

**Interactions of 4-chlorophenol and phenol with phosphatidylethanolamine monolayers in relation to antibacterial action**

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The introduction of a chlorine atom at the 4-position of the phenol aromatic ring leads to an increase in antibacterial activity. If the antibacterial mode of action of 4-chlorophenol and phenol involves disruption of the bacterial cytoplasmic membrane, 4-chlorophenol itself should be more effective than phenol in disrupting those interactions which contribute to the maintenance of the integrity of the membrane. In order to determine the effect of 4-chlorophenol and phenol on lipid-lipid interactions, which may well contribute to the maintenance of the integrity of a bacterial cytoplasmic membrane, the interactions of 4-chlorophenol and phenol with monolayers of 1,2-dipalmitoyl-L-3-phosphatidylethanolamine at the liquid-air interface has been studied using a film balance technique.

Monolayers of phosphatidylethanolamine were spread and compressed on a sub-phase which was basically a five-fold dilution of McIlvaine buffer, pH 6.0, adjusted to an ionic strength of 0.1 with sodium chloride. The sub-phase temperature was  $25 \pm 0.1^\circ$ . Precautions were taken to ensure that the monolayers attained equilibrium after spreading and during their subsequent compression (Proudfoot and Davdani 1973).

A sub-phase concentration of  $0.65 \times 10^{-3}$  mol dm<sup>-3</sup> 4-chlorophenol was found to expand the phosphatidylethanolamine monolayers. Compression of the expanded monolayers resulted in the 4-chlorophenol molecules being ejected from the monolayers, implying that the interaction of 4-chlorophenol with the phospholipid molecules is of a weak non-specific nature.

The presence of 4-chlorophenol molecules between adjacent phosphatidylethanolamine molecules in a monolayer will interfere with the close packing and hence total lateral cohesion between adjacent phosphatidylethanolamine molecules. In this respect, a sub-phase concentration of  $1.95 \times 10^{-3}$  mol dm<sup>-3</sup> 4-chlorophenol reduced the total lateral cohesion between adjacent phosphatidylethanolamine molecules in a monolayer sufficiently that loss of phosphatidylethanolamine molecules occurred from the monolayer. Kaye & Proudfoot (1971) have also shown phenol and a number of alkylated phenols to exert a similar effect on phosphatidylethanolamine monolayers.

4-Chlorophenol was found to be more effective than phenol in expanding monolayers of phosphatidylethanolamine and hence in reducing the total lateral cohesion between adjacent phosphatidylethanolamine molecules. This is probably a consequence of (1) the flat orientation of 4-chlorophenol molecules at the sub-phase-air interface facilitating their ability to interfere with the close packing of adjacent phosphatidylethanolamine molecules and (2) the increased surface activity of 4-chlorophenol compared to phenol.

## REFERENCES

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**Lysozyme—preservative interactions in eye preparations**

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The interaction of lysozyme with preservatives used in eye preparations is of importance, mainly because it is undesirable to inactivate the lysozyme present in the tear fluid which plays a major role in the hygiene of the eye. The possibility of lysozyme interactions occurring is greatest with the many contact lens solutions now on the market most of which contain two or three component preservative systems. Some of these solutions are carried into the eye on the contact lens or are instilled separately.

The disruption of Gram-negative cells by lysozyme in the presence of EDTA and tris is well known (Respaske, 1958). We have investigated certain preservatives to ascertain