

Background

- Myasthenia gravis (MG) is an autoimmune disorder of the neuromuscular junction and is caused by auto-antibodies directed at the muscle acetylcholine receptors.^{1,2}
- MG is associated with a considerable burden of disease; patients are significantly impacted in both the short and long term, and many experience inadequate disease control, poor quality of life, and fixed muscle weakness.³⁻⁶
- In a previous longitudinal claim-based study in France, we reported that 34.6% of the 6,354 patients with incident MG were admitted to intensive care at least once, and 44.3% were treated with intravenous immunoglobulin during follow-up.⁷
- Understanding the progression of disability in MG is important in informing treatment strategies and improving patient care, but such data are scarce, particularly in patients with early-onset MG (onset at age <50 years).⁸⁻¹¹

Objective

- To analyze the progression of disability and associated costs in patients with incident early-onset MG compared with the overall cohort with incident MG.

Methods

Study design and data source⁷

- This was a retrospective, longitudinal cohort study using the French national health insurance claims database (SNDS) from January 2013 to December 2020 (**Figure 1**).
- The index date was the date of the first healthcare reimbursement claim relating to MG documented during the study period.
- Patients were followed from the index date to the end of the study period (31 December 2020) or until death.
- A 3-year historical period dating from 1 January 2010 was also searched for previous MG-related claims.

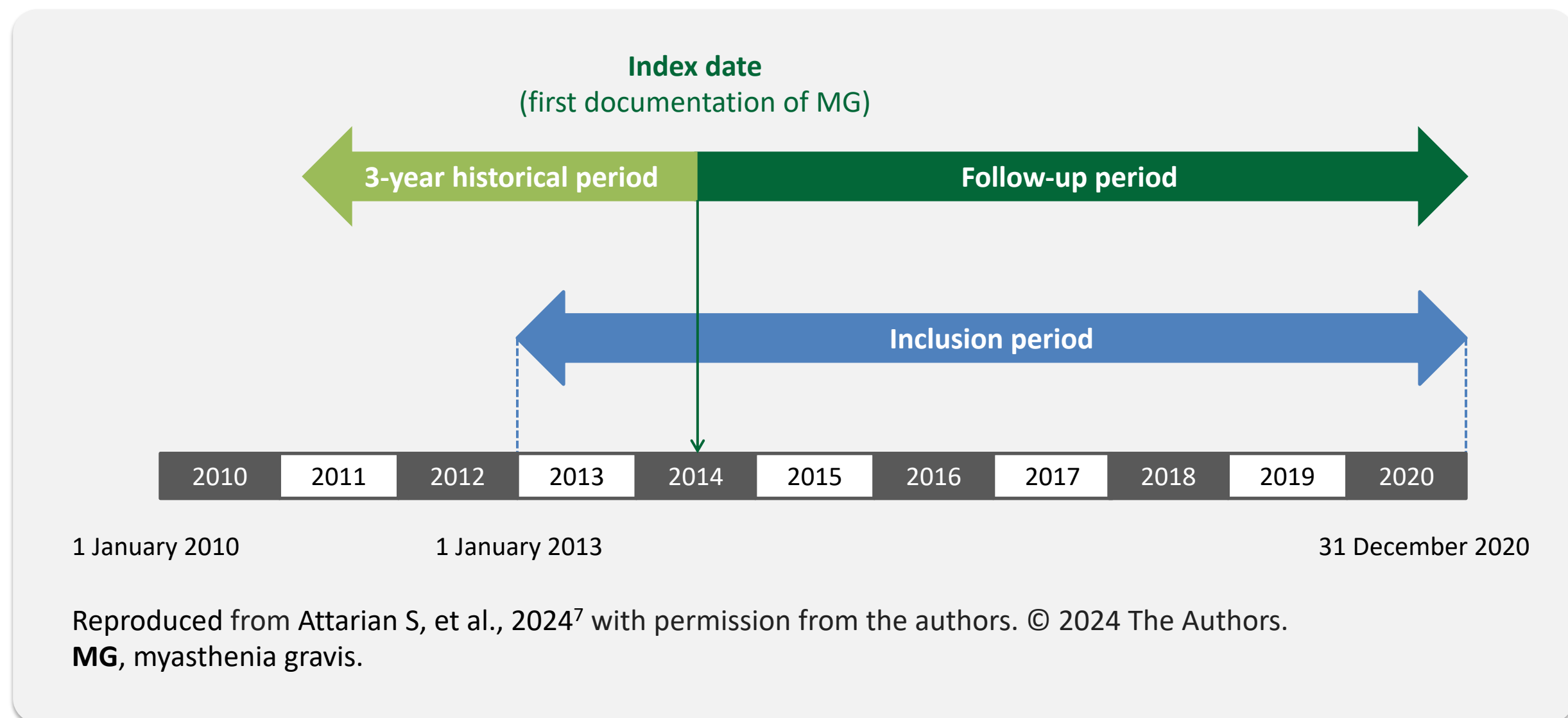


FIGURE 1. Study design⁷

Patient population

- Patients aged ≥18 years were included in the full study population according to the eligibility criteria previously described.⁷
- Claims for delivery of an acetylcholinesterase inhibitor prescribed by a gastroenterologist were excluded.⁷
- Patients with <12 months of follow-up were excluded.
- Incident patients were defined as all patients with a first MG-related claim during the inclusion period and no history of any MG-related claim during the historical period between 1 January 2010 and the index date.⁷
- Early-onset MG was defined as onset of MG before the age of 50 years.⁸⁻¹¹

Study outcomes

- Outcomes were assessed in the incident early-onset MG cohort and the overall incident MG cohort.
- Disability progression, as measured by disability status (eligibility for Disability Living Allowance), and use of sick leave were assessed.
- Costs (in Euros [€] at 2022 prices) associated with disability were evaluated over the follow-up period.

Statistical analysis

- Descriptive statistics were used to summarize the baseline characteristics and outcomes of the study populations.
- Predictors of disability progression were explored using a generalized estimating equation (GEE) model.

Results

Patient selection

- Among the 14,459 patients with MG included in the full study population, 6,354 patients had incident MG and 1,802 patients had incident early-onset MG (**Figure 2**).

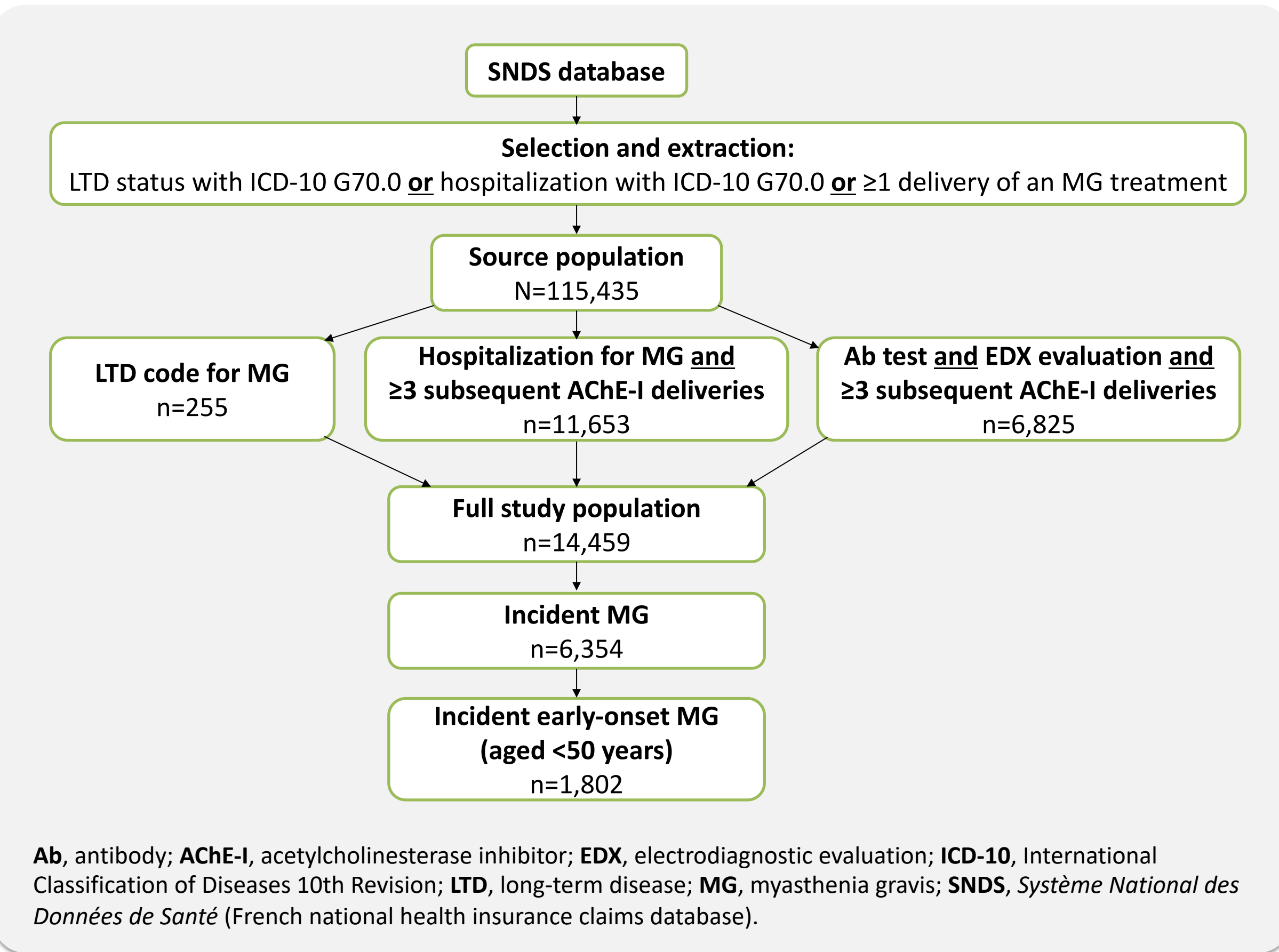


FIGURE 2. Patient flow

Baseline demographic and clinical characteristics

- Patients with incident early-onset MG were younger than those in the overall cohort with incident MG, and the proportion of female patients was higher (**Table 1**).
- Patients with incident early-onset MG had lower mean Charlson Comorbidity Index scores and had fewer comorbidities compared with the overall cohort with incident MG, including infections, asthma/chronic obstructive pulmonary disease, and cancer (**Table 2**).

TABLE 1. Baseline demographic characteristics

Characteristic	Incident early-onset MG (N=1,802)	Overall incident MG (N=6,354)
Age, years		
Mean (SD)	35.65 (8.94)	59.94 (18.34)
Median (IQR)	36 (28–44)	63 (47–74)
Distribution by age, n (%)		
18–40 years	1,149 (63.8)	1,149 (18.1)
41–65 years	653 (36.2)	2,310 (36.4)
>65 years	0 (0)	2,895 (45.6)
Sex, n (%)		
Male	611 (33.9)	4,424 (69.6)
Female	1,191 (66.1)	1,930 (30.4)
Follow-up		
Mean (SD), years	4.56 (2.19)	4.32 (2.23)

IQR, interquartile range; MG, myasthenia gravis; SD, standard deviation.

TABLE 2. Baseline clinical characteristics

Characteristic	Incident early-onset MG (N=1,802)	Overall incident MG (N=6,354)
Charlson Comorbidity Index^a		
Mean (SD)	0.30 (0.55)	2.73 (1.95)
0, n (%)	1,326 (73.6)	3,612 (56.8)
1–2, n (%)	466 (25.9)	2,461 (38.7)
3–4, n (%)	10 (0.6)	255 (4.0)
≥5, n (%)	0 (0)	26 (0.4)
Comorbidities, n (%)		
Infection	159 (8.8)	857 (13.5)
Depression	147 (8.2)	663 (10.4)
Anxiety	105 (5.8)	681 (10.7)
Asthma/COPD	95 (5.3)	601 (9.5)
Hypertension	90 (5.0)	1,505 (23.7)
Cancer	75 (4.2)	688 (10.8)
Cardiovascular disease	35 (1.9)	811 (12.8)

^aCharlson Comorbidity Index without age adjustment.

COPD, chronic obstructive pulmonary disease; MG, myasthenia gravis; SD, standard deviation.

Sick leave and disability

- Among patients with incident early-onset MG, 56.1% of patients took sick leave over the course of follow-up (39.8% occurring in the first year after diagnosis), and 16.5% transitioned to disability status (increasing from 4.1% in Year 1 to 16.7% in Year 5; data for overall follow-up not shown) (**Figure 3**).
- For the overall cohort with incident MG, 25.0% of patients took sick leave over the course of follow-up (18.8% in the first year), and 11.3% of the overall cohort transitioned to disability status (increasing from 5.6% in Year 1 to 11.0% in Year 5).

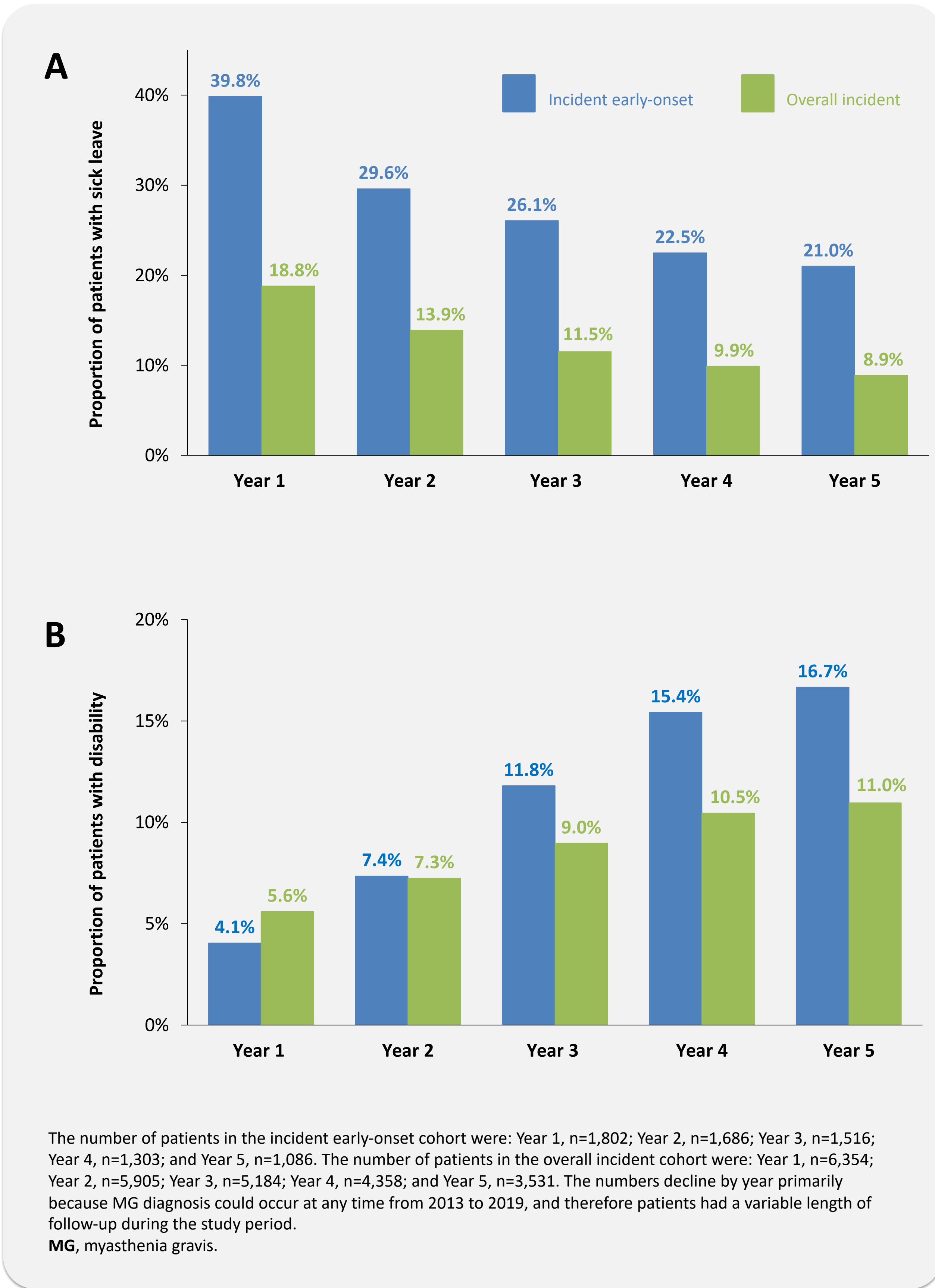


FIGURE 3. Evolution of (A) sick leave and (B) disability over time in incident early-onset MG vs overall incident MG

- Mean annual disability costs per patient in the incident early-onset cohort increased by 59% from Year 1 (€5,118) to Year 5 (€8,150; **Figure 4**).
- By contrast, mean annual disability costs per patient in the overall incident cohort increased by 39% from Year 1 (€5,432) to Year 5 (€7,563).

Conclusions

- Disability progression in incident early-onset MG patients is substantial, with rising costs over time and a notable shift from sick leave to disability.**
- Targeted, early interventions are essential to slow disability progression, optimize resource utilization, and improve long-term functional outcomes for this vulnerable patient group.**

Limitations

- Structured fields do not provide results specifying whether the electro-neuro-myogram result was normal or abnormal.
- The SNDS claims database does not provide medical information.

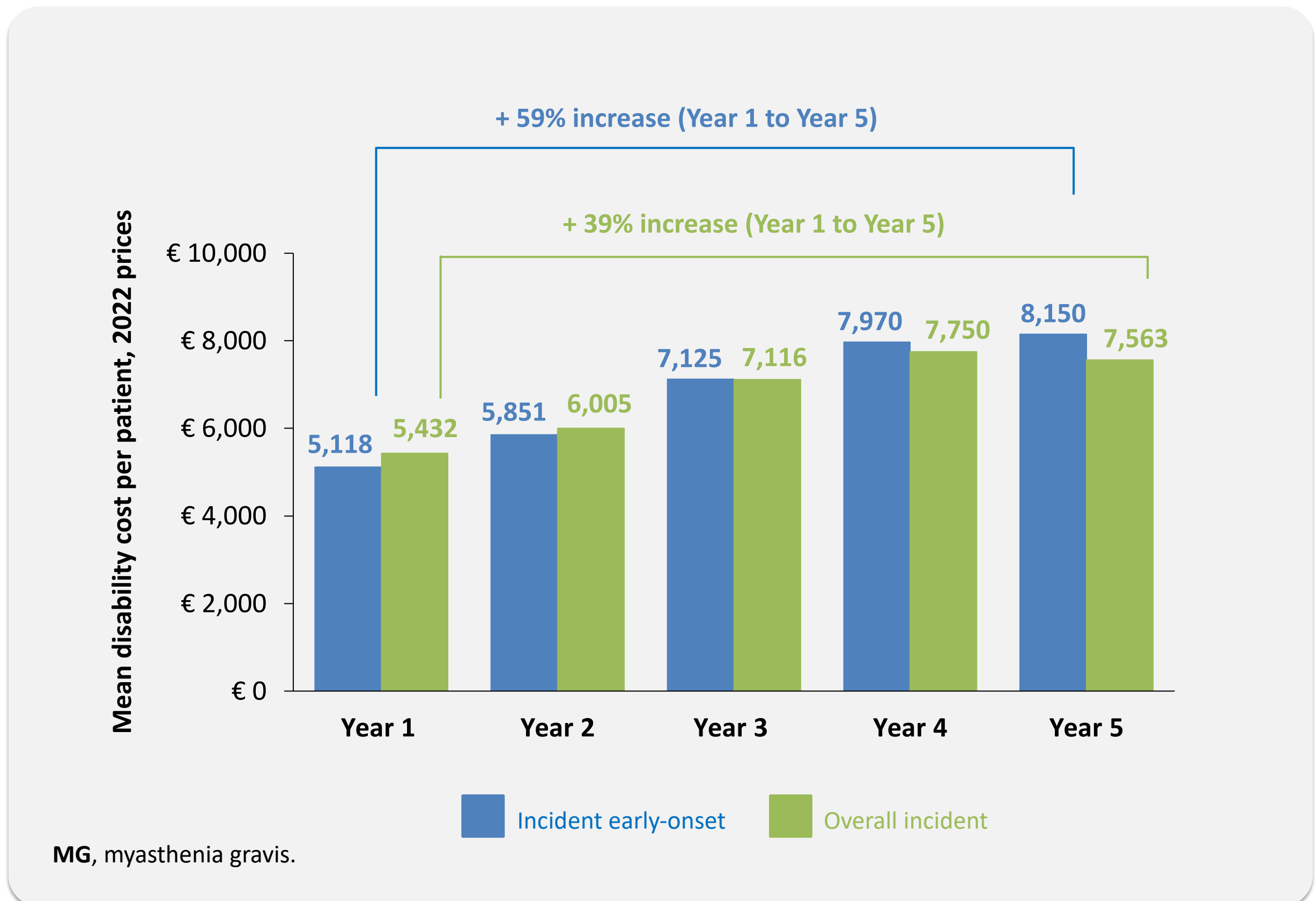


FIGURE 4. Evolution of mean costs of disability in incident early-onset MG vs overall incident MG

- Older age and longer duration of follow-up were significant drivers of disability progression (**Figure 5**).

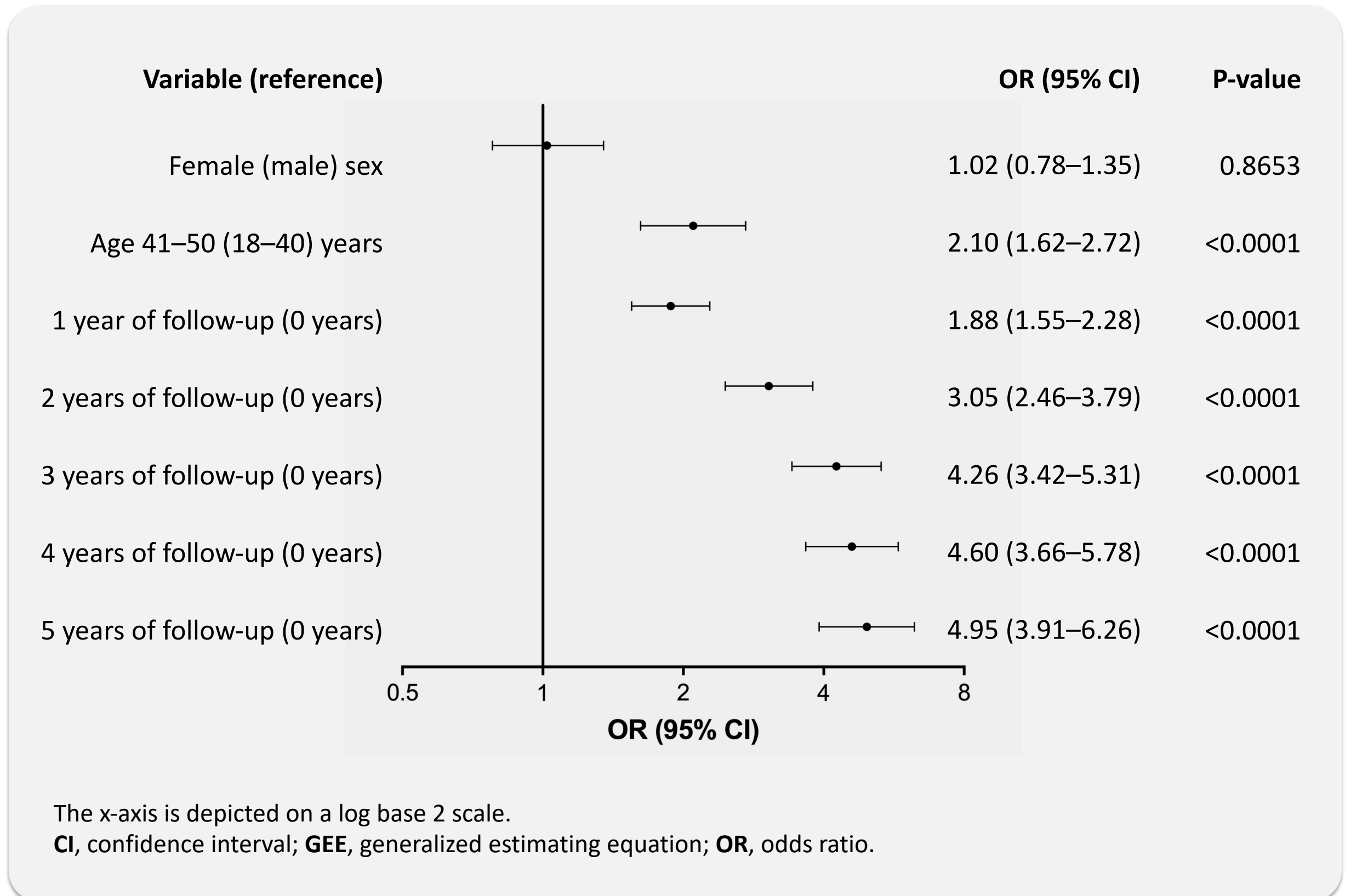


FIGURE 5. GEE model for disability progression

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Funding

This study was funded by Laboratoire argenx, Issy-les-Moulineaux, France.

Disclosures

CB and MC are employees of argenx. AE-I has received consulting and expert fees, lecture honoraria, support for participation in medical meetings, and honoraria for board participation from argenx. SA, J-PC, and GS have received honoraria from argenx for board participation or as a speaker at symposia. GS has received support from argenx to participate in medical meetings. GS has also received funding from argenx for the scientific policy of the French Society of Myology, paid to the French Society of Myology.

Acknowledgments

Medical writing support was provided by Mai Ping Tan, PhD, and Lisa Baker, PhD, of HEORpubs.