

period of extinction, the rats continued to run in response to the buzzer with a high degree of consistency until a point was reached where, relatively suddenly, no such responses were made. Inasmuch as the individual Vincent curves have essentially the same form found in the average curves, and inasmuch as the raw data reveal the same sudden termination of the conditioned response, the genuineness of the phenomenon seems well attested.

The extinction curves presented by Kleitman and Crisler and by Switzer are based on the magnitude of the conditioned response and not upon its frequency. Does the present conditioned locomotor response suffer a diminution in magnitude during the period of extinction so that the rat finally and gradually reaches a point where no movement is made to the buzzer? The present apparatus with its circular pathway permitted runs varying in magnitude from zero to an indefinitely large value. An examination of the records, however, shows that, although the lengths of the runs varied, there was no trend toward shorter and shorter runs. A curve relating the magnitude of the response to the amount of elapsed time would, therefore, have essentially the same form as the present curves, based on frequency of response. The suggestion is therefore apparent, from a comparison of the various curves found in the literature, that there is no one type of curve for extinction. Rather the character of the curve will depend upon many factors, including the type of response conditioned.

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DISSOCIATION OF THE PYRAMIDAL AND EXTRAPYRAMIDAL FUNCTIONS OF THE FRONTAL LOBE

SECTION of one pyramidal tract below its last large supra-segmental connection with the pons produces in cats a syndrome of deficit describable either as the specific reactions lost or impaired, or as a depression of phasic activity in general; or as a loss of excitation at the final moto-neurons in the cord. Spasticity, or other evidence of release, is absent. Stimulation of the motor cortex either immediately or months after such a section demonstrates the preservation within that cortex of inhibition, effective on tonic or clonic states present in the limbs, together with the abolition of the familiar motor function. An extrapyramidal type of motor activity is at the same time uncovered.

Repetition of this lesion in rhesus monkeys produces a similar, though graver syndrome of deficit, again without spasticity. And again, stimulation of the cortex brings inhibition to bear on activity present in the limbs after destruction of the characteristic

motor responses of the precentral gyrus. In the monkey, however, this inhibition is not confined, as in the cat, to the immediate motor region but is exercised by motor, premotor and prefrontal cortex, and from the second of these, most vigorously. From this region, for instance, tonic closure of the fingers into a grasp is most easily released. Moreover, although the fine type of movement characteristic of stimulation of the precentral gyrus is totally abolished by the lesion, the so-called adverse movements survive, and can be elicited not only from the premotor region but from the precentral and postcentral gyri, as well. Furthermore, after section of one or both pyramidal tracts at the level of the trapezoid body, with or without time for degeneration, epileptiform convulsions are easily set in train by stimulation of the motor, premotor and prefrontal regions, and even of spots in the parietal lobe. These involve all four limbs and face, show typical progression, tonic and clonic phases and after-exhaustion, resembling in essentials the clinical Jacksonian seizure.

One may conclude, therefore, that pyramidal and extra-pyramidal functions of the motor cortex and adjacent regions may be dissociated by pyramid section. The rapidly executed, fairly discrete movements, long familiar from cortical stimulation, are thus demonstrated to be mediated exclusively by the cortico-spinal tract. On the other hand, the integrity of this tract is not necessary to the exhibition of the larger movements called adverse, nor to the exercise of the very important inhibitory function of the cortex. Consequently, these, together with the epileptiform convulsions elicited, represent extrapyramidal activities of the cerebral cortex.

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THE NATURE OF THE BARBITURATE-PICROTOXIN ANTAGONISM¹

MALONEY, Fitch and Tatum² and Maloney and Tatum³ have shown that picrotoxin is a very effective antidote in acute barbiturate poisoning. We have had ample occasion to confirm their data in animals poisoned with large doses of intravenously administered barbiturates. Quoting only extreme cases, the results in Table 1 were obtained:

To a number of dogs and rabbits, we administered the minimum anesthetic doses of different barbiturates, waited for the onset of anesthesia and then recorded

¹ From the Department of Pharmacology and Materia Medica, Georgetown University, School of Medicine, Washington, D. C.

² Maloney, Fitch and Tatum, *Jour. Pharmacol. Exper. Therap.*, 41: 465, 1931.

³ Maloney and Tatum, *Jour. Pharmacol. Exper. Therap.*, 44: 337, 1932.

TABLE 1

Animal	Drug	Total dose of barbiturate, mgm per kgm	Total dose of picrotoxin, mgm per kgm	Time
Dog	Sodium barbital	1,650	20	Died in 1½ hrs.
Dog	Sodium phenobarbital	375	6	Died in 2 hrs.
Dog	Sodium pentobarbital	160	38	Died in 68 min.
Dog	Sodium pentobarbital	100	18	Recovered in 12 hrs.
Rabbit	Sodium pentobarbital	100	24	Recovered in 5½ hrs.

the animal's respiration and carotid blood pressure on a kymograph tracing. While the animal was thus under constant observation further doses (usually $\frac{1}{4}$ of the M.L.D. was used for a single injection) of the barbiturate originally administered were given intravenously. It was observed that these injections produced in each instance a sharp fall in blood pressure and a slowing or stoppage of respiration. The blood pressure fell after each additional injection. Death was due to a combination of circulatory collapse and respiratory failure.

The antidotal effect of different doses of picrotoxin

was manifested in four different ways: (a) Occasional rise in blood pressure; (b) prevention of the steep fall in blood pressure and hastening the recovery from the fall produced by intravenous barbiturate injection; (c) stimulation of respiration when stoppage was produced by barbiturates; (d) maintenance of respiration after barbiturate injection even after cardiac stoppage.

In several experiments, it was noticed that the action of small doses of adrenaline and ephedrine was especially marked following picrotoxin in the barbitalized animals. These drugs not only stimulated respiration with a simultaneous rise in blood pressure, but the hemodynamic effects they produced after picrotoxin was more pronounced and more sustained than in controls. In several cases, intravenous injections of adrenaline (a total of 0.1 cc of 1 to 20,000 solution) produced a gradual and sustained rise in blood pressure. These sustained rises with adrenaline and ephedrine may be explained as shifts in blood volume. Adrenaline and ephedrine then may be employed as effective antidotal agents together with picrotoxin in barbiturate poisoning and possibly in other instances of circulatory collapse and respiratory failure.

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SCIENTIFIC APPARATUS AND LABORATORY METHODS

PHYSIOLOGICAL STROBOSCOPE

IN SCIENCE for December 21, 1934, the authors described their physiological stroboscope and its application in the field of the mechanics of phonation. Refinement of the apparatus and the addition of cooperative instrumental units has followed the study of the stroboscope with a model larynx as the subject.

The heart of the instrument is a specially designed amplifier fed by a wide-frequency-range microphonic element. It was a simple step to add to the gaseous discharge lamp in the output a cathode ray oscillograph and a recording device with audio monitor. The chain of instrumental response is currently as follows: A tone originating in the artificial larynx is passed to the microphonic element and into the amplifier, where its electrical component is linearly increased in amplitude; thence into the amplifier's phasing and band pass filter section, where a reduction of the origin tone to its fundamental frequency is achieved and translated in the output into a flashing of the gaseous discharge lamp at a period synchronous with the frequency of the tone emitted by the artificial larynx. Simultaneously, as the lamp held near the vibrating cords of the larynx reveals them in stroboscopic immobility, the fluorescent screen of the

cathode ray oscillograph exhibits a wave analysis and the recording device registers and monitors the tone produced by the cordal configuration under observation.

With this apparatus revealing for the first time the tonal conformation of the cords during frequency transitions, and with a visual representation of the sound thus produced, a relationship is postulated wherein a particular tone is reproducible by the reestablishment of a predictable set of mechanical conditions in the larynx. Thus the isolation of individual factors responsible for vocal characteristics is accomplished.

Further study and interpretation of the apparatus operative with the artificial larynx will precede any report of observations made on clinical cases.

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A METHOD FOR IRRIGATING FUNGUS CULTURES

AN adequate study of smaller fungi must always include careful pure-culture work. Information re-