1.1 Samples Preparation

Metabolite extraction was performed as mentioned elsewhere [1]. Briefly,100 µL aliquot of plasma was mixed with 900 µL of an extraction solvent 1:1 acetonitrile (ACN): methanol (MeOH). Concurrently, quality control (QC) samples were generated by taking aliquots from all samples to verify system stability. The mixtures were agitated on a thermomixer (Eppendorf, CITY, Germany) at 600 rpm and kept at room temperature (RT) for one hour. Subsequently, the samples underwent centrifugation at 16000 rpm, at a temperature of 4ºC, for a duration of 10 minutes. After centrifugation, 950 µL of the resultant supernatant was transferred into a 1.5-ml Eppendorf tube and then subjected to complete evaporation using a SpeedVac system (Christ, Germany). The dried samples were reconstituted with 100 µL of a 50% mobile phase A and B (A: 0.1% Formic acid in dH2O, B: 0.1% Formic acid in 50% ACN: MeOH). This reconstitution was followed by brief vortexing and then introduced into the LC-MS system for analysis.