have compared EUS with PCD for the management of POPFC. Kwon et al. [30] have performed a retrospective study comparing outcomes of POPFC management using percutaneous (PD) versus endosonographic approach (EUSD). Clinical success was achieved in 79% of patients in the PD group compared to 100% in the EUSD group. Three patients in the PD group had recurrence following drain removal. The recurrence was later managed successfully with EUSD. Futagawa et al. [31] have compared the use of EUS guided trans-gastric drainage (EUS-GD) of POPFC versus PCD in 33 patients retrospectively. Twenty-one (63.6%) patients underwent PCD and 12 (36.4%) patients underwent EUS-GD. Technical success was achieved for 92% of patients in the EUS-GD group compared with 100% in the PCD group. There were no statistically significant differences in outcome measures between the two groups. A recent systematic review and meta-analyses by Mohan et al. [32] has confirmed findings of these isolated studies and inferred that in terms of clinical outcome, EUS guided drainage is a safe, effective, and preferable modality for the management of POPFC. However, regarding adverse events and technical success rate, EUS guided drainage is similar to PCD. Results of our retrospective series are consistent with published literature, further strengthening the evidence supporting minimally invasive management of POPFC using LAMS device such as Hot AXIOS™.

Our data also highlight some important points in the management of POPFC. Technical and clinical success were achieved in all patients. No major or minor adverse events were observed even after a long-term follow-up (364 days). All patients had successful resolution of intra-abdominal collections and retrieval of the stent by 6 weeks. Our experience demonstrated that EUS guided drainage using Hot AXIOS™ LAMS could be performed as early as three weeks from the detection of POPFC. This is not always possible as the development of a wall around the POPFC usually takes 4–6 weeks. The first line treatment of POPFC is often PCD under intervention radiology guidance.

Limitations of our study include its restriction to a single centre, the retrospective design, and the limited number of patients. It was impossible to have a comparative arm with this volume of patients. The current cost of Hot AXIOS™ LAMS might also be another limiting factor for consideration in the management of POPFC. However, compared to PCD, repeat
EUS guided drainage of infected post pancreatic resection fluid collections

surgery, and complications associated with trans-papillary drainage, EUS guided drainage using Hot AXIOS™ LAMS appears to be more economical. We conclude that EUS guided drainage of POPFC using Hot AXIOS™ LAMS is a safe and effective technique. Further large-scale prospective studies are needed determine the role of this technique in wider practice management of POPFC.

CONFLICT OF INTEREST

No potential conflict of interest relevant to this article was reported.

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Conceptualization: PC, BM, DT, NT. Data curation: DT, DK, AA, MW. Methodology: PC, BM, NT. Visualization: BM, DT, DK, AA, NT. Writing - original draft: PC, NT. Writing - review & editing: PC, BM, DK, AA, MW, NT.

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