(Biochem. Pharmacol., 37, 453 (1988))

Mouse Liver Dihydrodiol Dehydrogenases. Identity of the Predominant and a Minor Form with 17β -Hydroxysteroid Dehydrogenase and Aldehyde Reductase

HIDEO SAWADA,* AKIRA HARA, TOSHIHIRO NAKAYAMA, MAKOTO NAKAGAWA, YOSHIO INOUE, KAZUYA HASEBE, YUAN-PEI ZHANG

A major and a minor form of dihydrodiol dehydrogenese were co-purified with 17β -hydroxysteroid dehydrogenase and aldehyde reductase, respectively, from mouse liver cytosol. The major form of the enzyme showed higher affinity for 17β -hydroxysteroids and 17-ketosteroids than for xenobiotic alcohols. The dehydrogenase activities for hydroxysteroids and dihydrodiols exhibited identical heat stability and sensitivity to several inhibitors. The minor form of the enzyme reduced various aldehydes and was specifically inhibited by barbiturates and sorbinil.

(J. Biochem., 103, 1027 (1988))

Purification and Characterization of NADP-dependent 3α -Hydroxy-steroid Dehydrogenase from Mouse Liver Cyotosol.

AKIRA HARA, YOSHIO INOUE, MAKOTO NAKAGAWA, FUMIHARU NAGANEO, HIDEO SAWADA*

A monomeric 3α -hydroxysteroid dehydrogenese with a molecular weight of 34,000 was purified to apparent homogeneity from mouse liver cytosol. The enzyme catalyzed the reversible oxidation of the 3α -hydroxysteroids, reduced various carbonyl compounds, and was inhibited by SH-reagents, synthetic estrogens, anti-inflammatory drugs, prostaglandins, and Δ^4 -3-ketosteroids. Uncompetitive substrate inhibition with respect to NADPH was observed in the reverse reaction. Steady-state kinetic measurements in both directions indicated that the reaction proceeds through an ordered bi bi mechanism with NADP(H) binding to the free enzyme.

(Biochemistry (Life Sci. Adv.), 7, 121 (1988))

Purification of Cytosolic Estradiol 17 β -Dehydrogenases from Male and Female Mouse Liver.

Akira Hara, Yoshio Inoue, Hiroyuki Tanabe, Makoto Nakagawa, Toshihiro Nakayama, Hideo Sawada*

Three forms of 17β -hydroxysteroid dehydrogenase were purified from liver cytosol of male ddy mice. All forms of the enzyme were monomeric proteins with the same Mr of 36,000 but differing charges. The enzymes oxidized estradiol- 17β , 17β -hydroxyandrostanes, and nosteroidal alicyclic alcohols with NADP or NAD as a cofactor, and reduced estrone-3-sulfate, 17-ketoandrostanes and xenobiotic carbonyl compounds in the presence of NADPH. The three enzymes were not immunologically distinguished from one another. Female mouse liver cytosol exhibited the same level of 17β -hydroxy-steroid dehydrogenase as that in male liver cytosol, and also contained the three enzyme forms.