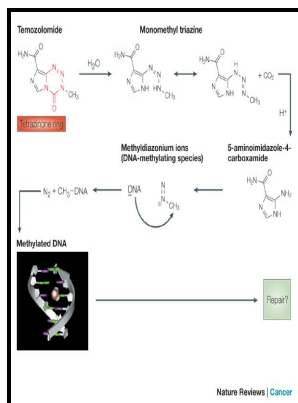


# Antitumour imidazotetrazines - probes for the major groove of DNA.

Aston University. Department of Pharmaceutical Sciences - Antitumor imidazo(5,1



Description: -

-Antitumour imidazotetrazines - probes for the major groove of DNA.

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## Antitumor imidazo(5,1

O 6 - 4-bromophenyl guanine improves the therapeutic index of temozolomide against A375M melanoma xenografts. It is well recognized that nucleic acids are the main target of both bleomycin and actinomycin.

## Antitumor imidazotetrazines. 38. New 8

Interestingly, at many GpT steps the relative extent of cleavage is significantly enhanced in the presence of actinomycin. Comparison of surface areas for the native and complexed B-DNA generated from d TCGCGA 2 duplexes showed only a small reduction in exposed DNA surface upon binding of either MTA or MSK, suggesting the absence of large helical rearrangements.

## Antitumor imidazotetrazines. 38. New 8

Binding of actinomycin to DNA involves insertion of the phenoxazone chromophore between consecutive base pairs coupled with fitting of its two bulky peptide rings into the minor groove of the double helix .,

## DNA and its associated processes as targets for cancer therapy

The mechanism of action of the mustards begins with the formation of an electrophilic aziridinium ion by displacement of chloride. Comparison of the susceptibility of the 160mer, 178mer and 265mer fragments to cleavage by bleomycin and deglycobleomycin in the absence and presence of actinomycin D 5 µM. O6-alkylguanine-DNA alkyltransferase in cutaneous T-cell lymphoma: implications for treatment with alkylating agents.

## Antitumor imidazotetrazines. 38. New 8

Leal JFM, Martinez-Diez M, Garcia-Hernandez V, Moneo V, Domingo A, Bueren-Calabuig JA, Negri A, Gago F, Guillen-Navarro MJ, Aviles P, Cuevas C, Garcia-Fernandez LF, Galmarini CM.



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