### Summary of evidence and Expert Committee recommendations

An application was prepared by Pharmacy Students: Sandeep Kishore, Maryam Shafaee, Mathew Price, and Rajesh Vedanthan, and Marcus Reidenberg, Professor of Pharmacology, Medicine and Public Health, New York, for the addition of bisoprolol with a square box to the Model List, replacing atenolol as the representative medicine of the class in Sections 12.1 to 12.3, and the inclusion of beta-blockers as a therapeutic class in Section 14.4 (Medicines used in heart failure). Listing is requested as a representative of its therapeutic class. The Committee noted that heart failure is an important global health issue and its prevalence is increasing worldwide due to both communicable and noncommunicable causes. Recent guidelines from the National Institute of Clinical Excellence (NICE), United Kingdom, and Heart Failure Society of America (HSFA) recommend beta-blockers for the treatment of heart failure and specifically cite metoprolol, bisoprolol, and carvedilol (1, 2). The Committee considered evidence from 3 RCTs (3-5) to support the efficacy and safety of the beta-blockers bisoprolol, metoprolol, and carvedilol in the treatment of heart failure, as well as 3 meta-analyses that found a reduction of 29% to 34% in the composite end-point of mortality or hospital admission with beta-blocker therapy in patients with heart failure (6-8). Mortality benefits have been shown in diverse patient groups, including the elderly (9), patients with diabetes (5) and without (10), patients with an ejection fraction above or below 25% (11) and patients not receiving rennin-angiotensin inhibitors (12); additionally bisoprolol can be used in patients with chronic obstructive pulmonary disease (13). The Committee noted that there is no high-quality evidence to support the use of atenolol for the treatment of heart failure. The Committee noted that there is evidence from clinical trials to support the efficacy and safety of bisoprolol for the treatment of heart failure and specifically cite metoprolol, bisoprolol, and carvedilol (1, 2). The Committee also took into consideration a meta-analysis (22) (5 studies, n=17671, follow-up 4.6 years) that suggested older hypertensive patients treated with atenolol have a significantly higher mortality when compared to patients treated with other classes of cardiovascular medicines. Cardiovascular mortality was also higher in the atenolol treated group than with other antihypertensive treatment, and strokes were more frequent with atenolol treatment. The Committee noted that on a cost per dose basis bisoprolol was cheaper than metoprolol and carvedilol. The Committee concluded that there was sufficient evidence of efficacy and safety compared to atenolol to support the request for bisoprolol to become the representative beta-blocker in sections 12.1 to 12.3 and also recommended,

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**Bisoprolol**

**Indication**
Heart failure

**ICD11 code**: BD0Z

**INN**
Bisoprolol

**Medicine type**
Chemical agent

**List type**
Core

**Formulations**
Oral > Solid: 1.25 mg ; 5 mg

**EML status history**
First added in 2011 (TRS 965)

**Sex**
All

**Age**
Adolescents and adults

**Therapeutic alternatives**
metoprolol (ATC codes: C07AB02)
carvedilol (ATC codes: C07AG02)

**Patent information**
Patents have expired in most jurisdictions
Read more about patents.

**Wikipedia**
Bisoprolol

**DrugBank**
Bisoprolol

**ATC codes**: C07AB07
based on evidence of efficacy, safety, and cost-effectiveness, that bisoprolol should be added to the Model List for the treatment of heart failure. Due to the similarities between bisoprolol and metoprolol in terms of efficacy, the Committee decided to add bisoprolol with a square box for this indication. It was noted that country programmes could choose between bisoprolol, metoprolol, or carvedilol, but that there were increasingly reasons not to select atenolol as the sole beta-blocker provided.

References: