Prevalence of Histologically Proven Acute Appendicitis and Incidental Carcinoid Tumour in the Practice of Surgical Pathology at BPKIHS

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Abstract
Background: Appendicitis is the most common cause of acute abdominal pain requiring surgical intervention.

Objectives: To determine the relative prevalence of histologically proven acute appendicitis in surgically resected specimens with clinical diagnosis of acute appendicitis and to find out the rate of occurrence of carcinoid tumour as an incidental histologic finding.

Methods:
Type of study- descriptive study
Study unit- all gross specimens received in the department of pathology over a period of twenty months (1.1.2006 to 31.8.2007)
Study sample- Histologic data on 515 appendicectomy samples (clinically diagnosed as appendicitis) of the total 7295 specimens received over a period of twenty months were retrieved from the archives of department of pathology, B. P. Koirala Institute of Health Sciences (BPKIHS).

Exclusion criteria- appendectomy incidental to another surgical procedure.

Results: Appendectomy specimens constituted 7.0% (n= 515; M:F 1.1:1) of all surgical pathologic specimens (n= 7295) at BPKIHS. Following is the breakup of histologic diagnosis: acute appendicitis with or without periappendicitis and gangrenous change (93.6%, n= 482), receding appendicitis (5.4%, n= 28), normal histology (1.0%, n= 5). Carcinoid tumours were detected incidentally in three cases (0.58%) out of the total number of 515 appendectomy specimens.

Conclusion: Analysis of data revealed
1. A relatively higher prevalence (6.99%) of histologically proven acute appendicitis in this Tertiary health care set up compared to similar data reported in the literature.
2. Rate of occurrence (0.58%) of carcinoid tumour as incidental finding is similar to that reported in the literature.

Keywords: Appendicectomy, Appendicitis, Appendix, Carcinoid Tumor.

Introduction
Appendicitis is the most common cause of acute abdomen and can affect all age groups. It is so common that a sharp onset of pain in the right lower abdomen is thought of as appendicitis even by a lay person. A clinical diagnosis is usually made by the surgeon. However, the pathologist plays an important role in confirming the diagnosis. Appendectomy is reported as one of the most frequently performed surgical procedures in the world1,2 and it's prevalence varies from 4.9% to 8.6%.2 The pathologic spectrum of the acutely inflamed appendix encompasses a wide range of infectious and non-infectious entities. Further, the incidental detection of a carcinoid tumour in appendicectomy specimens is emphasized.

Although the peak incidence of acute appendicitis occurs in the 15-24 year age group, 5-10% of all appendicitis occurs in the elderly (i.e. those over 60 years of age) and this accounts for 5% of all acute abdominal
conditions in the aged. The present study reviews the epidemiology, pathophysiology, the histology of cases presenting with acute appendicitis, with special reference to carcinoid tumor and the rate of negative appendicectomies.

**Aims:**

1. To determine the relative prevalence of histologically proven acute appendicitis in surgically resected specimens with clinical diagnosis of acute appendicitis
2. To find out the rate of occurrence of carcinoid tumor as an incidental histologic finding.

**Materials and Methods**

**Type of study:** Descriptive study

**Inclusion criteria:** Histologic data on all appendectomy specimens with a clinical diagnosis of acute appendicitis were retrieved from the archives of department of pathology, BPKIHS, during a period of twenty months (1.1.2006 – 31.8.2007) and analyzed retrospectively. Exclusion criteria included appendectomy incidental to another surgical procedure.

All the appendicectomy specimens were fixed in 10% formalin. Along with the histologic data, clinical details and findings on gross examination were also noted.

Three sections were taken from each of the appendicectomy specimen - comprising of one section from the tip, one each from proximal and mid one third. All sections were stained with hematoxylin and eosin and examined under the light microscope. Immunohistochemistry and special stains were used when required.

Data entry and analysis was done using MS Excel entry and EPI Info analysis. Chi² test was applied to test significant difference between the variables. P value less than 5% was considered as significant. Descriptive statistics like percentage, proportion were calculated and represented diagrammatically.

**Results**

The total number of specimens received for histopathologic evaluation in the department of pathology over a period of twenty months was 7295, out of which 515 were appendicectomy specimens which constituted 7.0% of all surgical pathologic specimens.

Out of the 515 appendicectomy specimens 294 were males and 221 females, with 1.1:1 ratio. (Fig. 1)

![Sex Distribution](image)

**Fig. 1: Sex Distribution**

The age ranged from 15 months to 81 years. The median age was 37 years. Mean age of male and female patients were 30.36 and 26.98 respectively. Ten percent (n = 53) of the total cases were in the above fifty age group, out of which 3.30% (n = 17) patients were of elderly (60 years and above) age group. The pediatric age group (upto 15 years) comprised 22.1% (n = 114) patients.

**Table 1: Age distribution**

<table>
<thead>
<tr>
<th>Age (years)</th>
<th>Number of Patients</th>
</tr>
</thead>
<tbody>
<tr>
<td>0-10</td>
<td>54</td>
</tr>
<tr>
<td>11-20</td>
<td>147</td>
</tr>
<tr>
<td>21-30</td>
<td>147</td>
</tr>
<tr>
<td>31-40</td>
<td>73</td>
</tr>
<tr>
<td>41-50</td>
<td>41</td>
</tr>
<tr>
<td>&gt;50</td>
<td>53</td>
</tr>
<tr>
<td><strong>Total</strong></td>
<td><strong>515</strong></td>
</tr>
</tbody>
</table>
Only five out of the 515 appendicectomy cases were cases of vermiform appendix (0.97%), the remainder 510 being appendicitis of varying severity. Therefore, the negative appendicectomy rate in our hospital was around 1% (0.97%) and the average annual rate of histologically proven appendicitis was calculated to be around 6.99%.

Carcinoid tumor was detected incidentally in three cases (0.58%).

Three cases of carcinoid tumour involved one female and two males, age ranging from 6 years to 38 years. In two of these specimens, tumour was located in the tip while the remaining one was located in the base. The tumor was less than one cm. in all specimens. Histology showed acute appendicitis with periappendicitis in two cases and gangrenous changes in one of them. The tumour was confined to the mucosa and submucosa and the resected base was free of tumour on microscopic examination.

Out of the five cases of vermiform appendix, one of a 12 year boy showed histologic feature of oxyuriasis appendix.

**Etiology**

Most of the acute appendicitis were due to noninfectious obstructive cause, contributed in maximum cases by the presence of fecolith (208/515), i.e., in 40% of cases. There was a single case of infectious obstructive aetiology contributed by presence of enterobius vermicularis. Tumours like carcinoid also contributed to the development of appendicitis in less than one percent (< 1%) of cases.

Thirty three cases showed perforation which were according to age. It showed relatively high rate of perforation in children less than ten years of age (11.1%) and in patients over 50 years (7.6%). Though a similar figure was also observed in second decade, this increase in rate of perforation could be attributed to the highest number of patients in this group.

**Table 2: Frequency of perforation in different age groups**

<table>
<thead>
<tr>
<th>Age (Years)</th>
<th>Total No. of Patients</th>
<th>No. of Perforation (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>0-10</td>
<td>54</td>
<td>6 (11.1)</td>
</tr>
<tr>
<td>11-20</td>
<td>147</td>
<td>11 (7.5)</td>
</tr>
<tr>
<td>21-30</td>
<td>147</td>
<td>9 (6.1)</td>
</tr>
<tr>
<td>31-40</td>
<td>73</td>
<td>3 (4.1)</td>
</tr>
<tr>
<td>41-50</td>
<td>41</td>
<td>0</td>
</tr>
<tr>
<td>&gt;50</td>
<td>53</td>
<td>4 (7.6)</td>
</tr>
</tbody>
</table>

When the cases were further divided into paediatric, elderly and as rest of the population, perforation peaks was identified in the paediatric and elderly age group.
When these values were compared statistically, it was observed that the perforations were significantly higher in both the paediatric (p=0.005) and elderly groups (p=0.007), as compared with rest of the population (16-60 years). However, the difference in the rate of perforation between the two groups was not significant.

The demographic data and religion of 90 patients were available. The highest number of patients had come from Sunsari (43), followed by Jhapa (18), Morang (11) Ilam (7) and Dhankutta (6). There were one patient each from Bhojpur, Saptari and Udaypur.

Most of the patients (75%) were Hindus, followed by Kiratis and Buddhists. Muslim community comprised of only 1% of cases.

Most of the patients presented with pain (46.6%), followed by vomiting (19.2%) and fever (9.7%).

**Discussion**

The first anatomic drawings of the appendix was sketched in 1492 when Leonardo da Vinci described an earlike structure, he termed the *orecchio* arising from the cecum. In 1543, Andreas Vesalius published the first illustration of an appendix. Although the anatomy of the appendix was clearly defined, its pathology and treatment remained controversial for the next 2-3 centuries until in 1886; Reginald Fitz, a professor of pathology described the appendix as the source of inflammation in acute typhlitis. It is Fitz who coined the term appendicitis and also described the signs and symptoms of acute and perforated appendicitis and was among the first to recommend early diagnosis and operative intervention.

Acute appendicitis affects all age groups. It is ranked either as the first or the second common cause of acute abdomen in late adulthood.

In the present study, maximum numbers of patients were in the second to third decade and there were around 10.29% of patients in the above fifty age group. Elderly patients who have appendicitis have a greater morbidity and mortality rate when compared with younger patients. It was estimated to be around 70%, as compared to 1% in the general population. As compared to younger age group, elderly patients have more underlying diseases and sluggish bodily physiological reactions resulting in a higher rate of morbidity and mortality. Therefore, appendicitis continues to be a challenging surgical problem in elderly patients. In our study also, the rate of perforation in appendix was 7.6% in patients over 50 years of age. This rate was second to the highest rate of 11.1% seen in the below 10 year age group, even though the number of patients in below 10 and above 50 was almost the same.

Therefore, the rate of perforation was highest in children below 10, followed by patients in the
above 50 age group. Redmond et al. have shown that the perforation rate is high in both children and elderly as in this study. However, the prognosis is graver in the elderly.\(^9\) The perforation rate in acute appendicitis in various studies, however, ranged from 21\% to 72\%.\(^3,10-13\)

Delay in presentation appears to be associated with risk of perforation. Some authors believe that the delay in presentation is multifactorial. Some of these elderly patients live alone and have difficulty in accessing medical care early, while others, with a higher pain threshold, would attribute the symptoms to indigestion or constipation, thus ignoring the initial symptoms until they worsen. Another major factor proposed in literature is probably the fear and anxiety associated with hospital admission in this age group.\(^3\)

Considering appendicitis as one of the early diagnostic possibilities in elderly patients with abdominal pain, followed by prompt management accordingly may improve the overall result.\(^6\)

Contrary to the above discussion few other studies, including ours, have stated that the perforation ratio is highest in younger children and then only in the elderly. Some of these studies have also shown a peak in the preschool children.\(^14,15,16,17\)

In our case, however, no such peak was observed in the preschool group and the cases were evenly distributed among all groups in children. When taken into account all patients up to 15 years old (114 appendicectomies with 13 perforations) the perforation rate was around 11.4\%, so this was also not different from patients under 10 years of age, therefore still maintaining a high perforation rate in children.

Though acute appendicitis can occur from the time of infancy to old age, the peak age of incidence is in the second and third decades of life,\(^14,15,18,19,20\) as also seen in our study where 294 patients were in the 2\(^{nd}\) to 3\(^{rd}\) decade.

History and clinical examination remains the mainstay of diagnosis in acute appendicitis. However, many surgical procedures are performed in which the appendix is subsequently found to be normal.

The negative appendicectomy rate reported in the literature varies; typical figures are between 7\% and 20\% in men and 20\% and 45\% in women.\(^14\) These figures are considered high taking into account the commonality of the disease.

Studies carried out around a decade back have quoted that the negative appendicectomy rate has remained largely unchanged over the last 70 years.\(^16\)

In our study, however, the negative appendicectomy rate was only 1\% with three males and two females. Studies done in the past show that the numbers of negative appendicectomies were larger in females than in males.\(^17,18,21\) The patients with negative appendicectomies, in our study, did not belong to a particular age group and were distributed widely between 2\(^{nd}\) to 5\(^{th}\) decades.

The inflammation in appendicitis can vary from case to case and histology may reveal an acute appendicitis with severe periappendicitis to gangrenous changes and perforation or simply a case of receeding appendicitis. Acute appendicitis with periappendicitis was the commonest histological picture in this study.

These various degrees of inflammation are said to be only due to a difference in the time of presentation i.e. patient's delay in seeking medical advice or post admission delay, and not as a result of any diagnostic dilemma.\(^5,22\) This stands true for our study as well as far as diagnostic dilemma is concerned, as the negative appendicectomy rate is very low in our case, compared to other studies. The time of presentation and delay in diagnosis, however cannot be addressed as details pertaining to this aspect are not included in this study.

There is marked variation in the etiopathogenesis of appendicitis. Although it
appears to be a result of mucosal injury, all cases of appendicitis do not show evidence of lumen obstruction by a fecolith.\textsuperscript{21}

There can, therefore, be a number of causes for luminal obstruction ranging from lymphoid hyperplasia due to a variety of causes, fecolith formation, foreign bodies and tumours including carcinoma, adenocarcinoma and Burkitt’s lymphoma. Luminal obstruction has been emphasized as an initiating event in the causation of acute appendicitis, irrespective of the various causes. Once there is obstruction, it leads to accumulation of mucosal secretion, inflammatory exudation which increases intraluminal pressure, which obstructs lymphatic drainage and thereby, develops edema and mucosal ulceration, distension of appendix increases and results in venous obstruction. At the end of this process, ischemic necrosis occurs at the wall of appendix. Once there is obstruction, it also causes stimulation of the visceral afferent nerve fibres and therefore, causing referred epigastric and periumbilical pain.\textsuperscript{4,24} Accepting these common causes and presentation, the present study also showed presence of fecolith in 40\% of cases and 46\% of the patients presented with pain. We would like to highlight the incidental detection of carcinoid tumour in three cases (0.58\%) out of the total number of 515 appendectomy specimens.

Appendiceal tumours are uncommon and most often present as acute appendicitis.\textsuperscript{25,26,27} Carcinoid is reported as the most frequent tumour of appendix;\textsuperscript{25,27} the prevalence rate varying from 0.3 to 0.9\%, 0.5 and 0.7\% in appendicectomy specimens\textsuperscript{25,27,28}

They are found in both sexes and at any age, the peak age being 30-50 years.\textsuperscript{27,28} Though carcinoid tumour is uncommon, it can be encountered several times during the career of a surgeon (1/200-300 appendicectomy).\textsuperscript{29}

The most common site of occurrence in the gastrointestinal tract is in the appendix where they are single and mainly at the tip, followed by the ileum where they are frequently multiple and then the rectum.\textsuperscript{27,28}

Carcinoid of the appendix occurs mainly in the tip (70\%), next 20\% being in the mid one third and only 10\% in the base.\textsuperscript{27,29}

The term carcinoid tumor was first described by Oberndorfer in 1907 in the mistaken belief that it was only locally invasive and did not metastasize.\textsuperscript{28}

Carcinoid tumors of the appendix are usually benign. Simple appendectomy is adequate treatment for appendiceal carcinoids, less than 1 cm in diameter. Tumours over 2 cm in diameter may exhibit malignant behaviour and right hemicolectomy is the choice of treatment in such cases. However, controversy exists concerning the clinical value of microscopic invasion of serosal surface, subserosal lymphatics, and/or mesoappendix for tumors smaller than 2 cm. Several authors suggested that these latter features should be considered reliable parameters of aggressive clinical behavior and, thus, recommended radical surgery (hemicolec tomy) in patients having at least one of these findings. These tumours are frequently discovered incidentally during an appendectomy for acute appendicitis.\textsuperscript{29,30}

In our study, one case of oxyuriasis appendix was seen in a vermiform appendix specimen. The emphasis lies that such patients should be treated with antihelminthic therapy first and an immediate surgical intervention is unwarranted.

**Conclusion:**
1. Public education regarding pain abdomen and acute appendicitis, specifically targeting those groups at risk, may provide a substantial and significant solution to the complicated appendix.
2. Carcinoid tumours of the appendix most often present as acute appendicitis. The value of histopathological analysis of every removed appendix is emphasized.

**References**


