angioplasty for renal artery stenosis occurring early after renal transplantation. We do suggest, however, that the procedure be seriously considered for patients with late and localized renal artery stenosis who have refractory hypertension or declining renal function that cannot be explained by other causes, or both. The procedure should, of course, be carried out only by angiographers experienced in the technique.

References

- 1. GRÜNTZIG A, VETTER W, MEIER B, et al: Treatment of renovascular hypertension with percutaneous transluminal dilatation of a renal-artery stenosis. *Lancet* 1: 801, 1978
- 2. MAHLER F, KRNETA A, HAERTEL M: Treatment of renovascular hypertension by transluminal renal artery dilatation. Ann Intern Med 90: 56, 1979

- 3. KATZEN BT, CHANG J, LUKOWSKY GH, et al: Percutaneous transluminal angioplasty for treatment of renovascular hypertension. *Radiology* 131: 53, 1979
- 4. MILLAN, VG, MAST WE, MADIAS NE: Nonsurgical treatment of severe hypertension due to renal-artery intimal fibroplasia by percutaneous transluminal angioplasty. N Engl J Med 300: 1371, 1979
- 5. WEINBERGER MH, YUNE HY, GRIM CE, et al: Percutaneous transluminal angioplasty for renal artery stenosis in a solitary functioning kidney. An alternative to surgery in the highrisk patient. Ann Intern Med 91: 684, 1979
- 6. ATHANASOULIS CA: Therapeutic applications of angiography (second of two parts). N Engl J Med 302: 1174, 1980
- 7. GRÜNFELD JP, KLEINKNECHT D, MOREAU JF, et al: Permanent hypertension after renal homotransplantation in man. *Clin Sci Mol Med* 48: 391, 1975

- 8. BACHY C, VAN YPERSELE DE STRIHOU C, ALEXANDRE GPJ, et al: Hypertension after renal transplantation. *Proc Eur Dial Transplant Assoc* 12: 461, 1976
- 9. RAO TKS, GUPTA SK, BUTT KMH, et al: Relationship of renal transplantation to hypertension in endstage renal failure. Arch Intern Med 138: 1236, 1978
- 10. SMELLIE WAB, VINIK M, HUME DM: Angiographic investigation of hypertension complicating human renal transplantation. Surg Gynecol Obstet 128: 963, 1969
- 11. MORRIS PJ, YADAV RVS, KINCAID-SMITH P, et al: Renal artery stenosis in renal transplantation. Med J Aust 1: 1255, 1971
- LACOMBE M: Arterial stenosis complicating renal allotransplantation in man: a study of 38 cases. Ann Surg 181: 283, 1975
- 13. SCHRAMEK A, ALDER O, HASHMONAI M, et al: Hypertensive crisis, erythrocytosis, and uraemia due to renalartery stenosis of kidney transplants. *Lancet* 1: 70, 1975

Increased requirement for thyroid hormone after a jejunoileal bypass operation

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A jejunoileal bypass operation can result in metabolic disturbances, various vitamin deficiencies and fatty degeneration or cirrhosis of the liver.¹ It has also recently been reported to cause thyroid hormone

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Reprint requests to: Dr. Robert Volpé, Department of medicine, The Wellesley Hospital, Rm. 112 D, Jones Building, 160 Wellesley St. E, Toronto, Ont. M4Y 1J3 malabsorption, at least in one patient.² We describe herein a patient in whom a fourfold increase in the daily dose of thyroxine (T_4) was necessary for euthyroidism after a jejunoileal bypass operation.

Case report

A severely obese woman (maximum weight 127 kg, height 154 cm) was found to be hypothyroid at the age of 25 years. She failed to lose weight taking levothyroxine, 0.2 mg/d orally, as hormone replacement therapy, but the symptoms and signs of hypothyroidism cleared. Her obesity remained a persistent serious problem. As a last resort a jejunoileal bypass operation was performed when she was 39 years old: the jejunum was divided 10 cm beyond the ligament of Treitz, and the proximal portion was anastomosed end to side to the terminal ileum 14 cm from the ileocecal valve.

Over the next 8 months she defecated about five times daily and lost 55 kg. The levothyroxine therapy was continued, but fatigue, cold intolerance, bloating, and dry hair and skin developed; she recognized these as symptoms of recurrent hypothyroidism. The serum levels of T_4 and triiodothyronine (T_3) were low and the level of thyrotropin was high (Table I). Liver function tests, including measurement of the serum levels of aspartate aminotransferase, alanine aminotransferase and albumin as well as the prothrombin time, gave normal results, but the serum alkaline phosphatase level was 121 IU/l (normal range 25 to 90 IU/l).

Despite an increased dose of T_4 the serum thyrotropin level remained high. The medication was changed to T_3 , and a daily dose of 300 μg (75 μg every 6 hours) was finally achieved. At this daily dose the serum level of T_3 was 100 ng/dl 3 hours after a dose of T_3

was given. Treatment with T_4 was resumed with the daily dose gradually increased; clinical and biochemical euthyroidism was finally achieved at the very high daily dose of 0.8 mg. She has subsequently felt well.

Discussion

Before the jejunoileal bypass operation the patient stayed euthyroid by taking 0.2 mg of T₄ daily; following the bypass procedure, however, she required as much as 300 μ g of T₃ or 0.8 mg of T₄ daily to remain euthyroid. This strongly suggests that thyroid hormone malabsorption was a consequence of the bypass procedure, although accelerated thyroid hormone degradation is another possible explanation. Azizi, Belur and Albano² recently documented impaired absorption of thyroid hormones labelled with radioactive isotopes in a patient who had undergone the same procedure.

It has been demonstrated that about 60% (range 35% to 90%) of orally administered T_4 is absorbed in healthy persons,³⁻⁵ and experiments in animals have shown that the time course for absorption is comparable in all segments of the small and large intestine.⁶ T_3 is absorbed more quickly⁷ and more completely⁸ than T_4 . Although no consistent effect on thyroid hormone absorption of either cirrhosis or hypothyroidism has been found,⁵ pancreatic steatorrhea impairs T_4 absorption,⁹ and the fecal excretion of T_4 is directly related to the fecal mass.¹⁰

Our patient required a fourfold increase in the daily dose of T₄ for euthyroidism after the jejunoileal bypass operation; this indicated serious impairment of thyroid hormone absorption. Therefore, in patients requiring thyroid hormone therapy who undergo this operation, and in patients with intact jejunoileal bypasses who subsequently become hypothyroid, a much higher daily dose of thyroid hormone for adequate replacement is to be expected. T₄ therapy seems preferable to T₃ as it avoids the large and rapid changes in serum levels of thyroid hormones, specifically T₃, related to the administration of a tablet several times a day.

References

- 1. DEWIND LT, PAYNE JH: Intestinal bypass surgery for morbid obesity. Long-term results. JAMA 236: 2298, 1976
- 2. AZIZI F, BELUR R, ALBANO J: Malabsorption of thyroid hormones after jejunoileal bypass for obesity. Ann Intern Med 90: 941, 1979
- 3. ODDIE TH, FISHER DA, ROGERS C: Whole-body counting of 131-Ilabeled thyroxine. J Clin Endocrinol Metab 24: 628, 1964
- 4. VAN MIDDLESWORTH L: Thyroxine requirement and the excretion of

Serum value [normal range] Thyroid Total T₄ Free thyroxine Total T₃ TSH (µg/dl) [4–11] hormone given, index* (ng/dl) (mIU/mI) Date daily dose [4-11] [90-210] [0-10] T₄, 0.1 mg Sept. 1971 2.5 1.9 Feb. 1972 T₄, 0.2 mg 6.5 6.1 Sept. 1977 Jejunoileal bypass operation May 1978 T₄, 0.2 mg 3.3 2.7 3.2 >80 -T₄, 0.4 mg 3.7 80 T₄, 0.4 mg July 1978 4.2 3.7 72 78 T₃ therapy begun† Sept. 1978 T₃, 10-20 μg 80 49 66 27 Oct. 1978 T₃, 40 μg 1‡ 1 T₃, 125 μg T₃, 200 μg Nov. 1978 1 1 1 Feb. 1979 1 Mar. 1979 T₃, 300 μg 1 1 100 3.5 Nov. 1979 T₄, 0.8 mg 8.3 9.4 105

Table I—Effect of jejunoileal bypass operation on the serum concentrations of thyroxine (T₄), triiodothyronine (T₃) and thyrotropin (TSH) in a hypothyroid patient taking T₄ or T₃ orally

*The total serum T_4 level times the T_3 uptake expressed as a percentage of the reference standard.

[†]T₃ was given in four divided doses a day.

 \ddagger low serum T₄ level is to be expected with T₃ therapy.

thyroxine metabolites, in *Clinical Endocrinology*, vol 1, ASTWOOD EB (ed), Grune, New York, 1960, p 103

- 5. HAYS MT: Absorption of oral thyroxine in man. J Clin Endocrinol Metab 28: 749, 1968
- 6. CHUNG SJ, VAN MIDDLESWORTH L: Absorption of thyroxine from the intestine of the rat. Am J Physiol 212: 97, 1967
- 7. Idem: Absorption of thyroxine from the small intestine of rats. *Endocrinology* 74: 694, 1964
- HAYS MT: Absorption of triiodothyronine in man. J Clin Endocrinol Metab 30: 675, 1970
- 9. HISS JM JR, DOWLING JT: Thyroxine metabolism in untreated and treated pancreatic steatorrhoea. J Clin Invest 41: 988, 1962
- 10. HÉROUX O, PETROVIC VM: Effect of high- and low-bulk diets on the thyroxine turnover rate in rats with acute and chronic exposure to different temperatures. Can J Physiol Pharmacol 47: 963, 1969



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LE DIAGNOSTIC PRÉNATAL. Cahier de Bioéthique. 2. Collection publiée par le Centre de bioéthique de l'Institut de recherches cliniques de Montréal. 281 pp. Illust. Les Presses de l'Université Laval, Québec, 1980. \$14, broché. ISBN 2-7637-6914-4

FAMILY MEDICINE. A Guidebook for Practitioners of the Art. David B. Shires and Brian K. Hennen. 512 pp. Illust. McGraw-Hill Book Company, New York, 1980. \$15.95, paperbound. ISBN 0-07-056920-7

INTERVENTIONAL DIAGNOSTIC AND THERAPEUTIC PROCEDURES. Barry T. Katzen. 155 pp. Illust. Springer–Verlag New York Inc., New York, 1980. \$22.50. ISBN 0-387-90390-9

LEAD TOXICITY. Edited by R.L. Singhal and J.A. Thomas. 514 pp. Illust. Urban & Schwarzenberg, Inc., Baltimore; the Macmillan Company of Canada Limited, Toronto, 1980. \$49.50. ISBN 0-8067-1801-3

LUNG DISEASE. State of the Art. 1978– 1979. Edited by John F. Murray. 463 pp. Illust. American Lung Association, New York, 1980. \$12.50. ISBN 0-915116-04-4

MANUAL OF ANTIMICROBIAL THER-APY AND INFECTIOUS DISEASES. Mickey S. Eisenberg, Clifton Furukawa and C. George Ray. 282 pp. Illust. W.B. Saunders Company Canada, Ltd., Toronto, 1980. \$15.55, spiralbound. ISBN 0-7216-3348-X

MANUAL OF ECHOCARDIOGRAPHIC TECHNIQUES. Betty J. Phillips and Vincent E. Friedewald, Jr. 276 pp. Illust. W.B. Saunders Company Canada, Ltd., Toronto, 1980. \$29.95. ISBN 0-7216-7219-1

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