

Amendment date: 12 April 2019

1 **1.3.1.1 Package Insert (clean)**
2

3 **SCHEDULING STATUS**

4 S1
5

6 **PROPRIETARY NAME AND DOSAGE FORM**

7 **Zymafluor® Tablets**

8 **Zymafluor® Drops**
9

10 **COMPOSITION**

11 Each tablet and every four drops contain 0,55 mg sodium fluoride equivalent to 0,25
12 mg fluoride.
13

14 **ZYMAFLUOR Drops** contains Benzoic acid 0,15 % m/v as preservative.
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16 **PHARMACOLOGICAL CLASSIFICATION**

17 A 24 Mineral substitutes, electrolytes
18

19 **PHARMACOLOGICAL ACTION**

20 The protective action of **ZYMAFLUOR** is due to its accumulation in the outer layers of
21 the dental enamel.

22 Before the teeth erupt, **ZYMAFLUOR** is carried to the developing teeth by the
23 bloodstream and thus permits effective fluoridation before the teeth appear. After
24 their eruption the teeth take up fluoride by way of direct contact with the fluoride
25 contained in the saliva.

Amendment date: 12 April 2019

26 **ZYMAFLUOR** tablets should thus not be swallowed whole but should be allowed to
27 dissolve slowly in the mouth.

28

29 **INDICATIONS**

30 Prophylaxis of dental caries.

31

32 The administration of **ZYMAFLUOR** is recommended solely for the prevention of
33 tooth decay and should be considered when water suppliers are fluoride deficient.

34

35 Special note: There is no convincing evidence that fluoride from any source reduces
36 the development of caries after the permanent teeth are formed (usually about age
37 14).

38

39 **CONTRA-INDICATIONS**

40 **ZYMAFLUOR** is contra-indicated in kidney disease.

41

42 Hypersensitivity to sodium fluoride.

43

44 **ZYMAFLUOR** should not be used until the fluoride concentration in the drinking water
45 supply is known.

46

47 Use must be especially avoided in regions in which fluoridation is practiced in the
48 form of addition of fluoride to table salt or the drinking water supply.

49

50 **WARNINGS:**

Amendment date: 12 April 2019

51 If the water contains more than 0,7 mg/l of fluoride, supplementation is not
52 recommended.

53

54 To avoid overdosage, allowance should be made for fluorides ingested from other
55 sources such as the diet or fluoride toothpastes.

56

57 **DOSAGE AND DIRECTIONS FOR USE**

58 If not otherwise prescribed by a doctor or dentist, the following dosages are
59 recommended as a guideline:

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Age	Drinking water containing less than 0,3 mg/l of fluoride	Drinking water containing 0,3 to 0,7 mg/l of fluoride	Drinking water containing more than 0,7 mg/l of fluoride
0 to 2 years	1 tablet or 4 drops daily	Nil	Nil
2 to 4 years	2 tablets or 8 drops daily	1 tablet or 4 drops daily	Nil
4 to 14 years and over	4 tablets or 16 drops daily	2 tablets or 8 drops daily	Nil

61

62 Administration should begin in very early childhood.

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Amendment date: 12 April 2019

64 Four drops or one tablet of 0,25 mg (which should be crushed and dissolved in a little
65 water) should be added to a bottle (not of milk) or solid meal or administered alone.

66

67 As soon as the age of the child permits, ZYMAFLUOR tablets should no longer be
68 swallowed, but sucked slowly in the mouth, between the cheek and gum, sometimes
69 on the left and sometimes on the right side. They are best taken in the evening
70 before bedtime, after brushing the teeth as in this way high fluoride concentrations in
71 the mouth can be maintained for a longer period.

72

73 Please consult your local Dentist, Pharmacist, Medical Doctor or Medical Officer of
74 Health to ascertain whether **ZYMAFLUOR** supplementation is necessary.

75

76 **SIDE EFFECTS AND SPECIAL PRECAUTIONS**

77 The major manifestations or signs of the chronic ingestion of excessive amounts of
78 fluoride are mottled enamel of the teeth and osteosclerosis (a phenomenon wherein
79 the density and calcification of bone are increased). Clinical signs are joint pain,
80 stiffness, limited movement, and in severe cases crippling deformity.

81

82 Don not exceed the recommended dose.

83

84 Keep out of the reach of young children.

85

86 **INTERACTIONS**

Amendment date: 12 April 2019

87 Absorption of fluoride is inhibited by calcium, magnesium or aluminium.

88 **ZYMAFLUOR** should therefore not be given with milk and dairy products, not with
89 antacids containing calcium, aluminium or magnesium salts.

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91 **KNOWN SYMPTOMS OF OVERDOSAGE AND PARTICULARS OF ITS**
92 **TREATMENT**

93 In acute poisoning, sodium fluoride taken by mouth is corrosive, forming hydrofluoric
94 acid in the stomach.

95

96 Initial symptoms are secondary to the local action of fluoride on the mucosa of the
97 gastro-intestinal tract. Ingestion of more than 250 mg of sodium fluoride may
98 frequently cause salivation, nausea, abdominal pain, vomiting and diarrhoea. (NB. ie.
99 **ZYMAFLUOR** tablets or 1 800 drops). Systemic symptoms are varied and severe.

100

101 Large doses cause thirst, perspiration, paralysis, muscular weakness and clonic
102 convulsions, followed by respiratory and cardiac failure; there may be renal failure;
103 death may occur within 2 to 4 hours.

104

105 The patient shows signs of increasing irritability of the nervous system, including
106 paraesthesias, a positive Chvostek sign, hyperactive reflexes, and tonic and clonic
107 convulsions. Metabolic and electrolyte disturbances may occur. Hypocalcaemia and
108 hypoglycaemia are frequent laboratory findings. The signs may be delayed for
109 several hours. Pain in various muscle groups may occur. The blood pressure falls,
110 presumably due to central vasomotor depression as well as a direct toxic action on
111 cardiac muscle. The respiratory centre is first stimulated and later depressed.

Amendment date: 12 April 2019

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113 It is stated that the lethal dose of sodium fluoride for adults is about 5 to 10 g,
114 whereas dangerous poisoning has been reported after oral doses of less than 1 g. In
115 children, 0,5 g of sodium fluoride may be fatal (NB. ie. Nearly 900 **ZYMAFLUOR**
116 tablets or 3 600 drops).

117

118 Ocular damage has also been reported.

119

120 *Treatment*

121 Contact your nearest hospital without delay.

122 Treatment is symptomatic and supportive.

123

124 The principles of the treatment are as follows:

125

1. Act quickly.

126

2. In acute poisoning, empty the stomach by aspiration and lavage with lime
127 water or a 1 % solution of calcium chloride or other calcium salt.

128

3. Calcium gluconate injection, 10 ml of a 10 % solution, may be given
129 intravenously to control convulsions and repeated every 4 to 6 ours if needed.

130

4. Morphine or pethidine may be given by injection if necessary, to control colic.

131

5. The circulation should be maintained with infusions of suitable electrolyte
132 solutions.

133

6. The respiration may require assistance.

134

7. Haemodialysis has been used.

135

8. Wash away vomitus, faeces and urine promptly to prevent external burns.

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Amendment date: 12 April 2019

137 **IDENTIFICATION**

138 **Tablets:** White, round, biconvex tablets: diameter approximately 4 mm, thickness
139 approximately 2,6 mm

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141 **Drops:** Clear, colourless and odourless solution

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143 **PRESENTATION**

144 **Tablets:** Plastic containers of 150 and 400 tablets.

145 **Drops:** Plastic containers of 20 ml

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147 **STORAGE INSTRUCTIONS**

148 Store at or below 25 °C. The container must be tightly closed. Store out of reach of
149 children.

150

151 **REFERENCE NUMBER**

152 Tablets: E/24/728

153 Drops: Y/24/394

154

155

156 **NAME AND BUSINESS ADDRESS OF THE HOLDER OF THE REGISTRATION**

157 **CERTIFICATE**

158 XIXIA PHARMACEUTICALS (PTY) LTD

159 Building 6

160 Greenstone Hill Office Park

161 Emerald Boulevard

Applicant: TAKEDA (PTY) LTD to XIXIA PHARMACEUTICALS (PTY) LTD
(Transfer of Applicancy)
Product Name: ZYMAFLUOR
Dosage form and strength: Each tablet contains 0,25 mg sodium fluoride

MODULE 1
1.3.1.1

Amendment date: 12 April 2019

162 Modderfontein

163 1645

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165 **DATE OF PUBLICATION OF THE PACKAGE INSERT**

166 07 June 1991