MOLECULAR AND KINETIC ANALYSIS OF THE GLUTATHIONE S-TRANSFERASES INVOLVED IN DDT RESISTANCE IN ANOPHELES GAMBIAE

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Resistance to the DDT in the malaria vector Anopheles gambiae is associated with increased metabolism by members of the glutathione S-transferase family of enzymes. There are two classes of glutathione S-transferases in insects, class I and class II. Biochemical analysis of partially purified enzymes suggests that the class I glutathione S-transferases are primarily involved in resistance. The resistance, although inherited as a single major gene effect also involves numerous glutathione S-transferase isozymes. We have isolated cDNAs from An. gambiae which code for several members of the class I family. Two of these enzymes have been expressed in E. coli and metabolism studies on the expressed proteins have demonstrated that these enzymes are able to metabolise DDT. Sequencing of the genomic DNA indicates that the class I genes are arranged sequentially on the chromosome. In situ hybridisation to the polytene chromosomes also shows that the GST class I family occurs at a single location in the genome.