makes it a potentially useful drug to deliberately enhance epileptogenic activity in patients undergoing electrocorticography as part of cortical resection of epileptogenic tissue for the treatment of refractory epilepsy and would be an acceptable alternative to methohexital, especially in those patients allergic to barbiturates.

REFERENCES


Precurarization Inhibits Maximal Ventilatory Effort

DAVID L. BRUCE, M.D.,* JOHN B. DOWNS, M.D.,† PRAMOD S. KULKARNI, M.B.,‡ LEVON M. CAPAN, M.D.§

Pretreatment with 3–5 mg d-tubocurarine (dTC) prevents succinylcholine- (SCh) induced muscle fascicu-

* Professor, Anesthesiology, New York University Medical Center.
† Associate Professor, Anesthesiology, Northwestern University Medical School; Medical Director, Anesthesiology and Pulmonary Medicine, Mercy Hospital, Urbana, Illinois.
‡ Fellow, Anesthesiology, Northwestern University Medical School.
§ Associate Professor, Clinical Anesthesiology, New York University Medical School.

Received from the Department of Anesthesiology, New York University Medical Center, New York, New York, Department of Anesthesiology, Northwestern University Medical School, and Department of Anesthesiology, Mercy Hospital, Urbana, Illinois. Accepted for publication May 2, 1984.

Address reprint requests to Dr. Bruce: Department of Anesthesiology, University of Mississippi Medical Center, 2500 N. State Street, Jackson, Mississippi 39216.

Key words: Larynx: muscles. Neuromuscular relaxants: d-tubocurarine. Ventilation: pattern.

This has been shown to attenuate the increased intragastric and intraocular pressures that usually accompany fasciculation and to reduce the incidence of post-SCh myalgia. Most patients today are given SCh before tracheal intubation, and many of these are pretreated with a nondepolarizing drug such as dTC. In the interval between injection of dTC and the induction agent, an occasional patient becomes dyspneic and, very rarely, one develops apnea and must be ventilated. In these circumstances, dyspnea may be due to respiratory muscle weakness, airway obstruction, or apprehension. Previous studies of low-dose dTC effect on muscle power of awake subjects employed doses of 0.1 mg/kg or greater. The usual dose of dTC used clinically to pretreat adults is 3 mg, which is 0.043 mg/kg for a 70-kg patient. We chose a comparable dose, 0.05 mg/kg, to study dTC effect on negative inspiratory pressure (NIP) and respiratory flow-volume loops of premedicated

Downloaded From: http://anesthesiology.pubs.asahq.org/pdfaccess.ashx?url=/data/journals/jasa/931419/ on 11/26/2018
patients. A somewhat similar study in volunteers given 0.014 mg/kg pancuronium was reported in 1980.11

METHODS

NIP measurements were made in 62 adult patients requiring surgical anesthesia, in a project approved by the Human Subjects Review Committee of NYU Medical Center. Patients were unselected for site or type of surgery, were of ASA physical status 1 or 2, and were premedicated according to preferences of the anesthesiologists scheduled to provide their care. Ten were not premedicated, eight were given oral diazepam preoperatively, two received only hydroxyzine, and the remainder received an opiate alone (8) or in combination with hydroxyzine (27) or pentobarbital (7). The patients' ages were from 19 to 62, with a mean of 33.4 years.

Preanesthetic preparations were made in a routine manner; then NIP was measured with a respiratory force meter (Boehringer) calibrated to register from 0 to 60 cmH2O, positive or negative. Multiple measurements were made until reproducible values were obtained. They were then given intravenous dTC, 0.05 mg/kg, without their knowledge, and NIP was measured 1, 2, and 3 min later. At each interval, patients were asked: "Are you comfortable?"; "Does your breathing feel normal to you?"; "Does your eyesight seem normal?". Anesthesia and operation then proceeded as usual without further investigative intervention. A second study subsequently was performed with healthy patients at Mercy Hospital, Urbana, Illinois, with approval of their institutional review board. Starting from their residual volumes, these patients inspired and expired maximally during spirometric flow-volume loop recordings (Hewlett-Packard Pulmonary Function System®). After stable baseline measurements were reproduced, 0.05 mg/kg dTC was given and the spirometry was repeated 1, 2, and 3 min later. These patients were not asked about symptoms. Anesthesia and operation then proceeded according to routine. The ratio of mid-expiratory to mid-inspiratory flow rates (MEF50/MIF50) were calculated from the airflow tracings.12 Statistical analyses of data were by two-way analysis of variance and Bonferroni t test.13 A P value less than 0.01 was considered significant.

RESULTS

Mean NIP values decreased after dTC, as shown in table 1. Analysis of individual changes showed that, at all three post-dTC intervals, these changes were highly significant statistically. However, not all patients had diminutions in NIP, as shown in table 2. Approximately one-third of the patients showed no change or an increase, i.e., a greater negative pressure generated, at these times. Of the 62 patients, 40 reported visual disturbances (usually diplopia), four had dyspnea at 2 and 3 min after dTC, and one had mild fasciculations after SCh. Two of these four were given no premedicants; the others had only hydroxyzine. Their NIP data are shown in table 3. All four of these patients also had diplopia, and none fasciculated after SCh.

Flow volume loops from the 15 patients in the second study showed an increased MEF50/MIF50 in 12, no change in 2, and a very slight decrease for 1. For the total group, the mean control ratio was 0.90 ± 0.32 (SD) and the ratio 2 or 3 min after dTC was 1.42 ± 0.65 (P < 0.01). Some tracings were technically unsatisfactory, requiring the selection of the better of the 2- or 3-min post-dTC records. A representative tracing is reproduced in figure 1, which shows a control expiratory to inspiratory ratio greater than usual, an insignificant reduction of vital capacity (VC) after dTC, and a distinct flattening of the inspiratory limb of the loop after dTC, causing the MEF50/MIF50 to increase from 1.41 to 2.32. This was the only case in which VC decreased after dTC.

DISCUSSION

Our patients were studied under clinical conditions to assess the incidence of problems from "precurarization." The effect of dTC undoubtedly would have been greater if the interval between this drug's injection and induction of anesthesia were allowed to be longer than 3 min. We chose not to do this, because our experience suggests that most clinicians wait no longer than 3 min. The study was deliberately designed to reflect the reality of routine practice. Because our patients were premedicated.

<table>
<thead>
<tr>
<th>Time, Postcurare, 0.05 mg/kg</th>
</tr>
</thead>
<tbody>
<tr>
<td>1 min</td>
</tr>
<tr>
<td>------</td>
</tr>
<tr>
<td>Decrease</td>
</tr>
<tr>
<td>None</td>
</tr>
<tr>
<td>Increase</td>
</tr>
</tbody>
</table>

| Total patients | 62  | 62  | 62  |
TABLE 3. NIP Values of Dyspneic Patients (NIP, cmH2O)

<table>
<thead>
<tr>
<th>Patient</th>
<th>Baseline</th>
<th>1 min</th>
<th>2 min</th>
<th>3 min</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>30</td>
<td>18*</td>
<td>20*</td>
<td>20*</td>
</tr>
<tr>
<td>2</td>
<td>80</td>
<td>75</td>
<td>60*</td>
<td>60*</td>
</tr>
<tr>
<td>3</td>
<td>22</td>
<td>26</td>
<td>23</td>
<td>16*</td>
</tr>
<tr>
<td>4</td>
<td>32</td>
<td>32</td>
<td>24*</td>
<td>22*</td>
</tr>
</tbody>
</table>

* Dyspneic.

sensitivities to the sensation of breathing impairment, just as there are large differences in pain threshold. We suspected that the site of this sensation would be the larynx rather than the thorax. If the striated muscle causing vocal cord abduction during inspiration were relaxed somewhat by a low dose of d'TC, a slightly smaller laryngeal inlet would result in changes in laryngeal inspiratory airflow that could be experienced as shortness of breath.

The flow volume loops confirmed our suspicion of a laryngeal effect of d'TC, 0.05 mg/kg. The only reasonable explanation for the flattened, reduced inspiratory airflow loops in 12 of 15 patients is that vocal cord abduction was impaired by the d'TC. Although VC was unaffected, and patient safety presumably was not compromised by d'TC pretreatment, a laryngeal effect of relaxants should not be considered unimportant. The occasional patient may perceive this change and suffer significant anxiety from it, just before hypnosis is induced, if he or she is ignored by the anesthetist.

Rao and Jacobs reported that 4 of 15 healthy volunteers suffered significant pulmonary dysfunction and dyspnea after 0.014 mg/kg pancuronium. It is difficult to compare their data with ours, because they studied volunteers over a longer time period than 3 min, but easy to agree with their conclusion, "ventilation should be monitored in all patients following pretreatment."

Those patients for whom such pretreatment is chosen should be monitored closely and reassured that visual difficulties they may experience are harmless and temporary. Their ventilation also should be observed closely to watch for the rare occurrence of the "mild myasthenic state," while reassuring them that they are breathing normally even if they feel they are not. Once a nondepolarizing relaxant has been given, it is imprudent to direct one's attention to anything or anyone other than the patient until hypnosis and a protected airway have been established.

The authors acknowledge the assistance given by Morris Antebi, M.D., and Isidore Lee, M.D.

REFERENCES

1. Jensen EC, Hansen PH: Objective measurement of succinylcholine induced fasciculations and the effect of pretreatment with pancuronium or gallamine. ANESTHESIOLOGY 51:159-160, 1979
4. Miller RD, Wray WL: Inhibition of succinylcholine-induced in-
creased intragastric pressure by nondepolarizing muscle relaxants and lidocaine. ANESTHESIOLOGY 34:185-188, 1971