

Clinical Pharmacokinetics of Ibuprofen

The First 30 Years

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Summary

Ibuprofen is a chiral nonsteroidal anti-inflammatory drug (NSAID) of the 2-arylpropionic acid (2-APA) class. A common structural feature of 2-APA NSAIDs is a sp^3 -hybridised tetrahedral chiral carbon atom within the propionic acid side chain moiety with the *S*-(+)-enantiomer possessing most of the beneficial anti-inflammatory activity. Ibuprofen demonstrates marked stereoselectivity in its pharmacokinetics. Substantial unidirectional inversion of the *R*-(-) to the *S*-(+) enantiomer occurs and thus, data generated using nonstereospecific assays may not be extrapolated to explain the disposition of the individual enantiomers.

The absorption of ibuprofen is rapid and complete when given orally. The area under the plasma concentration-time curve (AUC) of ibuprofen is dose-dependent. Ibuprofen binds extensively, in a concentration-dependent manner, to plasma albumin. At doses greater than 600mg there is an increase in the unbound fraction of the drug, leading to an increased clearance of ibuprofen and a reduced AUC of the total drug. Substantial concentrations of ibuprofen are attained in synovial fluid, which is a proposed site of action for nonsteroidal anti-inflammatory drugs.

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