Budget Impact Analysis of Cenobamate for Epilepsy Patients EE297 with Drug-Resistant Focal Onset Seizures in the Netherlands

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INTRODUCTION & OBJECTIVES

Epilepsy is one of the **most** prevalent chronic and severe neurological conditions, focal onset seizures (FOS) constitute the **most** common seizure type, representing up to 61% of the epilepsy population¹.

120 thousand

people has epilepsy **6** thousand

people are diagnosed per year



Several anti-seizure medications (ASM) have been developed for epilepsy treatment, however, 40% of patients with epilepsy are drug-resistant². The 3rd generation drugs were therefore launched.

| <u>RESULTS</u> | | | | | | | |
|---------------------------|--------------|--------------|--------------|---------------------|---------------|---------------|-----------------|
| Category | Year 1 | Year 2 | Year 3 | Year 4 | Year 5 | Total 5 years | |
| Cenobamate cost | € 733,039 | € 1,617,608 | € 2,520,196 | € 3,439,818 | € 4,375,646 | € 12,686,307 | |
| Incremental drug cost | € 151,697 | € 442,206 | € 738,784 | € 1,041,085 | € 1,348,824 | € 3,722,596 | |
| Subsequent treatment cost | € 47,261 | € 48,748 | € 50,182 | € 51,574 | € 52,931 | € 250,697 | 7 |
| Event management cost | -€ 843,257 | -€ 1,704,945 | -€ 2,583,956 | -€ 3,479,358 | -€ 4,390,374 | -€ 13,001,890 | Medical savin |
| Routine monitoring cost | -€ 76,754 | -€ 155,185 | -€ 235,193 | -€ 316 <i>,</i> 693 | -€ 399,614 | -€ 1,183,439 | €13,499,498 |
| Adverse event cost | € 65,741 | € 76,238 | € 86,882 | €97,671 | € 108,603 | € 435,135 | |
| Productivity cost | -€ 1,436,185 | -€ 2,903,762 | -€ 4,400,842 | -€ 5,925,838 | -€ 7,477,427 | -€ 22,144,054 | ➡Non-medical sa |
| TOTAL cost | -€ 2,214,206 | -€ 4,196,701 | -€ 6,344,143 | -€ 8,531,558 | -€ 10,757,058 | -€ 31,920,955 | ➡ Budget saving |

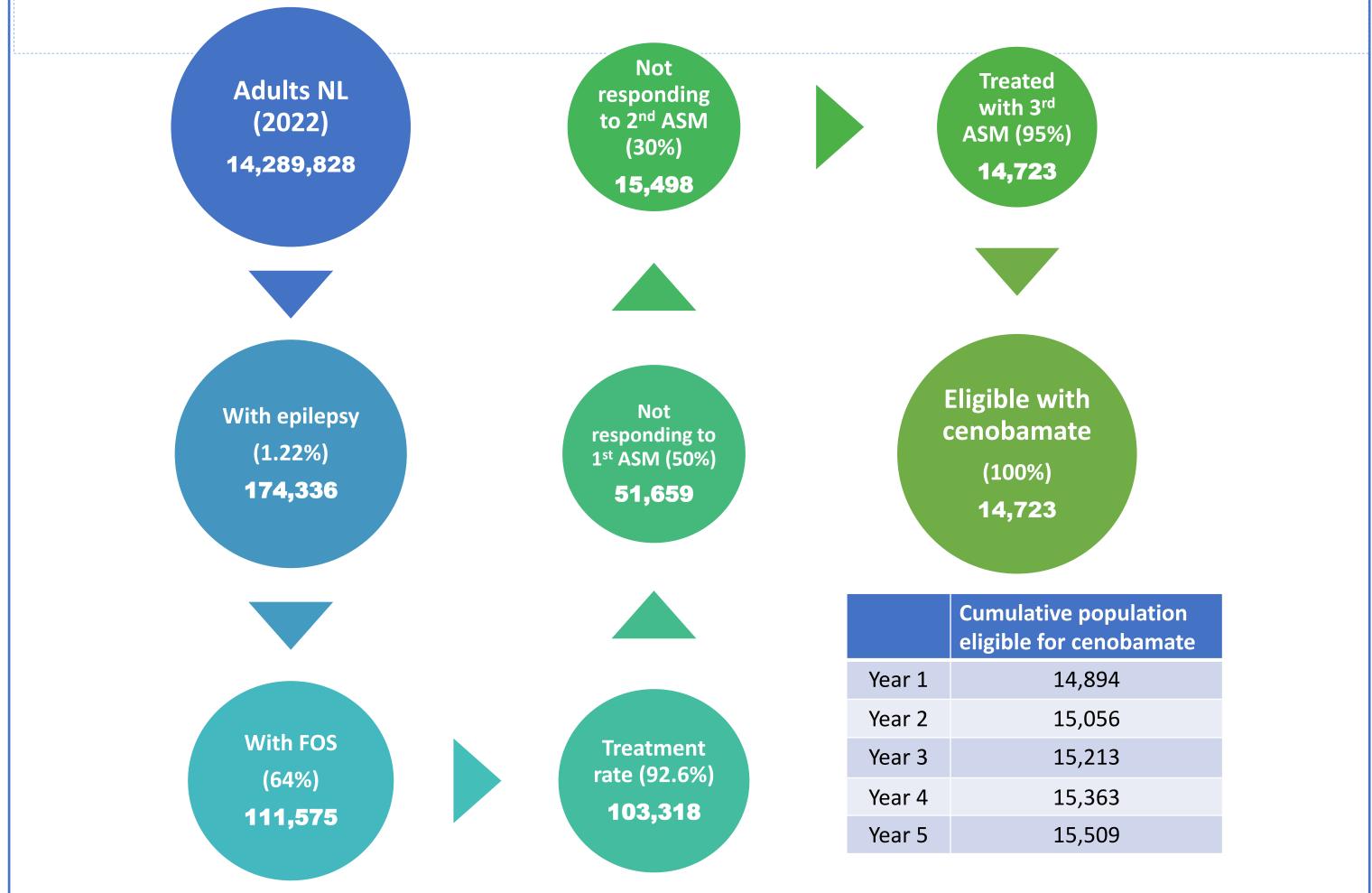
Although cenobamate adds a gross budget impact of €12,686,30, the Ο displacement of other drugs yields a total impact on the drug budget of

Cenobamate as a **novel therapy** was approved by the EMA in 2021 as **adjunctive treatment** of FOS with/without secondary generalization in adult epilepsy patients inadequately controlled despite a history of treatment with \geq 2 ASMs.

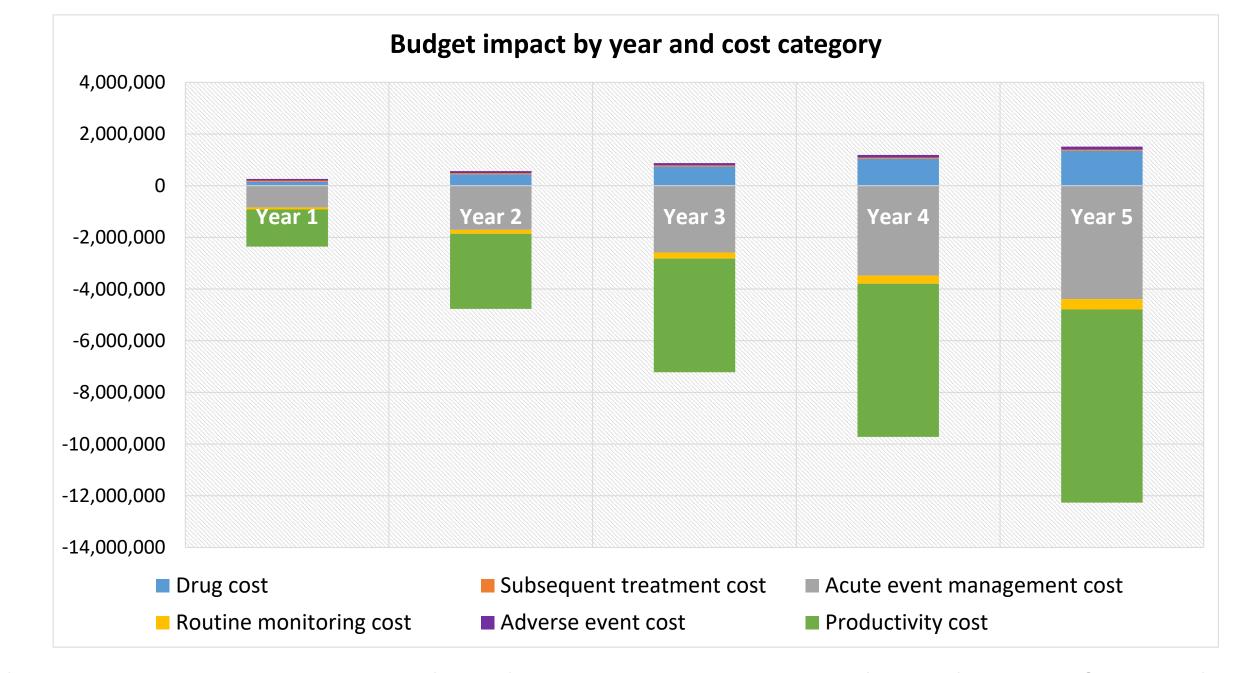
The clinical efficacy of cenobamate is well reported. The objective of this study was to explore the financial consequences of adopting cenobamate as a treatment alternative from a societal perspective in the Netherlands.

METHODS

A prevalence-based budget impact model³ was adapted to the Dutch setting with a **5-year** time horizon.

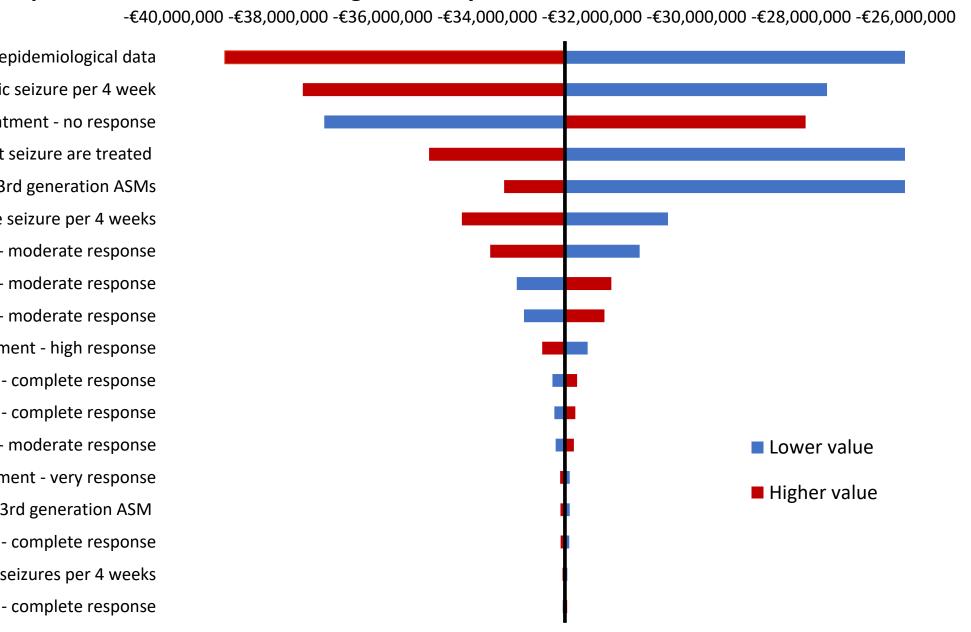


- €3,722,596 over 5 years;
- Adopting cenobamate resulted in a medical cost savings of €13,499,498 due to less resource use. Non-medical cost savings of €22,144,054 was associated with reductions in productivity due to focal epilepsy;
- Overall, savings generated at medical and non-medical cost level offset Ο the gross drug budget impact of cenobamate, resulting in a saving of €31,920,955 over 5 years.



The budget saving caused by the increase in market share of cenobamate was increasing overtime. Productivity and seizure event management costs are the top two contributors to the overall budget saving.

Impact on total health care budget over 5 years



The impact of changing epidemiological data Average number of focal to bilateral tonic-clonic seizure per 4 week Cenobamate proportion of patients responding to treatment - no response % patient with focal onset seizure are treated % patient treated with 3rd generation ASMs Average number of focal impaired aware seizure per 4 weeks Cenobamate proportion of patients responding to treatment - moderate response Lacosamide risk of treatment response relative to cenobamate - moderate response Brivaracetam risk of treatment response relative to cenobamate - moderate response Cenobamate proportion of patients responding to treatment - high response Brivaracetam risk of treatment response relative to cenobamate - complete response Lacosamide risk of treatment response relative to cenobamate - complete response Perampanel risk of treatment response relative to cenobamate - moderate response Cenobamate proportion of patients responding to treatment - very response % patient on 'other treatments' initiating a 3rd generation ASM Cenobamate proportion of patients responding to treatment - complete response Average number of focal aware seizureseizures per 4 weeks Perampanel risk of treatment response relative to cenobamate - complete response

The one-way sensitivity analysis confirms the robustness of our results.

CONCLUSION

Treatment with cenobamate is associated with both medical and non-medical cost savings, which offset the increase in drug budget and result in a significant

The model accounts for the eligible population, real-world market shares (from Kempenhaeghe & Maastricht UMC+), treatment effectiveness and resource use in two scenarios: cenobamate with constant market share versus cenobamate with **linearly increased** market share up to 20%.

| Scenario 1: Constant market share of cenobamate | | | | | | | | |
|---|---|------------|--------------|------------|-----------|-------|-------|--|
| Year | Current allocation of management treatments for epileptic patients with FOS | | | | | | | |
| | Cenobamate | Perampanel | Brivaracetam | Lacosamide | Resection | VNS | DBS | |
| Year O | 6.63% | 8.71% | 37.31% | 41.38% | 2.37% | 3.55% | 0.05% | |
| Year 1 | 6.63% | 8.71% | 37.31% | 41.38% | 2.37% | 3.55% | 0.05% | |
| Year 2 | 6.63% | 8.71% | 37.31% | 41.38% | 2.37% | 3.55% | 0.05% | |
| Year 3 | 6.63% | 8.71% | 37.31% | 41.38% | 2.37% | 3.55% | 0.05% | |
| Year 4 | 6.63% | 8.71% | 37.31% | 41.38% | 2.37% | 3.55% | 0.05% | |
| Year 5 | 6.63% | 8.71% | 37.31% | 41.38% | 2.37% | 3.55% | 0.05% | |
| Scenario 2: Linear increase in market share of cenobamate | | | | | | | | |

Proposed allocation of management treatments for epileptic patients with FOS Year

| | Cenobamate | Perampanel | Brivaracetam | Lacosamide | Resection | VNS | DBS |
|--------|------------|------------|--------------|------------|-----------|-------|-------|
| Year 0 | 6.63% | 8.71% | 37.31% | 41.38% | 2.37% | 3.55% | 0.05% |
| Year 1 | 9.30% | 8.45% | 36.17% | 40.12% | 2.37% | 3.55% | 0.05% |
| Year 2 | 11.98% | 8.18% | 35.03% | 38.85% | 2.37% | 3.55% | 0.05% |
| Year 3 | 14.65% | 7.91% | 33.89% | 37.58% | 2.37% | 3.55% | 0.05% |
| Year 4 | 17.33% | 7.65% | 32.74% | 36.32% | 2.37% | 3.55% | 0.05% |
| Year 5 | 20.00% | 7.38% | 31.60% | 35.05% | 2.37% | 3.55% | 0.05% |

Clinical inputs, such as treatment response, seizure reduction, and adverse events, were obtained from clinical trials. Costs for drugs, medical, and nonmedical expenses were sourced from national databases, MUMC+, and

| literature. | Drug | Medical | Non-medical | |
|---|---|--|---------------------------------------|--|
| | Cenobamte Perampanel Brivaracetam Lacosamide | Subsequent treatment Monitoring Seizure events Adverse events | Productivity loss | |
| Ioannou P, et al. Brain Behav. 20 Chen Z, et al. JAMA Neurol. 201 Vonck K, et al. Value in Health [| | Maastrich | nt Uni | |

potential budget saving of €32 million over 5 years in the Netherlands.

Limitation

- Not feasible to retrospectively obtain some resource use data, relevant estimations were based on Dutch expert opinion
- We conservatively assumed four neurologist visits per year for non-responders and two for responders, and additional three visits for treatment-related adverse event management
- Complex scenarios such as stopping and switching treatment, and the costs of pre-surgical evaluation were not taken into account
- The long-term efficacy and safety of cenobamate and its comparators remains uncertain

In the real-life setting, most medically refractory patients are on polypharmacy which makes it difficult to start cenobamate, and also some patients suffer far more comorbidities and are treated with multiple medications, all of these factors could interact with the safety and side effects of cenobamate which in our study is underestimated



