

Environmental Impact of Aflibercept for the Treatment of Neovascular Age-Related Macular Degeneration and Diabetic Macular Oedema in The Netherlands

S. Quist^{1,2}, J. Paulissen^{1,2}, R. Freriks^{2,3}

1. Department of Health Sciences, University of Groningen, University Medical Center Groningen, Groningen, The Netherlands. 2. Asc Academics, Groningen, The Netherlands. 3. Department of Economics, Econometrics & Finance, University of Groningen, Faculty of Economics & Business, Nettelbosje 2, Groningen, The Netherlands

Introduction

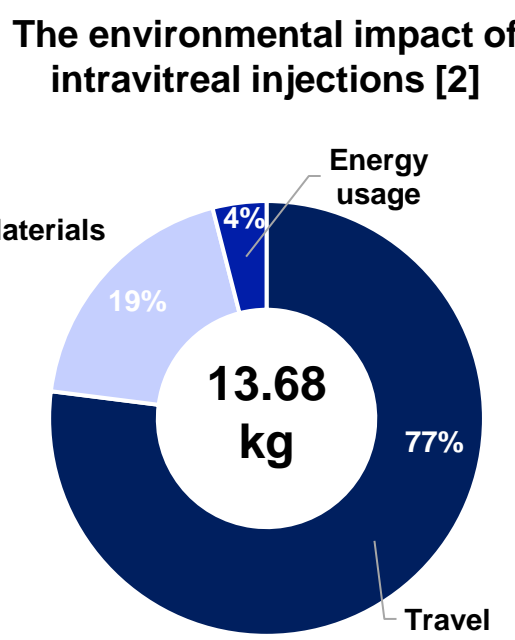
The environmental impact of the healthcare sector

- Healthcare plays a considerable role in carbon emissions and transitioning towards sustainable practices is essential for achieving climate goals.
- In the Netherlands, the healthcare sector contributes approximately 7% of global greenhouse gas emissions [1].
- The footprint of the healthcare system consists of medical procedures and the effects of the production of goods and services used in healthcare.



The environmental impact of anti-VEGF treatment

- With the increasing prevalence of neovascular age-related macular degeneration (nAMD) and diabetic macular oedema (DME), the use of anti-vascular endothelial growth factor (anti-VEGF) treatments through intravitreal injections has become more common.
- Each injection of anti-VEGF treatment is estimated to generate 13.68 kg of carbon dioxide equivalent (CO2eq) and therefore, this regimen strongly impacts the carbon footprint.
- Aflibercept 8mg allows treatment in extended intervals without compromising efficacy or safety and therefore can contribute to a lower burden on the environment.



Study aim

This study aims to calculate the environmental impact of the reduced treatment regimen of aflibercept 8mg, marking the initial step towards incorporating carbon analyses into the purview of HTA bodies.

Discussion and conclusions

Main findings

- Our study shows the substantial impact of the treatment regimens of anti-VEGFs on CO2eq emissions.
- Switching from the 8-weekly regimen of aflibercept 2mg to the 16-week dosing regimen of aflibercept 8mg contributes to the reduction of the carbon footprint within the healthcare system by reducing the CO2eq emission with 35.9% and 43.5%.



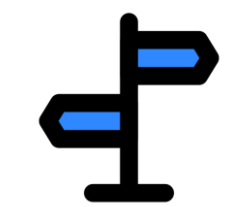
Discussion

- Analysis like this are an important first step in improving sustainability in healthcare.
- However, it is important to extend this analysis further by including the impact on medication and medication-related adverse events in the analysis.



Future steps

- To factor in medication's environmental impact in decision-making, we should create guidelines for measuring CO2eq emissions in treatments, evaluating environmental effects in cost-effectiveness analyses, and managing health-environment trade-offs.



Methods

Base case analysis

- We developed a model to estimate the CO2eq emissions related to treatment with aflibercept 2mg (using an 8-week dosing regimen) and aflibercept 8mg (using a 12-week [Q12] or 16-week [Q16] dosing regimen) in the total population of Dutch treatment-naïve nAMD and DME patients, over a two-year time horizon.
- The base-case analysis was performed based on the assumption that all treatment-naïve patients started with either aflibercept 2mg, aflibercept 8mg Q12, or aflibercept 8mg Q16.
- Input parameters were based on trial data and literature (Table 1). CO2eq emissions were based on travel, materials (i.e., production and waste), and energy usage.

Additional analyses

- In addition to the base analysis, we calculated the impact of a mix of aflibercept 2mg with aflibercept 8mg Q12 and aflibercept 8mg Q16 to simulate potential market shares.
- Moreover, we performed a sensitivity analysis in which we varied the injection frequency with a 25% interval.

Table 1. Overview of model inputs			
Parameter	Input		Source
Number of patients			
Incidence nAMD in the Netherlands	4,680		Zinnige zorg [3]
Incidence DME in the Netherlands	4,304		Calculated based on Vzinfo and Li et al. [4,5]
CO2eq emission per intravitreal injection, excluding anti-VEGF agent			
Travel	10.49 (77%)		Power et al. [2]
Materials	2.54 (19%)		
Energy usage	0.59 (4%)		
Number of injections			
	Year 1*	Year 2	
Aflibercept 2mg in nAMD	7.5	6.4	PULSAR trial [6]
Aflibercept 8 mg Q12 in nAMD	6.6	3.9	
Aflibercept 8 mg Q16 in nAMD	5.6	3.3	
Aflibercept 2mg in DME	8.6	6.4	PHOTON trial [7]
Aflibercept 8 mg Q12 in DME	6.5	3.8	
Aflibercept 8 mg Q16 in DME	5.4	3.0	
Note: *Including loading dose consisting of three, monthly injections			

Results

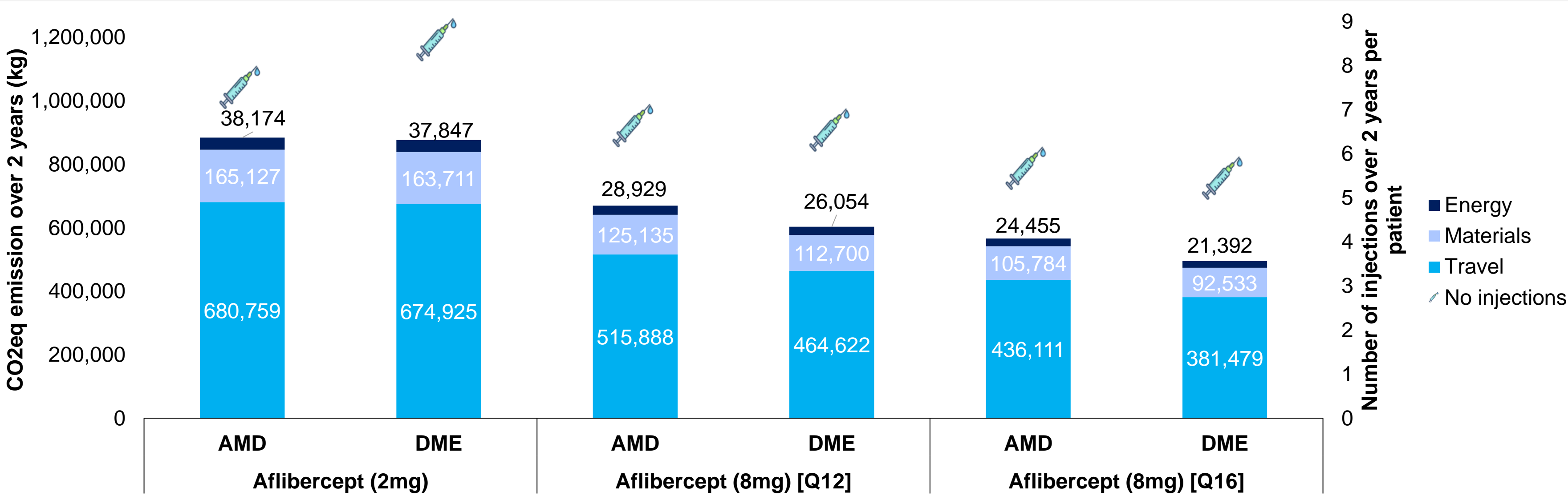


Figure 1. CO2eq emission over for total patient population over a 2-year time horizon and related number of injections (assuming a 100% market share)

Base case analysis

- Switching from aflibercept 2mg to aflibercept 8mg in treatment-naïve nAMD patients lowers the carbon emissions by 45.7 kg CO2eq (24.2%) and 63.5 kg CO2eq (35.9%) in a 12-week and 16-week dosing regimen, respectively (Figure 1).
- In treatment-naïve DME patients, the switch from aflibercept 2mg to aflibercept 8mg results in an even larger reduction, with carbon emissions decreasing by 67.9 kg CO2eq to 88.5 kg CO2eq (31.2% to 43.5%) (Figure 1).
- As expected, the CO2eq emission is directly related to the injection frequency.

Additional analyses

- Figure 2 shows the CO2eq emission with different market shares of aflibercept 2mg, aflibercept 8mg Q12, and aflibercept 8mg Q16. It shows a clear decrease towards a larger share of either regimens and how it decreases with a larger share of aflibercept 8mg.
- Figure 3 illustrates overlapping 25% distributions for the included treatments and regimens. The CO2eq emissions for aflibercept 2mg range from 132.3 to 237.1 kg, while those for aflibercept 8mg Q12 span from 107.8 to 179.7 kg, and for aflibercept 8mg Q16, they range from 91.1 to 151 kg CO2eq.

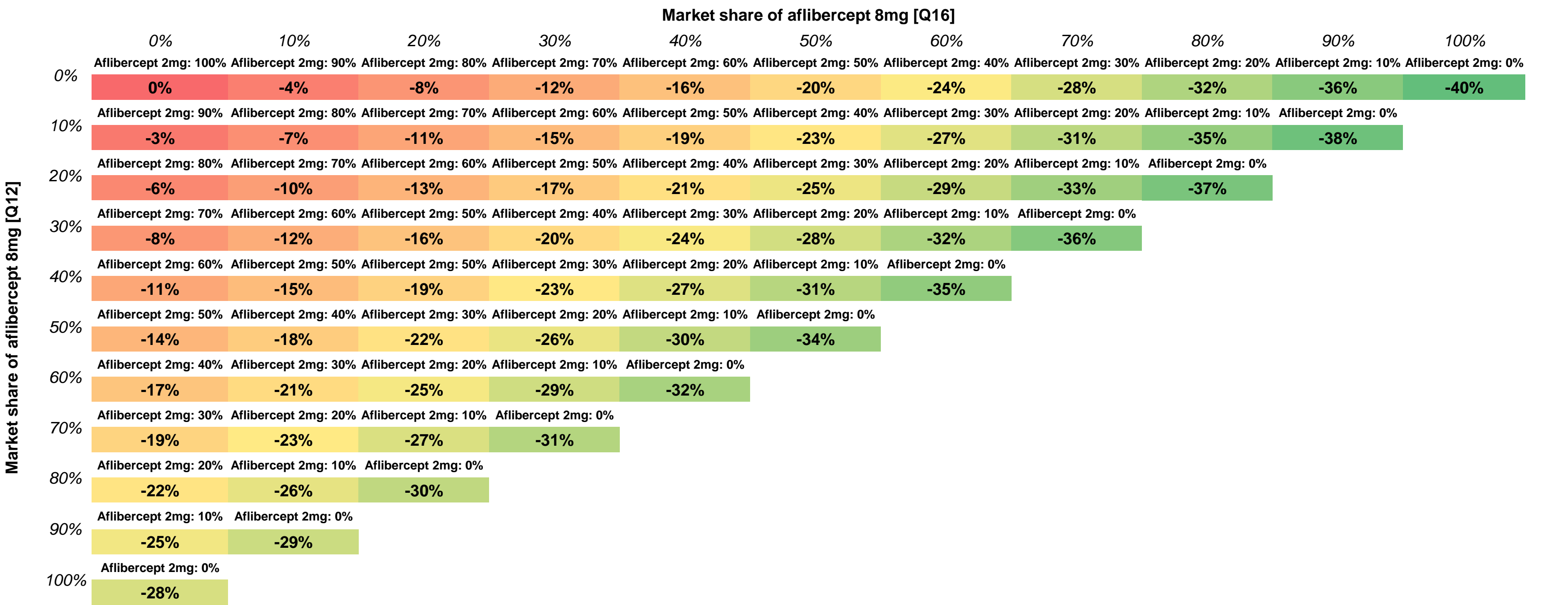


Figure 2. Two-way table presenting the reduction in CO2eq emission versus 100% aflibercept 2mg, using different market shares

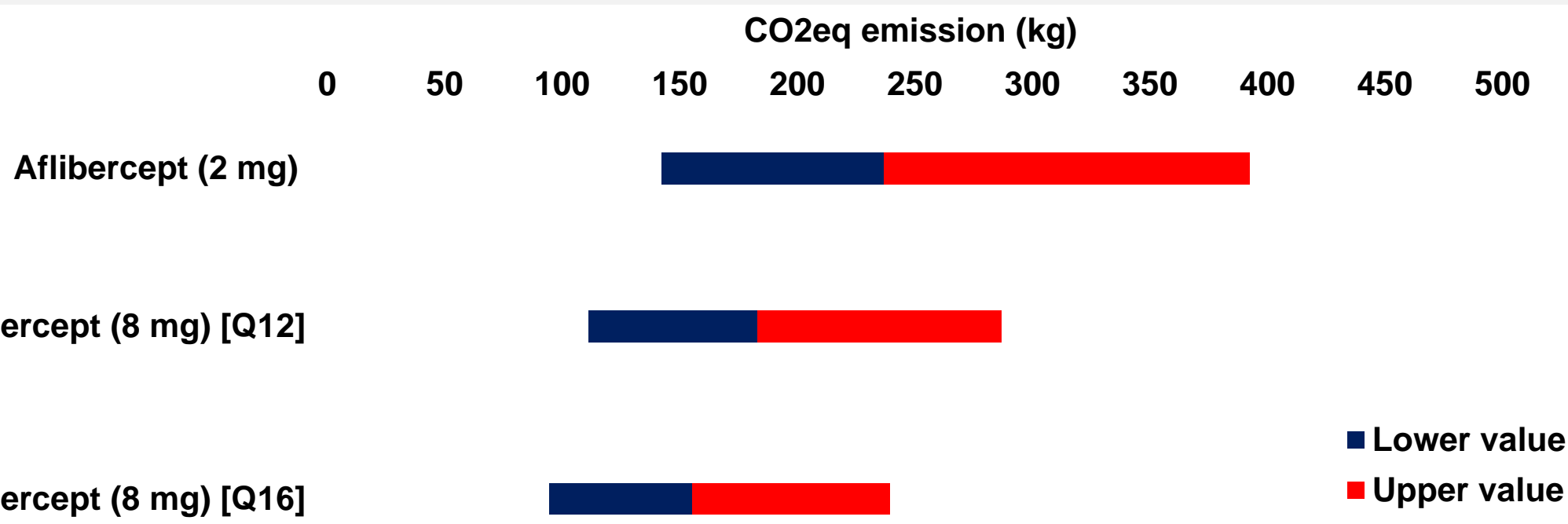


Figure 3. 25% sensitivity analysis of CO2eq emission (kg) of aflibercept 2mg and 8mg

References

- Dutch National institute for health and environment (2022): Het effect van de Nederlandse zorg op het milieu. Methode voor milieuoetadruk en voorbeelden voor een goede zorgomgeving
- Power B. et al. Analyzing the Carbon Footprint of an Intravitreal Injection. J Ophthalmic Vis Res. 2021 Jul 29;16(3):367-376.
- Dutch Healthcare Authority (2021). Zinnige Zorg - Verbetersignalement Leefstijdsgebonden maculadegeneratie [Sensible Care - Improvement Report Age-related macular degeneration]
- Vzinfo (2022) Diabetes mellitus
- Li, J.Q. et al. Prevalence, incidence and future projection of diabetic eye disease in Europe: a systematic review and meta-analysis. Eur J Epidemiol 35, 11–23 (2020)
- Power B. et al. Analyzing the Carbon Footprint of an Intravitreal Injection. J Ophthalmic Vis Res. 2021 Jul 29;16(3):367-376.
- Two-year pulsar trial results for aflibercept 8 mg demonstrate durable vision gains at extended dosing intervals in wet age-related macular degeneration (NCT: NCT04423718)
- Two-year results for aflibercept 8 mg from pivotal photon trial demonstrate durable vision gains at extended dosing intervals in diabetic macular Edema (NCT: NCT04429503)