Solvent (qs), e.g. water – if necessary (i.e. only the amount needed, if any)
Extra-granular fraction
10 to 50% (e.g. 20 to 40%) of excipient/diluent (preferably a non-soluble excipient/diluent e.g. microcrystalline cellulose, such as silicified microcrystalline cellulose)
1 to 5% (e.g. 2 to 4%) disintegrant (e.g. crospovidone, such as Polyplasdone XL)
0 to 3% (e.g. 0.1 to 1%) glidant (e.g. colloidal silicon dioxide, Aerosil 200)
0 to 5% (e.g. 1 to 3%) lubricant (e.g. sodium stearyl fumarate (Pruv)).

In another aspect of the present disclosure, there is provided a composition wherein the different parts of the composition, specifically the intra-granular and extra-granular fraction and binder portion, comprise the following ingredients by weight based on the total weight of the composition:
Intra-granular fraction
5 to 50% of bedaquiline fumarate as the active ingredient
10 to 50% of non-soluble excipient/diluent
1 to 5% disintegrant
0.1 to 5% glidant
Binder
1 to 10% binder or polymer
0.01 to 5% wetting agent or surfactant
Solvent (qs)
Extra-granular fraction
10 to 50% of excipient/diluent
1 to 5% disintegrant
0 to 3% glidant
0 to 5% lubricant.

The compositions (e.g. tablet compositions) of the disclosure mentioned herein may be charactersied in that specifically the intra-granular layer is absent any soluble excipient/diluent. Hence, the extra-granular fraction need not be absent any soluble excipient/diluent, although, preferably, it is the case that the extra-granular fraction is also absent any soluble excipient/diluent.

Claims

1. A dispersible composition comprising bedaquiline fumarate as the active ingredient and wherein the composition comprises an intra-granular and extra-granular layer in which the intra-granular layer comprises a non-soluble excipient/diluent and is charaterised in that the intra-granular layer is absent of a soluble excipient/diluent that is starch.

2. A dispersible composition as claimed in Claim 1, wherein the intra-granular layer is absent of any soluble excipient/diluent.

3. A dispersible composition as claimed in Claim 1 or Claim 2, wherein the non-soluble excipient/diluent in the intra-granular layer is microcrystalline cellulose.

4. A dispersible composition comprising bedaquiline fumarate as the active ingredient and wherein the composition comprises an intra-granular and extra-granular layer in which the intra-granular layer comprises a non-soluble excipient/diluent that is microcrystalline cellulose.

5. A dispersible composition as claimed in Claim 3, wherein the intra-granular layer is absent of a soluble excipient/diluent.

6. A dispersible composition as claimed in any one of the preceding claims, wherein the dispersible composition is a tablet.

7. A composition wherein the different parts of the composition, specifically the intra-granular and extra-granular fraction and binder portion, comprise the following ingredients by weight based on the total weight of the composition:

Intra-granular fraction

5 to 50% of bedaquiline fumarate as the active ingredient
10 to 50% of non-soluble excipient/diluent
1 to 5% disintegrant
0.1 to 5% glidant

Binder
1 to 10% binder or polymer
0.01 to 5% wetting agent or surfactant
Solvent (qs)
Extra-granular fraction
10 to 50% of excipient/diluent
1 to 5% disintegrant
0 to 3% glidant
0 to 5% lubricant.

8. A composition as claimed in Claim 7, wherein the intra-granular layer is absent of any soluble excipient/diluent; and/or, optionally, the extra-granular fraction is also absent of any soluble excipient/diluent.

9. A composition as claimed in Claim 7 or Claim 8, which consists of the following compositions of intra-granular fraction, binder and extra-granular fraction, by weight based on the total weight of the composition:
Intra-granular fraction
24.18% of bedaquiline fumarate as the active ingredient
29.82% of non-soluble excipient/diluent
3% disintegrant
2 % glidant
Binder
3% binder or polymer
0.2% wetting agent or surfactant
Solvent (qs)
Extra-granular fraction
32.3% of excipient/diluent
3% disintegrant
0.5 % glidant
2% lubricant.

10. A composition as claimed in Claim 7 or Claim 8, which consists of the following compositions of intra-granular fraction, binder and extra-granular fraction, by weight based on the total weight of the composition:
Intra-granular fraction
25% of bedaquiline fumarate as the active ingredient

30% of non-soluble excipient/diluent
3% disintegrant
2 % glidant
Binder
3% binder or polymer
0.2% wetting agent or surfactant
Solvent (qs)
Extra-granular fraction
30% of excipient/diluent
3% disintegrant
0.5 % glidant
2% lubricant.

11. A composition as claimed in any one of claims 7 to 10, where the composition is a tablet composition.

12. A composition as claimed in any of Claims 7, 9 and 10, wherein:
the non-soluble excipient/diluent is microcrystalline cellulose;
the disintegrant is crospovidone;
the glidant is colloidal silicon dioxide;
the binder or polymer is hypromellose 5 cps;
the wetting agent or surfactant is polysorbate 20;
the solvent (qs) is water;
(other) excipient/diluent (if applicable) is a non-soluble excipient/diluent microcrystalline cellulose;
the disintegrant is crospovidone;
the glidant is colloidal silicon dioxide; and/or
the lubricant is sodium stearyl fumarate.

13. A composition as claimed in any one of the preceding claims for use in the treatment of tuberculosis.

14. A composition as claimed in Claim 13, for use in the pediatric and/or geriatric population.

15. A composition as claimed in any one of claims 1 to 12 for use in combination with one or more other therapeutic agent(s) useful in the treatment of tuberculosis.

16. A combination comprising a composition as claimed in any one of Claims 1 to 12 and one or more other therapeutic agents useful in the treatment of tuberculosis.

17. A process for preparing a composition as claimed in any one of Claims 1 to 10which comprises:

(a) obtaining an intra-granular fraction using the intra-granular fraction components mentioned in any one of Claim 7 to 10;

(b) preparing a binder fraction, using the binder fraction components mentioned in any one of Claims 7 to 10;

(c) obtaining an extra-granular fraction using the extra-granular fraction components mentioned in any one of Claims 7 to 10,

and using those fractions to prepare a composition.

18. A composition obtainable by the process as defined in Claim 17.