



**Figure 3.10** (a) Generation of disulfide cross-links (cysteine), sulfenic, sulfonic, and cysteic acids from  $\text{H}_2\text{O}_2$  oxidation of Cys. (b) Proposed mechanism for metal-catalyzed oxidation of Cys.

Reproduced from [Li, Schoneich, and Borchardt \(1995a\)](#).

the pKa for the Cys residue appears to be decreased substantially, mainly by electrostatic interactions with surrounding Lys residues ([Griffiths et al., 2002](#)).

### Cys oxidation and mixed disulfides in mAbs

Intermolecular disulfide cross-linking has been shown to occur in several mAbs. At first glance this is surprising since a free Cys residue is generally required for formation of mixed disulfides, and all Cys residues in a mAb are paired (Figure 1.2 (c)) so that there should not be any free thiol. However, it has been shown that recombinantly produced