Crystal Structure of a Disintegrin Heterodimer from *Echis carinatus* at 1.9 Å resolution

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Disintegrins are a family of small proteins that bind to integrins specifically. Their binding site is characterized by the presence of Arg-Gly-Asp motif which indicates an RGD-dependant mode of interaction with integrins. The disintegrins interfere with the functions of integrins as antagonists. Disintegrin was isolated from the venom of Echis carinatus and crystallized in the tetragonal space group P4₃2₁2 with a=b=90.7Å and c=55.5Å. It exits as a heterodimer unlike the low resolution structure which existed as a homodimer with its two subunits related by a two fold crystallographic symmetry. It is interlinked by two disulfide bonds at the N-terminal region and contains 64 amino acid residues in each chain. Each monomer contains three pairs of six antiparallel β-strands and is stabilized by four disulphide bridges. It has been refined to an R-factor of 0.212 and R_{free} of 0.251 for all the data. The two chains of the dimer are anchored at N-terminal but diverge away at their C-termini exposing the Arg-Gly-Asp motif onto opposite directions, thus enhancing their binding efficiency. This is one of its unique features. The structural studies of disintegrins can provide a useful framework for the design of potent antagonists of integrins.

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