Three-year outcome of rivastigmine treatment in Alzheimer’s disease in a routine clinical setting.

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Three-Year Outcome of Rivastigmine Treatment in Alzheimer’s Disease in a Routine Clinical Setting

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Conclusions
Long-term rivastigmine treatment in AD yielded positive cognitive, global and functional outcomes in a routine clinical setting. Completers of the 3-year study tolerated higher doses of rivastigmine better than did the patients who dropped out from the study.

Background and objectives
Alzheimer’s disease (AD) is the most common diagnosis among the dementia disorders. AD is characterized by a progressive decline in cognitive and practical abilities and leads to major difficulties in the management of daily life after only a few years of illness. Currently, the main therapy used for mild-to-moderate AD is cholinesterase inhibitor (ChEI) treatment, which may delay disease progression.

Methods
The Swedish Alzheimer Treatment Study (SATS) is a prospective, open, non-randomized and multicentre study for evaluating ChEI therapy in clinical practice. In total, 269 outpatients with a clinical diagnosis of AD received rivastigmine. Patients were assessed using cognitive tests (MMSE and ADAS-cog), global performance (CIBIC) and instrumental ADL (IADL) at baseline, after 2 months (MMSE and CIBIC only) and every 6 months, for a total period of 3 years. The 2-year outcomes of this study have been published previously [1]. The outcomes were compared with mathematical models of change in ADAS-cog [2] and IADL scores [3, 4] in untreated patients with AD. These models take into consideration the rivastigmine-treated SATS patient scores at baseline. The rates of change were calculated for each individual and illustrated in the figures. The expected change in MMSE score based on earlier reported untreated historical patients AD was estimated at 2–4 points/year [5, 6]. Three groups of response were defined at each CIBIC interval: 1–3 indicated improvement, 4 indicated no change and 5–7 indicated worsening.

Results
The mean ± SD dose of rivastigmine was 6.2 ± 2.1 mg per day during the study. Three-year completers (n = 117, 44%) received higher mean doses than did drop-outs (7.2 ± 1.9 vs 5.5 ± 2.0 mg/day, p < 0.001).

Table 1. Baseline characteristics

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Mean ± SD</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sex (females)</td>
<td>48% ± 52%</td>
</tr>
<tr>
<td>Age at onset, years</td>
<td>71.1 ± 8.0</td>
</tr>
<tr>
<td>Duration of AD, years</td>
<td>2.9 ± 2.2</td>
</tr>
<tr>
<td>Education, years</td>
<td>9.4 ± 2.5</td>
</tr>
<tr>
<td>MMSE score, range 30–0</td>
<td>23.0 ± 4.3</td>
</tr>
<tr>
<td>ADAS-cog score, range 0–70</td>
<td>18.1 ± 8.8</td>
</tr>
<tr>
<td>IADL score, range 30–0</td>
<td>14.8 ± 5.2</td>
</tr>
</tbody>
</table>

*mean ± standard deviation (SD)

ADAS-cog, Alzheimer’s Disease Assessment Scale-cognitive subscale; IADL, Instrumental Activities of Daily Living; MMSE, Mini-Mental State Examination

Among the completers, 26% of patients exhibited global improvement or no changes after 3 years of rivastigmine treatment.

The mean change in MMSE score from the baseline remained above the baseline level for 6 months. After 3 years of rivastigmine treatment, the mean decline of MMSE score from the baseline was 2.8 points (95% CI, 2.1–3.6) (shaded area, 2–4 points/year: expected decline in untreated AD patients).

The mean change in MMSE score after 3 years of rivastigmine treatment was 5.7 points (95% CI, 3.7–7.7), which was significantly better than the score predicted by the Stern equation had these SATS patients remained untreated (13.1 points; 95% CI, 11.9–14.2, p < 0.001).

The mean change in ADAS-cog score after 3 years of rivastigmine treatment was 5.7 points (95% CI, 3.7–7.7), which was significantly better than the score predicted by the Green equation had these SATS patients remained untreated (13.1 points; 95% CI, 11.9–14.2, p < 0.001).

The mean change in IADL score after 3 years of rivastigmine treatment was 6.2 points (95% CI, 5.2–7.2), which was significantly better than the score predicted by the Green equation had these SATS patients remained untreated (17.6 points; 95% CI, 16.7–18.5, p < 0.001).

Figure 1. MMSE: mean change (95% CI) from baseline

Figure 2. ADAS-cog: mean change (95% CI) from baseline

Figure 3. CIBIC

Figure 4. IADL: mean change (95% CI) from baseline

References:
6. Vellas, B.J., Mann, J., Bertrand, X., Andrieu, S., Carney, D., de Blay, C., and Schröder, C. Mean ± standard deviation (SD)

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