Cyclooxygenase-2 (COX-2) inhibitors are a group of nonsteroidal anti-inflammatory drugs (NSAIDs) that may reduce the incidence of gastrointestinal side effects associated with nonselective NSAIDs. The COX-2 inhibitors celecoxib (Celebrex) and rofecoxib (Vioxx) were first listed on the Ontario Drug Benefit (ODB) formulary on Apr. 17, 2000, as “limited use” products. Limited use means that these products are covered as a benefit only if the prescribing physician completes a form indicating that previous NSAID therapy failed or was not tolerated by the patient or that the patient has a documented history of clinically significant gastrointestinal bleeding or ulcer. This paper provides information on the impact of the ODB listing of COX-2 inhibitors on the costs and quantity of overall NSAID prescriptions and describes some characteristics of patients who received COX-2 inhibitors as compared with those who received nonselective NSAIDs. The analysis uses claims routinely submitted by pharmacies for all prescription drugs provided under the universal ODB plan for elderly patients in Ontario, physician claims submitted to the Ontario Health Insurance Plan (OHIP) and census data.

We used ODB claims data for individuals aged 65 years and older to describe drug use and costs before and after the listing of COX-2 inhibitors. The combined number of prescription claims for COX-2 inhibitors and nonselective NSAIDs increased 68% between March and November 2000, from 44 to 74 per 1000 eligible individuals. Combined monthly expenditures on NSAIDs increased 153% over this period, from $2.3 million to $5.9 million. Time series analysis showed that COX-2 inhibitors were responsible for this growth (Fig. 1). COX-2 inhibitors accounted for 48% of prescriptions and 66% of expenditures for NSAIDs in the post-listing period (Apr. 18 to Nov. 30, 2000).

We performed a patient-level analysis of a cohort of individuals aged 70 years and older dispensed NSAIDs from Apr. 17, 2000, through July 17, 2000, using ODB claims for the preceding 5-year period to identify previous NSAID use. In addition, we used OHIP claims over the same period to identify previous gastrointestinal endoscopy or upper gastrointestinal radiological examinations as a marker of earlier significant upper gastrointestinal events or ulcers. Users of 3 categories of prescription NSAIDs (COX-2 inhibitors, diclofenac sodium and misoprostol, and other nonselective NSAIDs) were defined on the basis of an index
prescription for one of the 3 categories within 90 days following the listing of COX-2 inhibitors. We analyzed differences between groups using the Kruskall–Wallis test for continuous variables and the χ² test for categorical variables.

About three-quarters of the initial cohort of COX-2 inhibitor users did not have a prescription claim for a nonselective NSAID in the 120 days before their first COX-2 inhibitor prescription, and nearly 20% did not have a prescription for a nonselective NSAID in the previous 5 years. The COX-2 inhibitor users had a similar age distribution to that of the cohort using nonselective NSAIDs. A recent cost–effectiveness analysis concluded that COX-2 inhibitors are cost-effective relative to nonselective NSAIDs when used by very elderly patients (P.R.), Toronto, Ont.

The substantial impact of COX-2 inhibitors on drug expenditures combined with the need for their cost-effective use (by those who are most likely to benefit) will be an important challenge to drug-plan managers and policymakers. Further research on the clinical characteristics of COX-2 inhibitor users and the effect of COX-2 inhibitor use is warranted.

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References


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Table 1: Characteristics of patients aged ≥ 70 years who received different categories of NSAIDs in the 90 days following the initial listing of COX-2 inhibitors in Ontario

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>COX-2 inhibitors n = 37 690</th>
<th>Diclofenac sodium and misoprostol n = 20 893</th>
<th>Nonselective NSAIDs n = 30 453</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean age (and SD), yr</td>
<td>77.6 (5.7)</td>
<td>77.9 (5.9)</td>
<td>77.3 (5.7)</td>
<td>&lt; 0.01</td>
</tr>
<tr>
<td>Female</td>
<td>27 731 (73.6)</td>
<td>13 374 (64.0)</td>
<td>18 800 (61.7)</td>
<td>&lt; 0.01</td>
</tr>
<tr>
<td>At least 1 prescription for an NSAID in past 120d</td>
<td>9639 (25.6)</td>
<td>14 824 (71.0)</td>
<td>21 314 (70.0)</td>
<td>&lt; 0.01</td>
</tr>
<tr>
<td>At least 1 prescription for an NSAID in past 5 yr</td>
<td>30 797 (81.7)</td>
<td>19 516 (93.4)</td>
<td>28 882 (94.8)</td>
<td>&lt; 0.01</td>
</tr>
<tr>
<td>Upper GI diagnostic exam in past 5 yr</td>
<td>10 883 (28.9)</td>
<td>4215 (20.2)</td>
<td>5519 (18.1)</td>
<td>&lt; 0.01</td>
</tr>
</tbody>
</table>

Note: NSAIDs = nonsteroidal anti-inflammatory drugs, COX-2 = cyclooxygenase-2, SD = standard deviation, GI = gastrointestinal.

*Unless stated otherwise.