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iso,Structure of the mitochondrial
carboxylate carrier (SLC25A1)
complexed with isobutyric acid:
implications for proton binding in the
transport reaction. SLC25A1, the
mammalian isoform of the yeast
mitochondrial carriers, serves as an ion-
coupled electrogenic exchanger for
dicarboxylates and as a proton carrier
for acyl-coenzyme A. The X-ray crystal
structure of the complex of SLC25A1
with isobutyric acid (IBA) at 2.3-Å
resolution reveals a trimer in the
asymmetric unit with a three-fold
helical symmetry axis. Each protomer
consists of a long N-terminal alpha-helix

of 24 residues, a larger central alpha-helix of 57 residues, and a smaller C-terminal alpha-helix of 34 residues. In the asymmetric unit, three protein molecules are arranged in a trimer of dimers with a pseudo-twofold axis of 42.5 degrees. Amino acids at the two interprotomer interfaces contribute ion pairs to the binding site for IBA and regulate the transporter-mediated transport activity. The substrate-binding site of SLC25A1 contains two long parallel alpha-helices forming a trifurcated cap and a highly curved hydrophilic channel running parallel to the helical axes. The cap and channel form a D-like shape with a width of 10 Å that can accommodate a substrate. The two beta-barrels
