A Comparative in vitro Assessment of Suture Strength after Exposure to Pancreatic Enzyme Solution

Merei Ibrahim1, Nahm Christopher B1,2,3, Ofri Adam1, Samra Jaswinder1,2,4, Clarke Elizabeth2,5, Anubhav Mittal1,2

1 Australian Pancreatic Centre, Royal North Shore Hospital, St. Leonards, Australia
2 Sydney Medical School Northern, Faculty of Medicine and Health, University of Sydney, Australia
3 Bill Walsh Translational Cancer Research Laboratory, Kolling Institute, University of Sydney, Australia
4 Faculty of Medicine and Health Sciences, Macquarie University, Sydney, Australia
5 C Murray Maxwell Biomechanics Laboratory, Kolling Institute, University of Sydney, Australia

ABSTRACT

Background Post-operative pancreatic fistula (POPF) is a major cause of short-term morbidity following pancreatic resection. There is wide variation in the selection of suture type for pancreaticoenteric anastomoses resulting from a paucity of data pertaining to changes in suture strength after exposure to pancreatic enzymes. This study aimed to investigate the effect in vitro of a simulated pancreatic enzyme-rich environment on the tensile strengths of various suture materials.

Methods Ten replicates of five different suture materials (polydioxanone; polyester; polyglyconate; polypropylene and barbed polybutester) were tested for tensile strength after incubation for ten days in either a pancreatic enzyme solution (PES) or in a control normal saline solution.

Results Four of five suture materials (polydioxanone; polyester; polyglyconate; polypropylene and barbed polybutester) did not demonstrate any significant decrease in tensile strength after exposure to PES compared with normal saline. Polypropylene demonstrated a significant decrease in tensile strength after incubation in PES compared with normal saline (mean (SD), 33.4 (2.59) vs 36.51 (2.78) N; p=0.019).

Conclusion The tensile strengths of polydioxanone, polyester, polyglyconate and barbed polybutester are not significantly affected by exposure to pancreatic enzymes. The tensile strength of polypropylene suture is significantly decreased by exposure to pancreatic enzymes.

INTRODUCTION

Pancreatoduodenectomy (PD) remains the only potentially curative treatment for pancreatic ducal adenocarcinoma and other pancreatic tumours including neuroendocrine neoplasia and cystic neoplasms in the head or uncinate process of the pancreas. Mortality following PD has decreased to less than three percent over the last three decades but the rate of morbidity still remains high, ranging from 18% to 52% [1, 2, 3]. Postoperative pancreatic fistula (POPF) remains the Achilles’ heel of pancreatic resection, affecting between 13 to 41% of patients, and also leads to other frequent complications related to pancreatic resection including delayed gastric emptying (19-23%), intra-abdominal abscess formation (9-10%) and post-pancreatectomy haemorrhage (1-8%) [1, 2, 3].

In an effort to reduce the rate of POPF after PD, numerous anastomotic techniques have been described with variable success [4, 5, 6, 7, 8, 9]. The majority of these techniques involve a hand sewn, sutured technique. However, eight decades on from a case report of a fatal anastomotic leak due to disruption of the catgut suture used to construct the pancreaticoenteric anastomosis, the search for the ideal suture continues [10]. Currently, there is a paucity of data comparing the appropriateness of various types of suture material for the pancreaticoenteric anastomosis. In particular, little is known about the effect of enzyme-rich pancreatic fluid on the mechanical integrity of various suture materials utilised in pancreaticoenteric anastomosis. We hypothesised that different suture materials would have differing susceptibility to reduction in tensile strength after incubation in a solution of pancreatic enzymes. This study aimed to investigate the comparative loss of tensile strength of various suture materials following incubation in a solution of pancreatic enzymes.

METHODS

Preparation of suture materials

Five different suture materials were examined in this study, including: polyglyconate (MaxonTM, Medtronic,
conduct the analysis, the suture loop was mounted on the Instron 8874 as shown in Figure 2 before a tensile load was applied to failure at a constant displacement rate of 2 mm/s with a sampling rate of 50 Hz. The tensile strength was determined by measuring the maximum tensile force, expressed in Newtons, which the suture loop could withstand before breaking.

**Statistical analysis**

Differences in continuous variables between the PES and control groups were evaluated for each suture type using Student t-tests after checking for normal distribution. One-way ANOVA was also performed to evaluate differences in mean tensile strength between suture types for the two incubation groups. P-values<0.05 were considered statistically significant. Statistical analyses were performed with GraphPad Prism 7.00 (GraphPad Software Inc., La Jolla, CA, USA).

![Figure 1](image1.png)

**Figure 1.** Suture loops created using a cylindrical template and secured with 8 reef knots.

![Figure 2](image2.png)

**Figure 2.** Tensile load applied to suture loops attached to Instron 8874.
RESULTS

Enzymatic activity of freeze-thawed PES

The amylase and lipase activities of the PES solution are detailed in Table 1. Based on these results, it was decided that replenishment of PES every 48 hours would be sufficient to maintain a level of enzymatic activity consistent with that encountered in POPF.

The effect of PES on suture tensile strength

The mean tensile strength of each suture material is shown in Table 2 and Figure 3. For four of the five suture types (polyglyconate, polydioxanone, polyester and barbed polybutester), there was no significant difference in tensile strength after exposure to PES. Polypropylene was the only suture type found to be statistically significantly weaker in PES compared to normal saline (33.4 ± 2.59 vs 36.51 ± 2.78 N; p=0.019).

In both test and control environments, there was a significant difference in the mean tensile strength between suture types (p<0.001). The mean tensile strength of polyglyconate sutures was highest (control 63.71 N, test 55.27 N) followed by polydioxanone sutures (control 45.88 N, test 44.37 N). Barbed polybutester sutures, recorded the lowest mean tensile strength (control 34.15 N, test 31.24 N).

Table 1. Enzymatic activity of freeze-thawed PES measured at various intervals.

<table>
<thead>
<tr>
<th>Time (hours)</th>
<th>Amylase activity (U/L)</th>
<th>Lipase activity (U/L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>0</td>
<td>&gt;655 400</td>
<td>&gt;120 000</td>
</tr>
<tr>
<td>2</td>
<td>&gt;655 400</td>
<td>&gt;120 000</td>
</tr>
<tr>
<td>4</td>
<td>&gt;655 400</td>
<td>&gt;120 000</td>
</tr>
<tr>
<td>6</td>
<td>588 162</td>
<td>&gt;120 000</td>
</tr>
<tr>
<td>12</td>
<td>353 730</td>
<td>&gt;120 000</td>
</tr>
<tr>
<td>24</td>
<td>147 891</td>
<td>&gt;120 000</td>
</tr>
<tr>
<td>48</td>
<td>29 818</td>
<td>&gt;120 000</td>
</tr>
<tr>
<td>72</td>
<td>14 960</td>
<td>114 610</td>
</tr>
<tr>
<td>5 days</td>
<td>5 000</td>
<td>57 590</td>
</tr>
<tr>
<td>10 days</td>
<td>1 392</td>
<td>26 842</td>
</tr>
</tbody>
</table>

Table 2. Comparison of tensile strength (mean ± SD) of suture loops after 10 days of incubation in normal saline versus pancreatic enzyme solution.

<table>
<thead>
<tr>
<th>Suture Type</th>
<th>Saline (Newtons)</th>
<th>PES (Newtons)</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Polyglyconate</td>
<td>63.71 ± 8.89</td>
<td>55.27 ± 9.91</td>
<td>0.06</td>
</tr>
<tr>
<td>Polydioxanone</td>
<td>45.88 ± 4.44</td>
<td>44.37 ± 3.35</td>
<td>0.402</td>
</tr>
<tr>
<td>Polypropylene</td>
<td>36.51 ± 2.78</td>
<td>33.4 ± 2.59</td>
<td>0.019</td>
</tr>
<tr>
<td>Polyester</td>
<td>37.99 ± 2.54</td>
<td>37.78 ± 2.13</td>
<td>0.843</td>
</tr>
<tr>
<td>Barbed polybutester</td>
<td>34.15 ± 4.21</td>
<td>31.24 ± 3.43</td>
<td>0.107</td>
</tr>
</tbody>
</table>

PES, pancreatic enzyme solution. Bold figures denote p-values<0.05.

DISCUSSION

In this study, we have demonstrated a significant reduction in the tensile strength of polypropylene suture material after sustained exposure to pancreatic enzyme-rich solution, but no significant reduction in the tensile strengths of polyglyconate, polydioxanone, polyester and barbed polybutester suture materials.

Although the type of suturing technique used in pancreaticoenteric anastomosis has been studied extensively to date, little is known about the suitability of different suture material types for this use. According to an international survey, a wide variation currently exists in suture selection amongst pancreatic surgeons [11]. It was reported that absorbable monofilament sutures are the most popular choice for inner layer suturing. For outer layer suturing, however, both non-absorbable braided and absorbable monofilament sutures were popular choices. Due to the variability in suture selection, there has been a recent interest in comparing suture types in an effort to optimise suture choice. Whilst no randomised clinical trials comparing sutures have been conducted yet, one case-control study has reported an association between polyester sutures and a decreased risk of POPF development [12].

The focus of existing literature has been on evaluating the tensile strength of suture material after exposure to either animal or human pancreatic and bile fluid. Benchtop analyses of suture strength after exposure to pancreatic fluid have previously been performed, however little to no data is given regarding the actual enzymatic activity of this fluid. Moreover, the fluid is taken from patient drain samples, which is inherently non-standardised throughout the course of the study period [13, 14, 15]. The present study has improved on this by introducing a novel method by which a simulated pancreatic enzyme solution can be created with a standardised and measurable level of enzymatic activity.

In keeping with the present study, Muftuoglu et al. reported that the mean baseline tensile strength of polypropylene decreased by 23% following a 7-day incubation period in pancreatic fluid [13]. In contrast,
both Andrianello et al. and Karaman et al. reported no evidence of polydioxanone sutures weakening in comparison to baseline values following 20-day and 10-day incubation periods respectively [14, 15]. Similarly, electron microscopic analysis of polypropylene sutures did not reveal any signs of disintegration in the suture's architecture [15].

The development of clinically relevant POPF is multifactorial, and while reduction in the tensile strength of polypropylene was noted in our study, this may only be one of the contributing factors. Nonetheless, our findings suggest that further investigation is warranted comparing polypropylene sutures with other suture materials in a large clinical study to clarify this.

Studies have previously reported that polydioxanone sutures maintained the highest percentage of baseline strength [13, 14]. Polyester sutures were also demonstrated to retain almost all of their baseline strength following exposure to pancreatic fluid [14]. This is supported by our study which found no evidence of either polydioxanone or polyester sutures experiencing degradation by pancreatic enzymes. Further to this, there is also clinical evidence of a reduction in POPF with the use of polyester sutures for pancreaticoenteric anastomoses [12].

The present study also included barbed polybutester due to increasing use of this suture for pancreatic anastomoses in both open and robotic pancreaticoduodenectomies. Whilst it was associated with the lowest tensile strength overall, there was no significant decrease in tensile strength with exposure to pancreatic enzymes, and the breaking point was at forces much higher than that encountered in pancreaticojejunal anastomosis.

Our study has several limitations. The current analysis did not include other suture parameters that may be relevant to POPF development such as suture elongation and elasticity. This study only focused on the effect of pancreatic enzymes on sutures. Other post-operative factors such as pancreatitis or microbes, may also affect sutures in a way that was not investigated in the present study.

CONCLUSION

The tensile strengths of polyglyconate, polydioxanone, polyester and barbed polybutester sutures are not significantly affected by pancreatic enzymes. The tensile strength of polypropylene sutures is significantly decreased by exposure to pancreatic enzymes.

Conflicts of Interest

Some suture materials were provided in-kind by Medtronic (Minneapolis, MN, USA) for the conduct of this study.

This manuscript has not been published and is not under consideration elsewhere.

References