randomized controlled trials (RCTs) have clearly demonstrated that cholesterol-lowering statins are beneficial in the treatment and prevention of coronary heart disease. Although recent evidence indicates that major RCTs of statins have significantly influenced physicians’ prescribing behaviour,1 it is uncertain whether physicians have interpreted the results as a class-specific or a drug-specific effect. We investigated whether the release of 3 landmark RCTs of statins, namely, the Scandinavian Simvastatin Survival Study (4S),2 the West of Scotland (WOS) trial3 and the Cholesterol and Recurrent Events (CARE)4 trial, and the release of atorvastatin were associated with the market share of different statins in Canada.

Prescribing patterns were examined for July 1993 through June 1999, with the aim of identifying any immediate changes following the release of the results of the RCTs and the release of atorvastatin. An immediate change was defined as a significant shift in prescribing pattern (i.e., changes in the monthly proportion of prescriptions attributable to each statin within the class) from projected estimates within 5 months of the intervention being assessed. Although outcome-based studies examining atorvastatin had not been published during this observation period, atorvastatin had been shown to improve secondary end points, such as low-density lipoprotein levels, to a greater extent than the other drugs in its class.1 National estimates for monthly statin prescriptions dispensed in Canada were obtained from the Compuscript database of IMS Health Canada, which samples approximately two-thirds of all retail pharmacies in Canada. Time series analysis6 was conducted using interventional autoregressive integrated moving average (ARIMA) models to determine the impact of each intervention on the market share of the different statins.

The monthly number of statin prescriptions dispensed rose by 236% from about 166 000 in July 1993 to about 557 000 in June 1999. The release of the 4S study2 was associated with an observed positive shift in the market share of simvastatin ($p = 0.04$), an observed negative shift in the trend of pravastatin ($p = 0.03$) and no immediate changes in the market shares of fluvastatin ($p = 0.77$) and lovastatin ($p = 0.13$) (Fig. 1).

In contrast, the WOS study was associated with an observed positive shift in the market share of pravastatin ($p < 0.01$) and no significant observed changes in the market shares of fluvastatin ($p = 0.39$), lovastatin ($p = 0.65$) and simvastatin ($p = 0.81$) (Fig. 1).

The CARE study was associated with an observed positive shift in the market share of pravastatin ($p = 0.01$), a negative shift in simvastatin ($p = 0.01$) and no significant changes in the market shares of fluvastatin ($p = 0.46$) and lovastatin ($p = 0.90$) (Fig. 1).

The release of atorvastatin was associated with more gradual negative shifts in the market shares of simvastatin ($p = 0.01$) and lovastatin ($p < 0.01$) and a more immediate negative shift in the market share of pravastatin ($p = 0.04$). No significant effects were observed in the market share of fluvastatin ($p = 0.27$). The market share of atorvastatin increased from 0.9% in April 1997 to a top position of 39.2% in June 1999 ($p < 0.01$).

Although the overall prescribing of statins in Canada has increased dramatically over time, the publication of each major RCT was associated with a significant positive shift in the market share of the statin featured in each trial. These results suggest that physicians may have initially subscribed to a drug-specific rather than a class-effect belief when prescribing statins. A major limitation of this study was the lack of information about promotional expenditures and methods.7,8 These factors may have also accounted for changes in market share. The release of atorvastatin was associated with a relatively gradual, yet pronounced, shift in prescribing practice, which may in part be a reflection of competitive pricing or of favourable findings from clinical trials1 measuring surrogate end points, such as laboratory values. Drug prescribing is influenced by numerous factors. Our findings suggest that the
results of the landmark clinical trials of statins and the introduction of new agents may be among these factors.

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