ATROPINE POISONING TREATED BY FORCED DIURESIS

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A THIRTY-ONE-YEAR-OLD male patient was admitted to hospital three hours after swallowing 7.5 ml. of 1% atropine sulphate eyedrops in a suicidal attempt.

Case Report

He was found to be hallucinated, irritable, and tremulous. His pupils were widely dilated; muscle tone was increased, with exaggerated reflexes and bilateral ankle clonus. His tongue was dry and furred. His was increased, with exaggerated reflexes and his pupils, however, remained widely dilated for twenty-four hours, but gradually returned to normal during the subsequent twenty-four hours.

Discussion

Carter (1940) and Glaister (1957) describe patients who have recovered following consumption of similar quantities of atropine. Only symptomatic and supportive treatment have been available, however, and it seems agreed that chemical methods of treatment are ineffective (Goodman and Gilman, 1955; Graham, 1962).

Our results indicate that with forced diuresis 25.7 mg. of atropine sulphate, that is 34% of the ingested quantity, were excreted in the two hours following commencement of therapy or in the two hours after taking the drug. 30.9 mg. (41% of the ingested dose) were excreted in 10 hours following commencement of therapy, that is in the 14 hours after taking the drug. During this period the patient passed 5,800 ml. of urine. By comparison Tonnesen (1950) showed that humans, given doses of 2 to 5 mg. of atropine sulphate orally, excreted an average 12.7% of the ingested dose during the next two days.

We were able to detect atropine in the patient's urine for 24 hours after taking the drug, but no atropine could be detected chemically or by the cat's eye test (Sollmann, 1948; Krantz and Carter, 1961) after this.
Tentative Excretion. (Results expressed as (C₁₇H₂₃O₃N₂H₂SO₄)₀₂H₂O.)

<table>
<thead>
<tr>
<th>Time after commencement of diuresis</th>
<th>Volume of urine (ml)</th>
<th>Atropine concentration mg./100 ml</th>
<th>Atropine excreted mg.</th>
</tr>
</thead>
<tbody>
<tr>
<td>1st hour</td>
<td>500</td>
<td>2.3</td>
<td>11.3</td>
</tr>
<tr>
<td>2nd hour</td>
<td>1,360</td>
<td>1.1</td>
<td>14.4</td>
</tr>
<tr>
<td>3rd and 4th hours</td>
<td>1,400</td>
<td>0.2</td>
<td>2.8</td>
</tr>
<tr>
<td>5th and 6th hours</td>
<td>380</td>
<td>0.2</td>
<td>0.8</td>
</tr>
<tr>
<td>7th and 8th hours</td>
<td>825</td>
<td>0.1</td>
<td>1.1</td>
</tr>
<tr>
<td>9th and 10th hours</td>
<td>1,325</td>
<td>trace</td>
<td>0.5</td>
</tr>
<tr>
<td>11th—14th hours</td>
<td>1,490</td>
<td>trace</td>
<td>0.2</td>
</tr>
<tr>
<td>15th—18th hours</td>
<td>3,000</td>
<td>trace</td>
<td>&lt;0.1</td>
</tr>
<tr>
<td>Total</td>
<td>9,290</td>
<td></td>
<td>31.1</td>
</tr>
</tbody>
</table>

Subsequent specimens were tested chemically and biologically by the cat's eye test but showed no activity in the original solution or in the extracts prepared from them.

* Forced diuresis commenced four hours after ingestion of atropine.

Our results, therefore, are in agreement with the known rapid early excretion of atropine (Tonnesen, 1950). Possible explanations for the rapid fall in the excretion rate are the fixation of atropine in the tissues and the process of detoxication in the liver (Goodman and Gilman, 1955; Beckman, 1961). Although it is difficult to draw a strict comparison between Tonnesen's results and our own because of the difference in the amount of drug ingested, we feel that the quantity of atropine excreted has been significantly increased by producing a substantial diuresis in the early stages of the intoxication.

Summary

The case is described of a patient who took a large quantity of atropine (75 mg.) in a suicidal attempt. He recovered rapidly on a regime of forced diuresis. His excretion of atropine was estimated and it was shown that 41% of the ingested dose was excreted in the 10 hours following commencement of treatment, at the end of which time he had recovered from his intoxication apart from continued dilution of the pupils. It is concluded that forced diuresis is of value in the treatment of atropine poisoning.

Appendix: Method of estimating atropine

Transfer an appropriate volume of urine to a separator and add 2 ml. O.1 N HCl or sufficient to make acid. Dilute, if necessary, to about 50 ml. with water and extract with ether (50 ml.).

Run off the lower aqueous layer into a second separator, wash ether layer with water (10 ml.) and add the washing to the aqueous layer in the second separator.

Make the aqueous layer alkaline with ammonia and extract twice with ether (50, 40 ml.). Wash the bulked ether extracts with water (2 x 5 ml.), transfer the ether to a flask and evaporate to small volume (about 5 ml.). Transfer the concentrated ether quantitatively to an evaporating basin and evaporate to dryness.

Add 0.3 ml. fuming nitric acid to the residue and evaporate once more to dryness. Cool the basin and add dimethylformamide (cf. Freeman, 1955) (5.00 ml.) and ethanolic potassium hydroxide (0.50 N, 0.50 ml.). Compare the colour produced immediately with a blank of distilled water at 500 μ. Subtract the reading for a blank on normal urine and read off the amount of atropine from a curve obtained by treating known amounts of atropine sulphate with fuming nitric acid and continuing as described above from the words 'evaporate once more to dryness...

We would like to thank Mr. J. L. Paterson, Department of Pharmacy, the Royal College of Science and Technology, for performing the biological tests, and Dr. Alexander Brown for permission to publish the case report, and for helpful advice.

REFERENCES


