



## Introduction

Atorvastatin (ATS) is the gold-standard treatment for hypercholesterolemia and the prevention of cardiovascular illnesses caused by dyslipidemia around the world. Physiologically based pharmacokinetic (PBPK) models have been positioned as a significant tool for the study of complex pharmacokinetic (PK) processes and their extrapolation in specific sub-populations, leading to regulatory acceptance [1]. In recent years, several PBPK models of ATS have been produced, each addressing a distinct element of ATS's PK features. The goals of this study are to (i) outline the physicochemical and pharmacokinetic properties involved in the time-course of ATS, and (ii) assess the primary highlights and limits of the PBPK models of ATS that have been published thus far. Common features relating to the physicochemical characteristics of ATS are included in the PBPK models. However, the analyte tested, the kind and influence of transporters and metabolic enzymes, and the permeability value employed all varies significantly. This review also outlines significant processes (lactonization, P-gp contribution, ATS-Ca solubility, simultaneous management of numerous analytes, and experimental data in the target population) that would improve PBPK model prediction and make it a useful tool for ATS dose optimization [2-4].

Patients with chronic renal failure frequently have a secondary form of complicated dyslipidaemia, and lipid-lowering therapy may be beneficial. Although atorvastatin has been proven to effectively lower levels of atherogenic lipoproteins in patients with renal failure, there is a paucity of pharmacokinetic data in haemodialysis patients. Hypercholesterolaemic haemodialysis patients were given 40 mg (n=12) or 80 mg (n=11) atorvastatin once daily for two weeks, initially as a single dose and subsequently continuously. LC/MS/MS was used to determine plasma levels of atorvastatin and its active and inactive metabolites, and pharmacokinetic characteristics ( $C_{max}$ ,  $t_{max}$ , AUC,  $t_{1/2}$ ) were compared between single and multiple dosing, as well as between different dosages [5].

After single and 2-week multiple dosing, the pharmacokinetic characteristics of the parent drug atorvastatin acid were not significantly

loading method. Due to toxicity concerns, it is essential to explore a rapid and reliable method to effectively isolate and quantify the non-liposomal, namely, free CPT-11 and total CPT-11 in plasma. This study focuses on separation of non-liposomal CPT-11, evaluation of the