

Secondary Prevention of Ischemic Heart Disease: Closing the Gap

Menahem Fainaru MD

Medical Division, Maccabi Healthcare Services, Israel
Sackler Faculty of Medicine, Tel Aviv University, Ramat Aviv, Israel

Key words: ischemic heart disease, secondary prevention, statins, low density lipoprotein-cholesterol

IMAJ 2005;7:400–401

Compelling evidence has accumulated during the last few decades on ways to prevent or delay the occurrence of atherosclerotic cardiovascular disease and its complications. Big strides have been made in basic and clinical research towards prolonging life and improving its quality, enabling early identification and effective treatment of risk factors for cardiovascular disease and coronary heart disease in particular. These advances led, in the last 30 years of the 20th century, to an increase in life expectancy of 4 years in the United States, attributable to improvements in prevention and treatment of complications in patients suffering from cardiovascular disease [1]. One, if not the most important, approach that led to this achievement is the lowering of low density lipoprotein-cholesterol by HMG-CoA reductase inhibitors – namely statins, that have been available since the early 1980s. Several seminal randomized controlled studies have been conducted and published during the last decade, establishing beyond doubt the effectiveness of such treatment in both primary [2,3] and secondary prevention [4] of cardiovascular disease. Indeed, it is incomprehensible that it took so many years for them to be implemented. Dr. Claude Lenfant, former director of the National Heart Lung and Blood Institutes, in his Shattuck lecture: “Clinical Research to Clinical Practice – Lost in Translation?”, published in the *New England Journal of Medicine*, concentrates on this failure in translating sound research findings into medical practice and personal behavior [1]. He cites the U.S. National Committee for Quality Assessment report of 1999, which showed that only half to three-quarters of patients who had myocardial infarction were even being screened for elevated blood cholesterol levels, much less prescribed cholesterol-lowering agents. In the Israeli HOLEM study, conducted during the same year and published in this issue of IMAJ, the findings were similar [5].

These observations are supported also by studies conducted during the same period in nine European countries (EUROASPIRE II 1999-2000) [6], and by the U.S.-based HER study [7], which reported similar low rates of adherence to these guidelines, 63% and 53% respectively.

Since the HOLEM study, which was conducted during the late 1990s and early 2000s, new knowledge has been added, liberalizing the indication for statin treatment by lowering the threshold [8], starting therapy earlier [9], and intensifying the dose to achieve lower target values (below 70 mg/dl) [10,11]. In addition, the importance of the pleiotropic effects of statins has become evident, namely lowering inflammatory markers such as

C-reactive protein [12–14], strengthening further the indication for using statins to treat most, if not all, patients suffering from ischemic heart disease.

The main reasons for the existing gap between evidence-based clinical investigations and clinical practice reside in both the healthcare providers and the patients. It is the responsibility of the medical profession to ensure that patients are the beneficiaries of the tremendous progress being made in medicine today.

Although the situation has improved since the reports from the late 1990s, both worldwide and in Israel [15,16], much remains to be done. The management of acute coronary syndrome in the Israeli survey (ACSIS) has been surveyed prospectively by the Israel Heart Society and recorded bi-annually in all departments of cardiology in Israel since the early 1990s. Between 2000 and 2002, lipid-lowering therapy for patients at discharge increased from 47% to 65% respectively. These results show a marked improvement but are still far from the expected optimum, since early and intensive statin therapy results in early reduction in adverse cardiac events in patients with acute coronary syndromes, which is sustained over the years. These benefits result from a combination of early pleiotropic effects (modulation of inflammation, endothelial function, coagulation) and long-term effects on the atheroma size. Therefore, it is the responsibility of physicians in the hospitals (cardiologists and internists) and in the community (primary care physicians, the medical departments of the health maintenance organizations), as well as the Ministry of Health, to implement quality assurance measures, adhere to guidelines, and increase the prescription rate of statins in both secondary and primary prevention of cardiovascular disease. These should include measures to increase the compliance of physicians with established and available clinical practice guidelines, as well as the translation of the new knowledge into guidelines by policy makers and managers of the healthcare system.

In addition, it is the responsibility of the leading Israeli medical professional associations, such as the Israel Heart Society, to lead the way in translating new knowledge into practice guidelines. Consequently, it is the responsibility of healthcare organizations, including the Ministry of Health, to implement quality assurance surveillance to assess the quality of care and service, as was initiated last year by the Israel Ministry of Health (National Quality Measures). It is their joint responsibility to promote and assure that physicians abide and comply with these guidelines. Finally, it is the responsibility of all healthcare providers, including the

HMOs, to ensure that patients comply with the prescribed medication regimens by implementing health promotion programs.

In conclusion, in order to exploit the opportunities provided by novel biomedical research and to fulfill our patients' expectations, it is our obligation to initiate statin therapy as soon as possible and at the appropriate dosage to achieve the full benefit of this highly effective and miraculous drug, particularly for patients suffering from acute coronary syndromes [17].

References

1. Lenfant C. Shattuck Lecture: Clinical research to clinical practice – Lost in translation? *N Engl J Med* 2003;349:868–74.
2. Shepherd J, Cobbe SM, Ford I, et al. Prevention of coronary heart disease with pravastatin in men with hypercholesterolemia. West of Scotland Coronary Prevention Study Group. *N Engl J Med* 1995;333:1301–7.
3. Downs JR, Clearfield M, Weis S, et al. Primary prevention of acute coronary events with lovastatin in men and women with average cholesterol levels: results of AFCAPS/TexCAPS. Air Force/Texas Coronary Atherosclerosis Prevention Study. *JAMA* 1998;279:1615–22.
4. Scandinavian Simvastatin Survival Study Group. Randomised trial of cholesterol lowering in 4444 patients with coronary heart disease: the Scandinavian Simvastatin Survival Study (4S). *Lancet* 1994;344:1383–9.
5. Harats D, Leibovitz E, Maislos M, et al. Cardiovascular risk assessment and treatment to target low density lipoprotein levels in hospitalized ischemic heart disease patients: results of the HOLEM study. *IMAJ* 2005;7:355–9.
6. EUROASPIRE I and II group. Clinical reality of coronary prevention: a comparison of EUROASPIRE I and II in nine countries. *Lancet* 2001;357:995–1001.
7. Vittinghoff E, Shlipak MG, Varosy PD, et al. Risk factors and secondary prevention in women with heart disease: the Heart and Estrogen Progestin Replacement Study. *Ann Intern Med* 2003;138(2):81–9.
8. Heart Protection Study Collaborative Group. MRC/BHF Heart protection study of cholesterol lowering with simvastatin in 20,536 high-risk individuals: a randomised placebo-control trial. *Lancet* 2002;360:7–22.
9. de Lemos JA, Blazing MA, Wiviott SD, et al. Early intensive vs. a delayed conservative simvastatin strategy in patients with acute coronary syndromes. *JAMA* 2004;292(11):1307–16.
10. Cannon CP, Braunwald E, McCabe CH, et al. Intensive versus moderate lipid lowering with statins after acute coronary syndromes. *N Engl J Med* 2004;350:1495–504.
11. Grundy SM, Cleeman JJ, Merz NB, et al. Implications of recent clinical trials for the National Cholesterol Education Program Adult Treatment Panel III Guidelines. *Circulation* 2004;110:227–39.
12. Schonbeck U, Libby P. Inflammation, immunity and HMG-CoA reductase inhibitors: statins as antiinflammatory agents? [Review]. *Circulation* 2004;109(Suppl II):II18–26.
13. Ridker PM, Cannon CP, Morrow D, et al. C-reactive protein levels and outcomes after statin therapy. *N Engl J Med* 2005;352(1):20–8.
14. Davignon J. Beneficial cardiovascular pleiotopic effects of statins. *Circulation* 2004;109(Suppl III):III39–43.
15. Behar S, Battler A, Porath A, et al. A prospective national survey of management and clinical outcome of acute myocardial infarction in Israel, 2000. *IMAJ* 2003;5(4):249–54.
16. ACSIS 2002. Acute coronary syndromes – Israel 2002. Publication no. 230. ICDC Israel Center for Disease Control. April 2003.
17. Ray KK, Cannon CP. Intensive statin therapy in acute coronary syndromes: clinical benefits and vascular biology. *Curr Opin Lipidol* 2004;15:637–43.

Correspondence: Dr. M. Fainaru, Medical Director, Maccabi Healthcare Services, 27 Hamered Street, Tel Aviv 68125, Israel.
 Phone: (972-3) 514-3508
 Fax: (972-3) 514-3988
 email: mfainar@mac.org.il