



**SCHEME 8.8** Decision tree for impurities in drug substances. (Courtesy of Pharmquest Corporation, Mountain View, CA.)

available relevant information and performed in the species most likely to maximize the potential to detect the toxicity of an impurity. On a case-by-case basis, single-dose studies can be appropriate, especially for single-dose drugs. In general, a minimum duration of 14 days and a maximum duration of 90 days would be considered appropriate. Lower thresholds can be appropriate if the impurity is unusually toxic.