Investigations on the ethanol-induced flushing reaction: effects of propranolol and dipyridamole on acetaldehyde and prostacyclin metabolism

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Disulfiram, an aldehyde dehydrogenase (ALDH) inhibitor, induces a flushing reaction upon the ingestion of ethanol, exerting aversion against alcohol that has been used in the treatment of alcoholism. This unpleasant response has been associated with an accumulation of acetaldehyde, and more recently, with an increase in vascular prostacyclin (PGI2) production. To evaluate the possibility of evoking the flushing reaction with drugs less toxic than disulfiram, we studied the effects of propanolol and dipyridamole on ALDH and PGI2. Acetaldehyde oxidation rate was assessed by gas chromatography in mitochondria from rats treated with these drugs for seven days. Prostacyclin generation was determined in rat aortic rings incubated in Krebs-Ringer with these drugs separately and associated to acetaldehyde, and measured by radioimmunoassay of 6-keto-PGFa?. Propanolol inhibited acetaldehyde oxidation rate whereas dipyridamole did not. Furthermore, propanolol increased blood acetaldehyde levels w