

The present invention is based on the recognition, that the dosage form, which combines adequate drug stability, optimum drug release of both active ingredients, pharmacological efficacy and reliable manufacture for a combination of telmisartan and amlodipine best, is a bilayer tablet.

Summary of the invention

In accordance with the ~~present invention disclosure~~ problems associated with the preparation of a fixed dose combination drug comprising telmisartan and amlodipine can best be handled by means of a bilayer pharmaceutical tablet comprising a first layer of telmisartan, preferably in substantially amorphous form, in a dissolving tablet matrix and a second layer of amlodipine in a disintegrating or eroding tablet matrix.

Hence, in a first aspect, there is provided a pharmaceutical tablet comprising a first layer of telmisartan in a dissolving tablet matrix and a second layer of amlodipine in a disintegrating or eroding tablet matrix.

The tablet according to the ~~present invention disclosure~~ provides a largely pH-independent dissolution of the poorly water-soluble telmisartan, thereby facilitating dissolution of the drug at a physiological pH level, and adequate stability and drug release of amlodipine. The tablet structure also overcomes the stability problem caused by the incompatibility of amlodipine with basic constituents of the telmisartan formulation.

In a second aspect, there is provided a method for the manufacture of a tablet of the first aspect to treat hypertension either alone or in combination with the treatment or prevention of a condition selected from the group consisting of chronic stable angina, vasospastic angina, stroke, myocardial infarction, transient ischemic attack, congestive heart failure, cardiovascular disease, diabetes, insulin resistance, impaired glucose tolerance, pre-diabetes, type 2 diabetes mellitus, diabetic nephropathy, metabolic syndrome (syndrome X), obesity, dyslipidemia.

hypertriglyceridemia, elevated serum concentrations of C-reactive protein, elevated serum concentrations of lipoprotein(a), elevated serum concentration of homocysteine, elevated serum concentration of low-density lipoprotein (LDL)-
5 cholesterol, elevated serum concentration of lipoprotein-associated phospholipase (A2), reduced serum concentration of high density lipoprotein (HDL)-cholesterol, reduced serum concentration of HDL(2b)-cholesterol, reduced serum concentration of adiponectin, cognitive decline and dementia.

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Definitions

As used herein, the term "substantially amorphous" refers to a product comprising amorphous constituents in a proportion of at least 90%, preferably at least 95%, as determined by X-ray powder diffraction measurement.

15 The term "dissolving tablet matrix" refers to a pharmaceutical tablet base formulation having instant release (fast dissolution) characteristics that readily dissolves in a physiological aqueous medium.

The term "disintegrating or eroding tablet matrix" refers to a pharmaceutical tablet base formulation having instant release characteristics that readily disintegrates or
20 erodes in a physiological aqueous medium.