

Cost-effectiveness of Rasagiline Compared With Other First-Line Treatment Options for Early Parkinson's Disease in the United States

Raymond A Farkouh,¹ Michele R Wilson,¹ Marcy L Tarrants,² Jane Castelli-Haley,² Christophe Armand³

¹RTI Health Solutions, Research Triangle Park, NC, United States;

²Teva Neuroscience, Kansas City, MO, United States;

³H. Lundbeck A/S, Paris, France

BACKGROUND

- Parkinson's disease (PD) is a common disease that affects approximately 1 million people in the United States (US).¹
- The incidence of PD is approximately 60,000 new cases per year.²
- Pharmacologic interventions available for PD include:
 - Levodopa (LD)
 - Dopamine agonists (DA)
 - Selective irreversible monoamine oxidase type-B inhibitor
 - Once-daily rasagiline mesylate (rasagiline)³
- Dyskinesias (involuntary body movements) are linked to poor quality of life⁴ and higher health care costs.^{4,5,6}

OBJECTIVE

- Postponing the appearance of dyskinesias could be an effective strategy for reducing costs and improving the quality of life of patients with PD.
- The purpose of this study was to evaluate the cost-effectiveness of initiating first-line treatment of early PD with once-daily rasagiline monotherapy compared with initiating treatment with ropinirole XL, pramipexole, generic ropinirole, or first-line LD.

METHODS

Model Structure

- Markov model
- 5-year time horizon
- 6-month cycles
- US managed care perspective
- Costs and outcomes discounted at 3% per annum

Input Parameters

Patient Characteristics

- Early PD requiring a pharmacologic intervention
- Hoehn and Yahr stage 1.5⁷
- Average age of 61⁷

RESULTS

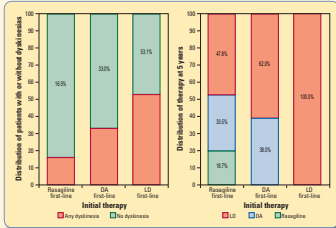
Base-Case Cost-Effectiveness Results

Table 4. Cost-effectiveness Results Over 5 Years of Early PD Treatment by First-Line Therapies

Result	Rasagiline	Generic Ropinirole	Rasagiline	Ropinirole XL	Rasagiline	Pramipexole	Rasagiline	LD First-Line Therapy
Total costs	\$82,339	\$82,099	\$82,339	\$90,468	\$82,821	\$83,336	\$85,566	\$88,666
Drug costs	\$8,040	\$3,862	\$11,267	\$12,921	\$8,521	\$8,188	\$11,267	\$3,909
Other resource costs	\$74,299	\$78,148	\$74,299	\$78,148	\$74,299	\$78,148	\$74,299	\$84,667
QALYs	3.45	3.32	3.45	3.32	3.45	3.32	3.45	3.21
Incremental costs		\$239		-\$4,902		\$516		-\$2,999
Incremental QALYs		0.13		0.13		0.13		0.24
Incremental cost per QALY		\$1,907						

QALY = quality-adjusted life-year.

Figure 2. Clinical Outcomes After 5 Years



Sensitivity Analysis Results

Figures 3 and 4 display sensitivity analyses of rasagiline as compared with generic ropinirole. Generic ropinirole was selected because it was the least cost-effective comparison. Results show that initiating treatment with rasagiline remained cost-effective or became cost-saving in nearly all sensitivity analyses. Probabilistic results show rasagiline was cost-effective (< \$50,000/QALY) in 66.6% of simulations.

Figure 3. One-Way Sensitivity Analysis Results: Rasagiline vs. Generic Ropinirole

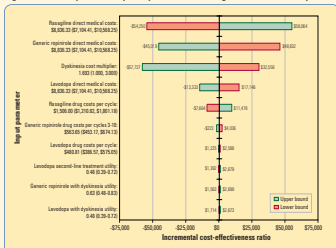
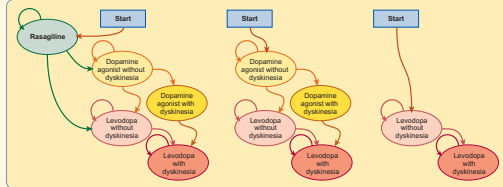


Figure 1. Markov Model: Early PD Treatment Pathways*



* All states may transition to death.

Transition Probabilities

Table 1. Health State Transition Probabilities (%)

Cycle	Rasagiline to DA	Rasagiline to LD	DA to LD	DA to DA With Dyskinesias	LD to LD With Dyskinesias	Any State to Death ¹⁰
1	4.03	2.42	10.20	2.74	7.37	1.22
2	15.05	7.77	10.20	0.87	6.82	1.28
3	20.00	7.33	10.20	1.09	6.10	1.34
4	13.21	2.83	10.20	3.09	6.75	1.41
5	11.63	5.81	10.20	0.68	11.14	1.47
6	14.93	4.48	10.20	5.73	2.19	1.53
7	18.87	3.77	10.20	3.16	11.54	1.59
8	5.56	0.00	10.20	7.79	10.51	1.65
9	3.23	9.68	10.20	5.18	9.72	1.72
10	7.69	7.69	10.20	2.01	7.62	1.78

Utility Weights

Table 2. Health State Utility Weights⁸

Utilities	VAS	SG
Rasagiline	0.83	0.85
DA	0.83	0.85
DA with dyskinesias	0.63	0.76
LD without dyskinesias, as first-line treatment	0.85	0.85
LD without dyskinesias, as second-line treatment	0.72	0.78
LD with dyskinesias	0.48	0.71

SG = standard gamble; VAS = visual analogue scale.

Figure 4. Probabilistic Sensitivity Analysis: Rasagiline vs. Generic Ropinirole

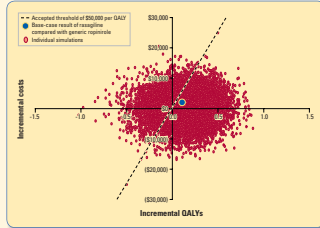


Table 5. Additional Sensitivity Analyses

Analysis Name	ICER Result
SG utility values	\$3,949
Rasagiline utility: lower bound	Generic ropinirole dominates
Rasagiline utility: upper bound	\$474
Generic ropinirole utility: lower bound	\$803
Generic ropinirole utility: upper bound	Generic ropinirole dominates
Transition probability: lower bound of DA to LD ¹	\$6,302
Transition probability: upper bound of DA to LD ¹	Rasagiline dominates
Transition probability: lower bound of DA without dyskinesias to DA with dyskinesias ¹	\$14,990
Transition probability: upper bound of DA without dyskinesias to DA with dyskinesias ¹	Rasagiline dominates
Transition probability: lower bound of LD without dyskinesias to LD with dyskinesias ¹	\$6,396
Transition probability: upper bound of rasagiline to DA ¹	\$5,966
Transition probability: upper bound of rasagiline to DA ¹	Rasagiline dominates
Transition probability: lower bound of rasagiline to LD ¹	\$176
Transition probability: upper bound of rasagiline to LD ¹	\$5,140

¹Lower bound calculated from the intent to treat population, where 82 out of 179 patients received LD¹. ²Upper bound calculated by assuming that all patients who withdrew prematurely from the clinical trial required LD; 94 withdrew and 86 received LD out of 179 patients over 5 years.¹

Costs

Health State Costs

- We assumed all nondyskinetic health states had the same nonpharmacy direct medical costs of \$8,836.¹¹
- Total nondyskinetic nonpharmacy direct medical costs were multiplied by 1.679 to obtain dyskinetic nonpharmacy direct medical costs of \$14,836.⁴

Pharmaceutical Costs

Table 3. Pharmaceutical Costs per 6-Month Cycle⁸

Treatment	Dosing	Cost per Cycle
Rasagiline ⁸	1 mg once daily	\$1,506.00
DAs		
Ropinirole XL¹³		
Cycle 1	8 mg per day	\$1,171.50
Cycle 2	12 mg per day	\$1,757.25
Cycles 3-10	16 mg per day	\$2,343.00
Pramipexole¹⁴		
Cycle 1	1.5 mg per day	\$410.02
Cycle 2	2.25 mg per day	\$615.03
Cycles 3-10	3 mg per day	\$820.04
Generic ropinirole¹⁵		
Cycle 1	9 mg per day	\$281.83
Cycle 2	12 mg per day	\$375.77
Cycles 3-10	18 mg per day	\$563.65
Coformulated carbidopa/LD ¹⁴	100 mg/400 mg per day	\$480.81

CONCLUSIONS

- Initiating treatment of early PD with rasagiline:
 - is predicted to be a cost-effective strategy when compared with initiating treatment with generic ropinirole
 - is predicted to be a cost-saving strategy when compared with initiating treatment with ropinirole XL, pramipexole, or LD
- Reduces the appearance of costly and uncomfortable dyskinesias by a relative 5% and 69% when compared with initiating therapy with a DA or LD, respectively, thus improving patients' quality of life.

REFERENCES

1. Olanow CW, Koller WC. An algorithm (decision tree) for the management of Parkinson's disease: treatment guidelines. *Neurology* 1998;50:S1-S7.
2. Aetlic (prescribing information). Available at: <http://www.aetlic.com/PrescribingInformation.pdf>. Accessed March 24, 2009.
3. Palmer CS, Schimer JK, Snyder E, Scott B. Patient preferences and utilities for off-time outcomes in the treatment of Parkinson's disease. *Qual Life* 2009;8:19-27.
4. Dodel RC, Singer M, Kohne-Welland R, Sauer T, Rathay B, Sobilo E, et al. The economic impact of Parkinson's disease: an estimation based on a 3-month prospective analysis. *Pharmacoeconomics* 1999;14:289-312.
5. Lefevre C, Wau S, Meunier-Marin E, Djurdjic M, Ziegler M. Cost of illness and disease severity in a cohort of French patients with Parkinson's disease. *Pharmacoeconomics* 1995;16:89-98.
6. Phechiev M, Clarke CE, Vemugopu P, Khakhriaeva B, Deschasseaux-Vincent B, Berbeaux G, et al. Effects of dyskinesias in Parkinson's disease on quality of life and health related costs: a prospective European study. *Eur J Neurol* 2005;12:956-62.
7. Hauser RA, Lew MF, Hurlig H, Ono WS, Wojcikzak J, Fitzer-Atlas C, et al. on behalf of the TEMPO Open Label Study Group. Long-term outcome of early versus delayed rasagiline treatment in early Parkinson's disease. *Mov Disord* 2009;24:52-71.
8. Rasoul O, Brooks DJ, Korczyn AD, De Deyn PP, Clarke CE, Lang AE. A five-year study of the incidence of dyskinesia in patients with early Parkinson's disease who were treated with ropinirole or levodopa. *096 Study Group*. *N Engl J Med* 2003;342:1484-91.
9. Clarke CE. Does levodopa therapy delay death in Parkinson's disease? A review of the evidence. *Mov Disord* 1995;10:50-4.
10. Kung H, Hoyert DL, Xu J, Murphy SL. Deaths: final data for 2006. *National Vital Statistics Reports* 56(10). Hyattsville, MD: National Center for Vital Statistics; 2008.
11. Orsini L, Castelli-Haley J, Kennedy S, House D. Healthcare utilization and expenditures among privately insured patients with Parkinson's disease in the U.S. Poster presented at the 8th Annual Meeting of the Movement Disorder Society's International Congress, Rome, Italy, June 13-17, 2004.
12. Red Book[®] for Windows, Version 6122. Vol. 50. Montvale, NJ: Thomson PDR; 2008.
13. Requip XL (prescribing information). Available at: http://us.gsk.com/products/astellas/us_requip.pdf. Accessed March 24, 2009.
14. Mirapex. Mirapex prescribing information 2009. Available at: <http://dbsdocs.bonhingtoningheim.com/BWebAccess/VWebServlet?docId=ras-remets&folderPath=Prescribing-Information/Ph/Mirapex/Mirapex>. Accessed September 18, 2009.
15. Hoerger TJ, Bala MV, Rowland C, Greer M, Christilles EA, Holloway RG. Cost effectiveness of pramipexole in Parkinson's disease in the US. *Pharmacoeconomics* 1996;14:541-57.

CORRESPONDING AUTHOR

Raymond Farkouh
Associate Director, Health Economics
RTI Health Solutions
200 Park Offices Drive
Research Triangle Park, NC 27709 United States
Telephone: +1.919.541.7322
Fax: +1.919.541.7222
E-mail: rfarkouh@rti.org

Presented at: ISPOR 12th Annual European Congress
October 24-27, 2009
Paris, France