A significant amount of inactive CaMKII forms molecular complex with $I_{to}$ channel Kv4.3 proteins. The coupled inactive CaMKII cannot be activated because the CaM binding sites are occupied by Kv4.3. The coupled inactive CaMKII will be dissociated from the Kv4.3-CaMKII units by $I_{to}$ channel blocker 4-AP (or reduction of Kv4.3 expression), leading to excessive CaMKII activation.