Phil McEwan¹, Martin Bøg², Mads Faurby³, Volker Foos¹, Ildiko Lingvay⁴, Christopher Lübker², Ryan Miller¹, Joshua Toliver³, Florian Yeates¹, A. Michael Lincoff⁵ ¹Health Economics and Outcomes Research Ltd., Cardiff, UK; ²Novo Nordisk A/S, Copenhagen Denmark; ³Novo Nordisk Inc., Plainsboro, NJ, USA; ⁴University of Texas Southwestern Medical Center, Dallas, TX, USA; ⁵Cleveland Clinic, Cleveland, OH, USA

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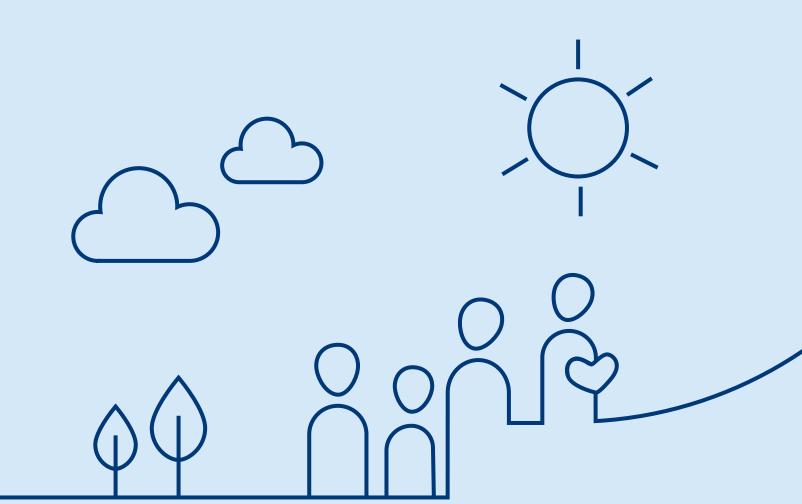
Treatment with semaglutide 2.4 mg was predicted to be cost-effective at a \$150,000 WTP threshold using US list price

6,091 fewer CV events per 100,000 subject

\$11,508

per subject savings due to delayed diabetes and avoided CV and CKD events \$140,512

ICER (\$/QALY)



Aim

SELECT is the only cardiovascular (CV) outcomes trial to demonstrate reductions in risk of major adverse CV events for a weight-management pharmacotherapy (semaglutide 2.4 mg) in addition to standard of care in subjects with overweight/obesity and CV disease without type 2 diabetes (T2D)[1]

We used SELECT data to assess the health economic impacts of participation in the SELECT trial beyond weight loss from a US payer perspective

Methods

- The cohort-level Markov state cost-effectiveness model (Figure 1) assesses clinical and health economic outcomes in the US setting for subjects with body mass index ≥ 27 kg/m² receiving onceweekly semaglutide (target dose 2.4 mg) compared with placebo, in addition to standard of care
- Baseline characteristics, treatment effects, adverse event rates, discontinuation, and survival equations were aligned to SELECT trial data and extrapolated to a lifetime horizon (39 years)
- Treatment costs were sourced from US list prices; costs and benefits were discounted at 3.0% per year. Cost-effectiveness was assessed against willingness-to-pay (WTP) thresholds described by the Institute for Clinical and Economic Review^[2] and the American Heart Association (AHA)/American College of Cardiology (ACC)[3]
- The average discount from list price for anti-obesity glucagonlike peptide-1 receptor agonists in the US has been estimated to be 41%;^[4] application of this discount to semaglutide 2.4 mg was explored in scenario analyses
- Probabilistic sensitivity analysis (PSA) sampled inputs from appropriate distributions around mean values over 1,000 iterations

MODEL STRUCTURE Alive states Diabetes stage Established CVD Secondary Tertiary CV events CV events MI HUA CR HHF **STROKE Death states*** CV Non-CV death death *Transitions to any death state can occur from any alive state **Abbreviations:** CKD, chronic kidney disease; CR, coronary revascularization; CV, cardiovascular; CVD, cardiovascular disease; HHF, hospitalization for heart failure, HUA, hospitalization for unstable

angina; MI, myocardial infarction

FIGURE 1: COST-EFFECTIVENESS

Key results

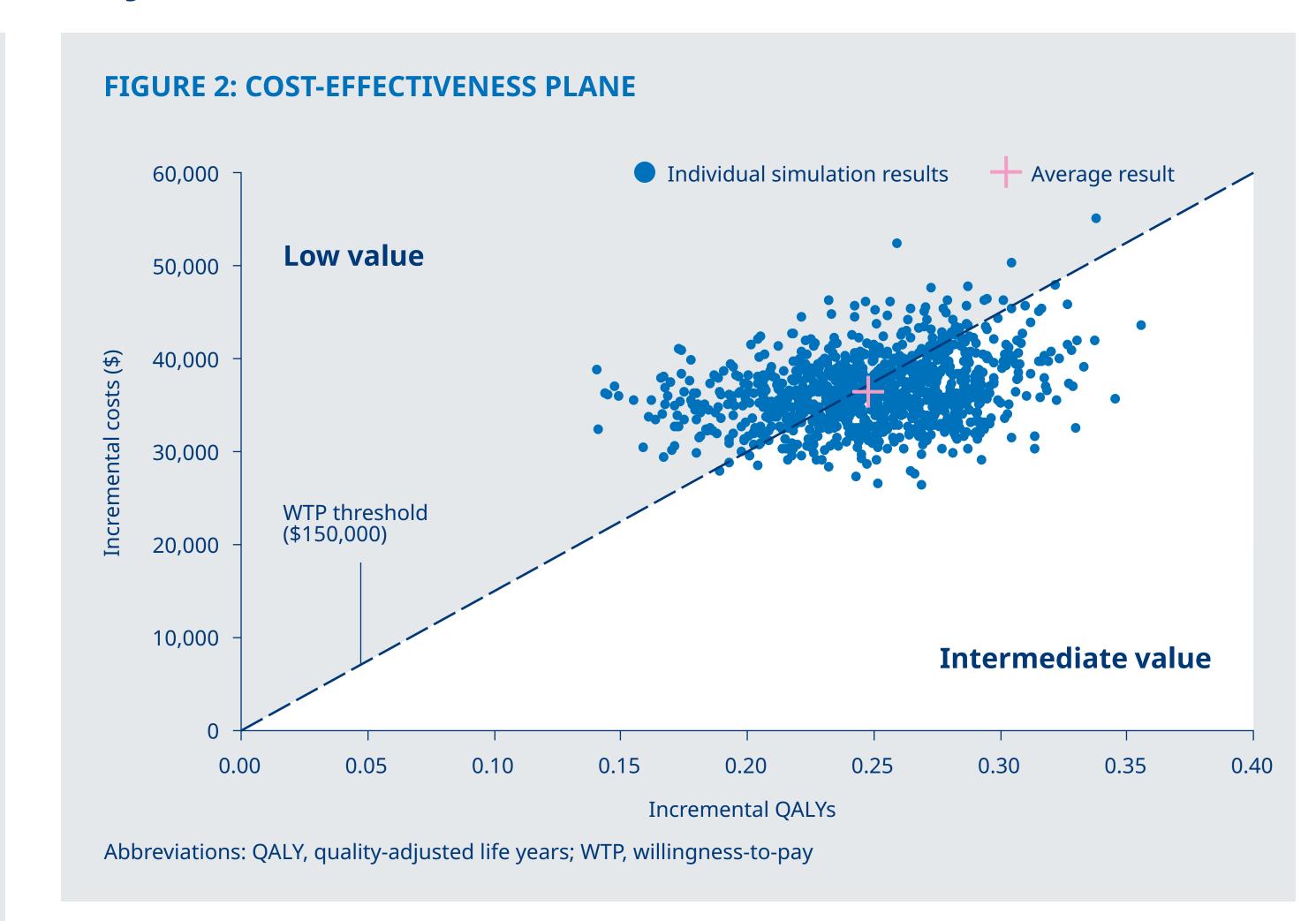


TABLE 1: COST-EFFECTIVENESS RESULTS (PER PATIENT)

ICER (\$/QALY)		\$140,512	\$64,514
Total QALYs	9.228	9.485	9.485
Total direct costs	\$203,367	\$239,509	\$219,961
Treatment costs [†]	\$0	\$47,679	\$28,130
Semaglutide 2.4 mg*	N/A	\$1,349	\$796
	Placebo	Base case (US list price)	Scenario (41% discount)

*Cost per 28 days,

[†]Mean semaglutide 2.4 mg treatment duration was 2.95 years Abbreviations: CV, cardiovascular; QALY, quality-adjusted life year

References: [1] Lincoff et al., New Engl. J Med. 2023;389(24):2221-2232. [2] https://icer.org/wp-content/uploads/2023/10/ ICER_2023_VAF_For-Publication_101723.pdf. [3] Anderson et al., Circulation. 2014;129(22):2229-2245. [4] Hernandez and Sullivan, Obesity (Silver Spring) 2024;32(3):472-475.

- Per 100,000 subjects, initiation of semaglutide 2.4 mg was predicted to avoid 6,091 total CV events: 2,464 non-fatal myocardial infarctions, 2,100 coronary revascularizations, 441 non-fatal strokes, and 505 CV deaths
- Initiation of semaglutide 2.4 mg treatment delayed the risk of progression to T2D
- Mean semaglutide 2.4 mg treatment duration was 2.95 years, resulting in per subject treatment costs of \$47,679. These were offset by cost savings associated with clinical events avoided, substantially driven by avoidance of T2D (\$8,835), chronic kidney disease (\$746), and CV events (\$1,926)
- Initiation of semaglutide 2.4 mg treatment was associated with 0.257 quality-adjusted life years (QALYs) gained, resulting in an incremental cost-effectiveness ratio (ICER) of \$140,512/QALY, i.e. cost-effective at a WTP threshold of \$150,000/QALY^[2]
- PSA indicated a 56.6% chance of cost-effectiveness at list price (Figure 2)
- In a scenario with a 41% discount on the list price of semaglutide 2.4 mg, the ICER decreased to \$64,514 (97.8% chance of cost-effectiveness at a \$100,000/QALY WTP threshold)

Summary

- Semaglutide 2.4 mg was predicted to be cost-effective at a \$150,000/QALY WTP threshold at US list price and at a \$100,000/ QALY WTP threshold at an estimated average discounted price
- Semaglutide 2.4 mg is likely to be an 'intermediate value' therapy according to the AHA/ACC value framework^[3]

Conclusion

- Treatment with semaglutide 2.4 mg represents a pivotal development in the clinical management of obesity and CV disease
- Treatment as per participation in SELECT was predicted to be cost-effective at a \$150,000 WTP threshold, forming an 'intermediate value' therapy at list price

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