

Cost-effectiveness of nirmatrelvir/ritonavir in Swedish adults, stratified by age, comorbidity and vaccination status, compared to no anti-viral treatment

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OBJECTIVE

Nirmatrelvir/ritonavir (NMV/r) is an antiviral agent indicated for adults at increased risk for progression to severe COVID-19, regardless of vaccination status, and is reimbursed in Sweden according to label. Both the pivotal clinical trial EPIC-HR [1] and real-world evidence (RWE) [i.a. 2,3,4] have shown that the NMV/r is effective in preventing hospitalization and death regardless of vaccination status and COVID-19 variant. This study assessed the cost-effectiveness of NMV/r for various risk groups, stratified by age, comorbidity, and vaccination status, compared to standard-of-care (SoC) in Sweden.

METHODS

A previously published and validated cost-effectiveness model was utilized and adapted to the Swedish setting [5]. The model used a short-term decision-tree (1 year) followed by a lifetime 2-state Markov model. The model accounted for symptom days, hospitalizations, intensive care unit admissions, quality of life, mortality, and treatment costs. It had a conservative approach by excluding the considerable, but still uncertain, costs related to post-acute covid syndrome. Overview of the model is shown in **Figure 1** and model inputs used in **Table 1**.

Figure 1. Model overview.

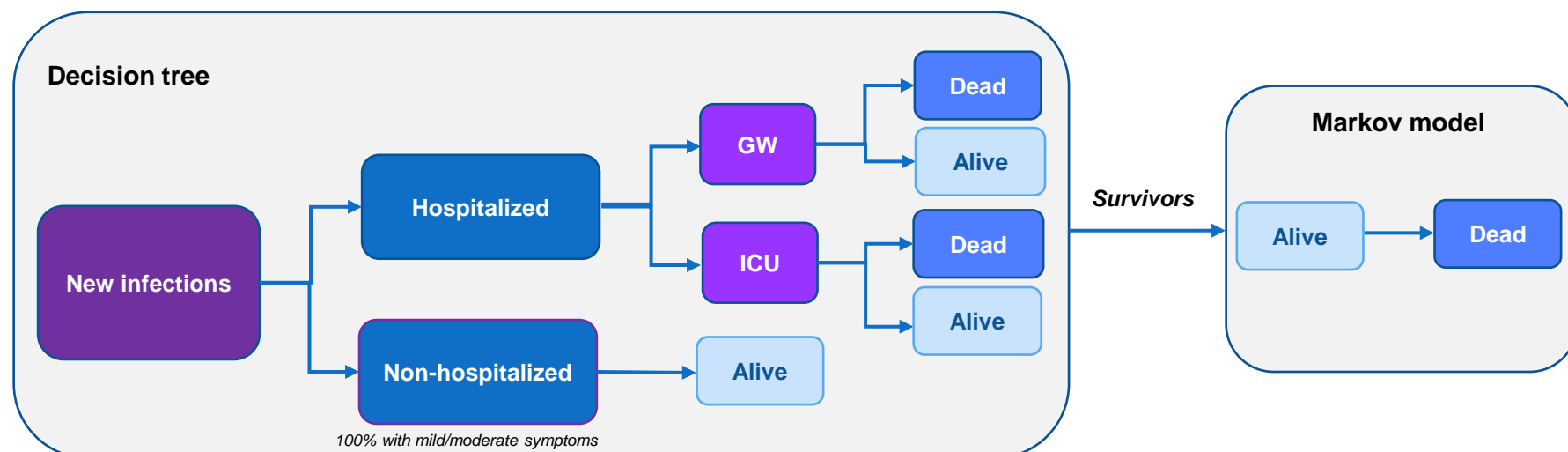


Table 1. Model inputs.

BASE CASE INPUTS	
Annual discount rate (costs & health benefits) [9]	3%
Medication cost per case – Paxlovid [10]	SEK 9,965
Cost per day at GW [11]	SEK 11,824
Cost per day at ICU [11]	SEK 71,473
Disutility, per day – Non-Hospitalized [12]	-0.290
Disutility, per day – Hospitalized [12]	-0.640
Disutility, total QALY loss – PACS [12]	-0.340
Proportion ICU [13]	2.40%
Proportion PACS in non-hospitalized [14]	5.7%
Proportion PACS in hospitalized [14]	17.5%
Symptom days in non-hospitalized [15]	6.87
Length of stay at GW (days) [16]	6
Length of stay at ICU (days) [16]	7
TREATMENT EFFECTIVENESS	
Reduction in number of hospitalizations and deaths, treatment within 5 days since the onset of symptoms [2]	79.6% (33.9-93.8)
Reduction in symptom days non-hospitalized [17]	20%
Reduction of length of stay, hospitalization [3]	30%
Reduction in proportion requiring ICU [3]	65%

The baseline absolute risks of hospitalization and death were collected from published data for 54 risk groups, unique in its granularity, detailing the risk for COVID-19 hospitalization and deaths in Sweden by age, vaccination status and comorbidity burden, based on nationwide data from the first quarter of 2022 [6]. The relative risk-reduction was set to 79.6%, based on RWE to reflect omicron vaccinated era [2]. Baseline utility for different age groups was sourced from Ara & Brazier [7].

Cost-effectiveness was assessed for a range of patient profiles (n=54) with different combinations of vaccination status, comorbidity profile and age. Vaccination status was categorized as 'vaccinated within past 180 days', 'vaccinated before past 180 days' and 'unvaccinated', comorbidity profiles assessed were 'no comorbidities', low- and high comorbidity status and age was varied in 10-year intervals from below 40 to above 80 years.

Quality-adjusted life years and costs were accumulated over the patients' life expectancy and the incremental cost-effectiveness ratio (ICER) was calculated. A willingness-to-pay threshold of SEK 500,000 per QALY was used to determine cost-effectiveness of NMV/r treatment. This is a commonly used threshold in Sweden for moderate conditions [8].

RESULTS

The ICERs for different patient profiles show significant variation, from almost nine million SEK to being dominant (i.e. cost saving with higher QALY gains vs SoC), see **Table 2**. NMV/r was a dominant treatment option in 12 risk groups. Utilizing a willingness to pay threshold of SEK 500,000 resulted in NMV/R being cost-effective in 30 risk groups. NMV/r was cost-effective in all patient groups older than 50 years with high comorbidity status. Only among patients younger than 40 years there were no risk groups in which NMV/r was cost-effective. NMV/r was cost-effective for all patients older than 70 years, except for those with no comorbidities that had been vaccinated in the prior 180 days.

Table 2. ICER for different patient profiles (in SEK): ICER below WTP level marked in bold

Vaccination status	Comorbidity burden	Age ^a					
		<40 years	40-49 years	50-59 years	60-69 years	70-79 years	80+ years
Vaccinated within past 180 days	none	7,659,802	6,028,560	9,003,026	4,432,556	580,257	53,463
	low	8,868,918	803,207	831,289	668,331	103,836	29,808
	high	8,146,790	156,407	140,263	44,585	16,588	3,062
Vaccinated before past 180 days	none	8,995,119	3,372,696	4,644,583	8,688,525	95,190	Dominant
	low	8,766,810	484,904	730,606	303,373	Dominant	Dominant
	high	8,300,464	8,223,517	104,539	3,397,720	Dominant	Dominant
Unvaccinated	none	5,720,717	2,505,151	1,675,717	303,373	Dominant	Dominant
	low	1,905,049	152,522	172,498	14,809	Dominant	Dominant
	high	6,083,939	16,006	20,142	Dominant	Dominant	Dominant

Notes:

ICER: Incremental cost-effectiveness ratio, SEK: Swedish krona, WTP: willingness to pay (500,000 SEK)

a. One specific age (not age range) had to be used to run the analysis, for <40 age 40 was used, for the intervals the midpoint, i.e. 45, 55, etc. was used, for 80+ age 85 was used to run the analysis.

CONCLUSION

This study suggests that utilization of nirmatrelvir/ritonavir is cost-effective compared to SoC for most patient groups, and especially for patients older than 70 years for whom it often is a cost-saving treatment. Even for several patient groups belonging to age cohorts as young as 40-49 years may NMV/r be considered a cost-effective treatment option.

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ACKNOWLEDGEMENTS

We acknowledge HEOR UK for their contributions to the model conceptualisation and development.

DISCLOSURES

This study was funded by Pfizer AB. C Gerdesköld, M Aldvén and F Nilsson are employed by Pfizer AB, Sweden. T Mugwagwa is employed by Pfizer Inc., UK.