

Nihon Doubutsu, 3 weeks old at the time of the operation for electrode implantation, were used. The animals were anesthetized with sodium pentobarbital (40 mg/kg, i.p.), and stainless steel electrodes (200 μ m in diameter) insulated for a small loop formed at the tip to avoid any injury to the dura and cortex, were placed gently on the dura.

The electrodes were implanted on the right frontal cortex (area 3), the left frontal cortex (area 3), the right occipital cortex (area 17) and the left occipital cortex (area 17). The reference electrode was implanted in sagittal line 7 mm anterior to the bregma (figure 1). The first EEG recording was made 3 days after implantation. During EEG recordings, rats remained free in a transparent cage.

The socket on the rat's head was connected to a electroencephalography by a flexible cable which did not hamper the rat's movement. Monopolar EEG were recorded from the frontal cortex and the occipital cortex in quiet wakefulness for 15 min by electroencephalography (Nihon Kodan ME-92D), and EEG from the left occipital cortex was analyzed by EEG frequency analyzer (Sanei-Sokki, 7p-11). And spectral energy distribution of each frequency band (δ , θ , α , β) was computed.

Results and discussion. The development of EEG in rats from 4 weeks old to 16 weeks old is shown in figure 1. In visual analysis, EEG consisted mainly of 5–6 Hz mixed with 3–4 Hz high voltage slow waves at 4–10 weeks. After 12 weeks, 8–9 Hz alpha waves occurred intermittently and were slowly increasing. The amplitude varied from 200 to 300 μ V at 4–5 weeks and became lower in older ages. After 8 weeks, they were around 100–200 μ V.

Spectral energy distribution analysis is shown in figure 2. Frequency components are divided into 4 bands, that is, delta (2–4 Hz), theta (4–8 Hz), alpha (8–13 Hz) and beta (13–30 Hz). Theta band had the highest power at 5–16 weeks. Especially after 11 weeks, theta band increased step by step and presented significantly higher peak than that of a 4-week-old. Delta band of 4-week-old rats occupied a large part among 4 frequency bands, but after 6 weeks it

was markedly decreased and theta, alpha bands were increasing. Beta bands were almost invariably exclusive of slightly significant peaks at 8–9 weeks.

Yoshii et al.² studied the EEG development of infant rats (2nd–21th day) acutely and reported that EEG of 21-day-old rats attained that of adults. Gramsbergen's³ report said that after the 18th day no further changes occurred in the power spectra during various sleep stages. Yoshii et al. made the experiment on acutely implanted rats. Gramsbergen analyzed the EEG in rats of only 9–30 days in various sleep stages.

We analyzed EEG in the wakeful state of a rat from 4th to 16th week. So our results were different from their results. In our results EEG of 4-week-old rats, slow components were remarkable, and this slow component developed into medium-fast component progressively with age until 16 weeks. Overholser et al.⁴ reported that the EEG of young rats of 4–8 weeks age showed the average frequency to be 30.8/sec. Timo-Iaria⁵ reported that ECoG recorded from all areas were generally in the range of 30–40 Hz. These results are much higher than those in our experiment. Their EEG seems to mingle with electromyogram. Our data of energy power spectra in the 4th week are in accord with that of Deza's 28th day EEG⁶. It is interesting to note that fast component (13–30 Hz) was not altered comparably for different ages.

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Formation of glomerulus-like structures by the olfactory nerve after neonatal bulbectomy

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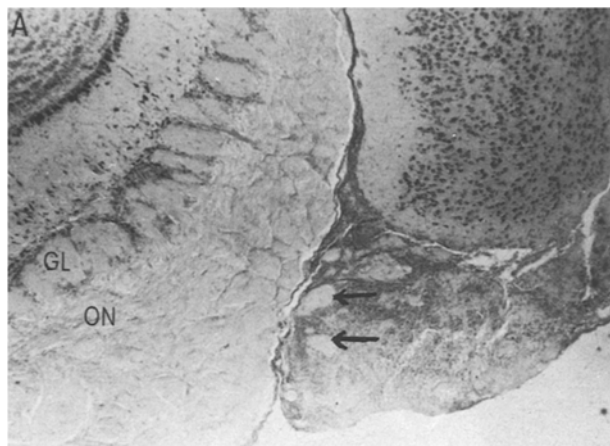
Summary. Evidence presented here suggests that glomerulus formation within the olfactory bulb of the rat, which is mostly a postnatal event, is directed by the olfactory nerve rather than by the influences of mitral and tufted cells.

The glomeruli (GL) of the olfactory bulb (OB) are round structures surrounded by short axon cells called periglomerular cells (PGLC). These structures contain the terminal dendritic branchings of the tufted and mitral cells (the 2nd order olfactory neurons), the axon terminals of the olfactory nerve (ON) and axons and dendrites of the PGLC². Formation of these structures is mostly postnatal and their average number shows a 10-fold increase from the 1st day after birth to 2 months of age³. Their average diameter also increases 2-fold during this period³. The underlying mechanisms responsible for the formation of these structures is as yet unknown. Altman⁴ has shown that cells originating in the proliferating zone of the subependymal layer around the lateral ventricle migrate to the OB, some of which surround the GL. It has been shown that the terminal branchings of the mitral and tufted cells are absent at birth and form postnatally⁵. It is therefore conceivable that the postnatal growth of these branchings

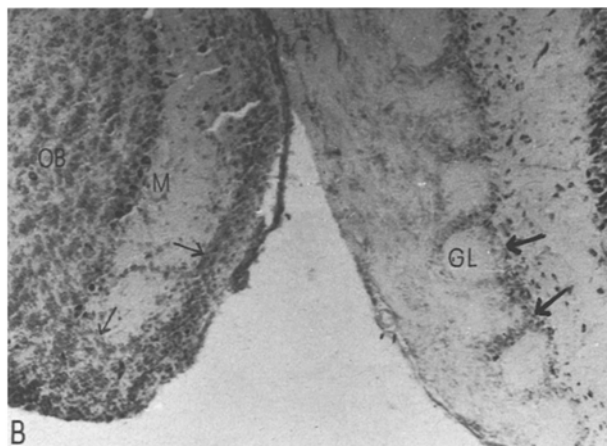
induces the formation of the GL. In this study, however, we report results that indicate that, under experimental conditions where mitral and tufted cells are absent due to bulbectomy but the regenerated ON is present, glomerulus-like structure can be formed.

Albino rats were monolaterally bulbectomized at 3–4 days of age under ether anesthesia. At 25 and 60 days of age, their brains were removed and fixed with Bouin fixative. Paraffin sections were cut with a thickness of 14 μ m and were stained with thionin for light microscopic studies.

Our findings reported here were essentially the same for both age groups. In most brains, the hemisphere ipsilateral to the lesion protruded and made contact with the regenerated ON. (The regenerative capacity of the ON has been well documented⁶). Light microscopic examination revealed that in specimens where the OB, the accessory olfactory bulb and the anterior olfactory nucleus were all removed, the plexus formed by the regenerated ON on the



A On the right, the regenerated nerve has formed glomerulus-like structures (arrows) underneath the protruded cortex in the absence of the olfactory bulb and retrobulbar structures. On the left, the intact olfactory (OB) bulb with normal glomeruli (GL) and the plexus formed by the olfactory nerve (ON) can be seen.



B Coronal section through the olfactory bulb. On the right, the control bulb (OB) is seen with glomeruli GL surrounded by periglomerular cells (arrows). On the left, a portion of the olfactory bulb (OB) which was spared from the lesion is seen to contain mitral (M) and periglomerular cells (arrows), but the glomeruli have not been formed.

ventral side of the protruded cortex, contained glomerulus-like structures (figure A). Inspection of serial sections revealed that the cells surrounding these structures formed a cell stream along the known migratory path of the postnatal bulbar cells, reaching the subependymal layer of the lateral ventricle. In 1 brain, glomerulus-like structures were seen to penetrate the brain along this path. In a few brains, however, the plexus formed by the ON was surrounded by cells without forming any glomerulus-like structures. In specimens where a portion of the anterior olfactory nucleus was spared, these glomerulus-like structures were seen to lie within the peripheral parts of this structure. In cases where a portion of the OB was spared but no indication of ON regeneration could be discerned by the light microscope, mitral and periglomerular cells occupied their appropriate position without forming any of these glomerulus-like structures (figure B). Although similar to the GL, these glomerulus-like structures form round bodies within the plexus of the ON and are surrounded by cells which appear to have the same origin as the PGLC, unlike the GL they lack the dendritic contribution from mitral and tufted cells, besides the synaptic organization within them is, as yet, unknown. Therefore, we prefer to call them glomerulus-like structures rather than referring to them as proper glomeruli.

In summary, findings described here show that glomerulus-like structures can be formed in the absence of mitral and

tufted cells as long as the regenerated ON is present. On the other hand, these structures do not form in the absence of the ON, even when mitral and tufted cells are present. Thus, although no definite conclusion can be reached about the mechanism of glomerular formation during normal growth, findings presented here indicate that formation of these postnatal structures are induced by the ON and are independent of mitral and tufted cells. Further, since the regenerated ON can form structures in the absence of its natural target, i.e., the OB, this system can serve as an interesting mammalian model for the study of neural plasticity and factors that guide the axons to their targets.

- 1 I am grateful to Miss Sekineh Hamed and Mr Davood Ahmadi for their assistance and to Dr Esmail Meisami for critical discussion of the topic. Reprint requests should be addressed to: 140 Flora Avenue, Apt. 133, Walnut Creek (California 94595, USA).
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Topographical distribution of ATP in rat brain

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Summary. Studies on the distribution of ATP in microdissected segments of the rat brain indicate that the nucleotide is concentrated in gray matter, and especially in the thalamus, hypothalamus, hippocampus, entorhinal cortex and sensorimotor cortex. These distribution studies in conjunction with previous neuropharmacological studies, support the concept that adenine nucleotides may function as intercellular mediators in various regions of the brain.

Adenosine triphosphate (ATP) has been proposed as a candidate for a neurotransmitter role in the peripheral nervous system¹, raising the possibility that it may also be a transmitter in the central nervous system (CNS). Previous

reports²⁻⁴ have described a powerful depressant action of adenine derivatives on the discharges of cerebral and cerebellar cortical neurones. This effect appears to result from the activation of an extracellularly located purine