Low prevalence of lipid lowering drug use in older men with established coronary heart disease

P H Whincup, J R Emberson, L Lennon, M Walker, O Papacosta, A Thomson

Objective: To determine the prevalence and correlates of lipid lowering drug use among older British men with established coronary heart disease (CHD).

Design: Cross sectional survey within a cohort study (British regional heart study) carried out at 20 years of follow up in 1998–2000.

Setting: General practices in 24 British towns.

Participants: 3689 men aged 60–75 years (response rate 76%).

Main outcome measures: Diagnoses of myocardial infarction and angina based on detailed review of general practice records. Lipid lowering drug use and blood cholesterol concentrations ascertained at 20 year follow up examination.

Results: Among 286 men with definite myocardial infarction, 102 (36%) were taking a lipid lowering drug (93 [33%] a statin); among 360 men with definite angina without myocardial infarction, 84 (23%) were taking a lipid lowering drug (78 [21%] a statin). Most men with documented CHD who were not receiving a lipid lowering drug had a total cholesterol concentration of 5.0 mmol/l or more (87% of those with myocardial infarction, 82% with angina). Fewer than half of men with CHD receiving a statin had a total cholesterol concentration below 5.0 mmol/l [45% of those with myocardial infarction and 47% of those with angina]. Only one third of the men taking a statin were receiving trial validated dosages. Among men with CHD, a history of revascularisation, more recent diagnosis, and younger age at diagnosis were associated with a higher probability of receiving lipid lowering drug treatment.

Conclusion: Among patients with established CHD, the prevalence of lipid lowering drug use remains low and statin regimens suboptimal. Major improvements in secondary prevention are essential if the benefits of statins are to be realised.
general practice records. Fewer than 1% of participants have been lost to follow up.

Between February 1998 and February 2000, all surviving men were invited to attend for a 20 year follow up at 60–79 years of age. Men who were still living in the same town as in 1978–1980 were invited to return to their original practice for reassessment; men who had moved from the original study town were given a choice of returning to their original town, of being measured in another (more convenient) town, or of travelling to a London assessment centre. Men were asked to attend for screening after a minimum six hour fast and with all their medications. At screening, men completed a study questionnaire, which was checked by research nurses. All current medications were recorded and verified either against the medication container or (if the container was not available) against a prescription record; any participant in whom the completeness of the medication record was uncertain was excluded from this analysis. Diabetes diagnosed by a doctor and current smoking habit, including number of cigarettes smoked per day, were recorded. Two seated blood pressure measurements were made with the Dinamap 1846 oscillometric blood pressure recorder (Critikon, Tampa, Florida, USA); overreading of systolic pressure by the instrument was corrected. A 12 lead ECG was recorded and analysed in accordance with Minnesota criteria (Professor P Macfarlane, Glasgow University) and a blood sample was taken for measurement of total cholesterol, using a Hitachi 747 (Hitachi, Tokyo, Japan) automated autoanalyser (Professor A F Winder, Department of Chemical Pathology, Royal Free Hospital).

In July 2000, the general practice records of all participants were reviewed in detail to identify all diagnoses and procedures related to CHD such as myocardial infarction, angina, coronary angiography, coronary artery bypass graft surgery, and percutaneous transluminal coronary angioplasty, including dates, that had occurred up to the time of examination. Before a practice diagnosis of myocardial infarction was accepted as definite, additional clinical information on standard World Health Organization criteria (requiring two of the following: (a) a history of severe chest pain, (b) changes in cardiac enzymes, and (c) characteristic electrocardiographic changes) was obtained from the general practitioner. Among patients without CHD diagnosed, absolute CHD risk was calculated by the standard Framingham equation, using age, diabetic status, systolic pressure, smoking status, total and high density lipoprotein cholesterol, and left ventricular hypertrophy.

### RESULTS

In all, 3689 men aged 60–75 years participated in the 20 year follow up examination and provided a full medication history (76% of those invited). Almost all (3495, 95%) had a study total cholesterol measurement. A total of 294 men (8.0%) were taking lipid lowering medication; of these 259 (88%) were taking statins, 31 (11%) fibrates, and 4 (1%) ion exchange resins. Of these 294 men, 208 (71%) had cardiovascular disease documented in their general practice records and a further 48 (16%) recalled a cardiovascular diagnosis.

#### Lipid lowering drug use in men with documented CHD

In all, 286 patients (7.8%) had definite myocardial infarction diagnosed, of which 33 (12%) were recurrent. Three hundred and sixty patients (9.8%) had angina diagnosed, of which 117 (33%) were recurrent. Men who had both of these conditions diagnosed were included in the myocardial infarction group. Table 1 shows the prevalences of lipid lowering drug use...
Control of blood total cholesterol among men with documented CHD receiving lipid lowering treatment

Among 186 men with CHD diagnosed who were on any lipid lowering treatment, 44% had a total cholesterol concentration below 5 mmol/l; this proportion was similar both among men with myocardial infarction and angina. When this analysis was restricted to 171 men with CHD diagnosed who were receiving statin treatment, 46% had a total cholesterol concentration below 5 mmol/l (45% of those with myocardial infarction, 47% of those with angina). Of patients with documented CHD and a presumed baseline total cholesterol of 5 mmol/l or above, 82% had a total cholesterol concentration of 5 mmol/l or more and 60% had one above 5.5 mmol/l.

Factors related to use of lipid lowering drug treatment in men with CHD

Table 1 presents factors related to the use of lipid lowering drug treatment in men with myocardial infarction or angina. A history of coronary revascularisation was related to increased lipid lowering drug use in both groups of men. Year of last diagnosis was strongly related to drug use in men with myocardial infarction, with prevalence of treatment being much higher among recently diagnosed cases; results of the year of first diagnosis (data not presented) were similar. Prevalence of lipid lowering drug use appeared to be slightly (though not significantly) higher in patients who were younger at last diagnosis, in patients who were of non-manual social class, and among men studied in Scotland and southern England compared with those in the Midlands and the north of England. Men who were cigarette smokers were slightly though not significantly less likely than non-smokers to receive lipid lowering drugs. Table 2 shows odds ratios after adjustment for these interrelated factors. For patients with either myocardial infarction or angina, a history of previous coronary revascularisation, younger age at last diagnosis, and more recent date of last diagnosis were strongly related to increased lipid lowering drug use. The weak and non-significant relation between social class, region, cigarette smoking, and lipid lowering drug use were little altered by adjustment for these other factors.

Lipid lowering drug use among men at high risk of CHD

Among 2868 men with no CHD diagnosis in whom a Framingham risk score could be calculated, 578 (20%) had a Framingham risk score consistent with an absolute CHD risk of 3.0% per year or more. However, of these men only 19 (3%) were receiving lipid lowering drug treatment of any kind.

DISCUSSION

These results suggest that most older men with established CHD are not receiving lipid lowering drug treatment, and that even those who receive treatment are in many cases receiving inadequate drug doses. The CHD ascertainment methods in this investigation were used specifically to avoid falsely low prevalence estimates of lipid lowering drug use. Cases in which there was any doubt about the completeness of the medication record were excluded. The CHD diagnoses on
which analyses were based were all known to the general practitioners and did not depend on patient recall, which is less specific than a diagnosis based on general practice records. Although the long term involvement of participating practices in the British regional heart study could have affected lipid lowering drug use in this study population, participation might have been expected to increase (rather than decrease) preventive activities, including prescription of lipid lowering drugs. However, the study has maintained a low profile in the participating practices and avoided making any specific recommendations about treatment practices to the general practitioners involved in the study. The prevalence of lipid lowering drug use in patients with established CHD (29%) is very similar to the estimates provided by Primatesa and Poulter in men aged 65–74 years in the Health Survey for England (25%) and from general practice statistics in men aged 65–74 years (29%). The overall proportion of patients with increased blood cholesterol concentrations controlled by lipid lowering drugs is also very similar to the estimate of 16% in the Health Survey for England. The present results show clearly that a large proportion of patients are taking inadequate doses of statins, in most cases well below the dosages shown to be effective in the randomised controlled trials of secondary prevention. Although these findings are only for men, recent studies suggest that the extent of secondary prevention in women is less complete. The lower rates of lipid lowering drug use in older patients in the present study (a group at particularly high CHD risk and who, up to 75 years of age, are known to benefit from statins) are also consistent with earlier reports. Our findings that the prevalence of lipid lowering drug use is lower among patients with angina than those with myocardial infarction, among those who have not undergone revascularisation, and among those whose condition was diagnosed less recently, are consistent with our earlier report on the use of aspirin in the secondary prevention of cardiovascular disease and with European experience. Our results are also consistent with the possibility of an adverse social gradient in lipid lowering drug use, though they also suggest that there is a particularly high prevalence of lipid lowering drug use in Scotland; this may result from the high local prevalence of lipid lowering drug use that there is particular interest in following treatment in this study (older patients, those with relatively longstanding CHD, particularly angina, and those not receiving invasive treatment) are likely to receive most or all of their clinical care in the primary care setting and will only be identified by systematic retrospective identification and management. The low current levels of lipid lowering drug use suggest that a strongly proactive approach to implementation of secondary prevention, both prospectively and retrospectively, will be essential. This will need to be based on clear guidance, giving appropriate priority to secondary prevention in relation to primary prevention, and providing clear recommendations on treatment regimens to ensure that patients receive optimal benefit. However, since the cost effectiveness of statin treatment for secondary prevention of CHD is greater than that of many procedures currently performed under the NHS, such action would be amply justified.

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Images in Cardiology

Intercoronary communication between the circumflex and right coronary arteries: distinct from coronary collaterals

A 49 year old man presented with atypical chest pain for five months. He had no risk factors for atherosclerosis. Exercise electrocardiography was stopped prematurely because of leg cramp. Coronary angiography was performed and revealed normal left ventricular function. Left and right coronary angiography did not show any evidence of luminal narrowing or occlusion of either coronary artery (below left). However, with the selective injection of the right coronary artery (RCA) in a left anterior oblique view, the RCA and distal and mid portion of left circumflex artery (CX) were simultaneously visualised. An intercoronary connection was also seen near the crux (below centre). The RCA and CX were then catheterised at the same time. There was no pressure damping during selective placement of the catheter tip in either coronary ostium. In the left anterior oblique view with cranial angulation, simultaneous injection of contrast material into both coronary ostia revealed interarterial continuity between the RCA and CX (below right: arrowheads show both antegrade and retrograde filling of CX lumen).

Intercoronary artery continuity or “coronary cascade” is a rare variant of the coronary circulation. Two types have been described: communication between the CX and the RCA in the posterior atrioventricular groove (as in this patient), and communication between left anterior descending and posterior descending artery in the distal interventricular groove. Compared with collaterals, intercoronary arterial connections are larger in diameter (> 1 mm), extramural, and straight. Furthermore, the structure of an intercoronary arterial connection is typical of an epicardial coronary artery, with a well defined muscular layer. Intercoronary arterial connections are thought to be congenital in origin. It is suggested that faulty embryological development allows the existing intercoronary channel to remain prominent and maintain a large calibre.

In our case selective injection of the left coronary artery did not demonstrate retrograde filling of the distal RCA. One may propose that both the force of injection and the velocity of flow in the left coronary artery would account for this discrepancy.

True prevalence of this entity is unknown but we identified only one patient among the 7086 angiograms performed at our laboratory during the last 10 years. The functional significance of this large anastomotic connection between normal coronary arteries is unclear but one may speculate that they have a potential role in protecting the myocardium should significant atherosclerosis develop in either of the parent arteries.

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