Toxicity of Thallium

Thallium is an extremely toxic and cumulative poison. Within three years of its discovery in 1861 a paper was published warning of its toxicity to animals and man. The degree of toxicity has been variously estimated as lying somewhere between that of lead and arsenic. But the recent trial of Graham Frederick Young for poisoning two of his workmates is said to be the first case of homicide by thallium in British legal history.

Like most elements soon after their discovery, the thallium salts have been tried for the treatment of certain chronic inflammatory diseases. They were also used as a temporary depilatory in cases of ringworm of the scalp. But their frequently poisonous effects led to thallium therapy being abandoned. The difficulty is that the therapeutic dose is too close to the lethal dose. E. Browning gives the therapeutic dosage as being 8 mg of thallium acetate per kg bodyweight, while 12 mg per kg is the minimum lethal dose for an adult. But in industry the use of thallium is increasing. Because of its high refractive index it is incorporated into the manufacture of optical lenses, in infra-red optical instruments, and in imitation precious jewellery. It is used in alloys along with silver and lead, serves as a catalyst in a number of organic reactions, and has also been included in fireworks. Recently thallium salts have been used in the crystals which serve as scintillation counters. But it is in the form of rodenticides and insecticides that thallium constitutes the main hazard to the general public.

Thallium salts can enter the body in three ways—by inhalation as a dust, by ingestion either from contaminated food from the hands, or by absorption through the skin. J. C. Munch examined the reports of 12 non-fatal cases of industrial poisoning and established that the main symptoms were fatigue, limb pains, and loss of hair. More severe poisoning results in peripheral neuritis, proteinuria, and joint pains, but occasionally neurological signs are the presenting factors. A striking example of non-industrial poisoning was described by Munch and colleagues in 1933 from California. At least 30 people accidentally ate barley grain impregnated with 1% thallium sulphate intended as a rodenticide. Six died of acute thalliotoxisnosis within 16 days and a seventh two months later; and 20 persons developed gastroenteritis, abdominal colic, peripheral neuritis, stavismus, disorientation, convulsions, joint pains, and alopecia, mostly within three days. A similar accident involving nine members of three families occurred, despite warnings, in Mexico in 1964.

Recently W. J. Bank and colleagues studied five patients with thallium intoxication, two of whom were children who had eaten poisoned bait. All of them developed neurological symptoms, and alopecia was present in four of the five. They stress the importance of including thallium in the differential diagnosis when early bizarre neurological signs present themselves. Furthermore they advocate the use of a simple screening test of urine for thallium salts.

The treatment of thallium poisoning poses some difficulties. The principle is to increase urinary and faecal excretion, though the rate of each is slow. Chelating agents have been proposed, but results with these have been inconsistent. In addition potassium chloride is useful, but Bank and colleagues point out that such a regimen is limited by the amount of thallium that can be released into the blood without worsening the symptoms. However, it is probably the better method of treatment of the two.

While thallium salts continue to be used as rodenticides the danger of accidental poisoning, especially to children, will continue to exist. So long as the hazard remains, its possibility should be borne in mind in all cases of unexplained neurological symptoms and peripheral joint pains. The early use of a screening test on the urine would appear to be far better than waiting to see if the hair falls out.

**Urinary Incontinence in Women**

The diagnosis of stress incontinence is often difficult. It requires differentiation from frequency, urgency, and urge incontinence, and even very rarely from true incontinence due to a small urinary fistula.

The abnormality underlying the symptom is not always clear, for even the famed loss of the posterior urethrovaginal angle, seen by cine-radiography, is not present in
every case. But loss of this angle is to be found in about 90% of lateral cysto-urethrograms of women with stress incontinence and therefore the most consistent abnormality detected. The cure of stress incontinence is usually by some form of surgery, most often by the vaginal route, and when this fails further measures include cysto-urethropexy and slings of some sort running under the bladder neck and fixed to the abdominal wall. The cure rate for these various procedures ranges, according to the surgeon, from about 50% to 90%, but it is well known that assessment of results is difficult and that the percentage of cures diminishes with time. Understanding of the disorder obviously leaves much to be desired.

Bladder function, and failure of function, must depend on the relationships between intravesical pressure and the resistance to the outflow of urine. Intravesical pressure is a compound of detrusor activity and intra-abdominal pressure. It is this last, suddenly raised by coughing, sneezing, and other varieties of straining, which is the proximate cause of stress incontinence. Resistance to outflow must depend on the arrangement of muscle at the internal meatus—the internal sphincter—and on the voluntary external sphincter (compressor urethrae), together with the length and bore of the urethra and the posterior urethro-vesical angle. The length of the urethra is not usually thought to be of great importance, since the lower half of it may be removed in some cancer operations without impairment of continence. This removes the compressor urethrae too, so its role in continence is probably minor, though when the stream of urine during voiding is checked it is the external sphincter which first stops the flow. In urge incontinence it is probably this muscle which is able to hold back the urine temporarily. Moreover, provided the woman with stress incontinence knows that the stress is imminent, she is often able to prevent the leakage of urine.

Against this confused background some new studies by A. S. Moolgaoker and colleagues are very welcome. Over the past four years they have investigated women with varieties of incontinence of urine resembling stress incontinence. Some of their patients had had previous operations for their urinary troubles. A most important finding was that the clinical history could be and often was misleading and that Bonney’s test was valueless. Their special investigations were the rate of urine flow, the measurement of urethral length, cine-urethrocytography, and pressure recordings during the cine-radiography. The methods were used on outpatients. On the basis of their findings they now recognize three major mechanisms of incontinence either alone or in combination. They are sphincter weakness, detrusor instability, and urethral narrowing.

Sphincter weakness is characterized by a normal return to basal intravesical pressure after coughing and straining, a normal flow rate of about 20 ml per second or above, a resting length of the urethra of 3 cm or more, the presence of contrast medium at least in the proximal part of the urethra on straining, an obliterated or partially obliterated posterior urethro-vesical angle, and a descent of the urethra on straining measured by its angular displacement of more than 20°. Obliteration of the posterior urethro-vesical angle was seen in 64% of cases.

With detrusor instability the intravesical pressure returns only slowly to normal after coughing and straining, the flow rate is normal, the bladder capacity is reduced, contrast medium is found in the whole or part of the urethra on straining, the urethral length is within normal limits. Furthermore, during voiding the patients cannot check the stream (in contrast with those with simple sphincter weakness) and the descent of the urethra on straining is less than 20°.

In patients with urethral narrowing, often following a previous operation, intravesical pressure returns to normal very quickly after coughing and straining, but the flow rate is reduced to under 10 ml per second. The patient has to strain to pass water, and there may be trabeculation of the bladder. Urethral length is normal, but it shows narrowing or strictures. Contrast medium does not enter the proximal urethra on straining and the descent of the urethra is less than 20°. There is often residual urine.

Sphincter weakness should respond to the usual surgical measures for clinical stress incontinence, though the authors recommend that a cystocele with a descent of more than 20° of the urethra on straining should have a vaginal repair and a sling operation. If the cystocele is small then a Marshall-Marchetti operation is proposed, and only if the angle of inclination is less than 20° should a vaginal approach alone be used. With detrusor instability, the bladder being of small capacity and in a state of high tone, the treatment should be medical. The patient is taught bladder drill and given bladder relaxants such as propantheline bromide 30 mg four times a day or a capsule of ephedrine sulphate 25 mg and amylobarbitone 50 mg three times a day. If the urethra is narrowed the problem is one of obstruction, a kind of chronic retention with overflow, so this is best dealt with by urethral dilatation or urethrotomy with an Otis urethrotome, incising the full length of the urethra.

First results suggest improvement over the present hit-or-miss methods based on clinical diagnosis and vaginal surgery. The techniques are elaborate, so that not all gynaecological and radiological departments may be able to cope with them, but the investigations certainly ought to be more widely available then they are at present. At least it is probable that no woman should be asked to undergo a second operation to cure her stress incontinence without full investigation on these lines.