

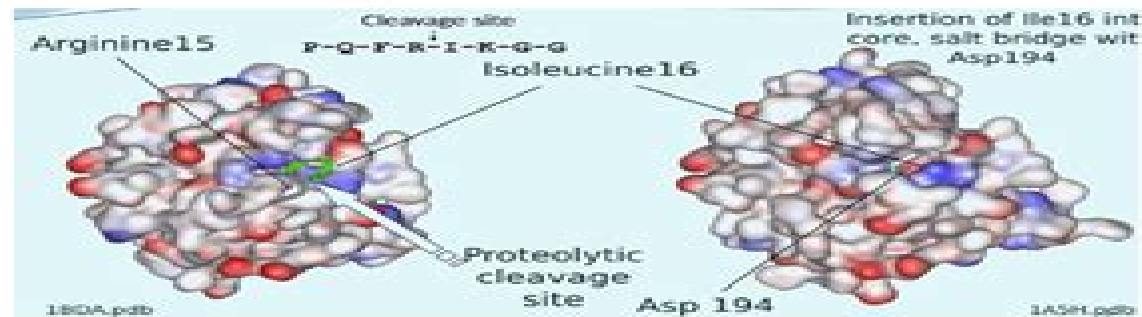
## Molecular Enzymology Lecture 2 (Serine Proteases: Mechanism)

### Serine Protease Activation Mechanism (Chymotrypsin)

- The limited proteolysis of the inactive, zymogen form of chymotrypsin, chymotrypsinogen, involves cleavage of the peptide bond between residue 15 and 16



- This cleavage creates a new N-terminus on the now exposed Ile-16 residue
- This N-terminus of Ile-16 is now able to form a salt bridge with the, highly conserved, Asp194 residue in the interior of the protein



- This conformational change puts the NH-groups of residues Gly193 and Ser195 into the correct orientation to form the oxyanion hole



- It also causes the shifting of the residue Met192, opening the substrate specificity pocket
- The four catalytic features of the enzyme are now formed, and the enzyme is active
- The zymogen form of the enzyme contains the catalytic triad and the non-specific binding pocket but NOT the oxyanion hole or the substrate specificity pocket

# Molecular Enzymology

**Ensheng Dong**



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