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**COVER:** Schematic superposition of the  $\alpha$ -helical isoprenyl-binding pocket of farnesyl transferase (orange), geranylgeranyl transferase-1 (green), and Rab geranylgeranyl transferase (yellow). A bulky residue (tryptophan) in farnesyl transferase shortens the pocket and sterically hinders binding of geranylgeranyl pyrophosphate (GGpp; red) but not farnesyl pyrophosphate (Fpp; blue). At the equivalent position, both geranylgeranyl transferases contain a smaller residue, which lengthens the pocket and allows binding of geranylgeranyl pyrophosphate. (See Leung et al., p. 467.)

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