P-0119

Genetical and Pathological Studies on the Mutant Mice as an Animal Model for Deafness Disease

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A new neurological mutant has been found in the ICR outbred strain mouse. Affected mice display profound deafness and a head-tossing and bidirectional circling behavior, showing an autosomal recessive mode of inheritance. It was, therefore, named cir/Kr with the gene symbol cir. The auditory tests identified clearly the hearing loss of the cir mice when compared to wild type mice. Pathological studies confirmed the developmental defects in the middle ear, cochlea, cochlear nerve, and semicircular canal areas, which were correlated to the abnormal behavior observed in the cir mice. Thus, cir mice may be useful as a model for studying inner ear abnormalities and deafness. We have constructed a genetic linkage map by positioning 14 microsatellite markers across the (cir) region and intraspecific backcross between cir and C57BL/6J mice. The cir mouse harbors an autosomal recessive mutation on mouse chromosome 9. The cir gene was mapped to a region between D9Mit116 and D9Mit38. Estimated distances between cir and D9Mit116, and between cir and D9Mit38 are 0.7 and 0.2 cM, respectively. The gene in order was defines: centromere-D9Mit182-D9Mit51/D9Mit79/D9Mit310-D9Mit212/D9Mit184-D9Mit116-cir-D9Mit38-D9Mit20-D9Mit243-D9Mit16-D9Mit55/D9Mit125- D9Mit281. The mouse map location of the cir locus appears to be in a region homologous to human 3q21. Our present date suggest that the nearest flanking marker D9Mit38 provides a useful anchor for the isolation of the cir gene in a yeast artificial chromosome contig.

(Key words) Neurological Mutant Mice, cir/Kr, Autosomal Recessive Mode, Microsatellite Markers.