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On the Visually Evoked Potentials in the Cerebral Visual, Somatosensory, Motor and Association Areas of Monkeys (Macaca Cyclopis)

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Seventeen unanesthetized monkeys (sixteen macaca cyclopis and one macaca mulatta) were immobilized by galamine triiodoethylate (Flaxedil) and masspotentials (EEG and evoked potentials) before and during flashing light stimulation to atropinized eyes were led by grossselectrodes from the cerebral visual (occipital cortex), association (superior and inferior parietal, superior frontal cortex), motor and somatosensory areas. Simultaneously, masspotentials of six loci and one tracing of stimulation signals were recorded onto 1/4 inch magnetic tape. These magnetic records were fed into the computer (ATAC—401 or ATAC—501—20) and summation average evoked potentials and autocorrelograms of background EEG were obtained.

It was found that there were the following three response components in the visually evoked potentials of the cerebral visual area, i.e., (i) sharp or relatively sharp initial biphasic positive-negative deflections (P1 N1, P2 N2, P3 N3, etc.), (ii) late biphasic slow components (p1 n1, p2 n2, p3 n3) and (iii) wavelets superimposed on the sharp or relatively sharp deflections and sometimes on the late slow components. Similar response components were also observed in areas other than the visual area.

Though the above three components were combined with each other in a variety of extents to show considerably complex configurations in the summation average evoked potentials in the visual area, their configurations could be classified into four groups. In the first group, the wavelets were superimposed on the sharp or relatively sharp biphasic deflections which were followed by the late slow components, wherein the first late slow negative deflection n1 was shown by moving average method to be masked in some instances by P3 N3 and P4 N4. In the second group, the wavelets were hardly observed so that the late slow components (p2 n2 and p3 n3) were evident. In the third group, negative deflections N2 and N3 were weak and in the fourth group, the peak latencies of each of the sharp or relatively sharp biphasic deflections, especially those of P3 and N3 were prolonged considerably, but the late biphasic slow components were observed in both of the above groups.
When the wavelets in the visual cortex are superimposed prominently on the sharp or relatively sharp deflections and on the late slow components of the evoked potentials to suggest high activity, not only was the fastwave in the background EEGs predominant, but also a regular oscillation of about 20c/s and irregular oscillations of slower frequencies were observed respectively in the autocorrelograms of the background EEGs in the visual and other cortices.

On the contrary, when the sharp or relatively sharp deflections in the evoked potential of the visual cortex were the least and most prolonged in their peak latencies, with complete absence of wavelets, slow waves were dominant in the background EEGs and slow irregular oscillations were revealed in their autocorrelograms to suggest lowest cerebral activity.

In the instance of intermediate patterns between the above two evoked potentials, various variations and alterations in the above two properties were observed in the EEGs and their autocorrelograms to present considerable complexity.

Three response components similar to those observed in the occipital cortex were also found to be elicited by flashing light stimulus in the superior and inferior parietal, post- and precentral and superior frontal gyri of the cerebral cortex, though their sizes were from one third to one tenth of those in the occipital cortex. The onset and/or peak latencies of their initial positive deflections was almost the same or slightly longer than that observed in the occipital cortex.

INTRODUCTION

In spite of the many studies on the visually evoked potentials in the cerebral cortex of cats (1, 3, 4, 5, 6, 11, 14, 16, 17, 18, 19, 20, 21, etc.), considerably less work is available on monkeys (Hughes 1964; Hughes and Mazurowski 1962; Doty and Kimura 1963; Doty et al. 1963, 1964; Cowey 1964), in which hardly any response configurations have been demonstrated and "rhythmical oscillatory activities" have chiefly been observed. Far fewer observations have been made on the visually evoked potentials in areas other than the visual areas, i.e. the mesial supracallosal area (Hughes123) and a portion of the superior temporal gyrus (Doty et al.9107). Recently, Bignall and Singer (1967)2 have found "polysensory areas" in various "nonprimary areas", wherein evoked potentials were elicited by flash, click and electric shock to skin. However, now observations were made on the response configurations. In this study, therefore, not only were time-patterns of summation average evoked potentials in the cerebral visual, somatosensory and other areas obtained, but also those in the visual area were studied in relation to the background EEG activity.
METHODS

Acute unanesthetized but immobilized experiments were carried out on seventeen adult monkeys (sixteen macaca cyclopis and one (NO.300-25) macaca mulatta) weighing from 2.4 to 5.6 Kg.

The trachea was canulated under ether anesthesia. After being mounted on Johnson's type stereotaxic instrument, an opening as small as possible was made in the skull. After ether had been expelled by artificial respiration (24/minute) with oxygen-mixed air, gallamine triiodoethylate (Flaxedil) was injected intraperitoneally, first a dose of 20 mg/kg and then at a rate of 10-20 mg/kg/hour to immobilize the animal. Silver ball tipped monopolar electrodes of 0.5 mm diameter were placed onto the dura or pia of the occipital, superior and inferior parietal, postcentral, precentral and superior frontal gyri of the cerebral cortices. Mono- and/or bipolar depth electrodes of steel or nicrome wire of 22 and 24 gauge insulated except at their tips were inserted into the lateral geniculate body, mesencephalic reticular formation, centrum medianum, etc. These were confirmed histologically after completion of the experiments by Klüver-Barrera's staining method (1953). The reference silver wire electrode was inserted into the muscle of the nape or the bilateral temporal region.

All wound areas and pressure points were locally anesthetized with 2% procain to remove pain. Both pupils were dilated with 0.5% atropin sulphate. Flashing light stimulation was delivered binocularly at intervals of 0.5, 0.7 or 1.0 f/sec. The flash of about 6000°K daylight, 1 Watt/sec and about 100us duration were from a stroboflash bulb (FT-100, Mazda) attached to a photic stimulator (PS-101, San'ei-Sokki Co.). To diffuse the flashing light, a frosted glass plate, a round iris opening and a cotton white curtain were placed in front of the flash bulb, located 50 cm from the eyes. The diameter of the iris opening used was 100 mm in fifteen monkeys and 30 mm in two monkeys. The relative intensity of the light on a logarithmic scale was 2.6 and 1.7, respectively, when the intensity through the iris of 5 mm diameter was taken as 0.

Ink-records of mass potentials (EEG's and evoked potentials) and stimulation signals were prepared on a 8-channel polygraph (RM-150, Nihon Kohden Co.) or a 8-channel electroencephalograph (EG-803, San'ei Sokki Co.) and, at the same time, recorded on 1/4 inch magnetic tape by the PWM and FM methods, respectively, through a 8-channel data recorder (SPRA-48, Shiroyama Tsushin Co.) while monitoring on four four-beam cathode ray oscilloscopes (VC-6, Nihon Kohden Co.) and a one-beam synchroscope (SS-5022, Iwasaki Tsushin Co.).

The summation averaged response time-patterns (crosscorrelograms of the stimulus and masspotentials) were obtained by feeding the magnetic tape records into the Digital Computer For Data Processing (ATAC-401, Nihon Kohden Co.). Positivity of responses was indicated by a downward deflection.

The autocorrelograms (averaged time-patterns) of the background EEGs of 2.6 seconds length (total delay) were obtained by the Computer For Data Processing (ATAC-501-20, Nihon Kohden Co.) from the magnetic EEG recording of 40 sec length (analysis time).
RESULTS

1) Photically evoked summated potentials in the visual cortex (occipital and lateral striate cortex).

Photically evoked potentials in the cerebral visual area (lateral striate cortex) of monkey were obtained over a time length of 500 msec after flash stimulus by thirty times summation (Fig. 1.1). Though the evoked potentials were considerably complex and highly variable in their configurations from animal to animal, the following three response components were observed in the evoked potentials: (i) Along with the primary positive deflection ($P_1$) with a peak latency of 25–60 msec, three to five sharp or relatively sharp initial biphasic components of 30–60 msec duration were observed until 150–250 msec after delivery of the flash ($P_1 N_1$, $P_2 N_2$, $P_3 N_3$, etc.). This was followed by (ii) two or three slow biphasic potentials ($p_1 n_1$, $p_2 n_2$, $p_3 n_3$) of about 150 msec duration which had longer peak latencies. In addition, (iii) about ten or fewer small, sharp deflection with a shorter duration of 10–25 msec were brought out which formed a time-pattern of wavelet superimposed on responses (i) and/or (ii) (Figs. 1.1 and 1.2). The peak latencies of every response components except those in the wavelets (iii) were more consistent than their amplitudes. The peak latencies of $P_1 N_1$ and $P_2 N_2$ were relatively consistent, whereas that of $P_3 N_3$ was inconsistent and those of the late slow components $p_1 n_1$, $p_2 n_2$ and $p_3 n_3$ were more variable. It was found that the slow biphasic potentials (ii) were masked in some instances by the initial biphasic potentials (i) since potential (ii) could be separated by taking the moving average over the length of response (i) (Fig. 1.3). Varying degrees of mixture of these three components were contained in each photically evoked potential of the seventeen monkeys to present complex configurations, which could be classified into the following four groups.

Group I. (Fig. 1.1; 300–24, -28, -29 and -30). The wavelets were superimposed on each of the sharp or relatively sharp initial biphasic components, $P_1 N_1$, $P_2 N_2$, etc. Further, in one case they were sustained even longer to be superimposed on the late slow component $p_2 n_2$ (Fig. 1.1, 300–24), and $P_1 N_1$ group, $P_2 N_2$ group, etc. were observed. In addition, $P_4 N_4$ and $P_5 N_5$ were often brought out following $P_3 N_3$, and the late biphasic slow components were also elicited. In this group, the earlier $n_1$ component preceding the $n_2$ component was masked by the $P_3 N_3$ and $P_4 N_4$ responses (Fig. 1.3). Group II. (Fig. 1.1, 300–16, -14). Almost none of the wavelets (iii) were brought out but the sharp or relatively sharp initial biphasic components (i) and the slow biphasic potentials (ii) were manifested consistently. Group III. (Fig. 1.1, 300–15, -13, -11). Though the components (i) and (ii) were elicited, $P_2 N_2$ and $P_3 N_3$ in the former components were not intense. Group IV. (Fig. 1.1, 300–9, -10, -12, -26 I, -25, -23 and -22). The peak latencies of the response components in the sharp or relatively sharp initial biphasic components (i) were longer than those in other groups, especially that of the response $P_3 N_3$ was considerably prolonged, and the components (ii) were also elicited.

2) Photically evoked averaged potentials in relation to the background EEGs
Fig. 1.1. Average photically evoked potentials in the cerebral visual cortex of mon-
Black circles in monkey brains indicate loci of lead electrodes. Each tracing of average evoked potentials was obtained by summations of thirty times.

Fig. 1.2. Peak latencies of each component in the photically evoked potentials in the cerebral visual cortex of sixteen monkeys.

Numerals on the left are numbers of monkeys. Abscissa: time of peak latency. Black and white circles: negative and positive deflections in the evoked potentials in Fig. 1.1 respectively. Shown from top to bottom are, group I (No. 24, 28, 29 and 30), group II (No. 16 and 14), group III (No. 15, 13, 11) and group IV (No. 9, 10, 12, 26-I, 25, 23 and 22). Broken and chain lines show respectively the dominant negative peaks and predominant positive peaks in each component.

Fig. 1.3. Slow response components masked by the sharp components.

By taking the moving average of the photically evoked potential (full line) twice, the sharp or relatively sharp responses (N₁, P₂N₂ and P₃N₃) were eliminated and two successive slow diphasic responses emerge. The former and latter were masked respectively by responses N₁, P₂N₂ and P₃N₃.
Fig. 2.1-1. Background EEGs of Group I visually evoked potentials in the visual cortex.

and their autocorrelograms.

As the above variability in the averaged evoked potentials seemed to be related more or less to background cerebral activity, background EEGs and their autocorrelograms were observed in relation to the evoked potentials in the visual cortex. In the animals showing Group I average evoked potentials (Fig. 1.1, 300-24, -28, -29 and -30), fast waves of 15–21 c/s were seen in the background EEGs of the visual areas, from which the evoked potentials also were obtained. In some animals, these waves of varying amplitude were superimposed on the slower EEG waves of 5–8 c/s of the visual and other areas, including the superior frontal (SF), pre-(Pr.C, motor area) and postcentral areas (Po.C, somatosensory area), and subcortical sites (optic radiation OR, lateral geniculate body GL, superior colliculus CS, midbrain reticular formation RET.MES). (Fig. 2.1–1).

In the instance of considerable wavelets in the average evoked potential (Fig. 1.1, 300–24), the fast wave was predominant in the

![Graph showing autocorrelograms of background EEGs](image)

**Fig. 2.1–2.** An example of autocorrelograms of background EEGs.

Analysis time 40 sec. Maximum delay 2.6 sec (see left top). Each autocorrelogram of EEG was calibrated by the height of pulse at the time origin of the autocorrelogram of pulse series over the same length of time as the EEG. Slow wave of 0.4 c/s is the artifact caused by artificial respiration. A regular oscillation of about 20 c/s was superimposed on a slow wave of about 2.2 c/s in the left occipital cortex (1–0 and bottom), while irregular oscillations were seen in other areas. This animal (300–24) showed the most prominent wavelets in the visually evoked potential of the occipital cortex (Fig. 1.1) and its background EEGs (Fig. 2.1–1) showed fast wave patterns of 20–21 c/s in the visual cortex (1–0), while low voltage fast waves of 24–25 c/s and fast waves were superimposed on slow waves of 5–8 c/s in subcortical sites (rOR, CS). The bottom autocorrelogram was recorded by the Eletronic Polyrecorcer (EPR-10A, Toa Elect. Ltd.). Calibration: 500μV.
background EEG (Fig. 2.1-1, l-O), and its autocorrelogram (Fig. 2.1-2) showed a regular oscillation of about 20 c/s frequency superimposed on a slow oscillation of about 2.2 c/s. On the other hand, in the case where the wavelets (iii) were entirely absent and in which the evoked sharp or relatively sharp responses (ii) were the weakest with the most prolonged peak latencies in the visual area (Fig. 1.1, 300-10), the background EEG in this area showed a pattern of irregular slow waves of about 1.4, 2.5 and 3 c/s and small waves of about 9 and 12.5 c/s (Fig. 2.2, left). The former waves were also seen in other cerebral areas (r-SP, l-SP), whereas slower fluctuations of about 1 c/s were observed in subcortical sites (VPM and RET. MES). In their autocorrelograms (Fig. 2.2,

right), the above irregular slow waves were seen in the occipital and superior parietal cortices while a slow wave of about 1c/s was noted in the subcortical sites. In the intermediate type between the above two average evoked potentials, a considerable variety of background EEG patterns were observed. In a representative animal (Fig. 1.1, 300-14), waves of about 6-7 c/s were manifested in the background EEGs of the oc-
Fig. 2.3. Background EEGs and their autocorrelograms of an animal (300-14) showing visually evoked potentials of Group II.

Background EEGs showed waves of 6–7 c/s in the cerebral cortex (Po.C, SP, O) and subcortical sites (OR, VPL), and further spindle-like waves of 12–14 c/s were seen in the somatosensory area (rPo.C). In their autocorrelograms regular oscillation of 6.6 c/s was manifested in all regions. See text. VPL: n. ventralis posterolateralis.

capital area (O), optic radiation adjacent to the lateral geniculate body (OR), nucleus ventralis posterolateralis (VPL), or nucleus paracentralis (Pc) while waves of 12–14 c/s were interposed in the above waves in the EEG of the postcentral area (Po.C.) or superimposed on that of the superior parietal area (SP) (Fig. 2.3, left). In their autocorrelograms (Fig. 2.3, right), a mixed pattern of about 3.3 and 6.6 c/s was exhibited in the occipital areas (O), and another wave of about 12 c/s was intermingled with these slow waves in the superior parietal areas (SP), whereas a regular wave of about 6.6 c/s was manifested in the subcortical sites (OR and VPL).

In the background EEGs of various areas of such monkeys that exhibited visually evoked potentials of Group III (Fig. 1.1, 300–13), irregular slow waves of 5–6 c/s were superimposed on large slow waves of 1–1.5 c/s (Fig. 2.4, left). Their autocorrelograms, on the average, demonstrated the above pattern (Fig. 2.4, right). On the contrary, the background EEGs (Fig. 2.5, left) which demonstrated visually evoked potentials of Group IV (Fig. 1.1, 300–25–15) may be a slow wave pattern of about 2 c/s, since the waves of about 10–30 c/s superimposed on the slow wave in the EEGs did not appear and only
Fig. 2.4. Background EEGs and their autocorrelograms of an animal (300-13) showing visually evoked potentials of Group III.

Irregular slow waves of 5–6 c/s were superimposed on large and slow waves of 1–1.5 c/s in background EEGs of all regions and irregular patterns were seen in their autocorrelograms. VPI: n. ventralis posteroinferior. THIP: tractus habenulo-interpeduncularis. b.p.: bipolar lead.

Fig. 2.5. Background EEGs and their autocorrelograms of an animal (300-25) showing visually evoked potentials of Group IV.

Background EEGs of cortical (O, SP, SF) and subcortical sites (CI, etc.) showed predominant large waves of 2c/s, upon which waves of 10–30 c/s were superimposed. In their autocorrelograms, regular waves of 2.2 c/s were manifested in rSF, ISP and ISF and irregular slow waves were depicted in other regions. The wave of 0.4 c/s (2.5 sec period) is the artifact caused by artificial respiration.
the wave of 2 c/s was seen in the autocorrelograms of the EEGs (Fig. 2.5, right).

3) Evoked potentials elicited by flash to the cerebral somatosensory, motor and association areas.

Although no sign was manifested in the electroencephalographic ink records during stimulation, summation averaging with the computer revealed evoked potentials in response to binocular single flash stimulus not only in the superior and inferior parietal cortex (SP, IP), which are considered as the association areas, but also even in the more rostral portion (superior frontal cortex, SF) (Figs. 3.1, 2,3,4,5,6). On

![Diagram](image)

Fig. 3.1. EEGs and photically evoked potentials in the motor cortex.

Though no significant deflection time-locked to the signals (stim.) of binocular flash stimulation of 1 f/sec was observed in EEGs of the motor cortices (r-Pr.C and i-Pr.C), clear cut average evoked potentials were manifested by summation of thirty times. CI: capsula interna. CC: corpus callosum.

the right of Fig. 3.1 are shown six evoked potentials successively obtained by summation of thirty responses in the motor cortex (Pr.C). Within 100 msec after the flash stimulus, two sharp positive-negative deflections followed by a deep positive swing were manifested. This was a fixed pattern with a slight variation in their magnitudes, whereas later responses were more variable both in their pattern and size. By summation of 180 responses, however, the later responses were converted to a relatively fixed pattern (Fig. 3.2). Initial sharp components (i(i) P1N1, P2N2 and P3) and late slow ones were found in the motor cortex. As depicted in Fig. 3.3, not only the late slow response, but also the wavelets composed of repetitive biphasic small deflections of
Six average evoked potentials, each of which were brought out in bilateral motor cortices by left and right monocular and binocular flashing light stimulation, were traced used the same base line. Top and bottom: thirty and one hundred eighty summations, respectively. Calibration of the bottom traces is two-fold of that of top traces.

The evoked potential in the visual cortex is of Group I and high wavelets (Fig. 1.1, 300–24). About 25 msec duration were also found superimposed on the slow response in the superior parietal (SP) and the motor cortex (Pr.C). These responses due to binocular stimulus were more prominent than those due to monocular stimulus as observed in the occipital cortex. Hence, it was found that the same three kinds of photically evoked potentials as those verified in the occipital cortex are elicited in the somatosensory, motor and association cortices, though less in size,
The duration of each component in the above three kinds of evoked responses was similar to that of each corresponding component in the evoked responses in the occipital cortex. Every one of the above three responses were brought out in varying extent to manifest a variety of patterns different from that of the response in the occipital cortex (Figs. 3.4, 5). However, they were less in their magnitudes being from one third to one tenth in amplitude and show considerably greater fluctuations. The onset and/or peak latencies of the initial positive deflection (P1) was almost the same or slightly longer than that observed in the occipital cortex (Figs. 3.4, 5). Though the above observations

Fig. 3.4. Visually evoked potentials in the motor, somatosensory, and superior parietal cortex in comparison with that in the visual cortex.
Fig. 3.5. An example of the visually evoked potentials in various cerebral cortices in comparison with that in the visual cortex.

Initial positive response (P1) was verified in every cortices. Their peak latencies were similar to or slightly longer than that of visual cortex.

were obtained by monopolar lead, summation average responses were also obtained by bipolar lead, wherein the paired electrodes were placed 3mm apart and parallel to the parieto-occipital sulcus. These also indicated that similar initial sharp components, which were composed of 3–5 biphasic deflections with durations of about 30 msec and onset latency of about 50 msec, and the two (late) slow components with peak latencies of about 130 and 260 msec and duration of about 130 and 240 msec respectively were also elicited. These slow components corresponded to the n₁ and n₂ components manifested by monopolar lead. These responses were distinct especially in the superior parietal cortex (SP, association cortex), and also were exhibited in the somatosensory (Po.C), motor (Pr.C) and superior frontal cortices (SF) (Fig. 3.6). Hence, the responses manifested in areas other than the occipital cortex were not induced by volume conduction of the responses in the occipital cortex but had been elicited and this suggests some cooperative integration of visual inflows in these areas.
DISCUSSION

Summation average response in the visual cortex and background EEG activities.

Though photically evoked cerebral potentials in monkey were complicated in their configuration (Fig. 1.1), as pointed out by Doty et al. and Bignall and Singer, the author was able to classify them into four groups, each of which seemed to be derived from a characteristic combination of three response components.

These three components were the first sharp or relatively sharp initial biphasic components (P1N1, P2N2, P3N3, etc.), the late biphasic slow components (p1n1, p2n2 and p3n3) and the wavelets. Wavelets occurring every 5-7 msec in the early portion and every 10-14 msec in the later portion of the evoked potential have been demonstrated by Hughes and Mazurowski. In addition, "the prominent rhythmical activity" of 5-30 msec duration noted by Hughes and "the oscillatory response" of 160-210/sec frequencies of Doty and Kimura and Doty et al. may be the same as the wavelets of the author, since they were brought out more markedly during higher brain activity and superimposed on the slower deflection with approximately the same duration as those of the wavelets of the author.

The more slowly developing positivity with peak latency of about 25-40 msec lasting about 50 msec and "the major response" with posi-
tive peak latencies of about 50–90, 120–140 msec and about 40–80 msec duration of Doty and Kimura\(^8\) and Doty et al.\(^{10}\) may correspond to the sharp or relatively sharp initial biphasic components. The more slow negativity noted by Hughes\(^{12}\) seemed to be the first negative deflection \(N_1\) in the above initial biphasic sharp component.

Though Hughes and Doty et al. have done on a study mainly on the wavelets, they have not mentioned the late slow components found in this paper.

*Photically evoked potential in the cerebral somatosensory, motor and association areas.*

Though their amplitudes were smaller (1/3–1/10), three types of response components similar to the three types of evoked potentials in the visual cortex, were observed by applying summation average method in the pre- (motor) and postcentral (somatosensory), superior and inferior parietal and superior frontal cortices. As these potentials were manifested not only monopolarly, but also bipolarly, volume conduction of the evoked potentials in the visual cortex can be ruled out.

Though Cowey (1964)\(^7\) and Doty et al. (1963\(^9\), 1964\(^{10}\)), have observed that a higher surface positive response and a lower response, respectively, are elicited usually about 10 msec later in the prestriate cortex than in the visual cortex, few observations have been carried out on visually evoked responses in other cerebral areas than the visual cortices. In the superior temporal area, large but considerably variable surface negative response only was demonstrated by Doty et al. (1963\(^9\), 1964\(^{10}\)).

Though Bignal and Singer (1967)\(^2\) have recently observed a visually evoked potential in the parietal, motor, prefrontal and superior temporal areas, almost all of these visually evoked potentials hitherto demonstrated by the above authors, however, were only the initially elicited potential, which presumably corresponds to the one in the initial sharp potential verified in this paper. As not only the configurations of the responers components but the onset and/or peak latencies of the initial positive deflection (\(P_1\)) were similar to those obtained in the visual cortex, there probably is direct projection to the cerebral somatosensory, motor and association cortices. Of course this projection would be through fewer afferent fibers than that to the cerebral visual cortex, because the response amplitudes in the former areas were considerably less than those in the latter.

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REFERENCES