

### 1.1.1 Absorption

Drug oral absorption is a complex transfer process across the intestinal lining which includes passive diffusion through the paracellular space or absorptive cells membranes, vesicular uptake (endocytosis/pinocytosis), and release at the basolateral space. Absorption depends initially on the dosage form dissolution in the aqueous contents of the gastrointestinal tract (GIT) to reach the blood (Le Ferrec *et al.* 2001a; Smith *et al.* 2012; Toutain and Bousquet-mélou 2004). Drug absorption from the GIT depends on various factors such as drug physicochemical properties (lipophilicity, size, molecular volume, stereochemistry, pK<sub>a</sub>, solubility, chemical stability, partition and structure), gastric and intestinal motility, gastric emptying, physicochemical properties of small intestine environment, intestinal pH and the Surface area available for absorption (Dressman and Reppas 2010; Le Ferrec *et al.* 2001b). Drug absorption takes place at different sites of the GIT which include duodenum, jejunum and ileum. Many biological *in vitro* methods are used as an assessment for the permeability of the GI mucosa to predict *in vivo* drug absorption such as everted rat intestinal sacs (ERIS) and *in situ* single pass intestinal perfusion (*in situ* SPIP) (Dressman and Reppas 2010).

*In situ* SPIP technique is used for the assessment of permeability of drugs through intestines while keeping intact blood supply to the intestinal tract throughout the experiment, making it a reliable tool for simulating real *in vivo* conditions following oral drug administration (Amidon *et al.* 1995; Amidon *et al.* 1988; Varma *et al.* 2004; Varma and Panchagnula 2005). Everted rat intestinal sac model is used for measuring absorption and permeability at different sites in the small intestine via passive diffusion and/or