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ACCEPTED MANUSCRIPT

Polymerase Bypass of N7-Guanine Monoadducts of Cisplatin, Diepoxybutane, and Epichlorohydrin

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Polymerase Bypass of N7-Guanine Monoadducts of Cisplatin, Diepoxybutane, and Epichlorohydrin

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Highlights

- DNA adducts of cisplatin, diepoxybutane, and epichlorohydrin have relevance in cancer therapy and carcinogenesis.
- We investigated the integrity of DNA synthesis from templates containing N7-guanine monoadducts of these agents.
- Two bacterial polymerases and recombinant human polymerase β successfully bypassed the lesions, although with reduced rate of DNA synthesis.
- The polymerases did not show significant misincorporation near the lesion.
- The N7-guanine monoadducts, although the principal reaction products of these agents, do not appear to contribute significantly to the mutational spectra of cisplatin, DEB, and ECH.

ABSTRACT

DNA oligonucleotides containing site-specific N7-guanine monoadducts of cisplatin, diepoxybutane, and epichlorohydrin were used as templates for DNA synthesis by two bacterial

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