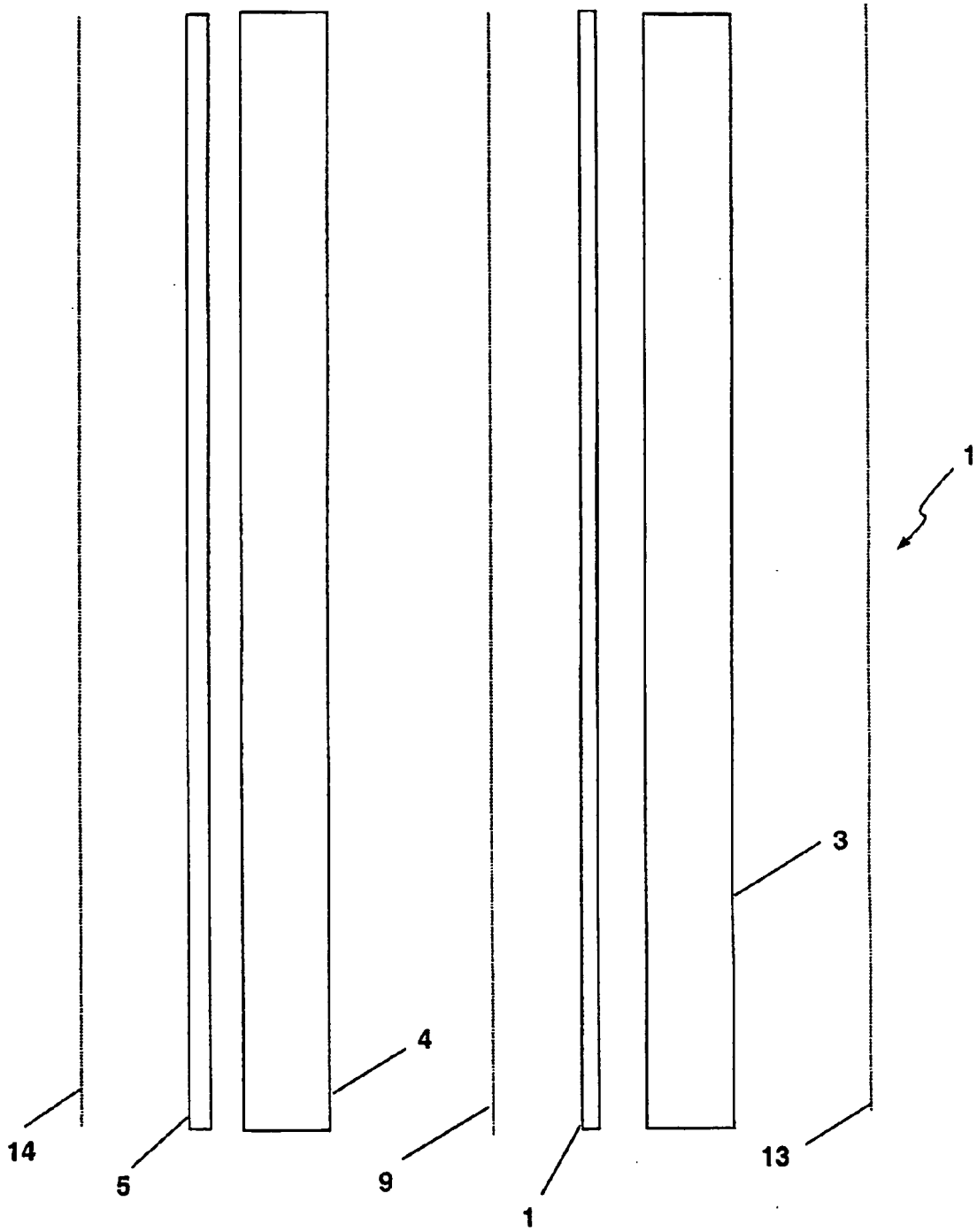**Figure 1**