Lawrence Berkeley National Laboratory

Recent Work

Title

7H-Benzo[c]fluorene DNA adduct formation in different human cells in culture

Permalink

https://escholarship.org/uc/item/081367n4

Journal

Proceedings American Association for Cancer Research Annual Meeting, 43

Authors

Goth-Goldstein, R. Russell, M. Parimoo, B. et al.

Publication Date

2002

LBNL-50479 ABSTRACT

7H-Benzo[c]fluorene DNA adduct formation in different human cells in culture

Goth-Goldstein, R., Russell, M., Parimoo B., and Weyand, E.H. Lawrence Berkeley Natl. Laboratory, Env. Energy Technology Div., Berkeley, CA 94720 and Rutgers-The State University of New Jersey, College of Pharmacy, Piscataway, NJ

7*H*-Benzo[c]fluorene (B[c]F) has been known for a long time as a component of complex mixtures such as coal tar or cigarette smoke. B[c]F has been identified recently as a potent lung tumorigen and a major DNA adduct-forming component of coal tar. investigated if human cells have the ability to form B[c]F:DNA adducts as detected in lungs of mice treated with B[c]F. MCF7 (human breast cancer), HepG2 (hepatoma) and Caco-2 (colon adenocarcinoma) cells were treated with increasing concentrations (0.2 - 10 μ g/ml) of B[c]F for 20 hours. Adduct formation was evaluated using ³²P-postlabeling. A dose response in DNA adduct formation was detected in all three cell lines. In MCF7 and HepG2 cells, two adducts were detected, one of them corresponded to an adduct observed in the lungs of mice treated with B[c]F. This adduct is derived from 3-hydroxy B[c]F while the second, slower migrating adduct, appears to be unique to human cells. In contrast, Caco-2 cells formed at least four adducts. Two of the three most predominant adducts correspond to the two adducts observed in MCF7 and HepG2 cells while the additional predominate and a minor adduct are derived from 3,4-dihydrodiol B[c]F. The adducts derived from 3,4-dihydrodiol B[c]F are similar to those observed in mouse lung and skin. The detection of B[c]F:DNA adducts clearly demonstrates that human cells have the capacity to metabolically activate B[c]F to derivatives that covalently modify DNA. Similarities in the types of B[c]F:DNA adducts detected also demonstrates that B[c]F activation is similar in both human cells and mouse tissue.

This research was supported by funds from EPRI.