Between February 26, 1981, and July 30, 1987, 36 patients underwent orthotopic liver transplantation for primary sclerosing cholangitis associated with ulcerative colitis. Three of the 36 recipients died within 3 mo because of graft nonfunction or surgical complications. The other 33 (92%) lived for at least 1 yr. Two of the 33 died after 12 and 14 mo, respectively, of recurrent cholangiocarcinoma that was not diagnosed before transplantation. Four other patients died of recurrent liver failure (three cases) or immunoblastic sarcoma (one case) after 14, 21, 36 and 44 mo. Twenty-seven (75%) of the 36 are still alive after 23 to 81 mo. The mean follow-up in the 31 patients admitted afterward was 26 (72.2%) were men. Of the 45 patients who had PSC associated with UC, 7 were excluded from the study because of previous right colectomy for a Duke's class B carcinoma (one patient), total colectomy for UC (three patients) and proctocolectomy for UC (three patients). The ages of the 36 recipients without previous colorectal surgery ranged from 22 to 63 yr (mean = 41 yr); 26 (72.2%) were men. UC duration before OLT was 3 to 30 yr (mean = 14 yr). PSC had been diagnosed 2 to 14 yr (mean = 6 yr) before transplantation. UC preceded PSC in all but two patients.

Three patients who died within 3 mo of transplantation from graft nonfunction or technical complications were excluded because meaningful colon studies were precluded. Two patients with previously undiagnosed cholangiocarcinomas in their native livers were also excluded. Transplantation was done successfully in these patients, but they died of tumor recurrence 12 and 14 mo later. Excluding these five cases, 31 (86%) of the 36 recipients were eligible for long-term study. Four of the 36 died of recurrent liver failure (three patients) or immunoblastic sarcoma (one patient) after 14, 21, 36 and 44 mo. Twenty-seven (75%) of the 36 are still alive after 23 to 81 mo. The mean follow-up in the 31 patients admitted...
diagnosed as having colorectal cancer 11 yr and 21 yr (Table 1). The corrected incidence of colorectal cancer was 2 (6.5%) of 31. The two patients with colorectal cancer were excluded because of early death or recurrent disease. There were no colorectal cancers in patients whose UC had been present for less than 10 yr (Table 1).

**RESULTS**

Two (5.6%) of the 36 patients with PSC and UC were diagnosed as having colorectal cancer 11 and 21 mo after OLT, respectively. Without the five patients who were excluded because of early death or recurrent cholangiocarcinoma, the corrected incidence of colorectal cancer was 2 (6.5%) of 31. The two patients with colorectal cancer both had histories of UC for 17 and 21 yr, respectively. There were no colorectal cancers in patients whose UC had been present for less than 10 yr (Table 1).

**Case Reports**

**Case 1:** A 48-yr-old white man with a 21-yr history of UC and PSC of 6 yr underwent uneventful OLT on July 28, 1986. Pretransplant colonoscopy on April 24, 1986, was normal except for colonic mucosal edema. Multiple biopsy samples did not show malignancy. The patient's course after transplantation was unremarkable except for an episode of easily reversible acute cellular rejection.

On July 1, 1987, she underwent colonoscopy for bloody diarrhea of 2 mo duration. Multiple colonic biopsy samples obtained at that time revealed severe dysplasia of the entire colon, for which total proctocolectomy with ileostomy was performed 11 mo after transplant. The surgical specimen contained an indurated ulcer measuring 3.0 × 2.5 cm in the midst of the transverse colon. The lesion was a well-differentiated adenocarcinoma with a prominent inflammatory cell infiltrate that extended through the colonic wall (Duke's class B) (Fig. 1). The patient remains well without evidence of recurrence 24 mo after proctocolectomy.

**DISCUSSION**

Risk factors for the development of colorectal carcinoma in patients with UC include prolonged duration or early onset of UC and extensive involvement along with severe dysplasia of the colonic mucosa (5-9). The incidence of colorectal cancer after 18 to 25 yr of UC has been reported to be from 1.4% to 34% (7-9). Our two patients are the first examples, to our knowledge, of this complication after OLT. Whether transplantation contributed to or accelerated the malignant process is open to speculation.

It has been well documented that immunosuppression after transplantation leads to the development of certain forms of cancer (12, 13). Kidney transplant recipients treated with azathioprine and prednisone have a higher than normal incidence of lymphoma, Kaposi's sarcoma and a variety of epithelial tumors, probably including colorectal carcinoma (12, 13). However, under immunosuppression with cyclosporine, incidence of carcinomas of the colon, rectum, lung, prostate and female breast has not increased much—if at all (14)—although the risk of lymphoma and sarcoma remains high.

In our two patients, pretransplant colorectal examinations, including colonoscopy and biopsies, revealed no evidence of neoplasm or dysplasia. These studies were done 9 and 3 mo before transplantation. Unfortunately, they were not repeated. Because the intervals between OLT and the development of colorectal cancer were only 11 and 21 mo in the two patients, respectively, it is tempting to speculate that malignant trans-
formation was stimulated by immunosuppression or other conditions of transplantation. However, there is no proof of this, and it is possible that the lesions were missed in the preoperative evaluation or developed between the examinations and the transplantation. Cancers arising in the colon or rectum involved with UC tend to be flat and to exhibit infiltrating growth. These qualities make it difficult, as in case 2, to establish the early diagnosis of colorectal cancer with the use of colonoscopy alone (6). Our patients had UC for 17 and 21 yr, respectively, increasing the risk of malignant degeneration.

It is clear that the changes caused by UC should be regarded as precancerous. Early detection of colorectal cancer in patients who have undergone OLT for PSC associated with UC should be attempted using semianual colonoscopy with multiple mucosal biopsies and double-contrast barium enema (15). Our patients were an object lesson in that failure to be assiduous led to a delay in diagnosis. Whether proctocolectomy should be considered prophylactically after liver transplantation is an unresolved issue.

REFERENCES

6. Collins RH, Feldman M, Fordtran JS. Colon cancer, dysplasia,