

EMBO

EUROPEAN MOLECULAR BIOLOGY ORGANIZATION

FIFTH EMBO ANNUAL SYMPOSIUM

NUCLEIC ACID-PROTEIN INTERACTIONS

12-15 October, 1979

Programme and Abstracts for the Poster Sessions

12, 13, 14, and 15 October

RELATIONSHIP BETWEEN PROTEIN AND NUCLEIC ACID SEQUENCES IMPLICATED IN SPECIFIC BINDING INTERACTIONS

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Arguments are summarised which support the existence of a certain correspondence (code) between the protein and nucleic acid sequences implicated in specific binding interactions. Recognition protein sites are assumed to contain pairs of antiparallel polypeptide chain segments which form a right-hand twisted antiparallel β -sheet, upon complex formation between a protein and DNA the sheet undergoes a structural transition allowing for the backbone amide groups in the two polypeptide chain segments to form hydrogen bonds with DNA base pairs. The binding is stereospecific in the sense that N-C-C' sequence in the polypeptide chain segments coincides with C3' - C5' direction in the adjacent polynucleotide chains. According to the code proposed by us the side chains of amino acid residues present in the outward-pointing positions in a β -sheet control hydrogen binding interactions between the peptide groups of recognition protein site and DNA base pairs. These residues can be divided in two groups those coding for AT pairs -(Ser, Thr, Asn, Gln, His, Cys, in some cases, presumably, Lys and Arg) and those coding for GC pairs (Gly, Ala, Leu, Val, Ile, Phe, Met, Trp, Tyr, Asp, Glu, in certain cases Lys and Arg).^{1,2} These rules are strongly supported by the fact that for a number of specific protein nucleic acid complexes there is a remarkable correspondence between the protein and nucleic acid sequences (lac repressor-lac operator, λ repressor- λ operator, Cro protein - λ operator,

CI protein and its interaction site in cY region of λ genome, ribosomal S8 protein and its interaction site on ribosomal 16S SNA). These rules are also supported by our observations that ϕ polypeptides bind to synthetic DNA's with defined sequences and exhibit base sequence preferences predicted by our code rules. For example, oligo (L-threonine) binds more strongly to poly(dA)*poly(dT) than to poly(dG)*poly(dC), whereas oligo (L-valine) exhibits an opposite order of binding preferences. The binding is accompanied by spectral changes in the UV spectral region 190-240 nm thereby indicating that amide groups of the oligopeptide molecules are implicated in the binding interactions. Oligopeptides in a single-stranded form exhibit a low affinity for BNA as compared with the affinity of dimeric species. Binding isotherms were obtained by a number of techniques and interpreted on the basis of statistical mechanical theory developed by us earlier.^{3,4}

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