Chyle Leakage and Early Enteral Feeding following Pancreatico-Duodenectomy: Management Options

H.Z. Malik    J. Crozier    L. Murray    R. Carter

West of Scotland Pancreatic Unit, Glasgow Royal Infirmary, Glasgow, UK

Introduction

Chylothorax following abdominal surgery is uncommon and its management is poorly described. Chyle is transported in lymphatics that drain the abdominal and retroperitoneal viscera and join ascending lymphatic trunks, which drain the lower extremities and the pelvis. As these trunks ascend they form a saccular dilatation called the cisterna chyli. The cisterna chyli lies in front of the first and second lumbar vertebrae and drains into the thoracic duct, which carries the lymph into the thoracic cavity. Disruption of these lymphatic pathways can result in both chylothorax as well as chylous ascites. A variety of mechanisms can result in a chyle leak ranging from congenital defects of the lymphatic system to malignancy or trauma [1–22]. However, chyle leaks are also a recognised complication of abdominal, thoracic and head and neck surgery [1–2, 4–9, 12–13, 15, 18].

In the published literature there have been numerous case reports as well as collective reviews of chylothorax [2–21]. However, there are relatively little data on chylous ascites. Leibovitch et al. [22] reviewed the literature on postoperative chylous ascites. The most common surgical cause of chylous ascites was found to be abdominal aortic surgery accounting for 81% of the cases of chylous ascites reported in the literature. Despite this, less than 1% of patients undergoing aortic surgery suffer from this complication. Other causes include retroperitoneal dissection for urological and gynaecological malignancy as
Pancreatico-Duodenectomy

Chyle Leakage following Pancreatice-Duodenectomy

A consecutive series of 105 patients undergoing resection for pancreatico-duodenal malignancy from January 1999 to January 2005 were reviewed. As part of our unit protocol all post-operative patients had their drain fluid analysed for amylase. The presence of a chyle leak was defined as drainage of \( > 600 \) ml of chylous (biochemically contained chylomicrons), amylase/bile-free fluid per day. Patients were identified and their case notes reviewed.

A standard pancreatico-duodenectomy was performed on all patients. Division of the jejunum at the level of the first jejunal arcade was performed with the vessels in the jejunal mesentery being ligated. Subsequent resection of the uncinate and head of pancreas in the perivascular plane producing \( 180^\circ \)skeletalisation of the superior mesenteric vein and artery. One patient, however, with a small primary within the pancreatic body required elective superior mesenteric vein resection with primary anastomosis to achieve surgical clearance. Reconstruction was by utilising a single retrocolic jejunal loop to create an end-to-side pancreatico jejuno-stomy, hepatico-jejunostomy and gastrojejuno-stomy. In those in whom early feeding was instituted, a naso-jejunal feeding tube was passed intra-operatively and placed beyond the most distal anastomosis, feed starting at \( 30 \) ml/h increasing to full enteral support by \( 48 \) h, before being reduced with the establishment of normal diet. The type of feed used was Peptisorb. This is a 1-kcal/ml semi-elemental oligopeptide feed. It has the lowest fat content of all available enteral feeds and the highest percentage of medium-chain triglycerides as the fat source.

We initially addressed our intra-operative technique to see if these leaks could be prevented. Intra-operative identification of mesenteric lymphatics can be difficult particularly in a fasted patient. In an effort to address this issue intra-operatively, we developed a technique of injecting 50 ml 10% Intralipid (Fresenius Kabi UK) into the jejunal lumen, just prior to its division. The lymphatic vessels become opalescent and targeted for ligation (fig. 1). Intra-operative measures used to address these lymphatics have included direct suture ligation; Ultracision (Johnson & Johnson Medical Ltd.); LigaSure (Tyco Healthcare UK Ltd.); and tissue glue – FloSeal (Baxter Healthcare UK Ltd.). However, none have proved universally successful and chyle leak occurrence appeared unrelated to intra-operative factors.

A SPSS version 7.5 statistics package was used to analyse the data with both the \( \chi^2 \) and \( t \) tests being used to analyse statistical differences between those who did and did not develop a chyle leak.

Patients and Methods

A number of techniques have been used in the management of chylous ascites. Alami et al. [24] reviewed the literature, consisting of 156 patients identified through case reports, and found that in two thirds of cases chylous ascites was managed conservatively. Conservative modalities used in the management of chylous ascites included repeated paracentesis; use of high-protein, low-fat enteral nutrition as well as total parenteral nutrition. Total parenteral nutrition has been shown to be highly effective in the management of chylous ascites with success rates from 60 to 100% having been reported [25–28]. More recently the use of somatostatin has also been advocated in the management of chylous ascites, although its mechanism of action remains unclear [29–33]. Spiro et al. [34], however, found that among patients with chylothorax the presence of a chyle leak of greater than 600 ml per day was unlikely to resolve with conservative management. In refractory cases surgery has been used to deal with the disruption in the lymphatic flow. Surgical options available include the use of peritoneovenous shunts or the direct ligation of the disrupted lymphatic channels [24].

There is little doubt that enteral nutrition carries advantages over total parenteral nutritional support in that it is easier to administer, more physiologic and less expensive. There may be preservation of gut barrier function and prevent structural alterations induced by starvation and injury. The rationale for feeding the gut early after surgery is that peristalsis of the small intestine recovers 6–8 h after surgical trauma, and that a moderate absorptive function is preserved even in absence of peristalsis. Moreover, the direct passage of food stuff in the gut lumen increases splanchic blood flow and stimulates gut immune system. There seems little effect on anastomotic healing and the use of early enteral nutrition has been shown to be well tolerated in the post-operative period in colorectal [35–36] and gynaecological surgery [37]. Following the suggestion that early enteral support could be beneficial in terms of reducing post-operative infectious complications and improving patient’s outcome, we modified our peri-operative regimen to allow the early introduction of enteral nutrition after pancreaticoduodenectomy, using a naso-jejunal feeding tube placed distal to the anastomoses intra-operatively. It was during this period of early feeding that chyle leak became problematic. The objective of this study is to review the management of patients identified as having chyle leaks, the diagnosis, intra-operative attempts at prevention and subsequent management strategy prior to resolution.
Results

A total of 7/105 (6.7%) patients were identified as having a significant chyle leak as defined above. Furthermore, the incidence of pancreatic anastomotic leak was 7% and the in-hospital mortality rate was 11% for the whole cohort of patients. Of the 105 patients, 38 underwent insertion of an enteral feeding tube at the time of surgery, with early nutritional support. Among patients who developed a chyle leak, 4 received early enteral nutrition. The clinical features of all patients are represented in table 1. There was no significant difference in the clinico-pathological features of patients that developed a chyle leak and those who did not (table 1). Of the patients who developed a chyle leak, all except 1 underwent a standard Whipple resection for pancreatic adenocarcinoma. One patient however underwent a pylorus-preserving pancreatico-duodenectomy for a duodenal carcinoma.

The recognition of a chyle leak usually occurred between the 5th and 9th (median 6 days) post-operative day. The maximum volume of chyle drained on a single day ranged from 1,400 to 3,000 ml, with the median being 1,900 ml. In all cases, the patients remained clinically well with no haemodynamic instability and there was no evidence of an inflammatory response with both the temperature and white blood cell count remaining within normal limits and only a marginal elevation in the C-reactive protein was noted in some of the patients (table 2).

Table 1. Clinico-pathological features

<table>
<thead>
<tr>
<th>Clinico-pathological features</th>
<th>Chyle leak</th>
<th>No chyle leak</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age*</td>
<td>61 (41–74)</td>
<td>62 (45–75)</td>
<td>0.512</td>
</tr>
<tr>
<td>Gender, females</td>
<td>5 (71%)</td>
<td>54 (55%)</td>
<td>0.429</td>
</tr>
<tr>
<td>T-stage</td>
<td></td>
<td></td>
<td>0.630</td>
</tr>
<tr>
<td>T1</td>
<td>2 (29%)</td>
<td>32 (33%)</td>
<td></td>
</tr>
<tr>
<td>T2</td>
<td>1 (14%)</td>
<td>34 (35%)</td>
<td></td>
</tr>
<tr>
<td>T3</td>
<td>5 (71%)</td>
<td>57 (58%)</td>
<td></td>
</tr>
<tr>
<td>N-stage N1</td>
<td>3 (43%)</td>
<td>56 (58%)</td>
<td>0.398</td>
</tr>
<tr>
<td>Early enteral nutrition</td>
<td>4 (57%)</td>
<td>34 (35%)</td>
<td>0.232</td>
</tr>
</tbody>
</table>

* Values represent median and range.

Table 2. Clinical features of patients with chyle leak

<table>
<thead>
<tr>
<th>Pattern of chyle leak</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Timing of chyle leak, days</td>
<td>6 (5–37)</td>
</tr>
<tr>
<td>Maximum volume, ml/24 h</td>
<td>1,900 (1,400–3,000)</td>
</tr>
<tr>
<td>Maximum CRP</td>
<td>28 (12–65)</td>
</tr>
<tr>
<td>Maximum WCC</td>
<td>10 (6–14)</td>
</tr>
<tr>
<td>Duration of TPN, days</td>
<td>7.5 (4–32)</td>
</tr>
</tbody>
</table>

Values represent median and range.
The patient whose chyle leak was managed surgically underwent a classical Whipple resection and was discharged on the 27th post-operative day. This delayed discharge was due to social factors and after the initial perioperative period this patient remained well in the ward until his discharge date. This patient in the early post-operative period had a persistent high volume drain loss that was low in amylase and lasted 8 days. Due to poor oral intake of fats this remained unrecognised as a chyle leak. The patient was subsequently readmitted 10 days following his discharge with ascites, which proved to be chylous on diagnostic tap. After failure of initial enteral feeding high in medium-chain fatty acids and somatostatin, total parenteral nutrition was instituted, again with a poor clinical response and this patient eventually required a peritoneo-venous shunt.

In all cases initial treatment consisted of conservative management with total parenteral nutrition, and clear oral fluids. Total parenteral nutrition was commenced between 1 and 5 days (median 2 days) after the demonstration of the chyle leak, 5 via a dedicated central venous catheter and one via a peripheral feeding catheter, consisting of a standard 2,500-ml bag containing 2,211 kcal of energy and 13.5 g of nitrogen per day. Parenteral nutrition was continued until the chyle leak had clinically resolved and the patient had tolerated a period of 24 h of enteral nutrition without recurrence of the leak. Overall, total parenteral nutrition was used for a median of 7.5 days (mean 10.7 days). Among these patients there were no complications related to the delivery of the total parenteral nutrition. Although these patients received somatostatin in the early postoperative period (up to day 5 post-operatively), none of these patients required additional therapy in the form of somatostatin in order to resolve the chyle leak.

**Discussion**

Post-operative chyle leaks are a recognised complication of abdominal surgery and have been traditionally difficult to manage with a small proportion of cases necessitating aggressive surgical intervention. Although this complication is thought to be relatively rare following abdominal surgery, we have described an overall incidence of 6.7% following surgery for pancreatico-duodenal malignancy. This incidence is difficult to explain. Over the period of this study there has been no change in the surgical technique employed, with the jejunal mesenteric vessels being suture ligated. More recently in an attempt to control the lymph leak, as identified by the use of intra-operative jejunal injection of intralipid, different techniques were employed including Ultracision and LigaSure as well as tissue glue within a limited number of cases, with little success. Furthermore, all patients underwent a standard pancreatico-duodenectomy without a radical lymphadenectomy. The mechanism of action leading to chyle leak within this group of patients may be due to the lipid content of the enteral feed keeping open the visceral lymphatic channels that have been divided as part of the standard resection thus leading to the persistent chyle leak. Chyle leak did occasionally occur despite a period of gut rest; however, it was during this period of early feeding that chyle leak became problematical, leading to the recognition that the likely source of this chyle was an earlier stimulation of the lymphatic drainage of the small intestine. This trend towards increased rate of chyle leaks amongst patients receiving enteral nutrition did not reach significance. However, as the incidence of chyle leak is relatively rare, this study was not powered to detect such a difference. We have since reverted to a more conservative policy, avoiding enteral intake in the immediate post-operative period, as the use of early nutrition was part of a recent change in our established protocol.

As a result of the demonstrated benefits in colorectal and gynaecological surgery, early restoration of enteral nutrition is being utilised following a wide range of surgical procedures, and overall there seems to be little detriment to introducing nutritional support by this route. A pancreatico-duodenectomy results in interruption of the mesenteric lymphatic plexuses, the hepatico-duodenal ligament and to a lesser degree the retro-peritoneum. The high rate of significant chyle leakage associated with pancreatico-duodenal resection must raise specific questions regarding the use of early re-feeding following pancreatic resection, which appear to be less relevant to other procedures.

When a chyle leak does occur, it appears to be associated with little morbidity. All the patients who developed a chyle leak remained clinically well and unlike chylolitho-rax did not develop an inflammatory response to the chyle leak; however, their discharge from hospital was delayed. In all those treated conservatively, the chyle leak abated following the introduction of total parenteral nutrition. Furthermore, the use of total parenteral nutrition was not associated with significant morbidity. One patient did however require surgical intervention for a chyle leak. This patient presented late with chylous ascites that failed to respond to conservative management.
In summary, compared to the existing literature we have reported a high incidence of chyle leak following surgery for pancreatico-duodenal malignancy. This incidence is partly explained by a policy of early enteral nutrition among these patients. Specific attempts at intra-operative ligation or peri-operative somatostatin failed to prevent chylous leakage. All those patients who developed a chyle leak remained clinically well and the majority of cases responded quickly to the cessation of oral intake and the introduction of total parenteral nutrition.

References

8. Perez J, Casal J, Rodriguez W: Always re-
14. Bhat A, Lowery GL: Chylous injury follow-