DNA Repair and Mutagenesis

I. Chemicals react with DNA like every other molecule

Evidence for repair Survival Assays

II. Repair of Specific Common Forms of DNA Damage

Deamination

Hypoxanthine, Xanthine, Uracil

Base Excision Repair

DNA Glycosylases, AP Endonucleases, Polymerase I, Ligase

Very Short Patch

Dcm, Vsr

Oxidation

8-oxoG (more than 30 other common forms as well)

MutM, MutY, MutT

Alkylation

N⁷-methylguanine, N³-methyladenine, O⁶-methylguanine, O⁴-methylthymine

Base Excision Repair

Direct Reversal- Ogt, Ada: Suicide Proteins

Adaptive Response

Photoproducts

Pyrimidine dimer, 6-4 photoproduct

Direct Reversal- Photolyase

Base Excision Repair (in some organisms)

Interstrand Crosslinks

Repaired by?

III. General Repair Pathways

Methyl Directed Mismatch Repair

Dam, MutS, MutL, MutH, exonucleases, MutU(UvrD), Polymerase III

Nucleotide Excision Repair

UvrA, UvrB, UvrC UvrD, Pol I Ligase transcription-coupled repair, Mfd

IV. When Replication is Blocked

Special Damage Recognizing Polymerases

UmuCD, DinB, PolB others

Recombination Proteins At Replication Forks, Replication coupled repair?

SOS Response- Inducible repair

RecA, LexA

Weigle Mutagenesis

V. Isolation of Repair Mutants, Mutagenesis Proteins