## Comparison of three treatment strategies in mild asthma: A carbon-utility analysis – Supporting information

eTable 1. Model inputs and willingness thresholds

Variable	Value	Distribution	Source
A. Model inputs			
Kilograms CO <sub>2</sub> equivalent emissions – mean (SE)			
Budesonide DPI + salbutamol pMDI	17.3 (3.3)	Gamma	1
Salbutamol pMDI	26.2 (3.3)	Gamma	1
Budesonide/formoterol DPI	1.1 (3.3)	Gamma	1
Proportion of well controlled days per year [GINA classification] (SD) <sup>a</sup>			
Budesonide DPI + salbutamol pMDI	0.453 (0.036)	Beta	2
Salbutamol pMDI	0.328 (0.051)	Beta	2
Budesonide/formoterol DPI	0.301 (0.065)	Beta	2
Proportion of partly controlled days per year [GINA classification] (SD) <sup>a</sup>			
Budesonide DPI + salbutamol pMDI	0.406 (0.046)	Beta	2
Salbutamol pMDI	0.451 (0.045)	Beta	2
Budesonide/formoterol DPI	0.522 (0.053)	Beta	2
Proportion of poorly controlled days per year [GINA classification] (SD) <sup>a</sup>			
Budesonide DPI + salbutamol pMDI	0.142 (0.031)	Beta	2
Salbutamol pMDI	0.222 (0.030)	Beta	2
Budesonide/formoterol DPI	0.177 (0.017)	Beta	2
Proportion of days under exacerbations not resulting in ED visits <sup>b</sup>			
Budesonide DPI + salbutamol pMDI	0.003	_c	2
Salbutamol pMDI	0.008	_c	2
Budesonide/formoterol DPI	0.004	_c	2
Proportion of days under exacerbations resulting in ED visits <sup>b</sup>			
Budesonide DPI + salbutamol pMDI	0.0003	_c	2
Salbutamol pMDI	0.0004	_c	2
Budesonide/formoterol DPI	0	_c	2
Utilities – mean (SD)			
Well controlled asthma [GINA classification]	0.969 (0.045)	Beta	3
Partly controlled asthma [GINA classification]	0.935 (0.065)	Beta	3
Poorly controlled asthma [GINA classification]	0.884 (0.088)	Beta	3
Exacerbations not resulting in ED visits	$0.864 (0.066^{d})$	Beta	4
Exacerbations resulting in ED visits	$0.764 (0.066^{d})$	Beta	4
B. Willingness thresholds (per capita kilograms CO <sub>2</sub> equivalent emission	ns in the healthca	are sector)	
China	250	NA	5
European Union	490	NA	5
United States	1720	NA	5

<sup>&</sup>lt;sup>a</sup> Considering days without exacerbations; <sup>b</sup> Considering that the effects of exacerbations last for seven days, as frequently assumed in health economic evaluation studies<sup>4, 6</sup>; <sup>c</sup> No data on uncertainty available; <sup>d</sup> Average of the standard-deviations for the utilities associated with the different levels of asthma control.

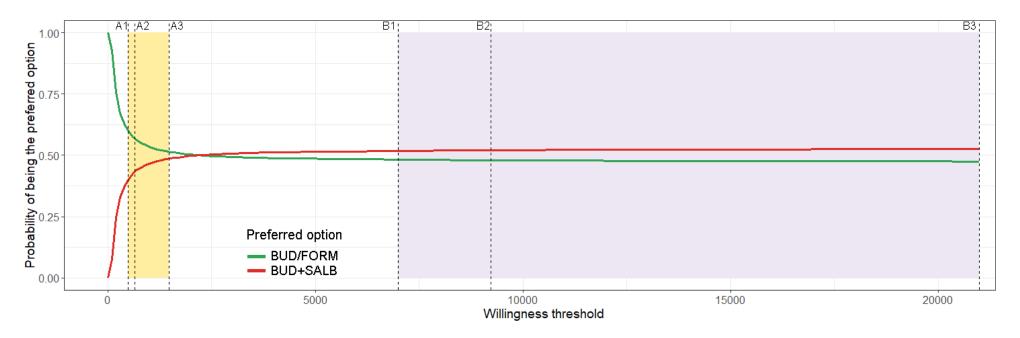
 $CO_2$ =Carbon dioxide; DPI=Dry powder inhaler; ED=Emergency department; GINA=Global Initiative for Asthma; NA=Not applicable; pMDI=pressurised metered-dose inhaler; SD=Standard-deviation; SE=Standard-error

eTable 2. Assessment of different willingness-to-emit thresholds considering the comparison between budesonide + salbutamol (BUD+SALB) versus budesonide/formoterol (BUD/FORM). Per capita kgCO<sub>2</sub>e emissions in the European Union are being considered.

Willingness-to-emit threshold [kgCO2e emissions/QALY gained]	Differenc e in utilities	Maximum difference in kgCO <sub>2</sub> e emissions for BUD+SALB to be the preferred option (% of per capita kgCO <sub>2</sub> e emissions in the healthcare sector)
Per capita kgCO <sub>2</sub> e emissions in the healthcare sector - 490	0.008	3.9 (0.8)
Per capita kgCO <sub>2</sub> e emissions in the healthcare sector adjusted to mean utilities in the general population <sup>a</sup> - 646	0.008	5.2 (1.1)
Two times per capita kgCO <sub>2</sub> e emissions in the healthcare sector - 980	0.008	7.8 (1.6)
Three times per capita kgCO <sub>2</sub> e emissions in the healthcare sector -1470	0.008	11.8 (2.4)
Per capita total kgCO <sub>2</sub> e emissions <sup>7</sup> - 7000	0.008	56.0 (11.4)
Per capita total kgCO <sub>2</sub> e emissions adjusted to mean utilities in the general population <sup>7 a</sup> - 9235	0.008	73.9 (15.1)
Two times per capita total kgCO <sub>2</sub> e emissions <sup>7</sup> - 14000	0.008	112.0 (22.9)
Three times per capita total kgCO <sub>2</sub> e emissions <sup>7</sup> - 21000	0.008	168.0 (34.3)

<sup>&</sup>lt;sup>a</sup> Portuguese values being considered as an example (0.758)<sup>8</sup>

eFigure 1. Results of the probabilistic sensitivity analysis considering different willingness-to-emit (WTE) thresholds. The yellow shade marks the probability of budesonide/formoterol (BUD/FORM) or budesonide + salbutamol (BUD+SALB) being the preferred option considering a set of WTE thresholds that ranges between 1 (A1) to 3 (A3) times the per capita kgCO<sub>2</sub>e emissions in the healthcare sector in the European Union (including an adjusted WTE threshold dividing A1 by the average utilities level in the general population – A2). The blue shade marks the probability of BUD/FORM or BUD+SALB being the preferred option considering a set of WTE thresholds that ranges between 1 (B1) to 3 (B3) times the per capita total kgCO<sub>2</sub>e emissions in the European Union (including an adjusted WTE threshold dividing B1 by the average utilities level in the general population – B2). For B1-B3, except for simulations in which BUD+SALB has been a dominated option, BUD+SALB is practically always the preferred option.



## References

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