

[P-34]**Mutational profiling of p53 gene in human breast epithelial cells (MCF10A) exposed to Benzo(a)pyrene(BaP) or N-Methyl-N'-nitro-N-nitrosoguanidine(MNNG)**

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Mutations of p53 tumor suppressor gene have been associated with exposure to carcinogens. Cultured human breast epithelial cells (MCF10A) were treated with Benzo(a)pyrene(BaP) or N-Methyl-N'-nitro-N-nitrosoguanidine(MNNG). MCF10A cells were grown in DMEM/F12 medium and treated for 6, 12, 24, 48 and 96h with BaP (1, 10 and 100 μ M) and MNNG (0.25, 2.5 and 25 μ M) dissolved in dimethylsulfoxide(DMSO). DNA was isolated from MCF10A treated with carcinogens and exons 5-9 of p53 gene were amplified with polymerase chain reaction. The PCR products of p53 gene were analyzed for mutations with direct sequencing. Point mutations induced BaP were higher in codon 129/exon 5, 225/exon 7 and codons 263, 282, 286/exon 8 than any other site. MNNG also induced various point mutations. The point mutations caused from MNNG were mainly at codon 129/exon 5. The p53 mutations were not affected by exposure dose and incubation time, but highly site-specific. The mutational profile of p53 in human normal cells can help identify particular carcinogens and define the biochemical mechanisms responsible for the genetic lesion in DNA that cause human cancer.