

Sequence of a cDNA specifying subunit VIIa of human cytochrome c oxidaseGian Maria Fabrizi¹, Rosario Rizzuto¹, Hirofumi Nakase¹, Shuji Mita¹, Margaret I. Lomax³, Lawrence I. Grossman⁴ and Eric A. Schon^{1,2*}

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Cytochrome c oxidase (COX; EC 1.9.3.1), the last component of the mitochondrial respiratory chain, catalyzes the transfer of electrons from reduced cytochrome c to molecular oxygen. In mammals, the apoprotein is composed of three large catalytic subunits, encoded by the mitochondrial genome, and by ten smaller, nuclear-encoded subunits, which may play a regulatory role (1); subunits VIa, VIIa, and VIII have been shown to have heart- and liver-specific isoforms in cows and pigs (2,3).

Using a bovine liver COX VIIa cDNA (4) as a probe, we isolated a full-length cDNA (clone pCOX7.22; sequence below) from a human adult endothelial cell cDNA library (a gift of M. Chao and D. Littman), specifying the liver-specific isoform of human COX VIIa. The deduced amino sequence of the mature polypeptide is 83% identical to beef liver COX VIIa (3,4) but is only 57% identical to beef heart COX VIIa (5). Human COX VIIa contains a 23-aa cleavable presequence for importation into mitochondria (underlined).

[GAATTCGG]AGTAACAGCCAAG (13)

ATGCTGGGAATCTGCTGCTCTCTGTCAGATTGGGCAGAGGAAGATAAGCAGCTGCTTCC (73)M L R N L L A L R Q I G Q R T I S T A S [-4]CGCAGGCATTTTAAAAATAAAGTTCCGGAGAAGCAAAAAGTTCAGGAGGATGATGAA (133)R R H F K N K V P E K Q K L F Q E D D E [17]ATTCACATGATCTAAAGGGTGGGGTAGCTGATGCCCTCCIGTATAGGCCACCATGATT (193)I P L Y L K G G V A D A L L Y R A T M I [37]CTTACAGTTGGTGGAAACAGCATATGCCATATATGAGCTGGCTGTGGCTTCATTTCCCAAG (253)L T V G G T A Y A I Y E L A V A S F P K [57]AAGCAGGAGTCACTTCAGTCATCCAGCAATCGCTTGGTTCAGTTTCATTCAGCTCTCTA (313)K Q E * [60]TGGACCAGTAATCTGATAAATAACCGAGCTCTTCTTTGGGGATCAATATTTATTGACTTG (373)TAGTAACTGCCACCAATAAAGCAGTCTTTACCATG-AAAAAAAAAAAAAAAA[CGAATTC] (408)

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Notes: (1) Kadenbach et al. (1987) *Curr. Top. Bioenerg.* 15, 113; (2) Kadenbach et al. (1983) *TIBS* 8, 398; (3) Yanamura et al. (1988) *Biochemistry* 27, 4909; (4) Seelan et al. (1989) *Nucl. Acids Res.*, in press; (5) Meinecke and Buse (1986) *Biol. Chem. Hoppe-Seyler* 367, 67. Supported by grants from NIH (NS11766 [E.A.S.], RR05384 [L.I.G.], and GM37086 [M.I.L.]); the Muscular Dystrophy Association; the Aaron Diamond Foundation; the Center for Molecular Biology, Wayne State University; and Dr. and Mrs. Libero Danesi.