

# Abstract

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## Differential effects of eicosapentaenoic acid and docosahexaenoic acid on human skin fibroblasts.

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**OBJECTIVE AND METHODS:** To better understand the mode of action of omega 3 fatty acids in cell membranes, human foreskin fibroblasts were grown in serum-free medium supplemented with 50 microM oleic acid linoleic acid, eicosapentaenoic acid (EPA) or docosahexaenoic acid (DHA), and the effects on membrane composition, fluorescence polarization and enzyme activities were followed.

**RESULTS:** The cells were enriched with EPA and DHA up to 7 and 13% of total lipids, respectively, of which > 95% was associated with phospholipids. In addition, the concentration of 22:5n-3 increased with both EPA and DHA to 7.5, and 2.1% of the total fatty acids, respectively. When compared to controls (oleic acid), cells treated with DHA showed a decrease in cholesterol, phospholipids, arachidonic acid (AA) and free cholesterol/phospholipid ratio ( $P < 0.05$ ). In the presence of EPA, only decreases in AA and cholesterol were significant ( $P < 0.05$ ). Membrane fluidity, assessed by fluorescence anisotropy, was increased 16% in cells enriched with DHA ( $P < 0.05$ ), but showed no change with EPA or linoleic acid. There was an increase in membrane-associated 5'-nucleotidase (+27%) and adenylate cyclase (+19%) activities ( $P < 0.05$ ), in DHA-enriched, but not in EPA-enriched cells, when compared with oleate controls.

**CONCLUSIONS:** The studies show that incorporation of DHA, but not EPA, into cell membranes of fibroblasts alters membrane biophysical characteristics and function. We suggest that these two major n-3 fatty acids of fish oils have differential effects on cell membranes, and this may be related to the known differences in their physiological effects.

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